

Cochrane Database of Systematic Reviews

Exercise training for adults undergoing maintenance dialysis (Review)

Bernier-Jean A, Beruni NA, Bondonno NP, Williams G, Teixeira-Pinto A, Craig JC, Wong G	

Bernier-Jean A, Beruni NA, Bondonno NP, Williams G, Teixeira-Pinto A, Craig JC, Wong G. Exercise training for adults undergoing maintenance dialysis. *Cochrane Database of Systematic Reviews* 2022, Issue 1. Art. No.: CD014653. DOI: 10.1002/14651858.CD014653.

www.cochranelibrary.com



TABLE OF CONTENTS

ABSTRACT	1
PLAIN LANGUAGE SUMMARY	2
SUMMARY OF FINDINGS	4
BACKGROUND	12
OBJECTIVES	12
METHODS	12
RESULTS	15
Figure 1	
Figure 2	
DISCUSSION	
AUTHORS' CONCLUSIONS	
ACKNOWLEDGEMENTS	
REFERENCES	
CHARACTERISTICS OF STUDIES	44
DATA AND ANALYSES	
Analysis 1.1. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 1: Death	
Analysis 1.2. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 2: Fatigue	
Analysis 1.3. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 3: HRQoL: Summary	
component scores	
Analysis 1.4. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 4: HRQoL: Individual domains	222
Analysis 1.5. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 5: Depression	226
Analysis 1.6. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 6: 6MWT	227
Analysis 1.7. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N reps/30 sec]	227
Analysis 1.8. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 8: Sit-To-Stand test [sit to 5	228
reps]	
Analysis 1.9. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 9: Systolic blood pressure	228
Analysis 1.10. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 10: Diastolic blood pressure	229
Analysis 1.11. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 11: Aerobic capacity (VO max or peak)	
Analysis 1.12. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 12: Albumin	230
Analysis 1.13. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 13: Blood lipids	231
Analysis 1.14. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 14: Body composition	232
Analysis 1.15. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 15: Body mass index	232
Analysis 1.16. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 16: Calcium	233
Analysis 1.17. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 17: C-reactive protein	233
Analysis 1.18. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 18: Dialysis adequacy: Kt/V .	234
Analysis 1.19. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 19: Energy intake	234
Analysis 1.20. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 20: Haemoglobin	235
Analysis 1.21. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 21: Left ventricular ejection fraction	235
Analysis 1.22. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 22: Left ventricular mass	236
index	226
Analysis 1.23. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 23: Maximum heart rate	236
Analysis 1.24. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 24: Muscular strength	237
Analysis 1.25. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 25: Phosphate	238
Analysis 1.26. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 26: Potassium	238
Analysis 1.27. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 27: Protein intake	239
Analysis 1.28. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 28: Parathyroid hormone	239
Analysis 1.29. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 29: Resting heart rate	239
Analysis 1.30. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 30: Timed up-and-go test	240



Analysis 2.1. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 1: Death
Analysis 2.2. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 2: Fatigue
Analysis 2.3. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 3: HRQoL: Summary
component scores
Analysis 2.4. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 4: HRQoL: Individual domains
Analysis 2.5. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 5: Depression
Analysis 2.6. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 6: 6MWT
Analysis 2.7. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N
reps/30 sec]
Analysis 2.8. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 8: Sit-To-Stand test [sit to 5 reps]
Analysis 2.9. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 9: Resting blood pressure
Analysis 2.10. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 10: Aerobic capacity (VO2 max or peak)
Analysis 2.11. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 11: Albumin
Analysis 2.12. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 12: Blood lipids
Analysis 2.13. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 13: Body composition
Analysis 2.14. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 14: Body mass index
Analysis 2.15. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 15: Calcium
Analysis 2.16. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 16: C-reactive protein
Analysis 2.17. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 17: Dialysis adequacy: Kt/V
Analysis 2.18. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 18: Energy intake
Analysis 2.19. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 19: Haemoglobin
Analysis 2.20. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 20: Heart rate
Analysis 2.21. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 21: Left ventricular
ejection fraction
Analysis 2.22. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 22: Left ventricular mass index
Analysis 2.23. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 23: Muscular strength
Analysis 2.24. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 24: Phosphate
Analysis 2.25. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 25: Potassium
Analysis 2.26. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 26: Protein intake
Analysis 2.27. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 27: Parathyroid
hormone
Analysis 2.28. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 28: Timed up-and-go test
Analysis 3.1. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 1: Fatigue
Analysis 3.2. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 2: HRQoL: Summary component scores
Analysis 3.3. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 3: HR-QoL: Individual domains
Analysis 3.4. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 4: Depression
Analysis 3.5. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 5: 6MWT
Analysis 3.6. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 6: Sit-To-Stand test [N
reps/30 sec]
Analysis 3.7. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N reps/30 sec]
Analysis 3.8. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 8: Albumin
Analysis 3.9. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 9: Blood lipids
Analysis 3.10. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 10: Body composition
Analysis 3.11. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 11: Body mass index



Analysis 3.12. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 12: Calcium	26
Analysis 3.13. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 13: CRP	26
Analysis 3.14. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 14: Dialysis adequacy: Kt/V	26
Analysis 3.15. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 15: Energy intake	268
Analysis 3.16. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 16: Haemoglobin	268
Analysis 3.17. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 17: Muscular strength	26
Analysis 3.18. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 18: Phosphate	26
Analysis 3.19. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 19: Potassium	27
Analysis 3.20. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 20: Protein intake	27
Analysis 3.21. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 21: PTH	27
Analysis 3.22. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 22: Timed up-and-go test	27
Analysis 4.1. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 1: HRQoL: Summary component scores	27
Analysis 4.2. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 2: HRQoL: Individual domains	27
Analysis 4.3. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 3: Depression	27
Analysis 4.4. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 4: 6MWT	27
Analysis 4.5. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 5: Sit-To-Stand test [N reps/30 sec]	27
Analysis 4.6. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 6: Sit-To-Stand test [sit to 5 reps]	27
Analysis 4.7. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 7: Resting blood pressure	27
Analysis 4.8. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 8: Aerobic capacity (VO2 max or peak)	27
Analysis 4.9. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 9: Albumin	27
Analysis 4.10. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 10: Blood lipids	27
Analysis 4.11. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 11: Body composition	27
Analysis 4.12. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 12: Body mass index	28
Analysis 4.13. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 13: Calcium	28
Analysis 4.14. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 14: CRP	28
Analysis 4.15. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 15: Dialysis adequacy: Kt/V	28
Analysis 4.16. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 16: Energy intake	28
Analysis 4.17. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 17: Haemoglobin	28
Analysis 4.18. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 18: Heart rate	28
Analysis 4.19. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 19: Muscular strength	28
Analysis 4.20. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 20: Phosphate	28
Analysis 4.21. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 21: Potassium	28



Analysis 4.22. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 22: Protein intake	282
Analysis 4.23. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 23: Timed up-and-go test	283
APPENDICES	283
HISTORY	298
CONTRIBUTIONS OF AUTHORS	298
DECLARATIONS OF INTEREST	298
SOURCES OF SUPPORT	298
DIFFERENCES BETWEEN PROTOCOL AND REVIEW	298
INDEX TERMS	299



[Intervention Review]

Exercise training for adults undergoing maintenance dialysis

Amelie Bernier-Jean^{1,2}, Nadim A Beruni³, Nicola P Bondonno^{4,5}, Gabrielle Williams², Armando Teixeira-Pinto⁶, Jonathan C Craig^{7,8}, Germaine Wong²

¹School of Public Health, Faculty of Medicine and Health, The University of Sydney, Sydney, Australia. ²Centre for Kidney Research, The Children's Hospital at Westmead, Westmead, Australia. ³Resident Support Unit, Western Sydney Local Health District, Westmead, Australia. ⁴School of Biomedical Sciences, The University of Western Australia, Royal Perth Hospital, Perth, Australia. ⁵School of Medical and Health Sciences, Edith Cowan University, Perth, Australia. ⁶Sydney School of Public Health, The University of Sydney, Sydney, Australia. ⁷College of Medicine and Public Health, Flinders University, Adelaide, Australia. ⁸Cochrane Kidney and Transplant, Centre for Kidney Research, The Children's Hospital at Westmead, Westmead, Australia

Contact: Amelie Bernier-Jean, ameliebjean@gmail.com.

Editorial group: Cochrane Kidney and Transplant Group. **Publication status and date:** New, published in Issue 1, 2022.

Citation: Bernier-Jean A, Beruni NA, Bondonno NP, Williams G, Teixeira-Pinto A, Craig JC, Wong G. Exercise training for adults undergoing maintenance dialysis. *Cochrane Database of Systematic Reviews* 2022, Issue 1. Art. No.: CD014653. DOI: 10.1002/14651858.CD014653.

Copyright © 2021 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

ABSTRACT

Background

Dialysis treatments weigh heavily on patients' physical and psychosocial health. Multiple studies have assessed the potential for exercise training to improve outcomes in adults undergoing dialysis. However, uncertainties exist in its relevance and sustainable benefits for patient-important outcomes. This is an update of a review first published in 2011.

Objectives

To assess the benefits and safety of regular structured exercise training in adults undergoing dialysis on patient-important outcomes including death, cardiovascular events, fatigue, functional capacity, pain, and depression. We also aimed to define the optimal prescription of exercise in adults undergoing dialysis.

Search methods

In this update, we conducted a systematic search of the Cochrane Kidney and Transplant Register of Studies up to 23 December 2020. The Register includes studies identified from CENTRAL, MEDLINE, EMBASE, the International Clinical Trials Register (ICTRP) Search Portal and ClinicalTrials.gov as well as kidney-related journals and the proceedings of major kidney conferences.

Selection criteria

Randomised controlled trials (RCTs) and quasi-RCTs of any structured exercise programs of eight weeks or more in adults undergoing maintenance dialysis compared to no exercise or sham exercise.

Data collection and analysis

Two authors independently assessed the search results for eligibility, extracted the data and assessed the risk of bias using the Cochrane risk of bias tool. Whenever appropriate, we performed random-effects meta-analyses of the mean difference in outcomes. The primary outcomes were death (any cause), cardiovascular events and fatigue. Secondary outcomes were health-related quality of life (HRQoL), depression, pain, functional capacity, blood pressure, adherence to the exercise program, and intervention-related adverse events.



Main results

We identified 89 studies involving 4291 randomised participants, of which 77 studies (3846 participants) contributed to the meta-analyses. Seven studies included adults undergoing peritoneal dialysis. Fifty-six studies reported aerobic exercise interventions, 21 resistance exercise interventions and 19 combined aerobic and resistance training within the same study arm. The interventions lasted from eight weeks to two years and most often took place thrice weekly during dialysis treatments. A single study reported death and no study reported long-term cardiovascular events. Five studies directly assessed fatigue, 46 reported HRQoL and 16 reported fatigue or pain through their assessment of HRQoL. Thirty-five studies assessed functional capacity, and 21 reported resting peripheral blood pressure. Twelve studies reported adherence to exercise sessions, and nine reported exercise-related adverse events. Overall, the quality of the included studies was low and blinding of the participants was generally not feasible due to the nature of the intervention.

Exercise had uncertain effects on death, cardiovascular events, and the mental component of HRQoL due to the very low certainty of evidence. Compared with sham or no exercise, exercise training for two to 12 months may improve fatigue in adults undergoing dialysis, however, a meta-analysis could not be conducted. Any exercise training for two to 12 months may improve the physical component of HRQoL (17 studies, 656 participants: MD 4.12, 95% CI 1.88 to 6.37 points on 100 points-scale; I² = 49%; low certainty evidence). Any exercise training for two to 12 months probably improves depressive symptoms (10 studies, 441 participants: SMD -0.65, 95% CI -1.07 to -0.22; I² = 77%; moderate certainty evidence) and the magnitude of the effect may be greater when maintaining the exercise beyond four months (6 studies, 311 participants: SMD -0.30, 95% CI 0.14 to -0.74; I² = 71%). Any exercise training for three to 12 months may improve pain (15 studies, 872 participants: MD 5.28 95% CI -0.12 to 10.69 points on 100 points-scale; I² = 63%: low certainty evidence) however, the 95% CI indicates that exercise training may make little or no difference in the level of pain. Any exercise training for two to six months probably improves functional capacity as it increased the distance reached during six minutes of walking (19 studies, 827 participants: MD 49.91 metres, 95% CI 37.22 to 62.59; I² = 34%; moderate certainty evidence) and the number of sit-to-stand cycles performed in 30 seconds (MD 2.33 cycles, 95% CI 1.71 to 2.96; moderate certainty evidence). There was insufficient evidence to assess the safety of exercise training for adults undergoing maintenance dialysis. The results were similar for aerobic exercise, resistance exercise, and a combination of both aerobic and resistance exercise.

Authors' conclusions

It is uncertain whether exercise training improves death, cardiovascular events, or the mental component of HRQoL in adults undergoing maintenance dialysis. Exercise training probably improves depressive symptoms, particularly when the intervention is maintained beyond four months. Exercise training is also likely to improve functional capacity. Low certainty evidence suggested that exercise training may improve fatigue, the physical component of quality of life, and pain. The safety of exercise training for adults undergoing dialysis remains uncertain.

PLAIN LANGUAGE SUMMARY

Exercise training for adults receiving dialysis treatments

What is the issue?

People undergoing dialysis treatments are at higher risk of cardiovascular disease and depression, have a lower quality of life and limited survival than the general population. Furthermore, many people undergoing dialysis have difficulty performing daily activities because they lack the physical capacity and strength to do so. Multiple trials have assessed the potential for exercise training to improve the condition of adults undergoing dialysis, but no consensus has been reached.

What did we do?

We searched the medical literature for all randomised trials that assessed structured exercise programs in people undergoing dialysis. We then assessed the quality of those studies and combined their results to draw conclusions regarding the effect of exercise training to improve aspects of physical and mental health that are important to patients undergoing dialysis.

What did we find?

We found 89 studies involving 4291 participants. The exercise training programs lasted from eight weeks to two years and most often took place three times a week during the dialysis treatment. We could not determine the impact of exercise training on death, cardiovascular events (such as a heart attack) or mental well-being. Moderate certainty evidence suggested that exercise training of any type is likely to improve depressive symptoms in adults undergoing dialysis, particularly when the exercise was maintained for longer than four months. Moderate quality evidence also suggested that exercise training may improve people's capacity to perform activities and tasks through the improvement of their capacity to walk and the strength and endurance of their legs. Exercise training may also improve fatigue and the physical aspects of quality of life, but the quality of the evidence was low. We could not conclude on the effect of exercise training on a person's mental well-being.

Conclusions

Exercise training for people undergoing maintenance dialysis is likely to improve depression and their capacity to perform activities and tasks. Exercise training may also improve fatigue and pain sightly. Exercise training may improve the physical aspects of quality of life,



but it is unclear whether it improves a person's mental well-being. It is unclear whether exercise training reduces the number of deaths or cardiovascular events.



Summary of findings 1. Any exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Any exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Patient or population: adults undergoing maintenance dialysis

Setting: all settings (e.g. during dialysis, pre- and post-dialysis; home exercise)

Intervention: any exercise

Comparison: no exercise or placebo exercise

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	No. of partici- pants	Certainty of the evidence	Comments
	Risk with no ex- ercise or place- bo exercise	Risk with any exercise	(35 % 5.1)	(studies)	(GRADE)	
Death (any cause Follow up: 3 years	159 per 1,000	151 per 1,000 (89 to 257)	RR 0.95 (0.56 to 1.62)	296 (1)	⊕⊝⊝⊝ VERY LOW ¹²³	-
Cardiovascular events	Not reported	Not reported	-	-	-	-
Fatigue Follow up: range 2 to 12 months	See comment	See comment	-	326 (6)	⊕⊕⊙⊝ LOW 47	A pooled estimate of the effect was not calculated because the included studies assessed different dimensions of fatigue. Based on the direction of the effect in the included studies, any exercise may reduce fatigue
HRQoL: Physical component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean physical component score ranged from 34 to 74 points	The mean physical component score was 4.1 points higher with exercise (1.9 to 6.4 higher)	-	656 (17)	⊕⊕⊝⊝ LOW 45	Any exercise may improve the physical component score of HRQoL
HRQoL: Mental component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean mental component score ranged from 38 to 76 points	The mean mental component score was 2.5 points higher with exercise (0.4 lower to 5.5 higher)	-	656 (17)	⊕⊝⊝⊝ VERY LOW 456	-

Pain Assessed: SF-36 Scale: 0 to 100 Follow up: range 3 to 12 months	The mean pain score ranged from 47 to 87 points	The mean pain score was 5.3 points higher with exercise (0.1 lower to 10.7 higher)	- 872 (15)	⊕⊕⊙⊝ LOW 4 5	Any exercise may reduce pain how- ever, the 95% CI indicates that ex- ercise training might make little or no difference in the level of pain
Depression Assessed: multiple sever- ity of depressive symp- toms scales Follow up: range 2 to 12 months		The SMD for depression was 0.62 SD lower with exercise (1.00 to 0.24 lower)	- 490 (11)	⊕⊕⊕⊝ MODERATE ⁵	A SD of 0.2 represents a small difference between groups^ Any exercise probably improves depression. The magnitude of the effect was greater after four months of exercise training (SMD -1.26, 95% CI -1.80 to -0.72)
Functional capacity Assessed: 6MWT Follow up: range 2 to 6 months	The mean 6MWT ranged from 290 to 495 metres	The mean 6MWT was 49.9 metres further with exercise (37.2 to 62.6 further)	- 827 (19)	⊕⊕⊕⊝ MODERATE ⁵	Any exercise probably improves functional capacity

^{*}The risk in the intervention group (and its 95% CI) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

^ Cohen's interpretation of effect size

CI: Confidence interval; RR: Risk ratio; 6MWT: 6-minute walking test; SMD: standardised mean difference

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect

Moderate certainty: We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different

Low certainty: Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect

Very low certainty: We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

¹ High risk of bias: significantly greater proportion of participants lost to follow-up in the exercise group compared to the control group

² Imprecision: based on a single study that was not powered for this outcome

 $^{^{3}}$ Indirectness: the outcome was assessed 2.5 years after the completion of the intervention

⁴ Indirectness: short interventions and short-term follow-up

⁵ High risk of bias in the included studies

⁶ Inconsistency: significant unexplained heterogeneity

⁷ Imprecision: outcome reported in few participants

Summary of findings 2. Aerobic exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Aerobic exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Patient or population: adults undergoing maintenance dialysis

Setting: all settings (e.g. during dialysis, pre- and post-dialysis; home exercise)

Intervention: aerobic exercise

Comparison: no exercise or placebo exercise

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	No. of partici- pants	Certainty of the evidence	Comments
	Risk with no exer- cise or placebo ex- ercise	Risk with Aerobic exercise	(43.7.5)	(studies)	(GRADE)	
Death (any cause) Follow up: 3 years	159 per 1,000	151 per 1,000 (89 to 257)	RR 0.95 (0.56 to 1.62)	296 (1)	⊕⊝⊝⊝ VERY LOW ¹²³	-
Cardiovascular events	Not reported	Not reported	-	-	-	-
Fatigue Follow up: range 2 to 12 months	See comment	See comment	-	221 (4)	⊕⊝⊝⊝ VERY LOW ¹⁴⁵	A pooled estimate of the effect was not calculated because the included studies assessed different dimensions of fatigue
HRQoL: Physical component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean physical component score ranged from 34 to 71 points	The mean physical component score was 6.0 points higher with aerobic exercise (1.3 lower to 10.7 higher)	-	306 (9)	⊕⊙⊙⊝ VERY LOW 456	-
HRQoL: Mental component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean mental component score ranged from 39 to 65 points	The mean mental component score was 3.3 points higher with aerobic exercise (0.9 lower to 7.6 higher)	-	306 (9)	⊕⊙⊙⊝ VERY LOW 4567	-

Pain Assessed: SF-36 Scale: 0 to 100 Follow up: range 3 to 12 months	The mean pain score ranged from 47 to 87 points	The mean pain score was 2.3 points higher with aerobic exercise (1.6 lower to 6.1 higher)	-	570 (8)	⊕⊕⊙⊝ LOW 4 6	Aerobic exercise may result in little to no difference in pain
Depression Assessed: multiple severity of depressive symptoms scales Follow up: range 2 to 12 months	-	The SMD for depression was 0.19 SD lower with aerobic exercise (0.89 lower to 0.52 higher)	-	127 (4)	⊕⊝⊝⊝ VERY LOW ^{5 6 7}	A SD of 0.2 represents a small difference between groups^
Functional capacity Assessed: 6MWT Follow up: range 2 to 6 months	The mean 6MWT ranged from 290 to 454 metres	The mean 6MWT was 53.0 metres further with aerobic exercise (33.8 to 72.2 further)	-	515 (10)	⊕⊕⊕⊝ MODERATE ⁶	Aerobic exercise probably improves functional capacity.

^{*}The risk in the intervention group (and its 95% CI) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

^ Cohen's interpretation of effect size

CI: Confidence interval; RR: Risk ratio; 6MWT: 6-minute walking test; SMD: standardised mean difference

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect

Moderate certainty: We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different

Low certainty: Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect

Very low certainty: We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

- ¹ High risk of bias: significantly greater proportion of participants lost to follow-up in the exercise group compared to the control group
- ² Imprecision: based on a single study that was not powered for this outcome
- ³ Indirectness: the outcome was assessed 2.5 years after the completion of the intervention
- ⁴ Indirectness: short interventions and short follow-up
- ⁵ Imprecision: outcome reported in few participants
- ⁶ High risk of bias in the included studies
- ⁷ Inconsistency: significant unexplained heterogeneity

Summary of findings 3. Resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Patient or population: adults undergoing maintenance dialysis





Cochrane Library

Setting: all settings (e.g. during dialysis, pre- and post-dialysis; home exercise)

Intervention: resistance exercise
Comparison: no exercise or placebo exercise

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	No. of partici- pants	Certainty of the evidence	Comments
	Risk with no exer- cise or placebo ex- ercise	Risk with resistance exercise	(33 % Ci)	(studies)	(GRADE)	
Death (any cause)	Not reported	Not reported	-	=	-	-
Cardiovascular events	Not reported	Not reported	-	-	-	-
Fatigue Assessed: Profile of Mood States score Follow up: 12 weeks	The mean fatigue score was 8.95 points	The mean fatigue score was 1.88 points lower with resistance exercise (4.14 lower to 0.38 higher)	-	68 (1)	⊕⊝⊝ VERY LOW ¹²	-
HRQoL: Physical component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean physical component score ranged from 46 to 74 points	The mean physical component score was 2.5 points higher with resistance exercise (1.3 lower to 6.3 higher)	-	176 (5)	⊕⊝⊝⊝ VERY LOW ² ³ ⁴	-
HRQoL: Mental component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean mental component score ranged from 38 to 76 points	the mean mental component score was 0.7 points lower with resistance exercise (5.9 lower to 4.6 higher)	-	176 (5)	⊕⊝⊝⊝ VERY LOW ² ³ ⁴ ⁵	-
Pain Assessed: SF-36 Scale: 0 to 100 Follow up: range 3 to 12 months	The mean pain score ranged from 60 to 82 points	The mean pain score was 10.7 points higher with resistance exercise (6.5 lower to 28.0 higher)	-	154 (5)	⊕⊝⊝⊝ VERY LOW 234	-
Depression Assessed: multiple severity of depressive symptoms scales Follow up: range 2 to 12 months	-	The SMD for depression was 0.52 SD lower with resistance exercise (0.92 to 0.12 lower)	-	99 (2)	⊕⊝⊝⊝ VERY LOW ²³⁴	A SD of 0.2 represents a small difference between groups^

						The evidence is very uncertain about the effect of resistance exercise on depression
Functional capacity Assessed: 6MWT Follow up: range 2 to 6 months	The mean 6MWT ranged from 407 to 495 metres	The mean 6MWT was 44.7 metres further with resistance exercise (27.0 to 62.4 further)	-	216 (7)	⊕⊕⊕⊝ MODERATE ²	Resistance exercise probably improves functional capacity

*The risk in the intervention group (and its 95% CI) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

^ Cohen's interpretation of effect size

CI: Confidence interval; RR: Risk ratio; 6MWT: 6-minute walking test; SMD: standardised mean difference

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect

Moderate certainty: We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different

Low certainty: Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect

Very low certainty: We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

- ¹ Imprecision: based on a single study that was not powered for this outcome
- ² High risk of bias in the included studies
- ³ Indirectness: short interventions and short follow-up
- ⁴ Imprecision: outcome reported in few participants
- ⁵ Inconsistency: significant heterogeneity

Summary of findings 4. Combined aerobic and resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Combined aerobic and resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Patient or population: adults undergoing maintenance dialysis

Setting: all settings (e.g. during dialysis, pre- and post-dialysis; home exercise)

Intervention: combined aerobic and resistance exercise

Comparison: no exercise or placebo exercise

Outcomes	Anticipated absolute effects* (95% CI)	Relative effect (95% CI)	No. of partici- pants	Certainty of the evidence	Comments
			(studies)	(GRADE)	

Cochrane Library

Trusted evidence.
Informed decisions.
Better health.

	Risk with no exer- cise or placebo ex- ercise)	Risk with combined aerobic and resistance exercise				
Death (any cause)	Not reported	Not reported	-	-	-	-
Cardiovascular events	Not reported	Not reported	-	=	-	-
Fatigue	Not reported	Not reported	-	-	-	-
HRQoL: Physical component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean physical component score ranged from 38 to 51	The mean physical component score was 4.4 points higher with combined exercise (1.9 higher to 6.8 higher)	-	228 (6)	⊕⊙⊙⊝ VERY LOW 123	The evidence is very uncertain about the effect of combined aerobic and resistance exercise on the physical component of HRQoL
HRQoL: Mental component score Assessed: SF-36 Scale: 0 to 100 Follow up: range 2 to 12 months	The mean mental component score ranged from 40 to 43	The mean mental component score was 2.6 points higher with combined exercise (1.7 lower to 6.9 higher)	-	228 (6)	⊕⊙⊙ VERY LOW ¹²³	-
Pain Assessed: SF-36 Scale: 0 to 100 Follow up: range 3 to 12 months	The mean pain score ranged from 68 to 83 points	The mean pain score was 4.0 points higher with combined exercise (2.5 lower to 10.5 higher)	-	161 (3)	⊕⊙⊙⊝ VERY LOW 123	-
Depression Assessed: multiple severity of depressive symptoms scales Follow up: range 2 to 12 months	-	The SMD for depression was 1.0 SD lower with combined exercise (1.7 lower to 0.3 lower)	-	214 (4)	⊕⊝⊝⊝ VERY LOW ²³	A SD of 0.2 represents a small difference between groups^ The evidence is very uncertain about the effect of combined aerobic and resistance exercise on depression
Functional capacity Assessed: 6MWT Follow up: range 2 to 6 months	The mean 6MWT ranged from 399 to 430 metres	The mean 6MWT was 53.6 metres further (39.4 to 67.9 further)	-	138 (6)	⊕⊕⊕⊝ MODERATE ¹²	Combined aerobic and resistance exercise probably improves functional capacity.

*The risk in the intervention group (and its 95% CI) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

^ Cohen's interpretation of effect size

CI: Confidence interval; RR: Risk ratio; 6MWT: 6-metre walking test; SMD: standardised mean difference

GRADE Working Group grades of evidence

High certainty: We are very confident that the true effect lies close to that of the estimate of the effect

Moderate certainty: We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different

Low certainty: Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of the effect

Very low certainty: We have very little confidence in the effect estimate: The true effect is likely to be substantially different from the estimate of effect

- ¹ Indirectness: short interventions and short follow-up
- ² High risk of bias in the included studies
- ³ Imprecision: outcome reported in few participants



BACKGROUND

Description of the condition

Kidney failure on dialysis is a debilitating condition weighing heavily on patients' physical and psychosocial health. Death is high, particularly in older age groups, with less than 50% surviving five years after initiation (ERA-EDTA 2017; USRDS 2017; ANZDATA 2019). In addition to the time and commitment for the treatment itself, dialysis is often accompanied by debilitating symptoms such as fatigue, pain, pruritus, cramping, sleep disturbances and sexual dysfunction. As a result, quality of life (QoL) for individuals undergoing dialysis is among the lowest of any chronic diseases (Wyld 2012).

Neuromuscular complications of chronic kidney disease (CKD) have long been described (Serratrice 1967; Tyler 1975). Multiple uraemic, hormonal, immunologic, mechanical and myocellular changes are likely to contribute to skeletal muscle wasting in dialysis patients (Fahal 2014). Furthermore, the transfer of oxygen to the muscle cells is impaired despite the correction of anaemia (Stray-Gundersen 2016). In consequence, people suffering from kidney failure have a severely impaired capacity to exercise, averaging 50% to 60% of the age-expected norm (Kaysen 2011; Painter 2017) and low self-reported physical functioning even amongst younger patients (DeOreo 1997; Painter 2005). Correspondingly, people with kidney failure have extremely low levels of physical activity and rank under the fifth percentile of healthy age-matched individuals (Cupisti 2017; Johansen 2010). Of note, low exercise capacity, low physical functioning and low levels of physical activity have all been associated with a higher risk of death in this population (DeOreo 1997; Johansen 2013; Knight 2003; Sietsema 2004).

Description of the intervention

Physical activity varies in its nature, intensity, frequency, and duration. Aerobic or cardiovascular exercise implies an increase in heart and respiratory rate such as running, cycling, walking, or swimming. Resistance exercise relates to activities leading to increased muscle strength, tone and bulk, such as repeated movements of the upper and lower limbs against gravity with weights or against elastic bands. The World Health Organization recommends that adults aged 18 to 64 years old perform a minimum of 150 minutes of moderate-intensity aerobic physical activity or 75 minutes of vigorous-intensity aerobic physical activity throughout the week to improve cardiorespiratory and muscular fitness (WHO 2010). Based on evidence in the general population, the 2005 KDOQI Clinical Practice Guidelines for Cardiovascular Disease in Dialysis Patients recommend working towards 30 minutes of moderate exercise most days for adults on dialysis (KDOQI 2005).

How the intervention might work

Exercise training has the potential to improve many outcomes that are important to patients receiving dialysis treatments. In the general population, physical activity may reduce the risk of death (any cause), coronary heart disease, high blood pressure, stroke, type 2 diabetes, metabolic syndrome, and colon and breast cancer (WHO 2010). Fatigue, a debilitating symptom affecting 55% to 97% of people receiving dialysis (Chang 2001; Jacobson 2019; Jhamb 2008; Yngman-Uhlin 2010), was improved after exercise training in people with cancer (Cramp 2012) and chronic fatigue

syndrome (Larun 2019). Exercise can also improve depression (Cooney 2013), which affects 23% to 39% of adults undergoing dialysis (Palmer 2013). Finally, the previous version of this review demonstrated exercise training is likely to improve physical fitness, physical functioning and health-related QoL (HRQoL) in adults with CKD (Heiwe 2011). Through better cardiorespiratory capacity and strength, exercise training may improve patients' capacity to perform their daily activities and ease the burden of dialysis treatments.

Why it is important to do this review

Patients, caregivers, and health professionals alike believe lifestyle interventions, including exercise, should be a top priority for research in CKD (Manns 2014; Tong 2015). However, most randomised controlled trials (RCTs) do not address patients' priorities or patient-important outcomes (Tong 2018). The previous version of this review found exercise training improved physical fitness, HRQoL, and some cardiovascular and nutritional parameters. However, the certainty of the evidence was low, and many important outcomes such as death, cardiovascular events, and fatigue could not be assessed. Numerous studies have since been published, but no consensus has emerged concerning the effects and safety of exercise training for adults undergoing maintenance dialysis.

Differences with the previous Cochrane review

In its previous form, the Cochrane review for exercise training in adults with CKD included studies performed in individuals at all stages of CKD, including kidney transplantation and earlier stages of CKD (Heiwe 2011). As the exercise interventions in adults undergoing dialysis differed significantly from those in adults not receiving dialysis, and because these populations differ in their needs, risk factors and coexisting diseases, an editorial decision was taken to divide the previously published review into three separate reviews. The current review will focus on RCTs of exercise interventions in adults undergoing maintenance haemodialysis (HD) or peritoneal dialysis (PD), and separate reviews to be published at a later time will focus on adults with CKD not undergoing dialysis and kidney transplant recipients.

OBJECTIVES

To assess the benefits and safety of regular structured exercise training in adults undergoing dialysis on patient-important outcomes including death, cardiovascular events, fatigue, functional capacity, pain, and depression. We also aimed to define the optimal prescription of exercise in adults undergoing dialysis.

METHODS

Criteria for considering studies for this review

Types of studies

We included all RCTs and quasi-RCTs (RCTs in which allocation to treatment was obtained by alternation, use of alternate medical records, date of birth or other predictable methods) evaluating a structured program of regular physical exercise training in adults undergoing dialysis.



Types of participants

Inclusion criteria

We included studies involving adults receiving maintenance HD or PD treatments.

Exclusion criteria

We excluded studies involving children, kidney transplant recipients or adults with CKD not undergoing dialysis.

Types of interventions

We included interventions consisting of a structured program of regular physical exercise lasting a minimum of eight weeks to ensure the intervention consisted of regular ongoing exercise training. Interventions consisting solely of the recommendation or promotion of physical activity were excluded. Interventions targeting a single muscle group for purposes other than improvement of the general fitness, such as respiratory muscle training or hand-forearm exercises for arteriovenous fistula maturation, were also excluded.

Eligible studies had to include a control group that did not partake in any significant exercise training. Sham exercises such as light stretching exercises were allowed. Co-interventions with exercise training were allowed if the co-interventions were also administered to the control group.

Types of outcome measures

While all outcomes were collected, this review focused on patient-important outcomes, which we identified using the SONG core-outcome set for adults undergoing HD (SONG-HD 2017). When the outcomes were measured at multiple time points within the same study, we included the results corresponding to the end of the intervention period in the meta-analyses. For long-term outcomes such as death and cardiovascular events, we also recorded outcome results that were measured after the completion of the intervention.

Primary outcomes

- Death (any cause)
- Cardiovascular events
- Fatigue

Secondary outcomes

- HRQoL
- Pain
- Depression
- Functional capacity
- Resting blood pressure: systolic blood pressure (SBP) and diastolic blood pressure (DBP)
- Adherence to the exercise program
- Adverse events related to the exercise program

Other outcomes

We also assessed exploratory outcomes that were either reported in the previous version of this review (Heiwe 2011) or were commonly reported across the included studies.

Haemoglobin

- · Dialysis adequacy
- Potassium
- Physical fitness (aerobic capacity, muscular strength)
- Measures from cardiac ultrasound (left ventricular ejection fraction, left ventricular mass index)
- Body mass indices (body mass index, muscle mass, fat mass)
- Nutritional measures (albumin, energy intake, protein intake)
- Blood lipids (total cholesterol, low-density lipoproteins (LDL), high-density lipoproteins (HDL), triglycerides)
- Bone health (calcium, phosphorus, parathyroid hormone)
- Markers of inflammation (C-reactive protein)

Search methods for identification of studies

Electronic searches

We searched the Cochrane Kidney and Transplant Register of Studies to 23 December 2020 through contact with the Information Specialist using search terms relevant to this review. The Cochrane Kidney and Transplant Specialised Register contains studies identified from the following sources.

- Monthly searches of the Cochrane Central Register of Controlled Trials (CENTRAL)
- 2. Weekly searches of MEDLINE OVID SP
- 3. Searches of kidney and transplant journals, and the proceedings and abstracts from major kidney and transplant conferences
- 4. Searching of the current year of EMBASE OVID SP
- 5. Weekly current awareness alerts for selected kidney and transplant journals
- 6. Searches of the International Clinical Trials Register (ICTRP) Search Portal and ClinicalTrials.gov.

Studies contained in the Register are identified through searches of CENTRAL, MEDLINE, and EMBASE based on the scope of Cochrane Kidney and Transplant. Details of search strategies, as well as a list of handsearched journals, conference proceedings and current awareness alerts, are available on the Cochrane Kidney and Transplant website under CKT Register of Studies.

See Appendix 1 for search terms used in strategies for this review.

Searching other resources

- Reference lists of review articles, relevant studies and clinical practice guidelines.
- Contacting relevant individuals/organisations seeking information about unpublished or incomplete studies.

Data collection and analysis

Selection of studies

Two authors independently screened the titles and abstracts from the electronic search and retained potentially eligible studies. Two authors then independently assessed the abstracts and, when necessary, the full published text and identified the studies to be included in the review.

Data extraction and management

Two authors independently extracted the data from each study using standardised data extraction forms. Studies in a non-English



language were translated into English. Results from multiple publications of the same study were grouped, and the primary study publication was used as the reference for the methods. One author performed the final data entry, and a second verified each entry using the independently collected extraction sheet. Disagreements were resolved by returning to the full published text, and a third author was available for persisting disagreements.

Assessment of risk of bias in included studies

The following items were assessed independently by two authors using the risk of bias assessment tool (Higgins 2011) (see Appendix 2).

- Was there adequate sequence generation (selection bias)?
- Was allocation adequately concealed (selection bias)?
- Was knowledge of the allocated interventions adequately prevented during the study?
- Participants and personnel (performance bias)
- · Outcome assessors (detection bias)
- Were incomplete outcome data adequately addressed (attrition bias)?
- Are reports of the study free of suggestion of selective outcome reporting (reporting bias)?
- Was the study apparently free of other problems that could put it at risk of bias?

Due to the nature of the intervention, we assumed that the studies that did not report whether the participants were blinded did not attempt to blind the participants.

Measures of treatment effect

We used the mean difference (MD) with 95% confidence intervals (CI) to measure the effect of exercise training on continuous outcomes. Where the included studies used different measuring scales, we used the standardised mean difference (SMD). For dichotomous outcomes, we used risk ratios (RR) with 95% CI to measure the effect of the intervention.

To assess whether the observed effect is clinically meaningful, we considered the following for each outcome measure.

- Anchor-based estimates of the minimal clinically important differences
- Distribution methods such as the standardised mean difference
- Definitions of a clinically meaningful effect that have been used in previous RCTs and systematic reviews of adults undergoing dialysis.

When an estimate of the minimal clinically important difference was not available for the kidney failure population, we used estimates established in populations with other debilitating chronic diseases.

Unit of analysis issues

This review included studies with non-standard designs such as cross-over RCTs, cluster RCTs, cluster step-wedge RCTs, factorial RCTs and studies with two or more intervention arms.

Cross-over RCTs

Cross-over RCTs were eligible for inclusion in the review. However, the exercise intervention administered in the first study period was likely to have carry-over effects into the subsequent study periods from long-lasting effects and behaviour changes arising from the intervention. Therefore, we planned to only include outcome data following the first treatment period, where the intervention was randomly allocated analogous to a two-arms parallel RCT. There was one cross-over RCT eligible for inclusion in the review.

Cluster RCTs

Cluster RCTs were eligible for inclusion in the review. To correct for the correlation between the individuals within a cluster, we divided the effective sample size by the design effect defined as 1+ICC(M-1), where M is the average cluster size and ICC the intracluster correlation coefficient. Two cluster RCTs were eligible for inclusion in the review and their published article provided the ICC used for sample size calculation.

Step-wedge RCTs

Step-wedge RCTs were eligible for inclusion in the review. We collected and analysed the results at the latest time point before the last group initiated the intervention. The last group which had not yet initiated the intervention was used as the control group, analogous to a parallel RCT.

Factorial RCTs

Factorial RCTs were eligible for inclusion in the review. We pooled the results from the arm receiving exercise and the alternative intervention together with the results of the arm receiving exercise only under the exercise group and pooled the results from the arm receiving only the alternative intervention together with the results of the arm receiving no intervention under the control arm.

Multi-arms RCTs

RCTs with more than two arms were eligible for inclusion in the review. One of the arms had to be a control group not undertaking any significant exercise training for the study to be included. We extracted the results from all arms meeting the inclusion criteria for the intervention. When two or more arms from the same study were relevant to a meta-analysis (e.g. an aerobic exercise arm and a resistance exercise arm both eligible for a meta-analysis of any exercise), we combined the results of each arm as if they were the same treatment arm. For subgroup analyses of continuous outcomes, if two or more arms from the same study were included in distinct subgroups but shared the same control group, we divided the sample size of the control group by the number of arms. At all times, we took special care not to include the same participants twice in either the treatment or the control group for all meta-analyses.

Dealing with missing data

We contacted the study authors by written correspondence whenever data was missing from the publication. We also contacted the authors of abstracts for which we could not identify a full-text publication. Whenever we suspected a report to be a secondary publication of another included study, we also contacted the authors for clarification.



When results were only provided in the form of graphs, we extracted, to the best of our abilities, the results from the graph and included them in meta-analyses. For continuous outcomes, when only the median and the range or only the median and the interquartile range were reported, we estimated the mean and the standard deviation (SD) using the method described by Wan 2014. For continuous outcomes, when the SD was not reported, we imputed the missing SD using the highest SD from the other studies included in the meta-analysis.

Assessment of heterogeneity

We first assessed the heterogeneity by visual inspection of the forest plot. We then quantified statistical heterogeneity using the I^2 statistic, which describes the percentage of total variation across studies that is due to heterogeneity rather than sampling error (Higgins 2003). A guide to the interpretation of I^2 values was as follows.

- 0% to 40%: might not be important
- 30% to 60%: may represent moderate heterogeneity
- 50% to 90%: may represent substantial heterogeneity
- 75% to 100%: considerable heterogeneity.

The importance of the observed value of I² depends on the magnitude and direction of treatment effects and the strength of evidence for heterogeneity (e.g. P-value from the Chi² test, or a confidence interval for I²) (Higgins 2011).

Assessment of reporting biases

In meta-analyses of 10 studies or more and in the absence of statistical heterogeneity, we used funnel plots whenever possible to assess for the potential small study bias (Higgins 2011).

Data synthesis

In the meta-analyses, we pooled the estimated effects of exercise training using the DerSimonian and Laird method for random effects (DerSimonian 1986).

Subgroup analysis and investigation of heterogeneity

We performed subgroup analyses to investigate the reasons behind heterogeneity. We performed the following subgroup analyse whenever there was evidence of significant heterogeneity in the effect of the intervention:

 Type of exercise (aerobic versus resistance versus combined aerobic and resistance)

- Duration of intervention (4 months or less versus longer)
- Intensity of the exercise intervention (light to moderate versus moderate versus moderate to vigorous versus unclear)
- Risk of bias (studies that blinded participants to treatment allocation versus those that didn't).

Sensitivity analysis

For each of the primary and secondary outcomes, we performed sensitivity analyses based on the risk of bias (study at higher risk of bias versus those at lower risk of bias).

Summary of findings and assessment of the certainty of the evidence

We have presented the main results of the review in 'Summary of findings' tables. These tables present key information concerning the quality of the evidence, the magnitude of the effects of the interventions examined, and the sum of the available data for the main outcomes (Schunemann 2011a). The 'Summary of findings' table also includes an overall grading of the evidence related to each of the main outcomes using the GRADE (Grades of Recommendation, Assessment, Development and Evaluation) approach (GRADE 2008; GRADE 2011). The GRADE approach defines the quality of a body of evidence as to the extent to which one can be confident that an estimate of effect or association is close to the true quantity of specific interest. The quality of a body of evidence involves consideration of the within-trial risk of bias (methodological quality), directness of evidence, heterogeneity, the precision of effect estimates and risk of publication bias (Schunemann 2011b). We have presented the following outcomes.

- · Death (any cause)
- Cardiovascular events
- Fatigue
- · HRQoL physical component score
- HRQoL mental component score
- Dain
- · Depression
- · Functional capacity 6MWT

RESULTS

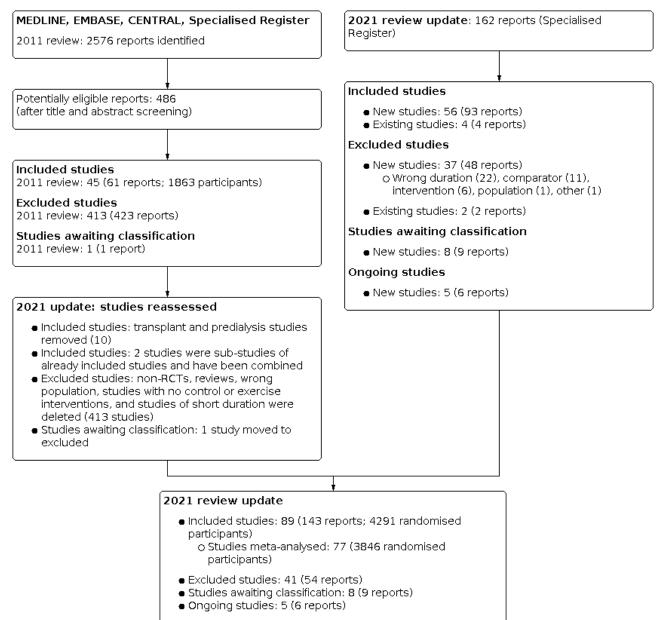
Description of studies

Results of the search

Figure 1 shows the number of studies screened and included in the 2011 review and in this current review.



Figure 1. Flow diagram showing study identification and selection



2011 review

The original literature search for the 2011 review identified 2576 reports. Sixty-one reports from 45 studies were included (Heiwe 2011). Studies were mainly excluded because they were not RCTs, did not involve an exercise intervention or did not involve a control group.

2021 update

The 2011 review has been divided into three independent reviews, one for adults undergoing dialysis, one for adults with CKD not undergoing dialysis and one for kidney transplant recipients. Of the 45 studies included in 2011, only studies involving participants undergoing dialysis were retained for this review update. We have confirmed with the authors that two studies were secondary publications of other already included studies, and have been

combined for this update (Harter 1985; Kouidi 1997). There are 33 studies remaining from the 2011 review (Akiba 1995; Carmack 1995; Chen 2010; Deligiannis 1999; Deligiannis 1999a, DePaul 2002; Frey 1999; Goldberg 1983; Harter 1985; Johansen 2006; Koh 2009; Konstantinidou 2002; Kopple 2007; Koufaki 2002; Koufaki 2003; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Kouidi 2008; Kouidi 2010; Lee 2001; Matsumoto 2007; Molsted 2004; Ouzouni 2009; Painter 2002a; Parsons 2004; PEAK 2006; Segura-Orti 2009; Toussaint 2008; Tsuyuki 2003; van Vilsteren 2005; Yurtkuran 2007).

We searched the Cochrane Kidney and Transplantation up to December 2020 and identified 162 new potentially eligible reports. After reviewing abstracts and full-text publications, we identified: 56 new included studies (93 reports); four reports of four previously included studies; 37 new excluded studies (48 reports); two reports of two previously excluded studies; and five ongoing studies (6



reports). Eight studies have been completed but are yet to publish results.

In total, for this 2021 update, we included 89 studies (143 reports) and excluded 41 studies (54 reports). There are five ongoing studies and eight studies awaiting classification which will be assessed in a future update of this review.

We contacted via email the authors of 25 studies (Abreu 2017; Abundis Mora 2017; Afshar 2010; Afshar 2011; Bennett 2013; Burrows 2018; Goldberg 1983, Harter 1985, IHOPE 2019; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Kouidi 2008; Kouidi 2010; Ma 2018; Marinho 2016; Mitsiou 2015; Miura 2015; Paluchamy 2018; Reboredo 2010; Rouchon 2016; Sheshadri 2020; Wilund 2010; Zhao 2017) and received unpublished data from two (Paluchamy 2018; Rouchon 2016).

Included studies

Details of each included study are provided in Characteristics of included studies and Appendix 3.

We included 89 studies (143 reports; 4291 randomised participants). There was one cross-over RCT (Toussaint 2008), one cluster RCT (CYCLE-HD 2016), one step-wedge cluster RCT (Bennett 2013), and three factorial RCTs (Johansen 2006; Mitsiou 2015; Painter 2002a). The remaining 83 studies were parallel-group RCTs. Sixteen studies had three arms (Afshar 2010; Amini 2016; AVANTE-HEMO 2020; Bennett 2013; Deligiannis 1999a; de Lima 2013; Dobsak 2012; Giannaki 2013a; IHOPE 2019; Koh 2009; McAdams-DeMarco 2018; McGregor 2018; Miura 2015; Pellizzaro 2013; Suhardjono 2019; Zhao 2017) and seven had four arms (Cho 2018; DIALY-SIZE 2016; Johansen 2006; Konstantinidou 2002; Kopple 2007, Mitsiou 2015; Painter 2002a). The remaining studies had two arms.

Nine studies were only published as abstracts (Abundis Mora 2017; Burrows 2018; CYCLE-HD 2016; Koufaki 2003; Jong 2004; Ma 2018; Mitsiou 2015; Miura 2015; Rouchon 2016). Twelve studies could not contribute to the meta-analyses (Abundis Mora 2017; Burrows 2018; Dashtidehkordi 2019; Harter 1985; Koufaki 2003; Kouidi 2003; Kouidi 2005; Ma 2018; McAdams-DeMarco 2018; Mitsiou 2015; Miura 2015; Mortazavi 2013) because they either did not report the number of participants in which the outcome was measured or did not report outcomes that were relevant to this review. Therefore, 77 studies (3846 randomised participants) contributed to the meta-analyses.

Twenty-six studies were conducted in Europe/UK (ACTINUT 2013; CYCLE-HD 2016; Deligiannis 1999; Deligiannis 1999a; Dobsak 2012; EXCITE 2014; Giannaki 2013a; Groussard 2015; Konstantinidou 2002; Koufaki 2002; Koufaki 2003; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Kouidi 2008; Kouidi 2010; Marinho 2016; McGregor 2018; Mitsiou 2015; Molsted 2004; Ouzouni 2009; Rouchon 2016; Samara 2016; Segura-Orti 2009; van Vilsteren 2005), 22 in North America (Abundis Mora 2017; AVANTE-HEMO 2020; Burrows 2018; Carmack 1995; Chen 2010; Cooke 2018; DePaul 2002; DIALY-SIZE 2016; Dong 2011; Frey 1999; Goldberg 1983; Harter 1985; IHOPE 2019; Johansen 2006; Kopple 2007; Martin-Alemany 2016; McAdams-DeMarco 2018; Olvera-Soto 2016; Parsons 2004; Painter 2002a; Sheshadri 2020; Wilund 2010), 17 in Asia (Akiba 1995; CHAIR 2015; Chang 2010; Cho 2018; Jong 2004; Lee 2001; Liao 2016; Ma 2018; Matsumoto 2007; Miura 2015; Paluchamy 2018; Song 2012a; Suhardjono 2019; Tsuyuki 2003; Uchiyama 2019; Wu 2014d; Zhao 2017), 10 in the Middle East (Afshar 2010; Afshar 2011; Amini 2016; Dashtidehkordi 2019; Makhlough 2012; Momeni 2014; Mortazavi 2013; Rahimimoghadam 2017; Rezaei 2015; Yurtkuran 2007), eight in South America (Abreu 2017; de Lima 2013; Fernandes 2019; Marchesan 2016; Martins do Valle 2020; Pellizzaro 2013; Reboredo 2010; Rosa 2018), four in Oceania (Bennett 2013; Koh 2009; PEAK 2006; Toussaint 2008), and two in Africa (Frih 2017a; Soliman 2015).

Participants

Three studies exclusively included participants on PD (Jong 2004; Rouchon 2016; Uchiyama 2019), and four others included participants either on maintenance HD or PD (EXCITE 2014; Koufaki 2002; Koufaki 2003; Sheshadri 2020) for a total of 151 included participants receiving PD. The remaining studies included participants on HD only. Exclusion criteria were diverse but often included any medical condition or physical incapacities precluding the participant from undertaking the exercise intervention, cognitive limitations, medical instability, and significant cardiac events in the months leading to the trial. Many studies relied on a convenience sample of prevalent HD patients with only 15 studies reporting a power and sample size calculation (AVANTE-HEMO 2020; Bennett 2013; Chang 2010; CYCLE-HD 2016; Dong 2011; EXCITE 2014; Giannaki 2013a; IHOPE 2019; Koh 2009; Rahimimoghadam 2017; Rezaei 2015; Sheshadri 2020; Song 2012a; Suhardjono 2019; Uchiyama 2019).

The number of participants randomised ranged from 11 and 296 participants (median = 38) and 30 (34%) studies randomised less than 30 participants (Abundis Mora 2017; ACTINUT 2013; Afshar 2010; Afshar 2011; Akiba 1995; Burrows 2018; CHAIR 2015; Cooke 2018; Dobsak 2012; Frey 1999; Giannaki 2013a; Goldberg 1983; Groussard 2015; Harter 1985; Koufaki 2003; Kouidi 2004a; Marchesan 2016; Marinho 2016; Martins do Valle 2020; McAdams-DeMarco 2018; Mortazavi 2013; Paluchamy 2018; Parsons 2004; Reboredo 2010; Rouchon 2016; Samara 2016; Segura-Orti 2009; Toussaint 2008; Tsuyuki 2003; Wilund 2010). The rate of attrition ranged from 0 to 49% (median 13%).

The participants mean age ranged from 30 to 72 years. In 15 studies, the participants' mean age was lower than 40 years old (Akiba 1995; AVANTE-HEMO 2020; CHAIR 2015; DIALY-SIZE 2016; Goldberg 1983; Harter 1985; Marinho 2016; Martin-Alemany 2016; Mortazavi 2013; Olvera-Soto 2016; Rahimimoghadam 2017; Tsuyuki 2003; Wu 2014d; Yurtkuran 2007; Zhao 2017) and older than 60 years in 12 (ACTINUT 2013; Bennett 2013; Chen 2010; EXCITE 2014; Frih 2017a; Groussard 2015; Liao 2016; Marchesan 2016; Miura 2015; PEAK 2006; Rouchon 2016; Uchiyama 2019).

The included studies involved predominantly males (62% of all the included participants). Three studies included only men (Afshar 2010; Afshar 2011, Frih 2017a) and six included more than 75% men (DIALY-SIZE 2016; EXCITE 2014; Rahimimoghadam 2017; Samara 2016; Sheshadri 2020; Wu 2014d). The average duration of dialysis across studies ranged from 1.8 to 6.0 years, and the average BMI across studies ranged from 20.1 to 31.2 kg/m². Eighteen studies had a mean participant's BMI above 25 (Chen 2010; Cooke 2018; de Lima 2013; Dobsak 2012; Dong 2011; EXCITE 2014; Giannaki 2013a; IHOPE 2019; Johansen 2006; Koh 2009; Koufaki 2002; Kouidi 1997; McAdams-DeMarco 2018; McGregor 2018; PEAK 2006; Rouchon 2016; Soliman 2015; Toussaint 2008).



Study comparisons

Within the 89 published studies, there were 100 different eligible exercise interventions. The characteristics of the included exercise interventions are detailed in Table 1 and in Characteristics of included studies. The interventions lasted between eight weeks and two years. In 49 studies (55%), the intervention lasted three months or less (Abreu 2017; Afshar 2010; Afshar 2011; Akiba 1995; Amini 2016; AVANTE-HEMO 2020; Bennett 2013; CHAIR 2015; Chang 2010; Cho 2018; Dashtidehkordi 2019; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Fernandes 2019; Frey 1999; Johansen 2006; Jong 2004; Koufaki 2002; Koufaki 2003; Lee 2001; Liao 2016; Makhlough 2012; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; McAdams-DeMarco 2018; McGregor 2018; Miura 2015; Momeni 2014; Olvera-Soto 2016; Paluchamy 2018; Parsons 2004; PEAK 2006; Pellizzaro 2013; Rahimimoghadam 2017; Rezaei 2015; Reboredo 2010; Rosa 2018; Rouchon 2016; Sheshadri 2020; Soliman 2015; Song 2012a; Suhardjono 2019; Toussaint 2008; Uchiyama 2019; van Vilsteren 2005; Wu 2014d; Yurtkuran 2007) whilst only 10 interventions lasted more than six months (Abundis Mora 2017; Goldberg 1983; Harter 1985; IHOPE 2019; Kouidi 2003; Kouidi 2005; Kouidi 2010; Ma 2018; Matsumoto 2007; Ouzouni 2009).

Aerobic exercise

Aerobic training was assessed in 56 (63%) studies (Abundis Mora 2017; ACTINUT 2013; Afshar 2010 Afshar 2011; Akiba 1995; Amini 2016; AVANTE-HEMO 2020; Carmack 1995; CHAIR 2015; Chang 2010; Cho 2018; Cooke 2018; CYCLE-HD 2016; Dashtidehkordi 2019; Deligiannis 1999a; de Lima 2013; DIALY-SIZE 2016; Dobsak 2012; EXCITE 2014; Fernandes 2019; Frey 1999; Giannaki 2013a; Goldberg 1983; Harter 1985; Groussard 2015; IHOPE 2019; Jong 2004; Koh 2009; Kopple 2007; Koufaki 2002; Koufaki 2003; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Lee 2001; Liao 2016; Makhlough 2012; Matsumoto 2007; McAdams-DeMarco 2018; McGregor 2018; Miura 2015; Momeni 2014; Mortazavi 2013; Painter 2002a; Paluchamy 2018; Parsons 2004; Reboredo 2010; Samara 2016; Sheshadri 2020; Suhardjono 2019; Toussaint 2008; Tsuyuki 2003; Wilund 2010; Wu 2014d; Zhao 2017).

The most common intervention consisted of stationary cycling on an ergometer in 46 studies (Abundis Mora 2017; ACTINUT 2013; Afshar 2010; Afshar 2011; Akiba 1995; AVANTE-HEMO 2020; Carmack 1995; Chang 2010; Cho 2018; Cooke 2018; CYCLE-HD 2016; Dashtidehkordi 2019; Deligiannis 1999a; de Lima 2013; DIALY-SIZE 2016; Dobsak 2012; Fernandes 2019; Frey 1999; Giannaki 2013a; Goldberg 1983; Harter 1985; Groussard 2015; IHOPE 2019; Koh 2009; Kopple 2007; Koufaki 2003; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Lee 2001; Liao 2016; Matsumoto 2007; McAdams-DeMarco 2018; McGregor 2018; Miura 2015; Momeni 2014; Mortazavi 2013; Painter 2002a; Paluchamy 2018; Parsons 2004; Reboredo 2010; Suhardjono 2019; Toussaint 2008; Wilund 2010; Wu 2014d), but also included chair-stand exercises (CHAIR 2015), walking (EXCITE 2014; Goldberg 1983; Harter 1985; Jong 2004; Koh 2009; Kouidi 1997; Lee 2001; Sheshadri 2020; Tsuyuki 2003), road cycling (Zhao 2017), and swimming (Samara 2016).

Duration of the aerobic training sessions varied between 10 and 90 minutes, with most intervention being between 20 and 40 minutes/sessions (ACTINUT 2013; Afshar 2010; Afshar 2011; Akiba 1995; AVANTE-HEMO 2020; Carmack 1995; Chang 2010; Cho 2018; CYCLE-HD 2016; Dashtidehkordi 2019; Deligiannis 1999a; Lee 2001; de Lima 2013; DIALY-SIZE 2016; Dobsak 2012; Fernandes 2019;

Frey 1999; Goldberg 1983; Groussard 2015; IHOPE 2019; Koh 2009; Koufaki 2003; Lee 2001; Liao 2016; Matsumoto 2007; Momeni 2014; Mortazavi 2013; Painter 2002a; Parsons 2004; Reboredo 2010; Samara 2016; Suhardjono 2019; Toussaint 2008; Tsuyuki 2003; Wilund 2010).

There was considerable heterogeneity on the method to assess the intensity of the exercise training: 19 studies used a version of the Borg scale of perceived exertion (ACTINUT 2013; Afshar 2011; Akiba 1995; AVANTE-HEMO 2020; Chang 2010; Cooke 2018; CYCLE-HD 2016; de Lima 2013; DIALY-SIZE 2016; IHOPE 2019; Koh 2009; Lee 2001; Liao 2016; Miura 2015; Mortazavi 2013; Reboredo 2010; Samara 2016; Wilund 2010; Wu 2014d); six used a percentage of the maximum heart rate (Deligiannis 1999a; Fernandes 2019; Frey 1999; Matsumoto 2007; Suhardjono 2019; Tsuyuki 2003); four used a percentage of the maximum load (Dobsak 2012; Giannaki 2013a; Groussard 2015; Parsons 2004); four using a percentage of the maximum oxygen consumption (Goldberg 1983; Harter 1985; Kopple 2007; Koufaki 2003); three using a combination of methods (Kouidi 1997; McGregor 2018; Painter 2002a); and the remaining studies did not report the method they used. Using the interpretation of each scale we classified five studies as light to moderate intensity (perceived as light to somewhat hard) (de Lima 2013; Dobsak 2012; Miura 2015; Mortazavi 2013; Parsons 2004), 23 studies as moderate (perceived as somewhat hard) (Abundis Mora 2017; ACTINUT 2013; Akiba 1995; AVANTE-HEMO 2020; Chang 2010; Deligiannis 1999; DIALY-SIZE 2016; Fernandes 2019; Giannaki 2013a; Goldberg 1983; Harter 1985; Groussard 2015; IHOPE 2019; Koh 2009; Kopple 2007; Kouidi 1997; Lee 2001; Matsumoto 2007; McGregor 2018; Reboredo 2010; Samara 2016; Suhardjono 2019; Tsuyuki 2003; Wilund 2010), and nine studies as moderate to vigorous (perceived as somewhat hard to hard) (Afshar 2010; Afshar 2011; Cooke 2018; CYCLE-HD 2016; Frey 1999; Koufaki 2002; Liao 2016; Painter 2002a; Wu 2014d).

Resistance exercise

Twenty-one (24%) studies assessed resistance training (Abreu 2017; Afshar 2010; AVANTE-HEMO 2020; Bennett 2013; Chen 2010; Cho 2018; de Lima 2013; DIALY-SIZE 2016; Dong 2011; Johansen 2006; Kopple 2007; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; Olvera-Soto 2016; PEAK 2006; Pellizzaro 2013; Rahimimoghadam 2017; Rosa 2018; Segura-Orti 2009; Song 2012a).

Twelve exercise programs focused solely on the lower body (Abreu 2017; Afshar 2010; Bennett 2013; Chen 2010; de Lima 2013; DIALY-SIZE 2016; Dong 2011; Johansen 2006; Kopple 2007; Marinho 2016; Pellizzaro 2013; Segura-Orti 2009) and eight exercised both the upper and lower limbs (AVANTE-HEMO 2020; Cho 2018; Martin-Alemany 2016; Martins do Valle 2020; Olvera-Soto 2016; PEAK 2006; Rosa 2018; Song 2012a). Eight studies used weights (Abreu 2017; Afshar 2010; Chen 2010; DIALY-SIZE 2016; Johansen 2006; Martin-Alemany 2016; Martins do Valle 2020; Pellizzaro 2013), three studies used resistance bands (AVANTE-HEMO 2020; Bennett 2013; Cho 2018), six studies used both (DIALY-SIZE 2016; Marinho 2016; Martin-Alemany 2016; Olvera-Soto 2016; Rosa 2018; Song 2012a) and two studies used a leg press machine (Dong 2011; Kopple 2007).

Eight studies defined the duration of the exercise session in terms of the time required to complete the prescribed number of repetitions (AVANTE-HEMO 2020; Bennett 2013; DIALY-SIZE 2016; Dong 2011; Johansen 2006; Marinho 2016; Martins do Valle 2020; Pellizzaro 2013). In 10 studies (Abreu 2017; Afshar 2010; AVANTE-



HEMO 2020; Martin-Alemany 2016; Olvera-Soto 2016; PEAK 2006; Rahimimoghadam 2017; Rosa 2018; Segura-Orti 2009; Song 2012a), the duration of the training sessions varied between 10 and 50 minutes. and in four studies the duration was not reported or unclear (Chen 2010; Cho 2018; de Lima 2013; Kopple 2007).

Eight studies defined the target level of intensity on the Borg scale of perceived exertion (Afshar 2010; AVANTE-HEMO 2020; DIALY-SIZE 2016; Martin-Alemany 2016; Martins do Valle 2020; PEAK 2006; Segura-Orti 2009; Song 2012a), one on the Omni scale of perceived exertion (Chen 2010), six as a percentage of the one, three or five-repetition maximum load (Abreu 2017; Dong 2011; Johansen 2006; Kopple 2007; Marinho 2016; Pellizzaro 2013), and six did not report the level of intensity (Bennett 2013; Cho 2018; de Lima 2013; Olvera-Soto 2016; Rahimimoghadam 2017; Rosa 2018). Using the interpretation of each scale we classified 12 studies as moderate (perceived as somewhat hard) (Abreu 2017; AVANTE-HEMO 2020; Chen 2010; DIALY-SIZE 2016; Dong 2011; Johansen 2006; Kopple 2007; Marinho 2016; Martin-Alemany 2016; Pellizzaro 2013; Segura-Orti 2009; Song 2012a), and three studies as moderate to vigorous (perceived as somewhat hard to hard) (Afshar 2010; Martins do Valle 2020; PEAK 2006).

Combined aerobic and resistance exercise

Nineteen (22%) studies assessed interventions that combined aerobic and exercises within the same treatment arm (Burrows 2018; Cho 2018; Deligiannis 1999; Deligiannis 1999a; DePaul 2002; DIALY-SIZE 2016; Frih 2017a; Konstantinidou 2002; Kopple 2007; Kouidi 2008; Kouidi 2010; Ma 2018; Marchesan 2016; Molsted 2004; Ouzouni 2009; Rouchon 2016; Suhardjono 2019; Uchiyama 2019; van Vilsteren 2005). These interventions consisted of a combination of the previously mentioned aerobic and resistance exercises in varying proportions. Cycling remained the most common aerobic exercise (14 studies: Burrows 2018; Cho 2018; Deligiannis 1999; Deligiannis 1999a; DePaul 2002; Frih 2017a; Konstantinidou 2002; Kopple 2007; Kouidi 2008; Kouidi 2010; Marchesan 2016; Ouzouni 2009; Rouchon 2016; Suhardjono 2019). The duration of the training sessions varied between 20 and 90 minutes. Five studies did not report a target intensity level (Cho 2018; Kouidi 2010; Ma 2018; Ouzouni 2009; Rouchon 2016), and the remaining studies used a combination of the previously mentioned scales. We classified one study as light to moderate intensity (perceived as light to somewhat hard) (Suhardjono 2019), 11 studies as moderate intensity (perceived as somewhat hard) (Burrows 2018; Deligiannis 1999; Deligiannis 1999a; DePaul 2002; DIALY-SIZE 2016; Frih 2017a; Konstantinidou 2002; Kopple 2007; Marchesan 2016; Uchiyama 2019; van Vilsteren 2005), and three as moderate to vigorous (perceived as somewhat hard to hard) (Kouidi 2008; Konstantinidou 2002; Molsted 2004).

Other exercise training

One study assessed a yoga intervention (Yurtkuran 2007). The sessions lasted 30 minutes, two times/week and were progressive and supervised. Three studies assessed range of movement exercises (Makhlough 2012; Rezaei 2015; Soliman 2015) which consist of movements of the body articulations in their range of movement without resistance. The sessions lasted between 15 and 30 minutes, three times/week, and while the intensity was not specified, based on their description, we classified them as light exercises.

Timing of exercise training in relation to dialysis sessions

In the majority of studies (65 studies; 73%), exercise training took place during dialysis (Abreu 2017; Abundis Mora 2017; ACTINUT 2013; Afshar 2010; Afshar 2011; Akiba 1995; AVANTE-HEMO 2020; Bennett 2013; Burrows 2018; Carmack 1995; Chang 2010; Chen 2010; Cho 2018; Cooke 2018; CYCLE-HD 2016; Dashtidehkordi 2019; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Dobsak 2012; Fernandes 2019; Frey 1999; Giannaki 2013a; Groussard 2015; IHOPE 2019; Johansen 2006; Koh 2009; Konstantinidou 2002; Kopple 2007; Koufaki 2002; Koufaki 2003; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Kouidi 2008; Kouidi 2010; Liao 2016; Ma 2018; Makhlough 2012; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; Marchesan 2016; McAdams-DeMarco 2018; McGregor 2018; Mitsiou 2015; Miura 2015; Momeni 2014; Mortazavi 2013; Olvera-Soto 2016; Ouzouni 2009; Painter 2002a; Paluchamy 2018; Parsons 2004; PEAK 2006; Pellizzaro 2013; Rosa 2018; Reboredo 2010; Segura-Orti 2009; Soliman 2015; Suhardjono 2019; Toussaint 2008; van Vilsteren 2005; Wilund 2010; Wu 2014d).

Exercise training took place before or after the dialysis sessions in nine studies (CHAIR 2015; Dong 2011; Lee 2001; Matsumoto 2007; PEAK 2006; Rosa 2018; Song 2012a; van Vilsteren 2005; Zhao 2017), and on non-dialysis days in eleven studies (Deligiannis 1999; Deligiannis 1999a; EXCITE 2014; Frih 2017a; Goldberg 1983; Harter 1985; Konstantinidou 2002; Kouidi 1997; Rahimimoghadam 2017; Samara 2016; Tsuyuki 2003). The timing of the exercise sessions was unclear in the remaining studies.

Supervision of exercise sessions

The exercise sessions were directly supervised by a physicians in 15 studies (ACTINUT 2013; Afshar 2010; CHAIR 2015; Deligiannis 1999; Deligiannis 1999a; Harter 1985; IHOPE 2019; Konstantinidou 2002; Kouidi 1997; Kouidi 2008; Kouidi 2010; Liao 2016; Ouzouni 2009; Suhardjono 2019; Tsuyuki 2003), by a kinesiologist or an exercise physiologist in 10 studies (Bennett 2013; Deligiannis 1999; DePaul 2002; DIALY-SIZE 2016; Harter 1985; Kouidi 1997; McGregor 2018; Ouzouni 2009; PEAK 2006; Rosa 2018), by an investigator or research personnel in 10 studies (Amini 2016; Dong 2011; IHOPE 2019; Johansen 2006; Kopple 2007; Matsumoto 2007; McAdams-DeMarco 2018; Painter 2002a; Song 2012a; Wilund 2010), by a physical education teacher in four studies (Deligiannis 1999; Deligiannis 1999a; Konstantinidou 2002; Marinho 2016), by a physiotherapist in four studies (Abreu 2017; Frih 2017a; Molsted 2004; Segura-Orti 2009), by an exercise trainer in four studies (Kouidi 1997; Kouidi 2008; Kouidi 2010; Samara 2016), and by other professionals in two studies (EXCITE 2014; Groussard 2015). A further eight interventions were described as supervised without further information (Chen 2010; Koh 2009; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Martins do Valle 2020; Olvera-Soto 2016; Reboredo 2010). The exercise sessions were unsupervised in six studies (Jong 2004; Koh 2009; Rezaei 2015; Sheshadri 2020; Toussaint 2008; Uchiyama 2019) and the remaining studies did not report whether the exercise intervention was supervised.

Tailoring

Twenty (22%) studies did not report tailoring of the intervention to the participant's physical capacity (Abreu 2017; Abundis Mora 2017; AVANTE-HEMO 2020; Amini 2016; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Ma 2018; McAdams-DeMarco 2018; Mitsiou 2015; Miura 2015; Momeni 2014; Olvera-Soto 2016; Rahimimoghadam 2017; Rezaei 2015; Rouchon 2016; Segura-Orti 2009; Soliman 2015; Toussaint



2008; Zhao 2017). In the remaining studies, the intervention was tailored to the participant's physical capacity through adjustment of the intensity level or adjustment of the duration of the exercise session or both.

Progression

In 50 (56%) studies, the intervention were progressive through time in term of either intensity, duration or the number of repetitions or steps to achieve (ACTINUT 2013; Afshar 2010; Akiba 1995; AVANTE-HEMO 2020; Bennett 2013; Burrows 2018; Chang 2010; Chen 2010; Cho 2018; CYCLE-HD 2016; Deligiannis 1999; Deligiannis 1999a; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Dobsak 2012; Dong 2011; EXCITE 2014; Frey 1999; Frih 2017a; Giannaki 2013a; Goldberg 1983; Harter 1985; Groussard 2015; IHOPE 2019; Johansen 2006; Koh 2009; Konstantinidou 2002; Kopple 2007; Kouidi 1997; Kouidi 2008; Kouidi 2010; Lee 2001; Liao 2016; Marchesan 2016; Olvera-Soto 2016; Ouzouni 2009; Painter 2002a; Parsons 2004; Pellizzaro 2013; Rosa 2018; Samara 2016; Segura-Orti 2009; Sheshadri 2020; Song 2012a; Suhardjono 2019; Uchiyama 2019; Wilund 2010; Wu 2014d; Yurtkuran 2007). In the remaining studies, the intervention either remained unchanged throughout the study period or was not sufficiently described to assess progression.

Structured exercise intervention versus no exercise or placebo exercise were included in this review:

- Aerobic exercise versus placebo/no exercise: Abundis Mora 2017; ACTINUT 2013; Afshar 2011; Akiba 1995; Amini 2016; Carmack 1995; CHAIR 2015; Chang 2010; Cooke 2018; CYCLE-HD 2016; Dashtidehkordi 2019; Dobsak 2012; EXCITE 2014; Fernandes 2019; Frey 1999; Giannaki 2013a; Goldberg 1983; Harter 1985; Groussard 2015; IHOPE 2019; Jong 2004; Koufaki 2003; Kouidi 1997; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Lee 2001; Liao 2016; Matsumoto 2007; McAdams-DeMarco 2018; McGregor 2018; Miura 2015; Momeni 2014; Mortazavi 2013; Painter 2002a; Paluchamy 2018; Parsons 2004; Reboredo 2010; Samara 2016; Sheshadri 2020; Toussaint 2008; Tsuyuki 2003; Wilund 2010; Wu 2014d; Zhao 2017
- Resistance exercise versus placebo/no exercise: Abreu 2017; Bennett 2013; Dong 2011; Johansen 2006; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; Olvera-Soto 2016; PEAK 2006; Pellizzaro 2013; Rahimimoghadam 2017; Rosa 2018; Segura-Orti 2009; Song 2012a
- Combined aerobic and resistance exercise versus placebo/no exercise: Burrows 2018; Chen 2010; Deligiannis 1999; DePaul 2002; Frih 2017a; Kouidi 2008; Kouidi 2010; Ma 2018; Marchesan 2016; Molsted 2004; Ouzouni 2009; Rouchon 2016; Uchiyama 2019; van Vilsteren 2005
- Aerobic exercise versus resistance exercise versus placebo/no exercise: Afshar 2010; AVANTE-HEMO 2020; Deligiannis 1999a; de Lima 2013
- Aerobic exercise versus combined aerobic and resistance exercise versus placebo/no exercise: Suhardjono 2019
- Aerobic exercise versus resistance exercise versus combined aerobic and resistance exercise versus placebo/no exercise: Cho 2018; DIALY-SIZE 2016; Kopple 2007
- Intra-HD combined aerobic and resistance exercise versus home-based aerobic exercise versus placebo/no exercise: Deligiannis 1999a
- Intra-HD aerobic exercise versus home-based aerobic exercise versus placebo/no exercise: Koh 2009

- Intra-HD combined aerobic and resistance exercise versus inter-HD rehabilitation centre-based combined aerobic and resistance exercise versus home-based combined aerobic and resistance exercise versus placebo/no exercise: Konstantinidou 2002
- Yoga versus placebo/no exercise: Yurtkuran 2007
- Range of motion exercise versus placebo/no exercise: Makhlough 2012; Rezaei 2015; Soliman 2015
- Undefined exercise versus placebo/no exercise: Mitsiou 2015

Co-interventions reported were dietary counselling (ACTINUT 2013; AVANTE-HEMO 2020), oral nutritional supplement (AVANTE-HEMO 2020; Dong 2011; IHOPE 2019; Martin-Alemany 2016), antidepressant medication (Zhao 2017), volume control (Burrows 2018), and erythropoietin (Konstantinidou 2002; Koufaki 2003; Kouidi 2005).

Study outcomes

The reported outcomes were numerous and disparate, which illustrate the broad spectrum of benefits that are expected from exercise training.

Death

One study reported death at the completion of the intervention which consisted of six months of home-based walking sessions and at a post-study follow-up, three years after randomisation (EXCITE 2014). Death was a secondary endpoint for which the study was not powered.

Cardiovascular events

No study reported cardiovascular events.

Fatigue

Six studies directly measured fatigue, each using different instruments including the revised Piper Fatigue Scale and Rhoten Fatigue Scale (Amini 2016), the Hemodialysis Fatigue Scale (Chang 2010), the Profile of Mood States (Johansen 2006), the Iowa Fatigue Scale (Soliman 2015), and a poorly defined visual analogue scale (Yurtkuran 2007). One study reported the fatigue domain of the Dialysis Symptom Index (Sheshadri 2020). Because these scales assess different dimensions of fatigue, we did not conduct a metanalysis.

A further 16 studies reported the vitality domain of either the Medical Outcomes Study 36-item Short-Form Health Survey (SF-36) or a version of the Kidney Disease Quality of Life (KDQOL) questionnaires (Abreu 2017; AVANTE-HEMO 2020; Dobsak 2012; EXCITE 2014; Koh 2009; Martin-Alemany 2016; Martins do Valle 2020; Matsumoto 2007; Paluchamy 2018; Parsons 2004; PEAK 2006; Pellizzaro 2013; Sheshadri 2020; van Vilsteren 2005; Wu 2014d; Zhao 2017). One study could not contribute to the meta-analysis because its results were not rescaled from 0 to 100 points (Paluchamy 2018) and another did not provide sufficient information to be included in the meta-analysis (Martins do Valle 2020).

Health-Related Quality of Life

Forty-six studies assessed HRQoL, 27 using the SF-36 questionnaire (Abreu 2017; ACTINUT 2013; CHAIR 2015; Chen 2010; DePaul 2002; Dobsak 2012; Frih 2017a; Giannaki 2013a; IHOPE 2019; Jong 2004; Johansen 2006; Koh 2009; Martins do Valle 2020; Matsumoto 2007;



Molsted 2004; Mortazavi 2013; Painter 2002a; Parsons 2004; PEAK 2006; Ouzouni 2009; Rosa 2018; Samara 2016; Segura-Orti 2009; Sheshadri 2020; Song 2012a; van Vilsteren 2005; Zhao 2017), three using the KDQOL questionnaire (Bennett 2013; Burrows 2018; Sheshadri 2020), nine using the KDQOL-Short Form (KDQOL-SF) which includes the SF-36 (AVANTE-HEMO 2020; de Lima 2013; EXCITE 2014; Martin-Alemany 2016; Paluchamy 2018; Pellizzaro 2013; Suhardjono 2019; Uchiyama 2019; Wu 2014d), one using the SF-12 (IHOPE 2019), one using the KDQOL-SF 36 which includes SF-12 (DIALY-SIZE 2016), two using the Spitzer Index (Kouidi 1997; Ouzouni 2009), one using the Scale of Life Satisfaction (Ouzouni 2009), one using questions from the Laupacis Kidney Disease Questionnaire (DePaul 2002) and one abstract that did not report the instrument (Kouidi 2005). Of the 39 that used either the SF-36, the SF-12 or a version of the KDQOL, 17 reported the summary physical and mental component scores (ACTINUT 2013; CHAIR 2015; Chen 2010; DIALY-SIZE 2016; Dobsak 2012; Frih 2017a; Giannaki 2013a; IHOPE 2019; Koh 2009; Molsted 2004; Ouzouni 2009; Rosa 2018; Samara 2016; Segura-Orti 2009; Song 2012a; Suhardjono 2019; Uchiyama 2019) and all could contribute to the meta-analysis.

Twenty studies reported the scores for at least one individual domain of the SF-36 questionnaire (Abreu 2017; AVANTE-HEMO 2020; CHAIR 2015; Dobsak 2012; EXCITE 2014; Johansen 2006; Jong 2004; Koh 2009; Martin-Alemany 2016; Martins do Valle 2020; Matsumoto 2007; Paluchamy 2018; Parsons 2004; PEAK 2006; Pellizzaro 2013; Sheshadri 2020; Uchiyama 2019; van Vilsteren 2005; Wu 2014d; Zhao 2017) and all but one (Paluchamy 2018), for which the results were not rescaled from 0 to 100 points, contributed to the meta-analysis.

Pain

One study reported pain on a 0 to 10 visual analogue scale (Yurtkuran 2007). Sixteen studies reported pain as a domain of the SF-36 questionnaire (Abreu 2017; AVANTE-HEMO 2020; Dobsak 2012; EXCITE 2014; Koh 2009; Martin-Alemany 2016; Martins do Valle 2020; Matsumoto 2007; Molsted 2004; Paluchamy 2018; Pellizzaro 2013; van Vilsteren 2005; Uchiyama 2019; Wu 2014d; Yurtkuran 2007; Zhao 2017) and all but one study (Paluchamy 2018), for which the results were not rescaled from 0 to 100 points, contributed to the meta-analysis.

Depression

Seventeen studies assessed depression (Carmack 1995; CYCLE-HD 2016; Frih 2017a; Giannaki 2013a; Goldberg 1983; Harter 1985; Johansen 2006; Kouidi 1997; Kouidi 2005; Kouidi 2010; Ma 2018; Ouzouni 2009; PEAK 2006; Rahimimoghadam 2017; Rezaei 2015; Sheshadri 2020; van Vilsteren 2005). Seven used the Beck Depression Index (Amini 2016; Goldberg 1983; Harter 1985; Kouidi 1997; Kouidi 2010; Ouzouni 2009; Rezaei 2015), three the Hospital Anxiety and Depression Scale (CYCLE-HD 2016; Frih 2017a; Kouidi 2010), two the Center for Epidemiologic Studies Depression Scale (CES-D) (Carmack 1995; Sheshadri 2020), two the Self-rating Depression Scale (Giannaki 2013a; van Vilsteren 2005), four used other instruments (Amini 2016; Johansen 2006; PEAK 2006; Rahimimoghadam 2017) and two did not report their instrument (Kouidi 2005; Ma 2018). Ten studies (Carmack 1995; Frih 2017a; Giannaki 2013a; Kouidi 1997; Kouidi 2010; Ouzouni 2009; Rahimimoghadam 2017; Rezaei 2015; Sheshadri 2020; van Vilsteren 2005) provided sufficient information to contribute to the metaanalysis using the standardised mean difference.

Functional capacity

Functional capacity was reported in 35 studies (ACTINUT 2013; AVANTE-HEMO 2020; Bennett 2013; CHAIR 2015; Cho 2018; Cooke 2018; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Dobsak 2012; EXCITE 2014; Fernandes 2019; Frih 2017a; Giannaki 2013a; Groussard 2015; IHOPE 2019; Johansen 2006; Koh 2009; Koufaki 2002; Liao 2016; Ma 2018; Martins do Valle 2020; Marchesan 2016; Mitsiou 2015; PEAK 2006; Pellizzaro 2013; Rosa 2018; Rouchon 2016; Samara 2016; Segura-Orti 2009; Song 2012a; Suhardjono 2019; Uchiyama 2019; Wilund 2010; Wu 2014d). We meta-analysed and reported the two most commonly reported tests.

Twenty-three studies reported result for the 6MWT which measures the distance in metres covered over six minutes and reflects aerobic capacity and endurance (ACTINUT 2013; AVANTE-HEMO 2020; CHAIR 2015; Cho 2018; DePaul 2002; DIALY-SIZE 2016; EXCITE 2014; Fernandes 2019; Frih 2017a; Groussard 2015; Koh 2009; Liao 2016; Ma 2018; Martins do Valle 2020; Marchesan 2016; Mitsiou 2015; PEAK 2006; Pellizzaro 2013; Rosa 2018; Rouchon 2016; Samara 2016; Segura-Orti 2009; Wu 2014d). Nineteen studies could be meta-analysed (ACTINUT 2013; CHAIR 2015; Cho 2018; DePaul 2002; DIALY-SIZE 2016; EXCITE 2014; Fernandes 2019; Frih 2017a; Koh 2009; Liao 2016; Martins do Valle 2020; Marchesan 2016; PEAK 2006; Pellizzaro 2013; Rosa 2018; Rouchon 2016; Samara 2016; Segura-Orti 2009; Wu 2014d).

Sixteen studies reported results for the sit-to-stand test which measures leg strength and endurance (AVANTE-HEMO 2020; Bennett 2013; Cho 2018; DIALY-SIZE 2016; EXCITE 2014; Frih 2017a; Giannaki 2013a; IHOPE 2019; Johansen 2006; Koufaki 2002; Marchesan 2016; Rosa 2018; Samara 2016; Segura-Orti 2009; Song 2012a; Wu 2014d). Eight reported the maximum number of sitto-stand cycles executed within 30 seconds (Bennett 2013; Cho 2018; DIALY-SIZE 2016; Giannaki 2013a; IHOPE 2019; Marchesan 2016; Rosa 2018; Song 2012a), and five reported the number of sitto-stand cycles executed within 60 seconds (Frih 2017a; Giannaki 2013a; Koufaki 2002; Segura-Orti 2009; Wu 2014d). To meta-analyse the results conjointly, we approximated the number of cycles executed within 30 seconds by dividing the results of the last five studies by two. Five studies reported the time in seconds required to execute five sit-to-stand cycles (AVANTE-HEMO 2020; EXCITE 2014; Giannaki 2013a; Johansen 2006; Koufaki 2002), and four studies reported the time in seconds required to execute 10 sitto-stand cycles (Frih 2017a; Samara 2016; Segura-Orti 2009; Wu 2014d). To combine these results within the same meta-analysis, we approximated the time to execute five cycles by dividing the results of the later four studies by two. All but one study (AVANTE-HEMO 2020) reported their results in a manner that was amenable to meta-analysis.

Resting blood pressure

Twenty-one studies assessed resting peripheral SBP and DBP (Cooke 2018; CYCLE-HD 2016; Deligiannis 1999a; DePaul 2002; Fernandes 2019; Frih 2017a; Goldberg 1983; IHOPE 2019; Koh 2009; Kouidi 2008; Liao 2016; McGregor 2018; Miura 2015, Molsted 2004; Ouzouni 2009; Paluchamy 2018; Soliman 2015; Toussaint 2008; Tsuyuki 2003; van Vilsteren 2005; Wilund 2010) and all but one (Miura 2015) provided the results in a form amenable to meta-analysis.



Adherence to the exercise intervention

Twelve (14%) studies reported the percentage of training sessions attended by the participants allocated to the intervention group (ACTINUT 2013; Chen 2010; Cooke 2018; IHOPE 2019; Kouidi 2008; Martins do Valle 2020; Molsted 2004; PEAK 2006; Reboredo 2010; Rosa 2018; Toussaint 2008; Uchiyama 2019).

Adverse events

Thirteen (15%) studies reported adverse events (AVANTE-HEMO 2020; CHAIR 2015; Chen 2010; Cho 2018; DIALY-SIZE 2016; EXCITE 2014; IHOPE 2019; Marinho 2016; McAdams-DeMarco 2018; PEAK 2006; Sheshadri 2020; Uchiyama 2019; Wu 2014d) of which three reported severe adverse events separately (CHAIR 2015; DIALY-SIZE 2016; EXCITE 2014). Nine studies specifically reported adverse events related to the intervention (AVANTE-HEMO 2020; CHAIR 2015; Chen 2010; Cho 2018; DIALY-SIZE 2016; IHOPE 2019; Sheshadri 2020; Uchiyama 2019; Wu 2014d) and were meta-analysed.

Other outcomes

Outcomes that were frequently reported but not identified as important to patients included: aerobic capacity (VO_2 max or

peak); maximum heart rate; muscular strength; body mass index; body composition (fat and lean mass); haemoglobin; serum albumin; blood lipids; serum potassium; serum calcium; serum phosphate; parathyroid hormone levels; C-reactive protein levels; left ventricular ejection fraction; and left ventricular mass index measured on cardiac ultrasonography. These outcomes were reported in Heiwe 2011 and have been retained for historical reference only.

Excluded studies

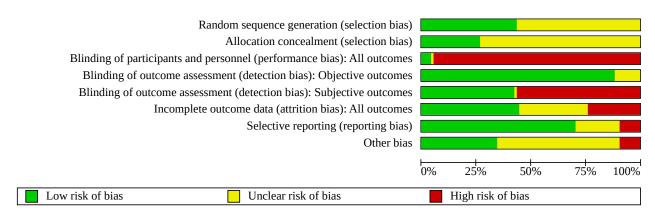
Forty-one studies were excluded. The reasons for exclusion were no control group or active control (11 studies); no intervention group (6 studies); duration < eight weeks (22 studies); wrong population (1 study); and co-interventions not the same in the control and intervention groups (1 study).

See Characteristics of excluded studies table.

Risk of bias in included studies

Figure 2 summarises the assessment of the risk of bias for the included studies, and Figure 3 provide the risk of bias assessment for individual studies.

Figure 2. Methodological quality graph: review authors' judgements about each methodological quality item presented as percentages across all included studies.



Allocation

Random sequence generation

The random sequence generation method was at low risk of bias in 39 studies (44%) (ACTINUT 2013; AVANTE-HEMO 2020; Bennett 2013; CHAIR 2015; Cho 2018; Cooke 2018; CYCLE-HD 2016; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Dong 2011; EXCITE 2014; Fernandes 2019; Frih 2017a; IHOPE 2019; Johansen 2006; Koh 2009; Kopple 2007; Koufaki 2002; Kouidi 2008; Makhlough 2012; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; McGregor 2018; Olvera-Soto 2016; Painter 2002a; Parsons 2004; PEAK 2006; Rahimimoghadam 2017; Rosa 2018; Rouchon 2016; Samara 2016; Segura-Orti 2009; Sheshadri 2020; Suhardjono 2019; Uchiyama 2019; Wu 2014d; Yurtkuran 2007), and not reported in the remaining 50 studies.

Allocation concealment

The method to conceal the treatment allocation was at low risk of bias in 24 studies (27%) (ACTINUT 2013; Bennett 2013; CHAIR

2015; Cho 2018; CYCLE-HD 2016; Dashtidehkordi 2019; de Lima 2013; DIALY-SIZE 2016; EXCITE 2014; Fernandes 2019; IHOPE 2019; Johansen 2006; Koh 2009; Koufaki 2002; Martins do Valle 2020; McGregor 2018; Molsted 2004; Painter 2002a; PEAK 2006; Rosa 2018; Sheshadri 2020; Toussaint 2008; Uchiyama 2019; Yurtkuran 2007) and not reported in the remaining 65 studies.

Blinding

Blinding of participants and investigators

While complete blinding of the participants to the exercise intervention is unlikely, we deemed the four studies that used a placebo or sham exercise were at low risk of bias (Chen 2010; DePaul 2002; Rosa 2018; Segura-Orti 2009). One study was judged to be at unclear risk of bias (Dashtidehkordi 2019), and the remaining 84 studies were judged to be at high risk of bias.



Blinding of outcome assessment

Objective outcomes

We considered the 6MWT, the Sit-To-Stand test, the Time-Up and Go test, muscular strength, blood pressure, heart rate, Kt/V, laboratory results, dietary intake, and cardiac ultrasound measures as objective outcomes that were less likely to be significantly affected by the lack of blinding of the assessors. With the exception of 10 abstracts (Abundis Mora 2017; Burrows 2018; Jong 2004; Koufaki 2003; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Ma 2018; Mitsiou 2015; Miura 2015) that we deemed at unclear risk; all studies were judged to be at low risk of bias.

Eight studies did not report any of the listed objective outcomes (Amini 2016; Chang 2010; Dashtidehkordi 2019; Matsumoto 2007; Mortazavi 2013; Rahimimoghadam 2017; Rezaei 2015; Wu 2014d).

Subjective outcomes

Fatigue, HRQoL, pain, and depression were considered subjective outcomes. Since the participants themselves assessed these outcomes, we deemed the four studies that used a placebo or sham exercise to be at low risk of bias for blinding of outcome assessment (Chen 2010; DePaul 2002; Rosa 2018; Segura-Orti 2009) as well as the 34 studies that did not report any subjective outcomes (Abundis Mora 2017; Afshar 2011; Akiba 1995; Cho 2018; Cooke 2018; Deligiannis 1999; Deligiannis 1999a; de Lima 2013; Dong 2011; Fernandes 2019; Groussard 2015; Harter 1985; Konstantinidou 2002; Kopple 2007; Koufaki 2003; Kouidi 2003; Kouidi 2004a; Kouidi 2008; Lee 2001; Liao 2016; Makhlough 2012; Marchesan 2016; Marinho 2016; McAdams-DeMarco 2018; McGregor 2018; Mitsiou 2015; Miura 2015; Momeni 2014; Olvera-Soto 2016; Reboredo 2010; Rouchon 2016; Toussaint 2008; Tsuyuki 2003; Wilund 2010). One abstract was judged as unclear (Burrows 2018), and the remaining 50 studies were judged to be at high risk of bias since the participants reported the outcomes and the participants were not blinded to treatment allocation.

Incomplete outcome data

We judged 40 (45%) studies to be at low risk of bias for incomplete outcome data (ACTINUT 2013; AVANTE-HEMO 2020; Chang 2010; Cho 2018; Dashtidehkordi 2019; de Lima 2013; DePaul 2002; DIALY-SIZE 2016; Fernandes 2019; Groussard 2015; Johansen 2006; Konstantinidou 2002; Koufaki 2002; Kouidi 1997; Kouidi 2008; Kouidi 2010; Liao 2016; Marchesan 2016; Marinho 2016; Martin-Alemany 2016; Martins do Valle 2020; Matsumoto 2007; Momeni 2014; Olvera-Soto 2016; Ouzouni 2009; Painter 2002a; Parsons 2004; Rahimimoghadam 2017; Rosa 2018; Samara 2016; Segura-Orti 2009; Sheshadri 2020; Song 2012a; Suhardjono 2019; Toussaint 2008; Uchiyama 2019; van Vilsteren 2005; Wilund 2010; Wu 2014d; Yurtkuran 2007) and 21 (23.5%) to be at high risk (Abreu 2017; Akiba 1995; Bennett 2013; Carmack 1995; CHAIR 2015; EXCITE 2014; Frey 1999; Frih 2017a; Harter 1985; IHOPE 2019; Koh 2009; Kopple 2007; Lee 2001; McAdams-DeMarco 2018; McGregor 2018; Molsted 2004; Pellizzaro 2013; Reboredo 2010; Rezaei 2015; Rouchon 2016; Soliman 2015). The remaining 28 studies provided insufficient information to permit judgement.

Selective reporting

Eight (9%) studies were at high risk of bias from selective reporting of outcomes (Lee 2001; Liao 2016; McAdams-DeMarco 2018; Olvera-Soto 2016; Painter 2002a; PEAK 2006; Pellizzaro 2013; Rezaei 2015).

Eighteen (20%) studies did not provide sufficient information to assess the risk of bias from selective reporting (Abundis Mora 2017; Afshar 2011; Akiba 1995; Burrows 2018; CYCLE-HD 2016; Harter 1985; Jong 2004; Koufaki 2003; Kouidi 2003; Kouidi 2004a; Kouidi 2005; Ma 2018; Mitsiou 2015; Miura 2015; Momeni 2014; Paluchamy 2018; Rouchon 2016; Zhao 2017). The remaining 63 studies were at low risk of bias from selective reporting.

Other potential sources of bias

We judge seven (8%) studies at high risk of bias because they received private funding without specifying whether the funders were involved in the conduction of the study (DePaul 2002; Groussard 2015; Johansen 2006; Koufaki 2002; Molsted 2004; Painter 2002a; PEAK 2006). A further study was judged at high risk of bias for discrepancies in the number of participants across the published article (Makhlough 2012). We judge 31 studies (34%) at low risk of other sources of bias because they reported either no funding or public funding (Abreu 2017; ACTINUT 2013; AVANTE-HEMO 2020; Bennett 2013; Chang 2010; Dashtidehkordi 2019; DIALY-SIZE 2016; Dobsak 2012; Dong 2011; Fernandes 2019; Giannaki 2013a; Goldberg 1983; Harter 1985; IHOPE 2019; Koh 2009; Kopple 2007; Marinho 2016; Martins do Valle 2020; McAdams-DeMarco 2018; McGregor 2018; Parsons 2004; Pellizzaro 2013; Rahimimoghadam 2017; Reboredo 2010; Rosa 2018; Segura-Orti 2009; Sheshadri 2020; Suhardjono 2019; Toussaint 2008; Wilund 2010; Wu 2014d). The remaining 50 studies did not report their source of funding.

Effects of interventions

See: Summary of findings 1 Any exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis; Summary of findings 2 Aerobic exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis; Summary of findings 3 Resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis; Summary of findings 4 Combined aerobic and resistance exercise versus no exercise or placebo exercise for adults undergoing maintenance dialysis

Primary outcomes

Death (any cause)

It is uncertain whether exercise training reduces the risk of death. EXCITE 2014 reported death three years after the intervention which consisted of six months of home-based walking exercise. Deaths were similar across the two groups (Analysis 1.1 (1 study, 296 participants): RR 0.95; 95% CI 0.56 to 1.62; very low certainty evidence). This study was not powered to assess death and the report did not specify whether there was missing data for this outcome at the three-year follow-up assessment.

Studies reporting adverse events (CHAIR 2015; Chen 2010; Cho 2018; DIALY-SIZE 2016; EXCITE 2014; Marinho 2016; McAdams-DeMarco 2018; PEAK 2006; Wu 2014d) did not report any deaths related to the exercise intervention during the duration of the study (range two to six months).

Cardiovascular events

No study reported cardiovascular events.



Fatigue

Fatigue was reduced after the exercise intervention in three studies that used fatigue-specific measures (Amini 2016; Soliman 2015; Yurtkuran 2007) and was reduced but did not reach statistical significance in two (Chang 2010; Johansen 2006). One study found similar results on the fatigue domain of the Dialysis Symptom Index across treatment groups after the exercise intervention (Sheshadri 2020) (Analysis 1.2). A sensitivity analysis based on the risk of bias could not be conducted due to the low number of studies. All the studies used aerobic training interventions and one used resistance training (Johansen 2006).

Exercise may improve vitality as assessed by the SF-36 questionnaire (Analysis 1.4.7 (16 studies, 940 participants): MD 4.47, 95% CI 0.79 to 8.15 points on a 100-point scale; $I^2 = 46\%$; low certainty evidence) where higher scores signify greater vitality. Considering a minimal clinically important difference for the individual scales of the SF-36 of two to five points (Eriksson 2016; Finkelstein 2018; Leaf 2009; Samsa 1999; Spinowitz 2019) or an SMD of 0.1 to 0.5 (Farivar 2004; Norman 2003; Samsa 1999), we judged the magnitude of the effect to be small. However, since vitality is an indirect measure of fatigue, the relevance of this result for the assessment of the outcome of fatigue is uncertain.

Health-Related Quality of Life

Physical component score

Exercise training of any type may increase the physical component of HRQoL slightly (Analysis 1.3.1 (17 studies, 656 participants): MD 4.12, 95% CI 1.88 to 6.37 points on 100 points-scale where higher scores signify a better QoL; I² = 49%; low certainty evidence). Considering a minimal clinically important difference for the physical component score of SF-36 of two to five points (Eriksson 2016; Erez 2016; Finkelstein 2018; Leaf 2009; Samsa 1999; Spinowitz 2019) or an SMD of 0.1 to 0.5 (Farivar 2004; Norman 2003; Samsa 1999) we estimated the size of the effect to be small. A sensitivity analysis including only the studies at low risk of bias (ACTINUT 2013; DIALY-SIZE 2016; Rosa 2018; Samara 2016; Segura-Orti 2009; Suhardjono 2019; Uchiyama 2019) led to a similar pooled estimate of the effect (7 studies, 309 participants: MD 4.33 points on 100 points-scale, 95% CI -0.11 to 8.76; I² = 67%).

It is uncertain whether aerobic, resistance, or combined aerobic and resistance exercise improved the physical component of QoL because the certainty of this evidence was very low (Analysis 2.3; Analysis 3.2; Analysis 4.1).

Mental component score

It is uncertain whether any exercise training improves the mental component of HR-QoL (Analysis 1.3 (17 studies, 656 participants): MD 2.53, 95% CI -0.40 to 5.47 points on 100 points-scale where higher scores signify a better QoL; I² = 73%; very low certainty evidence). There was evidence of significant heterogeneity in the effect of exercise training between studies that we could not explain with subgroups analyses based on the type, intensity or duration of exercise or based on the risk of bias. A sensitivity analysis including only the studies at low risk of bias (ACTINUT 2013; DIALY-SIZE 2016; Rosa 2018; Samara 2016; Segura-Orti 2009; Suhardjono 2019; Uchiyama 2019) led to a similar pooled estimate of the effect (7 studies, 309 participants: MD 3.04 points, 95% CI -2.91 to 8.98; I² = 67%).

It is also uncertain whether aerobic, resistance or combined aerobic and resistance exercise improves the mental component of HR-QoL as the certainty of the evidence was very low (Analysis 2.3; Analysis 3.2; Analysis 4.1).

The results of the meta-analyses for the individual domains of HR-QoL are available in Analysis 1.4 for any exercise, Analysis 2.4 for aerobic exercise, Analysis 3.3 for resistance exercise, and Analysis 4.2 for combined aerobic and resistance exercise.

Pain

Exercise training of any type may lead to lesser pain as assessed by the SF-36 or KDQOL questionnaires. However, the 95% CI indicates that exercise training might make little or no difference in the level of pain (Analysis 1.4.3 (15 studies, 872 participants): MD 5.28 95% CI -0.12 to 10.69 points on 100 points-scale where higher scores signify less pain; $I^2 = 63\%$; low certainty evidence). There was evidence of significant heterogeneity in the effect of exercise across studies. However, the heterogeneity was completely resolved by removing Pellizzaro 2013 which reported its results in figures only (pooled estimate after removing the study (14 studies. 844 participants): MD 2.80, 95% CI -0.30 to 5.91, $I^2 = 0\%$). Considering a minimal clinically important difference for each scale of the SF-36 of two to five points (Eriksson 2016; Finkelstein 2018; Leaf 2009; Samsa 1999; Spinowitz 2019) or an SMD of 0.1 to 0.5 (Farivar 2004; Norman 2003; Samsa 1999), we judged the magnitude of the effect to be small. A sensitivity analysis including only the studies at low risk of bias (AVANTE-HEMO 2020; Martin-Alemany 2016; Martins do Valle 2020; Parsons 2004; Uchiyama 2019; Wu 2014d) reported a similar pooled estimate of the effect (6 studies, 229 participants: MD 2.66 points on 100 points-scale, 95% CI -2.02 to 7.34; $I^2 = 0\%$).

Aerobic exercise training may make little or no difference to pain as assessed by the SF-36 questionnaire (Analysis 2.4.3 (8 studies, 570 participants): MD 2.26 points 95% CI -1.61 to 6.12 on 100 points-scale; $I^2 = 0\%$; low certainty evidence).

It is uncertain whether resistance exercise training and combined aerobic and resistance exercise training improves pain in adults undergoing dialysis because the certainty of this evidence was very low (Analysis 3.3.3; Analysis 4.2.3).

Depression

Exercise training of any type likely improves depression in adults undergoing dialysis (Analysis 1.5 (10 studies, 441 participants): SMD -0.65, 95% CI -1.07 to -0.22 where lower scores signify improved depressive symptoms; $I^2 = 77\%$; moderate certainty evidence). However, there was evidence of significant heterogeneity in the effect of exercise across studies. The heterogeneity was improved after stratifying the studies by the duration of the intervention (four months or less versus longer than four months). The magnitude of the effect was very large when the intervention lasted longer than four months (Analysis 1.5.2 (4 studies, 130 participants): SMD -1.26, 95% CI -0.72 to -1.80; $I^2 = 45\%$), while the 95% CI indicated that exercise training for four months or less may make little or no difference on depression (Analysis 1.5.1 (6 studies, 311 participants): SMD -0.30, 95% CI 0.14 to -0.74; $I^2 = 71\%$) (Test for subgroup differences: P = 0.007).

It is uncertain whether aerobic, resistance or combined aerobic and resistance exercise improves depressive symptoms because the certainty of this evidence is very low (Analysis 2.5; Analysis 3.4;



Analysis 4.3). A sensitivity analysis based on the risk of bias could not be conducted due to the low number of studies.

Functional capacity

6-Minute Walk Test

Exercise training of any type is likely to improve functional capacity as assessed by the 6MWT (Analysis 1.6 (19 studies, 827 participants): MD 49.91 metres, 95% CI 37.22 to 62.59; I² = 34%; moderate certainty evidence). Considering a previously reported minimal clinically important difference for the 6MWT ranging from 14.0 to 30.5 metres in patients with comorbidities and similar baseline results on the 6MWT (Bohannon 2017), we estimated the magnitude of the effect as moderate. A sensitivity analysis limited to the studies at low risk of bias (ACTINUT 2013; Cho 2018; DIALY-SIZE 2016; Fernandes 2019; Martins do Valle 2020; Rosa 2018; Segura-Orti 2009; Wu 2014d) did not significantly alter the pooled estimate of the effect (8 studies, 298 participants: MD 48.57 metres, 95% CI 34.23 to 62.92; I² = 0%).

Aerobic exercise (Analysis 2.6 (10 studies, 515 participants): MD 53.00 metres, 95% CI 33.84 to 72.17; $I^2 = 47\%$; moderate certainty evidence), resistance exercise (Analysis 3.5 (7 studies, 216 participants): MD 44.71 metres, 95% CI 27.00 to 62.43; $I^2 = 0\%$; moderate certainty evidence) or combined aerobic and resistance exercise (Analysis 4.4 (6 studies, 138 participants): MD 53.64 meters, 95% CI 39.36 to 67.91; $I^2 = 0\%$; moderate certainty evidence) are all likely to increase functional capacity.

Sit-To-Stand test

Exercise training of any type is likely to improve functional capacity and lower extremities strength as assessed by the 30- or 60-second STS test (Analysis 1.7 (12 studies, 478 participants): MD 2.36 repetitions in 30 seconds, 95% CI 1.73 to 2.98; $I^2 = 0\%$; moderate certainty evidence). We found no reference in the literature of the minimal clinically important difference for this test in adults undergoing dialysis. Judging from the results in another population with similar baseline results (Wright 2011) and the SMD of 0.63 (95% CI 0.35 to 0.91) we judged the size of the effect to be moderate. A sensitivity analysis limited to the studies at low risk of bias (Cho 2018; DIALY-SIZE 2016; Rosa 2018; Segura-Orti 2009; Wu 2014d) did not significantly alter the pooled estimate of the effect (5 studies, 219 participants: 2.79 repetitions in 30 seconds, 95% CI 1.73 to 3.86; $I^2 = 13\%$).

Exercise training is likely to improve functional lower extremities strength as assessed by the 5 to 10 repetitions STS test (Analysis 1.8 (8 studies, 508 participants) MD -1.74 seconds, 95% CI -2.25 to -1.22; $I^2=0\%$; moderate certainty evidence). Using a minimal clinically important difference of 4.2 seconds in adults with CKD (Wilkinson 2019) not on dialysis and an SMD of 0.53 (95% CI 0.30 to 0.75) we judged the size of the effect to be small. A sensitivity analysis based on the risk of bias could not be conducted for the 5 to 10 repetitions STS test due to the low number of studies. Taken together, the pooled estimates for these two versions of the STS test point to a positive effect of exercise training on lower extremities strength and physical functioning.

Aerobic (Analysis 2.7 (6 studies, 227 participants): MD 1.81 repetitions in 30 seconds, 95% CI 0.86 to 2.76; $I^2 = 0\%$; moderate certainty evidence) and resistance exercise (Analysis 3.6 (6 studies, 195 participants): MD 2.76 repetitions in 30 seconds, 95% CI 1.68 to 3.83; $I^2 = 0\%$, moderate certainty evidence) are both likely

to improve functional lower extremities strength as assessed by the 30- or 60-second STS test. Combined aerobic and resistance training may improve performance on the 30 or 60 seconds STS test (Analysis 4.5 (4 studies, 97 participants): MD 2.63 repetitions in 30 seconds, 95% CI 1.49 to 3.77; $I^2 = 9\%$; low certainty evidence). Aerobic exercise is likely to improve functional lower extremities strength as assessed by the 5 or 10 repetitions STS test (Analysis 2.8 (5 studies, 374 participants): MD -1.63 seconds, 95% CI -2.33 to -0.92, 2.33; $I^2 = 8\%$; moderate certainty evidence). It is uncertain whether resistance exercise or combined aerobic and resistance exercise improves the results of the 5 to 10 repetitions STS test because the certainty of this evidence is very low.

Peripheral resting blood pressure

The effect of exercise training on SBP and DBP was different across types of exercise (Test for subgroup differences P < 0.001 for both SBP and DBP). We will, therefore, present the results for each type of exercise separately and will not provide a pooled estimate for any exercise training.

It is uncertain whether aerobic exercise reduces SBP because the certainty of this evidence is very low (Analysis 1.9.1 (13 studies, 394 participants): MD -3.99 mm Hg, 95% CI -9.78 to 1.80; I^2 = 45%; very low certainty evidence). No study assessed the impact of resistance training alone on SBP. The evidence is very uncertain on the effect of combined aerobic and resistance training on SBP (Analysis 1.9.2 (7 studies, 282 participants): MD -8.69 mm Hg, 95% CI -13.69 to -3.69; I^2 = 57%; very low certainty evidence). The heterogeneity was entirely resolved after excluding a single study (Frih 2017a) (pooled estimate after excluding the study: MD -5.84 95% CI -9.94 to -1.74 mm Hg; I^2 = 0%).

It is uncertain whether aerobic exercise reduces DBP because the certainty of this evidence is very low (Analysis 1.10.1 (13 studies, 394 participants): MD 0.72 mm Hg, 95% CI -2.24 to 3.69; I^2 = 31%, very low certainty evidence). No study assessed the impact of resistance training alone on DBP. The evidence is very uncertain about the effect of combined aerobic and resistance training on DBP (Analysis 1.10.2 (7 studies, 282 participants): MD -4.45 mm Hg, 95% CI -5.98 to -2.91; I^2 = 0%; very low certainty evidence).

Adherence to the exercise intervention

Of the eleven studies that reported the percentage of training sessions attended by the participants allocated to the intervention, the lowest adherence was reported as a median of 60% (Cooke 2018), and the highest was a mean adherence of 88% (Kouidi 2008).

Exercise-related adverse events

It is uncertain whether exercise training is safe for adults undergoing maintenance dialysis because the certainty of this evidence is very low. Seven studies reported there were no exercise-related adverse events (AVANTE-HEMO 2020; Chen 2010; Cho 2018; DIALY-SIZE 2016; IHOPE 2019; Uchiyama 2019; Wu 2014d) within a total of 171 participants assigned to the exercise intervention. One study reported 6/26 participants assigned to the intervention presented exercise-related symptoms including shortness of breath, soreness, lower extremity pain, cramping and fatigue (Sheshadri 2020). Furthermore, two participants experienced chest pain during the intervention. One study reported that one of the six exercising participants presented with knee joint pain (CHAIR 2015).



Other outcomes

Meta-analysis for the outcomes that were frequently reported but not identified as important to patients, or previously included are available as forest plots in Analysis 1.11 to Analysis 1.30 for any exercise training, Analysis 2.10 to Analysis 2.28 for aerobic training, Analysis 3.8 to Analysis 3.22 for resistance training, and Analysis 4.8 to Analysis 4.23 for combined aerobic and resistance training but will not be discussed further.

DISCUSSION

Summary of main results

This review of the evidence supporting exercise training for adults undergoing maintenance dialysis included 89 studies involving randomising 4291 participants; 77 studies involving 3846 participants contributed to the meta-analyses. The exercise programs, a complex intervention, were heterogeneous and varied in type, intensity, duration, frequency of sessions and timing in relation to dialysis treatments. Interventions within subtypes of exercises (aerobic, resistance or a combination of the two) were more comparable; however, the duration of the intervention remained highly variable. Only one study had long-term follow-up after the completion of the study.

A single study reported death but was not sufficiently powered to assess it and no study reported long-term cardiovascular events. Compared to no or sham exercise, any exercise for two to 12 months may reduce fatigue in adults undergoing maintenance dialysis. Importantly, compared to no or sham exercise, any exercise training for two to 12 months is likely to significantly improve depression in adults undergoing maintenance dialysis, particularly when the intervention is sustained for longer than four months. Compared to no or sham exercise, any exercise training for two to six months is also likely to substantially improve functional capacity which has been associated with survival in people receiving dialysis treatments (DeOreo 1997; Knight 2003). Furthermore, compared to no or sham exercise, any exercise training for three to 12 months may increase the physical component of HRQoL. Compared to no or sham exercise, any exercise training for three to 12 months may lead to lesser pain. However, the 95% CI indicated that exercise training might make little or no difference in the level of pain. It is uncertain whether exercise training improves the mental component of HRQoL or resting blood pressure because the certainty of this evidence is very low.

Comparisons of one type of exercise to another were limited by the number of studies reporting patients-important outcomes. We observed a differential effect of the type of exercise training on resting blood pressure, but the certainty of the evidence was very low.

Overall completeness and applicability of evidence

The current review is a comprehensive assessment of the effects of structured exercise training in adults undergoing maintenance dialysis. While it includes a vast number of studies covering aerobic, resistance and combined aerobic and resistance training, many uncertainties remain. Only seven studies included participants undergoing PD. We therefore cannot conclude on the impact of exercise training in this population. Secondly, the inclusion criteria were often stringent, excluding patients with extensive comorbidities. Furthermore, participants had to be able to perform

some level of exercise from baseline, thereby excluding the frailest patients. The conclusions of the review therefore cannot be applied to the debilitated dialysis patient with a heavy burden of comorbidity, loss of autonomy, physical limitation, or cognitive decline.

Most exercise interventions were conducted during the dialysis treatments. While some patients might be fearful of exercising during dialysis, little is known about the effect and feasibility of home-based exercise training for this population.

The interventions were overall of short duration, with only 10 interventions lasting longer than six months. We may have observed effects of greater magnitude where the interventions were more sustained.

Patients-important outcomes were under-represented, with many studies focusing on biomarkers and measures of exercise capacity. A single study reported long-term outcomes and death but was insufficiently powered to do so. The long-term impacts of exercise training in adults undergoing dialysis, therefore, remain unknown at this point.

Finally, the second objective of the review, which is to inform the design of exercise interventions that maximise the benefits for adults undergoing dialysis, could not be achieved due to the low number of studies reporting patient-important outcomes. A network meta-analysis, including the studies that compared one exercise intervention to another without necessarily including a no or sham exercise control group, would better address this aim.

Quality of the evidence

In general, the quality of evidence was low to very low due to the high risk of bias, the short duration of the interventions and follow-up and the low number of participants in the included studies.

Regarding the internal validity of the included studies, a majority did not report the methods of randomisation and concealment of the allocation. Blinding of participants was generally not feasible in this review due to the nature of the intervention, and only four studies attempted to blind the participants using a sham intervention. Outcome assessors were also rarely blinded to treatment allocation, and a majority of the studies were at high or unclear risk of attrition bias. Overall, the quality of the included studies was low, and the certainty of the evidence for all the outcomes was downgraded by one level for the high risk of bias in the included studies.

The interventions were of short duration with more than half lasting three months or less. The reason behind the overall short duration of the interventions and the lack of long-term outcomes may be the complexity of the intervention, a lack of adherence to the exercise intervention or costs. Furthermore, imprecision was a significant issue with most included studies relying on small convenience samples.

Potential biases in the review process

We searched the Cochrane Kidney and Transplant Specialised Register, which includes trial registries and hand-searched conference abstracts (grey literature). However, some studies may have been reported only in exercise science conference proceedings or in conference proceedings in languages other



than English and, therefore, missed by the Cochrane Kidney and Transplant Specialised Register. Despite our efforts to estimate means and SD from medians and ranges and impute missing SD, some studies still reported insufficient information for their results to be included in the meta-analyses, which could lead to biases in the pooled estimates of effect. Finally, while the lack of blinding is likely to affect subjective outcomes more substantially than objective outcomes, the definition of objective versus subjective outcome is subject to interpretation, which could affect the level of certainty of the evidence presented in this review.

Agreements and disagreements with other studies or reviews

We have identified 17 systematic reviews of exercise training interventions relating to adults undergoing dialysis published in the past five years (Barcellos 2015; Bessa 2015; Chan 2016; Chung 2017; Clarkson 2019; Ferrari 2020; Gomes 2018; Heiwe 2014; Huang 2019; Pu 2019; Qiu 2017; Salhab 2019; Scapini 2019; Sheng 2014; Song 2018; Young 2018; Zhao 2019). They included between nine and 59 studies. The considerably larger number of studies included in the current review was probably due to broader inclusions criteria and our search of the grey literature. One review reported on fatigue and, like us, found an improvement with exercise training (Zhao 2019). Eleven reviews reported on the physical component of HRQoL, of which 10 found improvement with exercise (Barcellos 2015; Chan 2016; Chung 2017; Gomes 2018; Heiwe 2014; Huang 2019; Pu 2019; Salhab 2019; Thompson 1996; Zhao 2019) and one observed no effect (Young 2018). Ten reviews reported on the mental component of HRQoL, of which six found it unchanged by exercise training (Chan 2016; Chung 2017; Gomes 2018; Pu 2019; Sheng 2014; Young 2018) and three found it improved (Huang 2019; Salhab 2019; Zhao 2019). All of the five reviews that reported depression as an outcome found it improved with exercise training (Barcellos 2015; Gomes 2018; Pu 2019; Song 2018; Zhao 2019). Of the 10 reviews that reported the 6MWT, all but one that was focusing solely on resistance training (Chan 2016) concluded that exercise improved walking capacity (Chung 2017; Clarkson 2019; Ferrari 2020; Gomes 2018; Heiwe 2014; Huang 2019; Pu 2019; Sheng 2014; Young 2018). Two of the three studies that reported on the Sit-To-Stand test also concluded to improvement with exercise (Chan 2016; Clarkson 2019; Sheng 2014). Eight studies reported resting blood pressure, of which four observed improved SBP and DBP with exercise (Ferrari 2020; Pu 2019; Scapini 2019; Sheng 2014) and four did not observe a significant effect (Heiwe 2014; Huang 2019; Qiu 2017; Young 2018). No reviews reported death, cardiovascular events, or pain.

AUTHORS' CONCLUSIONS

Implications for practice

- Exercise training of any type for two to 12 months is likely to improve depressive symptoms in adults undergoing dialysis. Low certainty evidence suggests that extending the intervention for more than four months may provide additional benefits. There is no data as to whether the effect of exercise training on depressive symptoms persist beyond the duration of the intervention.
- Exercise training of any type for two to 12 months may reduce fatigue and improve the physical component of QoL in adults undergoing maintenance dialysis.

- Exercise training of any type for three to 12 months may reduce pain in adults undergoing maintenance dialysis slightly. However, the 95% CI indicates that exercise training might make little or no difference in the level of pain.
- Exercise training of any type for two to six months may increase patient functional capacity.
- Existing studies of exercise training in adults undergoing dialysis were not designed to assess long-term outcomes such as death and cardiovascular events.
- The level of certainty is very low for the effect of exercise training on mortality, the mental component of HR-QoL and resting blood pressure.
- There is little to no information on the effect of exercise training for adults undergoing PD.
- There is little to no information regarding the sustained effects of exercise training beyond the duration of the exercise program.
- Adverse effects of exercise training in adults undergoing dialysis are rarely reported and poorly defined. The evidence for the safety of exercise training in this population is therefore very uncertain.

Implications for research

- Studies of exercise training for adults undergoing dialysis should prioritise outcomes that are important to patients, their caregivers and health professionals, including death, cardiovascular events, fatigue, and pain.
- Long-term studies with extended follow-up periods are needed to assess critical outcomes, including death and cardiovascular disease, and to assess the persistence of the effect beyond the intervention. For long-term studies of an exercise intervention to be successful, strategies to enhance adherence to the interventions should be sought.
- Studies of exercise training for adults undergoing dialysis should put measures in place to minimise the effects of the lack of blinding in the participants, particularly for patient-reported outcomes.
- Studies should avoid convenience sampling and guide their recruitment on sample size and power calculations based on an estimate of a clinically relevant effect.
- Dialysis patients that are frail or with a heavy burden of comorbidities are an important subpopulation for which dedicated studies of exercise intervention should be considered.
- Studies of exercise training for adults undergoing dialysis must thoroughly assess and report adverse effects related to the intervention.

ACKNOWLEDGEMENTS

We want to thank...

- Susanne Heiwe and Stefan Jacobson who were the authors of the first version of this review
- Elisabeth Hodson for her guidance in conducting a Cochrane systematic review
- Fiona Russell for the administrative support
- Gail Higgins for her assistance with the search in The Cochrane Kidney and Transplant Specialised Register
- Yeoungjee Cho for translating one included article from Korean



- Angela Ju for her input as a content expert in the assessment of fatigue, depression and QoL in adults undergoing dialysis
- Jonathan Levesque for his input as a content expert in rehabilitation and the assessment of functional capacity.
- The authors are grateful to the following peer reviewers for their time and comments: Maristela Bohlke (Postgraduate Program in Health and Behaviour, Universidade Catolica de Pelotas, Brazil) and Dr Matt Hall (Nottingham University Hospitals NHS Trust, Nottingham, UK).



REFERENCES

References to studies included in this review

Abreu 2017 (published data only)

Abreu CC, Cardozo LF, Stockler-Pinto MB, Esgalhado M, Barboza JE, Frauches R, et al. Does resistance exercise performed during dialysis modulate Nrf2 and NF-kappaB in patients with chronic kidney disease? *Life Sciences* 2017;**188**:192-7. [MEDLINE: 28887058]

Abundis Mora 2017 {published data only}

Abundis Mora GJ, Garcia-Garcia G, Brambila DM, Jimenez Cornejo MC, Renoirte K, Chavez J. Effect of physical training on echocardiographic parameters during hemodialysis: a randomized clinical trial. [abstract no: SA-PO832]. *Journal of the American Society of Nephrology* 2017;**28**(Abstract Suppl):895. [EMBASE: 633698733]

ACTINUT 2013 {published data only}

Hristea D, Deschamps T, Paris A, Lefrancois G, Collet V, Savoiu C, et al. Combining intra-dialytic exercise and nutritional supplementation in malnourished older haemodialysis patients: towards better quality of life and autonomy. *Nephrology* 2016;**21**(9):785-90. [MEDLINE: 26890997]

Hristea D, Paris A, Lefrancois G, Volteau C, Savoiu C, Ozenne S, et al. Randomized controlled trial on the effects of a sixmonth intra-dialytic physical activity program and adequate nutritional support on protein-energy wasting, physical functioning and quality of life in chronic hemodialysis patients-ACTINUT [abstract no: SP655]. *Nephrology Dialysis Transplantation* 2014;**29**(Suppl 3):iii290-300. [EMBASE: 71492282]

Magnard J, Deschamps T, Cornu C, Paris A, Hristea D. Effects of a six-month intradialytic physical ACTIvity program and adequate NUTritional support on protein-energy wasting, physical functioning and quality of life in chronic hemodialysis patients: ACTINUT study protocol for a randomised controlled trial. *BMC Nephrology* 2013;**14**:259. [MEDLINE: 24279747]

Afshar 2010 {published data only}

Afshar R, Shegarfy L, Shavandi N, Sanavi S. Effects of aerobic exercise and resistance training on lipid profiles and inflammation status in patients on maintenance hemodialysis. *Indian Journal of Nephrology* 2010;**20**(4):185-9. [MEDLINE: 21206679]

Afshar 2011 {published data only}

Afshar R, Emany A, Saremi A, Shavandi N, Sanavi S. Effects of intradialytic aerobic training on sleep quality in hemodialysis patients: a randomized controlled trial [abstract no: PO02-061]. *Nephrology* 2010;**15**(Suppl 3):104. [EMBASE: 70467629]

Afshar R, Emany A, Saremi A, Shavandi N, Sanavi S. Effects of intradialytic aerobic training on sleep quality in hemodialysis patients. *Iranian Journal of Kidney Diseases* 2011;**5**(2):119-23. [MEDLINE: 21368391]

Akiba 1995 (published data only)

Akiba T, Matsui N, Marumo F. Erythropoietin can partially increase, but can not maintain exercise capacity without exercise training [abstract]. In: XIIth International Congress of Nephrology; 1993 June 13-18; Jerusalem, Israel. 1993:298.

Akiba T, Matsui N, Shinohara S, Fujiwara H, Nomura T, Marumo F. Effects of recombinant human erythropoietin and exercise training on exercise capacity in hemodialysis patients. *Artificial Organs* 1995;**19**(12):1262-8. [MEDLINE: 8967886]

Amini 2016 (published data only)

Amini E, Goudarzi I, Masoudi R, Ahmadi A, Momeni A. Effect of progressive muscle relaxation and aerobic exercise on anxiety, sleep quality, and fatigue in patients with chronic renal failure undergoing hemodialysis. *International Journal of Pharmaceutical & Clinical Research* 2016;**8**(12):1634-9. [EMBASE: 614181031]

AVANTE-HEMO 2020 {published data only}10251828

Martin-Alemany G, Espinosa-Cuevas ML, Perez-Navarro M, Wilund KR, Miranda-Alatriste P, Cortes-Perez M, et al. Effect of oral nutritional supplementation with and without exercise on nutritional status and physical function of adult hemodialysis patients: a parallel controlled clinical trial (AVANTE-HEMO Study). *Journal of Renal Nutrition* 2020;**30**(2):126-36. [MEDLINE: 31607547]

Bennett 2013 (published data only)

Bennett PN, Daly RM, Fraser SF, Haines T, Barnard R, Ockerby C, et al. The impact of an exercise physiologist coordinated resistance exercise program on the physical function of people receiving hemodialysis: a stepped wedge randomised control study. *BMC Nephrology* 2013;**14**:204. [MEDLINE: 24070232]

Bennett PN, Fraser S, Barnard R, Haines T, Ockerby C, Street M, et al. Effects of an intradialytic resistance training programme on physical function: a prospective stepped-wedge randomized controlled trial. *Nephrology Dialysis Transplantation* 2016;**31**(8):1302-9. [MEDLINE: 26715763]

Burrows 2018 {published data only}

Burrows B, Perez LM, Chan L, Harris AP, Barnes JL, Wilund K. The effect of intradialytic and home-based exercise on physical function and quality of life in hemodialysis patients [abstract no: FR-P0542]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):560. [EMBASE: 633735934]

Carmack 1995 {published data only}

* Carmack CL, Amaral-Melendez M, Boudreaux E, Brantley PJ, Franks JD, Jones GN, et al. Exercise as a component in the physical and psychological rehabilitation of hemodialysis patients. *International Journal of Rehabilitation & Health* 1995;**1**(1):13-23. [2010258777]

CHAIR 2015 (published data only)

Matsufuji S, Shoji T, Yano Y, Tsujimoto Y, Kishimoto H, Tabata T, et al. Effect of chair stand exercise on activity of daily living: a



randomized controlled trial in hemodialysis patients. *Journal of Renal Nutrition* 2015;**25**(1):17-24. [MEDLINE: 25194621]

Chang 2010 (published data only)

Chang Y, Cheng SY, Lin M, Gau FY, Chao YF. The effectiveness of intradialytic leg ergometry exercise for improving sedentary life style and fatigue among patients with chronic kidney disease: a randomized clinical trial. *International Journal of Nursing Studies* 2010;**47**(11):1383-8. [MEDLINE: 20537645]

Chen 2010 (published data only)

Chen J, Coomber S, Ng L, MacKinnon R, Moorthi R, Liangos O, et al. Effect of intradialytic low-intensity resistance exercise training on functional capacity, strength and quality of life in functionally impaired ESRD patients [abstract no: 069]. *Journal of Renal Nutrition* 2008;**18**(3 Suppl 1):S19.

Chen JL, Godfrey S, Ng TT, Moorthi R, Liangos O, Ruthazer R, et al. Effect of intra-dialytic, low-intensity strength training on functional capacity in adult haemodialysis patients: a randomized pilot trial. *Nephrology Dialysis Transplantation* 2010;**25**(6):1936-43. [MEDLINE: 20100734]

Cho 2018 {published data only}

Cho JH, Kim JC. Effect of intradialytic exercise on physical performance and echocardiographic findings in maintenance hemodialysis patients [abstract no: SA-OR076]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):99. [EMBASE: 633736859]

Cho JH, Kim JC. The effect of intradialytic exercise on daily physical activity and sleep quality in maintenance hemodialysis patients [abstract no: TH-PO1050]. *Journal of the American Society of Nephrology* 2016;**27**(Abstract Suppl):340A.

Cho JH, Lee JY, Lee S, Park H, Choi SW, Kim JC. Effect of intradialytic exercise on daily physical activity and sleep quality in maintenance hemodialysis patients. *International Urology & Nephrology* 2018;**50**(4):745-54. [MEDLINE: 29362960]

Cooke 2018 (published data only)

Cooke AB, Ta V, Iqbal S, Gomez YH, Mavrakanas T, Barre P, et al. The impact of intradialytic pedaling exercise on arterial stiffness: a pilot randomized controlled trial in a hemodialysis population. *American Journal of Hypertension* 2018;**31**(4):458-66. [MEDLINE: 29126178]

CYCLE-HD 2016 {published data only}11299707

Careless A, March D, Churchward D, Grantham C, Highton P, Tomlinson C, et al. Intradialytic exercise: a non-pharmacological solution to a uraemic problem? [abstract no: MP465]. Nephrology Dialysis Transplantation 2017;**32**(Suppl 3):iii599-600. [EMBASE: 617290883]

Graham-Brown MP, March DS, Churchward DR, Young HM, Dungey M, Lloyd S, et al. Design and methods of CYCLE-HD: improving cardiovascular health in patients with end stage renal disease using a structured programme of exercise: a randomised control trial. *BMC Nephrology* 2016;**17**(1):69. [MEDLINE: 27391774]

Graham-Brown MP, March DS, Hull KL, Wormleighton JV, Young R, McCann GP, et al. The effects of a 6-month structured programme of intradialytic cycling on cardiovascular remodelling, myocardial fibrosis, and aortic stiffness: results from the CYCLE-HD study [abstract no: SA-OR064]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):96. [EMBASE: 633771575]

March DS, Grantham CE, Graham-Brown MP, Young HM, Cooper N, Burton J. A six month program of intradialytic exercise is effective in reducing length of hospital stay in hemodialysis patients [abstract no: SA-PO787]. *Journal of the American Society of Nephrology* 2017;**28**(Abstract Suppl):882. [EMBASE: 633698553]

March DS, Hurt AW, Smith AC, Burton J. A 6-month program of intradialytic cycling results in a reduction in associated healthcare costs in patients receiving prevalent hemodialysis [abstract no: TH-PO256]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):181. [EMBASE: 633769797]

Tomlinson C, Churchward D, Grantham C, Young H, Highton P, Graham-Brown M, et al. A six month programme of intradialytic exercise improves resting heart rate in haemodialysis patients [abstract no: MP612]. *Nephrology Dialysis Transplantation* 2017;**32**(Suppl 3):iii658-9. [EMBASE: 617291352]

Dashtidehkordi 2019 {published data only}

Dashtidehkordi A, Shahgholian N, Attari F. Exercise during hemodialysis and health promoting behaviors: a clinical trial. *BMC Nephrology* 2019;**20**(1):96. [MEDLINE: 30890122]

Deligiannis 1999 {published data only}

* Deligiannis A, Kouidi E, Tourkantonis A. Effects of physical training on heart rate variability in patients on hemodialysis. American Journal of Cardiology 1999;**84**(2):197-202. [MEDLINE: 10426340]

Deligiannis 1999a {published data only}

* Deligiannis A, Kouidi E, Tassoulas E, Gigis P, Tourkantonis A, Coats A. Cardiac effects of exercise rehabilitation in hemodialysis patients. *International Journal of Cardiology* 1999;**70**(3):253-66. [MEDLINE: 10501340]

Deligiannis A, Kouidi E, Vassiliou S, Tourkantonis A. The effects of exercise training on the autonomic control of heart rate in hemodialysis patients [abstract]. *Nephrology Dialysis Transplantation* 1996;**11**(6):A175.

de Lima 2013 {published data only}

de Lima MC, de Lima Cicotoste C, da Silva Cardoso K, Forgiarini LA Jr, Monteiro MB, Dias AS. Effect of exercise performed during hemodialysis: strength versus aerobic. *Renal Failure* 2013;**35**(5):697-704. [MEDLINE: 23560491]

DePaul 2002 {published data only}

DePaul V, Moreland J, Eager T, Clase CM. The effectiveness of aerobic and muscle strength training in patients receiving hemodialysis and EPO: a randomized controlled trial. *American Journal of Kidney Diseases* 2002;**40**(6):1219-29. [MEDLINE: 12460041]



DIALY-SIZE 2016 {published data only}

Thompson S, Klarenbach S, Molzahn A, Lloyd A, Gabrys I, Haykowsky M, et al. Randomised factorial mixed method pilot study of aerobic and resistance exercise in haemodialysis patients: DIALY-SIZE! *BMJ Open* 2016;**6**(9):e012085. [MEDLINE: 27601500]

Thompson SE, Klarenbach S, Molzahn A, Haykowsky M, Lloyd A, Tonelli M. A randomized, factorial pilot study to evaluate the feasibility of an intradialytic exercise intervention (DIALY-SIZE!) [abstract no: PUB353]. *Journal of the American Society of Nephrology* 2015;**26**(Abstract Suppl):970A.

Dobsak 2012 (published data only)

Dobsak P, Homolka P, Svojanovsky J, Reichertova A, Soucek M, Novakova M, et al. Intra-dialytic electrostimulation of leg extensors may improve exercise tolerance and quality of life in hemodialyzed patients. *Artificial Organs* 2012;**36**(1):71-8. [MEDLINE: 21848929]

Dong 2011 (published data only)

Dong J, Sundell MB, Pupim LB, Wu P, Shintani A, Ikizler TA. Resistance exercise does not augment long-term benefits of intradialytic oral nutritional supplementation in chronic hemodialysis patients [abstract no: SA-FC329]. *Journal of the American Society of Nephrology* 2009;**20**(Abstract Suppl):77A.

Dong J, Sundell MB, Pupim LB, Wu P, Shintani A, Ikizler TA. The effect of resistance exercise to augment long-term benefits of intradialytic oral nutritional supplementation in chronic hemodialysis patients. *Journal of Renal Nutrition* 2011;**21**(2):149-59. [MEDLINE: 20580251]

EXCITE 2014 {published data only}

Baggetta R, Bolignano D, Torino C, Manfredini F, Aucella F, Barilla A, et al. Fitness for entering a simple exercise program and mortality: a study corollary to the exercise introduction to enhance performance in dialysis (EXCITE) trial. *Kidney & Blood Pressure Research* 2014;**39**(2-3):197-204. [MEDLINE: 25118055]

Baggetta R, D'Arrigo G, Torino C, ElHafeez SA, Manfredini F, Mallamaci F, et al. Effect of a home based, low intensity, physical exercise program in older adults dialysis patients: a secondary analysis of the EXCITE trial. *BMC Geriatrics* 2018;**18**(1):248. [MEDLINE: 30342464]

Mallamaci F, Manfredini F, Baggetta R, D'Arrigo G, Bolignano D, Bertoli S, et al. A personalized walking exercise program reduces the risk of hospitalization in dialysis patients [abstract no: 3410]. *Nephrology Dialysis Transplantation* 2017;**32**(Suppl 3). [CENTRAL: CN-01657758]

Mallamaci F, Manfredini F, Bolignano D, Bertoli S, Messa P, Zuccala A, et al. A personalized, low-intensity, easy to implement, home exercise program improves physical performance in dialysis patients: the exercise introduction to enhance performance in dialysis (EXCITE) trial [abstract no: FR-OR049]. *Journal of the American Society of Nephrology* 2014;**25**(Abstract Suppl):57A.

Manfredini F, Bolignano D, Rastelli S, Barilla A, Bertoli S, Ciurlino D, et al. Low intensity, home-based exercise improves physical capacity in dialysis patients: The EXCITE study (exercise

introduction to enhance performance in dialysis) [abstract]. *Nephrology Dialysis Transplantation* 2012;**27**(Suppl 2):ii125. [EMBASE: 70765654]

Manfredini F, Lamberti N, Malagoni AM, Felisatti M, Zuccala A, Torino C, et al. The role of deconditioning in the end-stage renal disease myopathy: physical exercise improves altered resting muscle oxygen consumption. *American Journal of Nephrology* 2015;**41**(4-5):329-36. [MEDLINE: 26067552]

Manfredini F, Mallamaci F, D'Arrigo G, Baggetta R, Bolignano D, Torino C, et al. Exercise in patients on dialysis: a multicenter, randomized clinical trial [Erratum in: J Am Soc Nephrol. 2018 Jul;29(7):2028; PMID: 29793961]. *Journal of the American Society of Nephrology* 2016;**28**(4):1259-68. [MEDLINE: 27909047]

Pomidori L, Lamberti N, Malagoni AM, Manfredini F, Pozzato E, Felisatti M, et al. Respiratory muscle impairment in dialysis patients: can minimal dose of exercise limit the damage? a preliminary study in a sample of patients enrolled in the EXCITE trial. *Journal of Nephrology* 2016;**29**(6):863-9. [MEDLINE: 27312989]

Torino C, Manfredini F, Bolignano D, Aucella F, Baggetta R, Barilla A, et al. Physical performance and clinical outcomes in dialysis patients: a secondary analysis of the EXCITE trial. *Kidney & Blood Pressure Research* 2014;**39**(2-3):205-11. [MEDLINE: 25118076]

Fernandes 2019 {published data only}

Fernandes AO, Sens YA, Xavier VB, Miorin LA, Alves VL. Functional and respiratory capacity of patients with chronic kidney disease undergoing cycle ergometer training during hemodialysis sessions: a randomized clinical trial. *International Journal of Nephrology* 2019;**2019**:7857824. [MEDLINE: 30805216]

Frey 1999 {published data only}

Frey S, Mir AR, Lucas M. Visceral protein status and caloric intake in exercising versus nonexercising individuals with endstage renal disease. *Journal of Renal Nutrition* 1999;**9**(2):71-7. [MEDLINE: 10089262]

Frih 2017a {published data only}

Frih B, Jaafar H, Mkacher W, Salah ZB, Hammami M, Frih A. The effect of interdialytic combined resistance and aerobic exercise training on health related outcomes in chronic hemodialysis patients: the Tunisian randomized controlled study. *Frontiers in Physiology* 2017;**8**(May):288. [EMBASE: 617015625]

Giannaki 2013a {published data only}

Giannaki CD, Hadjigeorgiou GM, Karatzaferi C, Maridaki MD, Koutedakis Y, Founta P, et al. A single-blind randomized controlled trial to evaluate the effect of 6 months of progressive aerobic exercise training in patients with uraemic restless legs syndrome. *Nephrology Dialysis Transplantation* 2013;**28**(11):2834-40. [MEDLINE: 23929523]

Goldberg 1983 (published data only)

Goldberg AP, Geltman EM, Hagberg JM, Gavin JR 3rd, Delmez JA, Carney RM, et al. Therapeutic benefits of exercise



training for hemodialysis patients. *Kidney International - Supplement* 1983;**16**:S303-9. [MEDLINE: 6588267]

Groussard 2015 {published data only}

Groussard C, Rouchon-Isnard M, Coutard C, Romain F, Malarde L, Lemoine-Morel S, et al. Beneficial effects of an intradialytic cycling training program in patients with end-stage kidney disease. *Applied Physiology, Nutrition, & Metabolism [Physiologie Appliquee, Nutrition et Metabolisme]* 2015;**40**(6):550-6. [MEDLINE: 25955722]

Harter 1985 (published data only)

Carney RM, Templeton B, Hong BA, Harter HR, Hagberg JM, Schechtman KB, et al. Exercise training reduces depression and increases the performance of pleasant activities in hemodialysis patients. *Nephron* 1987;**47**(3):194-8. [MEDLINE: 3317091]

Goldberg AP, Geltman EM, Gavin JR 3rd, Carney RM, Hagberg JM, Delmez JA, et al. Exercise training reduces coronary risk and effectively rehabilitates hemodialysis patients. *Nephron* 1986;**42**(4):311-6. [MEDLINE: 3960242]

* Harter HR, Goldberg AP. Endurance exercise training. An effective therapeutic modality for hemodialysis patients. *Medical Clinics of North America* 1985;**69**(1):159-75. [MEDLINE: 3883073]

IHOPE 2019 {published data only}

Jeong JH, Biruete A, Tomayko EJ, Wu PT, Fitschen P, Chung HR, et al. Results from the randomized controlled IHOPE trial suggest no effects of oral protein supplementation and exercise training on physical function in hemodialysis patients. *Kidney International* 2019;**96**(3):777-86. [MEDLINE: 31200945]

Wilund KR, Jeong JH, Fitschen PJ, Wu PT, Tomayko EJ, Chung HR, et al. Efficacy of intradialytic protein supplementation and exercise training [abstract no: MO069]. *Nephrology Dialysis Transplantation* 2016;**31**(Suppl 1):i58-9. [EMBASE: 72326051]

Johansen 2006 (published data only)

Johansen KL, Painter PL, Gordon P, Doyle J, Sakkas GK. Effects of resistance exercise training and anabolic steroid treatment among hemodialysis patients: results of the NEXT study [abstract no: SU-PO382]. *Journal of the American Society of Nephrology* 2004;**15**(Oct):617A. [CENTRAL: CN-00550563]

Johansen KL, Painter PL, Sakkas GK, Gordon P, Doyle J, Shubert T. Effects of resistance exercise training and nandrolone decanoate on body composition and muscle function among patients who receive hemodialysis: a randomized, controlled trial. *Journal of the American Society of Nephrology* 2006;**17**(8):2307-14. [MEDLINE: 16825332]

Jong 2004 (published data only)

* Jong KH, Jeong LS, Hyun YT, Suk PY, Young JM, Eun AS, et al. Effects of walking exercise for health status in patients on continuous ambulatory peritoneal dialysis [abstract no: SP504]. In: 41st Congress. European Renal Association European Dialysis and Transplantation Association; 2004 May 15-18; Lisbon, Portugal. 2004:188. [CENTRAL: CN-00509256]

Koh 2009 (published data only)

Koh KP, Fassett RG, Sharman JE, Coombes JS, Williams AD. Effect of intradialytic versus home-based aerobic exercise training on physical function and vascular parameters in hemodialysis patients: a randomized pilot study. *American Journal of Kidney Diseases* 2010;**55**(1):88-99. [MEDLINE: 19932545]

Koh KP, Fassett RG, Sharman JE, Coombes JS, Williams AD. Intradialytic versus home-based exercise training in hemodialysis patients: a randomised controlled trial. *BMC Nephrology* 2009;**10**:2. [MEDLINE: 19178747]

Konstantinidou 2002 {published data only}

* Konstantinidou E, Koukouvou G, Kouidi E, Deligiannis A, Tourkantonis A. Exercise training in patients with endstage renal disease on hemodialysis: comparison of three rehabilitation programs. *Journal of Rehabilitation Medicine* 2002;**34**(1):40-5. [MEDLINE: 11900261]

Kopple 2007 (published data only)

Kopple JD, Wang H, Casaburi R, Fournier M, Lewis MI, Taylor W, et al. Exercise in maintenance hemodialysis patients induces transcriptional changes in genes favoring anabolic muscle. *Journal of the American Society of Nephrology* 2007;**18**(11):2975-86. [MEDLINE: 17942969]

Koufaki 2002 (published data only)

Koufaki P, Mercer TH, Naish PF. Effects of exercise training on aerobic and functional capacity of end-stage renal disease patients. *Clinical Physiology & Functional Imaging* 2002;**22**(2):115-24. [MEDLINE: 12005153]

Naish PF, Koufaki P, Mercer T. Nutritional status is associated with physical functioning and quality of life in end stage renal disease patients [abstract no: A1755]. *Journal of the American Society of Nephrology* 2001;**12**(Program & Abstracts):341A. [CENTRAL: CN-00446891]

Koufaki 2003 (published data only)

* Koufaki P, Naish P, Mercer T. Exercise training augments EPOinduced changes in exercise tolerance and functional capacity of dialysis patients [abstract no: M497]. *Nephrology Dialysis Transplantation* 2003;**18**(Suppl 4):157. [CENTRAL: CN-00446168]

Kouidi 1997 {published data only}

Kouidi E, Iacovides A, Iordanidis P, Vassiliou S, Deligiannis A, Ierodiakonou C, et al. Exercise renal rehabilitation program: psychosocial effects. *Nephron* 1997;**77**(2):152-58. [MEDLINE: 9346380]

Kouidi E, Iacovides A, Iordinides P, Vassiliou S, Deligiannis A, Ierodiakonou C, et al. Exercise renal rehabilitation program (ERRP): psychosocial effects [abstract]. *Nephrology Dialysis Transplantation* 1995;**10**(6):1015. [CENTRAL: CN-00261120]

Kouidi 2003 (published data only)

* Kouidi E, Grekas D, Koukouvou G, Konstantinidou E, Kalevrosoglou J, Deligiannis A. The effects of exercise training on variables predisposing dialysis patients to cardiac death [abstract no: W557]. *Nephrology Dialysis Transplantation* 2003;**18**(Suppl 4):724. [CENTRAL: CN-00446169]



Kouidi 2004a {published data only}

* Kouidi E, Konstantinidou E, Kalevrosoglou J, Grekas D, Deligiannis A. Cardiac systolic and diastolic function in aging dialysis patients following exercise training [abstract no: MP366]. In: 41st Congress. European Renal Association. European Dialysis and Transplantation Association; 2004 May 15-18; Lisbon, Portugal. 2004:354. [CENTRAL: CN-00509286]

Kouidi 2005 (published data only)

* Kouidi E, Grekas DM, Ouzouni SG, Iacovides AI, Deligiannis AP. Psychosocial effects of exercise training during dialysis in uraemic patients under epoietin therapy [abstract no: MP338]. Nephrology Dialysis Transplantation 2005;**20**(Suppl 5):v314. [CENTRAL: CN-01658037]

Kouidi 2008 {published data only}

Kouidi E, Sioulis A, Koukouvou G, Konstantinidou E, Grekas D, Deligiannis A. The effects of exercise training on non-evasive variables for cardiac risk stratification in hemodialysis patients [abstract no: SP662]. *Nephrology Dialysis Transplantation* 2006;**21**(Suppl 4):iv238.

Kouidi EJ, Grekas DM, Deligiannis AP. Effects of exercise training on noninvasive cardiac measures in patients undergoing long-term hemodialysis: a randomized controlled trial. *American Journal of Kidney Diseases* 2009;**54**(3):511-21. [MEDLINE: 19646801]

Petraki M, Kouidi E, Grekas D, Deligiannis A. Effects of exercise training during hemodialysis on cardiac baroreflex sensitivity. *Clinical Nephrology* 2008;**70**(3):210-19. [MEDLINE: 18793562]

Kouidi 2010 (published data only)

Kouidi E, Grekas D, Iakovides A, Deligiannis A, Tourkantonis A. Depression, heart rate variability and exercise training in dialysis patients [abstract no: T388]. *Nephrology Dialysis Transplantation* 2002;**17**(Suppl 1):296.

Kouidi E, Karagiannis V, Grekas D, Iakovides A, Kaprinis G, Tourkantonis A, et al. Depression, heart rate variability, and exercise training in dialysis patients. *European Journal of Cardiovascular Prevention & Rehabilitation* 2010;**17**(2):160-7. [MEDLINE: 19745744]

Lee 2001 {published data only}

Lee YK, Kim C, Pyo JH, Kim CH, Ji JW. Endurance exercise training before hemodialysis: an effective therapeutic modality for end-stage renal disease patients. *Korean Journal of Nephrology* 2001;**20**(2):290-97.

Liao 2016 {published data only}

Liao MT, Liu WC, Lin FH, Huang CF, Chen SY, Liu CC, et al. Intradialytic aerobic cycling exercise alleviates inflammation and improves endothelial progenitor cell count and bone density in hemodialysis patients. *Medicine* 2016;**95**(27):e4134. [MEDLINE: 27399127]

Wu CC, Lu KC. Intradialytic aerobic cycling exercise improve inflammation status, endothelial progenitor cells and bone density in patients with end stage renal disease on maintenance hemodialysis [abstract no: TH-PO930]. *Journal of the American Society of Nephrology* 2015;**26**(Abstract Suppl):308a.

Ma 2018 {published data only}

Ma Y. The effects of exercise rehabilitation on physiological function and quality of life in maintain hemodialysis patients-a multicenter randomized controlled study [abstract no: SP401]. Nephrology Dialysis Transplantation 2018;33(Suppl 1):i482. [EMBASE: 622605253]

Makhlough 2012 (published data only)

Makhlough A, Ilali E, Mohseni R, Shahmohammadi S. Effect of intradialytic aerobic exercise on serum electrolytes levels in hemodialysis patients. *Iranian Journal of Kidney Diseases* 2012;**6**(2):119-23. [MEDLINE: 22388610]

Mohseni R, Emami Zeydi A, Ilali E, Adib-Hajbaghery M, Makhlough A. The effect of intradialytic aerobic exercise on dialysis efficacy in hemodialysis patients: a randomized controlled trial. *Oman Medical Journal* 2013;**28**(5):345-9. [MEDLINE: 24044062]

Marchesan 2016 (published data only)

Marchesan M, de Rosso Krug R, da Costa e Silva JR, Rodrigues Barbosa A, Rombaldi AJ. Physical exercise modifies the functional capacity of elderly patients on hemodialysis [Exercício físico modifica a aptidão funcional de idosos em hemodiálise]. *Fisioterapia em Movimento* 2016;**29**(2):351-9.

Marinho 2016 (published data only)

Marinho SM, Mafra D, Pelletier S, Hage V, Teuma C, Laville M, et al. In hemodialysis patients, intradialytic resistance exercise improves osteoblast function: a pilot study. *Journal of Renal Nutrition* 2016;**26**(5):341-5. [MEDLINE: 27113628]

Martin-Alemany 2016 {published data only}

Martin-Alemany G, Valdez-Ortiz R, Olvera-Soto G, Gomez-Guerrero I, Aguire-Esquivel G, Cantu-Quintanilla G, et al. The effects of resistance exercise and oral nutritional supplementation during hemodialysis on indicators of nutritional status and quality of life. *Nephrology Dialysis Transplantation* 2016;**31**(10):1712-20. [MEDLINE: 27510532]

Martins do Valle 2020 (published data only)

Martins do Valle F, Valle Pinheiro B, Almeida Barros AA, Ferreira Mendonca W, de Oliveira AC, de Oliveira Werneck G, et al. Effects of intradialytic resistance training on physical activity in daily life, muscle strength, physical capacity and quality of life in hemodialysis patients: a randomized clinical trial. *Disability & Rehabilitation* 2020;**42**(25):3638-44. [MEDLINE: 31034264]

Matsumoto 2007 {published data only}

Matsumoto Y, Furuta A, Furuta S, Miyajima M, Sugino T, Nagata K, et al. The impact of pre-dialytic endurance training on nutritional status and quality of life in stable hemodialysis patients (Sawada study). *Renal Failure* 2007;**29**(5):587-93. [MEDLINE: 17654322]

McAdams-DeMarco 2018 (published data only)

McAdams-DeMarco MA, Konel J, Warsame F, Ying H, Fernandez MG, Carlson MC, et al. Intradialytic cognitive and exercise training may preserve cognitive function. *KI Reports* 2018;**3**(1):81-8. [MEDLINE: 29340317]



McGregor 2018 (published data only)

McGregor G, Ennis S, Powell R, Hamborg T, Raymond NT, Owen W, et al. Feasibility and effects of intra-dialytic low-frequency electrical muscle stimulation and cycle training: a pilot randomized controlled trial. *PLoS ONE [Electronic Resource]* 2018;**13**(7):e0200354. [MEDLINE: 29995947]

Mitsiou 2015 (published data only)

Mitsiou M, Kouidi E, Liakopoulos V, Deligiannis A. Effects of music and exercise training during hemodialysis on the cardiac autonomic nervous system activity [abstract no: P0028]. *Nephrology Dialysis Transplantation* 2015;**30**(Suppl 3):iii15. [EMBASE: 72206309]

Mitsiou M, Kouidi EJ, Liakopoulos V, Deligiannis AP. Effects of music and exercise during hemodialysis on the cardiac autonomic nervous system activity [abstract no: TH-PO911]. *Journal of the American Society of Nephrology* 2015;**26**(Abstract Suppl):303A.

Miura 2015 (published data only)

Miura M, Hirayama A, Oowada S, Kohzuki M, Ito O. Effects of a renal rehabilitation exercise program in patients with CKD: controlled trial [abstract no: SP441]. *Nephrology Dialysis Transplantation* 2015;**30**(Suppl 3):iii524. [EMBASE: 72207767]

Miura M, Hirayama A, Owada S, Hirayama Y, Ito O, Kohzuki M. Multicenter trial of aerobic exercise in maintenance hemodialysis patients [abstract no: TH-PO931]. *Journal of the American Society of Nephrology* 2015;**26**(Abstract Suppl):308A.

Molsted 2004 (published data only)

Molsted S, Eidemak I, Sorensen HT, Kristensen JH. Five months of physical exercise in hemodialysis patients: effects on aerobic capacity, physical function and self-rated health. *Nephron* 2004;**96**(3):c76-81. [MEDLINE: 15056989]

Momeni 2014 (published data only)

Momeni A, Nematolahi A, Nasr M. Effect of intradialytic exercise on echocardiographic findings in hemodialysis patients. *Iranian Journal of Kidney Diseases* 2014;**8**(3):207-11. [MEDLINE: 24878943]

Mortazavi 2013 {published data only}

Mortazavi M, Vahdatpour B, Ghasempour A, Taheri D, Shahidi S, Moeinzadeh F, et al. Aerobic exercise improves signs of restless leg syndrome in end stage renal disease patients suffering chronic hemodialysis. *ScientificWorldJournal* 2013:628142. [MEDLINE: 24307876]

Mortazavi M, Vahdatpour B, Shahidi S, Ghasempour A, Taheri D, Dolatkhah S, et al. Aerobic exercise improves signs of restless leg syndrome in end stage renal disease patients suffering chronic hemodialysis [abstract no: FP627]. *Nephrology Dialysis Transplantation* 2012;**27**(Suppl 2):ii281. [EMBASE: 70766123]

Mortazavi M, Vahdatpour B, Shahidi S, Ghasempour A, Taheri S, Ghasami M. The effect of aerobic exercise on the symptoms of restless leg syndrome and quality of life in hemodialysis patients [abstract no: P407]. *Iranian Journal of Kidney Diseases* 2011;**5**(Suppl 2):63-4. [EMBASE: 70673915]

Olvera-Soto 2016 (published data only)

Olvera-Soto MG, Valdez-Ortiz R, Lopez Alvarenga JC, Espinosa-Cuevas ML. Effect of resistance exercises on the indicators of muscle reserves and handgrip strength in adult patients on hemodialysis. *Journal of Renal Nutrition* 2016;**26**(1):53-60. [MEDLINE: 26264173]

Ouzouni 2009 {published data only}

Ouzouni S, Kouidi E, Sioulis A, Grekas D, Deligiannis A. Effects of intradialytic exercise training on health-related quality of life indices in haemodialysis patients. *Clinical Rehabilitation* 2009;**23**(1):53-63. [MEDLINE: 19114437]

Painter 2002a {published data only}

Painter P, Moore G, Carlson L, Paul S, Myll J, Phillips W, et al. Effects of exercise training plus normalization of hematocrit on exercise capacity and health-related quality of life. *American Journal of Kidney Diseases* 2002;**39**(2):257-65. [MEDLINE: 11840365]

Paluchamy 2018 (published data only)

Paluchamy T, Vaidyanathan R. Effectiveness of intradialytic exercise on dialysis adequacy, physiological parameters, biochemical markers and quality of life - a pilot study. *Saudi Journal of Kidney Diseases & Transplantation* 2018;**29**(4):902-10. [MEDLINE: 30152428]

Parsons 2004 (published data only)

Parsons TL, Toffelmire EB, King-VanVlack CE. The effect of an exercise program during hemodialysis on dialysis efficacy, blood pressure and quality of life in end-stage renal disease (ESRD) patients. *Clinical Nephrology* 2004;**61**(4):261-74. [MEDLINE: 15125032]

PEAK 2006 (published data only)

Cheema B, Abas H, Smith B, O'Sullivan A, Chan M, Patwardhan A, et al. Progressive exercise for anabolism in kidney disease (PEAK): a randomized, controlled trial of resistance training during hemodialysis. *Journal of the American Society of Nephrology* 2007;**18**(5):1594-601. [MEDLINE: 17409306]

Cheema B, Abas H, Smith B, O'Sullivan A, Chan M, Patwardhan A, et al. Randomized controlled trial of intradialytic resistance training to target muscle wasting in ESRD: the Progressive Exercise for Anabolism in Kidney Disease (PEAK) study. *American Journal of Kidney Diseases* 2007;**50**(4):574-84. [MEDLINE: 17900457]

Cheema B, Abas H, Smith B, O'Sullivan AJ, Chan M, Patwardhan A, et al. Investigation of skeletal muscle quantity and quality in end-stage renal disease. *Nephrology* 2010;**15**(4):454-63. [MEDLINE: 20609098]

Cheema B, Sullivan A, Chan M, Patwardhan A, Kelly J, Gillan A, et al. A randomized controlled trial of progressive resistance training during maintenance hemodialysis treatment: The PEAK Study [abstract]. *Journal of Aging & Physical Activity* 2004;**12**(3):260.

Cheema BS, Abas H, Smith BC, O'Sullivan AJ, Chan M, Patwardhan A, et al. Effect of resistance training during



hemodialysis on circulating cytokines: a randomized controlled trial. *European Journal of Applied Physiology* 2011;**111**(7):1437-45. [MEDLINE: 21161265]

Cheema BS, O´Sullivan AJ, Patwardhan NA, Kelly J, Gillin A, Fiatarone Singh MA. Progressive resistance training during hemodialysis: rationale and method of a randomized-controlled trial. *Hemodialysis International* 2006;**10**(3):303-10. [MEDLINE: 16805893]

* Smith BC, Cheema BS, O'Sullivan AJ, Pang G, Lloyd BD, Patwardhan A, et al. Resistance training during hemodialysis reduces C-reactive protein. Results from a randomized controlled trial of progressive exercise for anabolism in kidney disease (the PEAK Study) [abstract no: P34]. *Journal of the American Geriatrics Society* 2005;**53**(Suppl 1):13-4. [CENTRAL: CN-00757895]

Pellizzaro 2013 (published data only)

Pellizzaro CO, Thome FS, Veronese FV. Effect of peripheral and respiratory muscle training on the functional capacity of hemodialysis patients. *Renal Failure* 2013;**35**(2):189-97. [MEDLINE: 23199095]

Rahimimoghadam 2017 {published data only}

Rahimimoghadam Z, Rahemi Z, Mirbagher AN, Sadat Z. Effects of Pilates exercise on general health of hemodialysis patients. Journal of Bodywork & Movement Therapies 2017;**21**(1):86-92. [MEDLINE: 28167196]

Reboredo 2010 {published data only}

Reboredo MM, Neder JA, Pinheiro BV, Henrique DM, Faria RS, Paula RB. Constant work-rate test to assess the effects of intradialytic aerobic training in mildly impaired patients with end-stage renal disease: a randomized controlled trial. *Archives of Physical Medicine & Rehabilitation* 2011;**92**(12):2018-24. [MEDLINE: 22133251]

Reboredo MM, Neder JA, Pinheiro BV, Henrique DM, Lovisi JC, Paula RB. Intra-dialytic training accelerates oxygen uptake kinetics in hemodialysis patients. *European Journal of Preventive Cardiology* 2015;**22**(7):912-9. [MEDLINE: 25038079]

Reboredo MM, Pinheiro B, Neder JA, Avila MP, Araujo E Ribeiro ML, de Mendonca AF, et al. Effects of aerobic training during hemodialysis on heart rate variability and left ventricular function in end-stage renal disease patients. *Jornal Brasileiro de Nefrologia* 2010;**32**(4):367-73. [MEDLINE: 21541451]

Rezaei 2015 {published data only}

Rezaei J, Abdi A, Rezaei M, Heydarnezhadian J, Jalali R. Effect of regular exercise program on depression in hemodialysis patients. *International Scholarly Research Notices Online* 2015:182030. [MEDLINE: 27347502]

Rosa 2018 (published data only)

Rosa CS, Nishimoto DY, Souza GD, Ramirez AP, Carletti CO, Daibem CG, et al. Effect of continuous progressive resistance training during hemodialysis on body composition, physical function and quality of life in end-stage renal disease patients: a randomized controlled trial. *Clinical Rehabilitation* 2018;**32**(7):899-908. [MEDLINE: 29504416]

Rouchon 2016 (published data only)

Rouchon MI, Coutard C, Matysiak M, Ravel P, Forte C, Boisseau N. High intensity interval training improves physical functioning, inflammation and quality of life in peritoneal dialysis patients [abstract no: MO068]. *Nephrology Dialysis Transplantation* 2016;**31**(Suppl 1):i58. [EMBASE: 72326050]

Samara 2016 {published data only}

Samara A, Kouidi E, Fountoulakis K, Alexiou S, Deligiannis A. The effects of aquatic exercise on functional capacity and health-related quality of life in hemodialysis patients [abstract no: MP398]. *Nephrology Dialysis Transplantation* 2016;**31**(Suppl 1):i472. [EMBASE: 72327249]

Segura-Orti 2009 {published data only}

Segura-Orti E, Kouidi E, Lison JF. Effect of resistance exercise during hemodialysis on physical function and quality of life: randomized controlled trial. *Clinical Nephrology* 2009;**71**(5):527-37. [MEDLINE: 19473613]

Sheshadri 2020 (published data only)

Sheshadri A, Kittiskulnam P, Delgado C, Sudore R, Lai JC, Johansen KL. Association of motivations and barriers with participation and performance in a pedometer-based intervention. *Nephrology Dialysis Transplantation* 2020;**35**(8):1405-11. [MEDLINE: 32437568]

Sheshadri A, Kittiskulnam P, Johansen KL. Pedometers and exercise in dialysis [abstract no: SA-PO879]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):966. [EMBASE: 633734947]

Sheshadri A, Kittiskulnam P, Lai JC, Johansen KL. Effect of a pedometer-based intervention on body composition in ESRD [abstract no: SA-OR065]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):96. [EMBASE: 633771639]

Sheshadri A, Kittiskulnam P, Lai JC, Johansen KL. Effect of a pedometer-based walking intervention on body composition in patients with ESRD: a randomized controlled trial. *BMC Nephrology* 2020;**21**(1):100. [MEDLINE: 32178648]

Sheshadri A, Kittiskulnam P, Lazar AA, Johansen KL. A walking intervention to increase weekly steps in dialysis patients: a pilot randomized controlled trial. *American Journal of Kidney Diseases* 2020;**75**(4):488-96. [MEDLINE: 31679747]

Soliman 2015 {published data only}

Soliman HM. Effect of intradialytic exercise on fatigue, electrolytes level and blood pressure in hemodialysis patients: a randomized controlled trial. *Journal of Nursing Education & Practice* 2015;**5**(11):16-28.

Song 2012a {published data only}

Song WJ, Sohng KY. Effects of progressive resistance training on body composition, physical fitness and quality of life of patients on hemodialysis. *Journal of Korean Academy of Nursing* 2012;**42**(7):947-56. [MEDLINE: 23377590]

Suhardjono 2019 {published data only}

Suhardjono VU, Tedjasukmana D, Setiati S. The effect of intradialytic exercise twice a week on the physical capacity,



inflammation, and nutritional status of dialysis patients: a randomized controlled trial. *Hemodialysis International* 2019;**23**(4):486-93. [MEDLINE: 31100199]

Toussaint 2008 (published data only)

Toussaint ND, Polkinghorne KR, Kerr PG. Impact of intradialytic exercise on arterial compliance and B-type natriuretic peptide levels in hemodialysis patients. *Hemodialysis International* 2008;**12**(2):254-63. [MEDLINE: 18394060]

Tsuyuki 2003 (published data only)

Tsuyuki K, Kimura Y, Chiashi K, Matsushita C, Ninomiya K, Choh K, et al. Oxygen uptake efficiency slope as monitoring tool for physical training in chronic hemodialysis patients. *Therapeutic Apheresis & Dialysis* 2003;**7**(4):461-7. [MEDLINE: 12887732]

Uchiyama 2019 {published data only}

Uchiyama K, Wakino S, Washida N, Morimoto K, Kasai T, Itoh H. Home-based aerobic and resistance exercise training on peritoneal dialysis patients: a randomized controlled trial [abstract no: FR-PO701]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):600. [EMBASE: 633735606]

Uchiyama K, Washida N, Morimoto K, Muraoka K, Kasai T, Yamaki K, et al. Home-based aerobic exercise and resistance training in peritoneal dialysis patients: a randomized controlled trial. *Scientific Reports* 2019;**9**(1):2632. [MEDLINE: 30796338]

van Vilsteren 2005 {published data only}

van Vilsteren MC, de Greef MH, Huisman RM. The effects of a low-to-moderate intensity pre-conditioning exercise programme linked with exercise counselling for sedentary haemodialysis patients in the Netherlands: results of a randomized clinical trial. *Nephrology Dialysis Transplantation* 2005;**20**(1):141-6. [MEDLINE: 15522901]

Wilund 2010 (published data only)

Wilund KR, Tomayko EJ, Wu P, Vallurupalli S, Kumar J, Lakshminarayanan B, et al. Effects of intradialytic exercise training on inflammation, oxidative stress and epicardial fat [abstract no: SA-FC383]. *Journal of the American Society of Nephrology* 2009;**20**(Abstract Suppl):89A.

Wilund KR, Tomayko EJ, Wu PT, Ryong Chung H, Vallurupalli S, Lakshminarayanan B, et al. Intradialytic exercise training reduces oxidative stress and epicardial fat: a pilot study. *Nephrology Dialysis Transplantation* 2010;**25**(8):2695-701. [MEDLINE: 20190243]

Wu 2014d {published data only}

Wu Y, He Q, Yin X, He Q, Cao S, Ying G. Effect of individualized exercise during maintenance haemodialysis on exercise capacity and health-related quality of life in patients with uraemia. *Journal of International Medical Research* 2014;**42**(3):718-27. [MEDLINE: 24781720]

Yurtkuran 2007 {published data only}

Yurtkuran M, Alp A, Yurtkuran M, Dilek K. A modified yogabased exercise program in hemodialysis patients: a randomized controlled study. *Complementary Therapies in Medicine* 2007;**15**(3):164-71. [MEDLINE: 17709061]

Zhao 2017 (published data only)

Zhao C, Ma H, Yang L, Xiao Y. Long-term bicycle riding ameliorates the depression of the patients undergoing hemodialysis by affecting the levels of interleukin-6 and interleukin-18. *Neuropsychiatric Disease & Treatment* 2017;**13**:91-100. [MEDLINE: 28096677]

References to studies excluded from this review

Al-Ali 2018a {published data only}

Al-Ali FS, Zhou H, Hamad A, Ibrahim RA, Talal T, Najafi B. Exercise games to improve balance and mobility in diabetic patients undergoing hemodialysis: a randomized controlled trial [abstract no: TH-PO771]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):321. [EMBASE: 633736974]

Aliasgharpour 2016 {published data only}

Aliasgharpour M, Abbasi Z, Pedram RS, Kazemnezhad A. The effect of stretching exercises on severity of restless legs syndrome in patients on hemodialysis. *Asian Journal of Sports Medicine* 2016;**7**(2):e31001. [MEDLINE: 27625757]

Alvares 2017 (published data only)

Alvares VR, Ramos CD, Pereira BJ, Pinto AL, Moyses RM, Gualano B, et al. Pneumatic compression, but not exercise, can avoid intradialytic hypotension: a randomized trial. *American Journal of Nephrology* 2017;**45**(5):409-16. [MEDLINE: 28407637]

Bogataj 2020 {published data only}

Bogataj S, Pajek J, Buturovic Ponikvar J, Hadzic V, Pajek M. Kinesiologist-guided functional exercise in addition to intradialytic cycling program in end-stage kidney disease patients: a randomised controlled trial [Erratum in: Sci Rep. 2020 Jun 23;10(1):10399; PMID: 32576882]. *Scientific Reports* 2020;**10**(1):5717. [MEDLINE: 32235852]

Bogataj S, Pajek J, Buturovic Ponikvar J, Pajek M. Functional training added to intradialytic cycling lowers low-density lipoprotein cholesterol and improves dialysis adequacy: a randomized controlled trial. *BMC Nephrology* 2020;**21**(1):352. [MEDLINE: 32811448]

Bogataj S, Pajek J, Ponikvar JB, Hadzic V, Pajek M. Author correction: Kinesiologist-guided functional exercise in addition to intradialytic cycling program in end-stage kidney disease patients: a randomised controlled trial. *Scientific Reports* 2020;**10**(1):10399. [MEDLINE: 32576882]

Bogataj S, Pajek M, Buturovic Ponikvar J, Pajek J. Outcome expectations for exercise and decisional balance questionnaires predict adherence and efficacy of exercise programs in dialysis patients. *International Journal of Environmental Research & Public Health* 2020;**17**(9):3175. [MEDLINE: 32370202]

Bohm 2014 {published data only}

Bohm C, Onyskie-Marcus J, Stewart K. A randomized controlled trial of aerobic exercise in patients with end stage renal disease (ESRD) during hemodialysis (HD) [abstract no: F-PO1723]. *Journal of the American Society of Nephrology* 2008;**19**(Abstracts Issue):495A.



Bohm C, Stewart K, Onyskie-Marcus J, Esliger D, Kriellaars D, Rigatto C. Effects of intradialytic cycling compared with pedometry on physical function in chronic outpatient hemodialysis: a prospective randomized trial. *Nephrology Dialysis Transplantation* 2014;**29**(10):1947-55. [MEDLINE: 25061127]

Bohm 2017 (published data only)

Bohm J, Monteiro MB, Andrade FP, Veronese F, Thome FS. Acute effects of intradialytic aerobic exercise on solute removal, blood gases and oxidative stress in patients with chronic kidney disease. *Jornal Brasileiro de Nefrologia* 2017;**39**(2):172-80. [MEDLINE: 28489182]

Brown 2018 {published data only}

Brown PD, Rowed K, Shearer J, Macrae JM, Parker K. Impact of intradialytic exercise intensity on urea clearance in hemodialysis patients. *Applied Physiology, Nutrition, & Metabolism [Physiologie Appliquee, Nutrition et Metabolisme]* 2018;**43**(1):101-4. [MEDLINE: 28961405]

Campos 2018 (published data only)

Campos NG, Marizeiro DF, Florencio AC, Silva IC, Meneses GC, Bezerra GF, et al. Effects of respiratory muscle training on endothelium and oxidative stress biomarkers in hemodialysis patients: a randomized clinical trial. *Respiratory Medicine* 2018;**134**:103-9. [MEDLINE: 29413495]

Castellino 1987 {published data only}

Castellino P, Bia M, DeFronzo RA. Metabolic response to exercise in dialysis patients. *Kidney International* 1987;**32**(6):877-83. [MEDLINE: 3323603]

Castellino P, Bia MJ, DeFronzo RA. Adrenergic modulation of potassium metabolism in uremia. *Kidney International* 1990;**37**(2):793-8. [MEDLINE: 2308263]

Chagolla 2018 (published data only)

Chagolla JM, Sanchez Cardenas M, Perez-Grovas HA. Exercise during dialysis ameliorates the decline in plasma volume by ultrafiltration [abstract no: SA-PO881]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):966. [EMBASE: 633735033]

CTRI/2018/02/012021 {published data only}

Raj PD, Bai RS. The effectiveness of intradialytic exercise on fatigue and quality of sleep among patients undergoing hemodialysis. http://ctri.nic.in/Clinicaltrials/showallp.php?mid1=21976&EncHid=&userName=intradialytic%20exercise%20on%20fatigue%20and%20quality%20of%20sleep (first received 21 February 2018).

De Villar 2016 {published data only}

De Villar LO, Dominguez BP, Gramage JM, Perez MJ, Garcia SA, Sala AB, et al. Comparison of intradialytic versus home-based exercise programmes on physical function, physical level and health related quality of life [abstract no: MP414]. *Nephrology Dialysis Transplantation* 2016;**31**(Suppl 1):i477-8. [EMBASE: 72327265]

Dias 2020 (published data only)

Dias EC, Orcy R, Antunes MF, Kohn R, Rombaldi AJ, Ribeiro L, et al. Intradialytic exercise with blood flow restriction: something to add to hemodialysis adequacy? Findings from a crossover study. *Hemodialysis International* 2020;**24**(1):71-8. [MEDLINE: 31612630]

Dungey 2013 {published data only}

Dungey M, Bishop NC, Young HMI, Burton JO, Smith AC. Effects of acute intradialytic exercise on blood pressure and circulating cytokines [abstract no: FR-PO400]. *Journal of the American Society of Nephrology* 2013;**24**(Abstract Suppl):457A.

Dungey 2015 {published data only}

Dungey M, Bishop NC, Young HM, Burton JO, Smith AC. The impact of exercising during haemodialysis on blood pressure, markers of cardiac injury and systemic inflammation-preliminary results of a pilot study. *Kidney & Blood Pressure Research* 2015;**40**(6):593-604. [MEDLINE: 26619202]

Martin N, Smith AC, Dungey MR, Young HM, Burton JO, Bishop NC. Exercise during hemodialysis does not affect the phenotype or prothrombotic nature of microparticles but alters their proinflammatory function. *Physiological Reports* 2018;**6**(19):e13825. [MEDLINE: 30294974]

Dziubek 2016 (published data only)

Dziubek W, Kowalska J, Kusztal M, Rogowski L, Golebiowski T, Nikifur M, et al. The level of anxiety and depression in dialysis patients undertaking regular physical exercise training--a preliminary study. *Kidney & Blood Pressure Research* 2016;**41**(1):86-98. [MEDLINE: 26872253]

Fontsere 2016 (published data only) 15802482

Fontsere Baldellou N, Mestres Alomar G, Yugueros Castellnou X, Lopez Alonso T, Yuguero Ortiz A, Riambau Alonso V. The effect of a postoperative exercise program on arteriovenous fistula maturity: a randomized controlled trial [abstract no: 38]. *Journal of Vascular Access* 2015;**15**(2):e27. [CENTRAL: CN-01657769]

Fontsere N, Mestres G, Yugueros X, Lopez T, Yuguero A, Bermudez P, et al. Effect of a postoperative exercise program on arteriovenous fistula maturation: a randomized controlled trial. *Hemodialysis International* 2016;**20**(2):306-14. [MEDLINE: 26486682]

Frih 2017 {published data only}

Frih B, Mkacher W, Bouzguenda A, Jaafar H, ALkandari SA, Ben SZ, et al. Effects of listening to Holy Qur'an recitation and physical training on dialysis efficacy, functional capacity, and psychosocial outcomes in elderly patients undergoing haemodialysis. *Libyan Journal of Medicine* 2017;**12**(1):1372032. [MEDLINE: 28891419]

Frih 2018 (published data only)

Frih B, Mkacher W, Jaafar H, Frih A, Ben Salah Z, El May M, et al. Specific balance training included in an endurance-resistance exercise program improves postural balance in elderly patients undergoing haemodialysis. *Disability & Rehabilitation* 2018;**40**(7):784-90. [MEDLINE: 28084833]



Fuhro 2018 (published data only)

Fuhro MI, Dorneles GP, Andrade FP, Romao PR, Peres A, Monteiro MB. Acute exercise during hemodialysis prevents the decrease in natural killer cells in patients with chronic kidney disease: a pilot study. *International Urology & Nephrology* 2018;**50**(3):527-34. [MEDLINE: 29134614]

Garcia Testal 2019 {published data only}

Garcia Testal A, Garcia Maset R, Hervas Marin D, Perez-Dominguez B, Royo M, Rico Salvador IS, et al. Influence of physical exercise on the dialytic adequacy parameters of patients on hemodialysis. *Therapeutic Apheresis & Dialysis* 2019;**23**(2):160-6. [MEDLINE: 30226299]

Giannaki 2015 (published data only) NCT00942253

Giannaki CD, Sakkas GK, Karatzaferi C, Hadjigeorgiou GM, Lavdas E, Kyriakides T, et al. Effect of exercise training and dopamine agonists in patients with uremic restless legs syndrome: a six-month randomized, partially double-blind, placebo-controlled comparative study. *BMC Nephrology* 2013;**14**:194. [MEDLINE: 24024727]

Giannaki CD, Sakkas GK, Karatzaferi C, Maridaki MD, Koutedakis Y, Hadjigeorgiou GM, et al. Combination of exercise training and dopamine agonists in patients with RLS on dialysis: a randomized, double-blind placebo-controlled study. *ASAIO Journal* 2015;**61**(6):738-41. [MEDLINE: 26262586]

Sakkas G, Giannaki C, Karatzaferi C, Hadjigeorgiou GM, Stefanidis I. Exercise training and dopamine agonists in hemodialysis patients with restless legs syndrome. A randomized double-blind placebo controlled study [abstract no: SP438]. *Nephrology Dialysis Transplantation* 2015;**30**(Suppl 3):iii523-4. [EMBASE: 72207765]

Hamad 2016 (published data only)

Hamad A, Ali-Ali FS, Zhou H, Talal T, Melo SNS, Elesnawi MA, et al. Therapeutic game-based exercise during hemodialysis to improve balance; a pilot randomized controlled trial [abstract no: TH-OR106]. *Journal of the American Society of Nephrology* 2016;**27**(Abstract Suppl):26A. [CENTRAL: CN-01657757]

Jeong 2018 (published data only)

Jeong JH, Biruete A, Fernhall B, Wilund KR. Effects of acute intradialytic exercise on cardiovascular responses in hemodialysis patients. *Hemodialysis International* 2018;**22**(4):524-33. [MEDLINE: 29745006]

Kirkman 2013 {published data only}

Kirkman D, Jibani M, Macdonald JH. Dialysis adequacy and solute removal: effect of intradialytic exercise [abstract no: FR-PO345]. *Journal of the American Society of Nephrology* 2012;**23**(Abstract Suppl):451A.

Kirkman DL, Roberts LD, Kelm M, Wagner J, Jibani MM, Macdonald JH. Interaction between intradialytic exercise and hemodialysis adequacy. *American Journal of Nephrology* 2013;**38**(6):475-82. [MEDLINE: 24296748]

Krase 2020 (published data only)

Krase AA, Flouris AD, Karatzaferi C, Giannaki CD, Stefanidis I, Sakkas GK. Separate and combined effects of cold dialysis and

intradialytic exercise on the thermoregulatory responses of hemodialysis patients: a randomized-cross-over study. *BMC Nephrology* 2020;**21**(1):524. [MEDLINE: 33267815]

Maheshwari 2012 (published data only)

Maheshwari V, Samavedham L, Rangaiah GP, Loy Y, Ling LH, Sethi S, et al. Comparison of toxin removal outcomes in online hemodiafiltration and intra-dialytic exercise in high-flux hemodialysis: A prospective randomized open-label clinical study protocol. *BMC Nephrology* 2012;**13**:156. [MEDLINE: 23176731]

Majchrzak 2008 (published data only)

Majchrzak KM, Pupim LB, Flakoll PJ, Ikizler TA. Resistance exercise augments the acute anabolic effects of intradialytic oral nutritional supplementation. *Nephrology Dialysis Transplantation* 2008;**23**(4):1362-9. [MEDLINE: 18065829]

Miura 2016 (published data only)

Miura M, Yoshizawa R, Hirayama A, Ito O, Kohzuki M, Owada S, et al. Comparing the effect of electric bicycle training and conventional exercise on physical function of end stage renal disease patients undergoing hemodialysis [abstract no: TH-PO1049]. *Journal of the American Society of Nephrology* 2016;**27**(Abstract Suppl):340A. [CENTRAL: CN-01657761]

Molsted 2013 (published data only) 72099857

Molsted S, Harrison AP, Eidemak I, Andersen JL. The effects of high-load strength training with protein- or nonprotein-containing nutritional supplementation in patients undergoing dialysis. *Journal of Renal Nutrition* 2013;**23**(2):132-40. [MEDLINE: 22959782]

Mora 2007 {published data only}

Mora FG, Mariscal L, Medel L, Cadena M, Magana F, Franco M, et al. Metabolism measurements in hemodiafiltration patients with exercise [abstract no: S-PO-0126]. In: 4th World Congress of Nephrology.19th International Congress of the International Society of Nephrology (ISN); 2007 Apr 21-25; Rio de Janeiro, Brazil. 2007:82. [CENTRAL: CN-01658052]

Moug 2004 (published data only)

Moug SJ, Grant S, Creed G, Boulton Jones M. Exercise during haemodialysis: West of Scotland pilot study. *Scottish Medical Journal* 2004;**49**(1):14-7. [MEDLINE: 15012046]

Orcy 2012 {published data only}

Orcy RB, Dias PS, Seus TL, Barcellos FC, Bohlke M. Combined resistance and aerobic exercise is better than resistance training alone to improve functional performance of haemodialysis patients - results of a randomized controlled trial. *Physiotherapy Research International* 2012;**17**(4):235-43. [MEDLINE: 22693148]

Orcy 2014 {published data only}

Orcy R, Antunes MF, Schiller T, Seus T, Bohlke M. Aerobic exercise increases phosphate removal during hemodialysis: a controlled trial. *Hemodialysis International* 2014;**18**(2):450-8. [MEDLINE: 24438516]



Pinto 2015 (published data only)

Pinto JS, Sarmento LA, Pereira da Silva AP, Cabral CM, Chiavegato LD. Effectiveness of conventional physical therapy and Pilates' method in functionality, respiratory muscle strength and ability to exercise in hospitalized chronic renal patients: a study protocol of a randomized controlled trial. *Journal of Bodywork & Movement Therapies* 2015;**19**(4):604-15. [MEDLINE: 26592217]

Sarmento LA, Pinto JS, da Silva AP, Cabral CM, Chiavegato LD. Effect of conventional physical therapy and Pilates in functionality, respiratory muscle strength and ability to exercise in hospitalized chronic renal patients: a randomized controlled trial. *Clinical Rehabilitation* 2017;**31**(4):508-20. [MEDLINE: 27178843]

Ribeiro 2019 (published data only)

Ribeiro HS, Cunha VA, Baiao VM, Duarte MP, Carvalho HL, Franca GD, et al. Intradialytic isometric handgrip training seems to be safe: a pilot study on hemodialysis patients [abstract no: TH-PO257]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):181. [EMBASE: 633769954]

Rossum 2019 (published data only)

Rossum KF, Thompson SE, Hancock EK, Riehl-Tonn V, Brar RS, Leon Mantilla SJ, et al. Timing of intradialytic exercise and its impact on intradialytic hypotension: a randomized crossover study [abstract no: SA-OR063]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):96. [EMBASE: 633771517]

Stray-Gundersen 2016 {published data only}

Stray-Gundersen J, Howden EJ, Parsons DB, Thompson JR. Neither hematocrit normalization nor exercise training restores oxygen consumption to normal levels in hemodialysis patients. *Journal of the American Society of Nephrology* 2016;**27**(12):3769-79. [MEDLINE: 27153927]

Sun 2002 (published data only)

Sun Y, Chen B, Jia Q, Wang J. The effect of exercise during hemodialysis on adequacy of dialysis. *Chung-Hua Nei Ko Tsa Chih [Chinese Journal of Internal Medicine]* 2002;**41**(2):79-81. [MEDLINE: 11940299]

Sun YB, Chen BL, Jia Q, Wang JM. Exercise therapy during hemodialysis to improve adequacy of dialysis randomized controlled trial. *Chinese Journal of Clinical Rehabilitation* 2003;**7**(27):3702-3. [EMBASE: 38636520]

Tao 2015 {published data only}

Tao X, Chow SK, Wong FK. A nurse-led case management program on home exercise training for hemodialysis patients: a randomized controlled trial. *International Journal of Nursing Studies* 2015;**52**(6):1029-41. [MEDLINE: 25840898]

Tao X, Chow SK, Wong FK. The effects of a nurse-supervised home exercise programme on improving patients' perceptions of the benefits and barriers to exercise: a randomised controlled trial. *Journal of Clinical Nursing* 2017;**26**(17-18):2765-75. [MEDLINE: 28278361]

Vrakas 2017 (published data only)

Vrakas S, Mameletzi D, Samaras T, Liakopoulos V, Kouidi E, Deligiannis A. The effects of intradialytic exercise plus music on anxiety [abstract no: SP567]. *Nephrology Dialysis Transplantation* 2017;**32**(Suppl 3):iii325. [EMBASE: 617290243]

References to studies awaiting assessment

Assawasaksakul 2018 (published data only)

Assawasaksakul N, Tiranathanagul K, Sirichana W, Kulaputana O, Eiam-Ong S, Praditpornsilpa K. The effects of intradialytic exercise on the improvement of daily physical activity in online hemodiafiltration patients [abstract no: SA-PO878]. *Journal of the American Society of Nephrology* 2018;**29**(Abstract Suppl):965. [EMBASE: 633734899]

Bennett 2019 (published data only)

Bennett PN, West MG, Hussein WF, Smith E, Reiterman M, Patel J, et al. The effect of a combined resistance and cardiovascular exercise program on peritoneal dialysis patients: a pilot randomized control trial [abstract no: TH-OR091]. *Journal of the American Society of Nephrology* 2019;**30**(Abstract Suppl):24. [EMBASE: 633771324]

Dong 2019 (published data only)

Dong ZJ, Zhang HL, Yin LX. Effects of intradialytic resistance exercise on systemic inflammation in maintenance hemodialysis patients with sarcopenia: a randomized controlled trial. *International Urology & Nephrology* 2019;**51**(8):1415-24. [MEDLINE: 31270740]

IMPCT 2020 {published data only}

McAdams-DeMarco MA, Chu NM, Steckel M, Kunwar S, Gonzalez Fernandez M, Carlson MC, et al. Interventions Made to Preserve Cognitive Function Trial (IMPCT) study protocol: a multi-dialysis center 2x2 factorial randomized controlled trial of intradialytic cognitive and exercise training to preserve cognitive function. *BMC Nephrology* 2020;**21**(1):383. [MEDLINE: 32883245]

Lopes 2019 {published data only}

Lopes LC, Mota JF, Prestes J, Schincaglia RM, Silva DM, Queiroz NP, et al. Intradialytic resistance training improves functional capacity and lean mass gain in individuals on hemodialysis: a randomized pilot trial. *Archives of Physical Medicine & Rehabilitation* 2019;**100**(11):2151-8. [MEDLINE: 31278924]

Maynard 2019 {published data only}

Maynard LG, de Menezes DL, Liao NS, de Jesus EM, Andrade NL, Santos JC, et al. Effects of exercise training combined with virtual reality in functionality and health-related quality of life of patients on hemodialysis. *Games for Health Journal* 2019;8(5):339-48. [MEDLINE: 31539293]

PEDAL 2021 {published data only}**83508514**

Greenwood SA, Koufaki P, Macdonald J, Bhandari S, Burton J, Dasgupta I, et al. The PrEscription of intraDialytic exercise to improve quAlity of Life in patients with chronic kidney disease trial: study design and baseline data for a



multicentre randomized controlled trial. *Clinical Kidney Journal* 2021;**14**(5):1345-55. [MEDLINE: 33959264]

Stringuetta Belik 2018 (published data only)

Oliviera E Silva VR, Stringuetta Belik F, Hueb JC, de Souza Goncalves R, Costa Teixeira Caramori J, Perez Vogt B, et al. Aerobic exercise training and nontraditional cardiovascular risk factors in hemodialysis patients: results from a prospective randomized trial. *Cardiorenal Medicine* 2019;**9**(6):391-9. [MEDLINE: 31597151]

Stringuetta Belik F, Oliveira E Silva VR, Braga GP, Bazan R, Perez Vogt B, Costa Teixeira Caramori J, et al. Influence of intradialytic aerobic training in cerebral blood flow and cognitive function in patients with chronic kidney disease: a pilot randomized controlled trial. *Nephron* 2018;**140**(1):9-17. [MEDLINE: 29879707]

References to ongoing studies

ACTRN12618000724279 {published data only}

Chojak-Fijalka K. Evaluation of the effectiveness of home-based physical training in patients undergoing haemodialysis [The effects of home-based physical training in patients undergoing haemodialysis on physical functioning, body composition, quality of life, fatigue and selected properties of blood]. www.anzctr.org.au/Trial/Registration/TrialReview.aspx? id=374398&isReview=true (first received 3 March 2018).

Cardoso 2019 (published data only)

Cardoso RK, Araujo AM, Del Vechio FB, Bohlke M, Barcellos FC, Oses JP, et al. Intradialytic exercise with blood flow restriction is more effective than conventional exercise in improving walking endurance in hemodialysis patients: a randomized controlled trial. *Clinical Rehabilitation* 2020;**34**(1):91-8. [MEDLINE: 31603002]

Cardoso RK, Araujo AM, Orcy RB, Bohlke M, Oses JP, Del Vecchio FB, et al. Effects of continuous moderate exercise with partial blood flow restriction during hemodialysis: a protocol for a randomized clinical trial. *MethodsX* 2019;**6**:190-8. [MEDLINE: 30740314]

Chan 2019 {published data only}

Chan KN, Chen Y, Lit Y, Massaband P, Kiratli J, Rabkin R, et al. A randomized controlled trial of exercise to prevent muscle mass and functional loss in elderly hemodialysis patients: Rationale, study design, and baseline sample. *Contemporary Clinical Trials Communications* 2019;**15**:100365. [MEDLINE: 31193611]

Clarkson 2017 {published data only}

Clarkson MJ, Fraser SF, Bennett PN, McMahon LP, Brumby C, Warmington SA. Efficacy of blood flow restriction exercise during dialysis for end stage kidney disease patients: protocol of a randomised controlled trial. *BMC Nephrology* 2017;**18**(1):294. [MEDLINE: 28893206]

NCT01721551 {published data only}

Sakkas GK. Sleep and training aspects in dialysis fatigue - exercise intervention (StandFirm). www.clinicaltrials.gov/ct2/show/NCT01721551 (first received 5 November 2012).

Additional references

ANZDATA 2019

Australia and New Zealand Dialysis and Transplant Registry. ANZDATA 42nd Report, Chapter 3: Mortality in end stage kidney disease. 2019. www.anzdata.org.au/wp-content/uploads/2019/09/ c03_mortality_2018_ar_2019_v1.0_20191202.pdf (accessed 24 June 2021).

Barcellos 2015

Barcellos FC, Santos IS, Umpierre D, Bohlke M, Hallal PC. Effects of exercise in the whole spectrum of chronic kidney disease: a systematic review. *Clinical Kidney Journal* 2015;**8**(6):753-65. [MEDLINE: 26613036]

Bessa 2015

Bessa B, de Oliveira Leal V, Moraes C, Barboza J, Fouque D, Mafra D. Resistance training in hemodialysis patients: a review. *Rehabilitation Nursing Journal* 2015;**40**(2):111-26. [MEDLINE: 24729123]

Bohannon 2017

Bohannon RW, Crouch R. Minimal clinically important difference for change in 6-minute walk test distance of adults with pathology: a systematic review. *Journal of Evaluation in Clinical Practice* 2017;**23**(2):377-81. [MEDLINE: 27592691]

Chan 2016

Chan D, Cheema BS. Progressive resistance training in endstage renal disease: systematic review. *American Journal of Nephrology* 2016;**44**(1):32-45. [MEDLINE: 27355619]

Chang 2001

Chang WK, Hung KY, Huang JW, Wu KD, Tsai TJ. Chronic fatigue in long-term peritoneal dialysis patients. *American Journal of Nephrology* 2001;**21**(6):479-85. [MEDLINE: 11799265]

Chung 2017

Chung YC, Yeh ML, Liu YM. Effects of intradialytic exercise on the physical function, depression and quality of life for haemodialysis patients: a systematic review and meta-analysis of randomised controlled trials. *Journal of Clinical Nursing* 2017;**26**(13-14):1801-13. [MEDLINE: 27532211]

Clarkson 2019

Clarkson MJ, Bennett PN, Fraser SF, Warmington SA. Exercise interventions for improving objective physical function in patients with end-stage kidney disease on dialysis: a systematic review and meta-analysis. *American Journal of Physiology - Renal Physiology* 2019;**316**(5):F856-72. [MEDLINE: 30759022]

Cooney 2013

Cooney GM, Dwan K, Greig CA, Lawlor DA, Rimer J, Waugh FR, et al. Exercise for depression. *Cochrane Database of Systematic Reviews* 2013, Issue 9. Art. No: CD004366. [DOI: 10.1002/14651858.CD004366.pub6]



Cramp 2012

Cramp F, Byron-Daniel J. Exercise for the management of cancer-related fatigue in adults. *Cochrane Database of Systematic Reviews* 2012, Issue 11. Art. No: CD006145. [DOI: 10.1002/14651858.CD006145.pub3]

Cupisti 2017

Cupisti A, D'Alessandro C, Finato V, Del Corso C, Catania B, Caselli GM, et al. Assessment of physical activity, capacity and nutritional status in elderly peritoneal dialysis patients. *BMC Nephrology* 2017;**18**(1):180. [MEDLINE: 28558794]

DeOreo 1997

DeOreo PB. Hemodialysis patient-assessed functional health status predicts continued survival, hospitalization, and dialysis-attendance compliance. *American Journal of Kidney Diseases* 1997;**30**(2):204-12. [MEDLINE: 9261030]

DerSimonian 1986

DerSimonian R, Laird N. Meta-analysis in clinical trials. Controlled Clinical Trials 1986;**7**(3):177-88. [MEDLINE: 3802833]

ERA-EDTA 2017

ERA-EDTA Registry. ERA-EDTA Registry Annual Report 2017. Amsterdam UMC, location AMC, Department of Medical Informatics, Amsterdam, the Netherlands, 2019. www.era-edta-reg.org/files/annualreports/AnnRep2017.pdf (accessed 24 June 2021).

Erez 2016

Erez G, Selman L, Murtagh FE. Measuring health-related quality of life in patients with conservatively managed stage 5 chronic kidney disease: limitations of the Medical Outcomes Study Short Form 36: SF-36. *Quality of Life Research* 2016;**25**(11):2799-809. [MEDLINE: 27522214]

Eriksson 2016

Eriksson D, Karlsson L, Eklund O, Dieperink H, Honkanen E, Melin J, et al. Health-related quality of life across all stages of autosomal dominant polycystic kidney disease. *Nephrology Dialysis Transplantation* 2016;**32**(12):2106-11. [MEDLINE: 27662885]

Fahal 2014

Fahal IH. Uraemic sarcopenia: aetiology and implications. *Nephrology Dialysis Transplantation* 2014;**29**(9):1655-65. [MEDLINE: 23625972]

Farivar 2004

Farivar SS, Liu H, Hays RD. Half standard deviation estimate of the minimally important difference in HRQOL scores? Expert Review of Pharmacoeconomics & Outcomes Research 2004;4(5):515-23. [MEDLINE: 19807545]

Ferrari 2020

Ferrari F, Helal L, Dipp T, Soares D, Soldatelli A, Mills AL, et al. Intradialytic training in patients with end-stage renal disease: a systematic review and meta-analysis of randomized clinical trials assessing the effects of five different training interventions. *Journal of Nephrology* 2020;**33**(2):251-66. [EMBASE: 2003942904]

Finkelstein 2018

Finkelstein FO, van Nooten F, Wiklund I, Trundell D, Cella D. Measurement properties of the Short Form-36 (SF-36) and the Functional Assessment of Cancer Therapy - Anemia (FACT-An) in patients with anemia associated with chronic kidney disease. *Health & Quality of Life Outcomes* 2018;**16**(1):111. [MEDLINE: 29855366]

Gomes 2018

Gomes Neto M, de Lacerda FF, Lopes AA, Martinez BP, Saquetto MB. Intradialytic exercise training modalities on physical functioning and health-related quality of life in patients undergoing maintenance hemodialysis: systematic review and meta-analysis. *Clinical Rehabilitation* 2018;**32**(9):1189-202. [MEDLINE: 29480025]

GRADE 2008

Guyatt GH, Oxman AD, Vist GE, Kunz R, Falck-Ytter Y, Alonso-Coello P, et al. GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. *BMJ* 2008;**336**(7650):924-6. [MEDLINE: 18436948]

GRADE 2011

Guyatt G, Oxman AD, Akl EA, Kunz R, Vist G, Brozek J, et al. GRADE guidelines: 1. Introduction-GRADE evidence profiles and summary of findings tables. *Journal of Clinical Epidemiology* 2011;**64**(4):383-94. [MEDLINE: 21195583]

Heiwe 2014

Heiwe S, Jacobson SH. Exercise training in adults with CKD: a systematic review and meta-analysis. *American Journal of Kidney Diseases* 2014;**64**(3):383-93. [MEDLINE: 24913219]

Higgins 2003

Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analysis. *BMJ* 2003;**327**(7414):557-60. [MEDLINE: 12958120]

Higgins 2011

Higgins JP, Green S (editors). Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011]. The Cochrane Collaboration. 2011. Available from www.cochrane-handbook.org.

Huang 2019

Huang M, Lv A, Wang J, Xu N, Ma G, Zhai Z, et al. Exercise training and outcomes in hemodialysis patients: systematic review and meta-analysis. *American Journal of Nephrology* 2019;**50**(4):240-54. [MEDLINE: 31454822]

Jacobson 2019

Jacobson J, Ju A, Baumgart A, Unruh M, O'Donoghue D, Obrador G, et al. Patient perspectives on the meaning and impact of fatigue in hemodialysis: a systematic review and thematic analysis of qualitative studies. *American Journal of Kidney Diseases* 2019;**74**(2):179-92. [MEDLINE: 30955947]

Jhamb 2008

Jhamb M, Weisbord SD, Steel JL, Unruh M. Fatigue in patients receiving maintenance dialysis: a review of definitions,



measures, and contributing factors. *American Journal of Kidney Diseases* 2008;**52**(2):353-65. [MEDLINE: 18572290]

Johansen 2010

Johansen KL, Chertow GM, Kutner NG, Dalrymple LS, Grimes BA, Kaysen GA. Low level of self-reported physical activity in ambulatory patients new to dialysis. *Kidney International* 2010;**78**(11):1164-70. [MEDLINE: 20811334]

Johansen 2013

Johansen KL, Kaysen GA, Dalrymple LS, Grimes BA, Glidden DV, Anand S, et al. Association of physical activity with survival among ambulatory patients on dialysis: the Comprehensive Dialysis Study. *Clinical Journal of The American Society of Nephrology: CJASN* 2013;**8**(2):248-53. [MEDLINE: 23124787]

Kaysen 2011

Kaysen GA, Larive B, Painter P, Craig A, Lindsay RM, Rocco MV, et al. Baseline physical performance, health, and functioning of participants in the Frequent Hemodialysis Network (FHN) trial. *American Journal of Kidney Diseases* 2011;**57**(1):101-12. [MEDLINE: 21184919]

KDOQI 2005

K/DOQI Workgroup. K/DOQI clinical practice guidelines for cardiovascular disease in dialysis patients. *American Journal of Kidney Diseases* 2005;**45**(4 Suppl 3):S1-153. [MEDLINE: 15806502]

Knight 2003

Knight EL, Ofsthun N, Teng M, Lazarus JM, Curhan GC. The association between mental health, physical function, and hemodialysis mortality. *Kidney International* 2003;**63**(5):1843-51. [MEDLINE: 12675862]

Larun 2019

Larun L, Brurberg KG, Odgaard-Jensen J, Price JR. Exercise therapy for chronic fatigue syndrome. *Cochrane Database of Systematic Reviews* 2019, Issue 10. Art. No: CD003200. [DOI: 10.1002/14651858.CD003200.pub8]

Leaf 2009

Leaf DE, Goldfarb DS. Interpretation and review of health-related quality of life data in CKD patients receiving treatment for anemia. *Kidney International* 2009;**75**(1):15-24. [MEDLINE: 18813284]

Manns 2014

Manns B, Hemmelgarn B, Lillie E, Dip SC, Cyr A, Gladish M, et al. Setting research priorities for patients on or nearing dialysis. *Clinical Journal of The American Society of Nephrology: CJASN* 2014;**9**(10):1813-21. [MEDLINE: 24832095]

Norman 2003

Norman GR, Sloan JA, Wyrwich KW. Interpretation of changes in health-related quality of life: the remarkable universality of half a standard deviation. *Medical Care* 2003;**41**(5):582-92. [MEDLINE: 12719681]

Painter 2005

Painter P. Physical functioning in end-stage renal disease patients: update 2005. *Hemodialysis International* 2005;**9**(3):218-35. [MEDLINE: 16191072]

Painter 2017

Painter PL, Agarwal A, Drummond M. Physical function and physical activity in peritoneal dialysis patients. *Peritoneal Dialysis International* 2017;**37**(6):598-604. [MEDLINE: 28970364]

Palmer 2013

Palmer S, Vecchio M, Craig JC, Tonelli M, Johnson DW, Nicolucci A, et al. Prevalence of depression in chronic kidney disease: systematic review and meta-analysis of observational studies. *Kidney International* 2013;**84**(1):179-91. [MEDLINE: 23486521]

Pu 2019

Pu J, Jiang Z, Wu W, Li L, Zhang L, Li Y, et al. Efficacy and safety of intradialytic exercise in haemodialysis patients: a systematic review and meta-analysis. *BMJ Open* 2019;**9**(1):e020633. [MEDLINE: 30670499]

Qiu 2017

Qiu Z, Zheng K, Zhang H, Feng J, Wang L, Zhou H. Physical exercise and patients with chronic renal failure: a meta-analysis. *BioMed Research International* 2017;**2017**:7191826. [MEDLINE: 28316986]

Salhab 2019

Salhab N, Karavetian M, Kooman J, Fiaccadori E, El Khoury CF. Effects of intradialytic aerobic exercise on hemodialysis patients: a systematic review and meta-analysis. *Journal of Nephrology* 2019;**32**(4):549-66. [MEDLINE: 30659520]

Samsa 1999

Samsa G, Edelman D, Rothman ML, Williams GR, Lipscomb J, Matchar D. Determining clinically important differences in health status measures: a general approach with illustration to the Health Utilities Index Mark II. *Pharmacoeconomics* 1999;**15**(2):141-55. [MEDLINE: 10351188]

Scapini 2019

Scapini KB, Bohlke M, Moraes OA, Rodrigues CG, Inacio JF, Sbruzzi G, et al. Combined training is the most effective training modality to improve aerobic capacity and blood pressure control in people requiring haemodialysis for end-stage renal disease: systematic review and network meta-analysis. *Journal of Physiotherapy* 2019;**65**(1):4-15. [MEDLINE: 30581137]

Schunemann 2011a

Schünemann HJ, Oxman AD, Higgins JP, Vist GE, Glasziou P, Guyatt GH. Chapter 11: Presenting results and 'Summary of findings' tables. In: Higgins JP, Green S (editors). Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011]. The Cochrane Collaboration, 2011. Available from www.cochrane-handbook.org.

Schunemann 2011b

Schünemann HJ, Oxman AD, Higgins JP, Deeks JJ, Glasziou P, Guyatt GH. Chapter 12: Interpreting results and drawing



conclusions. In: Higgins JP, Green S (editors). Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [updated March 2011]. The Cochrane Collaboration, 2011. Available from www.cochrane-handbook.org.

Serratrice 1967

Serratrice G, Toga M, Roux H, Murisasco A, de Bisschop G. Neuropathies, myopathies and neuromyopathies in chronic uremic patients [Neuropathies, myopathies et neuromyopathies chez des uremiques chroniques]. *Presse Medicale* 1967;**75**(37):1835-8. [MEDLINE: 4293231]

Sheng 2014

Sheng K, Zhang P, Chen L, Cheng J, Wu C, Chen J. Intradialytic exercise in hemodialysis patients: a systematic review and meta-analysis. *American Journal of Nephrology* 2014;**40**(5):478-90. [MEDLINE: 25504020]

Sietsema 2004

Sietsema KE, Amato A, Adler SG, Brass EP. Exercise capacity as a predictor of survival among ambulatory patients with endstage renal disease. *Kidney International* 2004;**65**(2):719-24. [MEDLINE: 14717947]

Song 2018

Song YY, Hu RJ, Diao YS, Chen L, Jiang XL. Effects of exercise training on restless legs syndrome, depression, sleep quality, and fatigue among hemodialysis patients: a systematic review and meta-analysis. *Journal of Pain & Symptom Management* 2018;**55**(4):1184-95. [MEDLINE: 29247753]

SONG-HD 2017

Evangelidis N, Tong A, Manns B, Hemmelgarn B, Wheeler DC, Tugwell P, et al. Developing a set of core outcomes for trials in hemodialysis: an international Delphi survey. *American Journal of Kidney Disease* 2017;**70**(4):464-75. [MEDLINE: 28238554]

Spinowitz 2019

Spinowitz B, Pecoits-Filho R, Winkelmayer WC, Pergola PE, Rochette S, Thompson-Leduc P, et al. Economic and quality of life burden of anemia on patients with CKD on dialysis: a systematic review. *Journal of Medical Economics* 2019;**22**(6):593-604. [MEDLINE: 30813807]

Thompson 1996

Thompson RT, Muirhead N, Marsh GD, Gravelle D, Potwarka JJ, Driedger AA. Effect of anaemia correction on skeletal muscle metabolism in patients with end-stage renal disease: 31P magnetic resonance spectroscopy assessment. *Nephron* 1996;**73**(3):436-41. [MEDLINE: 8832604]

Tong 2015

Tong A, Crowe S, Chando S, Cass A, Chadban SJ, Chapman JR, et al. Research priorities in CKD: report of a national workshop conducted in Australia. *American Journal of Kidney Diseases* 2015;**66**(2):212-22. [MEDLINE: 25943716]

Tong 2018

Tong A, Manns B, Wang AY, Hemmelgarn B, Wheeler DC, Gill J, et al. Implementing core outcomes in kidney disease: report of the Standardized Outcomes in Nephrology (SONG) implementation

workshop. *Kidney International* 2018;**94**(6):1053-68. [MEDLINE: 30360959]

Tyler 1975

Tyler HR. Neurological aspects of uremia: an overview. *Kidney International - Supplement* 1975;(2):188-93. [MEDLINE: 169421]

USRDS 2017

Saran R, Robinson B, Abbott KC, Agodoa LYC, Bhave N, Bragg-Gresham J, et al. USRDS 2017 Annual Data Report: Epidemiology of kidney disease in the United States. *American Journal of Kidney Diseases* 2018;**71**(3 Suppl 1):A7. [MEDLINE: 29477157]

Wan 2014

Wan X, Wang W, Liu J, Tong T. Estimating the sample mean and standard deviation from the sample size, median, range and/or interquartile range. *BMC Medical Research Methodology* 2014;**14**(1):135. [MEDLINE: 25524443]

WHO 2010

World Health Organization. Global recommendations on physical activity for health. 2010. www.who.int/publications/i/item/9789241599979 (accessed 24 June 2021).

Wilkinson 2019

Wilkinson TJ, Watson EL, Xenophontos S, Gould DW, Smith AC. The "Minimum Clinically Important Difference" in frequently reported objective physical function tests after a 12-week renal rehabilitation exercise intervention in nondialysis chronic kidney disease. *American Journal of Physical Medicine & Rehabilitation* 2019;**98**(6):431-7. [MEDLINE: 30362979]

Wright 2011

Wright AA, Cook CE, Baxter GD, Dockerty JD, Abbott JH. A comparison of 3 methodological approaches to defining major clinically important improvement of 4 performance measures in patients with hip osteoarthritis. *Journal of Orthopaedic & Sports Physical Therapy* 2011;**41**(5):319-27. [MEDLINE: 21335930]

Wyld 2012

Wyld M, Morton RL, Hayen A, Howard K, Webster AC. A systematic review and meta-analysis of utility-based quality of life in chronic kidney disease treatments. *PLoS Medicine* 2012;**9**(9):e1001307. [MEDLINE: 22984353]

Yngman-Uhlin 2010

Yngman-Uhlin P, Friedrichsen M, Gustavsson M, Fernström A, Edéll-Gustafsson U. Circling around in tiredness: perspectives of patients on peritoneal dialysis. *Nephrology Nursing Journal* 2010;**37**(4):407-13. [MEDLINE: 20830948]

Young 2018

Young HM, March DS, Graham-Brown MP, Jones AW, Curtis F, Grantham CS, et al. Effects of intradialytic cycling exercise on exercise capacity, quality of life, physical function and cardiovascular measures in adult haemodialysis patients: a systematic review and meta-analysis. *Nephrology Dialysis Transplantation* 2018;33(8):1436-45. [MEDLINE: 29608708]



Zhao 2019

Zhao QG, Zhang HR, Wen X, Wang Y, Chen XM, Chen N, et al. Exercise interventions on patients with end-stage renal disease: a systematic review. *Clinical Rehabilitation* 2019;**33**(2):147-56. [MEDLINE: 30789077]

References to other published versions of this review Heiwe 2001b

Heiwe S, Jacobson SH. Exercise training for adults with chronic kidney disease. *Cochrane Database of Systematic Reviews* 2001, Issue 3. Art. No: CD003236. [DOI: 10.1002/14651858.CD003236]

Heiwe 2011

Susanne Heiwe and Stefan H Jacobson. Exercise training for adults with chronic kidney disease. *Cochrane Database of Systematic Reviews* 05 October 2011, Issue 10. Art. No: CD003236. [DOI: 10.1002/14651858.CD003236.pub2]

* Indicates the major publication for the study

CHARACTERISTICS OF STUDIES

Characteristics of included studies [ordered by study ID]

Abreu 2017

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Brazil Setting: HD unit Inclusion criteria: > 18 years; without motor skill disorders; AV fistula for vascular access in the upper limb and who have been on maintenance dialysis for at least 6 months Number: exercise group (32); control group (29) Mean age ± SD: 46.4 ± 14.6 years Sex (M/F): 36/25 Exclusion criteria: patients with autoimmune diseases, cancer, infectious diseases, acquired immunodeficiency syndrome, uncontrolled hypertension, unstable angina, malignant arrhythmias, pregnancies, lower limb amputations; history of stroke; neurological or cardiovascular disease; under ca-
Interventions	tabolizing drugs; regularly exercises; smokers; complied with < 75% of the training Duration of intervention
	• 12 weeks
	 Type: resistance Description: lower limbs exercises Position: seated Material: ankle weights and resistance bands Location: HD unit Duration of training sessions: 30 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 60% of 1RM Supervised by: physiotherapist Mode of delivery: face-to-face Tailoring: not reported Modifications/progression: not reported



Abreu 2017 (Continued)

- · Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Usual care

Outcomes

- BMI
- · Waist circumference
- Albumin
- HCT (%)
- Hb (g/dL)
- Calcium (mg/dL)
- Phosphorus (mg/dL)
- Potassium (mg/dL)
- hs-CRP (mg/dL)
- GPx (nmol/min/mL)
- NF-κB expression
- Nitrite (μM)
- · Nrf2 expression
- SCr (mg/dL)
- Carbohydrates
- · Energy intake
- Energy intake
- Lipids intake
- Protein (g/kg/day)
- Arm muscular area
- Body pain
- QoL (SF-36)

Notes

Funding:

- Coordenação de Aperfeiçoamento de Pessoal de Nível Superior (CAPES)
- Fundação de Amparo à Pesquisa do Estado do Rio de Janeiro (FAPERJ)
- Conselho Nacional de Desenvolvimento Científico e Tecnológico (CNPq)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding



Abreu 2017 (Continued)		
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Plausible effect size among missing outcomes enough to induce clinically relevant bias in observed effect size
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Abundis Mora 2017

Study characteristics	
Methods	 Study design: parallel RCT Study duration: 8 months Study follow-up period: 35 weeks
Participants	 Country: Mexico Setting: HD unit Inclusion criteria: prevalent HD patients ≥ 18 years Number: exercise group (14); control group (14) Age: average age 41 years Sex: 64% males Exclusion criteria: amputation of lower limbs; motor sequelae of cerebral vascular event; vascular accesses in the lower extremities
Interventions	Duration of intervention • 35 weeks Exercise group
	 Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 135 min/week minutes Duration of warm-up/cool-down: not reported Frequency: not reported Timing in relation to dialysis treatments: during Intensity: moderate (scale not reported) Supervised by: not reported Mode of delivery: not reported Tailoring: not reported Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none



Abundis Mora 2017 (Continued)

Control group

Usual care

Outcomes

· ECG parameters

Notes

• Abstract-only publication: author contacted for full results

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

ACTINUT 2013

C+4	.,,	hara	cto	rictica	
Stua	v c	nara	ctei	ristics	

Methods

- Study design: parallel RCT
- Study duration: not reported
- Study follow-up period: 24 weeks

Participants

- · Country: France
- Setting: 2 outpatient HD units
- Inclusion criteria: adults aged > 18 years; minimum HD vintage of 3 months and stable; no recent hospitalizations; no acute or chronic medical conditions that would make exercise training potentially hazardous or primary outcomes impossible to assess. Patients who meet the following criteria for PEW, meeting at least 3 of the 4 listed categories and at least 1 test in each of the selected categories:
 - o Serum chemistry criteria: serum albumin level < 38 g/L, or serum prealbumin < 300 mg/L



ACTINUT 2013 (Continued)

- Body mass criteria: BMI < 23 kg/m², or unintentional weigh loss > 5% over 3 months or > 10% over
 6 months
- Muscle mass criteria: lean body mass estimated by bioimpedance spectroscopy lower than the 10th percentile of an age-matched normal population. This method is validated in dialysis patients
- Dietary intake criteria: unintentional low dietary protein intake < 1 g/kg of ideal weight/day for at least 2 months, unintentional low dietary energy intake < 30 kcal/kg of ideal weight/day for at least 2 months
- Informed consent of the patient
- Number: exercise group (10); control group (11)
- Mean age \pm SD (years): exercise group (68.5 \pm 14.0); control group (70.8 \pm 15.2)
- Sex (M/F): total (12/9)
- Exclusion criteria: contraindication or inability to perform the physical exercise; inadequate dialysis
 Kt/V < 1.2; presence of a cardiac pacemaker (incompatible with the BCM measures); systemic inflammation CRP > 20 mg/L; pregnancy; patient under guardianship; participation in another clinical interventional trial; unstable on dialysis

Interventions

Duration of intervention

· 24 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: 5 minutes/not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 3 on RPE (1 to 10)
- · Supervised by: nephrologists, nurses, specialist in adapted physical activities
- · Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: 5 minutes added monthly adding to 30 minutes
- · Strategies to enhance adherence: follow-up monthly
- Adherence to intervention (mean \pm SD of attended sessions): 88% \pm 17%
- Co-intervention: dietary counselling

Control group

• Usual care + dietary counselling

Outcomes

- · PEW remission
- BMI
- FTI
- LTI
- Bicarbonate
- Albumin
- Hb
- Calcium
- Phosphate
- CRP
- Compliance



ACTINUT 2013 (Continued)

- Energy intake
- Pre-albumin
- Normalised protein catabolic rate
- Protein intake
- 6MWT
- COP area
- Knee extension maximal strength
- QoL (SF-36)

Notes

Funding:

- University Hospital of Nantes
- ACTICLAN prize (Fresenius Kabi France)
- Region of Pays de la Loire, France

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Performed by an independent collaborator
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Afshar 2010

a			
Studv	chara	icteristic	:s

Methods

- Study design: parallel RCT (3 arms)
- Study duration: not reported
- Study follow-up period: 8 weeks



Afshar 2010 (Continued)

Participants

- · Country: Iran
- · Setting: HD unit
- Inclusion criteria: maintenance HD>3 months; age>20 years; good compliance with the dialysis treatment (not missing more than 2 dialysis sessions in the prior month); and absence of lower extremity dialysis graft
- Number: resistance group (7); aerobic group (7); control group (7)
- Mean age ± SD (years): resistance group (51.0 ± 16.4); aerobic group (50.7 ± 21.1) control group (53.0 ± 19.4)
- Sex (M/F): not reported
- Median HD vintage ± IQR (months): resistance group (24.9 ± 18.7); aerobic group (25.7 ± 7.61); control group (24.9 ± 15.4)
- Exclusion criteria: presence of active infection or inflammation, autoimmunity disorders, and malignancy; presence of severe muscle weakness or interfering skeletal deformity; history of repeated episodes of hypoglycaemia; cardiopulmonary contraindications to resistance exercise such as MI within prior 6 months, active angina, and uncompensated congestive heart failure; hospitalisation during the prior month; cerebrovascular accidents within prior 6 months; and history of prior regular exercise training.

Interventions

Duration of intervention

• 8 weeks

Resistance exercise group

- Type: resistance
- · Description: lower limbs exercises
- · Position: not reported
- Material: ankle weights
- · Location: HD unit
- Duration of training sessions: 10 to 30 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 15 to 17 on RPE (6 to 20)
- · Supervised by: physician
- Mode of delivery: face-to-face
- Tailoring: individualised load
- Modifications/progression: increase in the number of repetitions and the weights
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Aerobic exercise group

- Type: aerobic
- · Description: stationary cycling
- Position: recumbent
- Material: ergometer
- Location: HD unit
- Duration of training sessions: 10 to 30 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 16 on RPE (6 to 20)
- · Supervised by: not reported



Afshar 2010 (Continued)

- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- Albumin
- Hb
- HDL
- LDL
- Total cholesterol
- Triglyceride
- CRP

Notes

• Funding: nil

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement



Afshar 2011

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: Iran Setting: HD unit Inclusion criteria: maintenance HD > 3 months; aged > 20 years; good compliance with dialysis treatment (not missing more than 2 dialysis sessions in the prior month) Number: exercise group (14); control group (14) Mean age ± SD (years): exercise group (50.7 ± 21.1); control group (53.0 ± 19.4) Sex (M/F): men only Median HD vintage ± IQR (months): exercise group (25.7 ± 7.6); control group (24.9 ± 15.4) Mean BMI ± SD (kg/m²): exercise group (22.7 ± 2.98); control group (22.3 ± 2.18) Exclusion criteria: presence of active infection or inflammation; autoimmune disorders; malignancy; presence of psychiatric diseases; severe musculoskeletal disorders; poor controlled diabetes; uncontrolled heart failure or pulmonary diseases; hospitalisation during the prior month; using drugs that influence serum cytokines levels; vascular access in the lower extremity; BMI > 25 kg/m²
Interventions	Duration or intervention • 8 weeks Exercise group • Type: aerobic
	 Description: stationary cycling Position: recumbent Material: ergometer Location: HD unit Duration of training sessions: 10 to 30 minutes Duration of warm-up/cool-down: 5/5 minutes Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 12 to 16 on RPE (6 to 20) Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity
	 Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none Control group
	Usual care
Outcomes	LeptinCRPSleep quality score
Notes	Funding: not reported



Afshar 2011 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Akiba 1995

AKIDA 1995	
Study characteristics	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Japan Setting: HD unit Inclusion criteria: not reported Number: exercise group (10); control group (10) Mean age ± SD (years): exercise group (38.4 ± 9.5); control group (40.6 ± 10.8) Sex (M/F): exercise group (2/8); control group (7/3) Mean HD vintage ± SD (months): exercise group (73.8 ± 47.2); control group (68.3 ± 41.5) Exclusion criteria: not reported
Interventions	Duration of intervention 12 weeks Exercise group Type: aerobic Description: stationary cycling



Akiba 1995 (Continued)

- Position: not reported
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 on RPE
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increase in the duration and then in the workload
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Usual care

Outcomes

- Watt max
- VO₂ max
- HR max
- Maximum lactate level
- Hb

Notes

• Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Plausible effect size among missing outcomes enough to induce clinically relevant bias in observed effect size



Akiba 1995 (Continued)			
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement	
Other bias	Unclear risk	Insufficient information to permit judgement	

Amini 2016

Study characteristics	
Methods	 Study design: parallel group RCT (3 arms*) Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: Iran Setting: in-hospital HD unit Inclusion criteria: signing the informed consent form to participate in the study; history of undergoing regular HD for at least 12 months Number: exercise group (32); control group (35) Mean age (years): aerobic group (54.3); control group (55.2) Sex (M/F): exercise group (21/11); control group (21/14) Exclusion criteria: not reported
Interventions	Duration of intervention • 8 weeks Exercise group
	 Type: aerobic Description: not reported Position: not reported Material: not reported Location: HD unit Duration of training sessions: not reported Duration of warm-up/cool-down: not reported Frequency: not reported Timing in relation to dialysis treatments: not reported Intensity: not reported Supervised by: researcher Mode of delivery: face-to-face and then via recorded CDs Tailoring: not reported Modifications/progression: not reported Strategies to enhance adherence: checklist, researcher available by phone, follow-up every 2 week in person or via phone Adherence to intervention: not reported Co-intervention: none Control group Usual care
Outcomes	AnxietyFatigue



Amini 2016	(Continued)
------------	-------------

Sleep quality

Notes

- *PMR (progressive muscle relaxation) group was not analysed
- · Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	Trial described as "double blind" but no description of how the intervention, exercise, was blinded. Due to the nature of the intervention, it is unlikely that the participants were blinded to group allocation.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

AVANTE-HEMO 2020

Study characteristics

Methods

- Study design: parallel RCT (3 arms)
- Study duration: not reported
- Study follow-up period: 12 weeks

Participants

- Country: Mexico
- · Setting: HD unit
- Inclusion criteria: regular HD 2 or 3 times/week; any sex; age > 18 years; no previous exercise
- Number: aerobic group (15); resistance group (15); control group (15)
- Mean age \pm SD (years): aerobic group (32 \pm 10); resistance group (30 \pm 9); control group (27 \pm 8)
- Sex (M/F): aerobic group (7/8); resistance group (5/10); control group (9/6)
- Median HD vintage (IQR) (months): aerobic group (24, 4 to 36); resistance group (19, 8 to 36); control group (28, 8 to 48)
- Mean BMI±SD (kg/m²): aerobic group (19.7±3.1); resistance group (21.5±1.9); control group (19±1.8)



AVANTE-HEMO 2020 (Continued)

Exclusion criteria: amputation; hospitalisation in the last 3 months; 1 HD session/week; severe effort
angina (CC3) or stage 4 of the NYHA scale; pregnancy; severe dyspnoea; femoral fistula; arrhythmias;
precordial pain; orthopaedic or neurological compromises or cognitive alterations affecting study
participation; intolerance to oral nutritional supplement or intolerance/contraindications to the exercise routine according to the nephrologist and cardiologist evaluation

Interventions

Duration of interventions

• 12 weeks

Aerobic exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 to 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 to 13 on RPE (6 to 20)
- · Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increase in duration
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: Diet plan and nutritional supplement

Resistance exercise group

- Type: resistance
- Description: upper and lower limbs exercises
- Position: not reported
- Material: resistance bands
- · Location: HD unit
- Duration of training sessions: 40 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 to 13 on RPE (6 to 20)
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increase in frequency, intensity, type and time
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: Diet plan and nutritional supplement

Control group

• Diet plan and nutritional supplement

Outcomes

- Time up and go (sec)
- 6MWT



AVANTE-HEMO 2020 (Continued)

- Handgrip
- Sit-to-stand test
- Weight (kg)
- BMI (kg/m²)
- Midarm circumference (cm)
- Arm muscle circumference (mm)
- Arm muscle area (cm²)
- Fat mass (%)
- Triceps skinfold thickness (mm)
- Physical activity
- Hb (g/dL)
- · Total lymphocytes
- SCr (mg/dL)
- Albumin (g/dL)
- Phosphorus (mg/dL)
- Potassium (mmol/L)
- CRP (mg/L)
- HRQoL

e	9
	e

• Funding: National Kidney Foundation

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias



Bennett 2013

Stud	vc	har	act	eris	tics
JLUU	, .	uui	uct	ei is	ucs

Methods

- Study design: cluster step-wedge RCT (3 arms)
- · Study duration: not reported
- Study follow-up period: 48 weeks

Participants

- · Country: Australia
- Setting: community satellite HD clinics (15 sites; 5 clusters)
- Inclusion criteria: ESKD receiving HD; aged ≥ 18 years; able to understand and speak English; on HD > 12 weeks
- Number: group 1 (51); group 2 (61); group 3 (59)
- Mean age ± SD (years): 68.1 ± 12.6
- Sex (M/F): 107/64
- Median HD vintage (IQR): 44 months (26.0 to 85.5)
- Exclusion criteria: pregnancy; lower limb amputation; hospitalisation in the four weeks prior to study commencement; considered not suitable on medical grounds for the intervention

Interventions

Duration of intervention

- 12 weeks
- There were five clusters (clinics) in each of the 3 groups: the first group received 36 weeks of exercise
 training, the second group were followed for 12 weeks before receiving 24 weeks of exercise and the
 third group were followed for 24 weeks before receiving 12 weeks of exercise

Exercise group

- · Type: resistance
- · Description: lower body exercises
- · Position: seated
- · Material: resistance bands and tubing
- · Location: HD unit
- · Duration of training sessions: varied minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: not reported
- · Supervised by: exercise physiologist
- Mode of delivery: face-to-face
- · Tailoring: individualised not further defined
- Modifications/progression: increase in resistance of resistance bands
- · Strategies to enhance adherence: record cards reviewed weekly
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

Usual care

Outcomes

- DBP
- Falls and falls confidence
- Dialysis exercise adequacy
- Four-square step test
- Time up and go
- · Sit-to-stand test
- Community activity involvement



Bennett 2013 (Continued)

QoL

Notes

Funding:

- Alfred Deakin Postdoctoral Fellowship Scheme, Deakin University
- Centre for Nursing Research-Deakin University and Monash Health Partnership

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Allocation concealed at the time of participant consent
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Plausible effect size among missing outcomes enough to induce clinically relevant bias in observed effect size
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Burrows 2018

Ctudy	chara	ctoric	tica

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 24 weeks
Participants	 Country: USA Setting: HD unit Inclusion criteria: not reported Number: exercise group (9); control group (not reported) Mean age ± SD (years): not reported Sex (M/F): not reported Exclusion criteria: not reported



Burrows 2018 (Continued)

Interventions

Duration of intervention

• 24 weeks

Exercise group

- · Type: combined
- Description: stationary cycling and total body resistance and balance exercises
- Position: not reported
- Material: ergometer, resistance bands
- · Location: HD unit and at home
- Duration of training sessions: 15 to 30 intra-HD and 2 sessions at home of not reported duration minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 5 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: moderate
- · Supervised by: not reported
- Mode of delivery: not reported
- · Tailoring: individualised not further defined
- Modifications/progression: progressive not further defined
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: volume control

Control group

· Volume control

Outcomes

HRQoL (KDQOL)

Notes

- · Abstract-only publication
- Authors contacted for full results
- · Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias)	Unclear risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding



Burrows 2018	(Continued)
Subjective ou	tcomes

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Carmack 1995

Study characteristic	s
Methods	Study design: parallel RCTStudy duration: not reported
	Study follow-up period: 14 weeks
Participants	Country: USA Setting: UD units
	Setting: HD unitsInclusion criteria: not reported
	Number: exercise group (23); control group (25)
	 Mean age (range): 44.09 years (20 to 72)
	• Sex (M/F): 29/19
	 Mean HD vintage (range): 29.52 months (1 to 173)
	 Exclusion criteria: physical or mental impairment that precluded undergoing submaximal exercise tolerance tests and participating in an exercise programme; severe cardiac problems; leg vascular access; leg prosthesis

Interventions

Duration of intervention

• 10 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- Position: not reported
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 to 30 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: not reported
- Supervised by: not reported
- Mode of delivery: not reported
- · Tailoring: individualised not further defined
- Modifications/progression: not reported
- Strategies to enhance adherence: self-monitoring using report cards and letters to family members
- Adherence to intervention: not reported
- Co-intervention: none



Carmack	1995	(Continued)

Control group

· Not reported

Outcomes

- VO₂ peak
- Depression
- Stress appraisal measures
- Anxiety
- Frequency of physical complaints and symptoms

Notes

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

CHAIR 2015

Methods

- Study design: parallel RCT
- Study duration: not reported
- Study follow-up period: 12 weeks

Participants

- · Country: Japan
- Setting: outpatient dialysis
- Inclusion criteria: patients treated with HD; ≥ 60 years; ambulatory



CHAIR 2015 (Continued)

- Number: exercise group (12); control group (15)
- Median age, range (years): exercise group (69, 61 to 78); control group (69, 64 to 79)
- Sex (M/F): exercise group (8/4); control group (11/4)
- Median HD vintage, range) (years): exercise group (14, 6 to 22); control group (15, 6 to 79)
- Exclusion criteria: symptomatic ischaemic heart disease; symptomatic peripheral artery disease; arthritis; history of stroke with severe paralysis; chronic obstructive pulmonary disease; pregnancy; patient was judged as inappropriate for the study by the attending physician

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: aerobic
- · Description: chair-stand exercise
- · Position: seated-standing
- · Material: chair
- · Location: HD unit
- Duration of training sessions: 15 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: just before
- · Intensity: not reported
- Supervised by: physician and physical therapist
- · Mode of delivery: face-to-face
- Tailoring: individualised duration
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

• Passive stretch exercise with assistance by a physical therapist

Outcomes

- Serum albumin
- Hb
- Mini-Mental
- 6MWT
- · Isometric knee extensor strength
- FIM
- QoL

Notes

· Funding: nil

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Use of an external randomisation centre
Allocation concealment (selection bias)	Low risk	Central randomisation



CHAIR 2015 (Continued)		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Plausible effect size among missing outcomes enough to induce clinically relevant bias in observed effect size
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Chang 2010

Study characteristics	
Methods	 Study design: quasi-RCT Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: Taiwan Setting: HD units Inclusion criteria: ≥ 18 years; given their consent to participate in the study; on maintenance dialysis for at least 3 months Number: exercise group (36); control group (37) Mean age ± SD (years): exercise group (50.8 ± 10.7); control group (52.0 ± 8.7) Sex (M/F): exercise group (26/10); control group (24/11) Mean BMI ± SD (kg/m²): exercise group (22.3 ± 3.2); control group (22.0 ± 3.1) Mean HD vintage ± SD (months): exercise group (77.2 ± 46.9); control group (84.5 ± 49.9) Exclusion criteria: not reported
Interventions	Duration of intervention • 8 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: recumbent • Material: ergometer • Location: HD unit • Duration of training sessions: 10 to 30 minutes • Duration of warm-up/cool-down: 5/not reported minutes



Chang 2010 (Continued)

- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 to 13 on RPE (6 to 20)
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increase in time over the first 3 sessions
- · Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- IL-18 (pg/mL)
- IL-6 (pg/mL)
- QoL
- Depression severity (BDI)

Notes

• Funding: Taipei Medical University and Shin Kong Memorial Hospital fund

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias



Chen 2010

Study characteristics Methods · Study design: parallel RCT

- Study duration: 31 months
- Study follow-up period: 26 weeks

Participants · Country: USA

- Setting: HD units
- Inclusion criteria: ≥ 30 years; serum albumin < 4.2 g/dL; HD 3 times/week for at least 3 months with ≥80% compliance
- Number: exercise group (25); control group (25)
- Mean age \pm SD (years): exercise group (71.1 \pm 12.6); control group (66.9 \pm 13.4)
- Sex (M/F): exercise group (12/10); control group (11/11)
- Mean HD vintage \pm SD (years): exercise group (2.6 \pm 2.6); control group (4.8 \pm 5.2)
- Mean BMI \pm SD (kg/m²): exercise group (25.7 \pm 7.1); control group (27.7 \pm 7.8)
- Exclusion criteria: unstable cardiovascular disease or any uncontrolled chronic condition; cardiac surgery; retina laser therapy; MI; joint replacement or lower extremity fracture within the last 6 months; severe cognitive impairment; lower extremity amputation; or current strength training

Interventions

Duration of intervention

· 26 weeks

Exercise group

- · Type: resistance
- · Description: lower body exercises
- Position: seated and semi-recumbent
- Material: ankle weights
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: 5 to 10/5 minutes
- · Frequency: 2 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: moderate (6) on modified OMNI scale
- Supervised by: supervised not further defined
- · Mode of delivery: not reported
- · Tailoring: start with very low weights and increased according to individual capacity
- Modifications/progression: increase in ankle weights ad 20lbs
- Strategies to enhance adherence: not reported
- Adherence to intervention: mean (SD) of % of adherence to prescription 81% (15%)
- Co-intervention: none

Control group

· Stretching exercises with light resistance bands

Outcomes

- Muscular strength
- Physical performance
- Whole-body lean mass
- Whole-body fat mass
- Leisure-time physical activity
- **HROoL**
- Adherence to exercise



Chen 2010 (Continued)

Notes

Funding

- National Institute of Diabetes and Digestive and Kidney (NIDDK)
- USDA
- NIH General Clinical Research Center
- William B. Schwartz Nephrology Fund at Tufts Medical Center

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	Sham exercise in the control arm
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	Sham exercise in the control arm
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Cho 2018

Study	char	actor	ictics
Study	cnard	ıcter	istics

Methods	,
---------	---

- Study design: parallel RCT (4 arms)
- Study duration: not reported
- Study follow-up period: 12 weeks

Participants

- Country: South Korea
- Setting: HD unit
- Inclusion criteria: age ≥ 20 years; HD vintage ≥ 6 months; HD treatment 3 times/week; no hospitalisations during the previous 3 months, except for vascular access repair; no amputations or prostheses in upper and lower extremities; cognitive capacity sufficient for communication, able to ambulate and wear the physical activity monitor for 7 days; good compliance with the study protocol
- Number: aerobic group (15); resistance group (14); combination group (15); control group (13)



Cho 2018 (Continued)

- Mean age \pm SD (years): aerobic group (55.2 \pm 11.9); resistance group (52.9 \pm 8.8); combination group (50.0 \pm 14.3); control group (59.4 \pm 10.8)
- Sex (M/F): aerobic group (2/9); resistance group (6/4); combination group (8/4); control group (7/6)
- Mean BMI \pm SD (kg/m²): aerobic group (26.0 \pm 1.4); resistance group (22.8 \pm 1.2); combination group (23.5 \pm 0.8); control group (25.4 \pm 1.3)
- Mean HD vintage ± SD (months): aerobic group (54.8 ± 96.4); resistance group (47.6 ± 79.2); combination group (87.8 ± 70.5); control group (61.4 ± 36.5)
- Exclusion criteria: any acute infectious or other inflammatory illnesses; current malignancy except basal cell carcinoma; acute MI or unstable angina within the past 12 months; current heart or lung failure or severe liver disease; severe uncontrolled diabetes; severe retinal diseases, such as proliferative diabetic retinopathy and vitreous haemorrhage; and orthopaedic disorders exacerbated by activity

Interventions

Duration of intervention

• 12 weeks

Aerobic exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: not reported
- · Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: training load adjusted to performance
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Resistance exercise group

- Type: resistance
- Description: upper and lower limbs exercise with resistance bands
- Position: seated or supine
- Material: resistance bands and soft weights
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: not reported
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: "progression tailored to performance"
- Modifications/progression: increased resistance of the resistance bands
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none



Cho 2018 (Continued)

Combination (aerobic and resistance) exercise group

- · Type: combined
- Description: combination of aerobic and resistance exercise
- Position: seated or supine
- Material: ergometer, resistance bands and soft weights
- · Location: HD unit
- Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: not reported
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: combination of aerobic and resistance exercise
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Warm-up stretches

Outcomes

- Average total sleep time (min)
- Average wake after sleep onset (min)
- Average movement index (%)
- Average fragmentation index (%)
- Average sleep fragmentation index (%)
- Average sleep efficiency (%)
- Sit-to-stand test
- 6MWT
- LVEF
- LVMI
- Cardiac performance index

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Bloc randomisation; assumed computer-generated
Allocation concealment (selection bias)	Low risk	Sealed envelopes
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias)	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding



Cno 2018	(Continued)
Objective	e outcomes

Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Cooke 2018

Study characteristi	ics
---------------------	-----

Methods	

- Study design: parallel RCT
- · Study duration: not reported
- Study follow-up period: 16 weeks

Participants

- · Country: Canada
- · Setting: HD unit
- Inclusion criteria: stage 5 CKD; stable in-centre HD regimen for ≥ 12 weeks prior to recruitment; recent
 cardiac evaluation (< 1 year) showing sufficient cardiac function to undergo the exercise program
- Number: exercise group (15); control group (12)
- Mean age \pm SD (years): exercise group (58.2 \pm 17.2); control group (52.5 \pm 15.4)
- Sex (M/F): exercise group (7/3); control group (7/3)
- Mean BMI \pm SD (kg/m²): exercise group (25.6 \pm 4.3); control group (27.2 \pm 6.1)
- Exclusion criteria: any physical or psychological disability that would impact study participation; iPTH > 250 pmol/L within 30 days prior; dysrhythmia or severe cardiac disease or peripheral arterial disease; severe hyperkalaemia (> 6.5 mmol/L) for the last 2 weeks; active cancer; postdialytic SBP ≥ 160 mm Hg or DBP ≥100 mm Hg within 4 weeks prior; anticipated living donor kidney transplant or other planned major surgery over the study duration

Interventions

Duration of intervention

· 16 weeks

Exercise group

- Type: aerobic
- Description: stationary cycling
- · Position: not reported
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: to reach the intensity minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 16 on RPE (6 to 20)
- Supervised by: not reported



Cooke 2018 (Continued)

- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention (median (IQR) of attended sessions): 60% (42% to 79%)
- Co-intervention: none

Control group

• Usual care

Outcomes

- Adherence
- BMI (kg/m²)
- Waist:hip ratio
- Interdialytic weight gain (kg)
- Gait speed (m/sec)
- Grip strength (kg)
- Peripheral SBP (mm Hg)
- Peripheral DBP (mm Hg)
- Central SBP (mm Hg)
- Central DBP (mm Hg)
- Central pulse pressure (mm Hg)
- MAP (mm Hg)
- Carotid-femoral pulse wave velocity (m/sec)
- Augmentation index 75 (%)
- HR (bpm)

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation stratified by age and sex; assumed computer-generated
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement



Cooke 2018 (Continued)		
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

CYCLE-HD 2016

 Study design: cluster parallel RCT Study duration: not reported Study follow-up period: 30 weeks
 Country: UK Setting: HD units Inclusion criteria: prevalent HD patient (> 3 months), aged ≥ 18 years; able and willing to give informed consent Number: exercise group (65); control group (65) Mean age ± SD (years): exercise group (55.5 ± 15.5); control group (58.9 ± 14.9) Sex (M/F): exercise group (53/8); control group (42/23) Mean HD vintage, range (years): exercise group (1.2, 0.5 to 3.7); control group (1.3, 0.4 to 3.2) Exclusion criteria: unable to participate in current exercise programme due to perceived physical of psychological barriers; unable to undergo MRI scanning (metal implants, severe claustrophobia); unfit to undertake exercise according to the American College of Sports Medicine guidelines
Duration of intervention 26 weeks Exercise group Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 30 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 12 to 14 on RPE Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity Modifications/progression: duration and resistance adjusted to progress training Strategies to enhance adherence: regular visits Adherence to intervention: not reported Co-intervention: none



CYCLE-HD 2016 (Continued)

Outcomes

- DBP
- Mean BP
- SBP
- Cardiac index
- HR
- Stroke volume index
- Total peripheral resistance index
- Length of stay
- Adverse outcomes
- Anxiety
- Depression

Notes

- · Abstract-only data
- Authors contacted for full results
- Funding: NIHR in the United Kingdom (grant reference number CS-2013-13-014; JOB) and supported by Kidney Research UK

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Block randomisation; assumed computer-generated
Allocation concealment (selection bias)	Low risk	Cluster trial, dialysis shifts were randomised. Participants were assigned to a shift before inclusion in the study
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Dashtidehkordi 2019

Study	, ch	ara	cto	rict	ice
Stuuy	' CII	uı u	ctei	เวเ	LS

Methods
 Study design: parallel RCT



Dashtidehkordi 2019 (Continued)

- · Study duration: not reported
- Study follow-up period: 8 weeks

Participants

- · Country: Iran
- · Setting: HD units
- Inclusion criteria: aged between 18 and 65; history of at least 3 months of HD; no physical and mental
 disability, no known ischaemic heart disease; no MI and angina during the last 3 months; based on
 the patients' history, no acute pulmonary disease so that the patient needs oxygen therapy during
 dialysis; no history of stroke or transient ischaemic attacks over the past 3 months; no skeletal-muscle
 disorder that prevent the patient from exercising (pedalling the stationary bicycle); doing 3 sessions
 of 4-hour dialysis/week
- Number: exercise group (30); control group (30)
- Mean age (years): exercise group (51.2); control group (55.6)
- Sex (M/F): exercise group (3/24); control group (3/22)
- Mean HD vintage (years): exercise group (5.5); control group (4.5)
- Exclusion criteria: unwillingness to continue participating in the study; the presence of any disorder, including cardiovascular, pulmonary and musculoskeletal disorders during the study which may prevent the patient from exercise; not doing the exercises for 3 consecutive sessions and 6 non-consecutive sessions

Interventions

Duration of intervention

• 8 weeks

Exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 2 x 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: not reported
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reportedStrategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Stretching exercises

Outcomes

• Heath promoting behaviours

Notes

· Funding: nil

|--|



Dashtidehkordi 2019 (Continue	ed)	
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Low risk	"closed packets"
Blinding of participants and personnel (perfor- mance bias) All outcomes	Unclear risk	Stretching program in the control group but did not specify whether the participants were blinded
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Deligiannis 1999

Study characteristics	5
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 26 weeks
Participants	 Country: Greece Setting: not reported Inclusion criteria: undergoing HD Number: exercise group (30); control group (30) Mean age ± SD (years): exercise group (48.0 ± 12.0); control group (48.0 ± 11.0) Sex (M/F): exercise group (17/13); control group (15/15) Mean HD vintage ± SD (years): exercise group (6.3 ± 3.0); control group (6.2 ± 3.6) Exclusion criteria: documented MI during the previous 6 months; symptoms of angina or heart failure (NYHA class ≥ II); severe hypertension, DM, or any other disease that might interfere with autonomic regulation; sinus rhythm during a resting ECG; medication that might interfere with autonomic regulation (i.e. beta-blockers)
Interventions	Duration of intervention • 26 weeks Exercise group • Type: combined



Deligiannis 1999 (Continued)

- Description: bicycling and/or walking, callisthenics, steps, swimming, or ball games followed by a lowintensity resistance program
- · Position: not reported
- · Material: not reported
- · Location: not reported
- Duration of training sessions: 70 minutes
- Duration of warm-up/cool-down: 10/10 minutes
- Frequency: 3 to 4 times/week
- Timing in relation to dialysis treatments: on non-HD days
- Intensity: 60% to 70% of max HR
- Supervised by: physician, exercise physiologist, and physical education instructor
- · Mode of delivery: face-to-face
- Tailoring: individualised intensity
- · Modifications/progression: adjusted every 15 days to maintain intensity
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care

Outcomes

- HCT (%)
- Hb
- Urea (mg%)
- SCr (mg%)
- Potassium (mEq/L)
- Sodium (mEq/L)
- Calcium (mEq/L)
- Phosphate (mg%)
- 24-hour mean HR
- HR variability index
- Mean RR interval (sec)
- SDNN (sec)
- Sum of beats
- VO max
- Lactic acid

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding



Deligiannis 1999 (Continued)		
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Deligiannis 1999a

Deligiannis 1999a	
Study characteristics	;
Methods	 Study design: parallel RCT (3 arms) Study duration: not reported
	Study follow-up period: 26 weeks
Participants	Country: Greece
	Setting: not reported
	 Inclusion criteria: undergoing HD treatments
	 Number: exercise group 1 (16); exercise group 2 (10); control group (12)
	 Mean age ± SD (years): exercise group 1 (46.4 ± 3.9); exercise group 2 (51.4 ± 12.5); control group (50.2 ± 7.9)
	 Sex (M/F): exercise group 1 (11/5); exercise group 2 (8/2); control group (4/8)
	 Mean HD vintage ± SD (months): exercise group 1 (78.0 ± 62.0); exercise group 2 (62.0 ± 37.0); control group (79.0 ± 86.0)
	 Exclusion criteria: unstable hypertension; congestive heart failure; cardiac arrhythmias (III according to Lown); recent MI or unstable angina; DM; active liver disease; serious anaemia; peripheral vascula disease
Interventions	Duration of intervention
	• 26 weeks
	Exercise group 1
	Type: combined
	 Description: stationary cycling, callisthenics, steps and flexibility exercises
	Position: not reported
	Material: ergometer or treadmill
	Location: not reported
	Duration of training sessions: 50 to 70 minutes
	Duration of warm-up/cool-down: 10/10 minutes
	Frequency: 3 times/week
	Timing in relation to dialysis treatments: on non-HD days
	• Intensity: 50% to 70% of max HR



Deligiannis 1999a (Continued)

- Supervised by: physician and physical education teachers
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: Intensity adjusted
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Exercise group 2

- · Type: aerobic
- Description: cycling on ergometer + simple flexibility and muscular extension exercises
- · Position: not reported
- · Material: ergometer
- · Location: home
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 5 times/week
- · Timing in relation to dialysis treatments: not during
- Intensity: 50% to 60% of max HR
- Supervised by: physician and physical education teachers
- · Mode of delivery: face-to-face
- Tailoring: individualised not further defined
- Modifications/progression: program modified to physical adaptation
- · Strategies to enhance adherence: monthly follow-up at home
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care

Outcomes

- · HR (resting)
- SBP
- DBP
- Left ventricular internal dimension (diastole and systole)
- Intra-ventricular septal thickness
- Left ventricular posterior wall
- Left ventricular mass index
- HCT
- WBC
- Urea
- SCr
- Uric acid
- Glucose
- Potassium (mEq/L)
- Sodium (mEq/L)
- Calcium (mg%)
- Phosphorus (mg%)
- Fe (mg%)
- · Exercise time
- Maximal metabolic equivalents
- HR peak at exercise



Deligiannis 1999a (Continued)

- Maximum pulmonary ventilation
- VO₂ max
- · Lactic acid

Notes

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

de Lima 2013

- Study design: parallel RCT (3 arms)
- Study duration: not reported
- Study follow-up period: 8 weeks

Participants

- Country: Brazil
- Setting: HD unit
- Inclusion criteria: HD 3 times/week; aged 18 and 75 years; not practising any physical activity
- Number: resistance group (11); aerobic group (11); control group (11)
- Mean age \pm SD (years): resistance group (49.6 \pm 9.1); aerobic group (43.1 \pm 13.3); control group (43.5 \pm 11.1)
- Sex (M/F): resistance group (7/4); aerobic group (7/4); control group (6/5)
- Mean BMI \pm SD (kg/m²): resistance group (26.0 \pm 5.1); aerobic group (23.0 \pm 5.6); control group (27.4 \pm 3.7)



de Lima 2013 (Continued)

- Mean HD vintage ± SD (years): resistance group (5.4 ± 4.0); aerobic group (6.4 ± 4.4); control group (6.5 ± 4.2)
- Exclusion criteria: uncontrolled arterial hypertension; ischaemic cardiopathy; amputation; deep vein thrombosis; excessive pallor; severe dyspnoea; femoral fistula; arrhythmias; precordial pain; orthopaedic or neurological compromising; cognitive alterations affecting participation in the proposed protocol

Interventions

Duration of intervention

• 8 weeks

Resistance exercise group

- Type: resistance
- Description: lower limbs exercises
- · Position: seated
- Material: not reported
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- · Intensity: not reported
- · Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: individualised intensity
- · Modifications/progression: adjusted every 15 days to maintain intensity
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Aerobic exercise group

- Type: aerobic
- Description: stationary cycling
- · Position: seated
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 2 to 3 on RPE
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

Usual care

Outcomes

Prurit symptoms



de Lima 2013 (Continued)

- Hb
- Calcium
- Phosphorus
- Potassium
- FEV1
- FVC
- Maximal expiratory pressure
- Maximal inspiratory pressure
- Step test
- QoL

Notes

• Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Shuffling envelopes
Allocation concealment (selection bias)	Low risk	Quote: "envelops, without external marks"
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

DePaul 2002

Study characteristic	S
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	Country: Canada



DePaul 2002 (Continued)

- · Setting: HD unit
- Inclusion criteria: on HD for > 3 months; administered EPO for the treatment of anaemia; Hb level > 9.0 g/dL; able to maintain sitting and standing balance without assistance; ambulatory without assistance
- Number: exercise group (20); control group (18)
- Mean age ± SD (years): exercise group (55 ± 16); control group (54 ± 14)
- Sex (M/F): exercise group (10/10); control group (13/14)
- Exclusion criteria: ischaemic heart disease; recent MI < 6 months; uncontrolled hypertension; pericardial or pleural friction rub; aortic stenosis; active musculoskeletal lower-extremity problem; history of vertebral fraction caused by osteoporosis; patients who participated in team sports or formally organized exercise programs

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: combined
- · Description: stationary cycling + lower limbs strength training
- Position: seated
- Material: ergometer and response seated leg curl thigh extension pulley weight system
- · Location: HD unit
- Duration of training sessions: 20-varied minutes
- Duration of warm-up/cool-down: 2 minutes/not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during and just before or after
- Intensity: somewhat strong on RPE
- · Supervised by: kinesiologist
- · Mode of delivery: face-to-face
- Tailoring: individualised intensity and duration
- · Modifications/progression: intensity adjusted
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

• Progressive, no resistance, low-intensity, range-of-motion exercises

Outcomes

- Muscular strength
- 6MWT (metres)
- HRQoL
- Dialysis symptoms (Laupacis KDQ)

Notes

Funding:

- Father Sean O'Sullivan Research Centre, St Joseph's Healthcare, Hamilton
- Ortho Biotech/Janssen-Ortho Inc, North York, Ontario, Canada

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Random numbers table



DePaul 2002 (Continued)		
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	Sham exercise in the control arm
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	Sham exercise in the control arm
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Private funding. Funder's involvement not specified

DIALY-SIZE 2016

Study characteristic	s
Methods	 Study design: parallel RCT (4 arms) Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Canada Setting: HD unit Inclusion criteria: adults aged ≥18 years; dialysis-dependent for ≥3 consecutive months; receiving ≥3 dialysis treatments/week; mobile (any distance, walking aid permitted); at least one non-prosthetic limb; capable of providing consent Number: cycling group (8); weights group (7); combined group (8); control group (8) Median age, IQR (years): cycling group (66.9, 55.8 to 82.4); weights group (59.7, 45.9 to 81.4); combined group (60.3, 54.7 to 68.4); control group (49.3, 43.0 to 62.3) Sex (M/F): cycling group (8/0); weights group (6/1); combined group (3/5); control group (7/1) Median HD vintage, IQR (years): cycling group (3.7, 2.4 to 4.6); weights group (2.8, 2.0 to 4.0); combined group (2.9, 0.7 to 2.3); control group (3.3, 1.2 to 6.2) Median BMI, IQR (kg/m²): cycling group (23.6, 22.2 to 25.7); weights group (25.9, 24.6 to 29.9); combined group (25.3, 20.0 to 30.8); control group (24.2, 20.4 to 33.8) Exclusion criteria: currently enrolled in a clinical trial; missing an average of > 2 dialysis sessions/month; planned move or modality change within the next 4 months; currently enrolled in a structured exercise programme; scheduled hospitalisation for > 1 week; unstable during HD; any uncontrolled medical condition that would preclude participation in a low/moderate-intensity exercise program
Interventions	Duration of intervention • 12 weeks



DIALY-SIZE 2016 (Continued)

Aerobic exercise group

- Type: aerobic
- Description: stationary cycling
- Position: not reported
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 15 to 43 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 14 on RPE
- Supervised by: kinesiologist
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: weekly increase in duration by 2.5 minutes
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: 904 sessions completed over 1039 offered
- Co-intervention: none

Resistance exercise group

- · Type: resistance
- Description: lower limbs exercises
- Position: not reported
- Material: ankle weights and resistance bands
- · Location: HD unit
- Duration of training sessions: varied minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 to 14 on RPE
- Supervised by: kinesiologist
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: increase in weights or resistance to maintain intensity
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Combined aerobic and resistance exercise group:

- · Type: combined
- Description: all aerobic and resistance exercise groups
- Position: not reported
- Material: aerobic + resistance
- · Location: HD unit
- Duration of training sessions: varied minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 14 on RPE
- Supervised by: kinesiologist
- Mode of delivery: face-to-face



DIALY-SIZE 2016 (Continued)

- Tailoring: individualised intensity
- Modifications/progression: aerobic + resistance
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Stretching

Outcomes

- 6MWT
- Physical performance
- Strength (quadriceps)
- Sit-to-stand test
- QoL

Notes

Funding

- University Hospital Foundation
- Clinical Research Fellowship award from Alberta Innovates-Health Solutions

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Serial numbered, opaque, sealed envelopes
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Participants were blinded to aim and hypothesis but intervention was not blinded
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	Participants were blinded to aim and hypothesis. Lack of blinding on the intervention may still affect patient-reported outcomes.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias



Dobsak 2012

Study characteristics	•
Methods	 Study design: parallel RCT (3 arms*) Study duration: not reported Study follow-up period: 20 weeks
Participants	 Country: Czech Republic Setting: HD unit Inclusion criteria: at least 12 months of regular HD; clinically stable; optimised pharmacological treatment unchanged for 1 month before the start of the study Number: exercise group (11); control group (10) Mean age ± SD (years): exercise group (58.4 ± 7.2); control group (60.1 ± 0.0) Sex (M/F): exercise group (4/7); control group (4/6) Mean BMI ± SD (kg/m²): exercise group (28.6 ± 3.0); control group (26.9 ± 3.6) Mean HD vintage ± SD (years): exercise group (4.1 ± 2.1); control group (4.1 ± 2.3) Exclusion criteria: uncontrolled hypertension; venous thromboembolism; implanted cardiac pacemakers; unstable angina pectoris; heart failure; severe neurological diseases (epilepsy, multiple sclerosis, parkinsonism), severe orthopaedic complications (total hip or knee replacement); chronic bronchopulmonary disease; low urea clearance (Kt/V > 1.2)
Interventions	Duration of intervention • 20 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: not reported • Material: ergometer • Location: HD unit • Duration of training sessions: 20 to 40 minutes • Duration of warm-up/cool-down: 5/5 minutes • Frequency: 3 times/week • Timing in relation to dialysis treatments: during • Intensity: 30% to 60% of peak power • Supervised by: not reported • Mode of delivery: not reported • Tailoring: individualised intensity • Modifications/progression: increased duration at 5 weeks to 40 minutes • Strategies to enhance adherence: not reported • Adherence to intervention: not reported • Co-intervention: none Control group
Outcomes	 Usual care Walking test Strength (leg extensor) QoL
Notes	 * electrostimulation group not included in this review • Funding • grant IGA MZ CR No. NS/10096-4



Dobsak 2012 (Continued)

o grant MSM 0021622402

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Dong 2011

Study characteristic	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 24 weeks
Participants	 Country: USA Setting: HD unit Inclusion criteria: ≥ 18 years; HD > 3 months; adequate dose of dialysis (double pool Kt/V ≥ 1.2) on a 3 times/week HD program using a biocompatible HD membrane Number: exercise group (15); control group (17) Mean age ± SD (years): exercise group (46.5 ± 12.1); control group (40.2 ± 13.5) Sex (M/F): exercise group (9/6); control group (12/5) Mean BMI ± SD (kg/m²): exercise group (27.5 ± 6.3); control group (29.1 ± 6.4) Exclusion criteria: active inflammatory or infectious disease; pregnancy; hospitalisation within 1 month prior to the study; not capable of exercise due to cardiovascular disease or osteoarthritis
Interventions	Duration of intervention



Dong 2011 (Continued)

20 weeks

Exercise group

- · Type: resistance
- · Description: lower limbs exercises
- · Position: seated
- Material: pneumatic leg press machine
- · Location: HD unit
- Duration of training sessions: 3 sets of 12 repetitions minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: just before
- Intensity: 70% of 1RM
- Supervised by: study personnel
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: 1RM re-evaluated and training adjusted at 3 months and 6 months
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: nutritional supplementation

Control group

· Nutritional supplementation

Outcomes

- BMI (kg/m²)
- · Weight (kg)
- FM% from BIA
- FM% from anthropometry
- FM (kg)
- LBM (%)
- Leg LBM (%)
- LBM (kg)
- Leg LBM (kg)
- Waist/hip ratio
- Bicarbonate
- Serum albumin
- Total protein (g/dL)
- Hb
- Cholesterol (mmol/L)
- Glucose (mg/dl)
- CRP (mg/L)
- SCr (mg/dL)
- Dietary energy intake (kcal/kg/day)
- Pre-albumin (mg/dL)
- Dietary protein intake (g/kg/day)
- 1-RM (lb)

Notes

Funding

- National Institutes of Health
- · Diabetes Research Training Center
- · National Institute of Diabetes, Digestive and Kidney Diseases



Dong 2011 (Continued)

- Clinical Translational Science Award from the National Center for Research Resources
- Chinese Society of Nephrology
- International Society of Peritoneal Dialysis
- National Kidney Foundation
- Council of Renal Nutrition

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

EXCITE 2014

Study characteristics	
Methods	 Study design: parallel RCT Study duration: 15 months Study follow-up period: 6 months
Participants	 Country: Italy Setting: HD unit Inclusion criteria: stage G5 CKD Number: exercise group (151); control group (145) Mean age ± SD (years): exercise group (63.0 ± 13.0); control group (64.0 ± 14.0) Sex (M/F): exercise group (70/34); control group (103/20) Mean BMI ± SD (kg/m²): exercise group (26.0 ± 4.0); control group (27.0 ± 6.0)



EXCITE 2014 (Continued)

 Exclusion criteria: physical (e.g. amputation) or clinical (severe effort angina or stage 4 NYHA heart failure, any intercurrent illness requiring hospitalisation) limitations to mobility or a high degree of fitness, that is the ability to walk a distance of 0.550 m in 6 minutes during the standard walking test

Interventions

Duration of intervention

• 26 weeks

Exercise group

- · Type: aerobic
- · Description: home walking sessions
- Position: not applicable
- Material: not reported
- · Location: home
- Duration of training sessions: varied according to baseline level minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: on non-HD days
- Intensity: varied according to baseline 6MWT but described as low intensity
- Supervised by: prehabilitation team ensure training of dialysis personnel but exercise sessions were not directly supervised
- Mode of delivery: not reported
- · Tailoring: individualised not further defined
- · Modifications/progression: adjusted according to 6MWT
- Strategies to enhance adherence: encouragement by dialysis personnel
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Usual care

Outcomes

- · Need for medication
- Hospitalisations
- AV fistula events
- Adverse events
- Death
- FEV1 (L)
- FVC (L)
- Maximal inspiratory mouth pressure (kPa)
- Vital capacity (L)
- 6MWT
- · Sit-to-stand test
- · Lower extremity strength
- QoL

Notes

· Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Stratified. Assumed computer-generated



EXCITE 2014 (Continued)		
Allocation concealment (selection bias)	Low risk	Central allocation
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: Brazil Setting: HD unit Inclusion criteria: ≥ 18 years; undergoing HD > 6 months; clinically stable; no pulmonary, musculoskeletal, or neurological disease; agreed to participate in the study by signing the informed consensorm Number: exercise group (22); control group (22) Mean age ± SD (years): exercise group (44.3 ± 11.3); control group (42.6 ± 11.2) Sex (M/F): exercise group (8/12); control group (9/10) Mean BMI ± SD (kg/m²): exercise group (22.8 ± 2.8); control group (22.8 ± 1.9) Mean HD vintage ± SD (years): exercise group (6.7 ± 4.7); control group (7.2 ± 3.8) Exclusion criteria: need for urgent or elective surgical intervention during the protocol; decompensation of prior heart disease with arrhythmia and/or precordial pain; ischaemic cardiac event (< 3 months); significant valvular heart disease or dysrhythmia; continuous and/or night-time oxygen need for gait assistance devices or lower-limb orthoses
Interventions	Duration of intervention • 8 weeks Exercise group • Type: aerobic



Fernandes 2019 (Continued)

- · Description: stationary cycling
- Position: seated
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: 10/10 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during (1 hour after commencement of dialysis)
- Intensity: 50% to 70% of max HR
- · Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- FCV
- FEV
- · Peak expiratory flow
- Maximal inspiratory pressure
- Maximal expiratory pressure
- Peak flow
- SBP
- DBP
- HR
- Respiratory frequency
- Peripheral oxygen saturation
- Borg during 6MWT
- 6MWT
- HCT
- Hb
- SCr
- Urea
- Kt/V
- Albumin

Notes

• Funding: CAPES

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Sealed and opaque envelopes



Fernandes 2019 (Continued)		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Frey 1999

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: USA Setting: HD unit Inclusion criteria: 25 to 65 years undergoing HD Number: exercise group (5); control group (6) Mean age ± SD (years): exercise group (40.0 ± 11.0); control group (53.0 ± 13.0) Sex (M/F): exercise group (3/2); control group (3/3) Exclusion criteria: SBP > 160 mm Hg and DBP > 95 mm Hg at the beginning of the second hour of dialysis; average inter-dialytic weight gain > 3.5 kg between dialysis treatments; DM; unstable angina
Interventions	Duration of intervention • 8 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: seated • Material: multigym • Location: HD unit • Duration of training sessions: 45 minutes • Duration of warm-up/cool-down: 5/5 minutes • Frequency: 3 times/week • Timing in relation to dialysis treatments: during



Frey 1999 (Continued)

- Intensity: 60% to 80% of max HR
- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: duration increased by 3 min/day from week 5 to 8
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- Kilocalories
- Protein intake
- Serum prealbumin
- · Serum transferrin
- Albumin
- Kt/V

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Plausible effect size among missing outcomes enough to induce clinically relevant bias in observed effect size
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement



Frih 2017a

Stud	vc	har	act	eris	tics
JLUU	, .	uui	uct	ei is	ucs

Methods

- Study design: parallel RCT
- Study duration: 15 months
- Study follow-up period: 16 weeks

Participants

- Country: Tunisia
- · Setting: HD unit
- · Inclusion criteria: undergoing HD
- Number: exercise group (28); control group (22)
- Mean age \pm SD (years): exercise group (64.2 \pm 3.4); control group (65.2 \pm 3.1)
- Sex (M/F): all males
- Mean HD vintage ± SD (months): exercise group (72.7 ± 12.7); control group (73.6 ± 13.4)
- Mean BMI \pm SD (kg/m²): exercise group (25.4 \pm 2.8); control group (24.3 \pm 3.2)
- Exclusion criteria: chronic lung disease; ischaemic heart disease; uncontrolled arrhythmias or hypertension; haemodynamic instability or musculoskeletal disorders exacerbated by exercise; exercising regularly before starting the experiment

Interventions

Duration of intervention

· 16 weeks

Exercise group

- · Type: combined
- Description: upper and lower limbs strengthening exercises + cycling and treadmill walking
- Position: not applicable
- Material: ergometer, treadmill, multigym
- Location: multigym
- Duration of training sessions: 40 minutes
- Duration of warm-up/cool-down: 10/10 minutes
- Frequency: 4 times/week
- · Timing in relation to dialysis treatments: on non-HD days
- Intensity: 50% of 1-RM and 5 to 6 on RPE
- Supervised by: physiotherapy and physical training technicians
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: load increased by 5% of 1RM every month
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group:

· Usual care

Outcomes

- STS-10 (sec)
- Sit-to-stand test (60 sec)
- Handgrip strength
- TUG test (sec)
- 6MWT (metres)
- SBP
- DBP
- CRP



Frih 2017a (Continued)

- Hb (g/dL)
- Albumin (g/L)
- Total cholesterol
- HDL cholesterol
- · LDL cholesterol
- Mini nutritional assessment long-form score
- QoL (SF-36)
- Anxiety score
- Depression score

Notes

• Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Giannaki 2013a

Study characteristics	
Methods	 Study design: parallel RCT (3 arms) Study duration: 25 months Study follow-up period: 26 weeks
Participants	Country: GreeceSetting: HD unit



Giannaki 2013a (Continued)

- Inclusion criteria: dialysis for ≥ 3 months; adequate dialysis delivery; stable clinical condition; have RLS, no medication for RLS prior to the study
- Number: exercise group (12); control group (12)
- Mean age ± SD (years): exercise group (59.2 ± 11.8); control group (58.0 ± 10.7)
- Sex (M/F): exercise group (9/3); control group (8/4)
- Mean BMI \pm SD (kg/m²): exercise group (27.7 \pm 3.6); control group (26.5 \pm 4.4)
- Mean HD vintage \pm SD (months): Exercise group (24.0 \pm 15); control group (30 \pm 26)
- Exclusion criteria: diagnosed neuropathies or reasons for being in a catabolic state within 3 months prior to the start of the study; CRP > 3.0 mg/L; unable to exercise

Interventions

Duration of intervention

• 26 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- · Material: ergometer
- Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 60% to 65% of Wmax
- · Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: individualised intensity
- · Modifications/progression: adjusted intensity every monthly
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

Usual care

Outcomes

- Total body fat
- % leg fat
- Extramyocellular lipids
- · Subcutaneous adipose tissue
- Muscle percentage
- Total LBM
- RLS severity
- North Staffordshire Royal Infirmary walk test
- Gait speed test (fast walk)
- Gait speed test (normal walk)
- Muscle cross-sectional area
- Sit-to-stand test
- Depression
- · Daily sleepiness
- · Quality of sleep
- QoL



Giannaki 2013a (Continued)

Notes

Funding

- National and Community Funds of the Greek Ministry of Development-General Secretariat of Research and Technology
- European Social Fund

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome as- sessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Goldberg 1983

Study characteristics	;
-----------------------	---

М	et	hα	ds

- Study design: parallel RCT
- · Study duration: not reported
- Study follow-up period: not reported

Participants

- · Country: USA
- Setting: not reported
- Inclusion criteria: HD patients receiving treatments for 4 to 6 hours, 3 times/week using either a coil
 or hollow-fibre dialyser
- Number: exercise group (14); control group (11)
- Mean age \pm SD (years): exercise group (38.0 \pm 15.0); control group (37.0 \pm 12.0)
- Sex (M/F): exercise group (8/6); control group (7/4)
- Mean HD vintage ± SD (months): exercise group (22.2 ± 17.1); control group (40.1 ± 29.7)



Goldberg 1983 (Continued)

 Exclusion criteria: unstable angina pectoris; cardiac arrhythmias; haemodynamically significant valvular heart disease; congestive heart failure; poorly controlled hypertension; severe retinal disease; insulin-dependent DM; hypothyroidism

Interventions

Duration of intervention

· 26 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- · Material: ergometer
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 60% to 65% of Wmax
- Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: individualised intensity
- Modifications/progression: adjusted intensity every monthly
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care

Outcomes

- Graded exercise treadmill duration
- VO₂ max
- HR
- BP
- · Psychological function
- · Plasma triglyceride levels
- Plasma HGL cholesterol levels
- · Fasting plasma glucose
- Fasting plasma insulin
- Glucose disappearance
- Body weight
- Hb
- · Red cell mass
- HCT

Notes

· Funding: NIH

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement



Goldberg 1983 (Continued)		
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Groussard 2015

Study characteristics				
Methods	Study design: parallel RCTStudy duration: not reported			
	Study follow-up period: 24 weeks			
Participants	Country: France			
	Setting: HD unit			
	 Inclusion criteria: aged 20 to 85 years; dialysis for at least 2 years; consent of the patient's cardiologist Number: exercise group (10); control group (10) 			
	 Mean age ± SD (years): exercise group (66.5 ± 4.6); control group (68.4 ± 3.7) 			
	 Sex (M/F): exercise group (5/3); control group (7/3) 			
	 Mean HD vintage ± SD (months): exercise group (36.6 ± 8.2); Control group (41.2 ± 8.1) 			
	 Exclusion criteria: orthopaedic problems that prevented cycling during dialysis; participation in another study 			
Interventions	Duration of intervention			
	• 12 weeks			
	Exercise group			
	Type: aerobic			
	Description: stationary cycling			
	Position: seated			
	Material: ergometer			
	Location: HD unit			
	 Duration of training sessions: 15 to 30 minutes 			



Groussard 2015 (Continued)

- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 55% to 60% of Wpeak
- · Supervised by: professional team with expertise in physical activity
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: duration increased by 15 minutes over the first 2 weeks; intensity monitored and adapted
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care

Outcomes

- HDL (g/L)
- LDL (g/L)
- Ox-LDL (U/L)
- Total cholesterol (g/L)
- Triglycerides (g/L)
- GPx/g Hb
- GSH/GSSG
- · SOD/g Hb
- JOD/gill
- Peak power (W)
- Peak power (W/kg)VO₂ peak (L/min)
- VO₂ peak (mL/min/kg)
- 6MWT

Notes

Funding

- Amgen
- Baxter
- Hemotech
- Meditor
- Roche
- Association des Néphrologues Centre Auvergne

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding



Groussard 2015 (Continued)		
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Private funding. Funder's involvement not specified

Harter 1985		
Study characteristics		
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 26 weeks 	
Participants	 Country: USA Setting: not reported Inclusion criteria: HD patients receiving treatments for 4 to 6 hours, 3 times/week Number: exercise group (15); control group (12) Mean age ± SD (years): exercise group (40.0 ± 4.0); control group (36.0 ± 3.0) Sex (M/F): Exercise group (8/5); control group (7/5) Mean HD vintage ± SD (months): exercise group (23.0 ± 5.0); control group (40.0 ± 9.0) Exclusion criteria: unstable angina pectoris; cardiac arrhythmias; haemodynamically significan valvular heart disease; clinically significant or symptomatic cerebrovascular; peripheral vascular, o coronary atherosclerosis; congestive heart failure; poorly controlled hypertension; electrolyte imbal ance; severe retinal disease; insulin-dependent DM, hypothyroidism 	
Interventions	Duration of intervention • 52 weeks Exercise group • Type: aerobic • Description: cycling or walking • Position: not applicable • Material: ergometer, running track • Location: not reported • Duration of training sessions: 45 minutes • Duration of warm-up/cool-down: not reported • Frequency: not reported • Timing in relation to dialysis treatments: on non-HD days • Intensity: first 50% to 60%, then 70% to 80% of VO ₂ max • Supervised by: physician, nurse, exercise physiologist	



Harter 1985 (Continued)

- Mode of delivery: face-to-face
- Tailoring: individualised intensity and duration
- · Modifications/progression: progressive in duration and intensity
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care and nandrolone or placebo (factorial RCT)

Outcomes

- RDI
- Minnesota Multiphasix Personality Inventory
- Pleasant event schedule
- Unpleasant event schedule
- VO₂ peak
- Triglyceride
- HDL
- Plasma glucose level

Notes

- Funding: NIH
- We have contacted the author who has confirmed that the publications by Goldberg 1985 and Carney 1987 were reports of the same trial

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	'As-treated' analysis done with substantial departure of the intervention received from that assigned at randomisation
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Low risk	The study appears to be free of other sources of bias



IHOPE 2019

Study characteristics	S
Methods	 Study design: parallel RCT (3 arms*) Study duration: 5 years Study follow-up period: 52 weeks
Participants	 Country: USA Setting: HD units Inclusion criteria: receiving HD treatment ≥ 3 days/week, dialysis vintage ≥ 3 months, aged 30 to 80 years not currently receiving intradialytic oral nutritional supplementation or participating in intradialytic exercise Number: exercise + protein group (49); protein group (45) Mean age ± SD (years): exercise + protein group (53.7 ± 11.4), protein group (56.6 ± 13.0) Sex (M/F): exercise + protein group (29/20), protein group (23/22) Mean HD vintage ± SD (months): exercise + protein group (34.3 ± 34.8), protein group (45.6 ± 38.7) Mean BMI ± SD (kg/m): exercise + protein group (31.9 ± 8.3), protein group (30.6 ± 7.1) Exclusion criteria: persistent Hb levels < 10 g/dL; weight > 300 pounds; currently receiving any form of intradialytic protein supplementation (oral, enteral, or parenteral) or participating in any form of intradialytic exercise training; chronic obstructive pulmonary disease and decompensated chronic heart failure; on dialysis treatment for < 3 months (or enrolment may be postponed)
Interventions	Duration of intervention • 52 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: not reported • Material: ergometer • Location: HD unit • Duration of training sessions: 30 to 45 minutes • Duration of warm-up/cool-down: not reported • Frequency: 3 times/week • Timing in relation to dialysis treatments: during • Intensity: 12 to 14 on RPE • Supervised by: supervised by research staff • Mode of delivery: face-to-face • Tailoring: individualised intensity • Modifications/progression: progressive in duration • Strategies to enhance adherence: not reported • Adherence to intervention: 80% • Co-intervention: protein supplementation
	 Protein supplementation: 30 g whey protein supplement at each dialysis session
Outcomes	 Shuttle walk test Gait speed Sit-to-stand test (rep) TUG test (sec)



IHOPE 2019 (Continued)

- leg extension
- leg flexion
- BMI (kg/m²)
- Whole body lean mass (kg)
- Whole body fat percent (%)
- Leg lean mass (kg)
- Whole body BMD (g/cm²)
- Leg BMD (g/cm²)
- Hip BMD (g/cm²)
- BF
- Augmentation index at HR 75 bpm
- energy intake (Kcal/kg/day)
- protein intake (g/kg/day)
- Albumin (g/L)
- IL-6
- CRP (mg/L)
- central pulse wave velocity
- QoL: PhysicalQoL: Mental

Notes

- * Patients receiving non-nutritive beverage were not included in this review (44 participants)
- · Funding: NIH

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Performed by a research member that was not involved in data collection at that site
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias



Johansen 2006

Study characteristics	5			
Methods	Study design: factorial RCT (4 arms)			
	Study duration: unclear			
	Study follow-up period: 12 weeks			
Participants	Country: USA			
	Setting: outpatient HD units			
	 Inclusion criteria: adequate dialysis delivery with Kt/V 1.2 and good compliance with dialysis treatment (i.e., not missing more than 2 dialysis treatments in the month before enrolment) 			
	Number: exercise/exercise + nandrolone group (40); placebo/nandrolone group (39)			
	• Mean age \pm SD (years): exercise/exercise + nandrolone group (55.0 \pm 13.1); placebo/nandrolone group (31.9 \pm 13.6)			
	• Sex (M/F): exercise/exercise + nandrolone group (25/15); placebo/nandrolone group (24/15)			
	 Mean BMI ± SD (kg/m²): exercise/exercise + nandrolone group (27.6 ± 7.8); placebo/nandrolone group (26.3 ± 5.7) 			
	 Median HD vintage, range (months): exercise (33, 3.5 to 108); exercise + nandrolone group (14, 4 to 152); placebo group (25.5, 3 to 156); nandrolone group (40.0, 3 to 288) 			
	 Exclusion criteria: dialysis < 3 months; catabolic state; unable to give informed consent; active IV drug users; thigh dialysis graft; contraindications to resistance training such as MI within 6 months; active angina; uncompensated congestive heart failure; orthopaedic or musculoskeletal limitations 			
Interventions	Duration of intervention			
	• 12 weeks			
	Exercise group			
	Type: resistance			
	Description: lower limbs exercises			
	Position: not reported			
	Material: ankle weights			
	Location: HD unit			
	 Duration of training sessions: varied minutes 			
	 Duration of warm-up/cool-down: not reported 			
	Frequency: 3 times/week			
	Timing in relation to dialysis treatments: during			
	• Intensity: 60% of 3-RM			
	Supervised by: study personnel			
	Mode of delivery: face-to-face			
	Tailoring: individualised load			
	Modifications/progression: increase for 2 to 3 sets of 10 reps; weight also increased			
	Strategies to enhance adherence: not reported			
	Adherence to intervention: not reported			
	Co-intervention: none or anabolic steroid (factorial design) .			
	Control group			
	Usual care or usual care and anabolic steroid (factorial design)			
Outcomes	Weight Lean body mass			
	Lean body mass Tak m			
	• Fat mass			



Johansen 2006 (Continued)

- Muscle size (quadriceps muscle area)
- SC
- Muscular strength: knee extension 3RM (lb)
- Muscular strength: hip abduction 3RM (lb)
- Muscular strength: hip flexion 3RM (lb)
- Muscular strength: isokinetic knee extension at 90 degrees/s (Nm)
- Muscular strength: isokinetic knee extension at 120 degrees/s (Nm)
- Gait speed (cm/s)
- Stairs
- · Sit-to-stand test
- Accelerometry
- Human activity profile, maximum activity score
- Human activity profile, adjusted activity score
- SF-36 physical functioning
- Fatigue
- Anger

Notes

Funding

- National Institute of Diabetes and Digestive and Kidney Diseases
- Organon, Inc., Roseland, NJ

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Block randomisation. Assumed computer-based
Allocation concealment (selection bias)	Low risk	Performed independently from investigators and block sizes unknown
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Private funding. Funder's involvement not specified



Jong 2004

Methods	Study design: parallel RCT
	Study duration: 7 months
	Study follow-up period: 12 weeks
Participants	Country: Korea
	Setting: not reported
	Inclusion criteria: not reported, adults undergoing CAPD
	Number: exercise group (19); control group (17)
	Mean age (years): exercise group (48.8); control group (49.8) (40.7) (40.7)
	• Sex (M/F): exercise group (12/7); control group (11/6)
	Exclusion criteria: not reported
Interventions	Duration of intervention
	• 12 weeks
	Exercise group
	Type: aerobic
	Description: walking
	Position: not applicable
	Material: not applicable
	Location: at home
	Duration of training sessions: varied
	Duration of warm-up/cool-down: not reported
	Frequency: 2 to 4 times/week Timing in solution to disharity and a second solution to a
	Timing in relation to dialysis treatments: outside treatments
	Intensity: not reported Comparised by some
	Supervised by: none Made of delivery who are or focus to focus
	Mode of delivery: phone or face-to-face Tailoring: not reported.
	Tailoring: not reported Modifications (progression) not reported
	Modifications/progression: not reportedStrategies to enhance adherence: verbal persuasion
	Adherence to intervention: not reported
	Co interpreting your
	Control group • Usual care
	• Usual Care
Outcomes	Physical functioning (SF-36)
	• VO ₂ max
	Serum albumin
	• Cholesterol
	Triglyceride
	HDL cholesterol
	LDL cholesterol
	• HCT
	Serum urea
	• SCr
Notes	Abstract-only publication



Jong 2004 (Continued)

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Koh 2009

Study characteristics

ds

- Study design: parallel RCT (3 arms)
- Study duration: not reported
- Study follow-up period: 24 weeks

Participants

- Country: Australia
- · Setting: renal units
- Inclusion criteria: ≥ 18 years on stable adequate dialysis therapy; URR 70% for 3 months were eligible for inclusion
- Number: intra-HD exercise group (27); home exercise group (21); control group (22)
- Mean age \pm SD (years): intra-HD exercise group (52.3 \pm 10.9); home exercise group (52.1 \pm 13.6); control group (51.3 \pm 14.4)
- Sex (M/F): intra-HD intra-HD exercise group (10/5); home exercise group 2 (11/4); control group (8/8)
- Mean BMI ± SD (kg/m²): intra-HD exercise group (27.6 ± 7.2); home exercise group 2 (27.9 ± 4.9); control group (28.6 ± 7.3)
- Mean HD vintage \pm SD (months): intra-HD exercise group (32.1 \pm 26.7); home exercise group 2 (37.0 \pm 31.1); control group (25.8 \pm 22.2)



Koh 2009 (Continued)

Exclusion criteria: unstable angina, lower limb amputation, already meet or exceed the exercise recommendation of 120 minutes of moderate-intensity physical activity/week, participating in, or propose to participate in, another clinical intervention study within 30 days prior to study entry

Interventions

Duration of intervention

24 weeks

Intra-HD exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: ergometer
- Location: HD unit
- Duration of training sessions: 15 to 45 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 13 on RPE (6 to 20)
- Supervised by: supervised not further defined
- · Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- · Modifications/progression: increasing duration and resistance
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Home exercise group

- · Type: aerobic
- · Description: walking
- Position: not applicable
- Material: not reported
- Location: home
- Duration of training sessions: 15 to 45 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: not reported
- Intensity: 12 to 13 on RPE (6 to 20)
- · Supervised by: unsupervised
- · Mode of delivery: not reported
- · Tailoring: individualised intensity
- Modifications/progression: increase in duration from 15 to 45 minutes
- Strategies to enhance adherence: fortnightly phone calls
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- 6MWT
- Timed up-and-go test
- · Grip strength



Koh 2009 (Continued)

- Weekly physical activity
- SF-36
- HR
- SBP
- DBP
- Pulse pressure
- Central SBP
- Central DBP
- Central pulse pressure
- Mean arterial pressure
- Ejection duration
- · Time to reflection
- Pulse pressure amplification
- P1 height
- Augmentation
- Augmentation index
- Augmentation index at HR of 75 bpm
- Pulse wave velocity aortic
- Pulse wave velocity peripheral

Notes

• Funding: Renal Research Tasmania

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	External to the investigators
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	'As-treated' analysis done with substantial departure of the intervention received from that assigned at randomisation
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias



Konstantinidou 2002

Study characteristics

Methods

- Study design: parallel RCT (4 arms)
- · Study duration: not reported
- Study follow-up period: 26 weeks

Participants

- · Country: Greece
- Setting: renal units
- Inclusion criteria: undergoing regular HD with an artificial kidney for at least 6 months, 3 sessions/week, 4 hours each session
- Number: exercise group 1 (21); exercise group 2 (12); exercise group 3 (12); control group (13)
- Mean age ± SD (years): exercise group 1 (46.4 ± 13.9); exercise group 2 (48.3 ± 12.1); exercise group 3 (51.4 ± 12.5); control group (50.2 ± 7.9)
- Sex (M/F): exercise group 1 (11/5); exercise group 2 (8/2); exercise group 3 (8/2); control group (4/8)
- Mean HD vintage \pm SD (months): exercise group 1 (78.0 \pm 62.0); exercise group 2 (72.0 \pm 66.0); exercise group 3 (62.0 \pm 37.0); control group (79.0 \pm 86.0)
- Exclusion criteria: unstable hypertension; congestive heart failure (grade > II according to NYHA); cardiac arrhythmias (≥ III according to Lown); recent MI or unstable angina; persistent hyperkalaemia before dialysis; DM; active liver disease; bone disease that puts the patient at risk of fracture; arthritic or orthopaedic problems limiting exercise; peripheral vascular disease; undisciplined patients

Interventions

Duration of intervention

· 26 weeks

Exercise group 1

- · Type: combined
- Description: callisthenics, steps and flexibility exercises + stretching and low-weight resistance program
- · Position: not reported
- · Material: ergometer
- · Location: rehab centre
- Duration of training sessions: 30 to 40 minutes
- Duration of warm-up/cool-down: 10/10 minutes
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: on non-HD days
- Intensity: 60% to 70% of HR max
- Supervised by: sports physician, physical education teachers
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: progressive not further defined
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: EPO
- Duration: 26 weeks

Exercise group 2

- · Type: combined
- Description: stationary cycling + lower limbs exercises
- · Position: not reported
- Material: ergometer, resistance bands and weights
- · Location: HD unit



Konstantinidou 2002 (Continued)

- Duration of training sessions: 20 min cycling + resistance minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 70% of HR max
- · Supervised by: sports physician, physical education teachers
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: EPO

Exercise group 3

- · Type: combined
- · Description: stationary cycling
- · Position: not reported
- Material: ergometer
- · Location: home
- Duration of training sessions: 30 + resistance minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 5 times/week
- Timing in relation to dialysis treatments: not during, home-based
- Intensity: 50% to 60% of max HR
- · Supervised by: unsupervised
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: monthly home visits
- · Adherence to intervention: not reported
- Co-intervention: EPO

Control group

EPO

Outcomes

- Maximum HR
- VO₂ peak
- Exercise time
- Ventilation max
- VO₂ at anaerobic threshold
- Lactic acid
- Respiratory exchange ratio

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement



Konstantinidou 2002 (Continu	ed)	
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Kopple 2007

Study characteristics	5
Methods	 Study design: parallel RCT (4 arms) Study duration: not reported Study follow-up period: 20 weeks
Participants	 Country: USA Setting: not reported Inclusion criteria: clinically stable HD patients; aged 25 to 65 years; undergoing HD 3 times/week fo at least 6 months Number: endurance training (10); strength training (15); endurance + strength training (12); control group (14) Mean age ± SE (years): endurance training (46 ± 4); strength training (46 ± 3); endurance + strength training (43 ± 4); control group (41 ± 3) Sex (M/F): endurance training (6/4); strength training (9/6); endurance + strength training (7/5); control group (9/5) Mean HD vintage ± SE (months): endurance training (45.9 ± 14.1); strength training (51.9 ± 12.4); endurance + strength training (38.3 ± 5.8); control group (51.4 ± 21.0) Exclusion criteria: no history of hospitalisation or systemic infection for at least 3 months; active can cer other than basal cell carcinoma; severe heart, lung, or liver disease; poorly controlled hypertension; acute or chronic inflammatory disease including tuberculosis or acquired immunodeficiency disease; insulin-dependent diabetes; severe osteoporosis, neuropathy, or musculoskeletal disease; am putations involving the lower extremities; or a joint infirmity that would prevent participants from exercising
Interventions	Duration of intervention • 20 weeks



Kopple 2007 (Continued)

Endurance training group

- · Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 60 minutes
- Duration of warm-up/cool-down: 5 to 10/not reported minutes
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 50% of peak oxygen consumption
- · Supervised by: investigator
- · Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: increasing duration and attempt to go from interval to continuous training
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Strength training group

- · Type: resistance
- Description: lower limbs exercises
- · Position: NA
- Material: leg extension/leg curl and leg press/calf extension apparatus
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: just before
- Intensity: 70% of RM-5
- Supervised by: investigator
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: increasing resistance
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Combined exercise group

- · Type: combined
- Description: 50% of endurance + 50% of strength
- Position: recumbent
- Material: endurance and strength
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: just before and during
- Intensity: same as endurance and strength
- · Supervised by: investigator
- Mode of delivery: face-to-face



Kopple 2007 (Continued)

- Tailoring: endurance and strength
- Modifications/progression: endurance and strength
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- · Mean body mass
- Fat mass
- BMI
- Mid-thigh muscle area
- HI
- HCT
- Albumin
- CRP
- Protein intake
- · Energy intake
- Growth factors mRNA levels

Notes

Funding

- National Institutes of Health
- General Clinical Research Center

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported



Kopple 2007 (Continued)

Other bias Low risk The study appears to be free of other sources of bias

Koufaki 2002

Study characteristics	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: UK Setting: renal rehabilitation gym Inclusion criteria: undergoing CAPD or HD Number: exercise group (26); control group (22) Mean age ± SD (years): exercise group (57.8 ± 14.3); control group (51.0 ± 18.9) Sex (M/F): exercise group (13/5); control group (11/4) Mean BMI ± SD (kg/m²): exercise group (25.7 ± 3.3); control group (24.7 ± 3.5) Mean HD vintage ± SD (months): exercise group (41.4 ± 45.2); control group (53.4 ± 52.5) Exclusion criteria: evidence of recent MI (within 6 weeks); uncontrolled dysrhythmias; uncontrolled hypertension; unstable angina; severe uncontrolled DM; symptomatic left ventricular dysfunction or neurological disorder with a functional deficit; demonstrating an inter-dialytic weight ≥ 2.5 kg, predialysis potassium ≥ 5.5 mmol/L and urea clearance (Kt/V ≤ 1 mL/min/L)
Interventions	Duration of intervention 12 weeks Exercise group Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 18 to 40 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 90% of VO ₂ max Supervised by: not reported Mode of delivery: not reported Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not eported Co-intervention: none Control group
	Usual care
Outcomes	 VO₂ peak VO₂ peak/kg



Koufaki 2002 (Continued)

- VE peak
- Power output
- HR peak
- VO₂/HR
- Body mass
- BMI
- Self-reported physical activity level
- Hb
- Albumin
- TCO₂
- PTH
- Nutritional status
- Sit-to-stand test 5 cycles (sec)
- Sit-to-stand test in 60 sec
- · NSRI walk test

Notes

• Funding: Jansen-Cilag Ltd research scholarship

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Coin tossing
Allocation concealment (selection bias)	Low risk	Coin tossing
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Private funding. Funder's involvement not specified

Koufaki 2003

Study characteristics



Study design: parallel RCTStudy duration: not reported	
Study duration: not reported	
Study follow-up period: 12 weeks	
 Country: UK Setting: not reported Inclusion criteria: undergoing dialysis (HD or PD) Number: 12 (numbers per group not reported) Mean age ± SD: 47.8 ± 20.3 years Sex (M/F): all males Exclusion criteria: not reported 	
Duration of intervention	
12 weeks Exercise group	
 Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 40 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: at ventilatory threshold Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity Modifications/progression: increasing duration Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: EPO 	
 VO₂ peak Walk performance Hb Oxygen uptake at the ventilatory threshold Oxygen uptake kinetics 	
Abstract-only publicationFunding: not reported	
_	 Setting: not reported Inclusion criteria: undergoing dialysis (HD or PD) Number: 12 (numbers per group not reported) Mean age ± SD: 47.8 ± 20.3 years Sex (M/F): all males Exclusion criteria: not reported Duration of intervention 12 weeks Exercise group Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 40 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: at ventilatory threshold Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity Modifications/progression: increasing duration Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: EPO Control group EPO VO2 peak Walk performance Hb Oxygen uptake at the ventilatory threshold Oxygen uptake kinetics Abstract-only publication

Support for judgement

Exercise training for adults undergoing maintenance dialysis (Review)

Bias

Authors' judgement



Random sequence generation (selection bias) Allocation concealment (selection bias) Blinding of participants and personnel (performance bias) All outcomes Blinding of outcome assessment (detection bias) Objective outcomes Blinding of outcome assessment (detection bias) Subjective outcomes Low risk Insufficient information to permit judgement No patient-reported outcome Incomplete outcome data (attrition bias) All outcomes Selective reporting (reporting freporting bias) Other bias Unclear risk Insufficient information to permit judgement Insufficient information to permit judgement	Koufaki 2003 (Continued)		
Blinding of participants and personnel (performance bias) All outcomes Blinding of outcome assessment (detection bias) Objective outcomes Blinding of outcome assessment (detection bias) Subjective outcomes Low risk No patient-reported outcome Incomplete outcome data (attrition bias) All outcomes Selective reporting (reporting (reporting bias)) Selective reporting (reporting bias) Insufficient information to permit judgement	•	Unclear risk	Insufficient information to permit judgement
and personnel (performance bias) All outcomes Blinding of outcome assessment (detection bias) Objective outcomes Blinding of outcome assessment (detection bias) Subjective outcomes Low risk No patient-reported outcome Incomplete outcome data (attrition bias) All outcomes Selective reporting (reporting bias) Selective reporting (reporting bias) Insufficient information to permit judgement Insufficient information to permit judgement		Unclear risk	Insufficient information to permit judgement
Sessment (detection bias) Objective outcomes Blinding of outcome assessment (detection bias) Subjective outcomes Incomplete outcome data (attrition bias) All outcomes Selective reporting (reporting bias) Selective reporting (reporting bias) Insufficient information to permit judgement Insufficient information to permit judgement	and personnel (perfor- mance bias)	High risk	No blinding
sessment (detection bias) Subjective outcomes Incomplete outcome data (attrition bias) All outcomes Selective reporting (reporting bias) Insufficient information to permit judgement Insufficient information to permit judgement Insufficient information to permit judgement	sessment (detection bias)	Unclear risk	Insufficient information to permit judgement
(attrition bias) All outcomes Selective reporting (re- porting bias) Insufficient information to permit judgement	sessment (detection bias)	Low risk	No patient-reported outcome
porting bias)	(attrition bias)	Unclear risk	Insufficient information to permit judgement
Other bias Unclear risk Insufficient information to permit judgement		Unclear risk	Insufficient information to permit judgement
<u> </u>	Other bias	Unclear risk	Insufficient information to permit judgement

Kouidi 1997

Koulai 1997		
Study characteristics	5	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 26 weeks 	
Participants	 Country: Greece Setting: single centre Inclusion criteria: undergoing HD Number: exercise group (24); control group (12) Mean age ± SD (years): exercise group (49.6 ± 12.1); control group (52.8 ± 10.2) Sex (M/F): exercise group (11/9); control group (4/7) Mean BMI ± SD (kg/m²): exercise group (25.7 ± 3.3); control group (24.7 ± 3.5) Mean HD vintage ± SD (years): exercise group (5.9 ± 4.9); control group (6.2 ± 5.4) Exclusion criteria: symptomatic cardiovascular disease; DM; musculoskeletal limitation or other ical problems contraindicating participation in an exercise training program 	
Interventions	Duration of intervention • 26 weeks Exercise group • Type: aerobic	



Kouidi 1997 (Continued)

- Description: stationary cycling, walking or jogging, callisthenics, aerobics, swimming and/or game sports
- Position: not applicable
- Material: not reported
- · Location: not reported
- Duration of training sessions: 90 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 to 4 times/week
- Timing in relation to dialysis treatments: on non-HD days
- Intensity: 50% to 60% of VO_2 max or 60% to 70% of max HR
- Supervised by: physician, exercise physiologist, trainer
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: increasing intensity
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care

Outcomes

- Potassium
- Sodium
- Calcium
- Phosphorus
- VO₂ max
- HR
- BP
- HRQoL
- Severity of depression
- · Traits of personality

Notes

· Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding



Kouidi 1997 (Continued)		
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Kouidi 2003

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 52 weeks
Participants	 Country: Greece Setting: not reported Inclusion criteria: HD patients Number: exercise group (15), control group (15) Mean age ± SD (years): exercise group (50.6 ± 10.8); control group (51.3 ± 9.9) Sex (M/F): not reported Exclusion criteria: other systemic disease; clinical symptoms of heart disease
Interventions	Duration of intervention • 52 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: not reported • Material: not reported • Location: HD unit • Duration of training sessions: not reported • Duration of warm-up/cool-down: not reported • Frequency: 3 times/week • Timing in relation to dialysis treatments: during • Intensity: not reported • Supervised by: supervised not further defined • Mode of delivery: not reported • Tailoring: not reported • Modifications/progression: not reported • Strategies to enhance adherence: not reported



Cambral and	
Control group	
Usual care	
• VO ₂ peak	
• SDNN	
• LVEF	
·	
•	
• TWA	
	cation; authors were contacted for full results
Funding: not reporte	ed
Authors' judgement	Support for judgement
Unclear risk	Insufficient information to permit judgement
Unclear risk	Insufficient information to permit judgement
High risk	No blinding
Unclear risk	Insufficient information to permit judgement
Low risk	No patient-reported outcome
Unclear risk	Insufficient information to permit judgement
Unclear risk	Insufficient information to permit judgement
Unclear risk	Insufficient information to permit judgement
	Usual care VO ₂ peak SDNN LVEF LF/HF ECG late potentials TWA Abstract-only public Funding: not report Authors' judgement Unclear risk Unclear risk Unclear risk Unclear risk Unclear risk Unclear risk Unclear risk

Kouidi 2004a

Study characteristic	S
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 26 weeks
Participants	Country: Greece



Kouidi 2004a (Continued)

- · Setting: not reported
- Inclusion criteria: sedentary HD patients
- Number: exercise group (11); control group (10)
- Age range: 60 to 72 years
- Exclusion criteria: severe cardiovascular abnormalities; DM; active hepatitis

Interventions

Duration of intervention

• 26 weeks

Exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: not reported
- · Location: HD unit
- · Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- · Intensity: not reported
- · Supervised by: supervised not further defined
- Mode of delivery: not reported
- · Tailoring: not reported
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

Usual care

Outcomes

- VO₂ peak
- Peak torque
- · Ejection fraction
- Cardiac output index
- Transmittal flow
- Isovolemic relaxation time

Notes

- Abstract-only publication: authors were contacted, but full results could not me obtained
- · Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement



Kouidi 2004a (Continued)		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Kouidi 2005

ouldi 2005			
Study characteristics	;		
Methods	Study design: parallel RCT		
	Study duration: not reported		
	Study follow-up period: 10 months		
Participants	Country: Greece		
	Setting: not reported		
	 Inclusion criteria: undergoing HD treatments 		
	Number: exercise group (19); control group (14)		
	 Mean age ± SD: 48.8 ± 13.9 years 		
	• Sex (M/F): 27/6		
	Exclusion criteria: not reported		
Interventions	Duration of intervention		
	• 43.4 weeks		
	Exercise group		
	Type: aerobic		
	Description: stationary cycling		
	Position: not reported		
	Material: not reported		
	Location: HD unit		
	 Duration of training sessions: not reported 		
	 Duration of warm-up/cool-down: not reported 		
	Frequency: 3 times/week		
	 Timing in relation to dialysis treatments: during 		
	Intensity: not reported		



Kouidi 2005 (Continued)

- Supervised by: supervised not further defined
- Mode of delivery: not reported
- Tailoring: not reported
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: EPO

Control group

EPO

Outcomes

- VO₂ peak
- · Depression scores
- Ool
- Personality traits

Notes

- Abstract-only publication: authors were contacted, but full results could not me obtained
- · Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement



Kouidi 2008				
Study characteristics				
Methods	 Study design: parallel RCT Study duration: not reported 			
	Study follow-up period: 10 months			
Participants	Country: GreeceSetting: multicentre			
	 Inclusion criteria: undergoing HD treatments for at least 6 months 			
	 Number: exercise group (32); control group (31) 			
	 Mean age ± SD (years): exercise group (55 ± 9); control group (53 ± 6) 			
	 Sex (M/F): exercise group (18/12); control group (16/13) 			
	 HD vintage (mean ± SD years): exercise group (6.3 ± 3.7); control group (6.2 ± 3.9) 			
	 Exclusion criteria: bundle branch block; unstable hypertension; DM; severe congestive heart failure; recent MI; unstable angina 			
Interventions	Duration of intervention			
	• 43.4 weeks			
	Exercise group			
	Type: combined			
	 Description: stationary cycling + abdominal and lower limbs strength and flexibility exercises 			
	Position: seated			
	Material: ergometer, weights and elastic bands			
	Location: HD unit			
	Duration of training sessions: 90 minutes			
	 Duration of warm-up/cool-down: 10/10 minutes 			
	Frequency: 3 times/week			
	Timing in relation to dialysis treatments: during			
	Intensity: 60% to 70% of max HR			
	Supervised by: exercise trainers, physician			
	Mode of delivery: face-to-face			
	Tailoring: individualised intensity			
	Modifications/progression: increasing repetitions, duration, weights and resistance			
	Strategies to enhance adherence: not reported Adherence to intervention, not reported.			
	Adherence to intervention: not reported Co. intervention: none			
	Co-intervention: none Control group			
	Usual care			
Outcomes	Hb (g/dL)			
- accomes	Urea (mg/dL)			
	SCr (mg/dL)			
	Potassium (mEq/L)			
	Sodium (mEq/L)			
	Calcium (mEq/L)			
	Phosphorus (mg/dL)			
	Peak oxygen consumption			
	• LVMI			

• SD of the normal RR intervals



Kouidi 2008 (Continued)

- Mean RR interval
- LVEF
- Mean 24-hour HR
- LF/HF ratio
- Signal-averaged ECG
- Adherence with the exercise program
- Resting SBP and DBP

Notes

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Drawing of lots
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Kouidi 2010

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 52 weeks
Participants	 Country: Greece Setting: HD unit Inclusion criteria: underwent HD 3 times/week for 4 hours for at least 6 months Number: exercise group (25); control group (25)



Kouidi 2010 (Continued)

- Mean age ± SD (years): exercise group (46.3 ± 11.2); control group (45.8 ± 10.9)
- Sex (M/F): exercise group (14/10); control group (12/8)
- Mean HD vintage ± SD (years): exercise group (6.1 ± 4.6); control group (6.3 ± 4.9)
- Exclusion criteria: no history, clinical signs, or symptoms of psychiatric, neurological, cardiologic, or pulmonary disorders; absence of DM; no significant electrolytic instability or undisciplined patients; no musculoskeletal limitation or other medical problems contraindicating participation in an ET program

Interventions

Duration of intervention

• 52 weeks

Exercise group

- · Type: combined
- Description: stationary cycling and resistance exercises of the lower limbs
- · Position: not reported
- Material: ergometer, free weights and resistance bands
- · Location: HD unit
- Duration of training sessions: 60 to 90 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 70% of VO₂ max
- · Supervised by: physician and exercise trainer
- · Mode of delivery: face-to-face
- · Tailoring: individualised intensity and duration
- Modifications/progression: increasing duration and workload
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

Usual care

Outcomes

- Hb
- Calcium (mEq/L)
- Phosphorous (mg/dL)
- Potassium (mEq/L)
- LF/HF
- · Mean square successive difference (ms)
- pNN50 (ms)
- SDNN (ms)
- Urea (mg/dL)
- SCr (mg/dL)
- Sodium (mEq/L)
- VO₂ peak (ml/kg/min)
- · Hospital anxiety and depression scale
- Depression (BDI)

Notes

• Funding: not reported



Kouidi 2010 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Lee 2001

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Korea Setting: not reported Inclusion criteria: prevalent HD patients receiving dialysis 2 to 3 times/week Number: exercise group (25); control group (21) Mean age ± SD (years): exercise group (45 ± 12.8); control group (53.1 ± 14.2) Sex (M/F): exercise group (12/13); control group (9/12) Mean HD vintage ± SD (months): exercise group (37 ± 34.9); control group (41.7 ± 30.9) Exclusion criteria: not reported
Interventions	Duration of intervention 12 weeks Exercise group Type: aerobic Description: stationary cycling and walking



Lee 2001 (Continued)

- · Position: not reported
- Material: ergometer, treadmill
- · Location: HD
- Duration of training sessions: 5 to 30 minutes
- Duration of warm-up/cool-down: 5 to 10/not reported minutes
- Frequency: 2 to 4 times/week
- Timing in relation to dialysis treatments: just prior
- Intensity: 12 to 14 on RPE
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increasing intensity and duration
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- Serum lipids
- . Uh
- Physical work capacity (measured in the intervention group only)
- Physical fitness (measured in the intervention group only)

Notes

· Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	High risk	One or more outcomes of interest in the review are reported incompletely so that they cannot be entered in a meta-analysis



Lee 2001 (Continued)

Other bias Unclear risk Insufficient information to permit judgement

Study characteristic	s
Methods	Study design: parallel RCTStudy duration: not reported
	Study follow-up period: 12 weeks
Participants	Country: Taiwan
	Setting: HD unit
	 Inclusion criteria: receiving maintenance HD for at least 6 months with 3 times/week and 4 hours for each session
	Number: exercise group (20); control group (20)
	 Mean age ± SD (years): exercise group (62.0 ± 8.0); control group (62.0 ± 9.0)
	 Sex (M/F): exercise group (8/12); control group (9/11)
	• Mean BMI \pm SD (kg/m ²): exercise group (22.9 \pm 3.3); control group (23.7 \pm 4.2)
	 Mean HD vintage ± SD (months): exercise group (71.0 ± 46.0); control group (83.0 ± 71.0)
	 Exclusion criteria: presence of active infection or inflammation, autoimmune disorders, malignancy, psychiatric diseases, severe musculoskeletal disorders, poorly controlled DM or secondary hyperparathyroidism; uncontrolled heart failure or pulmonary diseases; hospitalisation during the previous month; use of drugs that influence serum cytokine levels; vascular access in the lower extremities; BMI > 25 kg/m²

Interventions

Duration of intervention

• 12 weeks

Exercise group

- Type: aerobic
- Description: stationary cycling
- Position: recumbent
- Material: ergometer
- Location: HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 12 to 15 on RPE
- · Supervised by: physician and rehabilitation nurse
- Mode of delivery: face-to-face
- Tailoring: individualised intensity and duration
- Modifications/progression: increasing duration
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care



Liao 2016 (Continued)

Outcomes

- · Pre HD SBP
- Pre HD DBP
- Pre HD HR
- iPTH (pg/mL)
- Calcium (mg/dL)
- tHcy (mol/L)
- hs-CRP (mg/dL)
- IL-6 (pg/mL)
- SCr (mg/dL)
- Albumin (g/dL)
- ALT (mu/L)
- Cholesterol (mg/dL)
- HCT (%)
- KT/V
- nPCR (g/kg/day)
- Mean BP (mm Hg)
- Weight (kg)
- BMI (kg/m²)
- Number of endothelial progenitor cells
- BMD femoral neck
- BMD spine
- 6MWT

Notes

Funding

- Research Fund of the Cardinal Tien Hospital
- TaoYuan Army Hospital
- Tri-Service General Hospital
- Ministry of Science and Technology

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias)	Low risk	No missing outcome data



Liao 2016 (Continued)

All outcomes

Selective reporting (reporting bias)	High risk	Not all of the study's pre-specified outcomes have been reported
Other bias	Unclear risk	Insufficient information to permit judgement

Ma 2018

 Study duration: not reported Study follow-up period: 104 weeks Country: China Setting: HD unit Inclusion criteria: maintenance HD > 3 months; aged 18 to 70 years; dialysis 3/week; Kt/V > 1.2 Number: total (132) Mean age ± SD: 55.2 ± 12.2 years Sex (M/F): 79/53 Median HD vintage (IQR): 44 months (2, 254) Exclusion criteria: cardiac function NYHA class IV; severe osteoarthrosis; walking distance < 200 m quiet condition of blood oxygen saturation < 90%; patients with limbs missing who cannot exercise 	Study characteristics	
Setting: HD unit Inclusion criteria: maintenance HD > 3 months; aged 18 to 70 years; dialysis 3/week; Kt/V > 1.2 Number: total (132) Mean age ± SD: 55.2 ± 12.2 years Sex (M/F): 79/53 Median HD vintage (IQR): 44 months (2, 254) Exclusion criteria: cardiac function NYHA class IV; severe osteoarthrosis; walking distance < 200 m quiet condition of blood oxygen saturation < 90%; patients with limbs missing who cannot exercise interventions Duration of intervention 104 weeks Exercise group Type: combined Description: aerobics, resistance, and flexibility training not further defined Position: not reported Material: not reported Location: HD unit Duration of training sessions: 20 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: not reported Supervised by: not reported Mode of delivery: not reported Mode of delivery: not reported Mode of delivery: not reported Modifications/progression: not reported Modefications/progression: not reported Adherence to intervention: not reported Co-intervention: none Control group	Methods	Study duration: not reported
 104 weeks Exercise group Type: combined Description: aerobics, resistance, and flexibility training not further defined Position: not reported Material: not reported Location: HD unit Duration of training sessions: 20 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: not reported Supervised by: not reported Mode of delivery: not reported Tailoring: not reported Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none 	Participants	 Setting: HD unit Inclusion criteria: maintenance HD > 3 months; aged 18 to 70 years; dialysis 3/week; Kt/V > 1.2 Number: total (132) Mean age ± SD: 55.2 ± 12.2 years Sex (M/F): 79/53 Median HD vintage (IQR): 44 months (2, 254) Exclusion criteria: cardiac function NYHA class IV; severe osteoarthrosis; walking distance < 200 m
Usual care	Interventions	 • 104 weeks Exercise group • Type: combined • Description: aerobics, resistance, and flexibility training not further defined • Position: not reported • Material: not reported • Location: HD unit • Duration of training sessions: 20 minutes • Duration of warm-up/cool-down: not reported • Frequency: 3 times/week • Timing in relation to dialysis treatments: during • Intensity: not reported • Supervised by: not reported • Mode of delivery: not reported • Modifications/progression: not reported • Strategies to enhance adherence: not reported • Adherence to intervention: note • Co-intervention: none
		Usual care

Outcomes

• Cardiopulmonary endurance index



Ma 2018 (Continued)	М	a 2	201	8	(Continued)
---------------------	---	-----	-----	---	-------------

- Hb
- Albumin
- Total cholesterol
- 6MWT
- · Anxiety score
- Depression score

Notes

- Abstract-only publication: authors contacted for full results
- Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Makhlough 2012

>	τu	a	V	CI	70	ır	a	C	τ	21	IS	U	C	S

Study characteristics	
Methods	 Study design: parallel RCT Study duration: 2 months Study follow-up period: 8 weeks
Participants	 Country: Iran Setting: HD unit Inclusion criteria: HD > 3 months Number: exercise group (25); control group (23)



Makhlough 2012 (Continued)

- Mean age ± SD (years): exercise group (53.3 ± 14.3); control group (56.2 ± 10.8)
- Sex (M/F): exercise group (18/7); control group (12/11)
- Mean HD vintage ± SD (months): exercise group (25.5 ± 10.7); control group (23.5 ± 13.6)
- Exclusion criteria: poorly controlled hypertension; uncompensated heart failure; cardiac arrhythmia
 requiring treatment; recent unstable angina; persistent hyperkalaemia before dialysis; significant
 valvular heart disease; MI within the past 6 months; significant cerebral or peripheral arteriosclerosis;
 bone disease with a risk of fracture; orthopaedic or musculoskeletal limitations; weight gains > 4 kg
 from Friday to Monday or from Saturday to Tuesday; recent significant change in the resting ECG; thirddegree atrioventricular heart block without pacemaker; severe aortic stenosis; suspected or known
 dissecting aneurysm; active or suspected myocarditis or pericarditis; thrombophlebitis or intracardiac thrombi; recent systemic or pulmonary embolus; acute infections

Interventions

Duration of intervention

8 weeks

Exercise group

- Type: range of motion
- Description: rotating the wrist, wrist up and down, ankle-twisting motion
- · Position: not reported
- · Material: not reported
- · Location: HD unit
- · Duration of training sessions: 15 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: not reported
- · Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: individualised duration
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- Serum phosphate (mg/dL)
- Serum calcium (mg/dL)
- · Serum potassium (mg/dL)
- Hb (g/dL)

Notes

- The total number of participants and the number per group do not add up. Percentages of men and women are not consistent with the number of participants per group
- Funding: Mazandaran University of Medical Sciences

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator



Makhlough 2012 (Continued)		
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Multiple errors and discrepancies in the reporting of the study

Marchesan 2016

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 24 weeks
Participants	 Country: Brazil Setting: HD unit Inclusion criteria: HD for at least 3 months Number: exercise group (8); control group (7) Mean age ± SD (years): exercise group (63 ± 4); control group (65 ± 5) Sex (M/F): exercise group (6/2); control group (5/2) Exclusion criteria: having a central catheter; < 60 years; severe heart disease, respiratory problems, and not having been released by the physician to participate in the program due to unstable clinical conditions; unstable medical conditions encompassed biochemical aspects; weight gain on the opposite dialysis day; uncontrolled anaemia; complications and hospitalisations in the last 6 months
Interventions	 Duration of intervention 24 weeks Exercise group Type: combined Description: stationary cycling and resistance exercises for upper and lower limbs, thorax, abdomen and the posterior region of the trunk Position: seated Material: ergometer and step



Marchesan 2016 (Continued)

- · Location: HD unit
- Duration of training sessions: 15 to 45 + resistance minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 60% to 70% of max HR and 3 to 4 on RPE (1 to 10)
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: increasing intensity and duration
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- 6MWT
- Sit-to-stand test (30 sec)
- Respiratory muscle strength test

Notes

• Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement



Marinho 2016

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 8 weeks
Participants	 Country: France Setting: HD unit Inclusion criteria: ≥ 18 years; treatment by maintenance dialysis for at least 3 months Number: exercise group (7); control group (7) Median age, IQR (years): exercise group (71.5, 58.5 to 87.2); control group (76.0, 59.0 to 83.0) Sex (M/F): exercise group (3/3); control group (3/4) Median BMI, IQR (kg/m²): exercise group (28.50, 21.1 to 35.8); control group (28.40, 20.8 to 35.2) Exclusion criteria: cancer; AIDS; autoimmune disease; taking catabolizing drugs or vitamin D recepto activator
Interventions	Duration of intervention8 weeksExercise groupType: resistance
	 Description: lower limbs exercises Position: not reported Material: resistance bands and ankle cuffs Location: HD unit Duration of training sessions: varied minutes Duration of warm-up/cool-down: not reported
	 Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 60% to 70% of 3-RM Supervised by: physical educator Mode of delivery: face-to-face
	 Tailoring: individualised intensity Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none
	Control group • Usual care
Outcomes	 CRP (mg/L) Calcium (mmol/L) Phosphate (m/dL) Potassium (mmol/L) SCr (mg/dL) Hb (g/dL) Lean mass (%)
	 BMI (kg/m²) Body fat (%)



Marinho 2016 (Continued)

- Sclerostin (ng/mL)
- BAP (U/L)
- BAP/PTH
- Leptin (ng/mL)
- PTH (pg/mL)
- 25 (OH) vitD (ng/mL)
- 1,25 (OH)2 vitD (pg/mL)

Notes

• Funding: Coordination of Improvement of Superior Education Personnel (CAPES)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Martin-Alemany 2016

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Mexico Setting: HD unit Inclusion criteria: regular HD 2 times/week; signed informed consent; any gender; > 18 years; no physical activity



Martin-Alemany 2016 (Continued)

- Number: exercise group (22); control group (22)
- Median age, IQR (years): exercise group (35, 24 to 41.5); control group (30, 24 to 47)
- Sex (M/F): exercise group (10/7); control group (11/8)
- Median BMI, IQR (kg/m²): exercise group (20.4, 19.4 to 23); control group (21, 18.3 to 22.1)
- Exclusion criteria: amputation; hospitalisation in the last 3 months; unsatisfactory attendance at HD
 sessions; pregnancy; excessive pallor; severe dyspnoea; femoral fistula; arrhythmias; precordial pain;
 orthopaedic or neurological compromises or cognitive alterations affecting their participation; intolerance to ONS; intolerance/contraindications to the exercise routine; infectious or cardiovascular
 complications during the study

Interventions

Duration of intervention

• 12 weeks

Exercise group

- Type: resistance
- Description: upper and lower limbs exercises
- · Position: seated and semi-recumbent
- Material: ankle weights and resistance springs
- Location: HD unit
- Duration of training sessions: 4 x 10 min with 3 min rest in between minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 2 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 13 on RPE
- · Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: oral nutritional supplementation

Control group

• Oral nutritional supplement

Outcomes

- · Weight
- BMI (kg/m²)
- Mid-arm circumference
- Arm muscle circumference
- Arm muscle area
- Triceps skinfold thickness (mm)
- FM% from anthropometry
- · Handgrip strength
- Resistance at 50 kHz
- Reactance at 50 kHz
- Phase angle (°)
- Hb (g/dL)
- Total lymphocyte count (cells/mm³)
- SCr (mg/dL)
- Albumin (g/dL)
- Phosphorus (mg/dL)
- Potassium (mmol/dL)



Martin-Alemany 2016 (Continued)

QoL

Notes • Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Martins do Valle 2020

Studv	cha	racto	rictica	
Stuav	cna	racte	ristics	5

Methods	

- Study design: parallel RCT
- Study duration: not reported
- Study follow-up period: 12 weeks

Participants

- Country: Brazil
- · Setting: HD unit
- Inclusion criteria: adult ESKD patients who were under chronic HD treatment, 3 times/week totalling 12 hours weekly, for at least 3 months
- Number: exercise group (12); control group (12)
- Mean age ± SD (years): exercise group (49.3 ± 12.4); control group (60.4 ± 10.6)
- Sex (M/F): exercise group (5/7); control group (8/4)
- Mean BMI \pm SD (kg/m²): exercise group (22.7 \pm 3.8); control group (23.2 \pm 5.1)
- Median HD vintage, IQR (years): exercise group (6.8, 11.4); control group (3.9, 12.5)
- Exclusion criteria: any limitation that prevents the physical tests; presence of severe and unstable comorbidities or hospitalisation in the 3 months prior to inclusion in the study (unstable angina; decom-



Martins do Valle 2020 (Continued)

pensated heart failure; MI in the last 6 months; uncontrolled arrhythmia; uncontrolled hypertension with SBP 200mm Hg and/or DBP 120mm Hg; uncontrolled DM; severe pneumopathies; acute systemic infection; neurological, musculoskeletal and disabling osteoarticular disturbances; or other conditions according to clinical judgment)

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: resistance
- Description: lower and upper limbs exercises
- · Position: seated or supine
- Material: ankle weights and dumbbells
- · Location: HD unit
- Duration of training sessions: varied minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 3 to 5 on RPE
- Supervised by: supervised not further defined
- Mode of delivery: not reported
- · Tailoring: individualised intensity
- · Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- Adherence to intervention (sessions attended): 80%
- · Co-intervention: none

Control group

· Stretching exercises

Outcomes

- Adherence
- Time spent in activities of daily living
- Physical activity in daily life (steps/day)
- 6MWT distance (m)
- Maximum voluntary isometric contraction (Kgf)
- QoL
- Hb (g/dL)
- Serum iron (µg/dL)
- Ferritin (ng/mL)
- Transferrin saturation index
- Adequacy of dialysis (Kt/V)
- · Albumin (g/dL)
- Sodium (mEq/L)
- Calcium (mg/dL)
- Potassium (mEq/L)
- Phosphorous (mg/dL)
- PTH (pg/mL)

Notes

Funding

- Fundação de Amparoa Pesquisa do Estado de Minas Gerais FAPEMIG
- Coordenacao de Aperfeicoamento de Pessoal de Nivel Superior Brasil (CAPES)



Martins do Valle 2020 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Sealed opaque envelopes
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	Study appears free of other biases

Matsumoto 2007

Study characteristic	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Japan Setting: HD unit Inclusion criteria: ≥ 30 years; HD 3 times/week (4 hours/dialysis) > 3 years Number: exercise group (22); control group (33) Mean age ± SD (years): exercise group (61 ± 10); control group (57 ± 8) Sex (M/F): exercise group (5/12); control group (15/17) Mean HD vintage ± SD (years): exercise group (12 ± 7); control group (13 ± 8) Exclusion criteria: chronic lung disease; current ischaemic heart disease; uncontrolled arrhythmias o hypertension; haemodynamic instability; inability to pedal a stationary cycle; Hb < 85 mmol/L; albu min > 40 mg/dL
Interventions	Duration of intervention

• 52 weeks



Matsumoto 2007 (Continued)

Exercise group

- Type: aerobic
- Description: stationary cycling
- Position: not reported
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: just before
- Intensity: 60% to 70% of max HR
- · Supervised by: study staff
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

Usual care

Outcomes

- Albumin
- HRQoL
- Creatinine generation rate

Notes

- Data extracted from figures
- Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias)	Low risk	Missing outcome data balanced in numbers across intervention groups



Matsumoto 2007 (Continued)

All outcomes

Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

McAdams-DeMarco 20 Study characteristics	
Methods	 Study design: parallel RCT (3 arms*) Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: USA Setting: HD unit Inclusion criteria: undergoing maintenance HD; ≥ 18 years; English speaking; able to provide informed consent Number: exercise group (6); control group (7) Mean age ± SD (years): exercise group (48.0 ± 7.0); control group (55.0 ± 9.7) Sex (M/F): exercise group (4/2); control group (7/0) Mean BMI ± SD (kg/m²): exercise group (32.0 ± 10.1); control group (30.4 ± 6.9) Exclusion criteria: angina pectoris; chronic lung disease; cerebral vascular disease; musculoskeletal or orthopaedic conditions limiting physical activity; lower or upper extremity amputation; decreased mental capacity; diagnosed dementia
Interventions	Duration of intervention • 12 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: seated • Material: ergometer

- Location: HD unit
- Duration of training sessions: ad tolerance minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: not reported
- Supervised by: research assistants
- Mode of delivery: face-to-face
- Tailoring: no
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- · Co-intervention: none

Control group



McAdams-DeMarco 2018 (Continued)

• Usual care

Outcomes

- Modified Mini Mental Status
- Trail Making Test A time
- Trail Making Test A-Trail Making Test B time (sec)
- Trail Making Test B time (sec)

Notes

- * Cognitive training group not included in this review
- Funding
 - o Johns Hopkins Faculty Innovation Fund
 - o National Institutes of Health
 - o Johns Hopkins Bloomberg School of Public Health Faculty Innovation Fund
 - o American Society of Nephrology Carl W. Gottschalk Research Scholar Grant
 - o Johns Hopkins University Claude D. Pepper Older Americans Independence Center
 - o National Institute on Aging

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	High risk	Not all of the study's pre-specified outcomes have been reported
Other bias	Low risk	The study appears to be free of other sources of bias

McGregor 2018

Study	chare	ectori	ictics

Methods

- Study design: parallel RCT (3 arms*)
- Study duration: 18 months



McGregor 2018 (Continued)

• Study follow-up period: 10 weeks

Participants

- · Country: UK
- · Setting: in-centre and satellite HD units
- Inclusion criteria: > 18 years, dialysis 3 time/week for 3 to 4 hours; dialysis vintage of > 3 months; URR
 > 65%; ability to complete dynamic exercise testing and training
- Number: exercise group (16); control group (18)
- Mean age (95% CI) (years): exercise group (52.1 (44.2; 59.9)); control group (54.3 (46.0; 62.5))
- Sex (M/F): exercise group (13/3); control group (11/7)
- Mean dialysis vintage (95% CI) (months): exercise group (48.1 (26.2; 70.0)); control group (49.3 (29.6; 69.0))
- Mean BMI (95% CI) (kg/m²): exercise group (29.2 (25.2; 33.2)); control group (27.5 (24.6; 30.37))
- Exclusion criteria: active malignant disease; ischaemic cardiac event (<3 months); significant valvular heart disease or dysrhythmia; planned kidney transplant during the study period; life expectancy of <6 months

Interventions

Duration of intervention

• 10 weeks

- Exercise groupType: aerobic
- · Description: stationary cycling
- · Position: semi-recumbent
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 50 to 60 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 40% to 60% of VO₂ reserve and 12 to 14 on RPE
- · Supervised by: clinical exercise physiologists
- · Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- · Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- HR rest (bpm)
- · HR peak (bpm)
- VO₂ AT (mL/kg/min)
- VO₂ peak (mL/kg/min)
- Respiratory exchange rate at VO₂ AT
- · Respiratory exchange rate at VO₂ peak
- Max. load (Watts)
- Leg strength (Newtons)
- LVMI (g/m²)
- LVED volume index (mL/m2)



McGregor 2018 (Continued)

- LV end diastolic volume index (mL/m²)
- LVEF (%)
- E/A ratio
- Mean E/e'
- Left atrium diameter (cm)
- SBP rest (mm Hg)
- DBP Rest (mm Hg)
- Pulse wave velocity
- Flow-mediated dilatation Delta (cm)
- Flow-mediated dilatation Delta (%)

Notes

- *Low-frequency electrical muscle stimulation group was not included in this review
- Funding: West Midlands Comprehensive Local Research Network

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Permuted stratified block randomisation. Assumed computer-generated.
Allocation concealment (selection bias)	Low risk	Performed independently by the trial statistician
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	Study appears free of other biases

Mitsiou 2015

Study chard	acteristics
-------------	-------------

Methods

- Study design: factorial RCT (4 arms)
- Study duration: not reported
- Study follow-up period: not reported



Mitsiou 2015 (Continued)

Pа	rti	cii	าล	nts

- · Country: Greece
- · Setting: not reported
- Inclusion criteria: HD patients free of other systemic disease
- Number: joint music + exercise training group (10); sole music program group (10); sole exercise training group (10); control group (10)
- Mean age \pm SD: 50 \pm 14.7 years
- Sex (M/F): not reported
- · Exclusion criteria: not reported

Interventions

Duration of intervention

• 26 weeks

Exercise group

- · Type: not reported
- · Description: not reported
- · Position: not reported
- Material: not reported
- · Location: not reported
- Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- · Frequency: not reported
- · Timing in relation to dialysis treatments: during
- · Intensity: not reported
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: not reported
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Music and usual care (factorial RCT)

Outcomes

- 6MWT
- Mean HR
- SD of NN intervals
- Root mean square of successive differences
- Components of the autoregressive power spectrum of the NN intervals $\,$

Notes

- Abstract-only publication: authors contacted for full results
- · Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement



Mitsiou 2015 (Continued)		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Miura 2015

Study characteristics	s
Methods	 Study design: parallel RCT (3 arms) Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Japan Setting: not reported Inclusion criteria: ESKD patients Number: exercise group (19); control group (10); electrical stimulation (6) Mean age ± SD: 70.2 ± 11.7 years Sex (M/F): 20/15 Exclusion criteria: not reported
Interventions	Duration of intervention 12 weeks Exercise group
	 Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: not reported Duration of training sessions: 60 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 11 to 13 on RPE



Miura 2015 (Continued)

- Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: not reported
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- Grip strength
- Quad muscle torque
- Workout time
- Activities
- · Dialysis efficacy
- HDL-cholesterol
- LDL-cholesterol
- CRP
- IL-6
- BP

Notes

- Abstract-only publication: authors contacted for full results
- · Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Unclear risk	Insufficient information to permit judgement
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement



Miura 2015 (Continued)

Other bias Unclear risk Insufficient information to permit judgement

Molsted 2004

Study characteristics	3
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 5 months
Participants	 Country: Denmark Setting: HD unit Inclusion criteria: > 18 years old; undergoing HD > 3 months Number: exercise group (22); control group (11) Median age, range (years): exercise group (59, 25 to 58); control group (48, 23 to 58) Sex (M/F): exercise group (14/8); control group (8/3) Mean dialysis vintage (years): exercise group (2); control group (1.5) Exclusion criteria: DM; symptomatic heart disease; orthopaedic limitations; severe peripheral polyneuropathy; dementia; participation in other studies with the risk of affecting the results; inability to speak either Danish or English; patients able to speak English were only excluded from the questionnaire
Interventions	Duration of intervention • 21.7 weeks
	Exercise group
	 Type: combined Description: step and circuit training, high and low impact aerobics and stationary cycling Position: not reported Material: ergometer, step Location: not reported Duration of training sessions: 50 minutes Duration of warm-up/cool-down: 10/10 minutes Frequency: 2 times/week Timing in relation to dialysis treatments: not reported Intensity: 14 to 17 on RPE Supervised by: physiotherapist Mode of delivery: face-to-face Tailoring: individualised intensity Modifications/progression: not reported Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none
	Control group
	Usual care
Outcomes	 HRQoL Physical functioning VO₂ max



Molsted 2004 (Continued)

- BP
- Lipids

Notes

Funding

- Roche A/S
- Janssen-Cilag A/S
- The Association of Danish Physiotherapists Research Foundation
- Danish Kidney Association
- Danish Society of Nephrology, Copenhagen Hospital Corporation
- Chr. Andersen and Ingeborg Andersen of the Schmidt Foundation
- Anna & Jakob Jakobsen's Foundation

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Low risk	Envelops
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	High risk	Private funding. Funder's involvement not specified

Momeni 2014

Study characteristics	
Methods	 Study design: parallel RCT Study duration: 8 months Study follow-up period: 12 weeks
Participants	Country: IranSetting: HD unit



Momeni 2014 (Continued)

- Inclusion criteria: > 18 years; dialysis duration > 3 months
- Number: total (40)
- Mean age ± SD: 43.1 ± 10.5 years
- Sex (M/F): 30/10
- Exclusion criteria: > 60 years; history of ischaemic heart disease; use of anti-arrhythmic agents; LVEF
 < 40% on ECG; inability of doing Intradialysis exercise; dyspnoea or chest pain during exercise; BP ≥
 160/100 mm Hg before exercise program

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: mini bike
- · Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: not reported
- · Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: not reported
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

Usual care

Outcomes

- HCT
- Hb
- Serum calcium (mg/dL)
- Serum phosphorus (mg/dL)
- · Serum potassium (mg/dL)
- E/A ratio
- Left atrial size (cm)
- Left ventricular end-diastolic diameter (cm)
- Left ventricular end-systolic diameter (cm)
- LVEF (%)
- Mitral valve maximum pressure
- Mitral valve minimum pressure gradient
- Mitral valve velocity time integral
- Right ventricular size (cm)
- Systolic pulmonary artery pressure (mm Hg)
- Blood urea nitrogen (mg/dL)
- SCr (mg/dL)
- · LVH severity



Momeni 2014 (Continued)

- · Diastolic dysfunction severity
- · Presence of pericardial effusion

Notes

• Funding: Shahrekord University of Medical Sciences

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Mortazavi 2013

Study characteristics

Methods

- Study design: parallel RCT
- Study duration: not reported
- Study follow-up period: 16 weeks

Participants

- Country: Iran
- · Setting: HD unit
- Inclusion criteria: HD for at least 3 months; dialysis at least 3 times/week; presence of RLS; ferritin > 100 ng/mL; transferrin saturation rate > 20%
- Number: exercise group (13); control group (13)
- Mean age \pm SD (years): exercise group (32.3 \pm 6.7); control group (47.1 \pm 13.1)
- Sex (M/F): 18/8



Mortazavi 2013 (Continued)

Exclusion criteria: musculoskeletal disorders which incapacitated them from physical activity; history
of ischaemic heart disease (recent MI or unstable angina); any catabolic process such as malignancies
opportunistic infections, and infections needing antibiotic therapy during the last 3 months

Interventions

Duration of intervention

• 16 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- Position: not reported
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 10 to 12 on RPE
- · Supervised by: not reported
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- · RLS questionnaire
- QoL

Notes

· Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes



Mortazavi 2013 (Continued)		
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Olvera-Soto 2016

 Study design: parallel RCT Study duration: 6 months Study follow-up period: 12 weeks
 Country: Mexico Setting: HD unit Inclusion criteria: > 18 years; on HD for at least 3 months, residents of Mexico City Number: exercise group (30); control group (31) Median age, IQR (years): exercise group (28.5, 23 to 46.5); control group (29, 19 to 38) Sex (M/F): exercise group (14/16); control group (19/12) Median HD vintage, IQR (years): exercise group (12, 5.75 to 37.75); control group (18, 8 to 39) Mean BMI ± SD (kg/m²): exercise group (21.8 ± 3.1); control group (21.1 ± 2.7) Exclusion criteria: not reported
Duration of intervention 12 weeks Exercise group Type: resistance Description: upper and lower limbs exercises Position: seated Material: weights and resistance bands Location: HD unit Duration of training sessions: 50 minutes Duration of warm-up/cool-down: not reported Frequency: 2 times/week Timing in relation to dialysis treatments: during Intensity: not reported Supervised by: supervised not further defined Mode of delivery: face-to-face Tailoring: not reported Modifications/progression: weights added on 3rd sessions



0	lvera-	Soto	2016	(Continued
---	--------	------	------	------------

• Co-intervention: none

Control group

Usual care

Outcomes

- Body fat (%)
- Dietary energy intake
- · Dietary protein intake
- Arm muscle circumference (mm)
- Arm muscular area
- Handgrip strength (kg)

Notes

• Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	High risk	The study report fails to include results for key outcomes that would be expected to have been reported for such a study
Other bias	Unclear risk	Insufficient information to permit judgement

Ouzouni 2009

Study characteristics			
Methods	 Study design: parallel RCT Study duration: 6 months Study follow-up period: 43 weeks 		
Participants	Country: Greece		



Ouzouni 2009 (Continued)

- · Setting: not reported
- Inclusion criteria: on maintenance HD 3 days/ week, 4 hours/session for at least 6 months prior to the study
- Number (randomised/analysed): exercise group (20/19); control group (15/14)
- Mean age ± SD (years): exercise group (47 ± 16); control group (51 ± 12)
- Sex (M/F): exercise group (14/5); control group (13/1)
- HD vintage ± SD (years): exercise group (7.7 ± 7.0); control group (8.6 ± 6.0)
- Exclusion criteria: unstable hypertension; heart failure (NYHA class > II); cardiac arrhythmias (> III according to Lown); recent MI or unstable angina; DM; active liver disease or orthopaedic problems limiting exercise

Interventions

Duration of intervention

• 43.4 weeks

Exercise group

- · Type: combined
- · Description: stationary cycling + abdominal and lower limbs exercises
- · Position: not reported
- Material: ergometer, weights and resistance bands
- · Location: HD unit
- Duration of training sessions: 20 min aerobic + varied for resistance minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 13 to 14 on RPE (6 to 20)
- · Supervised by: physician and exercise physiologist
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- · Modifications/progression: increasing duration, number of repetitions and weights added
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

Usual care

Outcomes

- HRQoL
- VO₂ peak
- Exercise time
- · Metabolic equivalents
- HR maximum
- BF
- Depression
- Double product
- Maximum pulmonary ventilation

Notes

Funding: not reported

Risk of bias

Bias Authors' judgement Support for judgement



Ouzouni 2009 (Continued)		
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Painter 2002a

Study characteristic	s
Methods	 Study design: factorial RCT (4 arms); stratified by age (< 50 years versus ≥ 50 years) Study duration: not reported Study follow-up period: 5 months
Participants	 Country: USA Setting: 5 HD units Inclusion criteria: ≥ 18 years; treated with HD for at least 3 months; mean HCT of 30% ± 3% for 4 weeks before study enrolment Exercise group 1: usual care HCT (30% to 33%) + exercise Exercise group 2: normalised HCT (40% to 42%) + exercise Number: exercise group 1 (10); exercise group 2 (12); control group 1 (14); control group 2 (12) Mean age ± SD (years): exercise group 1 (47.6 ± 11.9); exercise group 2 (43.5 ± 10.5); control group 1 (43.3 ± 9.8); control group 2 (50.1 ± 13.8) Sex (M/F): exercise group 1 (5/5); exercise group 2 (5/7); control group 1 (6/8); control group 2 (5/7) Mean HD vintage ± SD (months): exercise group 1 (23.1 ± 24.6); exercise group 2 (60.4 ± 80.0); control group 1 (61.8 ± 72.9); control group 2 (67.8 ± 54.4) Exclusion criteria: unstable hypertension; heart failure (NYHA class > II); cardiac arrhythmias (> III according to Lown); recent MI or unstable angina; DM; active liver disease or orthopaedic problems limiting exercise
Interventions	Duration of intervention • 21.7 weeks



Painter 2002a (Continued)

Exercise groups

- Type: aerobic
- Description: stationary cycling
- Position: not reported
- Material: ergometer
- · Location: HD unit
- Duration of training sessions: 10 to 30 minutes
- Duration of warm-up/cool-down: 10/not reported minutes
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 14 to 17 on RPE or 70% max HR
- Supervised by: study staff
- Mode of delivery: face-to-face
- Tailoring: individualised intensity
- Modifications/progression: increasing duration, addition of more intense intervals once 20 min was reached
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care or normalized HCT (factorial RCT)

Outcomes

- VO₂ peak
- · Physical functioning
- HRQoL
- HR maximum
- HCT
- Hb
- EPO dose
- Respiratory exchange ratio
- BP maximum

Notes

- The published article does not provide the results for HRQoL. The authors were contacted but could not provide the missing results
- Funding: Amgen

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Stratified by age. Assumed computer-generated
Allocation concealment (selection bias)	Low risk	Sealed envelopes
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias)	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding



Painter 2002a (Continued) Objective outcomes		
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	High risk	One or more outcomes of interest in the review are reported incompletely so that they cannot be entered in a meta-analysis
Other bias	High risk	Private funding. Funder's involvement not specified

Paluchamy 2018	
Study characteristics	s
Methods	 Study design: parallel RCT Study duration: 2 months Study follow-up period: not reported
Participants	 Country: India Setting: HD unit Inclusion criteria: undergoing HD Number: exercise group (10); control group (10) Age range: 51 to 70 years Sex (M/F): exercise group (9/1); control group (9/1) Exclusion criteria: symptomatic cardiovascular disease such as unstable angina, recent MI, congestive cardiac failure Grade II; body temperature more than 101°F; persistent hyperkalaemia before dialysis; active liver disease; musculoskeletal limitations; severe peripheral polyneuropathy; dementia or other mental disorders; on another exercise program; haemodynamically unstable during the dialysis treatment; lower limb amputation
Interventions	Duration of intervention • 12 weeks Exercise group • Type: aerobic • Description: stationary cycling • Position: recumbent • Material: ergometer • Location: HD unit • Duration of training sessions: 10 to 15 minutes • Duration of warm-up/cool-down: 5/not reported minutes • Frequency: 3 times/week
	 Timing in relation to dialysis treatments: during Intensity: not reported Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity and duration



Paluchamy 2018 (Continued)

- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- Kt/V
- SBP
- DBP
- Weight
- SCr
- blood urea
- Calcium
- Phosphate
- Potassium
- Hb
- QoL (KDQOL-SF)

Notes

- Unpublished results were provided by the authors
- The results for the individual domains of KDQOL-SF could not be included in the meta-analysis because they were not rescaled from 0 to 100
- Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement



Paluchamy 2018 (Continued)

Other bias Unclear risk Insufficient information to permit judgement

Parsons 2004

Study characteristics	s			
Methods	 Study design: parallel RCT Study duration: 2 months Study follow-up period: not reported 			
Participants	 Country: Canada Setting: HD unit Inclusion criteria: undergoing HD treatments Number: exercise group (6); control group (7) Mean age ± SD (years): exercise group (60 ± 17); control group (49 ± 25) Sex (M/F): exercise group (3/3); control group (4/3) Mean HD vintage ± SD (months): exercise group (25 ± 25); control group (49 ± 26) Exclusion criteria: cardiovascular, neurological or orthopaedic impairment which would preclude the ability to exercise during the 8-week protocol 			
Interventions	Duration of intervention			
	• 12 weeks			
	Exercise group			
	 Type: aerobic Description: stationary cycling Position: not reported Material: ergometer Location: HD unit Duration of training sessions: 45 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 405 to 50% of maximum load Supervised by: not reported Mode of delivery: not reported Tailoring: individualised intensity Modifications/progression: increased intensity at week 4 if improvement in Wmax Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none 			
	Usual care			
Outcomes	 Maximal work capacity Resting BP HRQoL Blood urea clearance 			



Parsons 2004 (Continued)	Dialysate urea clearance	
Notes	 Missing data: resting SBP and DBP post exercise training intervention for both the exercise group and the control group Funding Kidney Foundation of Canada 	

John Bedal Foundation at Ki

o John Bedal Foundation at Kingston General Hospital

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Stratified on multiple characteristics. Assumed computer-generated
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	Study appears free of other biases

PEAK 2006

F LAK 2000	
Study characteristics	
Methods	 Study design: parallel RCT Study duration: 57 months Study follow-up period: 12 weeks
Participants	 Country: Australia Setting: HD unit Inclusion criteria: > 18 years; on HD for > 3 months; without acute or chronic medical conditions precluding the intervention or collection of outcome measures; independent ambulation with or without an assistive device for > 50 min; adequately dialysed (Kt/V > 1.2) and stable during dialysis; cognition and English language sufficient to understand research procedures and provide written informed consent; willingness to be randomised and to undergo study protocols



PEAK 2006 (Continued)

- Number: exercise group (24); control group (25)
- Mean age ± SD (years): exercise group (60.2 ± 15.2); control group (66.3 ± 13.5)
- Sex (M/F): exercise group (9/4); control group (12/6)
- Median HD vintage, IQR (years): exercise group (3.9, 0.3 to 16.7); control group (1.2, 0.6 to 8.3)
- Mean BMI \pm SD (kg/m²): exercise group (27.9 \pm 6.7); control group (27.3 \pm 5.3)
- Exclusion criteria: cardiac instability; aortic stenosis; unstable cerebral aneurysms; psychological disorder/dementia; active malignancy; proliferative diabetic retinopathy; emphysema; multiple hernias; unstable HD; non-compliance to HD; hemiparesis

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: resistance
- Description: upper and lower limbs exercises
- · Position: seated or supine
- Material: ankle and free-weights dumbbells
- · Location: HD unit
- Duration of training sessions: 45 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: prior and during
- Intensity: 15 to 17 on RPE
- · Supervised by: exercise physiologist
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- Adherence to intervention (sessions attended): 75.9%
- Co-intervention: none

Control group

Usual care

Outcomes

- BMI (kg/m²)
- Body weight (kg)
- Regional fat estimates: subcutaneous mid-thigh fat (cm²)
- Regional fat estimates: total mid-thigh fat (cm²)
- · Waist circumference (cm)
- Serum albumin
- CRP
- IL-10b (pg/mL)
- IL-12a (pg/mL)
- IL-1 a (pg/mL)
- IL-6b (pg/mL)
- IL-8b (pg/mL)
- Lymphocytes (x 10⁹/L)
- Tumour necrosis factor a (pg/mL)
- WBC count (x 10⁹)
- SCr (mol/L)
- Adherence to exercise sessions



PEAK 2006 (Continued)

- Energy intake (20) (kcal/kg/day)
- Mini-nutritional assessment (19) (0 to 30)
- Protein catabolic rate (g/kg/day)
- Protein intake (20) (g/kg/day)
- Physical activity scale
- 6MWT
- Muscle attenuation (Hounsfield unit)
- Mid-arm circumference (cm)
- Mid-calf circumference (cm)
- Mid-thigh circumference (cm)
- Muscle cross-sectional area (cm²)
- Total strength (kg)
- Geriatric depression scale
- QoL
- Dialysis adequacy (Kt/V)

Notes

Funding

- University of Sydney Healthy Ageing Research Program
- · Australian Kidney Foundation
- National Health and Medical Research Council of Australia
- equipment donations from the Australian Barbell Company and SIMBEX Corporation

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Randomisation process independent from study team and use of opaque sealed envelopes
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	High risk	Not all of the study's pre-specified outcomes have been reported. One or more outcomes of interest in the review are reported incompletely so that they cannot be entered in a meta-analysis.
Other bias	High risk	Private funding. Funder's involvement not specified



Pellizzaro 2013		
Study characteristics	s	
Methods	 Study design: parallel RCT (3 arms*) Study duration: 3 months Study follow-up period: 10 weeks 	
Participants	 Country: Brazil Setting: HD unit Inclusion criteria: 18 and 70 years; on dialysis > 3 months; agree to participate by signing an informed consent form Number: exercise group (15); control group (15) Mean age ± SD (years): exercise group (48.9 ± 10.1); control group (51.9 ± 11.6) Sex (M/F): exercise group (7/7); control group (8/6) Mean BMI ± SD (kg/m²): exercise group (23.1 ± 2.6); control group (24.1 ± 3.6) Median HD vintage, IQR (months): exercise group (54.0, 10.7 to 120); control group (54.0, 12 to 78) Exclusion criteria: unstable angina; uncontrolled cardiac arrhythmia; decompensated heart failure SBP > 200 mm Hg, DBP > 120 mm Hg; acute pericarditis or myocarditis; decompensated DM (fasting serum glucose > 300 mg/dL); severe untreated mitral or aortic insufficiency/stenosis, severe lung conditions; acute systemic infection; severe bone disease; lower limb amputations; cognitive disorders unable to perform the proposed tests due to disabling musculoskeletal, bone, or joint disorders 	
Interventions	Duration of intervention 10 weeks Exercise group Type: resistance Description: knee extensions Position: seated Material: free leg weights Location: HD unit Duration of training sessions: varied minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: 50% of 1-RM Supervised by: not reported Mode of delivery: not reported Mode of delivery: not reported Modifications/progression: training load adjusted on 30th day Strategies to enhance adherence: not reported Adherence to intervention: not reported Co-intervention: none Control group Usual care	
2.1		
Outcomes	Serum albumin	

Hb

 Phosphorus • Potassium



Pellizzaro 2013 (Continued)

- hs-CRP
- Urea
- Kt/V
- Post-intervention FVC (L)
- Post-intervention PEmax (cmH₂O)
- Post-intervention Plmax (cmH₂O)
- 6MWT
- QoL

Notes

- *Respiratory muscle training group no included in this review
- Funding: Research Funding of Hospital de Clínicas de Porto Alegre (FIPE/HCPA)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	High risk	One or more primary outcomes is reported using measurements, analysis methods or subsets of the data (e.g. sub-scales) that were not pre-specified. The study report fails to include results for a key outcome that would be expected to have been reported for such a study.
Other bias	Low risk	The study appears to be free of other sources of bias

Rahimimoghadam 2017

Study characteristic	;
Methods	 Study design: parallel RCT Study duration: 3 months Study follow-up period: 8 weeks



Rahimimoghadam 2017 (Continued)

Participants

- Country: Iran
- · Setting: Hospital gym
- Inclusion criteria: 18 to 65 years; history of HD treatment 2 to 3 times/week for at least 6 months; physical ability to perform basic daily activities; Nephrologist's permission to practice the exercise
- Number: exercise group (25); control group (25)
- Mean age ± SD (years): exercise group (39.1 ± 2.2); control group (38.4 ± 1.8)
- Sex (M/F): exercise group (21/4); control group (20/5)
- Mean HD vintage ± SD (months): exercise group (32.2 ± 28.2); control group (45.5 ± 49.5)
- Exclusion criteria: 3 or more sessions of absence in exercises; being a habitual Pilates practitioner; detection of reduced exercise tolerance, including tachycardia, shortness of breath, and feeling too tired or weak; PD during the study; other concurrent clinical conditions, such as cardio-respiratory problems reported by physician and/or patients

Interventions

Duration of intervention

• 8 weeks

Exercise group

- Type: resistance
- Description: modified Pilates
- · Position: not applicable
- · Material: not reported
- · Location: hospital gym
- Duration of training sessions: 45 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: on non-HD days
- · Intensity: not reported
- Supervised by: qualified Pilates professionals
- Mode of delivery: face-to-face
- · Tailoring: not reported
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

Usual care

Outcomes

- Anxiety
- Depression
- · Physical symptoms
- Social dysfunction
- Total score of general health

Notes

• Funding: Kashan University of Medical Sciences

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Bloc randomisation. Assumed computer-generated



Rahimimoghadam 2017 (Continued)			
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding	
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes	
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding	
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data	
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported	
Other bias	Low risk	The study appears to be free of other sources of bias	

Reboredo 2010

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Brazil Setting: HD unit Inclusion criteria: adults undergoing HD who did not exercise on a regular basis for at least 6 months Number: exercise group (14); control group (14) Mean age ± SD (years): exercise group (49.6 ± 10.6); control group (43.5 ± 12.8) Sex (M/F): exercise group (4/7); control group (4/7) Mean HD vintage ± SD (months): exercise group (41.9 ± 42.4); control group (60.1 ± 54.4) Mean BMI ± SD (kg/m²): exercise group (22.6 ± 2.3); control group (22.9 ± 4.1) Exclusion criteria: DM; unstable angina; uncontrolled arterial hypertension (SBP ≥ 200 mm Hg and/ or DBP ≥ 120 mm Hg); use of antiarrhythmic drugs; severe pneumopathies; acute systemic infection; severe renal osteodystrophy; disabling neurological and muscle-skeletal disorders
Interventions	Duration of intervention 12 weeks Exercise group Type: aerobic Description: stationary cycling Position: not reported Material: horizontal ergometer



Reboredo 2010 (Continued)

- · Location: HD unit
- Duration of training sessions: 35 minutes
- Duration of warm-up/cool-down: 15/3 minutes
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: 4 to 6 on RPE (1 to 10)
- · Supervised by: supervised not further defined
- · Mode of delivery: not reported
- · Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention (mean ± SD sessions attended): 75.3% ± 15.2%
- · Co-intervention: none

Control group

· Usual care

Outcomes

- Serum albumin
- Hb
- Calcium (mg/dL)
- Phosphorus (mg/dL)
- Potassium (mEq/L)
- End systolic volume (mL)
- End diastolic volume (mL)
- LVMI (g/m²)
- Ejection fraction (%)
- Systolic volume (mL)
- HF (ms²)
- LF (ms²)
- LF/HF
- pNN50 (%)
- RMSSD (ms)
- SDNN index (ms)
- SCr (mg/dL)
- Adherence to exercise sessions
- Kt/V

Notes

Funding

- Fundação de Amparo à Pesquisa do Estado de Minas Gerais
- Coordenação de Aperfeiçoamento de Pessoal de Nível Superior
- IMEPEN Foundation

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement



Reboredo 2010 (Continued)		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Rezaei 2015

Study characteristics	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 10 weeks
Participants	 Country: Iran Setting: HD unit Inclusion criteria: aged 15 to 65 years, under treatment of HD for at least 3 months Number: exercise group (25); control group (26) Mean age ± SD (years): exercise group (44.0 ± 7.9); control group (42.6 ± 12.7) Sex (M/F): exercise group (21/4); control group (14/12) Mean HD vintage ± SD (months): exercise group (42.7 ± 38.9); control group (35.5 ± 27.0) Exclusion criteria: progressive cardiovascular or respiratory diseases; restricting musculoskeletal dis order; lacking the physical power to exercise; using any medicine or other procedures for treating de pression; not being under treatment of HD 2 or 3 times/week, not performing the exercise program fo 3 times continuously or 5 times alternatively; dissatisfaction for continuing collaboration; problem atic haemodynamic instability.
Interventions	 Duration of intervention 10 weeks Exercise group Type: range of movement Description: joints warming actions, stretching exercises, motions of lower back muscles and ab domen, and deep breathing exercises. Position: not reported Material: not reported Location: home



Rezaei 2015 (Continued)

- Duration of training sessions: 35 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: not during, home-based
- Intensity: not reported bur described as less than moderate
- Supervised by: unsupervised
- Mode of delivery: posters
- · Tailoring: not reported
- Modifications/progression: not reported
- Strategies to enhance adherence: phone calls and visits in dialysis
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes	•	Depression (BDI)
----------	---	------------------

Notes • Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	High risk	The study report fails to include results for key outcomes that would be expected to have been reported for such a study.
Other bias	Unclear risk	Insufficient information to permit judgement



Study characteristics	
Methods	Study design: parallel RCT
	Study duration: 21 months
	Study follow-up period: 12 weeks
Participants	Country: Brazil
	Setting: HD unit
	 Inclusion criteria: > 18 years; HD > 3 months; permission of the attending Nephrologist; independen ambulation for > 50 metres with or without an assistive device; cognition and willingness to be ran domly assigned into groups and to undergo the study protocols
	 Number: exercise group (30); control group (29)
	 Mean age ± SD (years): exercise group (54.49 ± 11.97); control group (57.10 ± 16.20)
	 Sex (M/F): exercise group (20/8); control group (15/9)
	 Mean BMI ± SD (kg/m²): exercise group (26.36 ± 4.48); control group (25.54 ± 3.95)
	 Mean HD vintage ± SD (years): exercise group (1.54 ± 1.26); control group (2.35 ± 1.66)
	 Exclusion criteria: acute or chronic medical conditions that would preclude exercise or the collection of the outcome measure data
Interventions	Duration of intervention
	• 12 weeks
	Exercise group
	Type: resistance
	 Description: upper and lower limbs exercises
	Position: not reported
	Material: weights and resistance bands
	Location: HD unit
	Duration of training sessions: 40 to 50 minutes
	Duration of warm-up/cool-down: not reported
	Frequency: 3 times/week
	Timing in relation to dialysis treatments: prior and during
	Intensity: not reported
	Supervised by: clinical exercise physiologists
	Mode of delivery: face-to-face Tailoring to individual in a description of the product of
	Tailoring: individualised resistance level Madifications / nysographics in property yesistance level to makintain Neff yengtitions.
	Modifications/progression: increasing resistance level to maintain N of repetitions Strategies to enhance adherence not repeted.
	Strategies to enhance adherence: not reported Adherence to intervention (many LSD sessions attended), upper limbs (67% L 18%), lower limbs (93%).
	 Adherence to intervention (mean ± SD sessions attended): upper limbs (67% ± 18%); lower limbs (83% ± 9%)
	Co-intervention: none
	Control group
	Very low-intensity exercise without load and progression and a breathing exercise
Outcomes	• BMI (kg/m²)
	Bone mineral content (kg)
	Total LBM (kg)
	Trunk lean mass (kg)
	Arm lean mass (kg)

Leg lean mass (kg)Total fat mass (kg)



Rosa 2018	(Continued)
-----------	-------------

- Total mass (kg)
- 6MWT (m)
- Sit-to-stand test (rep)
- Handgrip strength (kg/strength)
- Flexibility (cm)
- QoL

Notes

• Funding: nil

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Allocation concealment performed by researcher not involved in recruitment or assessment
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	Sham exercise in the control arm
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	Sham exercise in the control arm
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Rouchon 2016

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: France Setting: not reported Inclusion criteria: undergoing PD treatments Number: exercise group (9); control group (9) Loss to follow-up: exercise group (1); control group (5)



Rouchon 2016 (Continued)

- Mean age \pm SD (years): exercise group (66.94 \pm 2.89); control group (57.80 \pm 3,30)
- Sex M/F: exercise group (4/4); control group (0/4)
- Mean BMI \pm SD (kg/m²): exercise group (31.12 \pm 1.90); control group (26.01 \pm 1.59)
- Mean HD vintage \pm SD (years): exercise group (1.94 \pm 0.46); control group (2.1 \pm 0.8)
- · Exclusion criteria: not reported

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: combined
- Description: bicycle-HIIT sessions + upper and lower limbs exercises
- · Position: not reported
- Material: ergometer, weights
- · Location: not reported
- · Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: not reported/15 minutes
- · Frequency: 2 times/week
- · Timing in relation to dialysis treatments: PD patients only
- · Intensity: not reported
- · Supervised by: not reported
- · Mode of delivery: not reported
- · Tailoring: not reported
- Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

Usual care

Outcomes

- 6MWT
- VO₂ peak

Notes

- Abstract-only publication: unpublished results were provided by the authors
- Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Drawing of lots (provided by author)
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding



Rouchon 2016 (Continued)		
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement

Samara 2016

 Study design: parallel RCT Study duration: not reported Study follow-up period: 16 weeks
 Country: Greece Setting: outside the HD unit Inclusion criteria: HD 3 days/week, 4 hours/session for at least 6 months prior to the study Number: exercise group (16); control group (13) Mean age ± SD (years): exercise group (48.0 ± 11.3); control group (48.6 ± 15.4) Sex (M/F): exercise group (13/2); control group (11/1) Mean BMI ± SD (kg/m²): exercise group (24.23 ± 2.81); control group (25.46 ± 5.14) Exclusion criteria: no acute or chronic medical conditions that would affect the measured data; recen MI (within 6 weeks); malignant arrhythmias; unstable angina; Hb < 10 g/dL or inconstant throughou the study; receiving beta-blockers or other antiarrhythmic medication
Duration of intervention
16 weeks Exercise group
 Type: aerobic Description: swimming (freestyle, breaststroke, and backstroke) Position: not applicable Material: pool, foam tubes, buoyancy belts, paddles Location: pool Duration of training sessions: 20 to 40 minutes Duration of warm-up/cool-down: 10/10 minutes Frequency: 3 times/week Timing in relation to dialysis treatments: on non-HD days Intensity: 13 to 14 on RPE (6 to 20) Supervised by: specialised exercise trainer



Samara 2016 (Continued)

- Tailoring: individualised intensity
- Modifications/progression: increasing intensity
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- 6MWT
- Sit and reach (cm)
- Time up and go
- Handgrip (kg)
- Sit-to-stand test
- · QoL

Notes

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Drawing of lots
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Segura-Orti 2009

Study characteristics



Segura-Orti 2009 (Continued)

M	eτ	ho	as

- Study design: parallel RCT
- · Study duration: 18 months
- Study follow-up period: 24 weeks

Participants

- · Country: Spain
- · Setting: HD clinics
- Inclusion criteria: stable condition under their medication and undertaking HD sessions for at least 3 months
- Number: exercise group (19); control group (8)
- Mean age \pm SD (years): exercise group (53.5 \pm 18.0); control group (60.1 \pm 16.9)
- Sex (M/F): exercise group (11/6); control group (7/1)
- Mean BMI \pm SD (kg/m²): exercise group (24.6 \pm 2.6); control group (24.9 \pm 2.2)
- Mean HD vintage ± SD (months): exercise group (37.3 ± 34.9); control group (53.7 ± 42.0)
- Exclusion criteria: recent MI (6 weeks); uncontrolled hypertension; malignant arrhythmias; unstable angina and any disorder that could be exacerbated by activity

Interventions

Duration of intervention

24 weeks

Exercise group

- Type: resistance
- Description: lower limbs isotonic and isometric exercises
- · Position: not reported
- Material: ankle weights
- · Location: HD unit
- Duration of training sessions: 25 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 14 on RPE
- Supervised by: physiotherapist
- Mode of delivery: face-to-face
- · Tailoring: not reported
- Modifications/progression: progressive not further defined
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Cycling on the minimum possible workload

Outcomes

- Sit-to-stand test (10 seconds)
- Sit-to-stand test (60 repetitions)
- 6MWT
- HRQoL

Notes

• Funding: Universidad CEU Cardenal Herrera

Risk of bias

Bias

Authors' judgement Support for judgement



Segura-Orti 2009 (Continued)		
Random sequence generation (selection bias)	Low risk	Random numbers table
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	Sham exercise in the control arm
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	Sham exercise in the control arm
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Sheshadri 2020

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 24 weeks
Participants	 Country: USA Setting: HD units Inclusion criteria: ≥ 18 years; receiving in-centre HD or any form of PD; having telephone access; being ambulatory Number: exercise group (30), control group (30) Median age, IQR (years): exercise group (60, 53 to 66); control group (56, 51 to 65) Sex (M/F): exercise group (28/2); control group (19/11) Median BMI, IQR (kg/m²): exercise group (26.9, 25.3 to 32.9); control group (31.6, 26.7 to 34.6) Median HD vintage, IQR (months): exercise group (3.7, 1.5 to 7.2); control group (1.9, 0.95 to 4.7) Exclusion criteria: patients using wheelchairs or scooters
Interventions	Duration of intervention 12 weeks Exercise group Type: aerobic Description: walking and weekly steps goal



Sheshadri 2020 (Continued)

- Position: not applicable
- Material: pedometer
- · Location: not reported
- Duration of training sessions: not applicable
- Duration of warm-up/cool-down: not applicable
- Frequency: not applicable
- Timing in relation to dialysis treatments: outside treatments
- · Intensity: not reported
- · Supervised by: unsupervised
- · Mode of delivery: weekly phone counselling session
- Tailoring: based on baseline daily steps
- Modifications/progression: 10% of the previous week target
- Strategies to enhance adherence: weekly phone counselling
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Usual care and received a pedometer

Outcomes

- Physical function (SF-36 physical function scores, short physical performance battery)
- Endothelial function (reactive hyperemia index with peripheral arterial tonometry)
- HR variability (SDNN, LF/HF)
- Dialysis symptoms index
- SF-36 physical functioning and vitality score
- Centers for Epidemiologic Studies Depression Scale

Notes

Funding

- American Kidney Fund Clinical Scientist in Nephrology Fellowship
- Ruth L. Kirschstein National Research Service Award Individual Postdoctoral Fellowship
- International Society of Nephrology fellowship

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Block randomisation using computer generated program
Allocation concealment (selection bias)	Low risk	Sealed, opaque envelopes used to perform allocation concealment
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Nil blinding performed
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding



Sheshadri 2020 (Continued)		
Incomplete outcome data (attrition bias) All outcomes	Low risk	Low and equal rates of drop-out in both arms of treatment, unlikely to affect outcome
Selective reporting (reporting bias)	Low risk	All pre-specified outcome variables reported in body of text or supplementary material
Other bias	Low risk	Study appears to be free from other sources of bias

ioliman 2015	
Study characteristics	
Methods	 Study design: parallel RCT Study duration: 5 months Study follow-up period: 8 weeks
Participants	 Country: Egypt Setting: HD unit Inclusion criteria: aged > 18 years; minimum HD vintage of 3 months, receiving HD 3 times/week, fo 3 or 4 hours/session; stable on HD, Kt/V > 1.2; bicarbonate dialysis solution; unintentional low dietary protein intake < 1 g/kg of ideal weight/day for at least 2 months; unintentional low dietary energy intake < 30 kcal/kg of ideal weight/day for at least 2 months Number: exercise group (18); control group (12) Age: exercise group (61% between 40 and 60); control group (58% between 40 and 60) Sex (M/F): exercise group (8/10); control group (6/6) Mean BMI ± SD (kg/m²): exercise group (25.6 ± 4.3); control group (27.2 ± 5.7) Exclusion criteria: any acute or chronic medical conditions that would make exercise training potentially hazardous or primary outcomes impossible to assess; problematic AV fistula; uncontrolled hy pertension; congestive heart failure; arrhythmia requiring treatment; unstable angina; major valvula heart disease; MI; significant arteriosclerosis; a risk of fracture; musculoskeletal disorders; change in the resting ECG; severe aortic stenosis; suspected or known dissecting aneurysm; myocarditis; participation in another trial; inadequate dialysis Kt/V < 1.2; Hb < 10 g/dL; unstable on dialysis
Interventions	Duration of intervention • 8 weeks Exercise group
	 Type: range of movement Description: range of motion exercises Position: not reported Material: not reported Location: HD unit Duration of training sessions: 15 minutes Duration of warm-up/cool-down: not reported Frequency: 3 times/week Timing in relation to dialysis treatments: during Intensity: not reported Supervised by: not reported Mode of delivery: face-to-face and booklet

• Modifications/progression: not reported



So	liman	2015	(Continued)
----	-------	------	-------------

- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Usual care

Outcomes

- Fatigue (Iowa Fatigue Scale)
- Potassium
- Calcium
- Phosphate
- Hb
- Resting BP

Notes

• Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	High risk	Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Song 2012a

		_	
Study	chai	racte	ristics

Methods

- Study design: parallel RCT
- Study duration: not reported



Song 2012a (Continued)

• Study follow-up period: 12 weeks

Participants

- · Country: Korea
- · Setting: outpatient clinic
- Inclusion criteria: ≥ 18 years; on HD > 3 months; under the permission of their Nephrologist; ability
 to maintain a seated position; independent ambulation of 50 m or more, with or without an assistive
 device; adequately dialysed (most recent Kt/V = 1.2); stable during dialysis
- Number: exercise group (22); control group (22)
- Mean age ± SD (years): exercise group (52.1 ± 12.4); control group (54.6 ± 10.1)
- Sex (M/F): exercise group (8/12); control group (12/8)
- Mean HD vintage ± SD (months): exercise group (38.9 ± 26.1); control group (45.9 ± 56.2)
- · Exclusion criteria: not reported

Interventions

Duration of intervention

12 weeks

Exercise group

- · Type: resistance
- Description: upper and lower limbs exercises
- · Position: seated
- Material: ankle weights and resistance bands
- · Location: conference room adjacent to HD unit
- Duration of training sessions: 20 minutes
- Duration of warm-up/cool-down: 5/5 minutes
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: not during
- Intensity: 11 to 15 on RPE
- · Supervised by: investigator and research assistant
- · Mode of delivery: face-to-face
- Tailoring: individualised resistance level
- · Modifications/progression: ankle weights added at week 4
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- Body fat rate (%)
- Visceral fat area (cm²)
- · Skeletal muscle mass (kg)
- Waist circumference (cm)
- HDL cholesterol (mg/dL)
- LDL cholesterol (mg/dL)
- Total cholesterol (mg/dL)
- Triglyceride (mg/dL)
- Balance (sec)
- Shoulder flexibility (cm)
- Waist flexibility (cm)
- Arm muscle circumference (mm)
- · Grip strength (kg)
- Leg muscle strength (kg)



Song 2012a (Continued)

- · Sit-to-stand test
- QoL

Notes

· Funding: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Suhardjono 2019

Study	charact	eristics
JLUUV	ciiuiuci	ei istics

			-
М	etl	hο	ds

- Study design: parallel RCT (3 arms)
- · Study duration: 3 months
- Study follow-up period: 12 weeks

Participants

- Country: Indonesia
- · Setting: not reported
- Inclusion criteria: ≥ 18 years; given their consent to participate in the study; maintenance dialysis for at least 3 months
- Number: aerobic group (42); combined group (40); control group (41)
- Mean age \pm SD (years): aerobic group (49.78 \pm 11.65); combined group (46.38 \pm 14.19); control group (50.54 \pm 10.83)
- Sex (M/F): aerobic group (28/14); combined group (21/18); control group (18/21)
- Median HD vintage, range (months): aerobic group (48, 4 to 192); combined group (48, 6 to 204); control
 group (60, 5 to 240)



Suhardjono 2019 (Continued)

 Exclusion criteria: travelling on dialysis; being hospitalised for any reason within the past 3 months; having arrhythmias; being on dialysis for less than 2-week intervals; having a limited range of motion of extremities; being immobilized

Interventions

Duration of intervention

• 12 weeks

Aerobic exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: ergometer
- Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 2 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 40% to 60%, and then 60% to 80% of max HR
- · Supervised by: nephrologist, sports medicine doctor
- Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: increasing intensity
- · Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- Co-intervention: none

Combined aerobic + resistance exercise group

- · Type: combined
- Description: stationary cycling + ankle weightlifting
- Position: not reported
- Material: ankle weights
- Location: HD unit
- Duration of training sessions: not reported
- Duration of warm-up/cool-down: not reported
- Frequency: 2 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 11 to 13 on RPE
- Supervised by: nephrologist, sports medicine doctor
- · Mode of delivery: face-to-face
- · Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- Skeletal muscle index (kg/m²),
- Handgrip strength (kg)
- Gait speed (m/sec)



Suhardjono 2019 (Continued)

- Right lower extremity muscle strength (kg)
- Left lower extremity muscle strength (kg)
- CRP (g/dL)
- · Malnutrition-inflammation score
- QoL

Notes

- The results for CRP were not included in the meta-analysis because the reported numbers and unit of measure were implausible
- Funding: Universitas Indonesia

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Block randomisation. Assumed computer-generated
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Toussaint 2008

Study characteristics	
-----------------------	--

М	ethods	
141	cuious	•

- Study design: cross-over RCT
- · Study duration: not reported
- Study follow-up period: 24 weeks

Participants

- Country: Australia
- Setting: satellite HD unit
- Inclusion criteria: HD > 3 months; able to give informed consent; able and willing to commit to exercise regularly for 3 months
- Number: exercise group (9); control group (10)



Toussaint 2008 (Continued)

- Median age, range (years): exercise group (67, 60 to 83); control group (70, 28 to 77)
- Sex (M/F): exercise group (5/4); control group (4/6)
- Mean HD vintage ± SD (months): exercise group (35 ± 31); control group (72 ± 56)
- Mean BMI ± SD (kg/m²): exercise group (27 ± 4); control group (24 ± 4)
- Exclusion criteria: active or symptomatic cardiovascular or respiratory disease; musculoskeletal abnormalities that limited exercise ability

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: ergometer
- · Location: HD unit
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- · Frequency: 3 times/week
- Timing in relation to dialysis treatments: during
- Intensity: no target
- Supervised by: unsupervised
- · Mode of delivery: not reported
- · Tailoring: not reported
- · Modifications/progression: not reported
- · Strategies to enhance adherence: not reported
- Adherence to intervention (sessions attended); 88%
- Co-intervention: none

Control group

Usual care

Outcomes

- CRP
- Hb
- Calcium × phosphate product
- PTH
- Beta-2-microglobulin
- Homocysteine
- BF
- Albumin
- Augmentation index
- Brain-natriuretic peptide
- Pulse pressure
- Pulse wave velocity

Notes

- We only included the results at 3 months as we felt that the 1-month washout period was insufficient to eliminate the carry-over effect.
- Funding: National Health and Medical Research Grant

Risk of bias

Bias Authors' judgement Support for judgement



Toussaint 2008 (Continued)		
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Low risk	Sealed envelops
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Tsuyuki 2003

Study characteristics	s
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 5 months
Participants	 Country: Japan Setting: not reported Inclusion criteria: receiving regular HD Number: exercise group (17); control group (12) Mean age ± SD (years): exercise group (40.1 ± 11.9); control group (39.7 ± 10.7) Sex (M/F): exercise group (9/8); control group (5/7) Mean HD vintage ± SD (years): exercise group (2.1 ± 2.5); control group (2.7 ± 2.6) Exclusion criteria: hypertension (> 170/110 mm Hg); anaemia (HCT < 18%); weight gain (< 3.0 kg); heart disease; liver dysfunction; DM; chronic obstructive pulmonary disease
Interventions	Duration of intervention • 20 weeks Exercise group • Type: aerobic • Description: cycling, walking and jogging • Position: not reported



Tsuyuki 2003 (Continued)

- Material: ergometer
- · Location: not reported
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 2 to 3 times/week
- Timing in relation to dialysis treatments: on non-HD days
- Intensity: 50% to 60% of max HR
- Supervised by: medical supervision
- Mode of delivery: not reported
- Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- VO₂ peak
- HR
- BP
- Minute ventilation
- Carbon dioxide output
- Respiratory ratio
- · Tidal volume
- · Anaerobic threshold
- Hb
- HCT

Notes

- The authors were contacted during the previous version of this review for clarification on the methods, but without result.
- Funding: not reported

Bias	Authors' judgement	Support for judgement	
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement	
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding	
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding	
Blinding of outcome assessment (detection bias)	Low risk	No patient-reported outcome	



Tsuyuki 2003	(Continued)
Subjective or	utcomes

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Uchiyama 2019	
Study characteristics	
Methods	 Study design: parallel RCT Study duration: 17 months Study follow-up period: 3 months
Participants	 Country: Japan Setting: not reported Inclusion criteria: stable PD patients aged 20–90 years who had started with and undergone PD for at least 3 months Number: exercise group (24); control group (23) Mean age ± SD (years): exercise group (64.9 ± 9.2); control group (63.2 ± 9.5) Sex (M/F): exercise group (19/5); control group (16/7)

Sex (M/F): exercise group (19/5); control group (16/7) Mean BMI ± SD (kg/m²): exercise group (22.70 ± 3.50); control group (24.60 ± 4.10)

- Mean PD vintage ± SD (years): exercise group (3.6 ± 2.7); control group (4.0 ± 2.8)
- Exclusion criteria: uncontrolled hypertension (BP > 180/110 mm Hg); severe anaemia (Hb < 7 mg/dL); active and proliferative diabetic retinopathy; symptomatic coronary artery disease or cerebrovascular disease within 3 months before study recruitment; current heart failure (NYHA classes III and IV); symptomatic and fatal arrhythmia; significant valvular heart disease; difficulty walking without a walking aid owing to orthopaedic problems; a history of cerebrovascular disease; a history of peripheral artery disease

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: combined
- Description: walking, upper and lower limbs exercises
- Position: not applicable
- Material: resistance bands
- · Location: home
- Duration of training sessions: 20 to 30 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- Timing in relation to dialysis treatments: PD patients only
- Intensity: aerobic (40% to 60% of the peak HR and 11 to 13 on the Borg RPE); resistance (70% of 1RM)
- Supervised by: unsupervised
- Mode of delivery: face-to-face
- Tailoring: individualised intensity



Uchiyama 2019 (Continued)

- Modifications/progression: increasing duration
- Strategies to enhance adherence: weekly postcard
- Adherence to intervention: mean (SD) number of sessions attended: aerobic: 52% (40); resistance: 76% (37)
- · Co-intervention: none

Control group

· Usual care

Outcomes

- Incremental shuttle walking test (m)
- HR-QoL (KDQOL-SF)
- Handgrip strength (kg)
- Quadriceps strength (kg)
- BMI (kg/m²)
- Waist circumference (cm)
- Leg circumference (cm)
- Skeletal muscle mass index (kg/m²)
- Albumin (g/L)
- nPCR (g/kg/day)
- HbA1c (%)
- Total cholesterol (mmol/L)
- HDL cholesterol (mg/dL)
- Triglyceride (mg/dL)
- · Homeostasis model assessment of insulin resistance
- Renal Kt/V
- Ultrafiltration (mL/day)
- PD Kt/V
- CRP (mg/L)
- ANP (pg/mL)
- Brachial-ankle pulse wave velocity

Notes

· Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	External to the investigators
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias)	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding



Uchiyama 2019 (Continued)

Subjective outcomes

Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

van Vilsteren 2005

vali vitstereli 2005	
Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Netherlands Setting: single centre Inclusion criteria: prevalent HD patients Number (randomised/analysed): exercise group (60/53); control group (43/43) Mean age ± SD (years): exercise group (52 ± 15); control group (58 ± 16) Sex (M/F): exercise group (38/22); control group (30/13) Mean HD vintage ± SD (years): exercise group (3.22 ± 4.08); control group (3.90 ± 4.41) Exclusion criteria: severe cardiovascular disease; use of beta-blockers; unstable angina pectoris; orthopaedic complaints

Interventions

Duration of intervention

• 12 weeks

Exercise group

- · Type: combined
- Description: callisthenics, steps, flexibility and low weight resistance exercises + stationary cycling
- Position: not reported
- Material: multitrainer
- Location: HD unit
- Duration of training sessions: 20 to 30 minutes
- Duration of warm-up/cool-down: 20/5 to 10 minutes
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during and prior to HD
- Intensity: 60% of max HR
- · Supervised by: not reported
- Mode of delivery: not reported
- · Tailoring: individualised intensity
- Modifications/progression: not reported
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group



van Vilsteren 2005 (Continued)

• Usual care

Outcomes

- Muscle strength
- Physical functioning
- VO₂ peak
- HRQoL
- BP
- HR
- Cholesterol
- Depression
- Kt/V
- HCT
- Hb
- Behavioural change
- Mean body weight

Notes

- The results for VO_2 max could not be included in the meta-analysis because the number of participants in each group was not provided. The authors were contacted to obtain the missing information
- Funding: not reported

Bias	Authors' judgement	Support for judgement	
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement	
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding	
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding	
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding	
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups	
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported	
Other bias	Unclear risk	Insufficient information to permit judgement	



Wilund 2010

		_	
Stud	v cha	racto	ristics

Methods

- Study design: parallel RCT
- · Study duration: not reported
- Study follow-up period: 16 weeks

Participants

- Country: USA
- · Setting: HD unit
- Inclusion criteria: HD patients; 30 to 70 years, non-smoking; BMI < 35 kg/m²
- Number: exercise group (8); control group (9)
- Mean age ± SD (years): exercise group (60.8 ± 3.2); control group (59.0 ± 4.9)
- Sex (M/F): exercise group (3/4); control group (3/5)
- Mean HD vintage ± SD (months): exercise group (63.3 ± 8.7); control group (44.6 ± 12.2)
- BMI ± SD (kg/m²): exercise group (30.10 ± 2.40); control group (29.00 ± 2.00)
- Exclusion criteria: orthopaedic problems that prevented cycling during dialysis; chronic obstructive
 pulmonary disease, coronary heart failure or cardiovascular surgery (e.g. coronary bypass, valve replacement or angioplasty) in the past 6 months; did not get medical clearance from a primary care
 physician; participation in intradialytic exercise training for 6 months prior to recruitment in the study

Interventions

Duration of intervention

• 16 weeks

Exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: not reported
- · Material: ergometer
- Location: HD unit
- Duration of training sessions: 5 to 45 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 14 on RPE
- · Supervised by: study staff
- Mode of delivery: face-to-face
- Tailoring: individualised intensity and duration
- Modifications/progression: increasing duration 5 to 10 min/session
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

· Usual care

Outcomes

- BMI (kg/m²)
- · Serum albumin
- HCT
- ALP
- Calcium × phosphorous product
- Calcium (mg/dL)
- Phosphorous (mg/dL)
- Potassium (mEq/L)



Wilund 2010 (Continued)

- DBP
- SBP
- Epicardial fat thickness
- Left atrial volume index
- LVMI (g/m²)
- Myocardial performance index
- Relative wall thickness
- Cholesterol (mg/dL)
- CRP (mg/L)
- IL-6 (pg/mL)
- Fetuin-A (ng/mL)
- Blood urea nitrogen/creatinine ratio
- Thiobarbituric acid reactive substances (µmol/L)
- Shuttle walk distance

Notes

• Funding: College of Medicine, University of Illinois at Urbana–Champaign

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	Low risk	No patient-reported outcome
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Wu 2014d

Study	char	acte	ristics
-------	------	------	---------

Methods • Study design: parallel RCT



Wu 2014d (Continued)

- · Study duration: not reported
- Study follow-up period: 12 weeks

Participants

- · Country: China
- · Setting: HD unit
- Inclusion criteria: relatively stable disease; good compliance and co-operation with the doctor; no apparent cardiovascular complications (such as heart failure, severe arrhythmia, angina or cerebrovascular disease) or infection; no orthopaedic problems that would prevent cycling during dialysis; BP < 180/100 mm Hg; HD duration > 3 months
- Number: exercise group (34); control group (35)
- Median age, IQR (years): exercise group (45, 37 to 48); control group (44, 41 to 50)
- Sex (M/F): exercise group (27/5); control group (28/5)
- Mean HD vintage ± SD (months): exercise group (55.5 ± 37.3); control group (39.8 ± 29.7)
- Exclusion criteria: any chronic diseases not under control; retinal laser treatment; history of acute MI; joint replacement or fracture of the lower limb within the previous 6 months; severe cognitive disturbance

Interventions

Duration of intervention

12 weeks

Exercise group

- Type: aerobic
- · Description: stationary cycling
- · Position: recumbent
- · Material: ergometer
- Location: HD unit
- Duration of training sessions: 10 to 15 minutes
- Duration of warm-up/cool-down: 5 minutes/not reported
- Frequency: 3 times/week
- · Timing in relation to dialysis treatments: during
- Intensity: 12 to 16 on RPE
- · Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: individualised intensity and duration
- Modifications/progression: increasing intensity
- Strategies to enhance adherence: encouragements by study staff
- Adherence to intervention: not reported
- · Co-intervention: none

Control group

· Stretching exercises

Outcomes

- 6MWT
- Time to walk up and go test
- Grip strength
- Sit-to-stand test
- Time to perform 10 sit-to-stand manoeuvres
- QoL

Notes

Funding: nil



Wu 2014d (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No objective outcomes
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Low risk	The study appears to be free of other sources of bias

Study characteristics	
Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: 12 weeks
Participants	 Country: Turkey Setting: single centre Inclusion criteria: prevalent HD patients Number (randomised/analysed): exercise group (20/19); control group (20/18) Mean age ± SD (years): exercise group (38 ± 14); control group (41 ± 10) Sex (M/F): exercise group (9/11); control group (7/3) Median/mean HD vintage ± SD (months): 10.5/21.9 ± 14.2 (for all 40 patients) Exclusion criteria: unstable hypertension; arrhythmia or cardiac angina after 10 min of fast pedalling ischaemic cardiac pain; unstable angina; congestive heart failure grade II; significant cardiac valve disease; conduction abnormalities on the ECG; cerebrovascular disease; electrolyte imbalance; persistent hyperkalaemia before dialysis; DM; active liver disease; arthritic or orthopaedic problems limiting exercise: peripheral vascular disease; 'undisciplined patients'
Interventions	Duration of intervention • 12 weeks



Yurtkuran 2007 (Continued)

Exercise group

- Type: yoga
- Description: modified yoga exercise
- Position: seated, supine and standing
- Material: not reported
- · Location: not reported
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 2 times/week
- · Timing in relation to dialysis treatments: not reported
- Intensity: not reported
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: postures adapted
- Modifications/progression: increasing intensity and duration
- Strategies to enhance adherence: not reported
- · Adherence to intervention: not reported
- · Co-intervention: none

Control group

• Usual care

Outcomes

- Grip strength
- HDL cholesterol
- Triglyceride
- Pain
- Fatigue
- Sleep disturbance
- Urea
- SCr
- Calcium
- ALP
- Phosphorus
- Erythrocyte
- HCT

Notes

- 3 patients that missed 3 sessions and adhered poorly to the exercise instructions were excluded from the analyses
- Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer random number generator
Allocation concealment (selection bias)	Low risk	Concealed from the investigators
Blinding of participants and personnel (perfor- mance bias)	High risk	No blinding



Yurtkuran 20	07 (Continued)
--------------	-----------------------

ΛI	outcomes	
Αı	outcomes	•

Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	Blinding of outcome assessment ensured, and unlikely that the blinding could have been broken
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes (of interest to this review) were reported
Other bias	Unclear risk	Insufficient information to permit judgement

Zhao 2017

Study characteristics

Methods

- Study design: parallel RCT (3 arms*)
- Study duration: not reported
- Study follow-up period: 18 weeks

Participants

- Country: China
- · Setting: HD unit
- Inclusion criteria: HD had been performed for at least 3 months or at least 3 times within 1 week; ability
 to ride a bicycle
- Number: aerobic exercise + escitalopram group (63); aerobic exercise only group (63); escitalopram only control group (63)
- Median age, IQR (years): aerobic exercise + escitalopram group (52.9, 43.9 to 65.8); aerobic exercise only group (53.6, 44.5 to 66.3); escitalopram only control group (54.1, 42.3 to 68.7)
- Sex (M/F): aerobic exercise + escitalopram group (39/24); aerobic exercise only group (41/22); escitalopram only control group (40/23)
- Median HD vintage, IQR (months): aerobic exercise + escitalopram group (24.9, 13.8 to 35.7); aerobic exercise only group (25.3, 14.9 to 34.2); escitalopram only control group (24.7, 15.6 to 36.1)
- Exclusion criteria: opportunistic infections; medical therapy for other diseases during the last 3 months; SBP > 160 mm Hg and/or DBP > 110 mm Hg before and after HD and/or at hours 2 and 3 during HD; symptoms for interrupting the exercises, such as chest pain, dyspnoea, body temperature 38°C and cardiac arrhythmias; signs of neurological vertigo and/or imbalance; non-adherence to the exercise program and instability in haemodynamic parameters after exercises

Interventions

Duration of intervention

12 weeks

Exercise group

- Type: yoga
- · Description: modified yoga exercise
- · Position: seated, supine and standing
- Material: not reported



Zhao 2017 (Continued)

- · Location: not reported
- Duration of training sessions: 30 minutes
- Duration of warm-up/cool-down: not reported
- Frequency: 2 times/week
- Timing in relation to dialysis treatments: not reported
- · Intensity: not reported
- Supervised by: not reported
- · Mode of delivery: not reported
- Tailoring: postures adapted
- Modifications/progression: increasing intensity and duration
- Strategies to enhance adherence: not reported
- Adherence to intervention: not reported
- Co-intervention: none

Control group

• Escitalopram: 20 mg/day

Outcomes

- IL-18 (pg/mL)
- IL-6 (pg/mL)
- · QoL

Notes

- *Aerobic exercise only not included in our meta-analyses
- Funding: not reported

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Insufficient information to permit judgement
Allocation concealment (selection bias)	Unclear risk	Insufficient information to permit judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	No blinding
Blinding of outcome assessment (detection bias) Objective outcomes	Low risk	No blinding of outcome assessment (or not reported), but the outcome measurement is not likely to be influenced by lack of blinding
Blinding of outcome assessment (detection bias) Subjective outcomes	High risk	No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Insufficient information to permit judgement
Selective reporting (reporting bias)	Unclear risk	Insufficient information to permit judgement
Other bias	Unclear risk	Insufficient information to permit judgement



1RM - 1-repetition maximum test; 6MWT - 6 minute walk test; AIDS - acquired immune deficiency syndrome; ALP - alkaline phosphatase; AV - arteriovenous; BCM - body composition monitor; BDI - Beck Depression Index; BMD - bone mineral density; BMI - body mass index; BP - blood pressure; bpm - beats per minute; CAPD - continuous ambulatory peritoneal dialysis; COP - centre of foot pressure; CRP- C-reactive protein; DBP - diastolic blood pressure; DM - diabetes mellitus; ECG - echocardiograph; EPO - erythropoietin; ESKD - end-stage kidney disease; FEV - forced expiratory volume; FIM - functional independence measure; FM - fat mass; FTI - fat tissue index; FVC - forced vital capacity; Hb - haemoglobin; HCT - haematocrit; HD - haemodialysis; HDL - high-density lipoprotein; IL - Interleukin; iPTH - intact parathyroid hormone; HR - heart rate; IQR - interquartile range; LBM - lean body mass; LDL - low-density lipoprotein; LTI - lean tissue index; LVEF - left ventricular ejection fraction; LVMI - left ventricular mass index; MAP - mean arterial pressure; MI - myocardial infarction; MRI - magnetic resonance imaging; NSRI - North Staffordshire Royal Infirmary; NYHA - New York Heart Association; PD - peritoneal dialysis; PEW - protein-energy wasting; (HR)QoL - (health-related) quality of life; RCT- randomised controlled trial; RLS - restless leg syndrome; RPE - rating of perceived exertion; SBP - systolic blood pressure; SCr - serum creatinine; SD - standard deviation; SDNN - standard deviation of normal to normal R-R intervals; TUG - timed up-and-go; URR - urea reduction ratio; VO₂ max - maximum rate of oxygen consumption; WBC - white blood cell

Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion	
Al-Ali 2018a	Wrong comparator: control group also performing exercise	
Aliasgharpour 2016	Wrong intervention: intervention was stretching only	
Alvares 2017	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Bogataj 2020	Wrong comparator: control group also performing exercise	
Bohm 2014	Wrong comparator: control group also performing exercise	
Bohm 2017	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Brown 2018	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Campos 2018	Wrong intervention: intervention was not exercise training	
Castellino 1987	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Chagolla 2018	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
CTRI/2018/02/012021	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
De Villar 2016	Wrong comparator: control group also performing exercise	
Dias 2020	No control group not performing exercise; comparing exercise with and without blood flow restriction	
Dungey 2013	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Dungey 2015	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Dziubek 2016	Wrong comparator: control group also performing exercise	
Fontsere 2016	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Frih 2017	Wrong intervention: intervention was not exercise training	
Frih 2018	Wrong comparator: control group also performing exercise	



Study	Reason for exclusion	
Fuhro 2018	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Garcia Testal 2019	Wrong comparator: control group also performing exercise	
Giannaki 2015	Wrong intervention: intervention was not exercise training	
Hamad 2016	Wrong comparator: control group also performing exercise	
Jeong 2018	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Kirkman 2013	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Krase 2020	Wrong intervention duration: intervention for only 180 min; study aimed to investigate the thermoregulatory responses of cold dialysis and exercise	
Maheshwari 2012	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Majchrzak 2008	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Miura 2016	Wrong population: not chronic HD or PD	
Molsted 2013	Wrong intervention: intervention was not exercise training	
Mora 2007	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Moug 2004	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Orcy 2012	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Orcy 2014	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Pinto 2015	Wrong comparator: control group also performing exercise	
Ribeiro 2019	Wrong intervention duration: less than 8 weeks (during HD session)	
Rossum 2019	Wrong intervention duration: less than 8 weeks (4 weeks)	
Stray-Gundersen 2016	Wrong intervention: intervention was not exercise training	
Sun 2002	Wrong intervention duration: lasted less than 8 weeks (acute effects of exercise)	
Tao 2015	Wrong comparator: control group also performing exercise	
Vrakas 2017	Co-intervention other than exercise that was not offered to the control group	

HD - haemodialysis; PD - peritoneal dialysis

Characteristics of studies awaiting classification [ordered by study ID]

Assawasaksakul 2018

Methods	Study design: parallel RCT
	Study duration: unclearStudy follow-up period: unclear



Assawasaksakul 2018 (Continued)	
Participants	 Country: unclear Setting: HD unit Inclusion criteria: HD patients Number:12 (number per group not reported) Mean age ± SD: 53.1 ± 14.4 years Sex (M/F): not reported Dialysis vintage: not reported BMI: 23.23 ± 5.5 kg/m² Exclusion criteria: not reported
Interventions	 Intradialytic exercise group Trained with the customized exercise program to exercise, initiated by a multidisciplinary team, on a cycle ergometer within the first hour of HD Physical activity was measured in terms of the number of daily steps counted by a wrist-worn wearable triaxial accelerometer Control group Not reported
Outcomes	 Muscle mass Physical activity Hb Albumin Phosphate
Notes	Abstract-only publication

Bennett 2019

Methods	 Study design: parallel RCT Study duration: not reported Study follow-up period: not reported
Participants	 Country: USA Setting: single centre Inclusion criteria: PD patients Number (randomised/analysed): exercise group (18/13); control group (18/13) Age: not reported Sex: not reported Dialysis vintage: not reported BMI: not reported Exclusion criteria: not reported
Interventions	 Exercise group Monthly exercise physiologist consultation; exercise prescription (resistance and aerobic exercise program using exercise bands) and four phone calls over 12 weeks Control group Normal care



Bennett 2019 (Continued)

Οı.	itcor	nes

- · Physical function measured
- QoL
- · Adverse events

Notes

· Abstract-only publication

Dong 2019

Methods

- Study design: parallel RCT
- Study duration: May 2017 to July 2017
- Study follow-up period: 12 weeks

Participants

- · Country: China
- · Setting: HD unit
- Inclusion criteria: HD patients with sarcopenia; aged 18 to 80 years; stable dialysis time ≥ 3 months; no central system disease; can walk independently, no physical disability, muscle strength ≥ III; can communicate normally
- Number (randomised/analysed): exercise group (23/21); control group (22/20)
- Mean age, range (years): exercise group (59, 32.5 to 66.5); control group (62.5, 50.5 to 70.0)
- Sex (M/F): exercise group (9/12); control group (12/8)
- BMI: exercise group (19.96 ± 308); control group (20.49 ± 3.41)
- Exclusion criteria: pregnant woman; 3 months of bleeding or infection records; cannot perform BIA test, such as cardiovascular stent implantation, pacemaker installation, artificial joint replacement or amputation surgery; had other serious complications such as heart failure, serious infection, malignant tumours; patients with cognitive impairment and mental illness

Interventions

Exercise group

"In the first week, the ankle weight was 0 kg, and quadriceps training board was used to assist the patient in low intensity resistance training. According to the patient's tolerance, the ankle weight of + 0.5 kg (single foot) per week until it was +5 kg (one foot), with the angle of the training board reduced gradually (150°-90°) until it was removed. In the meantime, the untreated hand was holding the elastic ball for 10 × 10 performing each step of the upper limb resistance exercise. During the exercise, patients performed a 5-min warm-up followed by a 1-2 h bout of intradialytic resistance exercise: for the one-leg raise-and-down exercise, and upper limb bouncing ball movement which exerted pressure on the elastic ball and maximally maintained for 3-5 s for one cycle and then release it, both complete 10 × 10 cycles repeatedly"

Control group

· Usual care

Outcomes

- · Maximum grip strength
- · Daily pace
- Physical activity level
- Hb
- SCr
- Kt/V
- Albumin
- Protein decomposition rate
- CRP
- IL-6
- IL-10



Dong 2019 (Continued)

Notes	Funding source: nil

IMPCT 2020

Methods	 Study design: parallel block RCT; randomised by sex, race and dialysis centre Study duration: unclear Study follow-up period: unclear
Participants	 Country: USA Setting: multicentre (13 sites) Inclusion criteria: adults ≥ 18 years; ESKD patients undergoing HD 2 to 3 times/week; within 3 months to 3 years of initiating HD Number: 200, ~ 50 per group Exclusion criteria: pregnancy; angina pectoris; chronic lung disease requiring oxygen; musculoskeletal conditions; amputation; orthopaedic disorders exacerbated by physical activity; a femoral AV access; legally blind; hepatitis B infection requiring medical isolation, or current incarceration; inability to recognize numbers and letters
Interventions	 Exercise training Cognitive training Exercise + cognitive training Standard care
Outcomes	 Change in executive function Secondary measures of cognitive function ESKD-specific clinical functions (physical function, falls, hospitalisation, death, return to work) Patient-centred outcomes (e.g. HRQoL measures)
Notes	Computer-based allocation system

Lopes 2019	
Methods	 Study design: parallel RCT (3 arms) Study duration: unclear Study follow-up period: 12 weeks
Participants	 Country: Brazil Setting: unclear Inclusion criteria: HD patients aged 30 to 75 years, HD for at least 3 months, AV fistula. Kt/V ≥ 1.2; no mobility issues; medical consent from a nephrologist Number: ML group (16); HL group (14); control group (20) Mean age ± SD (years): ML group (56.2 ± 12.5); HL group (48.1 ± 10.8); control group (56.9 ± 12.4) Sex (M/F): ML group (9/7); HL group (8/6); control group (13/7) Mean dialysis vintage ± SD (months): ML group (72.1 ± 50.3); HL group (45.7 ± 39.3); control group (53.2 ± 44.1) Mean BMI ± SD (kg/m²): ML group (25.5 ± 5.1); HL group (24.5 ± 4.7); control group (26.3 ± 3.7) Exclusion criteria: undergoing regular exercise program; physical disability or severe orthopaedic problems; the history of a stroke in the past 6 months; a recent hospitalisation (< 3 months); diagnosis of acquired immunodeficiency syndrome



Lopes 2019 (Continued)			
Interventions	 Resistance training Moderate-load intradialytic group (ML) High-load intradialytic group (HL) Stretching exercise (control) 		
Outcomes	 Body composition Functional capacity Inflammatory markers 		
Notes			
Maynard 2019			
Methods	 Study design: parallel RCT Study duration: unclear Study follow-up period: 12 weeks 		
Participants	 Country: Brazil Setting: private HD centre Inclusion criteria: sedentary adults (≥ 18 years); on HD by AV fistula 3 times/week for at least 3 months Number (randomised/analysed): treatment group (22/20); control group (23/20) Mean age ± SD (years): treatment group (49 ± 15.2); control group (43.9 ± 11.7) Sex: treatment group (12/8); control group (10/10) Mean dialysis vintage ± SD (months): treatment group (62.7 ± 34.2); control group (55.95 ± 38.87) Mean BMI ± SD (kg/m²): treatment group (25.5 ± 5); control group (24.5 ± 4.5) Exclusion criteria: haemodynamic instability; diagnosed respiratory disorder; visual impairment, or musculoskeletal and/or neurological limitations that compromised the ability to perform the proposed exercises; absence from two consecutive sessions; withdrawal; or death were excluded from the final analysis 		
Interventions	 Wii Sports (2006) and Wii Fit Plus (2009) Control group 		
Outcomes	 HRQoL (KDQOL) Physical function Mental health Clinical parameters 		
Notes			
PEDAL 2021			
Methods	 Study design: parallel RCT Study duration: unclear Study follow-up period: not reported 		
Participants	Country: UKSetting: dialysis centres (10 sites)		



PEDAL 2021 (Continued)	
	 Inclusion criteria: adult patients > 18 years, treated as outpatients, undergoing in-centre (hospital unit, satellite unit) maintenance HD > 3 months
	Number: ~115 per group
	 Age: > 18 years

• Sex: males and females Exclusion criteria: expected survival on dialysis < 6 months; dialysis withdrawal was being considered; likely to receive a live-donor transplant or transfer to PD in the period of time; patients deemed to be clinically unstable by their treating physician; bilateral lower-limb amputations; dementia or severe cognitive impairment; unable to give informed consent; psychiatric disorders;

	pregnant			_	
Interventions	Resistance training w	vith progre	ession		

• Change in KDQOL-SF 1.3 physical capacity score (disease-specific QOL measure)

• Peak aerobic capacity

• Physical performance tests Anthropometric measures

Cardiovascular risk • Physical function questionnaires

 Biochemistry Medication

• Usual care

Safety of intervention

Notes · Protocol only

Outcomes

Methods	Study design: parallel RCT
Metrious	
	•
	Study follow-up period: 4 months
Participants	Country: Brazil
	Setting: HD centre
	 Inclusion criteria: > 18 years; maintenance HD for at least 3 months; stable medication; did not present contraindications for physical exercises
	 Number: intervention group (15); control group (15)
	 Mean age ± SD (years): intervention group (50 ± 17.2); control group (58 ± 15.0)
	 Sex (M/F): intervention group (7/8); control group (8/7)
	 Mean dialysis vintage ± SD (months): intervention group (26.0 ± 14.58); control group (21.0 ± 27.1)
	 Mean BMI ± SD: intervention group (25.7 ± 3.58); control group (26.7 ± 4.6)
	 Exclusion criteria: already physically active; previous diagnosis of coronary artery disease or a positive treadmill exercise test for coronary arterial disease; previous stroke; cancer; liver failure; infection inactivity; BP > 160 × 100 mm Hg at a treadmill test; inclusion in another concurrent trial
Interventions	Aerobic training
	Usual care
Outcomes	Physical activity
	Biochemistry
	 Cardiovascular outcomes



Stringuetta Belik 2018 (Continued)

Notes

AV - arteriovenous BIA - bioelectrical impedance analysis; BMI - body mass index; BP - blood pressure; CRP - C-reactive protein; ESKD - end-stage kidney disease; Hb - haemoglobin; HD - haemodialysis; HRQoL - health-related quality of life; Kt/V - dialysis capacity; M/F - male/female; PD - peritoneal dialysis; QoL - quality of life; RCT - randomised controlled trial; SCr - serum creatinine

Characteristics of ongoing studies [ordered by study ID]

Δc	T.) N 1	261	ደበበበ	7747	79

Study name	Evaluation of the effectiveness of home-based physical training in patients undergoing haemodialysis
Methods	 Study design: parallel RCT Study duration: unclear Study follow-up period: 26 weeks
Participants	 Country: Poland Setting: home Exclusion criteria: lack of logical contact with the patient
Interventions	Exercise group
	 Type: aerobic Description: stationary ergometric bicycle training Duration: 30 to 35 minutes Warm-up: 5 minutes Cool-down: 5 minutes Frequency: 3 times/week on non-dialysis days Intensity: 40% to 60% HR Supervised: partially Control group
	Not reported, assumed usual care
Outcomes	Exercise toleranceFunctional fitnessQoL
Starting date	10th August 2015
Contact information	
Notes	Trial registration information only

Cardoso 2019

Study name	Effects of continuous moderate exercise with partial blood flow restriction during hemodialysis: a protocol for a randomized clinical trial
Methods	Study design: parallel RCT (3 arms)Study duration: unclear



Cardoso 2019 (Continued)	Study follow-up period: 13 weeks
Participants	 Country: Brazil Setting: HD unit Exclusion criteria: diagnosis of coronary artery disease, presence of active infection or cancer; presence of musculoskeletal limitations preventing exercise performance; cognitive alterations making it impossible to understand the instructions of the exercises; SBP > 180 mm Hg or DBP > 105 mm Hg at rest; resting HR > 120 bpm
Interventions	Exercise group
	 Type: aerobic Description: stationary bicycle performed in the first 2 hours of HD Duration: min Warm-up: min Cool-down: min Frequency: times/week Intensity: weeks 1 to 6: HR between 60% and 63% of HRmax or 10 to 11 in the perceived subjective exertion (ranges from 6 to 20) Weeks 5 to 8: HR between 64% and 76% of HRmax or 12 to 13 in the subjective perception of effort scale Supervised: unclear Control group
	Not reported, no exercise assumed
Outcomes	 IL-6 IL-10 CRP Femoral quadriceps muscle thickness Catalase activity Superoxide dismutase activity Glutathione peroxidase activity Ankle-arm index Functional test Strength QoL
Starting date	Unknown
Contact information	rafaelorcy@gmail.com
Notes	Protocol published
Chan 2019	
Study name	A randomized controlled trial of exercise to prevent muscle mass and functional loss in elderly hemodialysis patients: rationale, study design, and baseline sample
Methods	 Study design: parallel RCT Study duration: unclear Study follow-up period: weeks



Chan 2019 (Continued)

_			•					
ν	а	rt	10	11	n	а	n	ts
	ч		1	. 1	ν	ч		w

- · Country: USA
- Setting: HD units
- Exclusion criteria: temporary vascular access; uncontrolled DM; active autoimmune disease; malignancy, severe obesity (BMI > 35); alcoholism or other recreational drug use; unstable cardiac disease (abnormal exercise test, angina, uncontrolled arrhythmias or MI within 3 months); peripheral vascular disease (claudication with exercise); medically unstable; currently active (> 2 hours/week of moderate-intensity exercise); have received anabolic, catabolic or cytotoxic medications in the past 3 months

Interventions

Exercise group

- · Type: aerobic and resistance
- Description: 12-week individualized exercise program combining supervised and home-based monitored exercise
- · Duration: 45 minutes
- Frequency: 7 times/week
- Intensity: 70% to 80% of HR reserve and 12 to 14 on the Borg perceived exertion scale
- · Supervised: partially

Outcomes

- VO₂ max
- · Chair raise test
- 6MWT
- Handgrip strength
- · Body composition
- HRQoL: measured using SF-36
- · Beeson cognitive test

Starting date

Contact	information

knchan@stanford.edu

Notes

Clarkson 2017

Study na	me
----------	----

Efficacy of blood flow restriction exercise during dialysis for end stage kidney disease patients: protocol of a randomised controlled trial

Methods

- Study design: parallel RCT (3 arms)
- · Study duration: unclear
- Study follow-up period: 12 Weeks

Participants

- Country: Australia
- Setting: HD Unit
- Exclusion criteria: do not understand English and are unable to complete or comprehend the surveys or study documents; within the previous 12 weeks they have participated in regular physical activity or sport (> 150 min/week) of moderate or greater intensity, or structured resistance training (> 1 session/week)
- symptomatic peripheral vascular disease; limb ischaemia; untreated symptomatic cardiovascular disease
- any other absolute contraindications to exercise training (such as musculoskeletal factors or neurological conditions) that may affect their ability to perform physical assessments or exercise training protocols in the present study; currently smokers; pregnancy; have required hospitalisa-



Clarkson 2017 (Continued)

tion for non-dialysis reasons in the 4 weeks prior to the study's commencement; also be deemed unable to exercise during individual dialysis sessions if they present with fluid overload (> 5% above dialysis base weight);, SBP > 180 mm Hg, DBP < 90 mm Hg

Interventions

Blood flow restriction group

- · Type: aerobic
- Description: stationary bicycle performed in the first 2 hours of HD
- Duration: 10 minutes of exercise, followed by 20 minutes of rest and a subsequent 10 minutes of exercise (20 minutes exercise in total)
- · Warm-up: unclear
- Cool-down: unclear
- · Frequency: unclear
- Intensity: 15 RPE, 60% of age-adjusted HRmax
- · Supervised: yes
- Co-Intervention: automated tourniquet system applied to patient thighs during exercise

Non-blood flow restriction group

- · Type: aerobic
- Description: stationary bicycle performed in the first 2 hours of HD
- · Duration: 20 minutes
- · Warm-up: unclear
- · Cool-down: unclear
- Frequency: unclear
- Intensity: 12 RPE, 50% of age-adjusted HRmax
- Supervised: yes
- · Co-Intervention: no

Control group

• Usual care, given exercise advice at end of study only

Outcomes

- · Lower limb muscle strength: 3RM
- · Sit-to-stand in 30 seconds
- TUG
- 6MWT
- Muscle cross-sectional area
- · Body composition
- Hb
- Albumin
- Potassium
- PTH
- Phosphate
- URR
- · Physical activity level
- · POS-S questionnaire for symptom-related QoL

Starting date

Contact information	stuart.warmington@deakin.edu.au	
Notes	Protocol Published	



NCT01721551	
Study name	Sleep and training aspects in dialysis fatigue - exercise intervention (StandFirm)
Methods	 Study design: parallel RCT Study duration: 32 months Study follow-up period: 39 Weeks
Participants	 Country: Greece Setting: unclear Exclusion criteria: unable to give informed consent; opportunistic infection in the last 3 months; malignancy; infection requiring intravenous antibiotics within 2 months prior to enrolment; myoskeletal contraindication to exercise; requirement for systemic anticoagulation; participating or participated in an investigational drug or medical device study within 30 days or 5 half-lives; pregnant or breastfeeding; female of childbearing potential who does not agree to remain abstinent or to use an acceptable contraceptive regimen; LDH > 300U/L; prolonged QT interval (as defined by QTc > 460 msec in males and > 470 msec in females) in screening ECG; known current alcohol or drug abuse; known or suspected hypersensitivity to the study medication or any of its ingredients
Interventions	 Exercise group Type: aerobic Description: recumbent cycle training during dialysis session Duration: 45 to 60 minutes Warm-up: unclear Cool-down: unclear Frequency: unclear Intensity: progressive from 30% to 40% of maximum exercise power to 60% to 70% of maximum exercise power Supervised: unclear Control group Assumed usual care, stated that patients will not participate in any type of systematic exercise training Co-Intervention No
Outcomes	FatigueBody compositionMuscle functionality
Starting date	November 2012
Contact information	
Notes	Trial Registry Document

6MWT - 6 minute walk test; BMI - body mass index; bpm - beats per minute; CRP - C-reactive protein; DBP - diastolic blood pressure; DM- diabetes mellitus; ECG - electrocardiograph; Hb - haemoglobin; HD - haemodialysis; HR - heart rate; IL - interleukin; MI - myocardial infarction; PTH - parathyroid hormone; (HR)QoL - (health-related) quality of life; RCT - randomised controlled trial; RPE - rating of perceived exertion; SBP - systolic blood pressure; TUG - timed up-and-go; URR - urea reduction ratio VO_2 max - maximum rate of oxygen consumption



DATA AND ANALYSES

Comparison 1. Any exercise versus control (no exercise/placebo exercise)

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.1 Death	1	296	Risk Ratio (M-H, Random, 95% CI)	0.95 [0.56, 1.62]
1.2 Fatigue	6		Std. Mean Difference (IV, Random, 95% CI)	Totals not selected
1.3 HRQoL: Summary component scores	17		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.3.1 Physical Component Score	17	656	Mean Difference (IV, Random, 95% CI)	-4.12 [-6.37, -1.88]
1.3.2 Mental Component Score	17	656	Mean Difference (IV, Random, 95% CI)	-2.53 [-5.47, 0.40]
1.4 HRQoL: Individual domains	20		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.4.1 Physical Function- ing	18	1040	Mean Difference (IV, Random, 95% CI)	-4.70 [-8.94, -0.47]
1.4.2 Role-physical	13	809	Mean Difference (IV, Random, 95% CI)	-3.75 [-13.73, 6.23]
1.4.3 Pain	15	872	Mean Difference (IV, Random, 95% CI)	-5.28 [-10.69, 0.12]
1.4.4 General health perceptions	14	834	Mean Difference (IV, Random, 95% CI)	-3.86 [-7.39, -0.33]
1.4.5 Emotional well-being	13	789	Mean Difference (IV, Random, 95% CI)	-4.24 [-8.00, -0.47]
1.4.6 Role-emotional	14	833	Mean Difference (IV, Random, 95% CI)	-8.08 [-11.26, -4.90]
1.4.7 Vitality	16	940	Mean Difference (IV, Random, 95% CI)	-4.47 [-8.15, -0.79]
1.4.8 Social function	15	851	Mean Difference (IV, Random, 95% CI)	-0.80 [-4.56, 2.96]
1.4.9 Symptoms	7	533	Mean Difference (IV, Random, 95% CI)	-6.07 [-12.07, -0.08]
1.4.10 Effects of kidney disease	5	409	Mean Difference (IV, Random, 95% CI)	-4.01 [-6.47, -1.55]
1.4.11 Burden of kidney disease	5	409	Mean Difference (IV, Random, 95% CI)	-0.06 [-2.64, 2.51]
1.4.12 Work status	4	362	Mean Difference (IV, Random, 95% CI)	-0.36 [-3.75, 3.03]
1.4.13 Cognitive function	5	409	Mean Difference (IV, Random, 95% CI)	-2.66 [-7.57, 2.25]
1.4.14 Quality of social interactions	5	409	Mean Difference (IV, Random, 95% CI)	-4.92 [-8.32, -1.51]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.4.15 Sexual function	4	362	Mean Difference (IV, Random, 95% CI)	-3.60 [-11.16, 3.96]
1.4.16 Sleep	6	437	Mean Difference (IV, Random, 95% CI)	-6.58 [-12.57, -0.60]
1.4.17 Social support	5	409	Mean Difference (IV, Random, 95% CI)	-3.98 [-7.07, -0.89]
1.4.18 Dialysis staff encouragement	5	409	Mean Difference (IV, Random, 95% CI)	-3.75 [-8.40, 0.90]
1.4.19 Patient satisfaction	5	409	Mean Difference (IV, Random, 95% CI)	-4.58 [-10.23, 1.06]
1.5 Depression	10	441	Std. Mean Difference (IV, Random, 95% CI)	0.65 [0.22, 1.07]
1.5.1 4 months or less	6	311	Std. Mean Difference (IV, Random, 95% CI)	0.30 [-0.14, 0.74]
1.5.2 More than 4 months	4	130	Std. Mean Difference (IV, Random, 95% CI)	1.26 [0.72, 1.80]
1.6 6MWT	19	827	Mean Difference (IV, Random, 95% CI)	-49.91 [-62.59, -37.22]
1.7 Sit-To-Stand test [N reps/30 sec]	12	478	Mean Difference (IV, Random, 95% CI)	-2.36 [-2.98, -1.73]
1.8 Sit-To-Stand test [sit to 5 reps]	8	508	Mean Difference (IV, Random, 95% CI)	1.74 [1.22, 2.25]
1.9 Systolic blood pressure	20		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.9.1 Aerobic	13	394	Mean Difference (IV, Random, 95% CI)	3.99 [-1.80, 9.78]
1.9.2 Combined aerobic and resistance	7	282	Mean Difference (IV, Random, 95% CI)	8.69 [3.69, 13.69]
1.9.3 Others	1	30	Mean Difference (IV, Random, 95% CI)	25.55 [14.95, 36.15]
1.10 Diastolic blood pressure	20		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.10.1 Aerobic	13	394	Mean Difference (IV, Random, 95% CI)	-0.72 [-3.69, 2.24]
1.10.2 Combined aerobic and resistance	7	282	Mean Difference (IV, Random, 95% CI)	4.45 [2.91, 5.98]
1.10.3 Others	1	30	Mean Difference (IV, Random, 95% CI)	13.42 [7.46, 19.38]
1.11 Aerobic capacity (VO max or peak)	14	407	Mean Difference (IV, Random, 95% CI)	-3.30 [-4.33, -2.28]
1.12 Albumin	23	767	Mean Difference (IV, Random, 95% CI)	-0.39 [-1.25, 0.47]
1.13 Blood lipids	12		Mean Difference (IV, Random, 95% CI)	Subtotals only



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.13.1 Total cholesterol [mmol/L]	12	439	Mean Difference (IV, Random, 95% CI)	0.22 [0.04, 0.39]
1.13.2 LDL cholesterol [mmol/L]	6	180	Mean Difference (IV, Random, 95% CI)	0.24 [-0.02, 0.51]
1.13.3 HDL cholesterol [mmol/L]	8	264	Mean Difference (IV, Random, 95% CI)	-0.07 [-0.18, 0.04]
1.13.4 Triglycerides [mmol/L]	8	264	Mean Difference (IV, Random, 95% CI)	0.09 [-0.25, 0.44]
1.14 Body composition	10		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.14.1 Fat mass [kg]	9	384	Mean Difference (IV, Random, 95% CI)	-0.04 [-0.10, 0.02]
1.14.2 Lean mass [kg]	7	313	Mean Difference (IV, Random, 95% CI)	-0.37 [-2.74, 1.99]
1.15 Body mass index	16	590	Mean Difference (IV, Random, 95% CI)	-0.12 [-0.55, 0.31]
1.16 Calcium	17	592	Mean Difference (IV, Random, 95% CI)	0.03 [-0.00, 0.06]
1.17 C-reactive protein	14	421	Mean Difference (IV, Random, 95% CI)	0.31 [-0.13, 0.74]
1.18 Dialysis adequacy: Kt/V	11	382	Mean Difference (IV, Random, 95% CI)	-0.08 [-0.16, 0.00]
1.19 Energy intake	7	316	Mean Difference (IV, Random, 95% CI)	-0.09 [-1.58, 1.40]
1.20 Haemoglobin	29	975	Mean Difference (IV, Random, 95% CI)	-0.06 [-0.18, 0.06]
1.21 Left ventricular ejection fraction	6	222	Mean Difference (IV, Random, 95% CI)	-1.45 [-3.60, 0.70]
1.22 Left ventricular mass index	6	215	Mean Difference (IV, Random, 95% CI)	-9.85 [-20.50, 0.80]
1.23 Maximum heart rate	8	275	Mean Difference (IV, Random, 95% CI)	-6.14 [-10.05, -2.24]
1.24 Muscular strength	16		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.24.1 Knee extension	8	316	Mean Difference (IV, Random, 95% CI)	-5.06 [-8.58, -1.54]
1.24.2 Handgrip	10	410	Mean Difference (IV, Random, 95% CI)	-4.16 [-6.61, -1.71]
1.25 Phosphate	20	672	Mean Difference (IV, Random, 95% CI)	0.05 [-0.07, 0.16]
1.26 Potassium	18	610	Mean Difference (IV, Random, 95% CI)	0.23 [-0.06, 0.51]
1.27 Protein intake	7	316	Mean Difference (IV, Random, 95% CI)	-0.02 [-0.10, 0.07]
1.28 Parathyroid hor- mone	5	129	Mean Difference (IV, Random, 95% CI)	0.39 [-10.90, 11.68]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.29 Resting heart rate	11	405	Mean Difference (IV, Random, 95% CI)	3.72 [1.89, 5.56]
1.30 Timed up-and-go test	6	285	Mean Difference (IV, Random, 95% CI)	1.63 [0.90, 2.36]

Analysis 1.1. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 1: Death

	Cont		Exerc			Risk Ratio	Risk Ratio
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Random, 95% CI
EXCITE 2014	22	145	24	151	100.0%	0.95 [0.56 , 1.62]	
Total (95% CI)		145		151	100.0%	0.95 [0.56 , 1.62]	
Total events:	22		24				
Heterogeneity: Not appl	icable						0.5 0.7 1 1.5 2
Test for overall effect: Z	= 0.17 (P =	0.86)					More with exercise More with control
Test for subgroup differen	ences: Not a	pplicable					

Analysis 1.2. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 2: Fatigue

		Control]	Exercise		Std. Mean Difference	Std. Mean	Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	IV, Random, 95% CI	IV, Rando	m, 95% CI	
Sheshadri 2020	2	1.6	26	2.3	1.6	27	-0.18 [-0.72 , 0.36	5] -	_	
Yurtkuran 2007	6.9	7.2	18	3.3	1.5	19	0.69 [0.02 , 1.39	5]		
Soliman 2015 (1)	29.75	5.19	12	14.44	5.29	18	2.84 [1.78 , 3.90)]		
Amini 2016	6.2	2.15	35	4.37	1.62	32	0.94 [0.44 , 1.45	5]	-	
Johansen 2006 (2)	8.95	4.71	33	7.07	4.78	35	0.39 [-0.09 , 0.87	7] .	+	
Chang 2010	45.5	19.66	35	41	20.09	36	0.22 [-0.24 , 0.69	-	+	
								-4 -2 (+ +	<u></u>
Footnotes								More with exercise	More with	•

⁽¹⁾ data has been verified

⁽²⁾ Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group



Analysis 1.3. Comparison 1: Any exercise versus control (no exercise/ placebo exercise), Outcome 3: HRQoL: Summary component scores

Study or Subgroup		Control]	Exercise			Mean Difference	Mean Difference
	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
1.3.1 Physical Componer	nt Score								
Giannaki 2013a	70.5	26.5	7	76.4	15.6	15	1.0%	-5.90 [-27.06, 15.26]	
ACTINUT 2013	59.87	21.37	9	84.7	13.32	7	1.5%	-24.83 [-41.93, -7.73]	
Koh 2009 (1)	55	25	15	51.5	22.71	30	1.9%	3.50 [-11.54 , 18.54]	
CHAIR 2015 (2)	33.58	18.05	11	51.6	7.41	6	2.7%	-18.02 [-30.22 , -5.82]	
Rosa 2018	74.43	18.07	24	72.02	20.36	28	3.5%	2.41 [-8.04 , 12.86]	
Molsted 2004 (3)	44.95	10.99	7	45.75	7.44	10	4.1%	-0.80 [-10.16, 8.56]	
Segura-Orti 2009	45.9	8.7	8	44.7	8.7	17	5.6%	1.20 [-6.11, 8.51]	
Song 2012a	64.2	12.2	20	72.5	9.8	20	6.0%	-8.30 [-15.16, -1.44]	
Chen 2010	50	11	22	54	12	22	6.1%	-4.00 [-10.80 , 2.80]	
DIALY-SIZE 2016 (4)	3.4	7.3	8	3.64	8.1	23	6.9%	-0.24 [-6.29 , 5.81]	
Samara 2016	43.9	8.8	12	49.9	6.6	15	7.0%	-6.00 [-12.00 , -0.00]	
Uchiyama 2019	38.2	9.2	23	41	8.1	24	8.3%	-2.80 [-7.76 , 2.16]	
Dobsak 2012	50.6	6.8	10	51.7	4.4	11	8.3%	-1.10 [-6.05 , 3.85]	
HOPE 2019	38.9	9.42	38	38.72	10.4	29	8.5%	0.18 [-4.65 , 5.01]	1
Suhardjono 2019 (5)	-3.45	12.45	38	6.52	8.32	73	9.1%	-9.97 [-14.36 , -5.58]	[
Ouzouni 2009	38.9	5.8	14	44.5	5.5	19	9.8%	-5.60 [-9.52 , -1.68]	
Frih 2017a	51	7	20	55.5	5.5	21	9.8%	-4.50 [-8.37 , -0.63]	
Subtotal (95% CI)	01	•	286	55.5	0.0	370	100.0%	-4.12 [-6.37 , -1.88]	<u> </u>
1.3.2 Mental Componen Giannaki 2013a		21.9	7	70.4	18 7	15	1 9%	-5 40 [-24 18 13 38]	_
Giannaki 2013a	65	21.9	7	70.4	18.7	15	1.9%	-5.40 [-24.18 , 13.38]	
Koh 2009 (1)	64	25	15	61.5	20.96	30	2.8%	2.50 [-12.21 , 17.21]	
ACTINUT 2013	52.07	16.11	9	74.3					
			,	74.3	10.61	7	3.2%	-22.23 [-35.37 , -9.09]	
Rosa 2018	76.08	19.15	24	78.02	10.61 16.44	7 28	3.2% 4.6%	-22.23 [-35.37 , -9.09] -1.94 [-11.73 , 7.85]	
Rosa 2018 Molsted 2004 (3)	76.08 51.55								
		19.15	24	78.02	16.44	28	4.6%	-1.94 [-11.73 , 7.85]	
Molsted 2004 (3)	51.55	19.15 10.26	24 7	78.02 54.1	16.44 6.47	28 10	4.6% 5.2%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04]	
Molsted 2004 (3) Song 2012a	51.55 60.8	19.15 10.26 12.4	24 7 20	78.02 54.1 69.4	16.44 6.47 13.7	28 10 20	4.6% 5.2% 5.4%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009	51.55 60.8 54.3	19.15 10.26 12.4 5.1	24 7 20 8	78.02 54.1 69.4 46.5	16.44 6.47 13.7 13.5	28 10 20 17	4.6% 5.2% 5.4% 5.9%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015	51.55 60.8 54.3 53.05	19.15 10.26 12.4 5.1 6.65	24 7 20 8 11	78.02 54.1 69.4 46.5 51.6	16.44 6.47 13.7 13.5 7.41	28 10 20 17 6	4.6% 5.2% 5.4% 5.9% 6.0%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016	51.55 60.8 54.3 53.05 39	19.15 10.26 12.4 5.1 6.65 10.4	24 7 20 8 11 12	78.02 54.1 69.4 46.5 51.6 53.3	16.44 6.47 13.7 13.5 7.41 6.9	28 10 20 17 6 15	4.6% 5.2% 5.4% 5.9% 6.0% 6.2%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4)	51.55 60.8 54.3 53.05 39 0.7	19.15 10.26 12.4 5.1 6.65 10.4 7.5	24 7 20 8 11 12 8	78.02 54.1 69.4 46.5 51.6 53.3 -2.36	16.44 6.47 13.7 13.5 7.41 6.9 8.75	28 10 20 17 6 15 23	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Duzouni 2009	51.55 60.8 54.3 53.05 39 0.7 40.1	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8	24 7 20 8 11 12 8	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8	16.44 6.47 13.7 13.5 7.41 6.9 8.75	28 10 20 17 6 15 23	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5% 6.8%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Duzouni 2009 IHOPE 2019	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63	24 7 20 8 11 12 8 14 38	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10	28 10 20 17 6 15 23 19	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5% 6.8% 7.1%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Duzouni 2009 IHOPE 2019 Chen 2010	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7 38	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63	24 7 20 8 11 12 8 14 38 22	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10	28 10 20 17 6 15 23 19 29	4.6% 5.2% 5.4% 5.9% 6.0% 6.5% 6.8% 7.1%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05] 1.00 [-4.32 , 6.32]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Duzouni 2009 IHOPE 2019 Chen 2010 Uchiyama 2019	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7 38 52.6	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63 9	24 7 20 8 11 12 8 14 38 22 23	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05 37 49.8	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10 10.81 9	28 10 20 17 6 15 23 19 29 22	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5% 6.8% 7.1% 7.1%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05] 1.00 [-4.32 , 6.32] 2.80 [-2.52 , 8.12]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Duzouni 2009 IHOPE 2019 Chen 2010 Uchiyama 2019 Dobsak 2012	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7 38 52.6 59.3	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63 9 9	24 7 20 8 11 12 8 14 38 22 23	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05 37 49.8 59.5	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10 10.81 9 9.6 5.5	28 10 20 17 6 15 23 19 29 22 24	4.6% 5.2% 5.4% 5.9% 6.0% 6.5% 6.8% 7.1% 7.19 7.5%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05] 1.00 [-4.32 , 6.32] 2.80 [-2.52 , 8.12] -0.20 [-4.96 , 4.56]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Ouzouni 2009 IHOPE 2019 Chen 2010 Uchiyama 2019 Dobsak 2012 Suhardjono 2019	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7 38 52.6 59.3 -0.56	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63 9 9 5.6 9.224	24 7 20 8 11 12 8 14 38 22 23 10	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05 37 49.8 59.5 2.67	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10 10.81 9 9.6 5.5 9.05	28 10 20 17 6 15 23 19 29 22 24 11 73	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5% 6.8% 7.1% 7.1% 7.5% 8.1% 8.6%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05] 1.00 [-4.32 , 6.32] 2.80 [-2.52 , 8.12] -0.20 [-4.96 , 4.56] -3.23 [-6.82 , 0.36] -8.50 [-11.11 , -5.89]	
Molsted 2004 (3) Song 2012a Segura-Orti 2009 CHAIR 2015 Samara 2016 DIALY-SIZE 2016 (4) Ouzouni 2009 IHOPE 2019 Chen 2010 Uchiyama 2019 Dobsak 2012 Suhardjono 2019 Frih 2017a	51.55 60.8 54.3 53.05 39 0.7 40.1 50.7 38 52.6 59.3 -0.56 42.5	19.15 10.26 12.4 5.1 6.65 10.4 7.5 6.8 11.63 9 9 5.6 9.224 4.5	24 7 20 8 11 12 8 14 38 22 23 10 38 20 286	78.02 54.1 69.4 46.5 51.6 53.3 -2.36 41.8 52.05 37 49.8 59.5 2.67 51	16.44 6.47 13.7 13.5 7.41 6.9 8.75 10 10.81 9 9.6 5.5 9.05	28 10 20 17 6 15 23 19 29 22 24 11 73 21	4.6% 5.2% 5.4% 5.9% 6.0% 6.2% 6.5% 6.8% 7.1% 7.1% 7.5% 8.1% 8.6%	-1.94 [-11.73 , 7.85] -2.55 [-11.14 , 6.04] -8.60 [-16.70 , -0.50] 7.80 [0.47 , 15.13] 1.45 [-5.66 , 8.56] -14.30 [-21.14 , -7.46] 3.06 [-3.25 , 9.37] -1.70 [-7.44 , 4.04] -1.35 [-6.75 , 4.05] 1.00 [-4.32 , 6.32] 2.80 [-2.52 , 8.12] -0.20 [-4.96 , 4.56] -3.23 [-6.82 , 0.36]	

- (1) two intervention arms pooled together in the exercise group
- (2) mean and standard deviation estimated from the median and range
- (3) mean and standard deviation estimated from the median and the range
- (4) three intervention arms pooled together in the exercise group
- (5) mean and standard deviation estimated from the median and range and both intervention arms pooled together



Analysis 1.4. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 4: HRQoL: Individual domains

Study or Subgroup	Mean	Control SD	Total	Mean	Exercise SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
1.4.1 Physical Functioning									
Parsons 2004	65.7	27.1	7	68.3	30.6	6	1.5%	-2.60 [-34.26, 29.06]	
Matsumoto 2007	48	36.0572	32	43	27.2293	17	3.6%	5.00 [-12.99, 22.99]	
Martins do Valle 2020	63.1	24.5	12	72.5	20.2	12	3.6%	-9.40 [-27.37 , 8.57]	
Molsted 2004 (1)	70	21.99	7	82.5	9.41	11	3.8%	-12.50 [-29.71 , 4.71]	
Zhao 2017	64.2	55.464	56	68.8	30.7879	59	4.0%	-4.60 [-21.11 , 11.91]	
Koh 2009 (2)	70	26	15	67.5	25.04	30	4.2%	2.50 [-13.42 , 18.42]	
Martin-Alemany 2016	59.7	26.4	19	71.3	22.3	17	4.2%	-11.60 [-27.51 , 4.31]	
AVANTE-HEMO 2020	84.44	22.7	13	76.79	14.02	21	5.0%	7.65 [-6.07 , 21.37]	- -
Johansen 2006 (3)	56.58	26.72	33	59.12	30.37	35	5.1%	-2.54 [-16.12 , 11.04]	<u> </u>
Sheshadri 2020	63.7	24.3	26	60.2	25.4	27	5.1%	3.50 [-9.88 , 16.88]	_
van Vilsteren 2005					28				-
	60.2	34.5	43	62.5		53	5.4%	-2.30 [-15.07 , 10.47]	-
Jong 2004	2.35	10.62	17	7.42	17.17	19	7.0%	-5.07 [-14.29 , 4.15]	
Abreu 2017	85	13	19	87	18	25	7.1%	-2.00 [-11.16 , 7.16]	+
Uchiyama 2019	73.2	13.9	23	76	15.7	24	7.4%	-2.80 [-11.27 , 5.67]	+
PEAK 2006	-1.8	17.6	25	7.6	11.8	24	7.5%	-9.40 [-17.76 , -1.04]	
Dobsak 2012	53.1	10	10	54.1	7.9	11	7.8%	-1.00 [-8.76 , 6.76]	+
EXCITE 2014	-2.7	27.4518	123	1.5	21.0824	104	8.5%	-4.20 [-10.52 , 2.12]	-
Wu 2014d	60.6	12.9	33	82.1	10	32	8.9%	-21.50 [-27.10 , -15.90]	+
Subtotal (95% CI)			513			527	100.0%	-4.70 [-8.94 , -0.47]	•
Heterogeneity: Tau ² = 44.10	; Chi ² = 43	3.01, df = 1	7 (P = 0.00)	05); $I^2 = 60$	0%				1
Test for overall effect: Z = 2	.18 (P = 0.	.03)							
1.4.2 Role-physical			_						
Molsted 2004 (1)	62.5	36.64	7	62.5	32.33	10	5.0%	0.00 [-33.74 , 33.74]	- + -
Parsons 2004	90.5	25.2	7	77.7	34.5	6	5.0%	12.80 [-20.52 , 46.12]	- •
Martin-Alemany 2016	68.8	41.2	19	65.6	40.7	17	6.3%	3.20 [-23.59 , 29.99]	
Koh 2009 (2)	48	44	15	37	39.82	30	6.3%	11.00 [-15.44 , 37.44]	
Matsumoto 2007	40	58.2462	32	44	35.0091	17	6.4%	-4.00 [-30.16 , 22.16]	
AVANTE-HEMO 2020	88.89	33.3	13	77.8	41.77	21	6.6%	11.09 [-14.34 , 36.52]	
Zhao 2017	54.4	65.3044	56	63.2	62.9545	59	7.0%	-8.80 [-32.26 , 14.66]	
van Vilsteren 2005	54.5	45.7	43	50	43	53	8.4%	4.50 [-13.41 , 22.41]	-
Abreu 2017	63	27	19	79	27	25	8.8%	-16.00 [-32.11 , 0.11]	
Uchiyama 2019	62.2	26.9	23	71.9	22	24	9.3%	-9.70 [-23.78, 4.38]	
EXCITE 2014	-9.2	53.2229	123	0.2	47.821	104	9.5%	-9.40 [-22.55, 3.75]	
Dobsak 2012	51.3	8.9	10	44.1	10.6	11	10.6%	7.20 [-1.15 , 15.55]	_
Wu 2014d	26.3	11.5	33	54.6	15.4	32	10.9%	-28.30 [-34.92 , -21.68]	_
Subtotal (95% CI)			400			409	100.0%	-3.75 [-13.73, 6.23]	
Heterogeneity: Tau ² = 227.1	3; Chi ² = 5	57.02, df =		0001): I ² =	79%			, 1	
Test for overall effect: $Z = 0$,	,,					
1.4.3 Pain		20.0	40	ED :	20.4	40	2.007	C CO [17.40	
Martins do Valle 2020	60	32.9	12	53.4	26.1	12	3.6%	6.60 [-17.16 , 30.36]	+
Parsons 2004	86.6	13.2	7	79.5	23.9	6	4.1%	7.10 [-14.38 , 28.58]	+-
Zhao 2017	52.7	54.6641	56	64.6	59.4176	59	4.3%	-11.90 [-32.75 , 8.95]	+
Koh 2009 (2)	57	31	15	62.5	29.98	30	4.8%	-5.50 [-24.51 , 13.51]	
Martin-Alemany 2016	65.4	34.7	19	77.3	21.2	17	5.0%	-11.90 [-30.47 , 6.67]	+
Matsumoto 2007	47	36.0572	32	46	23.3394	17	5.6%	1.00 [-15.71 , 17.71]	+
AVANTE-HEMO 2020	85	18.3	13	78.27	24.79	21	6.4%	6.73 [-7.81 , 21.27]	+-
Abreu 2017	82	23	19	85	19	25	7.2%	-3.00 [-15.74, 9.74]	-
Uchiyama 2019	67.5	24.4	23	73	19.1	24	7.3%	-5.50 [-18.06, 7.06]	<u>-</u>
Molsted 2004 (1)	82.5	13.92	7	90.5	11.93	11	7.3%	-8.00 [-20.49 , 4.49]	
Pellizzaro 2013	-15.5	16.7741	14	24	16.7741	14	7.3%	-39.50 [-51.93 , -27.07]	
van Vilsteren 2005	76.1	25.5	43	76.9	21	53	8.7%	-0.80 [-10.29 , 8.69]	
Dobsak 2012	55.7	10.7	10	57.6	10.9	11	8.8%	-1.90 [-11.15 , 7.35]	Ţ
EXCITE 2014	-3.2	34.1747	123	-1.1	30.8523	104	9.2%	-2.10 [-10.56 , 6.36]	†
Wu 2014d	-3.2 59	12.7	43	63	13.4	32	10.4%	-4.00 [-10.00 , 2.00]	<u>†</u>
	59	12./		0.3	13.4				₹
Subtotal (95% CI) Heterogeneity: Tau ² = 63.78	Cl 12		436	06) 12 -	20/	436	100.0%	-5.28 [-10.69 , 0.12]	♥



Analysis 1.4. (Continued)

1.4.4 General health perce	otions								
Molsted 2004 (1)	59	30.05	7	58.5	25.11	11	1.6%	0.50 [-26.25, 27.25]	
Parsons 2004	50.1	22.4	7	50.7	22.7	6	1.9%	-0.60 [-25.20 , 24.00]	
Zhao 2017	52.2	50.9911	56	65.1	62.0354	59	2.6%	-12.90 [-33.61 , 7.81]	
Koh 2009 (2)	48	27	15	39	23.42	30	4.0%	9.00 [-7.03, 25.03]	
Matsumoto 2007	44	33.2836	32	43	17.5045	17	4.9%	1.00 [-13.22 , 15.22]	
Martins do Valle 2020	50.5	13.3	12	52.7	19.7	12	5.3%	-2.20 [-15.65, 11.25]	
AVANTE-HEMO 2020	53.33	17.5	13	53.73	17.76	21	6.1%	-0.40 [-12.57, 11.77]	
Abreu 2017	71	21	19	78	17	25	6.6%	-7.00 [-18.56 , 4.56]	
Martin-Alemany 2016	51	14.1	19	44	17.9	17	7.4%	7.00 [-3.61 , 17.61]	
Uchiyama 2019	45.7	17.4	23	43.7	17.9	24	8.0%	2.00 [-8.09, 12.09]	
Dobsak 2012	42.5	9	10	50.9	8.7	11	11.1%	-8.40 [-15.99 , -0.81]	
van Vilsteren 2005	45.2	18.1	43	51.8	15.9	53	12.2%	-6.60 [-13.50 , 0.30]	_
Wu 2014d	34.6	9.3	33	48.1	15.8	32	13.1%	-13.50 [-19.83 , -7.17]	_
EXCITE 2014	-2.5	20.1687	123	0.8	19.5398	104	15.2%	-3.30 [-8.48 , 1.88]	
Subtotal (95% CI)			412			422		-3.86 [-7.39, -0.33]	
Heterogeneity: Tau ² = 14.33	Chi ² = 20	32 df = 13 (I ² = 36%			100.070	5100 [7155 ; 6155]	V
Test for overall effect: $Z = 2$			1 0.00),						
1.4.5 Emotional well-being Zhao 2017	55.3	61.2787	56	60.9	61.5759	59	2.5%	-5.60 [-28.06 , 16.86]	
Parsons 2004	84.3	16.9	- 50 - 7	80.7	19.8	6	3.0%	3.60 [-16.59 , 23.79]	
	84.3 70		12	80.7 65	19.8 29.6	12	3.0%		+
Martins do Valle 2020		16.7			29.6			5.00 [-14.23 , 24.23]	+
Matsumoto 2007	52 75 11	30.5099	32	54 76 09		17	4.9%	-2.00 [-17.32 , 13.32]	+
AVANTE-HEMO 2020	75.11 76	24.7	13 7	76.98	11.65	21	5.4%	-1.87 [-16.19 , 12.45]	+
Molsted 2004 (1)	76	17.59		84 76.0	10.04	11	5.4%	-8.00 [-22.32 , 6.32]	
Martin-Alemany 2016	65.6	17.3	19	76.8	19.4	17	7.0%	-11.20 [-23.26 , 0.86]	
Abreu 2017	78	22	19	86 CF	15	25	7.5%	-8.00 [-19.51 , 3.51]	
Dobsak 2012	63.4	14.9	10	65	9.2	11	8.3%	-1.60 [-12.32 , 9.12]	+
Uchiyama 2019	73.2	17.6	23	71.5	18.8	24	8.6%	1.70 [-8.71 , 12.11]	+
van Vilsteren 2005	79.4	15	43	76.2	18.9	53	13.8%	3.20 [-3.58 , 9.98]	+
Wu 2014d	54.2	14.1	33	68.2	12.8	32	14.3%	-14.00 [-20.54 , -7.46]	-
EXCITE 2014	-3.9	24.0904	123	1.2	19.5398	104	16.0%	-5.10 [-10.78 , 0.58]	-
Subtotal (95% CI)	Ch:2 - 10	20 36 - 12 6	397	T2 — D.40/		392	100.0%	-4.24 [-8.00 , -0.47]	•
Heterogeneity: $Tau^2 = 14.68$ Test for overall effect: $Z = 2$			P – 0.11);	1 34%					
44601 1									
1.4.6 Role-emotional Parsons 2004	71.4	30.4	7	50	44.7	6	0.6%	21.40 [-20.87 , 63.67]	
Martins do Valle 2020	54.1	43.3	12	75	38.8	12			
			7				0.9%	-20.90 [-53.80 , 12.00]	
Molsted 2004 (1)	75 60	36.64		83.33	21.55	10	1.1%	-8.33 [-38.58 , 21.92]	
Koh 2009 (2)	69 54.1	41	15 50	74	41.43	30	1.6%	-5.00 [-30.50 , 20.50]	
Zhao 2017	54.1	52.7802	56	61.8	70.3068	59	2.0%	-7.70 [-30.35 , 14.95]	
Matsumoto 2007	47	52.699	32	50	27.2293	17	2.0%	-3.00 [-25.38 , 19.38]	-
Abreu 2017	76	35	19	76	38	25	2.2%	0.00 [-21.67 , 21.67]	+
Martin-Alemany 2016	73.4	33.3	19	85.4	29.7	17	2.4%	-12.00 [-32.58 , 8.58]	+
van Vilsteren 2005	70.2	41.9	43	78.8	35	53	4.1%	-8.60 [-24.27 , 7.07]	-+
Uchiyama 2019	64.3	31.8	23	77.5	19.4	24	4.4%	-13.20 [-28.34 , 1.94]	
EXCITE 2014	-7.5	57.1446	123	-1.8	51.9278	104	5.0%	-5.70 [-19.90 , 8.50]	 +
AVANTE-HEMO 2020	92.59	22.2	13	95.92	11.32	21	6.0%	-3.33 [-16.33 , 9.67]	-
Dobsak 2012	57	14.7	10	59.3	10.9	11	8.1%	-2.30 [-13.46 , 8.86]	+
Wu 2014d	30.4	7.4	33	40	9.4	32	59.6%	-9.60 [-13.72 , -5.48]	
Subtotal (95% CI)			412			421	100.0%	-8.08 [-11.26 , -4.90]	•
Heterogeneity: Tau ² = 0.00;	$Chi^2 = 6.00$	o, df = 13 (P =	= 0.95); I ²	= 0%					*
Test for overall effect: $Z = 4$.98 (P < 0.	00001)							
1.4.7 Vitality									
Parsons 2004	62.9	14.1	7	46.7	30.3	6	1.7%	16.20 [-10.20 , 42.60]	
Molsted 2004 (1)	69	29.32	7	71.25	17.26	11	2.1%	-2.25 [-26.25 , 21.75]	
Pellizzaro 2013			14			14	2.1%		
	-10 50.4	28.5212		17.5 57.2	28.5212			-27.50 [-48.63 , -6.37]	
7haa 2017		58.47	56	57.2	44.2067	59	3.1%	-6.80 [-25.82 , 12.22]	
Zhao 2017			20		22 220 4	4.77	4.007	E 00 F 20 22 40 223	
Zhao 2017 Matsumoto 2007 Martin-Alemany 2016	47 68.1	30.5099 20	32 19	52 57.2	23.3394 26	17 17	4.3% 4.3%	-5.00 [-20.32 , 10.32] 10.90 [-4.38 , 26.18]	-



Analysis 1.4. (Continued)

Matsumoto 2007	47	30.5099	32	52	23.3394	17	4.3%	-5.00 [-20.32 , 10.32]	
Martin-Alemany 2016	68.1	20	19	57.2	26	17	4.3%	10.90 [-4.38 , 26.18]	
Koh 2009 (2)	52	23	15	51	23.31	30	4.8%	1.00 [-13.32 , 15.32]	+
AVANTE-HEMO 2020	72.78	22.6	13	58.22	15.91	21	4.9%	14.56 [0.52 , 28.60]	
Sheshadri 2020	52.7	26.3	26	55.6	25.8	27	4.9%	-2.90 [-16.93 , 11.13]	
Abreu 2017	67	21	19	74	22	25	5.6%	-7.00 [-19.79 , 5.79]	
Uchiyama 2019	54.8	20.3	23	57.5	20.3	24	6.3%	-2.70 [-14.31 , 8.91]	+
Dobsak 2012	50.2	13.3	10	52.2	5.1	11	8.6%	-2.00 [-10.78 , 6.78]	+
PEAK 2006	-7 56.1	14.1	25	2.8	16.3	24	8.9%	-9.80 [-18.35 , -1.25]	-
van Vilsteren 2005	56.1	17.4	43	66.1	15.3	53	11.0%	-10.00 [-16.63 , -3.37]	-
EXCITE 2014 Wu 2014d	-3.7	25.2109	123	8.0	13.3693	104 32	12.8%	-4.50 [-9.64 , 0.64]	=
Subtotal (95% CI)	42.5	7.8	33 465	52.3	8.5	475	14.2% 100.0%	-9.80 [-13.77 , -5.83] - 4.47 [-8.15 , -0.79]	-
Heterogeneity: Tau ² = 20.7	'5· Chi² = 27	95 df = 15		I2 = 46%		4/3	100.0 /0	-4.47 [-0.13 , -0.73]	▼
Test for overall effect: Z =			(1 0.02),	1 4070					
1.4.8 Social function									
Parsons 2004	80.3	20.3	7	77.1	35.7	6	1.3%	3.20 [-29.08, 35.48]	
Martins do Valle 2020	64	35.6	12	79.7	29.8	12	1.9%	-15.70 [-41.97 , 10.57]	
Zhao 2017	53.7	66.1989	56	66.4	53.764	59	2.6%	-12.70 [-34.81 , 9.41]	+
Koh 2009 (2)	73	30	15	68.5	27.42	30	3.7%	4.50 [-13.58 , 22.58]	-
Martin-Alemany 2016	76.4	25.5	19	79.8	29	17	3.8%	-3.40 [-21.33 , 14.53]	-
CHAIR 2015 (4)	41.68	14.78	11	30.01	17.5	6	4.4%	11.67 [-4.83 , 28.17]	+-
AVANTE-HEMO 2020	90.28	19.5	13	90.44	21	21	5.8%	-0.16 [-14.05 , 13.73]	+
Abreu 2017	76	26	19	91	19	25	5.8%	-15.00 [-28.86 , -1.14]	
Dobsak 2012	66.5	17.6	10	61.5	14	11	5.9%	5.00 [-8.69 , 18.69]	+
Uchiyama 2019	74.3	26	23	71.8	19.6	24	6.3%	2.50 [-10.71 , 15.71]	_
Molsted 2004 (1)	90.63	13.74 13.8681	7	90.63	11.77	11	7.0%	0.00 [-12.33 , 12.33]	+
Matsumoto 2007	59	13.0001	32	52	21.3944	17	8.0%	7.00 [-4.25 , 18.25]	 -
Mileteren 2005	74.1	25		71 C	10				
	74.1	25	43	71.6	19	53 104	10.7%	2.50 [-6.56 , 11.56]	†
van Vilsteren 2005 EXCITE 2014	-0.8	22.9699	123	-2.5	29.8239	104	14.2%	1.70 [-5.32 , 8.72]	+
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau ² = 13.1	-0.8 37 2; Chi² = 19	22.9699 8.9 .17, df = 14	123 33 423	-2.5 44.5	29.8239 11.7				‡ •
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z =	-0.8 37 2; Chi² = 19	22.9699 8.9 .17, df = 14	123 33 423	-2.5 44.5	29.8239 11.7	104 32	14.2% 18.6%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44]	•
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms	-0.8 37 2; Chi ² = 19 0.42 (P = 0.4	22.9699 8.9 3.17, df = 14 68)	123 33 423 (P = 0.16);	-2.5 44.5 I ² = 27%	29.8239 11.7	104 32 428	14.2% 18.6% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z =	-0.8 37 2; Chi² = 19	22.9699 8.9 .17, df = 14	123 33 423	-2.5 44.5	29.8239 11.7 14.8	104 32	14.2% 18.6% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96]	-
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42)	22.9699 8.9 .17, df = 14 68)	123 33 423 (P = 0.16);	-2.5 44.5 I ² = 27%	29.8239 11.7	104 32 428	14.2% 18.6% 100.0%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] -0.80 [-4.56 , 2.96] -6.50 [-16.76 , 3.76] -13.50 [-23.30 , -3.70]	-
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013	-0.8 37 2; Chi² = 19 0.42 (P = 0.	22.9699 8.9 .17, df = 14 68) 16.6 13.2288	123 33 423 (P = 0.16); 19 14	-2.5 44.5 I ² = 27% 76.6 13.5	29.8239 11.7 14.8 13.2288	104 32 428 17 14	14.2% 18.6% 100.0% 11.8% 12.2%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020	-0.8 37 2; Chi ² = 19 0.42 (P = 0. 70.1 0 76.87	22.9699 8.9 .17, df = 14 68) 16.6 13.2288 13.4	123 33 423 (P = 0.16); 19 14 13	-2.5 44.5 $I^{2} = 27\%$ 76.6 13.5 82.12	29.8239 11.7 14.8 13.2288 11.85	104 32 428 17 14 21	14.2% 18.6% 100.0% 11.8% 12.2% 12.9%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] -0.80 [-4.56 , 2.96] -6.50 [-16.76 , 3.76] -13.50 [-23.30 , -3.70] -5.25 [-14.12 , 3.62]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d	-0.8 37 2; Chi ² = 19 0.42 (P = 0. 70.1 0 76.87 78.7	22.9699 8.9 8.17, df = 144668) 16.6 13.2288 13.4 15.2	123 33 423 (P = 0.16); 19 14 13 23	-2.5 44.5 $I^2 = 27\%$ 76.6 13.5 82.12 79.5	29.8239 11.7 14.8 13.2288 11.85 11.4	104 32 428 17 14 21 24	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] -0.80 [-4.56 , 2.96] -6.50 [-16.76 , 3.76] -13.50 [-23.30 , -3.70] -5.25 [-14.12 , 3.62] -0.80 [-8.51 , 6.91]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005	-0.8 37 2; Chi ² = 19 0.42 (P = 0. 70.1 0 76.87 78.7 43.5	22.9699 8.9 8.17, df = 144668) 16.6 13.2288 13.4 15.2 8.8	123 33 423 (P = 0.16); 19 14 13 23 33	-2.5 44.5 $I^2 = 27\%$ 76.6 13.5 82.12 79.5 62.2	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6	104 32 428 17 14 21 24 32	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] -0.80 [-4.56 , 2.96] -6.50 [-16.76 , 3.76] -13.50 [-23.30 , -3.70] -5.25 [-14.12 , 3.62] -0.80 [-8.51 , 6.91] -18.70 [-24.29 , -13.11]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019	-0.8 37 2; Chi ² = 19 0.42 (P = 0. 70.1 0 76.87 78.7 43.5 23.9	22.9699 8.9 8.17, df = 144668) 16.6 13.2288 13.4 15.2 8.8 9.5	123 33 423 (P = 0.16); 19 14 13 23 33 43	-2.5 44.5 $I^2 = 27\%$ 76.6 13.5 82.12 79.5 62.2 23.5	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1	104 32 428 17 14 21 24 32 53	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6% 16.8%	1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] -0.80 [-4.56 , 2.96] -6.50 [-16.76 , 3.76] -13.50 [-23.30 , -3.70] -5.25 [-14.12 , 3.62] -0.80 [-8.51 , 6.91] -18.70 [-24.29 , -13.11] 0.40 [-3.35 , 4.15]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014	-0.8 37 2; Chi ² = 19 0.42 (P = 0. 70.1 0 76.87 78.7 43.5 23.9 -0.8	22.9699 8.9 .17, df = 14 (68) .16.6 13.2288 13.4 15.2 8.8 9.5 15.6868	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6% 16.8% 16.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi ² = 40 1.99 (P = 0.42)	22.9699 8.9 .17, df = 14 (68) .16.6 13.2288 13.4 15.2 8.8 9.5 15.6868	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6% 16.8% 16.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z =	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi ² = 40 1.99 (P = 0.42)	22.9699 8.9 .17, df = 14 (68) .16.6 13.2288 13.4 15.2 8.8 9.5 15.6868	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6% 16.8% 16.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z =	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi² = 40 1.99 (P = 0.42)	22.9699 8.9 1.17, df = 14 + 68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 15.6% 16.8% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08]	+ + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi² = 40 1.99 (P = 0.42) 03; Chi² = 40 04; Chi² = 40 05; Chi² = 40 06; Chi² = 40 07; Chi² = 40	22.9699 8.9 1.17, df = 14 + 68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 P < 0.0000	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 111.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08]	+ + + + + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi² = 40 1.99 (P = 0.42) 03: Chi² = 40 04: Chi² = 40 05: Chi² = 40 06: Chi² = 40 07: Chi² = 40	22.9699 8.9 1.17, df = 14 + 68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (105)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 111.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08]	+ + + + + + + + + + + + + + + + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 13; Chi ² = 40 1.99 (P = 0.42) 1586888 70.2 64.59 78.1	22.9699 8.9 1.17, df = 14 (68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (1005)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 5%	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87]	+ + + + + + + + + + + + + + + + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI)	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi² = 40 1.99 (P = 0.42) 1.99 (P = 0.42) 1.99 (P = 0.42) 1.99 (P = 0.43) 1.99 (P = 0.44) 1.99 (P = 0	22.9699 8.9 8.9 1.17, df = 14468 68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 259, df = 6 (105) 21.1 18.1 15.6 20.7289 5.7	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 5% 25 19.73 14.3 21.5966	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 16.6% 16.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74]	+ + + + + + + + + + + + + + + + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi² = 40 1.99 (P = 0.42) 1.99 (P = 0.43) 34.6 0; Chi² = 2.28	22.9699 8.9 8.9 1.17, df = 14+68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (I 05) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P =	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 5% 25 19.73 14.3 21.5966	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 111.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77]	+ + + + + + + + + + + + + + + + + + +
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi ² = 40 1.99 (P = 0.42) 4.59 78.1 -0.3 34.6 3; Chi ² = 2.28 3.20 (P = 0.42)	22.9699 8.9 8.9 1.17, df = 14+68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (I 05) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P =	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 5% 25 19.73 14.3 21.5966	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 111.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00 Test for overall effect: Z =	-0.8 37 2; Chi ² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi ² = 40 1.99 (P = 0.42) 4.59 78.1 -0.3 34.6 3; Chi ² = 2.28 3.20 (P = 0.42)	22.9699 8.9 8.9 1.17, df = 14+68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (I 05) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P =	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 5% 25 19.73 14.3 21.5966	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 111.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77]	**************************************
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00 Test for overall effect: Z =	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi² = 40 1.99 (P = 0.42) 43.5 43.5 43.5 23.9 20.8 34.6 35; Chi² = 2.28 3.20 (P = 0.43) 45; Chi² = 2.28 45; Chi² = 2.28 45; Chi² = 2.28 45; Chi² = 2.28	22.9699 8.9 1.17, df = 14 (68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (1005) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P = 0001)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211 0.68); I ² =	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 3% 25 19.73 14.3 21.5966 6.7	104 32 428 17 14 21 24 32 53 104 265	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 15.6% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77] -4.01 [-6.47, -1.55]	**************************************
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00 Test for overall effect: Z =	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 3; Chi² = 40 1.99 (P = 0.42) 4.59 78.1 -0.3 34.6 0; Chi² = 2.28 3.20 (P = 0.43) 4.3	22.9699 8.9 8.9 1.17, df = 14 (68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 1.59, df = 6 (10) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P = 0001)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 123 33 211 10.68); I ² =	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4 = 0%	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 3% 25 19.73 14.3 21.5966 6.7	104 32 428 17 14 21 24 32 53 104 265 17 21 24 104 32 198	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77] -4.01 [-6.47, -1.55]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00 Test for overall effect: Z = 1.4.11 Burden of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi² = 40 1.99 (P = 0.42) 03: Chi² = 2.28 03: Chi² = 2.28 03: Chi² = 2.28 03: Chi² = 3.20 (P = 0.42) 04: Chi² = 2.28 05: Chi² = 2.28 06: Chi² = 2.28 07: Chi² = 2.28	22.9699 8.9 8.9 1.17, df = 14 (68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 15.9, df = 6 (1005) 21.1 18.1 15.6 20.7289 5.7 3, df = 4 (P = 0001)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 31 23 31 268 2 < 0.0000 19 13 23 123 123 123 123 123 123	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4 = 0%	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 3% 25 19.73 14.3 21.5966 6.7	104 32 428 17 14 21 24 32 53 104 265 17 21 24 104 32 198	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 15.6% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77] -4.01 [-6.47, -1.55] -0.20 [-19.05, 18.65] -1.92 [-15.41, 11.57]	
EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 13.1 Test for overall effect: Z = 1.4.9 Symptoms Martin-Alemany 2016 Pellizzaro 2013 AVANTE-HEMO 2020 Uchiyama 2019 Wu 2014d van Vilsteren 2005 EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 51.9 Test for overall effect: Z = 1.4.10 Effects of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00 Test for overall effect: Z = 1.4.11 Burden of kidney d Martin-Alemany 2016 AVANTE-HEMO 2020 Uchiyama 2019	-0.8 37 2; Chi² = 19 0.42 (P = 0.42) 70.1 0 76.87 78.7 43.5 23.9 -0.8 03; Chi² = 40 1.99 (P = 0.42) 03: Chi² = 2.22 64.59 78.1 -0.3 34.6 0; Chi² = 2.22 3.20 (P = 0.42) 0; Chi² = 4.24	22.9699 8.9 1.17, df = 144 68) 16.6 13.2288 13.4 15.2 8.8 9.5 15.6868 20.59, df = 6 (I 20.7289 5.7 3, df = 4 (P = 001)	123 33 423 (P = 0.16); 19 14 13 23 33 43 123 268 2 < 0.0000 19 13 23 33 211 10.68); I ² =	-2.5 44.5 I ² = 27% 76.6 13.5 82.12 79.5 62.2 23.5 -0.6 1); I ² = 85 70 74.65 79.8 1.5 39.4 = 0%	29.8239 11.7 14.8 13.2288 11.85 11.4 13.6 9.1 12.3409 3% 25 19.73 14.3 21.5966 6.7	104 32 428 17 14 21 24 265 17 21 24 104 32 198	14.2% 18.6% 100.0% 11.8% 12.2% 12.9% 13.9% 16.8% 16.9% 100.0% 2.6% 3.6% 8.2% 19.7% 65.9% 100.0%	1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] -0.80 [-4.56, 2.96] -6.50 [-16.76, 3.76] -13.50 [-23.30, -3.70] -5.25 [-14.12, 3.62] -0.80 [-8.51, 6.91] -18.70 [-24.29, -13.11] 0.40 [-3.35, 4.15] -0.20 [-3.85, 3.45] -6.07 [-12.07, -0.08] 0.20 [-15.01, 15.41] -10.06 [-23.02, 2.90] -1.70 [-10.27, 6.87] -1.80 [-7.34, 3.74] -4.80 [-7.83, -1.77] -4.01 [-6.47, -1.55] -0.20 [-19.05, 18.65] -1.92 [-15.41, 11.57] -5.70 [-17.40, 6.00]	



Analysis 1.4. (Continued)

									٦
Wu 2014d Subtotal (95% CI)	27.6	5.91	33 211	26.8	6.29	32 198	75.2% 100.0%	0.80 [-2.17 , 3.77] - 0.06 [-2.64 , 2.51]	•
Heterogeneity: $Tau^2 = 0.00$; Test for overall effect: $Z = 0$,	0.80); I ²	= 0%					Ĭ
	J.05 (I 0.	30)							
1.4.12 Work status	50	25.4	10	20.0	20 F	21	1.00/	10 20 [15 41 25 01]	
AVANTE-HEMO 2020	50	35.4 41.2	13 19	39.8	39.5 36.7	21 17	1.8%	10.20 [-15.41 , 35.81]	-
Martin-Alemany 2016	27.5			25.3			1.8%	2.20 [-23.25 , 27.65]	
EXCITE 2014 Wu 2014d	-0.9 29.3	31.3735 8.97	123 33	0.3 29.7	22.1108	104 32	23.5%	-1.20 [-8.19 , 5.79]	1
	29.3	0.97		29.7	7.29		72.9%	-0.40 [-4.37 , 3.57]	
Subtotal (95% CI) Heterogeneity: Tau² = 0.00;	Chi2 - 0.7	E 4f = 2 (D =	188	- 00/		174	100.0%	-0.36 [-3.75 , 3.03]	•
Test for overall effect: $Z = 0$			0.00), 1	- 070					
1.4.13 Cognitive function									
AVANTE-HEMO 2020	24.46	27.3	13	24.29	28.42	21	5.7%	0.17 [-19.01 , 19.35]	
Martin-Alemany 2016	33	20.5	19	25.4	30	17	7.1%	7.60 [-9.38 , 24.58]	
EXCITE 2014	-6.4	30.8133	123	0.3	17.9972	104	25.9%	-6.70 [-13.15 , -0.25]	Ţ
Uchiyama 2019	92.4	9.5	23	90.3	10.7	24	28.4%	2.10 [-3.68 , 7.88]	
Wu 2014d	70.4	9.19	33	76.7	10.13	32	32.8%	-6.30 [-11.01 , -1.59]	
Subtotal (95% CI)	, 0.4	5.15	211	, 5.,	10.15	198	100.0%	-2.66 [-7.57 , 2.25]	7
Heterogeneity: Tau² = 13.38	R. Chi² = 7	69 df = 4 (D		2 = 1804		130	100.0 /0	-2.00 [-7.37 , 2.23]	₹
Test for overall effect: $Z = 1$			– 0.10), 1-	- 4070					
1.4.14 Quality of social int	teractions								
AVANTE-HEMO 2020	25.92	25.3	13	22.99	21.91	21	4.1%	2.93 [-13.71 , 19.57]	
Martin-Alemany 2016	33	22	19	30.8	27.3	17	4.3%	2.20 [-14.12 , 18.52]	
Uchiyama 2019	88.1	14.9	23	88.4	10.6	24	19.3%	-0.30 [-7.72 , 7.12]	
Wu 2014d	66.5	11.5	33	73.9	11.25	32	32.5%	-7.40 [-12.93 , -1.87]	
EXCITE 2014	-4.6	20.7289	123	2.1	16.9688	104	39.8%	-6.70 [-11.60 , -1.80]	
Subtotal (95% CI)	-4.0	20.7203	211	2.1	10.5000	198	100.0%	-4.92 [-8.32 , -1.51]	T
Heterogeneity: Tau ² = 1.31;	Chi2 - 43	1 df - 1 (D -		- Q0/ ₋		130	100.0 /0	-4.52 [-0.52 , -1.51]	▼
Test for overall effect: $Z = Z$			-10-0), -						
1.4.15 Sexual function									
AVANTE-HEMO 2020	8.33	25	13	34.57	45.49	21	8.5%	-26.24 [-49.97 , -2.51]	
Martin-Alemany 2016	88.3	23.3	19	96	12.7	17	22.6%	-7.70 [-19.79 , 4.39]	
EXCITE 2014	-2.1	46.5	123	-4.9	39.5938	104	24.7%	2.80 [-8.40 , 14.00]	1
Wu 2014d	15	10.37	33	15.7	9.39	32	44.2%	-0.70 [-5.51 , 4.11]	_
Subtotal (95% CI)	10	10.07	188	1017	0.00		100.0%	-3.60 [-11.16 , 3.96]	<u>.</u>
Heterogeneity: Tau ² = 27.68	R: Chi ² = 5	84 df = 3 (P		2 = 49%			1001070	5,00 [11,10 , 5,50]	\blacksquare
Test for overall effect: $Z = 0$			0.12), 1	4370					
1.4.16 Sleep									
Pellizzaro 2013	-15	20.2665	14	8.5	20.2665	14	9.8%	-23.50 [-38.51 , -8.49]	
Martin-Alemany 2016	63.9	25	19	67		17	10.1%	-3.10 [-17.67 , 11.47]	
Uchiyama 2019	60.9	18.1	23	56.6	16.7	24	14.9%	4.30 [-5.67 , 14.27]	
AVANTE-HEMO 2020	69.72	6.9	13	74.64	9.54	21	20.9%	-4.92 [-10.46 , 0.62]	
Wu 2014d	36.4	7.54	33	49.7	11.6	32	21.9%	-13.30 [-18.07 , -8.53]	
EXCITE 2014	0.7	19.0482	123	3.7	14.9119	104	22.3%	-3.00 [-7.42 , 1.42]	
Subtotal (95% CI)	0.7	-5.0.02	225	3.,		212	100.0%	-6.58 [-12.57, -0.60]	1
Heterogeneity: Tau² = 36.63	3· Chi² = 20	16 df = 5 (1		· I2 = 750/		-14	2000/0	5.55 [IZ.57 ; -0.00]	•
Test for overall effect: $Z = 2$			- 0.001)	, 1 - / 3 70	,				
1.4.17 Social support									
Martin-Alemany 2016	70	21.3	19	68.7	26.4	17	3.8%	1.30 [-14.49 , 17.09]	
Uchiyama 2019	81	16.9	23	80.7	18.7	24	9.1%	0.30 [-9.88 , 10.48]	
AVANTE-HEMO 2020	68.53	10.5	13	76.7	19.08	21	9.9%	-8.17 [-17.98 , 1.64]	
EXCITE 2014	-2	24.0904	123	-1.5	22.1108	104	25.8%	-0.50 [-6.52 , 5.52]	
Wu 2014d			33		9.41	32	51.3%	-6.09 [-10.31 , -1.87]	
	75.27	7.86	211	81.36	5.41	32 198			<u> </u>
Subtotal (95% CI)	Ch:2 - 4.0	- 4C - 4 C		_ 10/		196	100.0%	-3.98 [-7.07 , -0.89]	•
Heterogeneity: Tau² = 0.21; Test for overall effect: Z = 2			U.4U); I ² =	= 1%					
1 A 19 Dialycic staff ancou	ragament								I

-100

-50

Higher in exercise

100

Higher in control



Analysis 1.4. (Continued)

1est for overall effect: Z = 2.53 (P = 0.01) 1.4.18 Dialysis staff encouragement Uchiyama 2019 83 23 80.6 21.7 24 11.6% 2.40 [-8.82 , 13.62] Martin-Alemany 2016 80 17.4 19 79.7 12 17 14.0% 0.30 [-9.38, 9.98] AVANTE-HEMO 2020 80.56 12.6 13 83.29 15.65 21 14.2% -2.73 [-12.31 , 6.85] EXCITE 2014 -1.6 17.9277 123 1.1 4.1136 104 30.1% -2.70 [-5.97, 0.57] Wu 2014d 33 90.6 5.4 32 30.2% -9.50 [-12.73 , -6.27] 211 Subtotal (95% CI) 198 100.0% -3.75 [-8.40, 0.90] Heterogeneity: $Tau^2 = 15.94$; $Chi^2 = 12.32$, df = 4 (P = 0.02); $I^2 = 68\%$ Test for overall effect: Z = 1.58 (P = 0.11) 1.4.19 Patient satisfaction 63.5 3.20 [-10.86, 17.26] Martin-Alemany 2016 66.7 21.7 19 21.3 17 11.1% Uchiyama 2019 78 22.8 23 75.2 17.6 24 14.2% 2.80 [-8.88 , 14.48] AVANTE-HEMO 2020 52.22 13.4 13 58.77 21 18.2% -6.55 [-15.84, 2.74] 13.49 EXCITE 2014 -4.6 25.2109 123 -1.6 18.5114 104 26.3% -3.00 [-8.70 , 2.70] Wu 2014d 33 32 30.2% -11.10 [-15.17 , -7.03] 74.8 8.6 85.9 8.16 211 Subtotal (95% CI) 198 100.0% -4.58 [-10.23, 1.06] Heterogeneity: $Tau^2 = 23.13$; $Chi^2 = 10.51$, df = 4 (P = 0.03); $I^2 = 62\%$ Test for overall effect: Z = 1.59 (P = 0.11)

- (1) mean and standard deviation estimated from the median and the range
- (2) two intervention arms polled together in the exercise group
- (3) Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group
- (4) mean and standard deviation estimated from the median and range

Analysis 1.5. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 5: Depression

		Control]	Exercise			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
1.5.1 4 months or less									
Carmack 1995	5	5	11	6.8	8.2	10	8.7%	-0.26 [-1.12, 0.60]	
Frih 2017a	13	25.64	20	8.5	14.28	21	10.5%	0.21 [-0.40, 0.83]	
Rezaei 2015	26.11	13.72	25	12.64	11.07	25	10.7%	1.06 [0.47, 1.66]	
Rahimimoghadam 2017	10.4	2.4	25	8.6	3.06	25	10.9%	0.64 [0.07, 1.21]	
Sheshadri 2020	6.6	6.5	26	11.3	12.4	27	11.0%	-0.47 [-1.01, 0.08]	-
van Vilsteren 2005	41.4	9.6	43	37.2	8.3	53	12.0%	0.47 [0.06, 0.88]	
Subtotal (95% CI)			150			161	63.7%	0.30 [-0.14, 0.74]	•
Heterogeneity: Tau ² = 0.21;	Chi ² = 17.50,	df = 5 (P =	= 0.004); I ²	2 = 71%					•
Test for overall effect: $Z = 1$.34 (P = 0.18))							
1.5.2 More than 4 months									
Giannaki 2013a	43.71	11.17	7	35.84	6.38	15	8.1%	0.93 [-0.01, 1.88]	
Ouzouni 2009	19.4	4	14	11.7	3.6	19	8.7%	1.99 [1.13, 2.85]	
Kouidi 1997	21.3	11.9	11	13.7	9.5	20	9.4%	0.71 [-0.05 , 1.47]	
Kouidi 2010	22.1	6.24	20	14.61	4.15	24	10.1%	1.41 [0.74, 2.08]	
Subtotal (95% CI)			52			78	36.3%	1.26 [0.72, 1.80]	•
Heterogeneity: Tau ² = 0.13;	$Chi^2 = 5.44$, c	lf = 3 (P =	0.14); I ² =	45%					_
Test for overall effect: $Z = 4$	1.60 (P < 0.000	001)							
Total (95% CI)			202			239	100.0%	0.65 [0.22 , 1.07]	•
Heterogeneity: Tau ² = 0.35;	Chi ² = 38.89,	df = 9 (P ·	< 0.0001);	$I^2 = 77\%$. , .	—
Test for overall effect: $Z = 2$	2.99 (P = 0.003	3)	,						-4 -2 0 2 4
	es: Chi ² = 7.34	*							U -



Analysis 1.6. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 6: 6MWT

	(Control			xercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [metres]	SD [metres]	Total	Mean [metres]	SD [metres]	Total	Weight	IV, Random, 95% CI [metres]	IV, Random, 95% CI [metres]
Marchesan 2016 (1)	399.43	164	8	498.5	164	7	0.6%	-99.07 [-265.43 , 67.29]	
ACTINUT 2013	295.77	121.07	9	346.28	134.88	7	0.9%	-50.51 [-177.95 , 76.93]	
Martins do Valle 2020	494.8	66.9	12	457.3	155.6	12	1.6%	37.50 [-58.33 , 133.33]	
Coh 2009 (2)	452	144	16	509.5	121.07	28	2.1%	-57.50 [-141.10, 26.10]	
amara 2016	454.4	90.4	12	625.6	128.1	15	2.1%	-171.20 [-253.77, -88.63]	<u> </u>
EAK 2006	414.3	127.3	25	514.9	163.9	24	2.1%	-100.60 [-183.00 , -18.20]	<u> </u>
osa 2018	469.42	162.93	24	526.45	126.15	28	2.2%	-57.03 [-137.23, 23.17]	
ellizzaro 2013	407	116.7	14	475	74.1	14	2.7%	-68.00 [-140.41 , 4.41]	
ouchon 2016	400	65.99	4	420	11.96	8	3.2%	-20.00 [-85.20 , 45.20]	
HAIR 2015 (3)	317.5	81.6	11	307.5	54.62	6	3.2%	10.00 [-55.08, 75.08]	+
⁄u 2014d	359	132	33	441	135	32	3.2%	-82.00 [-146.93 , -17.07]	
ePaul 2002	430	80	14	464	94	15	3.4%	-34.00 [-97.40 , 29.40]	-
iao 2016	290	64.1	20	350	128.2	20	3.4%	-60.00 [-122.82 , 2.82]	
IALY-SIZE 2016 (4)	0.8	44	8	44.99	72.01	23	6.3%	-44.19 [-86.57 , -1.81]	-
egura-Orti 2009	20.6	36.6	8	48.5	60.8	17	7.3%	-27.90 [-66.35 , 10.55]	
ernandes 2019	325	59.8	19	386.9	19.38	20	10.5%	-61.90 [-90.10 , -33.70]	-
(ho 2018 (5)	-26	41	13	22.39	22.16	33	12.4%	-48.39 [-71.92 , -24.86]	-
rih 2017a	415.6	36.3	20	480.5	31.9	21	13.6%	-64.90 [-85.86 , -43.94]	-
XCITE 2014	2	44.82	123	39	35.99	104	18.9%	-37.00 [-47.52 , -26.48]	•
otal (95% CI)			393			434	100.0%	-49.91 [-62.59 , -37.22]	•
leterogeneity: Tau ² = 192	2.77; Chi ² = 27.33, df	= 18 (P = 0.07);	$I^2 = 34\%$						"
est for overall effect: Z	= 7.71 (P < 0.00001)							-	500 -250 0 250
est for subgroup differer	nces: Not applicable								ner with exercise Further with

- $(1) standard \ deviation \ imputed \ from \ the \ highest \ standard \ deviation \ of \ the \ other \ included \ studies$
- $\ensuremath{\text{(2)}}\ \text{two intervention arms pooled together in the exercise group}$
- (3) mean and standard deviation estimated from the median and interquartile range
- (4) three intervention arms pooled together in the exercise group
- (5) three interventions arms pooled together in the exercise group

Analysis 1.7. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N reps/30 sec]

	Control]	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Bennett 2013 (1)	7.8	3.1177	12	10.2	3.17	29	8.8%	-2.40 [-4.51 , -0.29)]
Cho 2018 (2)	-0.5	2.2	13	3.058	2.6	33	17.7%	-3.56 [-5.05 , -2.07	7] -
DIALY-SIZE 2016 (3)	1.4	4.3	8	1.287	2.64	23	3.9%	0.11 [-3.06 , 3.28	3]
Frih 2017a	10.85	2.05	20	13.3	1.75	21	28.7%	-2.45 [-3.62 , -1.28	3] 📲
Giannaki 2013a	17.25	3.2	7	17.84	4.68	15	3.5%	-0.59 [-3.94 , 2.76	5]
IHOPE 2019	10.7	5.6	38	11.7	4	29	7.4%	-1.00 [-3.30 , 1.30)] <u> </u>
Koufaki 2002	12.05	3.6	15	13.45	3.1	18	7.3%	-1.40 [-3.72 , 0.92	2]
Marchesan 2016 (4)	10	7.7	7	14.62	7.7	8	0.6%	-4.62 [-12.43 , 3.19	9]
Rosa 2018	11.79	2.93	24	15.18	6.07	28	6.1%	-3.39 [-5.93 , -0.85	5]
Segura-Orti 2009	0.3	3.55	8	2.55	2.9	17	4.9%	-2.25 [-5.07, 0.57	7] -
Song 2012a	7.1	7.6	20	8.2	7.7	20	1.7%	-1.10 [-5.84 , 3.64	1]
Wu 2014d	12.8	3.65	33	15.55	4.8	32	9.1%	-2.75 [-4.83 , -0.67	7]
Total (95% CI)			205			273	100.0%	-2.36 [-2.98 , -1.73	B] •
Heterogeneity: Tau ² = 0.	.00; Chi ² = 9	.29, df = 1	1 (P = 0.59); I ² = 0%					'
Test for overall effect: Z	= 7.37 (P <	0.00001)							-20 -10 0 10 20
Test for subgroup differen	ences: Not ap	plicable							More with exercise More with control

- (1) results from group 1 (24 weeks of intervention) and group 2 (12 weeks of intervention) were pooled together in the exercise group. The number of participants was correct
- (2) three interventions arms pooled together in the exercise group
- (3) three intervention arms pooled together in the exercise group
- (4) standard deviation imputed from the highest standard deviation of the other included studies



Analysis 1.8. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 8: Sit-To-Stand test [sit to 5 reps]

		Control]	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Johansen 2006 (1)	15.1	5.41	33	14.05	6.65	35	3.2%	1.05 [-1.82 , 3.92]	
Koufaki 2002	12.7	4.8	15	11	3.3	18	3.2%	1.70 [-1.17 , 4.57]	
Segura-Orti 2009	-0.65	1.75	8	-2.7	5.3	17	3.4%	2.05 [-0.75 , 4.85]	
Samara 2016	10.6	3.25	12	7.6	2.7	15	5.1%	3.00 [0.71, 5.29]	
Wu 2014d	12.6	3.6	33	10.75	3.4	32	9.2%	1.85 [0.15, 3.55]	
Giannaki 2013a	8.81	0.66	7	8.24	2.34	15	16.2%	0.57 [-0.71 , 1.85]	
EXCITE 2014	-0.6	5.04	123	-2.5	2.06	104	28.1%	1.90 [0.93, 2.87]	
Frih 2017a	15.5	1.55	20	13.5	1.45	21	31.5%	2.00 [1.08, 2.92]	-
Total (95% CI)			251			257	100.0%	1.74 [1.22 , 2.25]	•
Heterogeneity: Tau ² = 0	.00; Chi ² = 5.	.06, df = 7	(P = 0.65)	$I^2 = 0\%$					_
Test for overall effect: Z	Z = 6.60 (P <	0.00001)							-4 -2 0 2 4
Test for subgroup differ	ences: Not ap	plicable						Lon	ger with exercise Longer with contr

(1) Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

Analysis 1.9. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 9: Systolic blood pressure

	Control			I	exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mm Hg]	SD [mm Hg]	Total	Mean [mm Hg]	SD [mm Hg]	Total	Weight	IV, Random, 95% CI [mm Hg]	IV, Random, 95% CI [mm Hg	
1.9.1 Aerobic										
Liao 2016 (1)	138.8	16.7	20	96	64.2	20	3.2%	42.80 [13.73, 71.87]		
CYCLE-HD 2016	148.5	28.23	13	142.27	25.99	9	4.7%	6.23 [-16.66, 29.12]		
McGregor 2018	123.17	29.52	18	135.75	25.28	16	6.4%	-12.58 [-31.00, 5.84]		
Goldberg 1983	149	17	11	142	27	14	6.9%	7.00 [-10.35 , 24.35]	 -	
Koh 2009 (2)	136	29	16	140.5	25.34	30	7.1%	-4.50 [-21.36 , 12.36]		
Toussaint 2008	147.8	23.5	10	141.4	11.9	9	7.3%	6.40 [-10.11, 22.91]		
Wilund 2010	147.1	14.9	9	153	17.2	8	7.9%	-5.90 [-21.29 , 9.49]		
Tsuyuki 2003	130.8	23.3	12	141.5	16.4	17	8.0%	-10.70 [-26.02 , 4.62]		
Cooke 2018 (3)	0.4	9.89	10	-9.17	21.93	10	8.2%	9.57 [-5.34 , 24.48]	-	
Paluchamy 2018	148	14.77	10	137	16.349	10	9.0%	11.00 [-2.66 , 24.66]	-	
Deligiannis 1999a	144	10	6	143	17	10	9.3%	1.00 [-12.23 , 14.23]	_	
IHOPE 2019	148.9	23.3	38	132.4	27.9	29	9.7%	16.50 [3.93, 29.07]		
Fernandes 2019	143.16	16.68	19	140.5	11.9	20	12.4%	2.66 [-6.48, 11.80]	<u>_</u>	
Subtotal (95% CI)			192			202	100.0%	3.99 [-1.80 , 9.78]	.	
Heterogeneity: Tau ² = 4	48.74; Chi ² = 21.91, di	f = 12 (P = 0.04);	I ² = 45%						ľ	
Test for overall effect:	Z = 1.35 (P = 0.18)									
1.9.2 Combined aerob	oic and resistance									
Molsted 2004 (4)	149	24.75	8	132.5	19.46	11	4.9%	16.50 [-4.15, 37.15]		
DePaul 2002	153.1	20.2	14	146	19	15	8.6%	7.10 [-7.20 , 21.40]	<u> </u>	
Deligiannis 1999a	144	10	6	136	14	15	12.6%	8.00 [-2.69 , 18.69]		
van Vilsteren 2005	146	25	43	140	26.4	53	13.1%	6.00 [-4.31 , 16.31]	<u> </u>	
Kouidi 2008	133.7	14.9	21	128.9	13.2	22	16.0%	4.80 [-3.63, 13.23]	<u></u>	
Ouzouni 2009	139.3	9.1	14	135.3		19	18.5%	4.00 [-3.07 , 11.07]	_	
Frih 2017a	149.2	5.1	20	134.1	5.2	21	26.3%	15.10 [11.95 , 18.25]		
Subtotal (95% CI)			126			156	100.0%	8.69 [3.69 , 13.69]	<u> </u>	
Heterogeneity: Tau ² = 2	22.20; Chi ² = 13.84, di	f = 6 (P = 0.03); I	² = 57%					,		
Test for overall effect:										
1.9.3 Others										
Soliman 2015	143.33	16.14	12	117.78	11.66	18	100.0%	25.55 [14.95, 36.15]		
Subtotal (95% CI)			12			18	100.0%	25.55 [14.95 , 36.15]	🖚	
Heterogeneity: Not app	olicable									
Test for overall effect:)								
Test for subgroup differ	rences: Chi ² = 12.24, o	df = 2 (P = 0.002)	, I ² = 83.7	%					100 -50 0 50 her with exercise Higher w	

- (1) these data have been verified
- (2) Two intervention arms pooled together in the exercise group $% \left\{ 1,2,...,n\right\}$
- (3) Mean and standard deviation estimated from the median and interquartile range
- (4) mean and standard deviation estimated from the median and range



Analysis 1.10. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 10: Diastolic blood pressure

	Control			E	Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mm Hg]	SD [mm Hg]	Total	Mean [mm Hg]	SD [mm Hg]	Total	Weight	IV, Random, 95% CI [mm Hg]	IV, Random, 95% CI [mm Hg]	
1.10.1 Aerobic										
Paluchamy 2018	83	27.51	10	83	24.35	10	1.6%	0.00 [-22.77 , 22.77]		
Liao 2016 (1)	77.1	13.7	20	53.7	35.3	20	2.9%	23.40 [6.81, 39.99]		
Cooke 2018 (2)	3.5	10.32	10	-2.6	16.77	10	4.8%	6.10 [-6.10 , 18.30]	 	
Goldberg 1983	86	12	11	82	18	14	5.1%	4.00 [-7.80 , 15.80]	 -	
McGregor 2018	70.5	16.49	18	72.13	15.65	16	5.9%	-1.63 [-12.44, 9.18]		
Tsuyuki 2003	79	13.5	12	85.8	12.3	17	7.0%	-6.80 [-16.42 , 2.82]		
Koh 2009 (3)	75	15	16	78	13.15	30	8.1%	-3.00 [-11.73, 5.73]		
CYCLE-HD 2016	79.06	11.98	13	77.91	8.72	9	8.2%	1.15 [-7.50, 9.80]		
Wilund 2010	77.3	8.7	9	85.7	7.7	8	9.4%	-8.40 [-16.20 , -0.60]		
IHOPE 2019	78.5	12.5	38	76.6	16.6	29	10.3%	1.90 [-5.33 , 9.13]		
Toussaint 2008	72.8	9.4	10	77.2	5.7	9	10.9%	-4.40 [-11.31 , 2.51]		
Fernandes 2019	86.32	12.52	19	86	6.6	20	12.0%	0.32 [-6.01, 6.65]		
Deligiannis 1999a	82	3	6	83	8	10	13.9%	-1.00 [-6.51 , 4.51]	_	
Subtotal (95% CI)			192			202	100.0%	-0.72 [-3.69 , 2.24]	•	
Heterogeneity: Tau ² = 8	i.63; Chi ² = 17.33, df	= 12 (P = 0.14); I	² = 31%						Ĭ	
Test for overall effect: 2	Z = 0.48 (P = 0.63)									
1.10.2 Combined aero	bic and resistance									
Molsted 2004 (4)	86.25	18.83	8	79.25	17.89	11	0.8%	7.00 [-9.79 , 23.79]		
DePaul 2002	85.2	11.7	14			15	4.2%	3.50 [-4.02 , 11.02]	+-	
van Vilsteren 2005	79	12	43	80	14.9	53	8.1%	-1.00 [-6.38 , 4.38]	+	
Deligiannis 1999a (5)	82	3	6	79	8	15	10.6%	3.00 [-1.71 , 7.71]	 -	
Kouidi 2008	82.4	. 7	21	76.9	7.9	22	11.8%	5.50 [1.04, 9.96]		
Ouzouni 2009	85.2	4.6	14			19	13.2%	6.00 [1.78 , 10.22]		
Frih 2017a	78	3.4	20	73	3.6	21	51.2%	5.00 [2.86 , 7.14]	•	
Subtotal (95% CI)			126			156	100.0%	4.45 [2.91 , 5.98]	♦	
Heterogeneity: Tau ² = 0	.00; Chi ² = 5.44, df =	6 (P = 0.49); I ² =	0%						'	
Test for overall effect: 2	Z = 5.68 (P < 0.00001)								
1.10.3 Others										
Soliman 2015	90.42	8.11	12		8.24	18	100.0%	13.42 [7.46 , 19.38]	-	
Subtotal (95% CI)			12			18	100.0%	13.42 [7.46 , 19.38]	-	
Heterogeneity: Not app	licable									
Test for overall effect: 2	Z = 4.41 (P < 0.0001)									
Test for subgroup differ	rences: Chi ² = 19.60, o	df = 2 (P < 0.000)	1), I ² = 89.	8%				Hig	-50 -25 0 25 ther with exercise Higher with	

- (1) these data have been verified
- $\ensuremath{\text{(2)}}\ Mean\ and\ standard\ deviation\ estimated\ from\ the\ median\ and\ interquartile\ range$
- (3) Two intervention arms pooled together in the exercise group
 (4) mean and standard deviation estimated from the median and range



Analysis 1.11. Comparison 1: Any exercise versus control (no exercise/ placebo exercise), Outcome 11: Aerobic capacity (VO max or peak)

		Control			Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mL/kg/min]	SD [mL/kg/min]	Total	Mean [mL/kg/min]	SD [mL/kg/min]	Total	Weight	IV, Random, 95% CI [mL/kg/min]	IV, Random, 95% CI [mL/kg/min]	
Parsons 2004	55	26	7	58	44	6	0.1%	-3.00 [-43.13 , 37.13]		
Goldberg 1983	20	8	11	25	9	14	2.3%	-5.00 [-11.68 , 1.68]		
Deligiannis 1999a (1)	15.8	4.8	12	17.8	9.96	25	4.6%	-2.00 [-6.76 , 2.76]	-	
Painter 2002a (2)	19.83	5.91	25	21.11	9.8	23	4.8%	-1.28 [-5.91 , 3.35]		
Kouidi 1997	15.9	4.3	11	23.27	7.6	20	5.9%	-7.37 [-11.56 , -3.18]		
Molsted 2004 (3)	19.5	3.81	9	19.03	5.12	9	5.9%	0.47 [-3.70 , 4.64]	+	
McGregor 2018	15.93	5.168	18	20.71	6.831	16	6.1%	-4.78 [-8.89 , -0.67]		
Tsuyuki 2003	21.7	4.9	12	27	5.6	17	6.9%	-5.30 [-9.14 , -1.46]	-	
Koufaki 2002	18.8	4.9	15	19.9	6.3	18	7.0%	-1.10 [-4.92 , 2.72]	-	
Konstantinidou 2002 (4)	15.8	4.8	12	21.4	6.76	36	8.3%	-5.60 [-9.10 , -2.10]	-	
Carmack 1995	10.9	3.1	11	14.4	4.7	10	8.6%	-3.50 [-6.94 , -0.06]		
Jong 2004	22.86	5.46	17	25.23	4.33	19	9.6%	-2.37 [-5.61, 0.87]	-	
Akiba 1995	17.6	2.6	6	20	2.4	7	13.4%	-2.40 [-5.14, 0.34]	-	
Kouidi 2004a	-0.4	2.3	11	3.1	3.3	10	16.5%	-3.50 [-5.96 , -1.04]	-	
Total (95% CI)			177			230	100.0%	-3.30 [-4.33 , -2.28]	•	
Heterogeneity: Tau ² = 0.0		$(P = 0.43); I^2 = 2\%$								
Test for overall effect: Z =									-50 -25 0 25 50	
Test for subgroup differen	ces: Not applicable							Hig	ther with exercise Higher with control	

Footnotes

(1) two intervention arms pooled together in the exercise group

- (2) factorial design: two intervention arms pooled together in the exercise group and two control arms pooled together in the control group (3) mean and standard deviation estimated from the median and range
- $\left(4\right)$ three intervention arms pooled together in the exercise group

Analysis 1.12. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 12: Albumin

		Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
Frey 1999	37	7 7	6	44	5	5	1.2%	-7.00 [-14.11 , 0.11]	
CHAIR 2015 (1)	39.25	2.2	11	39	6.2	6	2.0%	0.25 [-4.88 , 5.38]	
Koufaki 2002	40.5	3.6	15	35.2	7.3	18	3.0%	5.30 [1.47, 9.13]	
Jong 2004	33	6.1	17	38.2	5.1	19	3.1%	-5.20 [-8.90 , -1.50]	
Matsumoto 2007	36.9	8.0435	32	38.6	4.4734	17	3.3%	-1.70 [-5.21 , 1.81]	
Reboredo 2010	41	. 5	11	39	3	11	3.3%	2.00 [-1.45 , 5.45]	
ACTINUT 2013	39.12	3.67	9	39.33	2.51	7	3.8%	-0.21 [-3.24 , 2.82]	
Dong 2011	42.1	2.2	12	41.5	4.4	10	3.8%	0.60 [-2.40 , 3.60]	- -
Afshar 2010 (2)	40) 3	7	40	3.4	14	4.0%	0.00 [-2.85 , 2.85]	
Martins do Valle 2020	39	3	12	37	4	12	4.0%	2.00 [-0.83 , 4.83]	
Uchiyama 2019	33.7	5.2	23	34.8	4.4	24	4.1%	-1.10 [-3.86 , 1.66]	
AVANTE-HEMO 2020	39	9 4	13	35.7	3.4	21	4.3%	3.30 [0.68, 5.92]	
Kopple 2007 (3)	39	3.7417	14	38	3.7	41	4.8%	1.00 [-1.26 , 3.26]	 -
Liao 2016	40.1	4.2	20	41.6	3	20	4.8%	-1.50 [-3.76 , 0.76]	
Martin-Alemany 2016	37	3.5	19	37	3.3	17	4.9%	0.00 [-2.22 , 2.22]	+
Fernandes 2019	34.6	4.1	19	37.9	2.5	20	5.0%	-3.30 [-5.44 , -1.16]	
IHOPE 2019	40.1	3.1	38	39.3	5.1	29	5.0%	0.80 [-1.30 , 2.90]	 -
Toussaint 2008	34.1	2	10	33.8	2.5	9	5.1%	0.30 [-1.75 , 2.35]	 -
Frih 2017a	40	2.6	21	40.4	3.7	20	5.2%	-0.40 [-2.37 , 1.57]	+
Abreu 2017	42	2 2	19	43	3	25	5.9%	-1.00 [-2.48, 0.48]	
PEAK 2006	-0.16	5 2.4	25	0.3	2.4	24	6.1%	-0.46 [-1.80 , 0.88]	+
Wilund 2010	38	0.6	9	39	1.5	8	6.4%	-1.00 [-2.11, 0.11]	-
Pellizzaro 2013	-2	2 1	14	1	1	14	6.8%	-3.00 [-3.74 , -2.26]	•
Total (95% CI)			376			391	100.0%	-0.39 [-1.25 , 0.47]	
Heterogeneity: Tau ² = 2.7	1; Chi ² = 79.17,	df = 22 (P <	0.00001)	; I ² = 72%					Ĭ
Test for overall effect: Z =	= 0.88 (P = 0.38)							-20 -10 0 10 20
Test for subgroup differer	nces: Not applica	able						Hi	gher with exercise Higher with cont

- (1) mean and standard deviation estimated from the median and range
- (2) Two intervention arms pooled together in the exercise group
- $\begin{tabular}{ll} (3) three intervention arms pooled together in the exercise group \\ \end{tabular}$



Analysis 1.13. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 13: Blood lipids

		Control			ercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
.13.1 Total cholesterol	l [mmol/f.]								
Molsted 2004 (1)	4.25	1.07	9	4.83	0.85	11	3.4%	-0.58 [-1.44, 0.28]	
Dong 2011	4.49	1.1	12	4.36	0.64	10	4.3%	0.13 [-0.61 , 0.87]	
Afshar 2010 (2)	3.4	0.812	7	3.329	0.722	14	4.6%	0.07 [-0.64 , 0.78]	
Lee 2001	5.08	0.7	21	4.38	0.98	19	6.8%	0.70 [0.17 , 1.23]	<u></u>
Uchiyama 2019	4.4	0.812	23	4.5	0.897	24	7.6%	-0.10 [-0.59 , 0.39]	
Wilund 2010	4.25		9	3.54	0.339	8	8.1%	0.71 [0.25 , 1.17]	
van Vilsteren 2005	4.6	1.2	43	4.6	1	53	8.4%	0.00 [-0.45 , 0.45]	
Liao 2016	4.71	0.58	20	4.456	0.776	20	8.9%	0.25 [-0.17 , 0.68]	<u> </u>
Song 2012a	4.192	0.67	20	3.85	0.678	20	9.1%	0.34 [-0.08, 0.76]	<u> </u>
Yurtkuran 2007	4.021	0.437	18	3.47	0.786	19	9.3%	0.55 [0.14, 0.96]	
Frih 2017a	4.1	0.6	20	3.9	0.4	21	11.7%	0.20 [-0.11 , 0.51]	<u> </u>
Groussard 2015	1.63	0.09	10	1.61	0.13	8	17.7%	0.02 [-0.09, 0.13]	Γ
Subtotal (95% CI)			212			227	100.0%	0.22 [0.04, 0.39]	
Heterogeneity: Tau ² = 0. Test for overall effect: Z		= 11 (P = 0.01); I ²	= 54%						
1.13.2 LDL cholesterol									
Molsted 2004 (1)	2.33	0.7	9	2.33	1.16	11	7.4%	0.00 [-0.82 , 0.82]	
Lee 2001	2.91	0.97	21	2.51	0.913	19	11.4%	0.40 [-0.18, 0.98]	+-
Afshar 2010 (2)	1.56	0.33	7	1.28	0.49	14	17.4%	0.28 [-0.07, 0.63]	 •
Song 2012a	2.102	0.55	20	2.02	0.54564	20	17.9%	0.08 [-0.26 , 0.42]	-
Frih 2017a	2.5	0.4	20	1.9	0.3	21	21.5%	0.60 [0.38, 0.82]	-
Groussard 2015	0.91	0.09	10	0.89	0.12	8	24.3%	0.02 [-0.08 , 0.12]	+
Subtotal (95% CI) Heterogeneity: Tau ² = 0.			87			93	100.0%	0.24 [-0.02 , 0.51]	*
1.13.3 HDL cholesterol		0.40	0	4.25	0.44		6.20/	0.00 [0.40 . 0.25]	
Molsted 2004 (1)	1.33		9	1.35	0.44	11	6.3%	-0.02 [-0.40 , 0.36]	+
Afshar 2010 (2)	0.82		7	0.912	0.29	14	9.1%	-0.09 [-0.38 , 0.20]	
Uchiyama 2019	1.182	0.367	23	1.543	0.538	24	10.2%	-0.36 [-0.62 , -0.10]	
Yurtkuran 2007	1.065432	0.447	18	1.13784	0.33877	19	10.5%	-0.07 [-0.33 , 0.18]	+
Frih 2017a	1.4	0.4	20	1.7	0.4	21	11.1%	-0.30 [-0.54 , -0.06]	
Song 2012a	1.1	0.4	20	1.07	0.26	20	13.0%	0.03 [-0.18 , 0.24]	+
Lee 2001	0.99	0.22	21	0.97	0.27	19	16.5%	0.02 [-0.13 , 0.17]	+
Groussard 2015 Subtotal (95% CI)	0.47	0.04	10	0.41	0.05	8	23.3% 100.0%	0.06 [0.02 , 0.10]	
			128 = 63%			136	100.0%	-0.07 [-0.18 , 0.04]	•
Heterogeneity: Tau ² = 0.		= 7 (P = 0.009); I ²	- 0370						
Heterogeneity: Tau ² = 0. Test for overall effect: Z	Z = 1.17 (P = 0.24)	= 7 (P = 0.009); I ²	- 0370						
Heterogeneity: Tau ² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m	Z = 1.17 (P = 0.24)			1.8677	0.71	14	9.4%	0.68 [-0.08 : 1.45]	
Heterogeneity: Tau ² = 0. Fest for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2)	Z = 1.17 (P = 0.24) amol/L] 2.55	0.903	7	1.8677 1.5	0.71 0.66	14 19	9.4% 10.1%	0.68 [-0.08 , 1.45] 0.78 [0.08 , 1.48]	
Heterogeneity: Tau ² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001	Z = 1.17 (P = 0.24) amol/L] 2.55 2.28	0.903 1.49	7 21	1.5	0.66	19	10.1%	0.78 [0.08 , 1.48]	
Heterogeneity: Tau ² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1)	Z = 1.17 (P = 0.24) amol/L] 2.55 2.28 1.135	0.903 1.49 0.428	7 21 9	1.5 1.54	0.66 0.99	19 11	10.1% 10.8%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a	z = 1.17 (P = 0.24) amol/L] 2.55 2.28 1.135 1.6699	0.903 1.49 0.428 1.2	7 21 9 20	1.5 1.54 1.34	0.66 0.99 0.71469	19 11 20	10.1% 10.8% 11.3%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94]	
Heterogeneity: Tau ² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m 47shar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019	z = 1.17 (P = 0.24) amol/L] 2.55 2.28 1.135 1.6699 1.305	0.903 1.49 0.428 1.2 1.009	7 21 9 20 23	1.5 1.54 1.34 1.838	0.66 0.99 0.71469 1.0105	19 11 20 24	10.1% 10.8% 11.3% 11.7%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Jchiyama 2019 Yurtkuran 2007	z = 1.17 (P = 0.24) http://linearchi.acm.com/L] 2.55 2.28 1.135 1.6699 1.305 2.2055	0.903 1.49 0.428 1.2 1.009 0.1859	7 21 9 20 23 18	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842	19 11 20 24 19	10.1% 10.8% 11.3% 11.7% 15.1%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.1.3.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019 Yurtkuran 2007 Frih 2017a	2 = 1.17 (P = 0.24) 1100I/L] 2.55 2.28 1.135 1.6699 1.305 2.2055 1.9	0.903 1.49 0.428 1.2 1.009 0.1859	7 21 9 20 23 18 20	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842 0.4	19 11 20 24 19 21	10.1% 10.8% 11.3% 11.7% 15.1% 15.6%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32] 0.50 [0.22 , 0.78]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019 Yurtkuran 2007 Frih 2017a Groussard 2015	z = 1.17 (P = 0.24) http://linearchi.acm.com/L] 2.55 2.28 1.135 1.6699 1.305 2.2055	0.903 1.49 0.428 1.2 1.009 0.1859	7 21 9 20 23 18	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842	19 11 20 24 19 21 8	10.1% 10.8% 11.3% 11.7% 15.1% 15.6% 16.0%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32] 0.50 [0.22 , 0.78] -0.37 [-0.61 , -0.13]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019 Yurtkuran 2007 Frih 2017a Groussard 2015 Subtotal (95% CI)	2 = 1.17 (P = 0.24) 1 = 0.24) 1 = 0.255 2.28 1.135 1.6699 1.305 2.2055 1.9 1.09	0.903 1.49 0.428 1.2 1.009 0.1859 0.5	7 21 9 20 23 18 20 10	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842 0.4	19 11 20 24 19 21	10.1% 10.8% 11.3% 11.7% 15.1% 15.6%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32] 0.50 [0.22 , 0.78]	
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019 Yurtkuran 2007 Frih 2017a Groussard 2015	2 = 1.17 (P = 0.24) 2.55 2.28 1.135 1.6699 1.305 2.2055 1.9 1.09	0.903 1.49 0.428 1.2 1.009 0.1859 0.5	7 21 9 20 23 18 20 10	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842 0.4	19 11 20 24 19 21 8	10.1% 10.8% 11.3% 11.7% 15.1% 15.6% 16.0%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32] 0.50 [0.22 , 0.78] -0.37 [-0.61 , -0.13]	+
Heterogeneity: Tau² = 0. Test for overall effect: Z 1.13.4 Triglycerides [m Afshar 2010 (2) Lee 2001 Molsted 2004 (1) Song 2012a Uchiyama 2019 Yurtkuran 2007 Frih 2017a Groussard 2015 Subtotal (95% CI) Heterogeneity: Tau² = 0.	2 = 1.17 (P = 0.24) 2.55 2.28 1.135 1.6699 1.305 2.2055 1.9 1.09	0.903 1.49 0.428 1.2 1.009 0.1859 0.5	7 21 9 20 23 18 20 10	1.5 1.54 1.34 1.838 2.2077	0.66 0.99 0.71469 1.0105 0.6842 0.4	19 11 20 24 19 21 8	10.1% 10.8% 11.3% 11.7% 15.1% 15.6% 16.0%	0.78 [0.08 , 1.48] -0.41 [-1.05 , 0.24] 0.33 [-0.28 , 0.94] -0.53 [-1.11 , 0.04] -0.00 [-0.32 , 0.32] 0.50 [0.22 , 0.78] -0.37 [-0.61 , -0.13]	

⁽¹⁾ mean and standard error estimated from the median and the range

⁽²⁾ Two intervention arms pooled together in the exercise group



Analysis 1.14. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 14: Body composition

		Control			Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
1.14.1 Fat mass [kg]									
Dong 2011	27.6	14.8	12	24.7	10.5	10	0.0%	2.90 [-7.71 , 13.51]	1
Giannaki 2013a	28.5	5.8	7	30.5	8.1	15	0.0%	-2.00 [-7.94, 3.94]	1
Chen 2010	33.1	10.1	22	29.6	9.8	22	0.0%	3.50 [-2.38, 9.38]	1 +
Johansen 2006 (1)	18.68	13.21	33	21.8	10.08	35	0.0%	-3.12 [-8.73 , 2.49]	1
Song 2012a	27.2	8.9	20	26	8.6	20	0.0%	1.20 [-4.22 , 6.62]	l
Martin-Alemany 2016	17.6	6.5	19	20.3	9	17	0.0%	-2.70 [-7.88, 2.48]	1
Rosa 2018	21.92	8.81	24	23.2	8.4	28	0.0%	-1.28 [-5.98 , 3.42]	l —
Kopple 2007 (2)	19.1	2.4	14	21.5	12.7738	37	0.0%	-2.40 [-6.70 , 1.90]	1
Sheshadri 2020	-0.04	0.1071	25	0	0.1071	24	99.9%	-0.04 [-0.10, 0.02]	l 💼
Subtotal (95% CI)			176			208	100.0%	-0.04 [-0.10 , 0.02]	· — — —
Heterogeneity: Tau ² = 0.00	0; Chi ² = 5.90), $df = 8 (I$	P = 0.66); I	$^{2} = 0\%$					
Test for overall effect: Z =	= 1.34 (P = 0.	18)							
1.14.2 Lean mass [kg]									
Dong 2011	56.2	9.9	12	47.4	7.1	10	8.8%	8.80 [1.68, 15.92]	ı
IHOPE 2019	58.6	15.9	38	61.5	13.1	29	9.2%	-2.90 [-9.85 , 4.05]	ı
Giannaki 2013a	45.4	5.1	7	46.7	8.3	15	12.5%	-1.30 [-6.95 , 4.35]	ı —
Chen 2010	46.3	8.7	21	47.9	9.9	21	12.6%	-1.60 [-7.24 , 4.04]	ı —
Johansen 2006 (1)	48.1	8.76	33	48.47	13.01	35	13.9%	-0.37 [-5.62 , 4.88]	l
Rosa 2018	44.04	8.23	24	47.55	9.49	28	15.6%	-3.51 [-8.33 , 1.31]	ı -
Song 2012a	22.5	5.2	20	22.2	3.7	20	27.3%	0.30 [-2.50, 3.10]	l
Subtotal (95% CI)			155			158	100.0%	-0.37 [-2.74, 1.99]	ı 🃥
Heterogeneity: Tau ² = 3.29	9; Chi ² = 9.01	l, df = 6 (I	P = 0.17); I	² = 33%					T
Test for overall effect: Z =	= 0.31 (P = 0.7	76)							
Test for subgroup differen	ices: Chi ² = 0	.08, df = 1	(P = 0.78)	$I^2 = 0\%$					-20 -10 0 10
								Н	igher with exercise Higher

- (1) Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group
- $\left(2\right)$ three intervention arms pooled together in the exercise group

Analysis 1.15. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 15: Body mass index

		Control		Exercise				Mean Difference	Mean Difference	
Study or Subgroup	Mean [kg/m²]	SD [kg/m ²]	Total	Mean [kg/m²]	SD [kg/m ²]	Total	Weight	IV, Random, 95% CI [kg/m²]	IV, Random, 95% CI [kg/m²]	
Marinho 2016 (1)	29.13	10.655	7	28.8	13.73	6	0.1%	0.33 [-13.20 , 13.86]		
Oong 2011	29.3	6.8	12	26.8	4.3	10	0.8%	2.50 [-2.18, 7.18]		
HOPE 2019	31.5	7.4	38	33.9	10.9	29	0.8%	-2.40 [-7.01, 2.21]		
Copple 2007 (2)	25.1	4.49	14	26.9	7.2993	37	1.6%	-1.80 [-5.13 , 1.53]		
Abreu 2017	24.1	4.9	19	23.8	4.5	25	2.1%	0.30 [-2.52 , 3.12]		
iao 2016	23.91	5.27	20	22.96	3.36	20	2.2%	0.95 [-1.79 , 3.69]		
Koufaki 2002	24.7	3.5	15	25.7	3.3	18	3.0%	-1.00 [-3.34 , 1.34]		
losa 2018	25.51	4.03	24	26.61	4.44	28	3.0%	-1.10 [-3.40 , 1.20]		
chiyama 2019	24.5	4.3	23	22.8	3.4	24	3.2%	1.70 [-0.52 , 3.92]		
Vilund 2010	30.3	2.5	9	28.3	1.8	8	3.7%	2.00 [-0.06 , 4.06]	-	
CTINUT 2013	20.93	2.57	9	20.89	1.19	7	4.3%	0.04 [-1.86 , 1.94]	+	
fartin-Alemany 2016 (1)	21.2	1.698	17	21.3	3.204	19	5.4%	-0.10 [-1.75 , 1.55]	+	
VANTE-HEMO 2020	19.8	1.7	13	21.47	2.1	21	7.8%	-1.67 [-2.96 , -0.38]	-	
Cooke 2018 (1)	0.2067	0.413	10	0.3333	1.015	10	16.0%	-0.13 [-0.81 , 0.55]	.	
heshadri 2020	0.5	0.7142	25	0.2	0.7142	24	21.8%	0.30 [-0.10, 0.70]		
EAK 2006	-0.1	0.5	25	0.3	0.5	24	24.1%	-0.40 [-0.68 , -0.12]	•	
otal (95% CI)			280			310	100.0%	-0.12 [-0.55 , 0.31]		
eterogeneity: Tau ² = 0.18	; Chi ² = 25.11, df	= 15 (P = 0.05);	$I^2 = 40\%$						Ī	
est for overall effect: Z =	0.55 (P = 0.59)								-20 -10 0 10 2	
est for subgroup difference	es: Not applicable	!							ner with exercise Higher with co	

- (1) mean and standard deviation estimated from the median and interquartile range
- (2) three intervention arms pooled together in the exercise group



Analysis 1.16. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 16: Calcium

	Control				Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]	
Kouidi 1997	2.15	0.25	11	2.2	0.7	20	0.7%	-0.05 [-0.39 , 0.29]		
Marinho 2016 (1)	2.2	0.367	7	2.2	0.19	6	0.8%	0.00 [-0.31, 0.31]		
Deligiannis 1999a (2)	2.02	0.2	12	2.206	0.62	25	1.1%	-0.19 [-0.45 , 0.08]		
de Lima 2013 (2)	2.32	0.22	11	2.324	0.53	21	1.2%	-0.00 [-0.27 , 0.26]		
Deligiannis 1999	2.17	0.15	30	2.2	0.7	30	1.3%	-0.03 [-0.29 , 0.23]		
Yurtkuran 2007	2.37	0.27	18	2.37	0.3	19	2.4%	0.00 [-0.18, 0.18]		
Kouidi 2010	2.22	0.32	20	2.17	0.25	24	2.8%	0.05 [-0.12, 0.22]		
Martins do Valle 2020	2.4	0.15	12	2.22	0.25	12	3.0%	0.18 [0.02, 0.34]		
Soliman 2015	1.876	0.24	12	1.87	0.19	18	3.1%	0.01 [-0.16, 0.17]		
Makhlough 2012	2.2	0.18	23	2.2	0.26	25	5.2%	0.00 [-0.13, 0.13]		
ACTINUT 2013	2.29	0.12	9	2.25	0.12	7	5.8%	0.04 [-0.08, 0.16]		
Wilund 2010	2.2	0.15	9	2.21	0.06	8	7.2%	-0.01 [-0.12, 0.10]		
Momeni 2014	2.27	0.15	20	2.24	0.14	20	10.2%	0.03 [-0.06, 0.12]		
Paluchamy 2018	2.13	0.1	10	2.09	0.09	10	11.8%	0.04 [-0.04, 0.12]		
Liao 2016	2.45	0.1	20	2.42	0.15	20	13.2%	0.03 [-0.05, 0.11]	-	
Kouidi 2008	2.1457	0.15	29	2.07	0.15	30	14.0%	0.08 [-0.00, 0.15]		
Abreu 2017	2.22	0.12	19	2.22	0.12	25	16.0%	0.00 [-0.07, 0.07]	+	
Total (95% CI)	0. Ch:2 = 0.20 df = 1	C (D = 0.00), 12 = 0	272			320	100.0%	0.03 [-0.00, 0.06]	*	
Heterogeneity: Tau ² = 0.0		o (P = 0.90); I ² = 0	17/0							
Test for overall effect: Z =	. ,								0.5 -0.25 0 0.25 0.5	
Test for subgroup differer	ices: ivot applicable							High	her with exercise Higher with contr	

(1) mean and standard deviation estimated from the median and the interquartile range

(2) two intervention arms pooled together in the exercise group

Analysis 1.17. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 17: C-reactive protein

		Control		F	Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mg/dL]	SD [mg/dL]	Total	Mean [mg/dL]	SD [mg/dL]	Total	Weight	IV, Random, 95% CI [mg/dL]	IV, Random, 95% CI [mg/dL]	
Marinho 2016 (1)	6.43	11.298	7	5.93	7.91	6	0.2%	0.50 [-9.99 , 10.99]		
Pellizzaro 2013 (1)	2.2	6.01	14	-3.73	6.75	14	0.8%	5.93 [1.20, 10.66]		
CTINUT 2013	4.99	5.96	9	1.75	1.62	7	1.1%	3.24 [-0.83 , 7.31]	 	
breu 2017	8.4	7.5	19	5.8	4.4	25	1.3%	2.60 [-1.19, 6.39]	 • •	
oussaint 2008	4.3	0.7	10	7.3	5.5	9	1.4%	-3.00 [-6.62 , 0.62]		
fshar 2010 (2)	4.14	3.87	7	1.575	1.47	14	2.0%	2.56 [-0.40 , 5.53]		
fshar 2011	4.1	3.9	7	0.93	0.66	14	2.0%	3.17 [0.26, 6.08]		
opple 2007 (3)	2.8	2.9933	14	4.363	5.67	43	3.0%	-1.56 [-3.87, 0.75]	-	
HOPE 2019	1.136	0.776	12	1.317	1.18	11	11.5%	-0.18 [-1.01 , 0.64]	.	
rih 2017a	4	1.4	20	4.1	1.2	21	11.8%	-0.10 [-0.90 , 0.70]	+	
Vilund 2010	6	0.67	9	4.9	0.69	8	13.6%	1.10 [0.45 , 1.75]	-	
VANTE-HEMO 2020 (4)	0.38	0.349	13	0.76	0.95	21	16.1%	-0.38 [-0.83 , 0.07]		
iao 2016	1.23	0.211	20	0.78	0.83	20	17.0%	0.45 [0.07, 0.83]		
Jchiyama 2019	0.3	0.5	23	0.14	0.25	24	18.4%	0.16 [-0.07 , 0.39]	+	
otal (95% CI)			184			237	100.0%	0.31 [-0.13, 0.74]		
leterogeneity: Tau ² = 0.26	; Chi ² = 38.01, df	= 13 (P = 0.000	3); I ² = 669	%					'	
est for overall effect: Z =	1.37 (P = 0.17)							_	20 -10 0 10	

Footnotes

- (1) mean and standard deviation estimated from the median and the interquartile range
- (2) Two intervention arms pooled together in the exercise group $% \left\{ 1,2,...,n\right\}$

Test for subgroup differences: Not applicable

- (3) three intervention arms pooled together in the exercise group
- (4) Mean and standard deviation estimated from the median and the interquartile range

Higher with exercise

Higher with control



Analysis 1.18. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 18: Dialysis adequacy: Kt/V

		Control]	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Reboredo 2010	1.8	0.7	11	2	0.8	11	1.5%	-0.20 [-0.83 , 0.43]	
Parsons 2004	1.22	0.21	7	1.27	0.31	6	5.4%	-0.05 [-0.34 , 0.24]	
Frey 1999	1.7	0.1	6	1.8	0.3	5	5.9%	-0.10 [-0.37, 0.17]	
Dobsak 2012	1.33	0.31	10	1.64	0.3	11	6.3%	-0.31 [-0.57 , -0.05]	
PEAK 2006	0.1	0.4	25	0.1	0.5	24	6.6%	0.00 [-0.25 , 0.25]	
Martins do Valle 2020	1.6	0.3	12	1.7	0.3	12	7.1%	-0.10 [-0.34, 0.14]	
Makhlough 2012	0.95	0.2	24	1.2	0.4	23	9.8%	-0.25 [-0.43 , -0.07]	_ -
Liao 2016	1.52	0.23	20	1.52	0.26	20	11.6%	0.00 [-0.15, 0.15]	-
Fernandes 2019	1.48	0.19	19	1.36	0.19	20	13.8%	0.12 [0.00, 0.24]	-
Paluchamy 2018	0.99	0.1265	10	1.15	0.0949	10	15.4%	-0.16 [-0.26 , -0.06]	
van Vilsteren 2005	1.23	0.2	43	1.26	0.2	53	16.6%	-0.03 [-0.11 , 0.05]	+
Total (95% CI)			187			195	100.0%	-0.08 [-0.16 , 0.00]	
Heterogeneity: Tau ² = 0.0	1; Chi ² = 22.1	17, df = 10	(P = 0.01)	; I ² = 55%					•
Test for overall effect: Z =	= 1.87 (P = 0.0	06)							-1 -0.5 0 0.5 1
Test for subgroup differen	ices: Not appl	icable						Hi	gher with exercise Higher with control

Analysis 1.19. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 19: Energy intake

		Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Weight	IV, Random, 95% CI [kCal/kg/d]	IV, Random, 95% CI [kCal/kg/d]
PEAK 2006	30.05	8.57	25	41.44	37.04	24	1.0%	-11.39 [-26.58 , 3.80]	
Dong 2011	27.6	11.9	12	26.5	7.1	10	3.4%	1.10 [-6.94, 9.14]	
ACTINUT 2013	27.53	8.42	9	30.09	6.64	7	4.1%	-2.56 [-9.94 , 4.82]	-
Kopple 2007 (1)	24.2	7.8575	14	25.96	18.2	43	4.8%	-1.76 [-8.58, 5.06]	-
Olvera-Soto 2016 (2)	23.77	13	31	23.17	9.65	30	6.8%	0.60 [-5.13, 6.33]	
IHOPE 2019	19.03	8.5	38	20.04	11.8	29	8.6%	-1.01 [-6.08 , 4.06]	4
Abreu 2017	30.9	3	19	30.6	2.9	25	71.4%	0.30 [-1.46 , 2.06]	•
Total (95% CI)			148			168	100.0%	-0.09 [-1.58 , 1.40]	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 3.24, df = 6	(P = 0.78); I ² = 0%							ľ
Test for overall effect: 2	Z = 0.12 (P = 0.90)								-50 -25 0 25 50
Test for subgroup differ	rences: Not applicable							Hig	gher with exercise Higher with cont

⁽¹⁾ three intervention arms pooled together in the exercise group

⁽²⁾ mean and standard deviation estimated from the median and interquartile range



Analysis 1.20. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 20: Haemoglobin

	(Control		E	xercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]	
Soliman 2015	9.19	0.89	12	11.04	8.24	18	0.1%	-1.85 [-5.69 , 1.99]		
Marinho 2016 (1)	11.2	2.756	7	10.467	2.098	6	0.2%	0.73 [-1.91, 3.38]		
Reboredo 2010	11.3	2.6	11	10.9	2.8	11	0.3%	0.40 [-1.86 , 2.66]		
Oong 2011	12.2	1.8	12	11.4	1.9	10	0.6%	0.80 [-0.76 , 2.36]	 	
Parsons 2004	11.1	1.7	7	11.7	0.7	6	0.8%	-0.60 [-1.98 , 0.78]	-	
Goldberg 1983	8.8	1.3	7	10	1.5	9	0.8%	-1.20 [-2.57 , 0.17]		
VANTE-HEMO 2020	10.6	1.8	13	10.61	2.23	21	0.8%	-0.01 [-1.38 , 1.36]		
aluchamy 2018	8.09	1.4546	10	8.35	1.6128	10	0.8%	-0.26 [-1.61 , 1.09]		
Martin-Alemany 2016 (1)	8.6	2.102	17	8.9	1.842	19	0.9%	-0.30 [-1.60 , 1.00]		
Abreu 2017	10.6	2.3	19	10.5	1.9	25	0.9%	0.10 [-1.17 , 1.37]		
Martins do Valle 2020	10.3	1.3	12	10.8	1.7	12	1.0%	-0.50 [-1.71, 0.71]		
Makhlough 2012	8.62	2.14	23	9.87	2.01	25	1.1%	-1.25 [-2.43, -0.07]		
Momeni 2014	9.8	1.7	20	10.03	2.07	20	1.1%	-0.23 [-1.40, 0.94]		
Copple 2007 (2)	12.5	1.87	14	12.77	1.09	37	1.4%	-0.27 [-1.31 , 0.77]		
rih 2017a	10	1.6	20	10.4	1.7	21	1.5%	-0.40 [-1.41, 0.61]		
ainter 2002a (3)	11.81	1.56	26	12.2	1.92	22	1.5%	-0.39 [-1.39, 0.61]	_	
CHAIR 2015 (4)	10.93	0.72	11	11.23	1.13	6	1.5%	-0.30 [-1.30 , 0.70]		
Oussaint 2008	12.03	1.41	10	12.09	0.59	9	1.7%	-0.06 [-1.02, 0.90]		
Koufaki 2002	12.2	1.4	15	12.1	1.3	18	1.8%	0.10 [-0.83 , 1.03]		
Fernandes 2019	10.25	1.66	19	10.7	1.14	20	1.9%	-0.45 [-1.35 , 0.45]	_	
e Lima 2013 (5)	11.1	1.2	11	10.82	1.04	21	2.2%	0.28 [-0.56 , 1.12]	<u>.</u>	
suyuki 2003	7.4	1.2	12	8	1	17	2.2%	-0.60 [-1.43, 0.23]	-	
Kouidi 2010	11.2	1.3	20	11.3	1.2	24	2.7%	-0.10 [-0.85, 0.65]		
ee 2001	8.7	1.3	21	8.2	1.1	19	2.7%	0.50 [-0.24 , 1.24]	-	
CTINUT 2013	11.21	0.66	9	10.92	0.69	7	3.4%	0.29 [-0.38, 0.96]	-	
Kouidi 2008	11	0.7	29	11	0.7	30	11.9%	0.00 [-0.36, 0.36]	.	
an Vilsteren 2005	7.57	0.8	43	7.52	0.8	53	14.7%	0.05 [-0.27, 0.37]		
Pellizzaro 2013	0.5	0.4	14	0.6	0.4	14	17.3%	-0.10 [-0.40 , 0.20]		
Afshar 2010 (6)	10.2	0.3	7	10.2	0.27	14	21.9%	0.00 [-0.26 , 0.26]	•	
Total (95% CI)			451			524	100.0%	-0.06 [-0.18 , 0.06]		
Heterogeneity: Tau ² = 0.00	; Chi ² = 18.94,	df = 28 (P =	0.90); I ² =	= 0%						
Test for overall effect: Z =	0.93 (P = 0.35)								-10 -5 0 5	
est for subgroup differenc	es: Not applica	ble							her with exercise Higher with	

Footnotes

- (1) mean and standard deviation estimated from the median and interquartile range
- (2) three intervention arms pooled together in the exercise group
- (3) factorial design: two intervention arms pooled together in the exercise group and two control arms pooled together in the control group
- (4) mean and standard deviation estimated from the median and range
- (5) two intervention arms pooled together in the exercise group
- (6) Two intervention arms pooled together in the exercise group $% \left\{ 1\right\} =\left\{ 1\right\} =\left\{$

Analysis 1.21. Comparison 1: Any exercise versus control (no exercise/ placebo exercise), Outcome 21: Left ventricular ejection fraction

		Control		1	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [%]	SD [%]	Total	Mean [%]	SD [%]	Total	Weight	IV, Random, 95% CI [%]	IV, Random, 95% CI [%]
Reboredo 2010	71.4	7.6	11	70.4	12	11	5.7%	1.00 [-7.39 , 9.39	9]
Kouidi 2008	54.6	17.3	29	60.3	13	30	6.4%	-5.70 [-13.53 , 2.13	3]
McGregor 2018	53.22	7.1186	18	54.22	9.8337	16	10.3%	-1.00 [-6.83 , 4.83	3]
Cho 2018 (1)	0.3	3.9	13	0.512	5.32	33	24.8%	-0.21 [-3.00 , 2.58	B] _ _
Momeni 2014	54.25	4.66	20	58.5	3.67	20	26.3%	-4.25 [-6.85 , -1.65	5] 🕳
Kouidi 2004a	0.2	3	10	-0.3	3	11	26.5%	0.50 [-2.07 , 3.0]	7]
Total (95% CI)			101			121	100.0%	-1.45 [-3.60 , 0.70	0]
Heterogeneity: Tau ² =	2.83; Chi ² = 8.9	1, df = 5 (P	= 0.11); I	$^{2} = 44\%$					•
Test for overall effect: $Z = 1.32$ ($P = 0.19$)									-20 -10 0 10 20
est for subgroup differences: Not applicable								Higher with exercise Higher with cont	

Footnotes

(1) Three interventions arms pooled together in the exercise group



Analysis 1.22. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 22: Left ventricular mass index

		Control		E	exrcise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/m²]	SD [g/m ²]	Total	Mean [g/m²]	SD [g/m ²]	Total	Weight	IV, Random, 95% CI [g/m²]	IV, Random, 95% CI [g/m²]
Reboredo 2010	131.3	48.4	11	120.9	26.6	11	8.4%	10.40 [-22.24 , 43.04]	
McGregor 2018	111.65	39.7958	18	139.2	45.6215	16	10.1%	-27.55 [-56.49 , 1.39	
Deligiannis 1999a (1)	137	35	12	147.6	40.22	25	12.2%	-10.60 [-35.91 , 14.71]	ı <u> </u>
Wilund 2010	127.4	18.17	9	154.4	25.83	8	15.1%	-27.00 [-48.48 , -5.52]	_
Cho 2018 (2)	-11.7	30.1	13	-0.44	21.1	33	18.7%	-11.26 [-29.14 , 6.62]	_
Kouidi 2008	137	11.9	29	138.3	10.1	30	35.5%	-1.30 [-6.94 , 4.34]	•
Total (95% CI)			92			123	100.0%	-9.85 [-20.50 , 0.80]	•
Heterogeneity: Tau ² = 74	4.81; Chi ² = 9.46,	df = 5 (P = 0.6)	09); I ² = 47	7%					· · · · · · · · · · · · · · · · · · ·
Test for overall effect: Z	= 1.81 (P = 0.07))							-100 -50 0 50 100
Test for subgroup differen	ences: Not applica	able							Higher in exercise Higher in control

Footnotes

- (1) two intervention arms pooled together in the exercise group
- $\ensuremath{\text{(2)}}\xspace Three interventions arms pooled together in the exercise group$

Analysis 1.23. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 23: Maximum heart rate

		Control		F	exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [bpm]	SD [bpm]	Total	Mean [bpm]	SD [bpm]	Total	Weight	IV, Random, 95% CI [bpm]	IV, Random, 95% CI [bpn	
McGregor 2018	122.5	27.107	18	126.88	23.6646	16	5.2%	-4.38 [-21.45 , 12.69]]	
Akiba 1995	136.3	19.5	6	155.4	8.6	7	5.4%	-19.10 [-35.95 , -2.25]	ı <u> </u>	
Koufaki 2002	127.2	24.4	15	129	22.7	18	5.8%	-1.80 [-18.00 , 14.40]]	
Painter 2002a (1)	128.72	27.44	25	137.78	24.68	23	7.0%	-9.06 [-23.81, 5.69]	1	
Tsuyuki 2003	155.8	20.7	12	164.2	10.2	17	9.5%	-8.40 [-21.08 , 4.28]	1	
Deligiannis 1999a (2)	139	12	12	144.4	16.58	25	17.3%	-5.40 [-14.80 , 4.00]	1	
Konstantinidou 2002 (3)	139	12	12	144.3	14.52	36	22.2%	-5.30 [-13.58 , 2.98]	1	
Ouzouni 2009	139.6	7.1	14	144.1	14.3	19	27.6%	-4.50 [-11.93 , 2.93]] —	
Total (95% CI)			114			161	100.0%	-6.14 [-10.05 , -2.24]	1 📥	
Heterogeneity: Tau ² = 0.0	0; Chi ² = 3.11, df	E = 7 (P = 0.87)); $I^2 = 0\%$						•	
Test for overall effect: Z =	= 3.08 (P = 0.002))							-50 -25 0 25	
Test for subgroup differer	nces: Not applicat	ole						Н	ligher with exercise Higher wi	

- (1) factorial design: two intervention arms pooled together in the exercise group and two control arms pooled together in the control group
- (2) two intervention arms pooled together in the exercise group $% \left\{ 1,2,...,n\right\}$
- (3) three intervention arms pooled together in the exercise group $% \left\{ 1,2,...,n\right\}$



Analysis 1.24. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 24: Muscular strength

	(Control		I	Exercise			Mean Difference	Mean Differen	ce
Study or Subgroup	Mean [kg]	SD [kg]	Total	Mean [kg]	SD [kg]	Total	Weight	IV, Random, 95% CI [kg	IV, Random, 95% (CI [kg]
1.24.1 Knee extension										
Dobsak 2012	85.36	18.32	10	104.68	14.23	11	4.9%	-19.32 [-33.45 , -5.1	9]	
Song 2012a	33.4	19.5	20	37.3	19	20	6.3%	-3.90 [-15.83, 8.0	[3]	
PEAK 2006	-2.4	13.8	25	15.2	15.4	24	10.1%	-17.60 [-25.80 , -9.4	0]	
DIALY-SIZE 2016 (1)	9.3	10.1	8	8.448	9.32	23	10.5%	0.85 [-7.12, 8.8	[2]	
Uchiyama 2019	24.9	10.5	23	24.2	10.7	24	13.5%	0.70 [-5.36, 6.7	[6]	
Johansen 2006 (2)	17.28	8.63	33	23.65	10.18	35	16.5%	-6.37 [-10.85 , -1.8	9] 🕳	
Chen 2010	12.1	6.1	22	15.8	5	22	18.8%	-3.70 [-7.00 , -0.4	0]	
ACTINUT 2013	7.87	2.19	9	10.56	3.49	7	19.4%	-2.69 [-5.64, 0.2	[6]	
Subtotal (95% CI)			150			166	100.0%	-5.06 [-8.58 , -1.5	i4] •	
Heterogeneity: Tau ² = 14.3	32; Chi ² = 20.6	68, df = 7 (P)	= 0.004);	$I^2 = 66\%$					"	
Test for overall effect: Z =	2.81 (P = 0.00)5)								
1.24.2 Handgrip										
Yurtkuran 2007	62.86	20.36	18	78.45	23.09	19	2.7%	-15.59 [-29.60 , -1.5	[8]	
Koh 2009 (3)	31	. 12	7	35.97	12.35	29	4.9%	-4.97 [-14.93 , 4.9	9]	
Samara 2016	32.3	9.9	12	37.2	14.7	15	5.4%	-4.90 [-14.21 , 4.4	1]	
Martin-Alemany 2016 (4)	20.867	9.701	17	23.5	10.813	19	8.7%	-2.63 [-9.33 , 4.0	7]	
Song 2012a	27.8	11.8	20	28.7	9	20	9.0%	-0.90 [-7.40 , 5.6	[0]	
Olvera-Soto 2016 (4)	19.6	8.56	31	22.07	14.77	30	9.8%	-2.47 [-8.55, 3.6	[1]	
Wu 2014d	28.6	9	33	37.8	12.9	32	11.2%	-9.20 [-14.62 , -3.7	[8]	
Cooke 2018 (4)	2	3.87	10	2.43	6.02	10	13.7%	-0.43 [-4.87 , 4.0	1]	
Uchiyama 2019	25.7	6.4	23	27.3	5.4	24	16.8%	-1.60 [-4.99, 1.7	9]	
Frih 2017a	30	5.2	20	37.4	4.8	21	17.8%	-7.40 [-10.47 , -4.3	3]	
Subtotal (95% CI)			191			219	100.0%	-4.16 [-6.61 , -1.7	[1]	
Heterogeneity: Tau ² = 6.31	; Chi ² = 16.57	, df = 9 (P =	0.06); I ²	= 46%					1	
Test for overall effect: Z =	3.33 (P = 0.00	009)								
									-100 -50 0	50 10
Footnotes										50 10 her with co
									J	,

⁽¹⁾ three intervention arms pooled together in the exercise group

⁽²⁾ Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

⁽³⁾ two intervention arms polled together in the exercise group

⁽⁴⁾ mean and standard deviation estimated from the median and interquartile range



Analysis 1.25. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 25: Phosphate

	(Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Marinho 2016 (1)	1.367	1.653	7	1.467	0.57	6	0.7%	-0.10 [-1.41 , 1.21]	
AVANTE-HEMO 2020 (2)	1.86	1.15	13	2.1	0.83	21	2.0%	-0.24 [-0.96, 0.48]	
Martins do Valle 2020	1.84	0.39	12	1.61	0.97	12	2.7%	0.23 [-0.36, 0.82]	
Reboredo 2010	1.91	0.61	11	1.58	0.55	11	3.4%	0.33 [-0.16, 0.82]	
Yurtkuran 2007	1.81	0.45	18	1.91	0.81	19	4.0%	-0.10 [-0.52 , 0.32]	
ACTINUT 2013	1.59	0.42	9	1.32	0.42	7	4.1%	0.27 [-0.14, 0.68]	
Martin-Alemany 2016	1.84	0.61	19	2.07	0.65	17	4.1%	-0.23 [-0.64, 0.18]	
Makhlough 2012	2.29	0.67	23	1.88	0.77	25	4.2%	0.41 [0.00, 0.82]	-
Kouidi 2010	2.1	0.52	20	2.13	0.55	24	5.2%	-0.03 [-0.35 , 0.29]	-
Paluchamy 2018	1.6855	0.4799	10	1.4853	0.143	10	5.3%	0.20 [-0.11, 0.51]	
Deligiannis 1999a (3)	1.94	0.42	12	2.05	0.49	25	5.3%	-0.11 [-0.42 , 0.20]	
de Lima 2013 (3)	1.81	0.29	11	1.93	0.56	21	5.5%	-0.12 [-0.41 , 0.17]	
Kouidi 1997	1.97	0.23	11	2.03	0.55	20	5.7%	-0.06 [-0.34 , 0.22]	
Soliman 2015	1.776	0.38	12	1.72	0.35	18	5.8%	0.06 [-0.21, 0.33]	-
Abreu 2017	2.1	0.42	19	2.2	0.48	25	5.8%	-0.10 [-0.37 , 0.17]	
Deligiannis 1999	1.97	0.36	30	2	0.55	30	6.2%	-0.03 [-0.27 , 0.21]	4
Wilund 2010	2.1	0.25	9	1.91	0.16	8	6.7%	0.19 [-0.01, 0.39]	-
Kouidi 2008	1.97	0.26	29	2	0.36	30	7.2%	-0.03 [-0.19 , 0.13]	4
Momeni 2014	1.69	0.27	20	1.78	0.03	20	7.7%	-0.09 [-0.21 , 0.03]	-
Pellizzaro 2013	0.19	0.06	14	-0.13	0.06	14	8.2%	0.32 [0.28, 0.36]	•
Total (95% CI)			309			363	100.0%	0.05 [-0.07, 0.16]	•
Heterogeneity: Tau ² = 0.04;		9 (P < 0.00001);	$I^2 = 79\%$						
Test for overall effect: $Z = 0$									-2 -1 0 1 2
Test for subgroup difference	es: Not applicable							High	ner with exercise Higher with control

- (1) mean and standard deviation estimated from the median and interquartile range
- (2) Mean and standard deviation estimated from the median and the interquartile range
- (3) two intervention arms pooled together in the exercise group $% \left\{ 1,2,...,n\right\}$

Analysis 1.26. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 26: Potassium

	•	Control		I	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Marinho 2016 (1)	4.5	2.848	7	5.067	1.62	6	1.1%	-0.57 [-3.04 , 1.91]	
Martin-Alemany 2016 (1)	5.93	1.213	17	5.1	0.481	19	5.1%	0.83 [0.21, 1.45]	
Parsons 2004	4.8	0.6	7	4.4	0.4	6	5.3%	0.40 [-0.15, 0.95]	-
Momeni 2014	5.47	0.95	20	5.4	0.81	20	5.3%	0.07 [-0.48 , 0.62]	
AVANTE-HEMO 2020	5.2	0.92	13	5.54	0.37	21	5.4%	-0.34 [-0.86, 0.18]	
Kouidi 1997	5.5	0.7	11	5.6	0.5	20	5.6%	-0.10 [-0.57 , 0.37]	-
Makhlough 2012	5.16	0.67	23	5.12	0.96	25	5.6%	0.04 [-0.43 , 0.51]	+
de Lima 2013	5.8	0.6	11	5.44	0.7	21	5.6%	0.36 [-0.10, 0.82]	
Paluchamy 2018	5.65	0.5692	10	5.18	0.4743	10	5.6%	0.47 [0.01, 0.93]	-
Martins do Valle 2020	5.3	0.4	12	5	0.6	12	5.8%	0.30 [-0.11, 0.71]	 -
Kouidi 2010	5.8	0.6	20	5.6	0.7	24	5.9%	0.20 [-0.18, 0.58]	 -
Soliman 2015	5.61	0.45	12	4.34	0.45	18	6.1%	1.27 [0.94, 1.60]	
Deligiannis 1999a (2)	5.4	0.4	12	5.6	0.58	25	6.1%	-0.20 [-0.52, 0.12]	-
Deligiannis 1999	5.7	0.7	30	5.8	0.5	30	6.1%	-0.10 [-0.41, 0.21]	4
Wilund 2010	4.9	0.3	9	4.9	0.25	8	6.2%	0.00 [-0.26, 0.26]	+
Kouidi 2008	5.3	0.4	29	5.6	0.6	30	6.3%	-0.30 [-0.56 , -0.04]	-
Abreu 2017	4.7	0.3	19	4.7	0.4	25	6.4%	0.00 [-0.21, 0.21]	+
Pellizzaro 2013	0.6	0.2	14	-0.5	0.2	14	6.5%	1.10 [0.95, 1.25]	•
Total (95% CI)			276			334	100.0%	0.23 [-0.06, 0.51]	•
Heterogeneity: Tau ² = 0.33; Chi ² = 206.50, df = 17 (P < 0.00001); I ² = 92%									
Test for overall effect: Z = Test for subgroup differenc	. ,							Hig	-4 -2 0 2 4 her with exercise Higher with cor

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range

(2) two intervention arms pooled together in the exercise group



Analysis 1.27. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 27: Protein intake

		Control		E	ercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/kg/d]	SD [g/kg/d]	Total	Mean [g/kg/d]	SD [g/kg/d]	Total	Weight	IV, Random, 95% CI [g/kg/d]	IV, Random, 95% CI [g/kg/d]
Kopple 2007 (1)	1.35	0.5987	14	1.02	0.68	43	4.6%	0.33 [-0.04 , 0.70]	
ACTINUT 2013	1.17	0.38	9	1.17	0.26	7	6.4%	0.00 [-0.31, 0.31]	
Dong 2011	1.1	0.4	12	1	0.3	10	7.3%	0.10 [-0.19, 0.39]	
Olvera-Soto 2016 (2)	1.03	0.4	31	1.01	0.62	30	8.9%	0.02 [-0.24, 0.28]	
IHOPE 2019	0.9	0.33	38	1	0.46	29	14.8%	-0.10 [-0.30 , 0.10]	
PEAK 2006	1.36	0.38	25	1.51	0.25	24	17.4%	-0.15 [-0.33 , 0.03]	
Abreu 2017	1.2	0.14	19	1.2	0.2	25	40.5%	0.00 [-0.10, 0.10]	+
Total (95% CI)			148			168	100.0%	-0.02 [-0.10 , 0.07]	
Heterogeneity: Tau ² = 0	.00; Chi ² = 6.91, df	= 6 (P = 0.33); I	2 = 13%						Ĭ
Test for overall effect: Z	t = 0.40 (P = 0.69)								-1 -0.5 0 0.5 1
Test for subgroup differ	ences: Not applicab	le						High	er with exercise Higher with control

Footnotes

- (1) three intervention arms pooled together in the exercise group
- (2) mean and standard deviation estimated from the median and interquartile range

Analysis 1.28. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 28: Parathyroid hormone

		Control		E	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [pmol/L]	SD [pmol/L]	Total	Mean [pmol/L]	SD [pmol/L]	Total	Weight	IV, Random, 95% CI [pmol/L]	IV, Random, 95% CI [pmol/L]
Martins do Valle 2020	56.28	37.28	12	80.64	84.74	12	4.3%	-24.36 [-76.74 , 28.02]	-
Toussaint 2008	42.7	57.2	10	37.5	45.9	9	5.4%	5.20 [-41.23 , 51.63]	_
Koufaki 2002	13.6	15.8	15	34.8	50.7	18	15.9%	-21.20 [-45.95 , 3.55]	-
Marinho 2016 (1)	28.11	28.06	7	16.37	14.93	6	16.6%	11.74 [-12.24, 35.72]	-
Liao 2016	30.2	9.42	20	25.74	7.92	20	57.7%	4.46 [-0.93 , 9.85]	•
Total (95% CI)			64			65	100.0%	0.39 [-10.90 , 11.68]	•
Heterogeneity: Tau ² = 49.	91; Chi ² = 5.49, df =	4 (P = 0.24); I ² =	27%						
Test for overall effect: Z =	= 0.07 (P = 0.95)								-500 -250 0 250 500
Test for subgroup differen	ices: Not applicable							Hig	ther with exercise Higher with control

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range

Analysis 1.29. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 29: Resting heart rate

	(Control		E	xercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [bpm]	SD [bpm]	Total	Mean [bpm]	SD [bpm]	Total	Weight	IV, Random, 95% CI [bpm]	IV, Random, 95% CI [bpm]
Liao 2016	71	10.5	20	51.9	34.3	20	1.4%	19.10 [3.38 , 34.82]	
Goldberg 1983	79	15	11	83	11	14	3.0%	-4.00 [-14.57 , 6.57]	
CYCLE-HD 2016	75.31	13.95	13	72.45	9.36	9	3.6%	2.86 [-6.88 , 12.60]	_ -
McGregor 2018	80.94	11.7854	16	74	14.6193	18	4.3%	6.94 [-1.95 , 15.83]	 -
Tsuyuki 2003	84.3	13.6	12	81.9	8.7	17	4.4%	2.40 [-6.34 , 11.14]	_ -
Koh 2009 (1)	75	12	16	70	10.38	30	7.0%	5.00 [-1.95 , 11.95]	 -
Ouzouni 2009	78.2	10.3	14	76.3	7.1	19	8.6%	1.90 [-4.37 , 8.17]	-
Deligiannis 1999a	81.8	8.5	12	77.74	9.43	25	9.2%	4.06 [-2.01 , 10.13]	 -
Cooke 2018 (2)	2.33	6.45	10	-3.767	4.73	10	13.7%	6.10 [1.14 , 11.05]	
Deligiannis 1999	76	7	30	75	9	30	20.3%	1.00 [-3.08, 5.08]	-
Kouidi 2008	78.4	8.1	29	73.7	6.3	30	24.5%	4.70 [0.99 , 8.41]	-
Total (95% CI)			183			222	100.0%	3.72 [1.89 , 5.56]	•
Heterogeneity: Tau ² = 0	.00; Chi ² = 9.67, d		 *						
Test for overall effect: Z	L = 3.97 (P < 0.000)		-50 -25 0 25 50						
Test for subgroup differ	ences: Not applica	Hig	her with exercise Higher with cor						

- (1) two intervention arms polled together in the exercise group
- (2) mean and standard deviation estimated from the median and interquartile range



Analysis 1.30. Comparison 1: Any exercise versus control (no exercise/placebo exercise), Outcome 30: Timed up-and-go test

	(Control		E	xercise			Mean Difference	Mean Di	fference
Study or Subgroup	Mean [sec]	SD [sec]	Total	Mean [sec]	SD [sec]	Total	Weight	IV, Random, 95% CI [sec]	IV, Random,	95% CI [sec]
Bennett 2013 (1)	11.7	79.3115	12	9.73	65.329	28	0.0%	1.97 [-49.01 , 52.95]	+	
Wu 2014d	30.2	8.3	33	27.3	7.3	32	3.5%	2.90 [-0.90 , 6.70]	4	
Samara 2016	6.6	2.5	12	4.6	1.4	15	16.3%	2.00 [0.42 , 3.58]		-
IHOPE 2019	8	3.6	38	6.2	1.7	29	21.7%	1.80 [0.50, 3.10]		-
Frih 2017a	15.2	1.9	20	12.9	1.6	21	27.7%	2.30 [1.22 , 3.38]		•
Koh 2009 (2)	6.1	1.5	16	5.54	1.8	29	30.7%	0.56 [-0.42 , 1.54]	•	•
Total (95% CI)			131			154	100.0%	1.63 [0.90 , 2.36]		•
Heterogeneity: Tau ² = 0	.20; Chi ² = 6.68,	df = 5 (P =	0.25); I ² =	25%						Y
Test for overall effect: Z	Z = 4.36 (P < 0.0)	001)							-20 -10 0	10 20
Test for subgroup differ	ences: Not appli	cable						Hi	gher with exercise	Higher with control

Footnote

(1) results from group 1 (24 weeks of intervention) and group 2 (12 weeks of intervention) were pooled together in the exercise group. The number of participants was corrected to account for (2) two intervention arms polled together in the exercise group

Comparison 2. Aerobic exercise versus control (no exercise/placebo exercise)

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2.1 Death	1	296	Risk Ratio (M-H, Random, 95% CI)	0.95 [0.56, 1.62]
2.2 Fatigue	4		Std. Mean Difference (IV, Random, 95% CI)	Totals not selected
2.3 HRQoL: Summary component scores	9		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.3.1 Physical Component Score	9	306	Mean Difference (IV, Random, 95% CI)	-6.00 [-10.71, -1.30]
2.3.2 Mental Component Score	9	306	Mean Difference (IV, Random, 95% CI)	-3.33 [-7.56, 0.90]
2.4 HRQoL: Individual do- mains	11		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.4.1 Physical Functioning	10	649	Mean Difference (IV, Random, 95% CI)	-2.87 [-10.12, 4.38]
2.4.2 Role-physical	8	560	Mean Difference (IV, Random, 95% CI)	-2.31 [-17.29, 12.67]
2.4.3 Pain	8	570	Mean Difference (IV, Random, 95% CI)	-2.26 [-6.12, 1.61]
2.4.4 General health perceptions	8	560	Mean Difference (IV, Random, 95% CI)	-5.38 [-10.32, -0.43]
2.4.5 Emotional well-being	7	515	Mean Difference (IV, Random, 95% CI)	-5.63 [-10.58, -0.67]
2.4.6 Role-emotional	8	560	Mean Difference (IV, Random, 95% CI)	-8.02 [-11.45, -4.58]
2.4.7 Vitality	9	613	Mean Difference (IV, Random, 95% CI)	-0.43 [-6.45, 5.60]



Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
2.4.8 Social function	9	577	Mean Difference (IV, Random, 95% CI)	0.94 [-4.48, 6.37]
2.4.9 Symptoms	3	317	Mean Difference (IV, Random, 95% CI)	-7.65 [-20.65, 5.35]
2.4.10 Effects of kidney disease	3	317	Mean Difference (IV, Random, 95% CI)	-4.27 [-6.87, -1.66]
2.4.11 Burden of kidney disease	3	317	Mean Difference (IV, Random, 95% CI)	0.05 [-2.63, 2.72]
2.4.12 Work status	3	317	Mean Difference (IV, Random, 95% CI)	-0.44 [-3.87, 2.99]
2.4.13 Cognitive function	3	317	Mean Difference (IV, Random, 95% CI)	-6.36 [-10.11, -2.60]
2.4.14 Quality of social interactions	3	317	Mean Difference (IV, Random, 95% CI)	-6.96 [-10.57, -3.36]
2.4.15 Sexual function	3	317	Mean Difference (IV, Random, 95% CI)	-0.87 [-7.23, 5.48]
2.4.16 Sleep	3	317	Mean Difference (IV, Random, 95% CI)	-6.44 [-13.46, 0.58]
2.4.17 Social support	3	317	Mean Difference (IV, Random, 95% CI)	-4.35 [-8.51, -0.19]
2.4.18 Dialysis staff en- couragement	3	317	Mean Difference (IV, Random, 95% CI)	-5.49 [-11.11, 0.13]
2.4.19 Patient satisfaction	3	317	Mean Difference (IV, Random, 95% CI)	-7.52 [-13.12, -1.92]
2.5 Depression	4	127	Std. Mean Difference (IV, Random, 95% CI)	0.19 [-0.52, 0.89]
2.6 6MWT	10	515	Mean Difference (IV, Random, 95% CI)	-53.00 [-72.17, -33.84]
2.7 Sit-To-Stand test [N reps/30 sec]	6	227	Mean Difference (IV, Random, 95% CI)	-1.81 [-2.76, -0.86]
2.8 Sit-To-Stand test [sit to 5 reps]	5	374	Mean Difference (IV, Random, 95% CI)	1.63 [0.92, 2.33]
2.9 Resting blood pressure	13		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.9.1 Systolic blood pressure	13	400	Mean Difference (IV, Random, 95% CI)	3.96 [-1.78, 9.70]
2.9.2 Diastolic blood pressure	13	400	Mean Difference (IV, Random, 95% CI)	-0.73 [-3.68, 2.22]
2.10 Aerobic capacity (VO2 max or peak)	12	326	Mean Difference (IV, Random, 95% CI)	-2.69 [-4.55, -0.82]
2.11 Albumin	15	429	Mean Difference (IV, Random, 95% CI)	-0.23 [-1.45, 0.99]
2.12 Blood lipids	5		Mean Difference (IV, Random, 95% CI)	Subtotals only



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2.12.1 Total cholesterol [mmol/L]	5	129	Mean Difference (IV, Random, 95% CI)	0.30 [-0.03, 0.63]
2.12.2 LDL cholesterol [mmol/L]	3	72	Mean Difference (IV, Random, 95% CI)	0.17 [-0.08, 0.42]
2.12.3 HDL cholesterol [mmol/L]	3	72	Mean Difference (IV, Random, 95% CI)	0.05 [0.01, 0.09]
2.12.4 Triglycerides [mmol/L]	3	72	Mean Difference (IV, Random, 95% CI)	0.23 [-0.59, 1.05]
2.13 Body composition	4		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.13.1 Fat mass [kg]	3	95	Mean Difference (IV, Random, 95% CI)	-0.04 [-0.10, 0.02]
2.13.2 Lean mass [kg]	2	89	Mean Difference (IV, Random, 95% CI)	-1.94 [-6.32, 2.45]
2.14 Body mass index	9	291	Mean Difference (IV, Random, 95% CI)	-0.17 [-0.78, 0.45]
2.15 Calcium	8	208	Mean Difference (IV, Random, 95% CI)	0.01 [-0.04, 0.06]
2.16 C-reactive protein	9	206	Mean Difference (IV, Random, 95% CI)	0.60 [-0.12, 1.32]
2.17 Dialysis adequacy: Kt/V	7	166	Mean Difference (IV, Random, 95% CI)	-0.07 [-0.20, 0.05]
2.18 Energy intake	4	118	Mean Difference (IV, Random, 95% CI)	-1.84 [-6.87, 3.20]
2.19 Haemoglobin	17	437	Mean Difference (IV, Random, 95% CI)	0.01 [-0.18, 0.21]
2.20 Heart rate	10		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.20.1 Resting	7	218	Mean Difference (IV, Random, 95% CI)	4.07 [0.49, 7.65]
2.20.2 Maximum	6	179	Mean Difference (IV, Random, 95% CI)	-6.54 [-12.01, -1.07]
2.21 Left ventricular ejec- tion fraction	5	141	Mean Difference (IV, Random, 95% CI)	-1.65 [-3.93, 0.62]
2.22 Left ventricular mass index	5	119	Mean Difference (IV, Random, 95% CI)	-14.47 [-26.25, -2.69]
2.23 Muscular strength	7		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.23.1 Knee extension	3	53	Mean Difference (IV, Random, 95% CI)	-5.94 [-13.95, 2.07]
2.23.2 handgrip	4	148	Mean Difference (IV, Random, 95% CI)	-4.65 [-9.44, 0.14]
2.24 Phosphate	9	214	Mean Difference (IV, Random, 95% CI)	0.05 [-0.07, 0.17]
2.25 Potassium	8	190	Mean Difference (IV, Random, 95% CI)	0.08 [-0.08, 0.24]
2.26 Protein intake	4	118	Mean Difference (IV, Random, 95% CI)	0.04 [-0.25, 0.33]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2.27 Parathyroid hor- mone	3	92	Mean Difference (IV, Random, 95% CI)	-2.69 [-20.31, 14.93]
2.28 Timed up-and-go test	4	204	Mean Difference (IV, Random, 95% CI)	1.38 [0.50, 2.26]

Analysis 2.1. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 1: Death

	Cont	rol	Exer			Risk Ratio	Risk Ratio			
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Random, 95% CI	M-H, Rando	om, 95% CI		
EXCITE 2014	22	145	24	151	100.0%	0.95 [0.56 , 1.62]	-	F		
Total (95% CI)		145		151	100.0%	0.95 [0.56 , 1.62]		•		
Total events:	22		24				T			
Heterogeneity: Not app	olicable						0.1 0.2 0.5 1	2 5 10		
Test for overall effect:	Z = 0.17 (P =	0.86)					Less with control	Less with exercise		
Test for subgroup diffe	rences: Not ar	onlicable								

Analysis 2.2. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 2: Fatigue

		Control]	Exercise		Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	IV, Random, 95% CI	IV, Random, 95% CI	
Chang 2010	45.5	19.66	35	41	20.09	36	0.22 [-0.24 , 0.69)]	
Amini 2016	6.2	2.15	35	4.37	1.62	32	0.94 [0.44, 1.45	5] —	
Soliman 2015	29.75	5.19	12	14.44	5.29	18	2.84 [1.78, 3.90	0]	
Sheshadri 2020	2	1.6	26	2.3	1.6	27	-0.18 [-0.72 , 0.36	5] - 	
								-4 -2 0 2 4	
								More with exercise More with control	



Analysis 2.3. Comparison 2: Aerobic exercise versus control (no exercise/ placebo exercise), Outcome 3: HRQoL: Summary component scores

		Control			Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
2.3.1 Physical Compor	nent Score									
Giannaki 2013a	70.5	26.5	7	76.4	15.6	15	3.9%	-5.90 [-27.06 , 15.26]		
ACTINUT 2013	59.87	21.37	9	84.7	13.32	7	5.5%	-24.83 [-41.93 , -7.73]		
Koh 2009 (1)	55	25	15	51.5	22.71	30	6.5%	3.50 [-11.54 , 18.54]		
CHAIR 2015 (2)	33.58	18.05	11	51.6	7.41	6	8.4%	-18.02 [-30.22 , -5.82]		
DIALY-SIZE 2016	3.4	7.3	8	5.2	9.3	8	12.3%	-1.80 [-9.99, 6.39]		
Samara 2016	43.9	8.8	12	49.9	6.6	15	14.9%	-6.00 [-12.00 , -0.00]		
Dobsak 2012	50.6	6.8	10	51.7	4.4	11	16.1%	-1.10 [-6.05, 3.85]		
Suhardjono 2019 (2)	-3.45	12.45	38	8.04	8.84	37	16.2%	-11.49 [-16.37 , -6.61]		
IHOPE 2019	38.9	9.42	38	38.72	10.4	29	16.2%	0.18 [-4.65, 5.01]	+	
Subtotal (95% CI)			148			158	100.0%	-6.00 [-10.71 , -1.30]	•	
Heterogeneity: Tau² = 2 Test for overall effect: 2	*	-	0 (1 0.0	,02),1 00	,,,					
Test for overall effect: 2	Z = 2.50 (P =	0.01)								
2.3.2 Mental Compone	ent Score									
Giannaki 2013a	65	21.9	7	70.4	18.7	15	4.0%	-5.40 [-24.18 , 13.38]		
Koh 2009 (1)	64	25	15	61.5	20.96	30	5.8%	2.50 [-12.21 , 17.21]		
ACTINUT 2013	52.07	16.11	9	74.3	10.61	7	6.8%	-22.23 [-35.37 , -9.09]		
DIALY-SIZE 2016	0.7	7.5	8	-2.3	10.7	8	10.3%	3.00 [-6.05 , 12.05]	- - -	
CHAIR 2015 (2)	53.05	6.65	11	51.6	7.41	6	12.5%	1.45 [-5.66 , 8.56]	-	
Samara 2016	39	10.4	12	53.3	6.9	15	12.9%	-14.30 [-21.14 , -7.46]	 -	
IHOPE 2019	50.7	11.63	38	52.05	10.81	29	14.7%	-1.35 [-6.75 , 4.05]	-	
Dobsak 2012	59.3	5.6	10	59.5	5.5	11	15.6%	-0.20 [-4.96 , 4.56]	+	
Suhardjono 2019 (2)	1.67	4.46	38	2.673	9.05	37	17.4%	-1.00 [-4.25 , 2.24]	+	
Subtotal (95% CI)			148			158	100.0%	-3.33 [-7.56 , 0.90]	•	
Heterogeneity: Tau ² = 2	23.94; Chi ² = 2	25.03, df =	= 8 (P = 0.0	002); I ² = 68	3%				•	
Test for overall effect: 2	Z = 1.54 (P =	0.12)								
Fastmatas									50 -25 0 25 5	
Footnotes								High	er with exercise Higher with co	

 $^{(1)\} two\ intervention\ arms\ polled\ together\ in\ the\ exercise\ group$

⁽²⁾ mean and standard deviation estimated from the median and range



Analysis 2.4. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 4: HRQoL: Individual domains

Study or Subgroup	Mean	Control SD	Total	Mean	Exercise SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
2.4.1 Physical Functioning	<u> </u>								
Parsons 2004	65.7	27.1	7	68.3	30.6	6	3.9%	-2.60 [-34.26, 29.06]	
Matsumoto 2007	48	36.0572	32	43	27.2293	17	7.8%	5.00 [-12.99 , 22.99]	
Zhao 2017	64.2	55.464	56	68.8	30.7879	59	8.4%	-4.60 [-21.11 , 11.91]	<u> </u>
Koh 2009 (1)	70	26	15	67.5	25.04	30	8.7%	2.50 [-13.42 , 18.42]	T
AVANTE-HEMO 2020	84.44	22.7	13	74.38	14.7	12	9.2%		<u> </u>
								10.06 [-4.82 , 24.94]	† -
Sheshadri 2020	63.7	24.3	26	60.2		27	9.9%	3.50 [-9.88 , 16.88]	
Jong 2004	2.35	10.62	17	7.42	17.17	19	12.1%	-5.07 [-14.29 , 4.15]	-• †
Dobsak 2012	53.1	10	10	54.1	7.9	11	12.8%	-1.00 [-8.76 , 6.76]	+
EXCITE 2014	-2.7	27.4518	123	1.5		104	13.5%	-4.20 [-10.52 , 2.12]	-
Wu 2014d	60.6	12.9	33	82.1	10	32	13.8%	-21.50 [-27.10 , -15.90]	+
Subtotal (95% CI)			332			317	100.0%	-2.87 [-10.12 , 4.38]	•
Heterogeneity: Tau ² = 91.35			9 (P < 0.00	001); I ² =	77%				
Test for overall effect: $Z = 0$)./8 (P = 1	0.44)							
2.4.2 Role-physical									
Parsons 2004	90.5	25.2	7	77.7	34.5	6	9.2%	12.80 [-20.52 , 46.12]	- - -
AVANTE-HEMO 2020	88.89	33.3	13	71.88	45.2	12	9.7%	17.01 [-14.32 , 48.34]	 • • • • • • • • • •
Koh 2009 (1)	48	44	15	37	39.82	30	11.0%	11.00 [-15.44, 37.44]	
Matsumoto 2007	40	58.2462	32	44	35.0091	17	11.1%	-4.00 [-30.16 , 22.16]	
Zhao 2017	54.4	65.3044	56	63.2		59	11.9%	-8.80 [-32.26 , 14.66]	
EXCITE 2014	-9.2	53.2229	123	0.2	47.821	104	14.9%	-9.40 [-22.55 , 3.75]	
Dobsak 2012	51.3	8.9	10	44.1	10.6	11	16.0%	7.20 [-1.15 , 15.55]	
Wu 2014d	26.3	11.5	33	54.6	15.4	32	16.3%		
	20.3	11.5		54.6	15.4			-28.30 [-34.92 , -21.68]	→
Subtotal (95% CI)	.c. cl ::	ED 45 30	289	0001	070/	271	100.0%	-2.31 [-17.29 , 12.67]	•
Heterogeneity: Tau² = 347.5 Test for overall effect: Z = 0			F / (P < 0.0	0001); 12 =	= 8/%				
	`	ĺ							
2.4.3 Pain		40.0	_	=0 =			2.00/	= 40 f 44 DD DD = 03	
Parsons 2004	86.6	13.2	7	79.5	23.9	6	3.2%	7.10 [-14.38 , 28.58]	 -
Zhao 2017	52.7	54.6641	56	64.6		59	3.4%	-11.90 [-32.75 , 8.95]	+
AVANTE-HEMO 2020	85	18.3	13	71.88	29.3	12	4.0%	13.12 [-6.21 , 32.45]	+
Koh 2009 (1)	57	31	15	62.5	29.98	30	4.1%	-5.50 [-24.51 , 13.51]	
Matsumoto 2007	47	36.0572	32	46	23.3394	17	5.3%	1.00 [-15.71 , 17.71]	—
Dobsak 2012	55.7	10.7	10	57.6	10.9	11	17.5%	-1.90 [-11.15 , 7.35]	+
EXCITE 2014	-3.2	34.1747	123	-1.1	30.8523	104	20.8%	-2.10 [-10.56, 6.36]	_
Wu 2014d	59	12.7	43	63	13.4	32	41.5%	-4.00 [-10.00 , 2.00]	_
Subtotal (95% CI)	33	121/	299	0.5	10	271		-2.26 [-6.12 , 1.61]	7
Heterogeneity: Tau ² = 0.00;	Chi ² = 4.	57, df = 7 ($I^2 = 0\%$		_,1	_50.070	, , ,,,,,,	T
Test for overall effect: $Z = 1$		•	. ,,						
2.4.4 General health perce	ntions								
2. 4.4 General nealth perce Parsons 2004	puons 50.1	22.4	7	50.7	22.7	6	3.6%	-0.60 [-25.20 , 24.00]	
Zhao 2017	52.2	50.9911	56	65.1	62.0354	59	4.9%	-12.90 [-33.61 , 7.81]	
Koh 2009 (1)	48	27	15	39	23.42	30	7.5%	9.00 [-7.03 , 25.03]	
							8.9%		 -
Matsumoto 2007	44 52.22	33.2836	32	43		17		1.00 [-13.22 , 15.22]	+
AVANTE-HEMO 2020	53.33	17.5	13	54.38	18.2	12	9.1%	-1.05 [-15.07 , 12.97]	+
Dobsak 2012	42.5	9	10	50.9		11	19.0%	-8.40 [-15.99 , -0.81]	-
<i>W</i> u 2014d	34.6	9.3	33	48.1	15.8	32	22.0%	-13.50 [-19.83 , -7.17]	+
EVCITE 2014	-2.5	20.1687	123	0.8	19.5398	104	25.0%	-3.30 [-8.48 , 1.88]	-
EXCITE 2014			289			271	100.0%	-5.38 [-10.32 , -0.43]	
			7 (P = 0.09); I ² = 43%	6				•
Subtotal (95% CI)	3; Chi² = 1	12.20, dt = 1							
Subtotal (95% CI) Heterogeneity: Tau ² = 18.53		,	•						l l
EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau ² = 18.53 Test for overall effect: Z = 2	2.13 (P = 0	,	·						
Subtotal (95% CI) Heterogeneity: Tau ² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being	2.13 (P = 0	0.03)		60.0	C1 = 7=0	F0	A 407	T (0 20 0C 40 0C)	
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017	2.13 (P = 0	61.2787	56	60.9		59	4.4%	-5.60 [-28.06 , 16.86]	_
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017 Parsons 2004	2.13 (P = 0 3 55.3 84.3	0.03) 61.2787 16.9	7	80.7	19.8	6	5.4%	3.60 [-16.59 , 23.79]	
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017	2.13 (P = 0	61.2787			19.8				-
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017 Parsons 2004 Matsumoto 2007	2.13 (P = 0 3 55.3 84.3	0.03) 61.2787 16.9	7	80.7	19.8	6	5.4%	3.60 [-16.59 , 23.79]	
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017 Parsons 2004	2.13 (P = 0 55.3 84.3 52	0.03) 61.2787 16.9 30.5099	7 32	80.7 54	19.8 23.3394	6 17	5.4% 8.7%	3.60 [-16.59 , 23.79] -2.00 [-17.32 , 13.32]	
Subtotal (95% CI) Heterogeneity: Tau² = 18.53 Test for overall effect: Z = 2 2.4.5 Emotional well-being Zhao 2017 Parsons 2004 Matsumoto 2007 AVANTE-HEMO 2020	2.13 (P = 0 55.3 84.3 52 75.11	0.03) 61.2787 16.9 30.5099 24.7	7 32 13	80.7 54 73	19.8 23.3394 12.4 9.2	6 17 12	5.4% 8.7% 8.8%	3.60 [-16.59 , 23.79] -2.00 [-17.32 , 13.32] 2.11 [-13.04 , 17.26]	



Analysis 2.4. (Continued)

W. 2014d	540								1
Wu 2014d	54.2	14.1	33	68.2	12.8	32	27.0%	-14.00 [-20.54 , -7.46]	
EXCITE 2014	-3.9	24.0904	123	1.2	19.5398	104	30.6%	-5.10 [-10.78, 0.58]	_
Subtotal (95% CI)			274			241	100.0%	-5.63 [-10.58, -0.67]	
Heterogeneity: Tau ² = 12.4	7: Chi ² = 8	.62. df = 6 (P	P = 0.20);	$I^2 = 30\%$				• / •	Y
Test for overall effect: Z = :									
2.4.6 Role-emotional	71.4	20.4	7	Ε0.	44.7	C	0.70/	21 40 [20 97 - 62 67]	
Parsons 2004	71.4	30.4	7	50	44.7	6	0.7%	21.40 [-20.87 , 63.67]	- -
Koh 2009 (1)	69	41	15	74	41.43	30	1.8%	-5.00 [-30.50 , 20.50]	
Zhao 2017	54.1	52.7802	56	61.8	70.3068	59	2.3%	-7.70 [-30.35 , 14.95]	+
Matsumoto 2007	47	52.699	32	50	27.2293	17	2.4%	-3.00 [-25.38 , 19.38]	
EXCITE 2014	-7.5	57.1446	123	-1.8	51.9278	104	5.9%	-5.70 [-19.90 , 8.50]	
AVANTE-HEMO 2020	92.59	22.2	13	100	0.001	12	8.1%	-7.41 [-19.48 , 4.66]	
Dobsak 2012	57	14.7	10	59.3	10.9	11	9.5%	-2.30 [-13.46, 8.86]	
Wu 2014d	30.4	7.4	33	40	9.4	32	69.5%	-9.60 [-13.72, -5.48]	-
Subtotal (95% CI)			289			271	100.0%	-8.02 [-11.45 , -4.58]	<u> </u>
Heterogeneity: Tau ² = 0.00;	; Chi ² = 3.8	30, df = 7 (P =	= 0.80); I ²	= 0%					•
Test for overall effect: Z =			,						
2.4.7 Vitality		4	_					1000510	
Parsons 2004	62.9	14.1	7	46.7	30.3	6	4.1%	16.20 [-10.20 , 42.60]	+-
Zhao 2017	50.4	58.4669	56	57.2	44.2067	59	6.7%	-6.80 [-25.82 , 12.22]	-+
Matsumoto 2007	47	30.5099	32	52	23.3394	17	8.7%	-5.00 [-20.32 , 10.32]	
AVANTE-HEMO 2020	72.78	22.6	13	49.38	14	12	9.2%	23.40 [8.78, 38.02]	-
Koh 2009 (1)	52	23	15	51	23.31	30	9.4%	1.00 [-13.32, 15.32]	
Sheshadri 2020	63.7	24.3	26	60.2	15.4	27	12.0%	3.50 [-7.50 , 14.50]	
Dobsak 2012	50.2	13.3	10	52.2	5.1	11	14.0%	-2.00 [-10.78 , 6.78]	
EXCITE 2014	-3.7	25.2109	123	0.8	13.3693	104	17.4%	-4.50 [-9.64 , 0.64]	
Wu 2014d	42.5	7.8	33	52.3	8.5	32	18.4%	-9.80 [-13.77 , -5.83]	
Subtotal (95% CI)	42.3	7.0	315	32.3	0.5	298	100.0%	-0.43 [-6.45, 5.60]	•
								-05 [-05 , 5.00]	•
	1. Chi2 = 2	6 08 df = 8 0). I2 – 600	4	250			Ť
Heterogeneity: Tau ² = 47.3); I ² = 699	6	250			Ĭ
); I ² = 699	%	250			
Heterogeneity: Tau ² = 47.3); I ² = 699	6	250			
Heterogeneity: $Tau^2 = 47.3$ Test for overall effect: $Z = 47.3$); I ² = 699 77.1	6 35.7	6	2.6%	3.20 [-29.08 , 35.48]	
Heterogeneity: Tau ² = 47.3 Test for overall effect: Z = 0 2.4.8 Social function	0.14 (P = 0	0.89)	P = 0.001					3.20 [-29.08 , 35.48] -12.70 [-34.81 , 9.41]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017	80.3 53.7	20.3 66.1989	P = 0.001	77.1	35.7 53.764	6	2.6% 5.1%	-12.70 [-34.81 , 9.41]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020	80.3 53.7 90.28	20.3 66.1989 19.5	7 56 13	77.1 66.4 85.94	35.7 53.764 26.2	6 59 12	2.6% 5.1% 7.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1)	80.3 53.7 90.28 73	20.3 66.1989 19.5 30	7 56 13 15	77.1 66.4 85.94 68.5	35.7 53.764 26.2 27.42	6 59 12 30	2.6% 5.1% 7.0% 7.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2)	80.3 53.7 90.28 73 41.68	20.3 66.1989 19.5 30 14.78	7 56 13 15	77.1 66.4 85.94 68.5 30.01	35.7 53.764 26.2 27.42 17.5	6 59 12 30 6	2.6% 5.1% 7.0% 7.0% 8.1%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012	80.3 53.7 90.28 73 41.68 66.5	20.3 66.1989 19.5 30 14.78 17.6	7 56 13 15 11	77.1 66.4 85.94 68.5 30.01 61.5	35.7 53.764 26.2 27.42 17.5	6 59 12 30 6	2.6% 5.1% 7.0% 7.0% 8.1% 10.6%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007	80.3 53.7 90.28 73 41.68 66.5	20.3 66.1989 19.5 30 14.78 17.6 13.8681	7 56 13 15 11 10 32	77.1 66.4 85.94 68.5 30.01 61.5 52	35.7 53.764 26.2 27.42 17.5 14 21.3944	6 59 12 30 6 11	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014	80.3 53.7 90.28 73 41.68 66.5 59	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699	7 56 13 15 11 10 32 123	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239	6 59 12 30 6 11 17	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d	80.3 53.7 90.28 73 41.68 66.5	20.3 66.1989 19.5 30 14.78 17.6 13.8681	7 56 13 15 11 10 32 123 33	77.1 66.4 85.94 68.5 30.01 61.5 52	35.7 53.764 26.2 27.42 17.5 14 21.3944	6 59 12 30 6 11 17 104 32	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI)	80.3 53.7 90.28 73 41.68 66.5 59 -0.8	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9	7 56 13 15 11 10 32 123 33 300	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) COHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) COHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 4	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300 (P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 20.9% 25.2% 100.0%	-12.70 [-34.81, 9.41] 4.34 [-13.88, 22.56] 4.50 [-13.58, 22.58] 11.67 [-4.83, 28.17] 5.00 [-8.69, 18.69] 7.00 [-4.25, 18.25] 1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] 0.94 [-4.48, 6.37]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300 (P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300 (P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 I I ² = 41%	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 EXCITE 2014	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (7 56 13 15 11 10 32 123 33 300 P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 I I ² = 41%	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7' Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI)	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73)	7 56 13 15 11 10 32 123 300 P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7' Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI)	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73)	7 56 13 15 11 10 32 123 300 P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7' Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.1	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868	7 56 13 15 11 10 32 123 300 P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7¹ Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.1² Test for overall effect: Z = 1	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0 76.87 43.5 -0.8	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868	7 56 13 15 11 10 32 123 300 P = 0.09);	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.T Test for overall effect: Z = 1 2.4.10 Effects of kidney di	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi² = 1.15 (P = 0 isease	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2	7 56 13 15 11 10 32 123 33 300 P = 0.09); 13 33 123 169 (P < 0.00	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81, 9.41] 4.34 [-13.88, 22.56] 4.50 [-13.58, 22.58] 11.67 [-4.83, 28.17] 5.00 [-8.69, 18.69] 7.00 [-4.25, 18.25] 1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] 0.94 [-4.48, 6.37] -3.88 [-13.72, 5.96] -18.70 [-24.29, -13.11] -0.20 [-3.85, 3.45] -7.65 [-20.65, 5.35]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7' Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d SUCTE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120' Test for overall effect: Z = 1 2.4.10 Effects of kidney di AVANTE-HEMO 2020	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi² = 1.15 (P = 0 64.59 64.59	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 169 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 12 = 41% 80.75 62.2 -0.6 001); I ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0%	-12.70 [-34.81, 9.41] 4.34 [-13.88, 22.56] 4.50 [-13.58, 22.58] 11.67 [-4.83, 28.17] 5.00 [-8.69, 18.69] 7.00 [-4.25, 18.25] 1.70 [-5.32, 8.72] -7.50 [-12.56, -2.44] 0.94 [-4.48, 6.37] -3.88 [-13.72, 5.96] -18.70 [-24.29, -13.11] -0.20 [-3.85, 3.45] -7.65 [-20.65, 5.35]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7' Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 120.1' Test for overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi ² = 1.15 (P = 0 64.59 -0.3	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 123 123 123 123 123 123	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); I ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 30.2% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 1 2.4.8 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 1 2.4.9 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.7 Test for overall effect: Z = 1 2.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d EXCITE 2014 Subtotal effect: Z = 2 2.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi² = 1.15 (P = 0 64.59 64.59	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 169 (P < 0.000 13 123 33 33 30 (P = 0.09); 15 15 169 (P < 0.000 13 123 33 33 33 33 33 33 33 33 33 33 33 33 3	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 12 = 41% 80.75 62.2 -0.6 001); I ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74] -4.80 [-7.83 , -1.77]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.2 Test for overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal effect: Z = 22.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI)	0.14 (P = 0 80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi ² = 1 1.15 (P = 0 64.59 -0.3 34.6	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2 0.25)	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 139 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); 1 ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 30.2% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.7 Test for overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 120.7 Test for overall effect: Z = 2.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00, Test for Overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00,	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1.0.34 (P = 0 64.59 -0.3 34.6 64.59 -0.3 34.6 ; Chi ² = 1.2 (Chi ² = 1.2	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2 0.25) 18.1 20.7289 5.7	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 139 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); 1 ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74] -4.80 [-7.83 , -1.77]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.7 Test for overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 120.7 Test for overall effect: Z = 2.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00, Test for Overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 0.00,	80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi ² = 1.0.34 (P = 0 64.59 -0.3 34.6 64.59 -0.3 34.6 ; Chi ² = 1.2 (Chi ² = 1.2	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2 0.25) 18.1 20.7289 5.7	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 139 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); 1 ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74] -4.80 [-7.83 , -1.77]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 47.3	0.14 (P = 0 80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi² = 1.15 (P = 0 64.59 -0.3 34.6 ; Chi² = 1.2	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2 0.25) 18.1 20.7289 5.7	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 139 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); 1 ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74] -4.80 [-7.83 , -1.77]	
Heterogeneity: Tau² = 47.3 Test for overall effect: Z = 42.48 Social function Parsons 2004 Zhao 2017 AVANTE-HEMO 2020 Koh 2009 (1) CHAIR 2015 (2) Dobsak 2012 Matsumoto 2007 EXCITE 2014 Wu 2014d Subtotal (95% CI) Heterogeneity: Tau² = 23.7 Test for overall effect: Z = 42.49 Symptoms AVANTE-HEMO 2020 Wu 2014d EXCITE 2014 Subtotal (95% CI) Heterogeneity: Tau² = 120.2 Test for overall effect: Z = 42.410 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal effect: Z = 22.4.10 Effects of kidney di AVANTE-HEMO 2020 EXCITE 2014 Wu 2014d Subtotal (95% CI)	0.14 (P = 0 80.3 53.7 90.28 73 41.68 66.5 59 -0.8 37 7; Chi² = 1 0.34 (P = 0 76.87 43.5 -0.8 36; Chi² = 1.15 (P = 0 64.59 -0.3 34.6 ; Chi² = 1.2	20.3 66.1989 19.5 30 14.78 17.6 13.8681 22.9699 8.9 3.58, df = 8 (0.73) 13.4 8.8 15.6868 29.65, df = 2 0.25) 18.1 20.7289 5.7	7 56 13 15 11 10 32 123 33 300 (P = 0.09); 13 123 123 123 139 (P < 0.00)	77.1 66.4 85.94 68.5 30.01 61.5 52 -2.5 44.5 1 ² = 41% 80.75 62.2 -0.6 001); 1 ² =	35.7 53.764 26.2 27.42 17.5 14 21.3944 29.8239 11.7 11.7 13.6 12.3409 93%	6 59 12 30 6 11 17 104 32 277	2.6% 5.1% 7.0% 8.1% 10.6% 13.5% 20.9% 25.2% 100.0% 34.2% 35.5% 100.0%	-12.70 [-34.81 , 9.41] 4.34 [-13.88 , 22.56] 4.50 [-13.58 , 22.58] 11.67 [-4.83 , 28.17] 5.00 [-8.69 , 18.69] 7.00 [-4.25 , 18.25] 1.70 [-5.32 , 8.72] -7.50 [-12.56 , -2.44] 0.94 [-4.48 , 6.37] -3.88 [-13.72 , 5.96] -18.70 [-24.29 , -13.11] -0.20 [-3.85 , 3.45] -7.65 [-20.65 , 5.35] -8.47 [-22.23 , 5.29] -1.80 [-7.34 , 3.74] -4.80 [-7.83 , -1.77]	

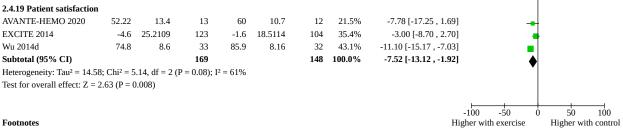


Analysis 2.4. (Continued)

2.4.11 Burden of kidney d		10.0	10	C 4 O C	17.0	10	2.20/	7.00 [22.00 . 0.01	
AVANTE-HEMO 2020	56.96	19.6	13	64.86	17.9	12	3.3%	-7.90 [-22.60 , 6.80]	-+
EXCITE 2014	0.4	28.0121	123	2.6	24.1676	104	15.5%	-2.20 [-8.99 , 4.59]	1
Wu 2014d	27.6	5.91	33 160	26.8	6.29	32	81.2%	0.80 [-2.17, 3.77]	•
Subtotal (95% CI)	. Ch:2 = 4.7	70 46. 272	169	- 00/		148	100.0%	0.05 [-2.63 , 2.72]	•
Heterogeneity: $Tau^2 = 0.00$ Test for overall effect: $Z =$			= 0.41); 12	= 0%					
2.4.12 Work status									
AVANTE-HEMO 2020	50	35.4	13	37.5	44.3	12	1.2%	12.50 [-19.10 , 44.10]	-
EXCITE 2014	-0.9	31.3735	123	0.3	22.1108	104	24.1%	-1.20 [-8.19 , 5.79]	+
Wu 2014d	29.3	8.97	33	29.7	7.29	32	74.7%	-0.40 [-4.37 , 3.57]	
Subtotal (95% CI)			169			148	100.0%	-0.44 [-3.87 , 2.99]	•
Heterogeneity: Tau² = 0.00 Test for overall effect: Z =			= 0.71); I ²	? = 0%					
2.4.13 Cognitive function									
AVANTE-HEMO 2020	24.46	27.3	13	27.51	33.3	12	2.5%	-3.05 [-27.03 , 20.93]	
EXCITE 2014	-6.4	30.8133	123	0.3	17.9972	104	33.9%	-6.70 [-13.15 , -0.25]	
Wu 2014d	70.4	9.19	33	76.7	10.13	32	63.7%	-6.30 [-11.01 , -1.59]	<u> </u>
Subtotal (95% CI)	, U. -1	5.15	169	, 5.7	10.10	148	100.0%	-6.36 [-10.11 , -2.60]	
Heterogeneity: Tau ² = 0.00	· Chi ² = 0.0	08 df = 2 (D		2 = 0%		170	100.0 /0	0.00 [10.11 , -2.00]	▼
Test for overall effect: Z =			0.30J, I*	- 0/0					
2.4.14 Quality of social in	teractions								
AVANTE-HEMO 2020	25.92	25.3	13	31.66	23.3	12	3.6%	-5.74 [-24.79 , 13.31]	
Wu 2014d	66.5	11.5	33	73.9	11.25	32	42.4%	-7.40 [-12.93 , -1.87]	-
EXCITE 2014	-4.6	20.7289	123	2.1	16.9688	104	54.0%	-6.70 [-11.60 , -1.80]	
Subtotal (95% CI)			169			148	100.0%	-6.96 [-10.57 , -3.36]	•
Heterogeneity: Tau ² = 0.00	; Chi ² = 0.0	05, df = 2 (P	= 0.97); I ²	? = 0%					*
Test for overall effect: Z =	3.79 (P = 0	0.0002)							
2.4.15 Sexual function AVANTE-HEMO 2020	8.33	25	13	29.69	42.2	12	5.1%	-21.36 [-48.83 , 6.11]	
EXCITE 2014	-2.1	46.5	123	-4.9	39.5938	104	25.2%	2.80 [-8.40 , 14.00]	
Wu 2014d	15	10.37	33	15.7	9.39	32	69.7%	-0.70 [-5.51 , 4.11]	_
Subtotal (95% CI)			169			148	100.0%	-0.87 [-7.23 , 5.48]	
Heterogeneity: Tau ² = 9.07	: Chi ² = 2.5	55. df = 2 (P	= 0.28); I ²	= 21%				, ,	Y
Test for overall effect: Z =			,						
2.4.16 Sleep									
AVANTE-HEMO 2020	69.72	6.9	13	72.5	8.4	12	31.1%	-2.78 [-8.83 , 3.27]	+
Wu 2014d	36.4	7.54	33	49.7	11.6	32	34.1%	-13.30 [-18.07 , -8.53]	-
EXCITE 2014	0.7	19.0482	123	3.7	14.9119	104	34.8%	-3.00 [-7.42 , 1.42]	-
Subtotal (95% CI)			169			148	100.0%	-6.44 [-13.46 , 0.58]	•
Heterogeneity: $Tau^2 = 31.7$ Test for overall effect: $Z =$			P = 0.003); I ² = 839	%				
2.4.17 Social support									
AVANTE-HEMO 2020	68.53	10	13	77.08	21.7	12	9.0%	-8.55 [-21.98 , 4.88]	_
EXCITE 2014	-2	24.0904	123	-1.5	22.1108	104	35.1%	-0.50 [-6.52 , 5.52]	4
Wu 2014d	75.27	7.86	33	81.36	9.41	32	55.9%	-6.09 [-10.31 , -1.87]	_
Subtotal (95% CI)			169			148	100.0%	-4.35 [-8.51 , -0.19]	
Heterogeneity: Tau ² = 3.42	; Chi ² = 2.5	59, df = 2 (P	= 0.27); I ²	2 = 23%					•
Test for overall effect: Z =	2.05 (P = 0	0.04)							
2.4.18 Dialysis staff encou	_								
AVANTE-HEMO 2020	80.56	12.6	13	82.81	16.3	12	16.1%	-2.25 [-13.74 , 9.24]	+
EXCITE 2014	-1.6	17.9277	123	1.1	4.1136	104	41.9%	-2.70 [-5.97 , 0.57]	•
Wu 2014d	81.1	7.7	33	90.6	5.4	32	42.0%	-9.50 [-12.73 , -6.27]	-
Subtotal (95% CI)			169			148	100.0%	-5.49 [-11.11 , 0.13]	lack
Heterogeneity: Tau ² = 16.8			P = 0.01);	$I^2 = 77\%$					·
Test for overall effect: Z =	1.91 (P = 0	0.06)							
2.4.19 Patient satisfaction	I 52.22	13./	12	60	10.7	12	21 5%	-7 78 f-17 25 1 691	
ауамти-нимит лілі	5, ,,	1 3 7	1.4	ы	111.7	1)	11 50%	_/ /X I_I / /S I KUI	-1



Analysis 2.4. (Continued)



- (1) two intervention arms polled together in the exercise group
- (2) mean and standard deviation estimated from the median and range

Analysis 2.5. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 5: Depression

		Control		1	Exercise			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Giannaki 2013a	43.71	11.17	7	35.84	6.38	15	21.7%	0.93 [-0.01 , 1.88	
Carmack 1995	5	5	11	6.8	8.2	10	23.4%	-0.26 [-1.12, 0.60]
Kouidi 1997	21.3	11.9	11	13.7	9.5	20	25.3%	0.71 [-0.05 , 1.47	1
Sheshadri 2020	6.6	6.5	26	11.3	12.4	27	29.6%	-0.47 [-1.01 , 0.08	···
Total (95% CI)			55			72	100.0%	0.19 [-0.52 , 0.89	
Heterogeneity: Tau ² = 0	0.35; Chi ² = 10	0.16, df =	3 (P = 0.02)	2); I ² = 70%					
Test for overall effect: 2	Z = 0.52 (P =	0.60)							$\begin{array}{c ccccccccccccccccccccccccccccccccccc$
Test for subgroup differ	ences: Not ap	plicable						H	Higher with exercise Higher with control

Analysis 2.6. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 6: 6MWT

	(Control		E	xercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [metres]	SD [metres]	Total	Mean [metres]	SD [metres]	Total	Weight	IV, Random, 95% CI [metres]	IV, Random, 95% CI [metres]
ACTINUT 2013	295.77	121.07	9	346.28	134.88	7	2.1%	-50.51 [-177.95 , 76.93]	
Koh 2009 (1)	452	144	16	509.5	121.07	28	4.5%	-57.50 [-141.10 , 26.10]	
Samara 2016	454.4	90.4	12	625.6	128.1	15	4.5%	-171.20 [-253.77 , -88.63]	<u> </u>
DIALY-SIZE 2016	0.8	44	8	42.3	88.8	8	6.1%	-41.50 [-110.17, 27.17]	
CHAIR 2015 (2)	317.5	81.6	11	307.5	54.62	6	6.7%	10.00 [-55.08, 75.08]	
Wu 2014d	359	132	33	441	135	32	6.7%	-82.00 [-146.93 , -17.07]	
Liao 2016	290	64.1006	20	350	128.2012	20	7.0%	-60.00 [-122.82 , 2.82]	-
Fernandes 2019	325	59.8	19	386.9	19.38	20	17.8%	-61.90 [-90.10 , -33.70]	-
Cho 2018	-26	41	13	25	29	11	17.9%	-51.00 [-79.11 , -22.89]	-
EXCITE 2014	2	44.8193	123	39	35.9943	104	26.7%	-37.00 [-47.52 , -26.48]	•
Total (95% CI)			264			251	100.0%	-53.00 [-72.17 , -33.84]	•
Heterogeneity: Tau ² = 3	29.93; Chi ² = 16.89,	df = 9 (P = 0.05)	$I^2 = 47\%$						*
Test for overall effect: 2	Z = 5.42 (P < 0.00001)							-500 -250 0 250 500
Test for subgroup differ	ences: Not applicable	2						Furt	ther with exercise Further with control

- (1) two intervention arms polled together in the exercise group
- (2) mean and standard deviation estimated from the median and range



Analysis 2.7. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N reps/30 sec]

		Control		1	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% CI
Giannaki 2013a	17.25	3.2	7	17.84	4.68	15	8.1%	-0.59 [-3.94 , 2.70	51 -
DIALY-SIZE 2016	1.4	4.3	8	1	2.2	8	8.1%	. ,	-
Koufaki 2002	12.05	3.6	15	13.45	3.1	18	16.8%	-1.40 [-3.72 , 0.93	2]
IHOPE 2019	10.7	5.6	38	11.7	4	29	17.1%	-1.00 [-3.30 , 1.30	0]
Wu 2014d	12.8	3.65	33	15.55	4.8	32	21.0%	-2.75 [-4.83 , -0.6	7]
Cho 2018	-0.5	2.2	13	2.3	2.2	11	29.0%	-2.80 [-4.57 , -1.03	3]
Total (95% CI)			114			113	100.0%	-1.81 [-2.76 , -0.8	6]
Heterogeneity: Tau ² = 0. Test for overall effect: Z		,	(P = 0.44)	$I^2 = 0\%$					-10 -5 0 5 10
Test for subgroup differen	ences: Not ap	plicable							More with exercise More with control

Analysis 2.8. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 8: Sit-To-Stand test [sit to 5 reps]

		Control]	Exercise			Mean Difference	Mean	Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Rand	lom, 95% CI	
Koufaki 2002	12.7	4.8	15	11	3.3	18	5.9%	1.70 [-1.17 , 4.57]		
Samara 2016	10.6	3.25	12	7.6	2.7	15	9.1%	3.00 [0.71, 5.29]		
Wu 2014d	12.6	3.6	33	10.75	3.4	32	16.0%	1.85 [0.15 , 3.55]		
Giannaki 2013a	8.81	0.66	7	8.24	2.34	15	26.7%	0.57 [-0.71 , 1.85]	-	
EXCITE 2014	-0.6	5.0422	123	-2.5	2.0568	104	42.4%	1.90 [0.93 , 2.87]	-	
Total (95% CI)			190			184	100.0%	1.63 [0.92 , 2.33]	•	
Heterogeneity: Tau ² = 0	0.06; Chi ² = 4.	36, df = 4	(P = 0.36)	; I ² = 8%						*	
Test for overall effect: Z	Z = 4.51 (P <	0.00001)							-10 -5	0 5	10
Test for subgroup differ	ences: Not ap	plicable						L	onger with exercise	Longer wi	ith control



Analysis 2.9. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 9: Resting blood pressure

	(Control		E	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mm Hg]	SD [mm Hg]	Total	Mean [mm Hg]	SD [mm Hg]	Total	Weight	IV, Random, 95% CI [mm Hg]	IV, Random, 95% CI [mm Hg]
2.9.1 Systolic blood p	ressure								
Liao 2016	138.8	16.7	20	96	64.2	20	3.2%	42.80 [13.73, 71.87]	
CYCLE-HD 2016	148.5	28.23	13	142.27	25.99	9	4.7%	6.23 [-16.66, 29.12]	
McGregor 2018	123.17	29.5201	18	135.75	25.2786	16	6.3%	-12.58 [-31.00, 5.84]	
Goldberg 1983	149	17	11	142	27	14	6.8%	7.00 [-10.35 , 24.35]	
Coh 2009 (1)	136	29	16	140.5	25.34	30	7.0%	-4.50 [-21.36, 12.36]	
Oussaint 2008	147.8	23.5	10	141.4	11.9	9	7.2%	6.40 [-10.11, 22.91]	
Wilund 2010	147.1	14.9	9	153	17.2	8	7.8%	-5.90 [-21.29, 9.49]	
suyuki 2003	130.8	23.3	12	141.5	16.4	17	7.9%	-10.70 [-26.02 , 4.62]	
Cooke 2018 (2)	0.4	9.89	10	-9.167	21.93	10	8.1%	9.57 [-5.34, 24.48]	-
aluchamy 2018	148	14.7678	10	137	16.349	10	8.9%	11.00 [-2.65, 24.65]	-
HOPE 2019	148.9	23.3	38	132.4	27.9	29	9.6%	16.50 [3.93, 29.07]	
Deligiannis 1999a	144	10	12	143	17	10	10.1%	1.00 [-10.96, 12.96]	
ernandes 2019	143.16	16.68	19	140.5	11.9	20	12.3%	2.66 [-6.48 , 11.80]	
ubtotal (95% CI)			198			202	100.0%	3.96 [-1.78, 9.70]	L
leterogeneity: Tau ² =	47.91; Chi ² = 21.95, di	f = 12 (P = 0.04):	I ² = 45%						Y
est for overall effect.	()								
.9.2 Diastolic blood p	pressure								
.9.2 Diastolic blood p		27.5118	10	83	24.3495	10	1.6%	0.00 [-22.77 , 22.77]	
.9.2 Diastolic blood p	pressure 83 77.1	13.7	20	53.7	35.3	10 20	2.8%	0.00 [-22.77 , 22.77] 23.40 [6.81 , 39.99]	
. 9.2 Diastolic blood p aluchamy 2018 .iao 2016	pressure 83	13.7			35.3				
.9.2 Diastolic blood paraluchamy 2018 iao 2016 Cooke 2018 (3)	pressure 83 77.1	13.7 10.32	20	53.7	35.3 16.77	20	2.8%	23.40 [6.81, 39.99]	
9.2 Diastolic blood p Paluchamy 2018 .iao 2016 Cooke 2018 (3) Goldberg 1983	Pressure 83 77.1 3.5	13.7 10.32 12	20 10	53.7 -2.6	35.3 16.77 18	20 10	2.8% 4.8%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30]	
.9.2 Diastolic blood paluchamy 2018 iao 2016 Cooke 2018 (3) Goldberg 1983 AcGregor 2018	93 77.1 3.5 86	13.7 10.32 12 16.4894	20 10 11	53.7 -2.6 82	35.3 16.77 18 15.6513	20 10 14	2.8% 4.8% 5.1%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80]	
.9.2 Diastolic blood paluchamy 2018 iao 2016 Cooke 2018 (3) Goldberg 1983 AcGregor 2018 Suyuki 2003	983 77.1 3.5 86 70.5	13.7 10.32 12 16.4894 13.5	20 10 11 18	53.7 -2.6 82 72.13	35.3 16.77 18 15.6513 12.3	20 10 14 16	2.8% 4.8% 5.1% 5.8%	23.40 [6.81, 39.99] 6.10 [-6.10, 18.30] 4.00 [-7.80, 15.80] -1.63 [-12.44, 9.18]	
.9.2 Diastolic blood p aluchamy 2018 iao 2016 cooke 2018 (3) ioldberg 1983 dcGregor 2018 suyuki 2003 coh 2009 (4)	93 77.1 3.5 86 70.5 79	13.7 10.32 12 16.4894 13.5	20 10 11 18 12	53.7 -2.6 82 72.13 85.8	35.3 16.77 18 15.6513 12.3 13.15	20 10 14 16 17	2.8% 4.8% 5.1% 5.8% 7.0%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82]	
.9.2 Diastolic blood p laluchamy 2018 i.ao 2016 Gooke 2018 (3) Goldberg 1983 4cGregor 2018 Suyuki 2003 fon 2009 (4) EYCLE-HD 2016	83 77.1 3.5 86 70.5 79	13.7 10.32 12 16.4894 13.5 15	20 10 11 18 12 16	53.7 -2.6 82 72.13 85.8 78	35.3 16.77 18 15.6513 12.3 13.15 8.72	20 10 14 16 17 30	2.8% 4.8% 5.1% 5.8% 7.0% 8.0%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73]	
.9.2 Diastolic blood p aluchamy 2018 iao 2016 Cooke 2018 (3) Goldberg 1983 AcGregor 2018 Suyuki 2003 Coh 2009 (4) YYCLE-HD 2016 Vilund 2010	83 77.1 3.5 86 70.5 79 75 79.06	13.7 10.32 12 16.4894 13.5 15 11.98 8.7	20 10 11 18 12 16	53.7 -2.6 82 72.13 85.8 78 77.91	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7	20 10 14 16 17 30 9	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73] 1.15 [-7.50 , 9.80] -8.40 [-16.20 , -0.60] 1.90 [-5.33 , 9.13]	
.9.2 Diastolic blood p aluchamy 2018 .iao 2016 .cooke 2018 (3) .coldberg 1983 .c.Gregor 2018 .suyuki 2003 .coug (4) .cyCLE-HD 2016 .viyULD 2010 .cyCLE-UD 2010 .cyCLE-UD 2010	83 77.1 3.5 86 70.5 79 75 79.06 77.3	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5	20 10 11 18 12 16 13	53.7 -2.6 82 72.13 85.8 78 77.91 85.7	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6	20 10 14 16 17 30 9	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73] 1.15 [-7.50 , 9.80] -8.40 [-16.20 , -0.60]	
.9.2 Diastolic blood p aluchamy 2018 iao 2016 cooke 2018 (3) ioldberg 1983 fcGregor 2018 suyuki 2003 coh 2009 (4) YYCLE-HD 2016 VIYCLE-HD 2019 ioussaint 2008	83 77.1 3.5 86 70.5 79 75 79.06 77.3 78.5	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5 9.4	20 10 11 18 12 16 13 9 38	53.7 -2.6 82 72.13 85.8 78 77.91 85.7	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6 5.7	20 10 14 16 17 30 9 8 29	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3% 10.3%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73] 1.15 [-7.50 , 9.80] -8.40 [-16.20 , -0.60] 1.90 [-5.33 , 9.13]	
.9.2 Diastolic blood p laluchamy 2018 liao 2016 looke 2018 (3) loidberg 1983 lcGregor 2018 lsuyuki 2003 loh 2009 (4) lYCLE-HD 2016 vilund 2010 HOPE 2019 loussaint 2008 lemandes 2019	93 77.1 3.5 86 70.5 79 75 79.06 77.3 78.5 72.8	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5 9.4	20 10 11 18 12 16 13 9 38	53.7 -2.6 82 72.13 85.8 78 77.91 85.7 76.6	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6 5.7	20 10 14 16 17 30 9 8 29	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3% 10.3%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 18.30] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73] 1.15 [-7.50 , 9.80] -8.40 [-16.20 , -0.60] 1.90 [-5.33 , 9.13] -4.40 [-11.31 , 2.51]	
.9.2 Diastolic blood p taluchamy 2018 .iao 2016 .cooke 2018 (3) .coldberg 1983 .dcGregor 2018 .suyuki 2003 .coh 2009 (4) .YCLE-HD 2016 .Vilund 2010 .HOPE 2019 .coussaint 2008 .ernandes 2019 .deligiannis 1999a	93 83 77.1 3.5 86 70.5 79.06 77.3 78.5 72.8 86.32	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5 9.4	20 10 11 18 12 16 13 9 38 10	53.7 -2.6 82 72.13 85.3 78 77.91 85.7 76.6 77.2	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6 5.7	20 10 14 16 17 30 9 8 29 9	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3% 10.3% 12.0% 14.5%	23.40 [6.81 , 39.99] 6.10 [-6.10 , 18.30] 4.00 [-7.80 , 15.80] -1.63 [-12.44 , 9.18] -6.80 [-16.42 , 2.82] -3.00 [-11.73 , 5.73] 1.15 [-7.50 , 9.80] -8.40 [-16.20 , -0.60] 1.90 [-5.33 , 9.13] -4.40 [-11.31 , 2.51] 0.32 [-6.01 , 6.65]	
n.9.2 Diastolic blood paraluchamy 2018 diauchamy 2018 diao 2016 Cooke 2018 (3) Goldberg 1983 AcGregor 2018 Suyuki 2003 Coh 2009 (4) CYCLE-HD 2016 Vilund 2010 HOPE 2019 Coussaint 2008 Pernandes 2019 Poletigiannis 1999a Subtotal (95% CI)	93 83 77.1 3.5 86 70.5 79.06 77.3 78.5 72.8 86.32	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5 9.4 12.52	20 10 11 18 12 16 13 9 38 10 19	53.7 -2.6 82 72.13 85.3 78 77.91 85.7 76.6 77.2	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6 5.7	20 10 14 16 17 30 9 8 29 9	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3% 10.3% 12.0% 14.5%	23.40 [6.81, 39.99] 6.10 [-6.10, 18.30] 4.00 [-7.80, 15.80] -1.63 [-12.44, 9.18] -6.80 [-16.42, 2.82] -3.00 [-11.73, 5.73] 1.15 [-7.50, 9.80] -8.40 [-16.20, -0.60] 1.90 [-5.33, 9.13] -4.40 [-11.31, 2.51] 0.32 [-6.01, 6.65] -1.00 [-6.24, 4.24]	
Test for overall effect: 1.9.2 Diastolic blood p Paluchamy 2018 1.10 2016 1.10 2016 1.10 2018 1.10 2018 1.10 2018 1.10 2018 1.10 2018 1.10 2019 1	77.1 3.5 86 70.5 79 75 79.06 77.3 78.5 72.8 86.32 82 8.50; Chi ² = 17.33, df	13.7 10.32 12 16.4894 13.5 15 11.98 8.7 12.5 9.4 12.52	20 10 11 18 12 16 13 9 38 10 19	53.7 -2.6 82 72.13 85.3 78 77.91 85.7 76.6 77.2	35.3 16.77 18 15.6513 12.3 13.15 8.72 7.7 16.6 5.7	20 10 14 16 17 30 9 8 29 9	2.8% 4.8% 5.1% 5.8% 7.0% 8.0% 8.1% 9.3% 10.3% 12.0% 14.5%	23.40 [6.81, 39.99] 6.10 [-6.10, 18.30] 4.00 [-7.80, 15.80] -1.63 [-12.44, 9.18] -6.80 [-16.42, 2.82] -3.00 [-11.73, 5.73] 1.15 [-7.50, 9.80] -8.40 [-16.20, -0.60] 1.90 [-5.33, 9.13] -4.40 [-11.31, 2.51] 0.32 [-6.01, 6.65] -1.00 [-6.24, 4.24]	

- (1) two intervention arms polled together in the exercise group
- (2) mean and standard deviation estimated from the median and interquartile range
- (3) Mean and standard deviation estimated from the median and interquartile range
- (4) Two intervention arms polled together in the exercise group $% \left\{ 1,2,...,n\right\}$

Analysis 2.10. Comparison 2: Aerobic exercise versus control (no exercise/ placebo exercise), Outcome 10: Aerobic capacity (VO2 max or peak)

Study or Subgroup	Mean [mL/kg/min]	Control SD [mL/kg/min] Total		Exercise I Mean [mL/kg/min] SD [mL/kg/min]			Weight	Mean Difference IV, Random, 95% CI [mL/kg/min]	Mean Difference IV, Random, 95% CI [mL/kg/min]
Parsons 2004	55	26	7	58	44	6	0.2%	-3.00 [-43.13 , 37.13]	
Goldberg 1983	20	8	11	25	9	14			
Painter 2002a (1)	19.83		25	21.11		23			- - <u>-</u> -
Deligiannis 1999a	15.8	4.8	12	9	5.3	10			T_
Kouidi 1997	15.9	4.3	11	23.27	7.6	20			
McGregor 2018	15.93		18	20.71	6.831	16			
Tsuyuki 2003	21.7	4.9	12	27	5.6	17			
Koufaki 2002	18.8	4.9	15	19.9	6.3	18	9.1%	-1.10 [-4.92 , 2.72]	_
Carmack 1995	10.9	3.1	11	14.4	4.7	10	9.9%	-3.50 [-6.94, -0.06]	-
Jong 2004	22.86	5.46	17	25.23	4.33	19	10.2%	-2.37 [-5.61, 0.87]	
Akiba 1995	17.6	2.6	6	20	2.4	7	11.3%	-2.40 [-5.14, 0.34]	_
Kouidi 2004a	-0.4	2.3	10	3.1	3.3	11	11.9%	-3.50 [-5.92 , -1.08]	•
Total (95% CI)			155			171	100.0%	-2.69 [-4.55 , -0.82]	•
Heterogeneity: Tau ² = 6 Test for overall effect: Z Test for subgroup differ		$(P = 0.002); I^2 = 62\%$	Ď					Hiį	-50 -25 0 25 50 gher with exercise Higher with con

 $(1) factorial \ design: two \ intervention \ arms \ pooled \ together \ in \ the \ exercise \ group \ and \ two \ control \ arms \ pooled \ together \ in \ the \ control \ group$



Analysis 2.11. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 11: Albumin

	Favo	urs exercise		E	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
Frey 1999	37	7	6	44	5	5	2.4%	-7.00 [-14.11 , 0.1	1]
CHAIR 2015 (1)	39.25	2.2	11	39	6.2	6	3.8%	0.25 [-4.88 , 5.38	B]
Koufaki 2002	40.5	3.6	15	35.2	7.3	18	5.5%	5.30 [1.47, 9.13	3]
AVANTE-HEMO 2020	39	4	13	34	5.5	12	5.5%	5.00 [1.20 , 8.86	0]
Afshar 2010	40	3	7	40	4	7	5.7%	0.00 [-3.70 , 3.70	0]
ong 2004	33	6.1	17	38.2	5.1	19	5.7%	-5.20 [-8.90 , -1.50	0]
Matsumoto 2007	36.9	8.0435	32	38.6	4.4734	17	6.0%	-1.70 [-5.21 , 1.8	1]
teboredo 2010	41	. 5	11	39	3	11	6.1%	2.00 [-1.45 , 5.4	5]
ACTINUT 2013	39.12	3.67	9	39.33	2.51	7	6.8%	-0.21 [-3.24 , 2.83	2]
Copple 2007	39	3.7417	14	38	3.2	10	7.3%	1.00 [-1.79 , 3.79	9]
iao 2016	40.1	4.2	20	41.6	3	20	8.4%	-1.50 [-3.76 , 0.76	6]
Fernandes 2019	34.6	4.1	19	37.9	2.5	20	8.6%	-3.30 [-5.44 , -1.10	6]
HOPE 2019	40.1	3.1	38	39.3	5.1	29	8.7%	0.80 [-1.30 , 2.96	0]
Coussaint 2008	34.1	. 2	10	33.8	2.5	9	8.8%	0.30 [-1.75 , 2.3	5]
Wilund 2010	38	0.6	9	39	1.5	8	10.7%	-1.00 [-2.11 , 0.1	1] -
Total (95% CI)			231			198	100.0%	-0.23 [-1.45 , 0.99	9]
Heterogeneity: Tau ² = 3.3	32; Chi ² = 40.18	B, df = 14 (P	= 0.0002);	$I^2 = 65\%$					Ĭ
Test for overall effect: Z	= 0.37 (P = 0.7	1)							-20 -10 0 10 2
Test for subgroup differe	nces: Not appli	cable						I	Higher with exercise Higher with co

(1) mean and standard deviation estimated from the median and range

Analysis 2.12. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 12: Blood lipids

	Control			I	exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L	
2.12.1 Total cholesterol	[mmol/L]									
Afshar 2010	3.4	0.812	7	3.377	0.88	7	10.1%	0.02 [-0.86, 0.91]		
Lee 2001	5.08	1.57	21	4.38	0.98	19	11.6%	0.70 [-0.10 , 1.50]	<u> </u>	
Wilund 2010	4.25	0.61	9	3.54	0.339	8	21.1%	0.71 [0.25 , 1.17]		
Liao 2016	4.71	0.58	20	4.456	0.776	20	22.6%	0.25 [-0.17, 0.68]		
Groussard 2015	1.63	0.09	10	1.61	0.13	8	34.7%	0.02 [-0.09 , 0.13]	•	
Subtotal (95% CI)			67			62	100.0%	0.30 [-0.03, 0.63]		
Heterogeneity: Tau ² = 0.	.08; Chi ² = 11.30, df	= 4 (P = 0.02); I ² =	65%							
Test for overall effect: Z	= 1.76 (P = 0.08)									
2.12.2 LDL cholesterol	[mmol/L]									
Lee 2001	2.91	0.97	21	2.51	0.91	19	14.1%	0.40 [-0.18, 0.98]		
Afshar 2010	1.56	0.33	7	1.245	0.25	7	31.5%	0.31 [0.01, 0.62]	L	
Groussard 2015	0.91	0.09	10	0.89	0.12	8	54.4%	0.02 [-0.08, 0.12]	<u>.</u>	
Subtotal (95% CI)			38			34	100.0%	0.17 [-0.08, 0.42]	—	
Heterogeneity: Tau ² = 0.	.03; Chi ² = 4.57, df =	2 (P = 0.10); I ² =	56%							
Test for overall effect: Z	= 1.30 (P = 0.19)									
2.12.3 HDL cholesterol	[mmol/L]									
Afshar 2010	0.82	0.332	7	0.99	0.32	7	1.4%	-0.17 [-0.51 , 0.17]	<u>-</u>	
Lee 2001	0.99	0.22	21	0.97	0.27	19	7.0%	0.02 [-0.13, 0.17]	_	
Groussard 2015	0.47	0.04	10	0.41	0.05	8	91.5%	0.06 [0.02, 0.10]	•	
Subtotal (95% CI)			38			34	100.0%	0.05 [0.01, 0.09]	T	
Heterogeneity: Tau ² = 0.	.00; Chi ² = 1.92, df =	2 (P = 0.38); I ² =	0%						ľ	
Test for overall effect: Z	= 2.59 (P = 0.010)									
2.12.4 Triglycerides [m	mol/L]									
Afshar 2010	2.55	0.903	7	2.09	0.7	7	28.6%	0.46 [-0.39 , 1.31]		
Lee 2001	2.28	1.49	21	1.5	0.66	19	31.6%	0.78 [0.08, 1.48]		
Groussard 2015	1.09	0.13	10	1.46	0.33	8	39.8%	-0.37 [-0.61 , -0.13]		
Subtotal (95% CI)			38			34	100.0%	0.23 [-0.59 , 1.05]		
Heterogeneity: Tau ² = 0.	.42; Chi ² = 11.69, df	= 2 (P = 0.003); I ²	= 83%							
	I = 0.55 (P = 0.58)									



Analysis 2.13. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 13: Body composition

		Control]	Exercise			Mean Difference		Mean Di	ifference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI		IV, Randoi	m, 95% CI	
2.13.1 Fat mass [kg]												
Giannaki 2013a	28.5	5.8	7	30.5	8.1	15	0.0%	-2.00 [-7.94 , 3.94]			
Kopple 2007	19.1	2.4	14	21.7	8.85	10	0.0%	-2.60 [-8.23, 3.03]		_	
Sheshadri 2020	-0.04	0.1071	25	0	0.1071	24	100.0%	-0.04 [-0.10, 0.02]			
Subtotal (95% CI)			46			49	100.0%	-0.04 [-0.10 , 0.02]	_		
Heterogeneity: Tau ² = 0	.00; Chi ² = 1.	21, df = 2	(P = 0.55)	$I^2 = 0\%$								
Test for overall effect: Z	Z = 1.32 (P =	0.19)										
2.13.2 Lean mass [kg]												
IHOPE 2019	58.6	15.9	38	61.5	13.1	29	39.8%	-2.90 [-9.85 , 4.05]			
Giannaki 2013a	45.4	5.1	7	46.7	8.3	15	60.2%	-1.30 [-6.95 , 4.35]	_		
Subtotal (95% CI)			45			44	100.0%	-1.94 [-6.32 , 2.45	1			
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	12, df = 1	(P = 0.73)	$I^2 = 0\%$								
Test for overall effect: Z			. /									
									+	-10 (
								т.	-20	-10 (th exercise) 10 Higher w	20

Analysis 2.14. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 14: Body mass index

	•	Control		E	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [kg/m²]	SD [kg/m ²]	Total	Mean [kg/m²]	SD [kg/m ²]	Total	Weight	IV, Random, 95% CI [kg/m²]	IV, Random, 95% CI [kg/m²]
IHOPE 2019	31.5	7.4	38	33.9	10.9	29	1.7%	-2.40 [-7.01 , 2.21]	
Liao 2016	23.91	5.27	20	22.96	3.36	20	4.4%	0.95 [-1.79 , 3.69]	· · · · · ·
Koufaki 2002	24.7	3.5	15	25.7	3.3	18	5.7%	-1.00 [-3.34 , 1.34]	_
Wilund 2010	30.3	2.5	9	28.3	1.8	8	7.0%	2.00 [-0.06 , 4.06]	
ACTINUT 2013	20.93	2.57	9	20.89	1.19	7	7.9%	0.04 [-1.86 , 1.94]	
AVANTE-HEMO 2020	19.8	1.7	13	21	2.6	12	9.0%	-1.20 [-2.94, 0.54]	· ·
Kopple 2007	25.1	1.2	14	26.6	1.8	10	13.4%	-1.50 [-2.78, -0.22]	_
Cooke 2018 (1)	0.2067	0.413	10	0.3333	1.015	10	22.9%	-0.13 [-0.81 , 0.55]	.
Sheshadri 2020	0.5	0.7142	25	0.2	0.7142	24	27.9%	0.30 [-0.10 , 0.70]	•
Total (95% CI)			153			138	100.0%	-0.17 [-0.78 , 0.45]	•
Heterogeneity: Tau ² = 0.3	1; Chi ² = 15.11, df	= 8 (P = 0.06);	$I^2 = 47\%$						Ţ
Test for overall effect: Z =	= 0.53 (P = 0.60)								-10 -5 0 5 10
Test for subgroup differer	nces: Not applicabl	e						H	igher with exercise Higher with cont

(1) mean and standard deviation estimated from the median and interquartile range

Analysis 2.15. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 15: Calcium

	Control			E	exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]	
Kouidi 1997	2.15	0.25	11	2.2	0.7	20	2.2%	-0.05 [-0.39 , 0.29]		
de Lima 2013	2.32	0.22	11	2.15	0.32	11	4.6%	0.17 [-0.06 , 0.40]		
Deligiannis 1999a	2.02	0.2	12	2.22	0.15	10	9.5%	-0.20 [-0.35 , -0.05]		
ACTINUT 2013	2.29	0.12	9	2.25	0.12	7	12.7%	0.04 [-0.08, 0.16]		
Wilund 2010	2.2	0.15	9	2.21	0.06	8	14.6%	-0.01 [-0.12 , 0.10]	-	
Momeni 2014	2.27	0.15	20	2.24	0.14	20	17.6%	0.03 [-0.06, 0.12]		
Paluchamy 2018	2.13	0.1	10	2.09	0.09	10	18.9%	0.04 [-0.04, 0.12]	 	
Liao 2016	2.45	0.1	20	2.42	0.15	20	19.9%	0.03 [-0.05, 0.11]	-	
Total (95% CI)			102			106	100.0%	0.01 [-0.04, 0.06]		
Heterogeneity: Tau ² = 0	.00; Chi ² = 11.14, df =	= 7 (P = 0.13); I ² =	37%						Ť	
Test for overall effect: Z	Z = 0.38 (P = 0.71)								-0.5 -0.25 0 0.25 0.5	
Test for subgroup differ	ences: Not applicable								ther with exercise Higher with control	



Analysis 2.16. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 16: C-reactive protein

	Control			F	xercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mg/dL]	SD [mg/dL]	Total	Mean [mg/dL]	SD [mg/dL]	Total	Weight	IV, Random, 95% CI [mg/dL]	IV, Random, 95% CI [mg/dL]
ACTINUT 2013	4.99	5.96	9	1.75	1.62	7	2.8%	3.24 [-0.83 , 7.31]	
Kopple 2007	2.8	2.9933	14	2.5	6.0083	10	2.8%	0.30 [-3.74 , 4.34]	
Toussaint 2008	4.3	0.7	10	7.3	5.5	9	3.4%	-3.00 [-6.62 , 0.62]	
Afshar 2010	4.14	3.87	7	0.88	0.59	7	4.9%	3.26 [0.36, 6.16]	_
Afshar 2011	4.1	3.9	14	0.93	0.66	14	8.0%	3.17 [1.10, 5.24]	
IHOPE 2019	1.136	0.776	12	1.317	1.18	11	17.8%	-0.18 [-1.01 , 0.64]	4
AVANTE-HEMO 2020 (1	0.38	0.349	13	0.843	1.2	12	19.0%	-0.46 [-1.17, 0.24]	-
Wilund 2010	6	0.67	9	4.9	0.69	8	19.5%	1.10 [0.45 , 1.75]	-
Liao 2016	1.23	0.211	20	0.78	0.83	20	21.8%	0.45 [0.07, 0.83]	<u>-</u>
Total (95% CI)			108			98	100.0%	0.60 [-0.12 , 1.32]	•
Heterogeneity: Tau ² = 0.59); Chi ² = 28.05, df	= 8 (P = 0.0005)); I ² = 71%	1					
Test for overall effect: Z =	1.63 (P = 0.10)								-10 -5 0 5 10
Test for subgroup differen	ces: Not applicable	е						Hi	gher with exercise Higher with control

_ . .

Analysis 2.17. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 17: Dialysis adequacy: Kt/V

		Control		1	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Reboredo 2010	1.8	0.7	11	2	0.8	11	3.4%	-0.20 [-0.83 , 0.43]
Parsons 2004	1.22	0.21	7	1.27	0.31	6	10.6%	-0.05 [-0.34 , 0.24]
Frey 1999	1.7	0.1	6	1.8	0.3	5	11.5%	-0.10 [-0.37 , 0.17]
Dobsak 2012	1.33	0.31	10	1.64	0.3	11	12.1%	-0.31 [-0.57 , -0.05	ı <u> </u>
Liao 2016	1.52	0.23	20	1.52	0.26	20	18.8%	0.00 [-0.15, 0.15]]
Fernandes 2019	1.48	0.19	19	1.36	0.19	20	21.1%	0.12 [0.00, 0.24]]
Paluchamy 2018	0.99	0.1265	10	1.15	0.0949	10	22.5%	-0.16 [-0.26 , -0.06	J -
Total (95% CI)			83			83	100.0%	-0.07 [-0.20 , 0.05	
Heterogeneity: Tau ² = 0	0.02; Chi ² = 17	7.13, df =	6 (P = 0.00)	9); I ² = 659	%				•
Test for overall effect:	Z = 1.14 (P =	0.26)							-1 -0.5 0 0.5 1
Test for subgroup differ	rences: Not ap	plicable						H	ligher with exercise Higher with contr

Analysis 2.18. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 18: Energy intake

		Control		I	Exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Weight	IV, Random, 95% CI [kCal/kg/d]	IV, Random, 95% CI [kCal/kg/d]		
Frey 1999	45.76	10.68	6	66.64	18.05	5	6.9%	-20.88 [-38.86 , -2.90	1		
ACTINUT 2013	27.53	8.42	9	30.09	6.64	7	25.1%	-2.56 [-9.94 , 4.82	·]		
Kopple 2007	24.2	7.8575	14	22.4	6.01	10	32.8%	1.80 [-3.75 , 7.35	i]		
IHOPE 2019	19.03	8.5	38	20.04	11.8	29	35.2%	-1.01 [-6.08 , 4.06	i) <u>+</u>		
Total (95% CI)			67			51	100.0%	-1.84 [-6.87 , 3.20	ı 📥		
Heterogeneity: Tau ² = 1	12.06; Chi ² = 5.84, df = 3	3 (P = 0.12); I ² = 49 ⁴	%						٦		
Test for overall effect: 2	Z = 0.72 (P = 0.47)								-50 -25 0 25 5		
est for subgroup differences: Not applicable								I	Higher with exercise Higher with co		

⁽¹⁾ Mean and standard deviation estimated from the median and the interquartile range



Analysis 2.19. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 19: Haemoglobin

	(Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
Reboredo 2010	11.3	2.6	11	10.9	2.8	11	0.8%	0.40 [-1.86 , 2.66]	
AVANTE-HEMO 2020	10.6	1.8	13	9.8	2.3	12	1.5%	0.80 [-0.83 , 2.43]	
Parsons 2004	11.1	1.7	7	11.7	0.7	6	2.0%	-0.60 [-1.98 , 0.78]	
Goldberg 1983	8.8	1.3	7	10	1.5	9	2.0%	-1.20 [-2.57 , 0.17]	
Paluchamy 2018	8.09	1.4546	10	8.35	1.6128	10	2.1%	-0.26 [-1.61 , 1.09]	
Kopple 2007	12.5	1.87	14	12.9	1.26	10	2.5%	-0.40 [-1.65 , 0.85]	
Momeni 2014	9.8	1.7	20	10.03	2.07	20	2.8%	-0.23 [-1.40 , 0.94]	
Painter 2002a (1)	11.81	1.56	26	12.2	1.92	22	3.8%	-0.39 [-1.39 , 0.61]	
CHAIR 2015 (2)	10.93	0.72	11	11.23	1.13	6	3.9%	-0.30 [-1.30 , 0.70]	
Toussaint 2008	12.03	1.41	10	12.09	0.59	9	4.2%	-0.06 [-1.02 , 0.90]	
Koufaki 2002	12.2	1.4	15	12.1	1.3	18	4.5%	0.10 [-0.83 , 1.03]	
Fernandes 2019	10.25	1.66	19	10.7	1.14	20	4.8%	-0.45 [-1.35 , 0.45]	
de Lima 2013	11.1	1.2	11	10.3	0.9	11	4.9%	0.80 [-0.09 , 1.69]	
Tsuyuki 2003	7.4	1.2	12	8	1	17	5.6%	-0.60 [-1.43 , 0.23]	
Lee 2001	8.7	1.3	21	8.2	1.1	19	7.0%	0.50 [-0.24 , 1.24]	 -
ACTINUT 2013	11.21	0.66	9	10.92	0.69	7	8.6%	0.29 [-0.38 , 0.96]	
Afshar 2010	10.2	0.3	7	10.1	0.3	7	39.0%	0.10 [-0.21 , 0.41]	+
Total (95% CI)			223			214	100.0%	0.01 [-0.18, 0.21]	•
Heterogeneity: $Tau^2 = 0.00$; $Chi^2 = 15.31$, $df = 16$ ($P = 0.50$); $I^2 = 0\%$									
Test for overall effect: Z	= 0.14 (P = 0.89	9)						-	4 -2 0 2 4
Test for subgroup differences: Not applicable								High	er with exercise Higher with contr

(2) mean and standard deviation estimated from the median and range

Analysis 2.20. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 20: Heart rate

	Favo	urs exercise		F	xercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [bpm]	SD [bpm]	Total	Mean [bpm]	SD [bpm]	Total	Weight	IV, Random, 95% CI [bpm]	IV, Random, 95% CI [b]	pm]
2.20.1 Resting										
Liao 2016	71	10.5	20	51.9	34.3	20	5.1%	19.10 [3.38 , 34.82]	ı	_
Goldberg 1983	79	15	11	83	11	14	10.8%	-4.00 [-14.57 , 6.57]	ı _	
CYCLE-HD 2016	75.31	13.95	13	72.45	9.36	9	12.7%	2.86 [-6.88 , 12.60]	ı ——	
McGregor 2018	80.94	11.7854	16	74	14.6193	18	15.0%	6.94 [-1.95 , 15.83]	ı	
Tsuyuki 2003	84.3	13.6	12	81.9	8.7	17	15.5%	2.40 [-6.34 , 11.14]	1 📥	
Deligiannis 1999a	81.8	8.5	12	78.4	10.5	10	17.8%	3.40 [-4.69 , 11.49]	ı -	
Koh 2009	75	12	16	70	10.7	30	23.1%	5.00 [-2.02 , 12.02]	l +	
Subtotal (95% CI)			100			118	100.0%	4.07 [0.49, 7.65]	♠	
Heterogeneity: Tau ² =	1.63; Chi ² = 6.44, c	lf = 6 (P = 0.3)	8); I ² = 7%	5					\ \	
Test for overall effect:	Z = 2.23 (P = 0.03))								
2.20.2 Maximum										
McGregor 2018	122.5	27.107	18	126.88	23.6646	16	10.3%	-4.38 [-21.45 , 12.69]	ı 	
Akiba 1995	136.3	19.5	6	155.4	8.6	7	10.5%	-19.10 [-35.95 , -2.25]	ı <u> </u>	
Koufaki 2002	127.2	24.4	15	129	22.7	18	11.4%	-1.80 [-18.00 , 14.40]	1	
Painter 2002a (1)	128.72	27.44	25	137.78	24.68	23	13.8%	-9.06 [-23.81, 5.69]	ı 	
Tsuyuki 2003	155.8	20.7	12	164.2	10.2	17	18.6%	-8.40 [-21.08 , 4.28]	ı <u> </u>	
Deligiannis 1999a	139	12	12	142	10	10	35.4%	-3.00 [-12.19 , 6.19]	ı _ _	
Subtotal (95% CI)			88			91	100.0%	-6.54 [-12.01 , -1.07]	ı 📤	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 3.29, c	lf = 5 (P = 0.6	6); I ² = 0%	,					•	
Test for overall effect:	Z = 2.34 (P = 0.02))								
									-50 -25 0 25	
Footnotes								H	igher with exercise Higher	with

⁽¹⁾ factorial design: two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

⁽¹⁾ factorial design: two intervention arms pooled together in the exercise group and two control arms pooled together in the control group



Analysis 2.21. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 21: Left ventricular ejection fraction

		Control		1	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [%]	SD [%]	Total	Mean [%]	SD [%]	Total	Weight	IV, Random, 95% CI [%	IV, Random, 95% CI [%]
Reboredo 2010	71.4	7.6	11	70.4	12	11	6.4%	1.00 [-7.39 , 9.3	39]
McGregor 2018	53.22	7.1186	18	54.22	9.8337	16	11.7%	-1.00 [-6.83 , 4.8	33]
Cho 2018	0.3	3.9	13	2.5	5.2	11	21.3%	-2.20 [-5.93 , 1.5	53]
Momeni 2014	54.25	4.66	20	58.5	3.67	20	30.2%	-4.25 [-6.85 , -1.6	65]
Kouidi 2004a	0.2	3	10	-0.3	3	11	30.5%	0.50 [-2.07 , 3.0)7]
Total (95% CI)			72			69	100.0%	-1.65 [-3.93 , 0.6	52]
Heterogeneity: Tau ² =	2.70; Chi ² = 7.0	3, df = 4 (P)	= 0.13); 1	$I^2 = 43\%$					
Test for overall effect:	Z = 1.42 (P = 0)	.15)							-10 -5 0 5 10
Test for subgroup diffe	Test for subgroup differences: Not applicable								Higher with exercise Higher with control

Analysis 2.22. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 22: Left ventricular mass index

	Control			F	exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [g/m²]	SD [g/m ²]	Total	Mean [g/m²]	SD [g/m ²]	Total	Weight	IV, Random, 95% CI [g/m²]	IV, Random, 95% CI [g/m²]		
Reboredo 2010	131.3	48.4	11	120.9	26.6	11	12.0%	10.40 [-22.24 , 43.04	4]		
McGregor 2018	111.65	39.7958	18	139.2	45.6215	16	14.9%	-27.55 [-56.49 , 1.39	9]		
Deligiannis 1999a	137	35	12	147	27	10	18.1%	-10.00 [-35.93 , 15.93	3]		
Wilund 2010	127.4	18.17	9	154.4	25.83	8	24.9%	-27.00 [-48.48 , -5.52	2]		
Cho 2018	-11.7	30.1	13	-1.5	16.7	11	30.1%	-10.20 [-29.31 , 8.9	1]		
Total (95% CI)			63			56	100.0%	-14.47 [-26.25 , -2.69	9]		
Heterogeneity: Tau ² = 2	24.90; Chi ² = 4.63,	df = 4 (P = 0.3)	33); I ² = 14	4%					•		
Test for overall effect: 2	Z = 2.41 (P = 0.02))							-100 -50 0 50 100		
Test for subgroup differences: Not applicable								1	Higher with exercise Higher with control		

Analysis 2.23. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 23: Muscular strength

		Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [kg]	SD [kg]	Total	Mean [kg]	SD [kg]	Total	Weight	IV, Random, 95% CI [kg	IV, Random, 95% CI [kg]
2.23.1 Knee extension									
Dobsak 2012	85.36	18.32	10	104.68	14.23	11	20.2%	-19.32 [-33.45 , -5.1	9]
DIALY-SIZE 2016	9.3	10.1	8	11.6	10.7	8	29.0%	-2.30 [-12.50 , 7.9	0]
ACTINUT 2013	7.87	2.19	9	10.56	3.49	7	50.8%	-2.69 [-5.64, 0.2	6]
Subtotal (95% CI)			27			26	100.0%	-5.94 [-13.95 , 2.0	7]
Heterogeneity: Tau ² = 3	0.62; Chi ² = 5.	14, df = 2 (I	P = 0.08;	$I^2 = 61\%$					_
Test for overall effect: 2	Z = 1.45 (P = 0.	15)							
2.23.2 handgrip									
Koh 2009 (1)	31	12	7	35.97	12.35	29	15.9%	-4.97 [-14.93 , 4.9	9]
Samara 2016	32.3	9.9	12	37.2	14.7	15	17.5%	-4.90 [-14.21 , 4.4	1]
Wu 2014d	28.6	9	33	37.8	12.9	32	31.0%	-9.20 [-14.62 , -3.7	8]
Cooke 2018 (2)	2	3.87	10	2.43	6.02	10	35.6%	-0.43 [-4.87 , 4.0	1]
Subtotal (95% CI)			62			86	100.0%	-4.65 [-9.44 , 0.1	4]
Heterogeneity: Tau ² = 1	1.62; Chi ² = 6.0	08, df = 3 (I	P = 0.11); 1	[2 = 51%					•
Test for overall effect: 2	Z = 1.90 (P = 0.	06)							
									-50 -25 0 25
Footnotes									-50 -25 0 25 Higher with exercise Higher wit

 $^{(1)\} two\ intervention\ arms\ polled\ together\ in\ the\ exercise\ group$

⁽²⁾ mean and standard deviation estimated from the median and interquartile range



Analysis 2.24. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 24: Phosphate

	Control				exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
AVANTE-HEMO 2020 (1)) 1.86	1.15	13	1.93	0.87	12	2.0%	-0.07 [-0.87 , 0.73]	
Reboredo 2010	1.91	0.61	11	1.58	0.55	11	5.0%	0.33 [-0.16, 0.82]	
ACTINUT 2013	1.59	0.42	9	1.32	0.42	7	6.6%	0.27 [-0.14, 0.68]	
de Lima 2013	1.81	0.29	11	1.74	0.55	10	7.5%	0.07 [-0.31 , 0.45]	
Deligiannis 1999a	1.94	0.42	12	2.07	0.36	10	9.6%	-0.13 [-0.46, 0.20]	
Paluchamy 2018	1.6855	0.4799	10	1.4853	0.143	10	10.4%	0.20 [-0.11, 0.51]	
Kouidi 1997	1.97	0.23	11	2.03	0.55	20	12.2%	-0.06 [-0.34 , 0.22]	
Wilund 2010	2.1	0.25	9	1.91	0.16	8	18.5%	0.19 [-0.01, 0.39]	
Momeni 2014	1.69	0.27	20	1.78	0.03	20	28.1%	-0.09 [-0.21 , 0.03]	-
Total (95% CI)			106			108	100.0%	0.05 [-0.07 , 0.17]	
Heterogeneity: Tau ² = 0.01	1; Chi ² = 11.61, df =	8 (P = 0.17); I ² =	31%						
Test for overall effect: Z =							-1 -0.5 0 0.5 1		
Test for subgroup differen	ces: Not applicable							Higl	ner with exercise Higher with control

Footnote

 $(1) \ Mean \ and \ standard \ deviation \ estimated \ from \ the \ median \ and \ the \ interquartile \ range$

Analysis 2.25. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 25: Potassium

	Control			I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Deligiannis 1999a	5.4	0.4	12	5.5	1	10	5.9%	-0.10 [-0.76 , 0.56]	
AVANTE-HEMO 2020	5.2	0.92	13	5.5	0.74	12	6.1%	-0.30 [-0.95 , 0.35]	-
Parsons 2004	4.8	0.6	7	4.4	0.4	6	8.6%	0.40 [-0.15, 0.95]	
Momeni 2014	5.47	0.95	20	5.4	0.81	20	8.7%	0.07 [-0.48 , 0.62]	
de Lima 2013	5.8	0.6	11	5.6	0.7	11	8.7%	0.20 [-0.34 , 0.74]	
Kouidi 1997	5.5	0.7	11	5.6	0.5	20	11.8%	-0.10 [-0.57, 0.37]	
Paluchamy 2018	5.65	0.5692	10	5.18	0.4743	10	12.3%	0.47 [0.01, 0.93]	
Wilund 2010	4.9	0.3	9	4.9	0.25	8	37.9%	0.00 [-0.26 , 0.26]	-
Total (95% CI)			93			97	100.0%	0.08 [-0.08, 0.24]	
Heterogeneity: Tau ² = 0.0	0; Chi ² = 6.79, df = 1	7 (P = 0.45); I ² = 0	%						
Test for overall effect: Z = 0.97 (P = 0.33)									-1 -0.5 0 0.5 1
Test for subgroup differer	nces: Not applicable							Higl	her with exercise Higher with contr

Analysis 2.26. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 26: Protein intake

	(Control		Exercise				Mean Difference	Mean Difference		
Study or Subgroup	Mean [g/kg/d]	SD [g/kg/d]	Total	Mean [g/kg/d]	SD [g/kg/d]	Total	Weight	IV, Random, 95% CI [g/kg/d]	IV, Random, 95%	% CI [g/kg/d]	
Frey 1999	1.91	0.34	6	2.6	1.35	5	5.1%	-0.69 [-1.90 , 0.52]		-	
Kopple 2007	1.35	0.5987	14	0.91	0.32	10	26.5%	0.44 [0.07, 0.81]	-	-	
ACTINUT 2013	1.17	0.38	9	1.17	0.26	7	30.1%	0.00 [-0.31 , 0.31]	•		
IHOPE 2019	0.9	0.33	38	1	0.46	29	38.2%	-0.10 [-0.30 , 0.10]	+		
Total (95% CI)			67			51	100.0%	0.04 [-0.25 , 0.33]	•		
Heterogeneity: Tau ² = 0	0.05; Chi ² = 7.62, df	= 3 (P = 0.05); I	$^{2} = 61\%$						[
Test for overall effect: 2	Z = 0.29 (P = 0.77)								-4 -2 0	2 4	
Test for subgroup differ	rences: Not applicab	le						Hi	gher with exercise	Higher with control	



Analysis 2.27. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 27: Parathyroid hormone

	Control				Exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [pmol/L]	SD [pmol/L]	Total	Mean [pmol/L]	SD [pmol/L]	Total	Weight	IV, Random, 95% CI [pmol/L]	IV, Random, 95	% CI [pmol/L]	
Toussaint 2008	42.7	57.2	10	37.5	45.9	9	11.7%	5.20 [-41.23 , 51.63	B]	•——	
Koufaki 2002	13.6	15.8	15	34.8	50.7	18	28.2%	-21.20 [-45.95 , 3.55	i]	-	
Liao 2016	30.2	9.42	20	25.74	7.92	20	60.0%	4.46 [-0.93 , 9.85	5]		
Total (95% CI)			45			47	100.0%	-2.69 [-20.31 , 14.93	3]		
Heterogeneity: Tau ² = 1	127.03; Chi ² = 3.95, d	f = 2 (P = 0.14);	$I^2 = 49\%$								
Test for overall effect:	Z = 0.30 (P = 0.76)								-100 -50 0	50 100	
Test for subgroup diffe	rences: Not applicable	e						H	Higher with exercise	Higher with contr	

Analysis 2.28. Comparison 2: Aerobic exercise versus control (no exercise/placebo exercise), Outcome 28: Timed up-and-go test

	Control			I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [sec]	SD [sec]	Total	Mean [sec]	SD [sec]	Total	Weight	IV, Random, 95% CI [sec]	IV, Random, 95% CI [sec]
Wu 2014d	30.2	8.3	33	27.3	7.3	32	5.0%	2.90 [-0.90 , 6.70]	
Samara 2016	6.6	2.5	12	4.6	1.4	15	22.9%	2.00 [0.42 , 3.58]	
IHOPE 2019	8	3.6	38	6.2	1.7	29	30.1%	1.80 [0.50, 3.10]	_
Koh 2009 (1)	6.1	1.5	16	5.54	1.8	29	42.0%	0.56 [-0.42 , 1.54]	-
Total (95% CI)			99			105	100.0%	1.38 [0.50 , 2.26]	•
Heterogeneity: Tau ² = 0	0.23; Chi ² = 4.16	df = 3 (P =	0.24); I ² =	28%					•
Test for overall effect: 2	Z = 3.08 (P = 0.0)	002)							-10 -5 0 5 10
Test for subgroup differ	rences: Not appli	icable						High	ner with exercise Higher with control

Footnotes

(1) two intervention arms polled together in the exercise group

Comparison 3. Resistance exercise versus control (no exercise/placebo exercise)

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
3.1 Fatigue	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
3.2 HRQoL: Summary component scores	5		Mean Difference (IV, Random, 95% CI)	Subtotals only
3.2.1 Physical Component Score	5	176	Mean Difference (IV, Random, 95% CI)	-2.52 [-6.32, 1.29]
3.2.2 Mental Component Score	5	176	Mean Difference (IV, Random, 95% CI)	0.68 [-4.57, 5.94]
3.3 HR-QoL: Individual domains	7		Mean Difference (IV, Random, 95% CI)	Subtotals only
3.3.1 Physical functioning	6	243	Mean Difference (IV, Random, 95% CI)	-5.28 [-10.09, -0.46]
3.3.2 Role-physical	3	102	Mean Difference (IV, Random, 95% CI)	-8.13 [-21.33, 5.07]
3.3.3 Pain	5	154	Mean Difference (IV, Random, 95% CI)	-10.74 [-27.96, 6.47]
3.3.4 General health perceptions	4	126	Mean Difference (IV, Random, 95% CI)	-0.05 [-6.43, 6.33]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size	
3.3.5 Emotional well-being	4	126	Mean Difference (IV, Random, 95% CI)	-7.22 [-13.98, -0.46]	
3.3.6 Role-emotional	4	126	Mean Difference (IV, Random, 95% CI)	-4.25 [-14.62, 6.12]	
3.3.7 Vitality	5	179	Mean Difference (IV, Random, 95% CI)	-5.17 [-15.18, 4.85]	
3.3.8 Social function	4	126	Mean Difference (IV, Random, 95% CI)	-9.28 [-17.08, -1.47]	
3.3.9 Symptoms	3	86	Mean Difference (IV, Random, 95% CI)	-9.25 [-15.19, -3.30]	
3.3.10 Effects of kidney disease	2	58	Mean Difference (IV, Random, 95% CI)	-4.87 [-16.82, 7.08]	
3.3.11 Burden of kidney disease	2	58	Mean Difference (IV, Random, 95% CI)	3.35 [-9.05, 15.75]	
3.3.12 Work status	2	58	Mean Difference (IV, Random, 95% CI)	4.30 [-14.99, 23.59]	
3.3.13 Cognitive function	2	58	Mean Difference (IV, Random, 95% CI)	6.31 [-6.73, 19.36]	
3.3.14 Quality of social interactions	2	58	Mean Difference (IV, Random, 95% CI)	8.30 [-3.74, 20.34]	
3.3.15 Sexual function	2	58	Mean Difference (IV, Random, 95% CI)	-14.16 [-35.63, 7.31]	
3.3.16 Sleep	3	86	Mean Difference (IV, Random, 95% CI)	-10.70 [-20.99, -0.40]	
3.3.17 Social support	2	58	Mean Difference (IV, Random, 95% CI)	-4.41 [-13.92, 5.09]	
3.3.18 Dialysis staff encouragement	2	58	Mean Difference (IV, Random, 95% CI)	-1.10 [-8.72, 6.52]	
3.3.19 Patient satisfaction	2	58	Mean Difference (IV, Random, 95% CI)	-1.07 [-10.75, 8.60]	
3.4 Depression	2	99	Std. Mean Difference (IV, Random, 95% CI)	0.52 [0.12, 0.92]	
3.5 6MWT	7	216	Mean Difference (IV, Random, 95% CI)	-44.71 [-62.43, -27.00]	
3.6 Sit-To-Stand test [N reps/30 sec]	6	196	Mean Difference (IV, Random, 95% CI)	-2.76 [-3.83, -1.68]	
3.7 Sit-To-Stand test [N reps/30 sec]	2	93	Mean Difference (IV, Random, 95% CI)	1.56 [-0.44, 3.57]	
3.8 Albumin	9	268	Mean Difference (IV, Random, 95% CI)	-0.27 [-1.59, 1.05]	
3.9 Blood lipids	3		Mean Difference (IV, Random, 95% CI) Subtotals		
3.9.1 Total cholesterol [mmol/L]	3	76	Mean Difference (IV, Random, 95% CI)	0.26 [-0.07, 0.58]	



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
3.9.2 LDL cholesterol [mmol/L]	2	54	Mean Difference (IV, Random, 95% CI)	0.12 [-0.17, 0.41]
3.9.3 HDL cholesterol [mmol/L]	2	54	Mean Difference (IV, Random, 95% CI)	0.02 [-0.16, 0.19]
3.9.4 Triglycerides [mmol/L]	2	54	Mean Difference (IV, Random, 95% CI)	0.54 [-0.00, 1.07]
3.10 Body composition	7		Mean Difference (IV, Random, 95% CI)	Subtotals only
3.10.1 Fat mass [kg]	7	291	Mean Difference (IV, Random, 95% CI)	-0.69 [-2.94, 1.56]
3.10.2 Lean mass [kg]	5	224	Mean Difference (IV, Random, 95% CI)	0.16 [-2.95, 3.28]
3.11 Body mass index	8	267	Mean Difference (IV, Random, 95% CI)	-1.00 [-1.98, -0.01]
3.12 Calcium	4	102	Mean Difference (IV, Random, 95% CI)	0.04 [-0.08, 0.16]
3.13 CRP	6	153	Mean Difference (IV, Random, 95% CI)	-0.22 [-0.58, 0.14]
3.14 Dialysis adequacy: Kt/V	2	73	Mean Difference (IV, Random, 95% CI)	-0.05 [-0.23, 0.12]
3.15 Energy intake	5	208	Mean Difference (IV, Random, 95% CI)	0.17 [-1.45, 1.80]
3.16 Haemoglobin	10	254	Mean Difference (IV, Random, 95% CI)	-0.11 [-0.29, 0.07]
3.17 Muscular strength	8		Mean Difference (IV, Random, 95% CI)	Subtotals only
3.17.1 knee extension	6	238	Mean Difference (IV, Random, 95% CI)	-6.09 [-10.68, -1.50]
3.17.2 handgrip	3	137	Mean Difference (IV, Random, 95% CI)	-2.01 [-5.71, 1.69]
3.18 Phosphate	7	188	Mean Difference (IV, Random, 95% CI)	-0.06 [-0.36, 0.24]
3.19 Potassium	7	188	Mean Difference (IV, Random, 95% CI)	0.32 [-0.23, 0.86]
3.20 Protein intake	5	208	Mean Difference (IV, Random, 95% CI)	-0.01 [-0.09, 0.07]
3.21 PTH	2	37	Mean Difference (IV, Random, 95% CI)	1.51 [-30.38, 33.39]
3.22 Timed up-and-go test	1		Mean Difference (IV, Random, 95% CI)	Totals not selected

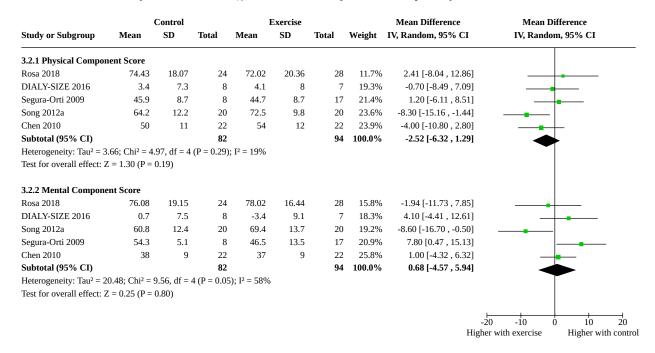


Analysis 3.1. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 1: Fatigue

		Control]	Exercise		Mean Difference	Mean I	Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	IV, Random, 95% CI	IV, Rande	om, 95% CI
Johansen 2006 (1)	8.95	4.71	33	7.07	4.78	35	1.88 [-0.38 , 4.14	.]	
								-10 -5	0 5 10
Footnotes							I	Higher with exercise	Higher with control

⁽¹⁾ Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

Analysis 3.2. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 2: HRQoL: Summary component scores





Analysis 3.3. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 3: HR-QoL: Individual domains

Study or Subgroup	Mean	Control SD	Total	Mean	Exercise SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
3.3.1 Physical functioning									
Martins do Valle 2020	63.1	24.5	12	72.5	20.2	12	7.2%	-9.40 [-27.37, 8.57]	
Martin-Alemany 2016	59.7	26.4	19	71.3	22.3	17	9.2%	-11.60 [-27.51 , 4.31]	
AVANTE-HEMO 2020	84.44	22.7	13	80	13.2	9	10.2%	4.44 [-10.61 , 19.49]	
Johansen 2006 (1)	56.58	26.72	33	59.12	30.37	35	12.6%	-2.54 [-16.12 , 11.04]	
Abreu 2017	85	13	19	87	18	25	27.6%	-2.00 [-11.16 , 7.16]	
PEAK 2006	-1.8	17.6	25	7.6	11.8	24	33.2%	-9.40 [-17.76 , -1.04]	
	-1.0	17.0		7.0	11.0				-
Subtotal (95% CI)	CI :2 D.O.	0 16 5 00	121	00/		122	100.0%	-5.28 [-10.09 , -0.46]	◆
Heterogeneity: $Tau^2 = 0.00$; Test for overall effect: $Z = 2$			= 0.55); 12	= 0%					
3.3.2 Role-physical									
AVANTE-HEMO 2020	88.89	33.3	13	85.71	37.8	9	17.9%	3.18 [-27.44, 33.80]	
Martin-Alemany 2016	68.8	41.2	19	65.6	40.7	17	23.1%	3.20 [-23.59, 29.99]	
Abreu 2017	63	27	19	79	27	25	59.0%	-16.00 [-32.11 , 0.11]	
Subtotal (95% CI)		=-	51				100.0%	-8.13 [-21.33 , 5.07]	
Heterogeneity: Tau ² = 9.35;	Chi ² = 2.1	2 df = 2 (D		= 6%		31		2.22 [21.00 ; 5.07]	
Test for overall effect: $Z = 1$			- 0.55), 1	- 070					
3.3.3 Pain	60	22.0	10	E0.4	20.4	10	16.70/	C CO [17 10 20 20]	
Martins do Valle 2020	60	32.9	12	53.4	26.1	12	16.7%	6.60 [-17.16 , 30.36]	- - -
Martin-Alemany 2016	65.4	34.7	19	77.3	21.2	17	19.0%	-11.90 [-30.47 , 6.67]	+
AVANTE-HEMO 2020	85	18.3	13	86.79	14.6	9	21.1%	-1.79 [-15.57 , 11.99]	+
Abreu 2017	82	23	19	85	19	25	21.5%	-3.00 [-15.74, 9.74]	-
Pellizzaro 2013	-15.5	16.7741	14	24	16.7741	14	21.7%	-39.50 [-51.93 , -27.07]	
Subtotal (95% CI)			77			77	100.0%	-10.74 [-27.96, 6.47]	
				01), 12 = 0	1%				
Test for overall effect: $Z = 1$ 3.3.4 General health perception	.22 (P = 0.		I (P < 0.00	01), 1 64	•70				
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020	.22 (P = 0.		13 12 19	52.86 52.7 78	18.2 19.7 17	9 12 25	16.9% 21.4% 28.4%	0.47 [-14.76 , 15.70] -2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56]	-
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017	.22 (P = 0. ptions 53.33 50.5	17.5 13.3	13 12	52.86 52.7	18.2 19.7	12	21.4%	-2.20 [-15.65 , 11.25]	-
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016	22 (P = 0. ptions 53.33 50.5 71	17.5 13.3 21	13 12 19	52.86 52.7 78	18.2 19.7 17	12 25	21.4% 28.4%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56]	-
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI)	.22 (P = 0. ptions 53.33 50.5 71 51	17.5 13.3 21 14.1	13 12 19 19	52.86 52.7 78 44	18.2 19.7 17	12 25 17	21.4% 28.4% 33.3%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau ² = 2.54;	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1	17.5 13.3 21 14.1 9, df = 3 (P	13 12 19 19	52.86 52.7 78 44	18.2 19.7 17	12 25 17	21.4% 28.4% 33.3%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being	ptions 53.33 50.5 71 51 Chi² = 3.1'	17.5 13.3 21 14.1 9, df = 3 (P	13 12 19 19 63 = 0.36); I ²	52.86 52.7 78 44 = 6%	18.2 19.7 17 17.9	12 25 17 63	21.4% 28.4% 33.3% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020	.22 (P = 0. ptions 53.33 50.5 71 51 .02 (P = 0.	22) 17.5 13.3 21 14.1 9, df = 3 (P 99)	13 12 19 19 63 = 0.36); I ²	52.86 52.7 78 44 = 6%	18.2 19.7 17 17.9	12 25 17 63	21.4% 28.4% 33.3% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020	.22 (P = 0. ptions 53.33 50.5 71 51 .02 (P = 0. 70 75.11	17.5 13.3 21 14.1 9, df = 3 (P 99)	13 12 19 19 63 = 0.36); I ²	52.86 52.7 78 44 = 6%	18.2 19.7 17 17.9	12 25 17 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0.	17.5 13.3 21 14.1 9, df = 3 (P 99)	13 12 19 19 63 = 0.36); I ² 12 13	52.86 52.7 78 44 = 6% 65 82.29 76.8	18.2 19.7 17 17.9 29.6 8.5 19.4	12 25 17 63	21.4% 28.4% 33.3% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016	.22 (P = 0. ptions 53.33 50.5 71 51 .02 (P = 0. 70 75.11	17.5 13.3 21 14.1 9, df = 3 (P 99)	13 12 19 19 63 = 0.36); I ²	52.86 52.7 78 44 = 6%	18.2 19.7 17 17.9	12 25 17 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35]	**************************************
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0.	17.5 13.3 21 14.1 9, df = 3 (P 99)	13 12 19 19 63 = 0.36); I ² 12 13	52.86 52.7 78 44 = 6% 65 82.29 76.8	18.2 19.7 17 17.9 29.6 8.5 19.4	12 25 17 63 12 9	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00;	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 70 75.11 65.6 78 Chi² = 1.9	22) 17.5 13.3 21 14.1 9, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 19 63	52.86 52.7 78 44 = 6% 65 82.29 76.8 86	18.2 19.7 17 17.9 29.6 8.5 19.4	12 25 17 63 12 9 17 25	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 70 75.11 65.6 78 Chi² = 1.9	22) 17.5 13.3 21 14.1 9, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 70 75.11 65.6 78 Chi² = 1.9 .09 (P = 0.	22) 17.5 13.3 21 14.1 9, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 19 63	52.86 52.7 78 44 = 6% 65 82.29 76.8 86	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 70 75.11 65.6 78 Chi² = 1.9	22) 17.5 13.3 21 14.1 9, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 70 75.11 65.6 78 Chi² = 1.9 .09 (P = 0.	22) 17.5 13.3 21 14.1 9, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1² .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0.	17.5 13.3 21 14.1 9, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0.	22) 17.5 13.3 21 14.1 19, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3	13 12 19 19 63 = 0.36); I ² 12 13 19 19 63 = 0.58); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00;	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2	17.5 13.3 21 14.1 19, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 12 19 13 63	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2	17.5 13.3 21 14.1 19, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 12 19 13 63	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 3.3.7 Vitality	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2	17.5 13.3 21 14.1 19, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 12 19 13 63	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15	12 25 17 63 12 9 17 25 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Avantin-Alemany 2016 Abreu 2017 Martin-Alemany 2016 Avantin-Alemany 2016 Avantin-Alemany 2016 Avantin-Alemany 2016 Avantin-Alemany 2016 Avantin-Alemany 2016 Avantin-Alemany 2016 Avantin-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 3.3.7 Vitality Pellizzaro 2013	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2 .80 (P = 0.	22) 17.5 13.3 21 14.1 19, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P 42)	13 12 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 12 19 19 13 63 = 0.52); I ²	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0% 75 76 85.4 90.49 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15 38.8 38.29.7 16.2	12 25 17 63 12 9 17 25 63 12 25 17 9 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0% 22.9% 25.4% 41.8% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-55.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15] -4.25 [-14.62 , 6.12]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 3.3.7 Vitality Pellizzaro 2013 Martin-Alemany 2016	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2 .80 (P = 0.	22) 17.5 13.3 21 14.1 9, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P 42)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 14 19	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0% 75 76 85.4 90.49 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15 38.8 38 29.7 16.2	12 25 17 63 12 9 17 25 63 12 25 17 9 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0% 22.9% 25.4% 41.8% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15] -4.25 [-14.62 , 6.12]	
Test for overall effect: Z = 1 3.3.4 General health percel AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 3.3.7 Vitality Pellizzaro 2013 Martin-Alemany 2016 AVANTE-HEMO 2020 3.3.7 Vitality Pellizzaro 2013 Martin-Alemany 2016 AVANTE-HEMO 2020	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2 .80 (P = 0.	22) 17.5 13.3 21 14.1 19, df = 3 (P 99) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 14 19 13	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0% 75 76 85.4 90.49 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15 38.8 38 29.7 16.2 28.5212 26 9.5	12 9 17 63 12 9 17 25 63 12 25 17 9 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0% 22.9% 25.4% 41.8% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15] -4.25 [-14.62 , 6.12] -27.50 [-48.63 , -6.37] 10.90 [-4.38 , 26.18] 2.78 [-10.98 , 16.54]	
Heterogeneity: Tau² = 315.8 Test for overall effect: Z = 1 3.3.4 General health percey AVANTE-HEMO 2020 Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 Subtotal (95% CI) Heterogeneity: Tau² = 2.54; Test for overall effect: Z = 0 3.3.5 Emotional well-being Martins do Valle 2020 AVANTE-HEMO 2020 Martin-Alemany 2016 Abreu 2017 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 2 3.3.6 Role-emotional Martins do Valle 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 Subtotal (95% CI) Heterogeneity: Tau² = 0.00; Test for overall effect: Z = 0 3.3.7 Vitality Pellizzaro 2013 Martin-Alemany 2016 AVANTE-HEMO 2020 AVANTE-HEMO 2020 Abreu 2017 Martin-Alemany 2016 AVANTE-HEMO 2020 AVANTE-HEMO 2020 Abreu 2017 PEAK 2006	.22 (P = 0. ptions 53.33 50.5 71 51 Chi² = 3.1¹ .02 (P = 0. 75.11 65.6 78 Chi² = 1.9 .09 (P = 0. 54.1 76 73.4 92.59 Chi² = 2.2 .80 (P = 0.	22) 17.5 13.3 21 14.1 9, df = 3 (P 999) 16.7 24.7 17.3 22 9, df = 3 (P 04) 43.3 35 33.3 22.2 8, df = 3 (P 42)	13 12 19 19 63 = 0.36); I ² 12 13 19 63 = 0.58); I ² 14 19	52.86 52.7 78 44 = 6% 65 82.29 76.8 86 = 0% 75 76 85.4 90.49 = 0%	18.2 19.7 17 17.9 29.6 8.5 19.4 15 38.8 38 29.7 16.2	12 25 17 63 12 9 17 25 63 12 25 17 9 63	21.4% 28.4% 33.3% 100.0% 12.4% 21.7% 31.4% 34.5% 100.0% 22.9% 25.4% 41.8% 100.0%	-2.20 [-15.65 , 11.25] -7.00 [-18.56 , 4.56] 7.00 [-3.61 , 17.61] -0.05 [-6.43 , 6.33] 5.00 [-14.23 , 24.23] -7.18 [-21.71 , 7.35] -11.20 [-23.26 , 0.86] -8.00 [-19.51 , 3.51] -7.22 [-13.98 , -0.46] -20.90 [-53.80 , 12.00] 0.00 [-21.67 , 21.67] -12.00 [-32.58 , 8.58] 2.10 [-13.95 , 18.15] -4.25 [-14.62 , 6.12]	

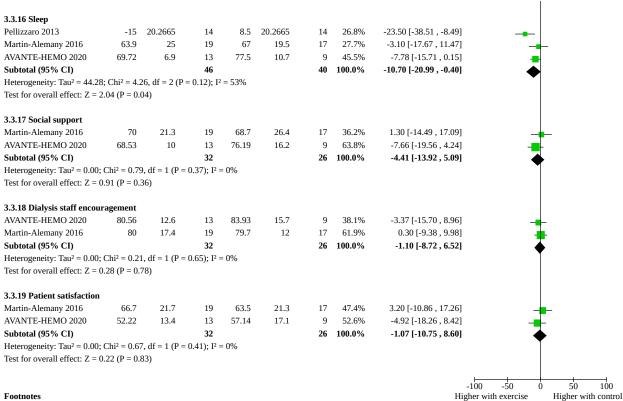


Analysis 3.3. (Continued)

,,0.00.000	,								
Abreu 2017	6/	21	19	/4	22	25	21.4%	-/.00 [-19./9, 5./9]	 +
PEAK 2006	-7	14.1	25	2.8	16.3	24	26.5%	-9.80 [-18.35 , -1.25]	-
Subtotal (95% CI)			90			89	100.0%	-5.17 [-15.18 , 4.85]	•
Heterogeneity: $Tau^2 = 79.69$			= 0.03);]	$I^2 = 64\%$					
Test for overall effect: $Z = 1$	1.01 (P = 0.3)	1)							
3.3.8 Social function									
Martins do Valle 2020	64	35.6	12	79.7	29.8	12	8.8%	-15.70 [-41.97 , 10.57]	
Martin-Alemany 2016	76.4	25.5	19	79.8	29	17	18.9%	-3.40 [-21.33 , 14.53]	
Abreu 2017	76	26	19	91	19	25	31.7%	-15.00 [-28.86 , -1.14]	
AVANTE-HEMO 2020	90.28	19.5	13	96.43	9.4	9	40.6%	-6.15 [-18.40 , 6.10]	
Subtotal (95% CI)			63			63	100.0%	-9.28 [-17.08 , -1.47]	
Heterogeneity: Tau ² = 0.00;	Chi ² = 1.55	df = 3 (P =		= 0%		0.5	100.070	3120 [17100 ; 1717]	
Test for overall effect: $Z = 2$			0.07), 1	070					
3.3.9 Symptoms									
	76.87	13.4	13	83.94	12.5	9	29.5%	7.07 [10.01 2.07]	
AVANTE-HEMO 2020		16.6						-7.07 [-18.01 , 3.87]	
Martin-Alemany 2016	70.1		19	76.6	14.8	17	33.6%	-6.50 [-16.76 , 3.76]	
Pellizzaro 2013	0	13.2288	14	13.5	13.2288	14	36.8%	-13.50 [-23.30 , -3.70]	
Subtotal (95% CI)			46			40	100.0%	-9.25 [-15.19 , -3.30]	◆
Heterogeneity: Tau ² = 0.00;			0.56); I ² =	= 0%					
Test for overall effect: $Z = 3$	0.05 (P = 0.0	02)							
3.3.10 Effects of kidney dis									
AVANTE-HEMO 2020	64.59	18.1	13	76.79	23.8	9	40.9%	-12.20 [-30.60 , 6.20]	
Martin-Alemany 2016	70.2	21.1	19	70	25	17	59.1%	0.20 [-15.01 , 15.41]	
Subtotal (95% CI)			32			26	100.0%	-4.87 [-16.82 , 7.08]	
Heterogeneity: Tau ² = 2.71;	Chi ² = 1.04	, df = 1 (P =	0.31); I ² =	= 4%					7
Test for overall effect: $Z = 0$			**						
		,							
3.3.11 Burden of kidney di		20.4	10	44 -	20.0	17	49 20/	0.20 [10.05 10.05]	
Martin-Alemany 2016	44.3	26.4	19	44.5	30.8	17	43.3%	-0.20 [-19.05 , 18.65]	
AVANTE-HEMO 2020	56.96	19.6	13	50.9	19.2	9	56.7%	6.06 [-10.40 , 22.52]	-
Subtotal (95% CI)			32			26	100.0%	3.35 [-9.05 , 15.75]	•
Heterogeneity: Tau ² = 0.00; Test for overall effect: Z = 0			0.62); I ² =	= 0%					
3.3.12 Work status AVANTE-HEMO 2020	50	35.4	13	42.86	34.4	9	42.5%	7.14 [-22.45 , 36.73]	
Martin-Alemany 2016	27.5	41.2	19	25.3	36.7	17	57.5%	2.20 [-23.25 , 27.65]	
Subtotal (95% CI)	CI 12	10	32	001		26	100.0%	4.30 [-14.99 , 23.59]	•
Heterogeneity: Tau ² = 0.00; Test for overall effect: Z = 0			0.80); I ² =	= 0%					
	, 2.0	,							
3.3.13 Cognitive function	24.40	27.2	17	20	21.4	0	41.00/	4.46 [15.02 24.05]	
AVANTE-HEMO 2020	24.46	27.3	13	20	21.4	9	41.0%	4.46 [-15.93 , 24.85]	-
Martin-Alemany 2016	33	20.5	19	25.4	30	17	59.0%	7.60 [-9.38 , 24.58]	-
Subtotal (95% CI)			32			26	100.0%	6.31 [-6.73 , 19.36]	.
Heterogeneity: $Tau^2 = 0.00$; Test for overall effect: $Z = 0$			0.82); I ² =	= 0%					ľ
10. 0.cian chect. Z = 0	(1 - 0.0	•,							
3.3.14 Quality of social int		05.0	40	44 10	40.0	_	40.607	14.40 [4.05 00.00]	
AVANTE-HEMO 2020	25.92	25.3	13	11.43	13.8	9	49.6%	14.49 [-1.95 , 30.93]	H
Martin-Alemany 2016	33	22	19	30.8	27.3	17	50.4%	2.20 [-14.12 , 18.52]	-
Subtotal (95% CI)			32			26	100.0%	8.30 [-3.74 , 20.34]	4
Heterogeneity: $Tau^2 = 5.67$; Test for overall effect: $Z = 1$			0.30); I ² =	= 8%					
3.3.15 Sexual function									
AVANTE-HEMO 2020	8.33	25	13	41.07	51.4	9	25.8%	-32.74 [-68.97 , 3.49]	
Martin-Alemany 2016	88.3	23.3	19	96	12.7	17	74.2%	-7.70 [-19.79 , 4.39]	
Subtotal (95% CI)	30.3	20.0	32	50	14./	26	100.0%	-14.16 [-35.63 , 7.31]	
	6. Chi2 = 1	65 df = 1 (P		12 = 200/		20	100.070	-14.10 [-33.03 , /.31]	
Heterogeneity: $Tau^2 = 123.6$ Test for overall effect: $Z = 1$			- 0.20);	1- – 39%					
	- , 3.2	,							
3.3.16 Sleep									



Analysis 3.3. (Continued)



(1) Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

Analysis 3.4. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 4: Depression

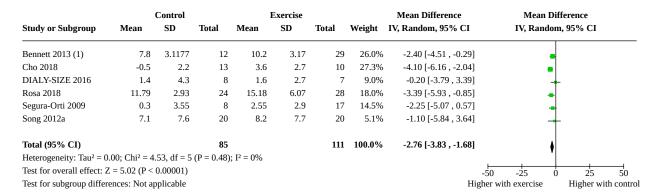
		Control		1	Exercise			Std. Mean Difference	!	Std. Me	an Difference	2
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	[IV, Ran	dom, 95% C	I
Rahimimoghadam 2017	10.4	2.4	25	8.6	3.06	25	49.7%	0.64 [0.07 , 1.2	1]			
PEAK 2006	1	2.9	25	-0.3	3.6	24	50.3%	0.39 [-0.17 , 0.9	6]		+-	
Total (95% CI)			50			49	100.0%	0.52 [0.12, 0.9	2]			
Heterogeneity: Tau ² = 0.00;	$Chi^2 = 0.38, c$	lf = 1 (P =	0.54); I ² =	0%								
Test for overall effect: $Z = 2$.53 (P = 0.01)	1							-2	-1	0 1	<u>l</u>
Test for subgroup difference	s: Not applica	ible							Higher w	ith exercise	Higher	high control



Analysis 3.5. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 5: 6MWT

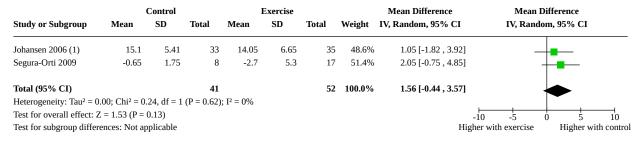
		Control		E	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [metres]	SD [metres]	Total	Mean [metres]	SD [metres]	Total	Weight	IV, Random, 95% CI [metres]	IV, Random, 95% CI [metres]
Martins do Valle 2020	494.8	66.9	12	457.3	155.6	12	3.4%	37.50 [-58.33 , 133.33]	
PEAK 2006	414.3	127.3	25	514.9	163.9	24	4.6%	-100.60 [-183.00 , -18.20]	
Rosa 2018	469.42	162.93	24	526.45	126.15	28	4.9%	-57.03 [-137.23, 23.17]	
Pellizzaro 2013	407	116.7	14	475	74.1	14	6.0%	-68.00 [-140.41 , 4.41]	
DIALY-SIZE 2016	0.8	44	8	54.9	52.9	7	12.7%	-54.10 [-103.75 , -4.45]	
Segura-Orti 2009	20.6	36.6	8	48.5	60.8	17	21.2%	-27.90 [-66.35 , 10.55]	
Cho 2018	-26	41	13	20	21	10	47.1%	-46.00 [-71.81 , -20.19]	-
Total (95% CI)			104			112	100.0%	-44.71 [-62.43 , -27.00]	•
Heterogeneity: Tau ² = 0.00); Chi ² = 5.96, df = 6	(P = 0.43); I ² = 0)%						~
Test for overall effect: Z =	4.95 (P < 0.00001)								-200 -100 0 100 200
Test for subgroup differen	ces: Not applicable								Higher in exercise Higher in control

Analysis 3.6. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 6: Sit-To-Stand test [N reps/30 sec]



(1) results from group 1 (24 weeks of intervention) and group 2 (12 weeks of intervention) were pooled together in the exercise group. The number of participants was correcte

Analysis 3.7. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 7: Sit-To-Stand test [N reps/30 sec]



Footnotes

(1) Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group



Analysis 3.8. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 8: Albumin

	(Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
AVANTE-HEMO 2020	39	4	13	38	4.4	9	7.5%	1.00 [-2.60 , 4.60]	ı .
Afshar 2010	40	3	7	40	3	7	8.6%	0.00 [-3.14 , 3.14]	ı —
Dong 2011	42.1	2.2	12	41.5	4.4	10	9.0%	0.60 [-2.40, 3.60]	ı _
Martins do Valle 2020	39	3	12	37	4	12	9.5%	2.00 [-0.83 , 4.83]	ı • • • • • • • • • • • • • • • • • •
Kopple 2007	39	3.7417	14	38	4	15	9.6%	1.00 [-1.82 , 3.82]	ı -
Martin-Alemany 2016	37	3.5	19	37	3.3	17	11.4%	0.00 [-2.22, 2.22]	ı <u>—</u>
Abreu 2017	42	2	19	43	3	25	13.9%	-1.00 [-2.48, 0.48]	ı _
PEAK 2006	-0.16	2.4	25	0.3	2.4	24	14.4%	-0.46 [-1.80, 0.88]	l <u>-</u>
Pellizzaro 2013	-2	1	14	1	1	14	16.0%	-3.00 [-3.74 , -2.26]	-
Total (95% CI)			135			133	100.0%	-0.27 [-1.59 , 1.05]	
Heterogeneity: Tau ² = 2.7	70; Chi ² = 33.66,	df = 8 (P < 0)	0.0001); I ²	? = 76%					Ť
Test for overall effect: Z	= 0.40 (P = 0.69)								-10 -5 0 5
Test for subgroup differen	nces: Not applica	ible						Н	igher with exercise Higher wi

Analysis 3.9. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 9: Blood lipids

	(Control			Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]	
3.9.1 Total cholesterol	[mmol/L]									
Afshar 2010	3.4	0.812	7	3.28	0.58	7	19.5%	0.12 [-0.62, 0.86]		
Dong 2011	4.49	1.1	12	4.36	0.64	10	19.5%	0.13 [-0.61, 0.87]		
Song 2012a	4.192	0.67	20	3.85	0.678	20	61.0%	0.34 [-0.08, 0.76]	 	
Subtotal (95% CI)			39			37	100.0%	0.26 [-0.07, 0.58]		
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.40, df =	2 (P = 0.82); I ² =	0%						•	
est for overall effect:	Z = 1.55 (P = 0.12)									
3.9.2 LDL cholesterol	[mmol/L]									
Afshar 2010	1.56	0.33	7	1.32	0.68	7	26.9%	0.24 [-0.32 , 0.80]		
Song 2012a	2.102	0.55	20	2.02	0.54564	20	73.1%	0.08 [-0.26 , 0.42]		
Subtotal (95% CI)			27			27	100.0%	0.12 [-0.17 , 0.41]	-	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.22, df =	1 (P = 0.64); I ² =	0%							
Test for overall effect:	Z = 0.84 (P = 0.40)									
3.9.3 HDL cholesterol	[mmol/L]									
Afshar 2010	0.82	0.332	7	0.83	0.26	7	30.9%	-0.01 [-0.32 , 0.30]		
Song 2012a	1.1	0.4	20	1.07	0.26	20	69.1%	0.03 [-0.18, 0.24]	•	
Subtotal (95% CI)			27			27	100.0%	0.02 [-0.16 , 0.19]	•	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.04, df =	1 (P = 0.83); I ² =	0%							
Test for overall effect:	Z = 0.20 (P = 0.84)									
3.9.4 Triglycerides [m	mol/L]									
Afshar 2010	2.55	0.903	7	1.65	0.7	7	36.3%	0.90 [0.05, 1.75]		
Song 2012a	1.6699	1.2	20	1.34	0.71469	20	63.7%	0.33 [-0.28, 0.94]	+-	
Subtotal (95% CI)			27			27	100.0%	0.54 [-0.00 , 1.07]		
Heterogeneity: Tau ² = 0	0.02; Chi ² = 1.14, df =	1 (P = 0.28); I ² =	13%							
Test for overall effect:	Z = 1.96 (P = 0.05)									
									-2 -1 0 1	
								High	her with exercise Higher with	



Analysis 3.10. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 10: Body composition

		Control]	Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
3.10.1 Fat mass [kg]										
Dong 2011	27.6	14.8	12	24.7	10.5	10	4.5%	2.90 [-7.71 , 13.51]		
Kopple 2007	19.1	2.4	14	23.1	18.2	15	5.9%	-4.00 [-13.30 , 5.30]		
Chen 2010	33.1	10.1	22	29.6	9.8	22	14.6%	3.50 [-2.38, 9.38]		
Johansen 2006 (1)	18.68	13.21	33	21.8	10.08	35	16.1%	-3.12 [-8.73 , 2.49]		
Song 2012a	27.2	8.9	20	26	8.6	20	17.2%	1.20 [-4.22, 6.62]		
Martin-Alemany 2016	17.6	6.5	19	20.3	9	17	18.8%	-2.70 [-7.88, 2.48]		
Rosa 2018	21.92	8.81	24	23.2	8.4	28	22.9%	-1.28 [-5.98, 3.42]		
Subtotal (95% CI)			144			147	100.0%	-0.69 [-2.94, 1.56]	—	
Heterogeneity: Tau ² = 0.0	0; Chi ² = 4.70	0, $df = 6$ (F	P = 0.58); I	$^{2} = 0\%$					Y	
Test for overall effect: Z =	= 0.60 (P = 0.5	55)								
3.10.2 Lean mass [kg]										
Dong 2011	56.2	9.9	12	47.4	7.1	10	13.0%	8.80 [1.68, 15.92]		
Chen 2010	46.3	8.7	21	47.9	9.9	21	17.4%	-1.60 [-7.24 , 4.04]		
Johansen 2006 (1)	48.1	8.76	33	48.47	13.01	35	18.8%	-0.37 [-5.62 , 4.88]		
Rosa 2018	44.04	8.23	24	47.55	9.49	28	20.5%	-3.51 [-8.33, 1.31]		
Song 2012a	22.5	5.2	20	22.2	3.7	20	30.4%	0.30 [-2.50, 3.10]		
Subtotal (95% CI)			110			114	100.0%	0.16 [-2.95, 3.28]	—	
Heterogeneity: $Tau^2 = 6.2$	7; Chi ² = 8.28	3, df = 4 (F)	P = 0.08); I	² = 52%					T	
Test for overall effect: Z =	= 0.10 (P = 0.9	92)								
Test for subgroup differer	nces: Chi² = 0	.19, df = 1	(P = 0.66)), $I^2 = 0\%$				Ui	-20 -10 0 10 20 gher with exercise Higher with cor	
								пі	gner with exercise — Higher with co	

 $(1) Factorial \ design, two \ intervention \ arms \ pooled \ together \ in \ the \ exercise \ group \ and \ two \ control \ arms \ pooled \ together \ in \ the \ control \ group$

Analysis 3.11. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 11: Body mass index

			E	exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [kg/m²]	SD [kg/m ²]	Total	Mean [kg/m²]	SD [kg/m ²]	Total	Weight	IV, Random, 95% CI [kg/m²]	IV, Random, 95% CI [kg/m²]	
Marinho 2016 (1)	29.13	10.655	7	28.8	13.73	6	0.5%	0.33 [-13.20 , 13.86]		
Dong 2011	29.3	6.8	12	26.8	4.3	10	3.8%	2.50 [-2.18 , 7.18]		
Abreu 2017	24.1	4.9	19	23.8	4.5	25	8.3%	0.30 [-2.52 , 3.12]		
Rosa 2018	25.51	4.03	24	26.61	4.44	28	10.8%	-1.10 [-3.40 , 1.20]	-	
Martin-Alemany 2016 (1)	21.2	1.698	17	21.3	3.204	19	15.0%	-0.10 [-1.75 , 1.55]	+	
Kopple 2007	25.1	1.2	14	27.7	2.5	15	16.9%	-2.60 [-4.01 , -1.19]	-	
AVANTE-HEMO 2020	19.8	1.7	13	22.1	1	9	19.3%	-2.30 [-3.43 , -1.17]	-	
PEAK 2006	-0.1	0.5	25	0.3	0.5	24	25.4%	-0.40 [-0.68 , -0.12]	•	
Total (95% CI)			131			136	100.0%	-1.00 [-1.98 , -0.01]	•	
Heterogeneity: Tau ² = 0.98	; Chi ² = 20.87, df	= 7 (P = 0.004);	$I^2 = 66\%$						"	
Test for overall effect: Z =	1.98 (P = 0.05)								-20 -10 0 10 2	
Test for subgroup differenc	es: Not applicable							1	Higher in exercise Higher in contr	

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range



Analysis 3.12. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 12: Calcium

	(Control		F	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
de Lima 2013	2.32	0.22	11	2.52	0.65	10	7.1%	-0.20 [-0.62 , 0.22	
Marinho 2016 (1)	2.2	0.367	7	2.2	0.19	6	12.0%	0.00 [-0.31 , 0.31	1
Martins do Valle 2020	2.4	0.15	12	2.22	0.25	12	28.8%	0.18 [0.02 , 0.34	.]
Abreu 2017	2.22	0.12	19	2.22	0.12	25	52.1%	0.00 [-0.07 , 0.07	1 🛊
Total (95% CI)			49			53	100.0%	0.04 [-0.08 , 0.16	
Heterogeneity: Tau ² = 0.0	1; Chi ² = 4.96, df = 3	$(P = 0.17); I^2 = 40$	1%						
Test for overall effect: Z =	0.62 (P = 0.54)								-1 -0.5 0 0.5 1
Test for subgroup differen	ces: Not applicable							I	Higher with exercise Higher with control

Footnotes

(1) mean and standard deviation estimated from the median and the interquartile range

Analysis 3.13. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 13: CRP

		Control		E	xercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mg/dL]	SD [mg/dL]	Total	Mean [mg/dL]	SD [mg/dL]	Total	Weight	IV, Random, 95% CI [mg/dL]	IV, Random, 95% CI [mg/dL]
Marinho 2016 (1)	6.43	11.298	7	5.93	7.91	6	0.1%	0.50 [-9.99 , 10.99]	
Abreu 2017	8.4	7.5	19	5.8	4.4	25	0.9%	2.60 [-1.19, 6.39]	ı
Pellizzaro 2013 (1)	2.67	4.86	14	2.267	4.86	14	1.0%	0.40 [-3.20 , 4.00]	ı <u>—</u>
Afshar 2010	4.14	3.87	7	2.27	1.79	7	1.3%	1.87 [-1.29, 5.03]	ı
Kopple 2007	2.8	2.9933	14	4.2	5.52	18	1.4%	-1.40 [-4.39 , 1.59]	ı <u>-</u>
AVANTE-HEMO 2020 (2)	0.38	0.349	13	0.643	0.48	9	95.3%	-0.26 [-0.63 , 0.10]	ı 💼
Total (95% CI)			74			79	100.0%	-0.22 [-0.58 , 0.14]	
Heterogeneity: Tau ² = 0.00	0; Chi ² = 4.59, df =	5 (P = 0.47); I ²	= 0%						1
Test for overall effect: Z =	1.20 (P = 0.23)								-20 -10 0 10 20
Test for subgroup difference	ces: Not applicable	2						Н	igher with exercise Higher with contro

Footnotes

- (1) mean and standard deviation estimated from the median and the interquartile range
- (2) Mean and standard deviation estimated from the median and the interquartile range

Analysis 3.14. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 14: Dialysis adequacy: Kt/V

		Control]	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
PEAK 2006	0.1	0.4	25	0.1	0.5	24	47.1%	0.00 [-0.25 , 0.25]	
Martins do Valle 2020	1.6	0.3	12	1.7	0.3	12	52.9%	-0.10 [-0.34 , 0.14]	
Total (95% CI)			37			36	100.0%	-0.05 [-0.23 , 0.12]	
Heterogeneity: Tau ² = 0.0	00; $Chi^2 = 0.31$	l, df = 1 (P	e = 0.58); I	$^{2} = 0\%$					
Test for overall effect: Z =	= 0.59 (P = 0.5)	55)						-(0.5 -0.25 0 0.25 0.5
Test for subgroup differen	nces: Not appl	icable						High	er with exercise Higher with control



Analysis 3.15. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 15: Energy intake

	Control			Exercise				Mean Difference	Mean Difference		
Study or Subgroup	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Weight	IV, Random, 95% CI [kCal/kg/d]	IV, Random, 95% CI [kCal/kg/d]		
PEAK 2006	30.05	8.57	25	41.44	37.04	24	1.1%	-11.39 [-26.58 , 3.80]			
Kopple 2007	24.2	7.8575	14	26.9	25.46	18	1.7%	-2.70 [-15.16, 9.76]			
Dong 2011	27.6	11.9	12	26.5	7.1	10	4.1%	1.10 [-6.94, 9.14]			
Olvera-Soto 2016 (1)	23.77	13	31	23.17	9.65	30	8.0%	0.60 [-5.13, 6.33]	_		
Abreu 2017	30.9	3	19	30.6	2.9	25	85.0%	0.30 [-1.46 , 2.06]	•		
Total (95% CI)			101			107	100.0%	0.17 [-1.45 , 1.80]			
Heterogeneity: Tau ² = 0	0.00; Chi ² = 2.52, df = 4	(P = 0.64); I ² = 0%							Ĭ		
Test for overall effect: Z	Z = 0.21 (P = 0.84)								-50 -25 0 25 50		
Test for subgroup differ	ences: Not applicable							Hig	ther with exercise Higher with control		

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range

Analysis 3.16. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 16: Haemoglobin

		Control		E	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
Marinho 2016 (1)	11.2	2.756	7	10.467	2.098	6	0.5%	0.73 [-1.91 , 3.38	
Dong 2011	12.2	1.8	12	11.4	1.9	10	1.4%	0.80 [-0.76 , 2.36	i) ——
AVANTE-HEMO 2020	10.6	1.8	13	11.7	1.7	9	1.5%	-1.10 [-2.58 , 0.38	i]
Martin-Alemany 2016 (1)	8.6	2.102	17	8.9	1.842	19	1.9%	-0.30 [-1.60 , 1.00]
Abreu 2017	10.6	2.3	19	10.5	1.9	25	2.0%	0.10 [-1.17 , 1.37	1
Martins do Valle 2020	10.3	1.3	12	10.8	1.7	12	2.2%	-0.50 [-1.71, 0.71]
Kopple 2007	12.5	1.87	14	12.5	0.77	15	3.0%	0.00 [-1.05 , 1.05	i]
de Lima 2013	11.1	1.2	11	11.4	0.9	11	4.2%	-0.30 [-1.19, 0.59	·]
Pellizzaro 2013	0.5	0.4	14	0.6	0.4	14	37.4%	-0.10 [-0.40 , 0.20	ı) <u>+</u>
Afshar 2010	10.2	0.3	7	10.3	0.2	7	46.0%	-0.10 [-0.37 , 0.17	n 📥
Total (95% CI)			126			128	100.0%	-0.11 [-0.29 , 0.07	n 👃
Heterogeneity: Tau ² = 0.00	; Chi ² = 4.23, c	f = 9 (P = 0.	90); I ² = 0	1%					T
Test for overall effect: Z =	1.22 (P = 0.22))							-4 -2 0 2 4
Test for subgroup differenc	es: Not applica	able						H	Higher with exercise Higher with cor

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range



Analysis 3.17. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 17: Muscular strength

		Control		Exercise				Mean Difference	Mean Difference	
Study or Subgroup	Mean [kg]	SD [kg]	Total	Mean [kg]	SD [kg]	Total	Weight	IV, Random, 95% CI [kg]	IV, Random, 95% CI [kg]	
3.17.1 knee extension										
Dong 2011	239.04	139	12	263.99	66.69	10	0.3%	-24.95 [-113.80, 63.90]	·	
Song 2012a	33.4	19.5	20	37.3	19	20	10.4%	-3.90 [-15.83, 8.03]	l —	
PEAK 2006	-2.4	13.8	25	15.2	15.4	24	16.5%	-17.60 [-25.80 , -9.40]	-	
DIALY-SIZE 2016	9.3	10.1	8	8.9	5.5	7	16.8%	0.40 [-7.70, 8.50]	-	
Johansen 2006 (1)	17.28	8.63	33	23.65	10.18	35	26.3%	-6.37 [-10.85 , -1.89]		
Chen 2010	12.1	6.1	22	15.8	5	22	29.7%	-3.70 [-7.00 , -0.40]	ı	
Subtotal (95% CI)			120			118	100.0%	-6.09 [-10.68 , -1.50]	· •	
Heterogeneity: Tau ² = 15.6	62; Chi ² = 11.9	1, df = 5 (P	= 0.04); I ²	2 = 58%					1	
Test for overall effect: Z =	2.60 (P = 0.00	9)								
3.17.2 handgrip										
Martin-Alemany 2016 (2)	20.867	9.701	17	23.5	10.813	19	30.5%	-2.63 [-9.33 , 4.07]	ı 📥	
Song 2012a	27.8	11.8	20	28.7	9	20	32.4%	-0.90 [-7.40, 5.60]		
Olvera-Soto 2016 (2)	19.6	8.56	31	22.07	14.77	30	37.0%	-2.47 [-8.55, 3.61]	ı 📥	
Subtotal (95% CI)			68			69	100.0%	-2.01 [-5.71 , 1.69]	. ♦	
Heterogeneity: Tau ² = 0.00	0; $Chi^2 = 0.17$,	df = 2 (P =	0.92); I ² =	0%					Ĭ	
Test for overall effect: Z =	1.06 (P = 0.29)								
									-100 -50 0 50	
Footnotes								Н	igher with exercise Higher with	

⁽¹⁾ Factorial design, two intervention arms pooled together in the exercise group and two control arms pooled together in the control group

Analysis 3.18. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 18: Phosphate

	(Control		I	Exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]	
Marinho 2016 (1)	1.367	1.653	7	1.467	0.57	6	4.3%	-0.10 [-1.41 , 1.21]		
AVANTE-HEMO 2020 (2)	1.86	1.15	13	2.34	0.76	9	8.7%	-0.48 [-1.28, 0.32]		
Martins do Valle 2020	1.84	0.39	12	1.61	0.97	12	12.1%	0.23 [-0.36, 0.82]		
Martin-Alemany 2016	1.84	0.61	19	2.07	0.65	17	15.9%	-0.23 [-0.64, 0.18]		
de Lima 2013	1.81	0.29	11	2.13	0.52	10	17.0%	-0.32 [-0.69, 0.05]		
Abreu 2017	2.1	0.42	19	2.2	0.48	25	19.3%	-0.10 [-0.37 , 0.17]		
Pellizzaro 2013	0.19	0.06	14	-0.13	0.06	14	22.6%	0.32 [0.28, 0.36]	•	
Total (95% CI)			95			93	100.0%	-0.06 [-0.36 , 0.24]		
Heterogeneity: Tau ² = 0.10;	Chi ² = 30.74, df = 6	6 (P < 0.0001); I ²	= 80%						Ť	
Test for overall effect: Z = 0	0.37 (P = 0.71)								-2 -1 0 1	
Test for subgroup difference	es: Not applicable							High	her with exercise Higher with	

Footnotes

- (1) mean and standard deviation estimated from the median and interquartile range
- (2) mean and standard deviation estimated from the median and the interquartile range

⁽²⁾ mean and standard deviation estimated from the median and interquartile range



Analysis 3.19. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 19: Potassium

		Control		I	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Marinho 2016 (1)	4.5	2.848	7	5.067	1.62	6	3.8%	-0.57 [-3.04 , 1.91]	
AVANTE-HEMO 2020	5.2	0.92	13	5.6	0.58	9	14.6%	-0.40 [-1.03, 0.23]	-
Martin-Alemany 2016 (1)	5.93	1.213	17	5.1	0.481	19	14.7%	0.83 [0.21, 1.45]	
e Lima 2013	5.8	0.6	11	5.6	0.7	10	15.2%	0.20 [-0.36, 0.76]	-
fartins do Valle 2020	5.3	0.4	12		0.6	12	16.4%	0.30 [-0.11, 0.71]	 -
breu 2017	4.7	0.3	19	4.7	0.4	25	17.6%	0.00 [-0.21, 0.21]	.
ellizzaro 2013	0.6	0.2	14	-0.5	0.2	14	17.8%	1.10 [0.95 , 1.25]	•
otal (95% CI)			93			95	100.0%	0.32 [-0.23, 0.86]	
Heterogeneity: Tau ² = 0.43	; Chi ² = 90.68, df = 6	6 (P < 0.00001); I ²	= 93%						
est for overall effect: Z =	1.14 (P = 0.25)								-4 -2 0 2
est for subgroup difference	oc. Not applicable							Hial	her with evercise Higher with c

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range

Analysis 3.20. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 20: Protein intake

		Control		F	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [g/kg/d]	SD [g/kg/d]	Total	Mean [g/kg/d]	SD [g/kg/d]	Total	Weight	IV, Random, 95% CI [g/kg/d]	IV, Random, 95% CI [g/kg/d]
Kopple 2007	1.35	0.5987	14	1.08	0.93	18	2.3%	0.27 [-0.26 , 0.80]	
Dong 2011	1.1	0.4	12	1	0.3	10	7.4%	0.10 [-0.19, 0.39]	
Olvera-Soto 2016 (1)	1.03	0.4	31	1.01	0.62	30	9.2%	0.02 [-0.24 , 0.28]	
PEAK 2006	1.36	0.38	25	1.51	0.25	24	19.6%	-0.15 [-0.33, 0.03]	
Abreu 2017	1.2	0.14	19	1.2	0.2	25	61.5%	0.00 [-0.10 , 0.10]	•
Total (95% CI)			101			107	100.0%	-0.01 [-0.09 , 0.07]	•
Heterogeneity: Tau ² = 0	.00; Chi ² = 4.02, df	= 4 (P = 0.40); I	2 = 1%						Ţ
Test for overall effect: Z	L = 0.35 (P = 0.73)								-1 -0.5 0 0.5 1
Test for subgroup differ	ences: Not applicab	le						High	ner with exercise Higher with con-

Footnotes

(1) mean and standard deviation estimated from the median and interquartile range

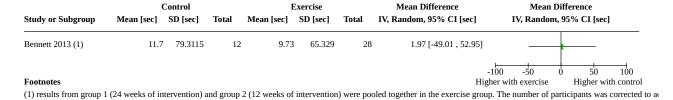
Analysis 3.21. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 21: PTH

		Control		I	Exercise			Mean Difference	Mean Dif	ference	
Study or Subgroup	Mean [pmol/L]	SD [pmol/L]	Total	Mean [pmol/L]	SD [pmol/L]	Total	Weight	IV, Random, 95% CI [pmol/L]	IV, Random, 95	% CI [pmol	L]
Martins do Valle 2020	56.28	37.28	12	80.64	84.74	12	28.3%	-24.36 [-76.74 , 28.02]	-		
Marinho 2016 (1)	28.11	28.06	7	16.37	14.93	6	71.7%	11.74 [-12.24 , 35.72]	•	I	
Total (95% CI)			19			18	100.0%	1.51 [-30.38 , 33.39]	•	•	
Heterogeneity: Tau ² = 219	9.68; Chi ² = 1.51, df =	= 1 (P = 0.22); I ²	= 34%						Ĭ		
Test for overall effect: Z =	= 0.09 (P = 0.93)								-500 -250 0	250	500
Test for subgroup differer	nces: Not applicable							Hi	igher with exercise	Higher wi	th control

Footnote

(1) mean and standard deviation estimated from the median and interquartile range

Analysis 3.22. Comparison 3: Resistance exercise versus control (no exercise/placebo exercise), Outcome 22: Timed up-and-go test





Comparison 4. Combined aerobic and resistance exercise versus control (no exercise/placebo exercise)

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
4.1 HRQoL: Summary component scores	6		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.1.1 Physical Component Score	6	228	Mean Difference (IV, Random, 95% CI)	-4.38 [-6.82, -1.94]
4.1.2 Mental Component Score	6	228	Mean Difference (IV, Random, 95% CI)	-2.58 [-6.91, 1.74]
4.2 HRQoL: Individual domains	3		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.2.1 Physical Functioning	3	161	Mean Difference (IV, Random, 95% CI)	-4.07 [-10.60, 2.47]
4.2.2 Role-physical	3	160	Mean Difference (IV, Random, 95% CI)	-3.86 [-14.38, 6.66]
4.2.3 Pain	3	161	Mean Difference (IV, Random, 95% CI)	-3.98 [-10.46, 2.49]
4.2.4 General health perceptions	3	161	Mean Difference (IV, Random, 95% CI)	-3.67 [-9.24, 1.90]
4.2.5 Emotional well-being	3	161	Mean Difference (IV, Random, 95% CI)	1.29 [-3.99, 6.57]
4.2.6 Role-emotional	3	160	Mean Difference (IV, Random, 95% CI)	-10.68 [-20.92, -0.43]
4.2.7 Vitality	3	161	Mean Difference (IV, Random, 95% CI)	-7.88 [-13.48, -2.28]
4.2.8 Social function	3	161	Mean Difference (IV, Random, 95% CI)	1.83 [-4.56, 8.22]
4.2.9 Symptoms	2	143	Mean Difference (IV, Random, 95% CI)	0.17 [-3.20, 3.54]
4.2.10 Effects of kidney disease	1	47	Mean Difference (IV, Random, 95% CI)	-1.70 [-10.27, 6.87]
4.2.11 Burden of kidney disease	1	47	Mean Difference (IV, Random, 95% CI)	-5.70 [-17.40, 6.00]
4.2.12 Cognitive function	1	47	Mean Difference (IV, Random, 95% CI)	2.10 [-3.68, 7.88]
4.2.13 Quality of social interactions	1	47	Mean Difference (IV, Random, 95% CI)	-0.30 [-7.72, 7.12]
4.2.14 Sleep	1	47	Mean Difference (IV, Random, 95% CI)	4.30 [-5.67, 14.27]
4.2.15 Social support	1	47	Mean Difference (IV, Random, 95% CI)	0.30 [-9.88, 10.48]
4.2.16 Dialysis staff encouragement	1	47	Mean Difference (IV, Random, 95% CI)	2.40 [-8.82, 13.62]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
4.2.17 Patient satisfaction	1	47	Mean Difference (IV, Random, 95% CI)	2.80 [-8.88, 14.48]
4.3 Depression	4	214	Std. Mean Difference (IV, Random, 95% CI)	0.97 [0.25, 1.68]
4.4 6MWT	6	138	Mean Difference (IV, Random, 95% CI)	-53.64 [-67.91, -39.36]
4.5 Sit-To-Stand test [N reps/30 sec]	4	97	Mean Difference (IV, Random, 95% CI)	-2.63 [-3.77, -1.49]
4.6 Sit-To-Stand test [sit to 5 reps]	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
4.7 Resting blood pressure	7		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.7.1 Systolic blood pressure	7	288	Mean Difference (IV, Random, 95% CI)	8.69 [3.78, 13.59]
4.7.2 Diastolic blood pressure	7	288	Mean Difference (IV, Random, 95% CI)	4.42 [2.90, 5.94]
4.8 Aerobic capacity (VO2 max or peak)	3	93	Mean Difference (IV, Random, 95% CI)	-4.29 [-8.98, 0.39]
4.9 Albumin	3	116	Mean Difference (IV, Random, 95% CI)	-0.22 [-1.61, 1.16]
4.10 Blood lipids	4		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.10.1 Total cholesterol [mmol/L]	4	204	Mean Difference (IV, Random, 95% CI)	0.03 [-0.21, 0.27]
4.10.2 LDL cholesterol [mmol/L]	2	61	Mean Difference (IV, Random, 95% CI)	0.44 [-0.09, 0.96]
4.10.3 HDL cholesterol [mmol/L]	3	108	Mean Difference (IV, Random, 95% CI)	-0.27 [-0.44, -0.10]
4.10.4 Triglycerides [mmol/L]	3	108	Mean Difference (IV, Random, 95% CI)	-0.11 [-0.86, 0.64]
4.11 Body composition	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
4.11.1 Fat mass [kg]	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
4.12 Body mass index	2	73	Mean Difference (IV, Fixed, 95% CI)	-0.42 [-1.37, 0.53]
4.13 Calcium	4	190	Mean Difference (IV, Random, 95% CI)	0.06 [-0.01, 0.12]
4.14 CRP	3	117	Mean Difference (IV, Random, 95% CI)	0.09 [-0.27, 0.46]
4.15 Dialysis adequacy: Kt/V	1		Mean Difference (IV, Random, 95% CI)	Totals not selected



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
4.16 Energy intake	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
4.17 Haemoglobin	5	266	Mean Difference (IV, Random, 95% CI)	-0.02 [-0.24, 0.20]
4.18 Heart rate	5		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.18.1 Resting	4	179	Mean Difference (IV, Random, 95% CI)	3.05 [0.70, 5.40]
4.18.2 Maximum	3	90	Mean Difference (IV, Random, 95% CI)	-5.37 [-11.10, 0.35]
4.19 Muscular strength	3		Mean Difference (IV, Random, 95% CI)	Subtotals only
4.19.1 Knee extension	2	63	Mean Difference (IV, Random, 95% CI)	1.60 [-3.66, 6.87]
4.19.2 Handgrip	2	88	Mean Difference (IV, Random, 95% CI)	-4.55 [-10.23, 1.14]
4.20 Phosphate	4	190	Mean Difference (IV, Random, 95% CI)	-0.04 [-0.15, 0.08]
4.21 Potassium	4	190	Mean Difference (IV, Random, 95% CI)	-0.15 [-0.36, 0.06]
4.22 Protein intake	1		Mean Difference (IV, Random, 95% CI)	Totals not selected
4.23 Timed up-and-go test	1		Mean Difference (IV, Random, 95% CI)	Totals not selected



Analysis 4.1. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 1: HRQoL: Summary component scores

		Control		1	Exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
4.1.1 Physical Compo	nent Score								
Molsted 2004 (1)	44.95	10.99	7	45.75	7.44	10	6.1%	-0.80 [-10.16, 8.56]	
DIALY-SIZE 2016	3.4	7.3	8	1.7	7.4	8	9.7%	1.70 [-5.50 , 8.90]	
Uchiyama 2019	38.2	9.2	23	41	8.1	24	17.3%	-2.80 [-7.76 , 2.16]	
Suhardjono 2019 (2)	-3.45	12.45	38	4.97	7.56	36	18.9%	-8.42 [-13.09, -3.75]	
Ouzouni 2009	38.9	5.8	14	44.5	5.5	19	23.8%	-5.60 [-9.52 , -1.68]	
Frih 2017a	51	7	20	55.5	5.5	21	24.2%	-4.50 [-8.37, -0.63]	
Subtotal (95% CI)			110			118	100.0%	-4.38 [-6.82 , -1.94]	•
Heterogeneity: Tau ² = 2	2.51; Chi ² = 6.	91, df = 5	(P = 0.23)	$I^2 = 28\%$					•
Test for overall effect: 2	Z = 3.52 (P =	0.0004)							
4.1.2 Mental Compone	ent Score								
Molsted 2004 (1)	51.55	10.26	7	54.1	6.47	10	12.1%	-2.55 [-11.14 , 6.04]	
DIALY-SIZE 2016	0.7	7.5	8	-1.5	5.9	8	15.0%	2.20 [-4.41 , 8.81]	
Ouzouni 2009	40.1	6.8	14	41.8	10	19	16.4%	-1.70 [-7.44 , 4.04]	
Uchiyama 2019	52.6	9	23	49.8	9.6	24	17.1%	2.80 [-2.52 , 8.12]	
Suhardjono 2019	-2.85	12	38	2.673	9.05	36	18.0%	-5.52 [-10.35 , -0.70]	
Frih 2017a	42.5	4.5	20	51	4	21	21.3%	-8.50 [-11.11 , -5.89]	
Subtotal (95% CI)			110			118	100.0%	-2.58 [-6.91 , 1.74]	
Heterogeneity: Tau ² = 2	21.06; Chi ² = 2	21.33, df =	5 (P = 0.0	007); I ² = 7	77%				
Test for overall effect: 2	Z = 1.17 (P =	0.24)							
									-20 -10 0 10 20
Footnotes								Hig	gher with exercise Higher with cont

- (1) mean and standard deviation estimated from the median and the range
- (2) mean and standard deviation estimated from the median and range



Analysis 4.2. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 2: HRQoL: Individual domains

Study or Subgroup	Mean	Control SD	Total	Mean	Exercise SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
4.2.1 Physical Functioni	ng								
Molsted 2004 (1)	70	21.99	7	82.5	9.41	11	14.4%	-12.50 [-29.71 , 4.71]	
van Vilsteren 2005	60.2	34.5	43	62.5	28	53	26.1%	-2.30 [-15.07 , 10.47]	
Uchiyama 2019	73.2	13.9	23	76	15.7	24	59.5%	-2.80 [-11.27 , 5.67]	
=	75.2	15.5	73	70	13.7	88	100.0%		
Subtotal (95% CI)	0. Ch:2 - 1	00 46 - 2		T2 - 00/		00	100.0 76	-4.07 [-10.60 , 2.47]	
Heterogeneity: Tau² = 0.0 Test for overall effect: Z =			(P = 0.58);	12 = 0%					
4.2.2 Role-physical									
Molsted 2004 (1)	62.5	36.64	7	62.5	32.33	10	9.7%	0.00 [-33.74, 33.74]	
van Vilsteren 2005	54.5	45.7	43	50	43	53	34.5%	4.50 [-13.41 , 22.41]	
Uchiyama 2019	62.2	26.9	23	71.9	22	24	55.8%	-9.70 [-23.78 , 4.38]	_
Subtotal (95% CI)	02.2	20.5	73	71.3		87	100.0%	-3.86 [-14.38 , 6.66]	
	10. Ch:2 = 1	EE 46 - 3		12 - 00/		67	100.0 76	-3.00 [-14.30 , 0.00]	
Heterogeneity: Tau ² = 0.0 Test for overall effect: Z =			(P – 0.40),	1 - 070					
4.2.3 Pain									
Uchiyama 2019	67.5	24.4	23	73	19.1	24	26.6%	-5.50 [-18.06 , 7.06]	
Molsted 2004 (1)	82.5	13.92	7	90.5	11.93	11	26.9%	-8.00 [-20.49 , 4.49]	
van Vilsteren 2005	76.1	25.5	43	76.9	21	53	46.6%	-0.80 [-10.29, 8.69]	
Subtotal (95% CI)			73			88	100.0%	-3.98 [-10.46 , 2.49]	<u> </u>
Heterogeneity: $Tau^2 = 0.0$	0; Chi ² = 0	.89, df = 2		$I^2 = 0\%$,	
Test for overall effect: Z =			, J.O.1),	0,3					
4.2.4 General health per	_								
Molsted 2004 (1)	59	30.05	7	58.5	25.11	11	4.3%	0.50 [-26.25, 27.25]	
Uchiyama 2019	45.7	17.4	23	43.7	17.9	24	30.5%	2.00 [-8.09 , 12.09]	—
Ochryania 2013									i i
•	45.2	18.1	43	51.8	15.9	53	65.2%	-6.60 [-13.50 , 0.30]	
van Vilsteren 2005 Subtotal (95% CI)		18.1	43 73	51.8	15.9	53 88	65.2% 100.0%	-6.60 [-13.50 , 0.30] - 3.67 [-9.24 , 1.90]	
van Vilsteren 2005 Subtotal (95% CI)	45.2		73		15.9				•
van Vilsteren 2005	45.2 00; Chi² = 2	.00, df = 2	73		15.9				•
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau ² = 0.0 Test for overall effect: Z =	45.2 00; Chi² = 2 = 1.29 (P =	.00, df = 2	73		15.9				•
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0	45.2 00; Chi² = 2 = 1.29 (P =	.00, df = 2	73						•
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1)	45.2 00; Chi ² = 2 = 1.29 (P = ing	0.20) df = 2	73 (P = 0.37);	$I^2 = 0\%$		88	100.0%	-3.67 [-9.24 , 1.90]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019	45.2 00; Chi² = 2 = 1.29 (P = ing	0.20) df = 2 0.20)	73 (P = 0.37); 7 23	I ² = 0%	10.04 18.8	11 24	13.6% 25.8%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005	45.2 00; Chi² = 2 = 1.29 (P = ing 76 73.2	.00, df = 2 0.20) 17.59 17.6	73 (P = 0.37); 7 23 43	I ² = 0% 84 71.5	10.04 18.8	11 24 53	13.6% 25.8% 60.6%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98]	+
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI)	45.2 90; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4	17.59 17.6 15	73 (P = 0.37); 7 23 43 73	84 71.5 76.2	10.04 18.8	11 24	13.6% 25.8% 60.6%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0	45.2 90; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4 90; Chi ² = 1	17.59 17.6 15	73 (P = 0.37); 7 23 43 73	84 71.5 76.2	10.04 18.8	11 24 53	13.6% 25.8% 60.6%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005	45.2 90; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4 90; Chi ² = 1	17.59 17.6 15	73 (P = 0.37); 7 23 43 73	84 71.5 76.2	10.04 18.8	11 24 53	13.6% 25.8% 60.6%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional	45.2 90; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4 90; Chi ² = 1	17.59 17.6 15	73 (P = 0.37); 7 23 43 73	84 71.5 76.2	10.04 18.8	11 24 53	13.6% 25.8% 60.6%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1)	45.2 10; Chi ² = 2 = 1.29 (P = 1.	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63)	73 (P = 0.37); 7 23 43 73 (P = 0.38);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$	10.04 18.8 18.9	11 24 53 88	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005	45.2 10; Chi ² = 2 = 1.29 (P = 1.	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63)	73 (P = 0.37); 7 23 43 73 (P = 0.38);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33	10.04 18.8 18.9	11 24 53 88	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07]	*
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8	10.04 18.8 18.9 21.55 35	11 24 53 88	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94]	*
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005	45.2 10; Chi ² = 2 1.29 (P = 1.29 (.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5	10.04 18.8 18.9 21.55 35	11 24 53 88 10 53 24	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z =	45.2 10; Chi ² = 2 1.29 (P = 1.29 (.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5	10.04 18.8 18.9 21.55 35	11 24 53 88 10 53 24	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z =	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0%	-3.67 [-9.24, 1.90] -8.00 [-22.32, 6.32] 1.70 [-8.71, 12.11] 3.20 [-3.58, 9.98] 1.29 [-3.99, 6.57] -8.33 [-38.58, 21.92] -8.60 [-24.27, 7.07] -13.20 [-28.34, 1.94] -10.68 [-20.92, -0.43]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) Van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1)	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24, 1.90] -8.00 [-22.32, 6.32] 1.70 [-8.71, 12.11] 3.20 [-3.58, 9.98] 1.29 [-3.99, 6.57] -8.33 [-38.58, 21.92] -8.60 [-24.27, 7.07] -13.20 [-28.34, 1.94] -10.68 [-20.92, -0.43]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) Van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43]	** ** ** ** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43]	** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005	45.2 10; Chi ² = 2 = 1.29 (P = 1.29	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43]	*** *** *** *** *** ** ** ** *
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0	45.2 10; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4 10; Chi ² = 1 = 0.48 (P = 75 70.2 64.3 10; Chi ² = 0 = 2.04 (P = 69 54.8 56.1 10; Chi ² = 1	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37]	** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z =	45.2 10; Chi ² = 2 = 1.29 (P = ing 76 73.2 79.4 10; Chi ² = 1 = 0.48 (P = 75 70.2 64.3 10; Chi ² = 0 = 2.04 (P = 69 54.8 56.1 10; Chi ² = 1	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37]	** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z =	45.2 10; Chi ² = 2 = 1.29 (P = ing) 76 73.2 79.4 10; Chi ² = 1 = 0.48 (P = 75 70.2 64.3 10; Chi ² = 0 = 2.04 (P = 69 54.8 56.1 10; Chi ² = 1 = 2.76 (P = 75)	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4 .37, df = 2	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73 (P = 0.50);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1 $I^{2} = 0\%$	10.04 18.8 18.9 21.55 35 19.4	11 24 53 88 10 53 24 87 11 24 53 88	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37] -7.88 [-13.48 , -2.28]	** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.8 Social function Uchiyama 2019	45.2 10; Chi ² = 2 1.29 (P = ing) 76 73.2 79.4 10; Chi ² = 1 10.48 (P = 75 70.2 64.3 10; Chi ² = 0	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4 .37, df = 2	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73 (P = 0.50);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1 $I^{2} = 0\%$ 71.8	10.04 18.8 18.9 21.55 35 19.4 17.26 20.3 15.3	111 244 533 88 110 24 87 111 244 533 88	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37] -7.88 [-13.48 , -2.28]	** ** ** ** ** ** ** ** ** **
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.8 Social function Uchiyama 2019 Molsted 2004 (1)	45.2 10; Chi ² = 2 1.29 (P = 1.29 (.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4 .37, df = 2 0.006)	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73 (P = 0.50);	$I^{2} = 0\%$ 844 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1 $I^{2} = 0\%$ 71.8 90.63	10.04 18.8 18.9 21.55 35 19.4 17.26 20.3 15.3	11 24 53 88 10 53 24 87 11 24 53 88	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37] -7.88 [-13.48 , -2.28]	
van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.5 Emotional well-bei Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.6 Role-emotional Molsted 2004 (1) van Vilsteren 2005 Uchiyama 2019 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.7 Vitality Molsted 2004 (1) Uchiyama 2019 van Vilsteren 2005 Subtotal (95% CI) Heterogeneity: Tau² = 0.0 Test for overall effect: Z = 4.2.8 Social function Uchiyama 2019	45.2 10; Chi ² = 2 1.29 (P = ing) 76 73.2 79.4 10; Chi ² = 1 10.48 (P = 75 70.2 64.3 10; Chi ² = 0	.00, df = 2 0.20) 17.59 17.6 15 .93, df = 2 0.63) 36.64 41.9 31.8 .20, df = 2 0.04) 29.32 20.3 17.4 .37, df = 2	73 (P = 0.37); 7 23 43 73 (P = 0.38); 7 43 23 73 (P = 0.91); 7 23 43 73 (P = 0.50);	$I^{2} = 0\%$ 84 71.5 76.2 $I^{2} = 0\%$ 83.33 78.8 77.5 $I^{2} = 0\%$ 71.25 57.5 66.1 $I^{2} = 0\%$ 71.8	10.04 18.8 18.9 21.55 35 19.4 17.26 20.3 15.3	111 244 533 88 110 24 87 111 244 533 88	13.6% 25.8% 60.6% 100.0% 11.5% 42.7% 45.8% 100.0%	-3.67 [-9.24 , 1.90] -8.00 [-22.32 , 6.32] 1.70 [-8.71 , 12.11] 3.20 [-3.58 , 9.98] 1.29 [-3.99 , 6.57] -8.33 [-38.58 , 21.92] -8.60 [-24.27 , 7.07] -13.20 [-28.34 , 1.94] -10.68 [-20.92 , -0.43] -2.25 [-26.25 , 21.75] -2.70 [-14.31 , 8.91] -10.00 [-16.63 , -3.37] -7.88 [-13.48 , -2.28]	



Analysis 4.2. (Continued)

van Vilsteren 2005	74.1	25	43	71.6	19	53	49.8%	2.50 [-6.56 , 11.56]	_
Subtotal (95% CI)	Ch:2 - 0.1	2 4f = 2 (D	73 - 0.04), I	2 – 00/		88	100.0%	1.83 [-4.56 , 8.22]	•
Heterogeneity: Tau² = 0.00; Fest for overall effect: Z = 0			- 0.94); 1	0%					
rest for overall effect. Z = c	7.50 (F = 0.	.37)							
1.2.9 Symptoms									
Jchiyama 2019	78.7	15.2	23	79.5	11.4	24	19.1%	-0.80 [-8.51 , 6.91]	
van Vilsteren 2005	23.9	9.5	43	23.5	9.1	53	80.9%	0.40 [-3.35 , 4.15]	•
Subtotal (95% CI)			66			77	100.0%	0.17 [-3.20, 3.54]	→
Heterogeneity: $Tau^2 = 0.00$;	$Chi^2 = 0.0$	8, df = 1 (P	= 0.78); I	$^{2} = 0\%$					Ĭ
Test for overall effect: $Z = 0$	0.10 (P = 0.	.92)							
1.2.10 Effects of kidney dis	sease								
Jchiyama 2019	78.1	15.6	23	79.8	14.3	24	100.0%	-1.70 [-10.27 , 6.87]	_
Subtotal (95% CI)			23				100.0%	-1.70 [-10.27, 6.87]	
Heterogeneity: Not applicab	ole							(, ,	
Test for overall effect: $Z = 0$		70)							
1044 D. J. (111 J.									
1.2.11 Burden of kidney d i Jchiyama 2019	sease 42.4	19.2	23	48.1	21.7	24	100.0%	-5.70 [-17.40 , 6.00]	
Subtotal (95% CI)			23	. 311			100.0%	-5.70 [-17.40 , 6.00]	
Heterogeneity: Not applicat	ole							[-//// , 5////]	
Test for overall effect: $Z = 0$.34)							
1.2.12 Cognitive function									
•	02.4	0.5	22	00.2	10.7	24	100.00/	2 10 [2 00 7 00]	
Jchiyama 2019	92.4	9.5	23	90.3	10.7	24	100.0%	2.10 [-3.68 , 7.88]	The state of the s
Subtotal (95% CI)	.l.		23			24	100.0%	2.10 [-3.68 , 7.88]	•
Heterogeneity: Not applicat		40)							
Test for overall effect: $Z = 0$	0.71 (P = 0.	.48)							
1.2.13 Quality of social int	eractions								
Jchiyama 2019	88.1	14.9	23	88.4	10.6	24	100.0%	-0.30 [-7.72 , 7.12]	-
Subtotal (95% CI)			23			24	100.0%	-0.30 [-7.72 , 7.12]	•
Heterogeneity: Not applicat	ole								Ţ
Test for overall effect: $Z = 0$	0.08 (P = 0.00)	.94)							
1.2.14 Sleep									
Jchiyama 2019	60.9	18.1	23	56.6	16.7	24	100.0%	4.30 [-5.67 , 14.27]	
Subtotal (95% CI)			23				100.0%	4.30 [-5.67 , 14.27]	
Heterogeneity: Not applicab	ole								
Test for overall effect: $Z = 0$		40)							
12.15 Social cupport									
1.2.15 Social support Jchiyama 2019	81	16.9	23	80.7	18.7	24	100.0%	0.30 [-9.88 , 10.48]	
Subtotal (95% CI)	01	10.5	23	30.7	10.7		100.0%	0.30 [-9.88, 10.48]	
Heterogeneity: Not applicat	ماه		23			2-4	100.0 /0	0.50 [-5.00 ; 10.40]	
Fest for overall effect: $Z = 0$.95)							
1.2.16 Dialysis staff encour	-	17.4	22	00.0	24 5	2.4	100.007	2.40 [0.02 42.02]	<u> </u>
Jchiyama 2019	83	17.4	23	80.6	21.7	24	100.0%	2.40 [-8.82 , 13.62]	—
Subtotal (95% CI)	1.		23			24	100.0%	2.40 [-8.82 , 13.62]	—
Heterogeneity: Not applicat Fest for overall effect: Z = 0		.68)							
	•	•							
1.2.17 Patient satisfaction	=-	00.5	0.5		4= 5		400.05	0.00 5.000	\perp
Jchiyama 2019	78	22.8	23	75.2	17.6	24	100.0%	2.80 [-8.88 , 14.48]	—
Subtotal (95% CI)			23			24	100.0%	2.80 [-8.88 , 14.48]	•
Heterogeneity: Not applicat		C (1)							
Test for overall effect: $Z = 0$	0.47 (P = 0.	.64)							
									-50 -25 0 25 5

(1) mean and standard deviation estimated from the median and the range



Analysis 4.3. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 3: Depression

		Control			Exercise			Std. Mean Difference	Std. Mea	n Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Rand	om, 95% CI	
Ouzouni 2009	19.4	4	14	11.7	3.6	19	21.6%	1.99 [1.13 , 2.8	5]		
Kouidi 2010	22.1	6.24	20	14.61	4.15	24	24.6%	1.41 [0.74, 2.0	8]		
Frih 2017a	13	25.6402	20	8.5	14.2796	21	25.4%	0.21 [-0.40, 0.8	3] .	_	
van Vilsteren 2005	41.4	9.6	43	37.2	8.3	53	28.4%	0.47 [0.06, 0.8	8]	-	
Total (95% CI)			97			117	100.0%	0.97 [0.25 , 1.6	8]		
Heterogeneity: Tau ² = 0.	.43; Chi ² = 1	6.56, df = 3	3 (P = 0.00)	09); I ² = 82	2%						
Test for overall effect: Z	= 2.63 (P =	0.009)							-4 -2	0 2	4
Test for subgroup differen	ences: Not ap	oplicable							Higher with exercise	Higher wit	h control

Analysis 4.4. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 4: 6MWT

	Control		Exercise				Mean Difference	Mean Difference	
Study or Subgroup	Mean [metres]	SD [metres]	Total	Mean [metres]	SD [metres]	Total	Weight	IV, Random, 95% CI [metres]	IV, Random, 95% CI [metres]
Marchesan 2016 (1)	399.43	164	8	498.5	164	7	0.7%	-99.07 [-265.43 , 67.29]	
Rouchon 2016	400	65.987	4	420	11.9614	8	4.8%	-20.00 [-85.20 , 45.20]	ı <u> </u>
DePaul 2002	430	80	14	464	94	15	5.1%	-34.00 [-97.40 , 29.40]	ı <u>-</u>
DIALY-SIZE 2016	0.8	44	8	39	76.8	8	5.4%	-38.20 [-99.53 , 23.13]	ı
Cho 2018	-26	41	13	22	12	12	37.6%	-48.00 [-71.30 , -24.70]	
Frih 2017a	415.6	36.3	20	480.5	31.9	21	46.4%	-64.90 [-85.86 , -43.94]	•
Total (95% CI)			67			71	100.0%	-53.64 [-67.91 , -39.36]	ı •
Heterogeneity: Tau ² = 0	0.00; Chi ² = 3.26, df =	5 (P = 0.66); I ²	= 0%						•
Test for overall effect:	Z = 7.36 (P < 0.00001)							-500 -250 0 250 50
Test for subgroup differ	rences: Not applicable	2							Higher in exercise Higher in contr

(1) standard deviation imputed from the highest standard deviation of the other included studies

Analysis 4.5. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 5: Sit-To-Stand test [N reps/30 sec]

	Control			1	Exercise			Mean Difference	Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, F	Random, 9	5% CI	
Marchesan 2016 (1)	10	7.7	7	14.62	7.7	8	2.1%	-4.62 [-12.43 , 3.19	9]			
DIALY-SIZE 2016	1.4	4.3	8	1.4	3.5	8	8.4%	0.00 [-3.84 , 3.84	4]	_		
Cho 2018	-0.5	2.2	13	3.3	3.1	12	25.2%	-3.80 [-5.92 , -1.68	8]	-		
Frih 2017a	10.85	2.05	20	13.3	1.75	21	64.3%	-2.45 [-3.62 , -1.28	8]			
Total (95% CI)			48			49	100.0%	-2.63 [-3.77 , -1.49	9]	•		
Heterogeneity: Tau ² = 0	0.17; Chi ² = 3.	.31, df = 3	(P = 0.35)	; I ² = 9%						•		
Test for overall effect: 2	Z = 4.52 (P <	0.00001)							-20 -10	0	10	20
Test for subgroup differ	ences: Not ap	plicable						1	Higher with exerc	ise I	ligher wi	th control

Footnotes

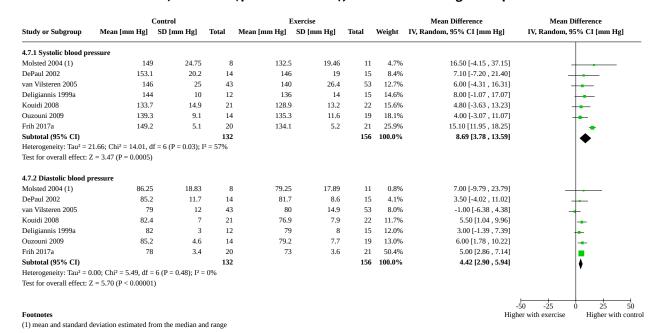
 $(1) \, standard \, deviation \, imputed \, from \, the \, highest \, standard \, deviation \, of \, the \, other \, included \, studies$



Analysis 4.6. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 6: Sit-To-Stand test [sit to 5 reps]

	Control]	Exercise		Mean Difference	Mean Difference			
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	IV, Random, 95% CI	IV, Rando	m, 95% CI		
Frih 2017a	15.5	1.55	20	13.5	1.45	21	2.00 [1.08 , 2.92]		-		
								-4 -2	0 2 4		
							Hı	gher with exercise	Higher with control		

Analysis 4.7. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 7: Resting blood pressure



Analysis 4.8. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 8: Aerobic capacity (VO2 max or peak)

		Control]	Exercise			Mean Difference		Mean D	ifference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl		IV, Rando	m, 95% Cl	I
Deligiannis 1999a	15.8	4.8	12	23.7	7.7	15	30.8%	-7.90 [-12.65 , -3.1	5]			
Molsted 2004 (1)	19.5	3.81	9	19.03	5.12	9	33.2%	0.47 [-3.70 , 4.64	4]		-	
Konstantinidou 2002 (2)	15.8	4.8	12	21.4	6.76	36	36.0%	-5.60 [-9.10 , -2.10	0]	-		
Total (95% CI)			33			60	100.0%	-4.29 [-8.98 , 0.39	9]		•	
Heterogeneity: Tau ² = 12.	67; Chi ² = 7.	.76, df = 2	(P = 0.02)	; I ² = 74%						_		
Test for overall effect: Z =	= 1.80 (P = 0	.07)							-20	-10	0 10	20
Test for subgroup differen	ces: Not app	licable						1	Higher w	ith exercise	Higher	with control

Footnotes

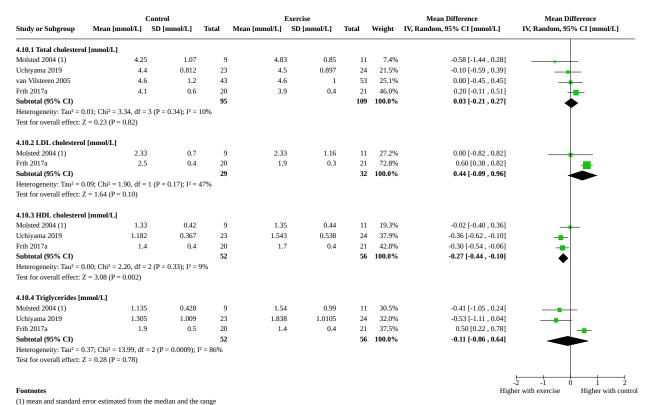
- (1) mean and standard deviation estimated from the median and range
- (2) three intervention arms pooled together in the exercise group



Analysis 4.9. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 9: Albumin

	(Control		E	exercise			Mean Difference		Mean Dif	ference	
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV.	, Random, 9	5% CI [g/	L]
Uchiyama 2019	33.7	5.2	23	34.8	4.4	24	25.2%	-1.10 [-3.86 , 1.6	6]			
Kopple 2007	39	3.7417	14	38	3.7	14	25.2%	1.00 [-1.76 , 3.7	6]		_	
Frih 2017a	40	2.6	21	40.4	3.7	20	49.6%	-0.40 [-2.37 , 1.5	7]	-		
Total (95% CI)			58			58	100.0%	-0.22 [-1.61 , 1.10	6]		-	
Heterogeneity: Tau ² =	0.00; Chi ² = 1.18,	, df = 2 (P =	0.56); I ² =	= 0%								
Test for overall effect:	Z = 0.32 (P = 0.7)	(5)							-4	-2 0	2	4
Test for subgroup diffe	rences: Not appli	icable							Higher with	exercise	Higher v	with contro

Analysis 4.10. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 10: Blood lipids



Analysis 4.11. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 11: Body composition

Study or Subgroup	Mean	Control SD	Total	Exercise Mean SD		Total	Mean Difference IV, Random, 95% CI	Mean Di IV, Randon	
4.11.1 Fat mass [kg] Kopple 2007	19.1	2.4	14	19.4	2.9	37	-0.30 [-1.87 , 1.27		
								-2 -1 0 Higher in exercise	1 2 Higher in control



Analysis 4.12. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 12: Body mass index

	Control		Exercise				Mean Difference	Mean Difference		
Study or Subgroup	Mean [kg/m²]	SD [kg/m ²]	Total	Mean [kg/m²]	SD [kg/m ²]	Total	Weight	IV, Fixed, 95% CI [kg/m²]	IV, Fixed, 95%	CI [kg/m²]
Uchiyama 2019	24.5	4.3	23	22.8	3.4	24	18.4%	1.70 [-0.52 , 3.92]		
Kopple 2007	25.1	1.2	14	26	1.5	12	81.6%	-0.90 [-1.96 , 0.16]	-	
Total (95% CI)			37			36	100.0%	-0.42 [-1.37 , 0.53]		.
Heterogeneity: Chi ² = 4	4.29, df = 1 (P = 0.0	4); I ² = 77%								
Test for overall effect: 2	Z = 0.86 (P = 0.39)								-4 -2 0	2 4
Test for subgroup differ	rences: Not applicab	ole						Hi	gher with exercise	Higher with control

Analysis 4.13. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 13: Calcium

	Control		Exercise				Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Deligiannis 1999a	2.02	0.2	12	2.2	0.8	15	2.5%	-0.18 [-0.60 , 0.24]	
Deligiannis 1999	2.17	0.15	30	2.2	0.7	30	6.8%	-0.03 [-0.29 , 0.23]	
Kouidi 2010	2.22	0.32	20	2.17	0.25	24	15.0%	0.05 [-0.12 , 0.22]	
Kouidi 2008	2.1457	0.15	29	2.07	0.15	30	75.8%	0.08 [-0.00 , 0.15]	=
Total (95% CI)			91			99	100.0%	0.06 [-0.01, 0.12]	•
Heterogeneity: Tau ² = 0	0.00; Chi ² = 1.90, df =	3 (P = 0.59); $I^2 = 0$	0%						
Test for overall effect: 2	Z = 1.71 (P = 0.09)								-1 -0.5 0 0.5 1
Test for subgroup differ	rences: Not applicable							High	her with exercise Higher with control

Analysis 4.14. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 14: CRP

		Control		I	Exercise			Mean Difference	Mean Dif	ference	
Study or Subgroup	Mean [mg/dL]	SD [mg/dL]	Total	Mean [mg/dL]	SD [mg/dL]	Total	Weight	IV, Random, 95% CI [mg/dL]	IV, Random, 959	% CI [mg/dL]	
Kopple 2007	2.8	2.9933	14	5.8	8.13	15	0.7%	-3.00 [-7.40 , 1.40]	1	_	_
Frih 2017a	4	1.4	20	4.1	1.2	21	17.5%	-0.10 [-0.90 , 0.70]] 🗼		
Uchiyama 2019	0.3	0.5	23	0.14	0.25	24	81.8%	0.16 [-0.07 , 0.39]] 🙀		
Total (95% CI)			57			60	100.0%	0.09 [-0.27 , 0.46]	1		
Heterogeneity: Tau ² = 0	0.03; Chi ² = 2.33, df	$= 2 (P = 0.31); I^{2}$	$^{2} = 14\%$								
Test for overall effect:	Z = 0.50 (P = 0.61)								-10 -5 0	5 10	0
Test for subgroup differ	rences: Not applicab	le						H	ligher with exercise	Higher with cor	ntrol

Analysis 4.15. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 15: Dialysis adequacy: Kt/V

	•	Control		1	Exercise		Mean Difference	Mean D	ifference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	IV, Random, 95% CI	IV, Rando	om, 95% CI	
van Vilsteren 2005	1.23	0.2	43	1.26	0.2	53	-0.03 [-0.11 , 0.05]		
							H	-0.2 -0.1 ligher with exercise	0 0.1 Higher w	0.2 ith control



Analysis 4.16. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 16: Energy intake

		Control		I	exercise		Mean Difference	Mean Di	fference
Study or Subgroup	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	Mean [kCal/kg/d]	SD [kCal/kg/d]	Total	IV, Random, 95% CI [kCal/kg/d]	IV, Random, 95%	6 CI [kCal/kg/d]
Kopple 2007	24.2	7.8575	14	27.2	8.52	15	-3.00 [-8.96 , 2.96]	1	
							н	-10 -5 0	5 10

Analysis 4.17. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 17: Haemoglobin

	Control		F	exercise			Mean Difference	Mean Difference	
Study or Subgroup	Mean [g/L]	SD [g/L]	Total	Mean [g/L]	SD [g/L]	Total	Weight	IV, Random, 95% CI [g/L]	IV, Random, 95% CI [g/L]
Kopple 2007	12.5	1.87	14	13	1.39	12	3.0%	-0.50 [-1.76 , 0.76	6]
Frih 2017a	10	1.6	20	10.4	1.7	21	4.7%	-0.40 [-1.41, 0.63	1]
Kouidi 2010	11.2	1.3	20	11.3	1.2	24	8.6%	-0.10 [-0.85 , 0.65	5]
Kouidi 2008	11	0.7	29	11	0.7	30	37.5%	0.00 [-0.36 , 0.36	5]
van Vilsteren 2005	7.57	0.8	43	7.52	0.8	53	46.2%	0.05 [-0.27 , 0.37	7] —
Total (95% CI)			126			140	100.0%	-0.02 [-0.24 , 0.20	0)
Heterogeneity: Tau ² = 0	0.00; Chi ² = 1.34	l, df = 4 (P =	0.85); I ² =	= 0%					Ĭ
Test for overall effect: 2	Z = 0.17 (P = 0.8)	36)							-2 -1 0 1 2
Test for subgroup differ	rences: Not appl	icable						1	Higher with exercise Higher with control

Analysis 4.18. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 18: Heart rate

	(Control			Exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [bpm]	SD [bpm]	Total	Mean [bpm]	SD [bpm]	Total	Weight	IV, Random, 95% CI [bpm]	IV, Random, 95% CI [bpm]		
4.18.1 Resting											
Deligiannis 1999a	81.8	8.5	12	77.3	9	15	12.6%	4.50 [-2.12 , 11.12]	+-		
Ouzouni 2009	78.2	10.3	14	76.3	7.1	19	14.1%	1.90 [-4.37 , 8.17]			
Deligiannis 1999	76	7	30	75	9	30	33.2%	1.00 [-3.08, 5.08]			
Kouidi 2008	78.4	8.1	29	73.7	6.3	30	40.1%	4.70 [0.99, 8.41]			
Subtotal (95% CI)			85			94	100.0%	3.05 [0.70, 5.40]	•		
Heterogeneity: Tau ² = 0	.00; Chi ² = 2.04, df	= 3 (P = 0.56)); I ² = 0%						•		
Test for overall effect: Z	Z = 2.54 (P = 0.01)										
4.18.2 Maximum											
Konstantinidou 2002	139	12	4	145.23	16.26	26	18.5%	-6.23 [-19.55 , 7.09]			
Deligiannis 1999a	139	12	12	146	20	15	22.1%	-7.00 [-19.19, 5.19]			
Ouzouni 2009	139.6	7.1	14	144.1	14.3	19	59.4%	-4.50 [-11.93 , 2.93]			
Subtotal (95% CI)			30			60	100.0%	-5.37 [-11.10 , 0.35]			
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.14, df	= 2 (P = 0.93)); I ² = 0%						•		
Test for overall effect: Z	Z = 1.84 (P = 0.07)										
									-20 -10 0 10		



Analysis 4.19. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 19: Muscular strength

		Control			Exercise			Mean Difference	Mean Difference		
Study or Subgroup	Mean [kg]	SD [kg]	Total	Mean [kg]	SD [kg]	Total	Weight	IV, Random, 95% CI [kg]	IV, Random, 95% CI [k	g]	
4.19.1 Knee extension											
DIALY-SIZE 2016	9.3	10.1	8	4.9	11.6	8	24.4%	4.40 [-6.26 , 15.06	5]		
Uchiyama 2019	24.9	10.5	23	24.2	10.7	24	75.6%	0.70 [-5.36 , 6.76	5]		
Subtotal (95% CI)			31			32	100.0%	1.60 [-3.66, 6.87	7]		
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.3	5, df = 1 (P	= 0.55); I ²	= 0%							
Test for overall effect: 2	Z = 0.60 (P = 0.	55)									
4.19.2 Handgrip											
Uchiyama 2019	25.7	6.4	23	27.3	5.4	24	49.2%	-1.60 [-4.99 , 1.79	9]		
Frih 2017a	30	5.2	20	37.4	4.8	21	50.8%	-7.40 [-10.47 , -4.33	3]		
Subtotal (95% CI)			43			45	100.0%	-4.55 [-10.23 , 1.14	4]		
Heterogeneity: Tau ² = 1	4.10; Chi ² = 6.	18, df = 1 (l	P = 0.01);	$I^2 = 84\%$							
Test for overall effect: 2	Z = 1.57 (P = 0.	12)									
									-20 -10 0 10		
									Higher in exercise Higher i	in con	

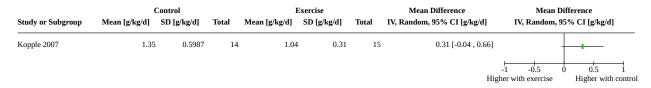
Analysis 4.20. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 20: Phosphate

	Control		Exercise				Mean Difference	Mean Difference	
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Deligiannis 1999a	1.94	0.42	12	2.03	0.58	15	9.5%	-0.09 [-0.47 , 0.29]	
Kouidi 2010	2.1	0.52	20	2.13	0.55	24	13.4%	-0.03 [-0.35, 0.29]	
Deligiannis 1999	1.97	0.36	30	2	0.55	30	24.4%	-0.03 [-0.27 , 0.21]	
Kouidi 2008	1.97	0.26	29	2	0.36	30	52.8%	-0.03 [-0.19 , 0.13]	—
Total (95% CI)			91			99	100.0%	-0.04 [-0.15 , 0.08]	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.09, df =	3 (P = 0.99); I ² =	0%						\neg
Test for overall effect: 2	Z = 0.60 (P = 0.55)								-0.5 -0.25 0 0.25 0.5
Test for subgroup differ	rences: Not applicable							Hig	ther with exercise Higher with contro

Analysis 4.21. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 21: Potassium

		Control		E	exercise			Mean Difference	Mean Difference
Study or Subgroup	Mean [mmol/L]	SD [mmol/L]	Total	Mean [mmol/L]	SD [mmol/L]	Total	Weight	IV, Random, 95% CI [mmol/L]	IV, Random, 95% CI [mmol/L]
Kouidi 2010	5.8	0.6	20	5.6	0.7	24	19.9%	0.20 [-0.18 , 0.58]	
Deligiannis 1999a	5.4	0.4	12	5.7	0.5	15	23.2%	-0.30 [-0.64 , 0.04]	
Deligiannis 1999	5.7	0.7	30	5.8	0.5	30	26.0%	-0.10 [-0.41, 0.21]	
Kouidi 2008	5.3	0.4	29	5.6	0.6	30	30.9%	-0.30 [-0.56 , -0.04]	
Total (95% CI)			91			99	100.0%	-0.15 [-0.36 , 0.06]	
Heterogeneity: Tau ² = 0	.02; Chi ² = 5.29, df =	3 (P = 0.15); I ² = 4	43%						
Test for overall effect: Z	Z = 1.38 (P = 0.17)								-1 -0.5 0 0.5 1
Test for subgroup differ	ences: Not applicable							Hig	her with exercise Higher with control

Analysis 4.22. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 22: Protein intake





Analysis 4.23. Comparison 4: Combined aerobic and resistance exercise versus control (no exercise/placebo exercise), Outcome 23: Timed up-and-go test

Study or Subgroup	Mean [sec]	Control SD [sec]	Total	Mean [sec]	Exercise SD [sec]	Total	Mean Difference IV, Random, 95% CI [sec]		Mean I IV, Random			İ
Frih 2017a	15.2	1.9	20	12.9	1.6	21	2.30 [1.22 , 3.38]				<u> </u>	
								⊢ -4 Higher	-2 in exercise	0	2 Higher in	4 control

APPENDICES

Appendix 1. Electronic search strategies

Database	Search terms								
CENTRAL	1. exercise:ti,ab,kw								
	2. (physical next (training or activity or fitness or rehabilitation)):ti,ab,kw								
	3. (resistance next (training or program*)):ti,ab,kw								
	(strength* and (muscle* or program* or training)):ti,ab,kw								
	5. kinesiotherapy:ti,ab,kw								
	6. {or #1-#5}								
	7. (uremi* or uraemi*):ti,ab,kw								
	8. renal replacement therapy:ti,ab,kw								
	9. dialysis:ti,ab,kw								
	10.(hemodialysis or haemodialysis):ti,ab,kw								
	11.(kidney transplant* or renal transplant*):ti,ab,kw								
	12.(predialysis or pre-dialysis):ti,ab,kw								
	13.renal insufficiency:ti,ab,kw								
	14.MeSH descriptor: [Renal Insufficiency, Chronic] explode all trees								
	15.((kidney or renal) next (failure or disease)):ti,ab,kw								
	16.(CKD or CKF or CRD or CRF or ESRD or ESKD or ESRF or ESKF):ti,ab,kw								
	17.{or #7-#16}								
	18.{and #6, #17}								
MEDLINE	1. exp Exercise/								
	2. Physical Exertion/								
	3. exp Physical Fitness/								
	4. exp Exercise Therapy/								
	5. Exercise Test/								
	6. exp Exercise Movement Techniques/								
	7. exercise.tw.								
	8. (resistance training or resistance program\$).tw.								
	9. (physical fitness or physical rehabilitation).tw.								
	10.(strength\$ and (muscle or program\$ or training)).tw.								
	11.or/1-10								
	12.Kidney Diseases/								
	13.exp Renal Replacement Therapy/								
	14.Renal Insufficiency/								
	15.exp Renal Insufficiency, Chronic/								
vercise training for adul	Its undergoing maintenance dialysis (Review)	28							



(Continued)

- 16. Diabetic Nephropathies/
- 17.exp Hypertension, Renal/
- 18.dialysis.tw.
- 19.(hemodialysis or haemodialysis).tw.
- 20.(hemofiltration or haemofiltration).tw.
- 21.(hemodiafiltration or haemodiafiltration).tw.
- 22.(kidney disease* or renal disease* or kidney failure or renal failure).tw.
- 23.(ESRF or ESKF or ESRD or ESKD).tw.
- 24.(CKF or CKD or CRF or CRD).tw.
- 25.(CAPD or CCPD or APD).tw.
- 26.(predialysis or pre-dialysis).tw.
- 27.or/12-26
- 28.and/11,27

EMBASE

- 1. exp exercise/
- 2. exp "physical activity, capacity and performance"/
- 3. exp kinesiotherapy/
- 4. exp exercise test/
- 5. exercise.tw.
- 6. (resistance training or resistance program\$).tw.
- 7. (physical fitness or physical rehabilitation).tw.
- 8. (strength\$ and (muscle or program\$ or training)).tw.
- 9. or/1-8
- 10.exp renal replacement therapy/
- 11.kidney disease/
- 12.chronic kidney disease/
- 13.kidney failure/
- 14.chronic kidney failure/
- 15.mild renal impairment/
- 16.stage 1 kidney disease/
- 17.moderate renal impairment/
- 18. severe renal impairment/
- 19.end stage renal disease/
- 20.renal replacement therapy-dependent renal disease/
- 21.diabetic nephropathy/
- 22.kidney transplantation/
- 23.renovascular hypertension/
- 24.(hemodialysis or haemodialysis).tw.
- 25.(hemofiltration or haemofiltration).tw.
- 26. (hemodia filtration or haemodia filtration).tw.
- 27.dialysis.tw.
- 28.(CAPD or CCPD or APD).tw.
- 29.(kidney disease* or renal disease* or kidney failure or renal failure).tw.
- 30.(CKF or CKD or CRF or CRD).tw.
- 31.(ESRF or ESKF or ESRD or ESKD).tw.
- 32.(predialysis or pre-dialysis).tw.
- 33.((kidney or renal) adj (transplant* or graft* or allograft*)).tw.
- 34.or/10-33
- 35.and/9,34



Appendix 2. Risk of bias assessment tool

Potential source of bias

Assessment criteria

Random sequence generation

Selection bias (biased allocation to interventions) due to inadequate generation of a randomised sequence

Low risk of bias: Random number table; computer random number generator; coin tossing; shuffling cards or envelopes; throwing dice; drawing of lots; minimisation (minimisation may be implemented without a random element, and this is considered to be equivalent to being random).

High risk of bias: Sequence generated by odd or even date of birth; date (or day) of admission; sequence generated by hospital or clinic record number; allocation by judgement of the clinician; by preference of the participant; based on the results of a laboratory test or a series of tests; by availability of the intervention.

Unclear: Insufficient information about the sequence generation process to permit judgement.

Allocation concealment

Selection bias (biased allocation to interventions) due to inadequate concealment of allocations prior to assignment

Low risk of bias: Randomisation method described that would not allow investigator/participant to know or influence intervention group before eligible participant entered in the study (e.g. central allocation, including telephone, web-based, and pharmacy-controlled, randomisation; sequentially numbered drug containers of identical appearance; sequentially numbered, opaque, sealed envelopes).

High risk of bias: Using an open random allocation schedule (e.g. a list of random numbers); assignment envelopes were used without appropriate safeguards (e.g. if envelopes were unsealed or non-opaque or not sequentially numbered); alternation or rotation; date of birth; case record number; any other explicitly unconcealed procedure.

Unclear: Randomisation stated but no information on method used is available.

Blinding of participants and personnel

Performance bias due to knowledge of the allocated interventions by participants and personnel during the study Low risk of bias: No blinding or incomplete blinding, but the review authors judge that the outcome is not likely to be influenced by lack of blinding; blinding of participants and key study personnel ensured, and unlikely that the blinding could have been broken.

High risk of bias: No blinding or incomplete blinding, and the outcome is likely to be influenced by lack of blinding; blinding of key study participants and personnel attempted, but likely that the blinding could have been broken, and the outcome is likely to be influenced by lack of blinding.

Unclear: Insufficient information to permit judgement

Blinding of outcome assessment

Detection bias due to knowledge of the allocated interventions by outcome assessors.

Low risk of bias: No blinding of outcome assessment, but the review authors judge that the outcome measurement is not likely to be influenced by lack of blinding; blinding of outcome assessment ensured, and unlikely that the blinding could have been broken.

High risk of bias: No blinding of outcome assessment, and the outcome measurement is likely to be influenced by lack of blinding; blinding of outcome assessment, but likely that the blinding could have been broken, and the outcome measurement is likely to be influenced by lack of blinding.

Unclear: Insufficient information to permit judgement

Incomplete outcome data

Attrition bias due to amount, nature or handling of incomplete outcome data.

Low risk of bias: No missing outcome data; reasons for missing outcome data unlikely to be related to true outcome (for survival data, censoring unlikely to be introducing bias); missing outcome data balanced in numbers across intervention groups, with similar reasons for missing data across groups; for dichotomous outcome data, the proportion of missing outcomes compared with observed event risk not enough to have a clinically relevant impact on the intervention effect estimate; for continuous outcome data, plausible effect size (difference in means or standardised difference in means) among missing outcomes not enough to have a clinically relevant impact on observed effect size; missing data have been imputed using appropriate methods.



(Continued)

High risk of bias: Reason for missing outcome data likely to be related to true outcome, with either imbalance in numbers or reasons for missing data across intervention groups; for dichotomous outcome data, the proportion of missing outcomes compared with observed event risk enough to induce clinically relevant bias in intervention effect estimate; for continuous outcome data, plausible effect size (difference in means or standardized difference in means) among missing outcomes enough to induce clinically relevant bias in observed effect size; 'as-treated' analysis done with substantial departure of the intervention received from that assigned at randomisation; potentially inappropriate application of simple imputation.

Unclear: Insufficient information to permit judgement

Selective reporting

Reporting bias due to selective outcome reporting

Low risk of bias: The study protocol is available and all of the study's pre-specified (primary and secondary) outcomes that are of interest in the review have been reported in the pre-specified way; the study protocol is not available but it is clear that the published reports include all expected outcomes, including those that were pre-specified (convincing text of this nature may be uncommon).

High risk of bias: Not all of the study's pre-specified primary outcomes have been reported; one or more primary outcomes is reported using measurements, analysis methods or subsets of the data (e.g. sub-scales) that were not pre-specified; one or more reported primary outcomes were not pre-specified (unless clear justification for their reporting is provided, such as an unexpected adverse effect); one or more outcomes of interest in the review are reported incompletely so that they cannot be entered in a meta-analysis; the study report fails to include results for a key outcome that would be expected to have been reported for such a study.

Unclear: Insufficient information to permit judgement

Other bias

Low risk of bias: The study appears to be free of other sources of bias.

Bias due to problems not covered elsewhere in the table

High risk of bias: Had a potential source of bias related to the specific study design used; stopped early due to some data-dependent process (including a formal-stopping rule); had extreme baseline imbalance; has been claimed to have been fraudulent; had some other problem.

Unclear: Insufficient information to assess whether an important risk of bias exists; insufficient rationale or evidence that an identified problem will introduce bias.

Appendix 3. Characteristics of included interventions

Trial name	Type of exercise	Description of exercise	Materials	Intensity class	Who provid- ed/supervised	Maximum duration	Frequen- cy (time/ week)	Timing in relation to HD ses- sions	Duration of inter- vention (week)
Abreu 2017	resistance	lower limbs exercises	ankle weights and resistance bands	moderate	physiotherapist	30	3	during	12
Abundis Mora 2017	aerobic	stationary cycling	ergometer	moderate	not reported	135/week	not report- ed	during	35
ACTINUT 2013	aerobic	stationary cycling	ergometer	moderate	physician, nurse, exercise physiol- ogist	35	3	during	24
Afshar 2010 (A)	resistance	lower limbs exercises	ankle weights	moderate to vigorous	physician	40	3	during	8
Afshar 2010 (B)	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	40	3	during	8
Afshar 2011	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	40	3	during	8
Akiba 1995	aerobic	stationary cycling	ergometer	moderate	not reported	30	3	during	12
Amini 2016	aerobic	not reported	not reported	not reported	researcher	not report- ed	not report- ed	not report- ed	8
AVANTE-HE- MO 2020 (A)	aerobic	stationary cycling	ergometer	moderate	not reported	30	3	during	12
AVANTE-HE- MO 2020 (B)	resistance	upper and lower limbs exer- cises	resistance bands	moderate	not reported	40	3	during	12
Bennett 2013	resistance	lower body exercises	resistance bands and tub- ing	not reported	exercise physiol- ogist	varied	3	during	12

(Continued)									
Burrows 2018	combined	stationary cycling and to- tal body resistance and bal- ance exercises	ergometer + resistance bands	moderate	not reported	30 min in- traHD + home ses- sions	5	during	24
Carmack 1995	aerobic	stationary cycling	ergometer	not reported	not reported	30	3	during	10
CHAIR 2015	aerobic	chair-stand exercise	chair	not reported	physician and physiotherapist	15	3	just before	12
Chang 2010	aerobic	stationary cycling	ergometer	moderate	not reported	35	3	during	8
Chen 2010	resistance	lower body exercises	ankle weights	moderate	supervised not further defined	not report- ed	2	during	26
Cho 2018 (A)	aerobic	stationary cycling	ergometer	not reported	not reported	30	3	during	12
Cho 2018 (B)	resistance	upper and lower limbs exer- cises	resistance bands and soft weights	not reported	not reported	not report- ed	3	during	12
Cho 2018 (C)	combined	combination of A and B	ergometer, resistance bands and soft weights	not reported	not reported	not report- ed	3	during	12
Cooke 2018	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	varied	3	during	16
CYCLE-HD 2016	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	30	3	during	26
Dashtidehkor- di 2019	aerobic	stationary cycling	ergometer	not reported	not reported	60	3	during	8
de Lima 2013 (A)	resistance	lower limbs exercises	not reported	not reported	not reported	not report- ed	3	during	8
de Lima 2013 (B)	aerobic	stationary cycling	ergometer	light to moderate	not reported	20	3	during	8
Deligiannis 1999	combined	bicycling and/or walking, callisthenics, steps, swim- ming, or ball games fol-	not reported	moderate	physician, exer- cise physiologist,	90	3-4	on non-HD days	26

ter commencement of dialysis)

Cochrane Library

Trusted evidence.
Informed decisions.
Better health.

(Continued)		lowed by resistance program			and physical edu- cation instructor				
Deligiannis 1999a (A)	combined	stationary cycling, callis- thenics, steps and flexibility exercises	ergometer or treadmill	moderate	physician and physical educa- tion teachers	90	3	on non-HD days	26
Deligiannis 1999a (B)	aerobic	stationary cycling and ex- tension exercises	ergometer	moderate	physician and physical educa- tion teachers	30	5	not during	26
DePaul 2002	combined	stationary cycling and lower limbs strength training	ergometer and Response Seat- ed Leg Curl Thigh Extension pulley weight system	moderate	kinesiologist	varied	3	during and just before or after	12
DIALY-SIZE 2016(A)	aerobic	stationary cycling	ergometer	moderate	kinesiologist	53	3	during	12
DIALY-SIZE 2016 (B)	resistance	lower limbs exercises	ankle weights and resistance bands	moderate	kinesiologist	varied	3	during	12
DIALY-SIZE 2016 (C)	combined	all of (A) and (B)	A + B	moderate	kinesiologist	varied	3	during	12
Dobsak 2012	aerobic	stationary cycling	ergometer	light to moderate	not reported	50	3	during	20
Dong 2011	resistance	lower limbs exercises	pneumatic leg press machine	moderate	study personnel	varied	3	just before	20
EXCITE 2014	aerobic	walking	-	light to moderate	not supervised	varied	3	on non-HD days	26
Fernandes 2019	aerobic	stationary cycling	ergometer	moderate	not reported	50	3	during (1 hour af-	8

(Continued)									
Frey 1999	aerobic	stationary cycling	multigym	moderate to vigorous	not reported	55	3	during	8
Frih 2017a	combined	upper and lower limbs exer- cises and stationary cycling and walking	ergometer, treadmill, multigym	moderate	physiotherapist and trainer	60	4	on non-HD days	16
Giannaki 2013a	aerobic	stationary cycling	ergometer	moderate	not reported	not report- ed	3	during	26
Goldberg 1983	aerobic	cycling or walking	ergometer, run- ning track	moderate	not reported	80	3	on non-HD days	52 ± 16
Harter 1985	aerobic	cycling or walking	ergometer, run- ning track	moderate	physician, nurse, exercise physiol- ogist	45	not report- ed	on non-HD days	52
Groussard 2015	aerobic	stationary cycling	ergometer	moderate	"professional team with exper- tise in physical activity"	40	3	during	12
IHOPE 2019	aerobic	stationary cycling	ergometer	moderate	research staff	45	3	during	52
Johansen 2006	resistance	lower limbs exercises	ankle weights	moderate	study personnel	varied	3	during	12
Koh 2009 (A)	aerobic	stationary cycling	ergometer	moderate	supervised not further defined	45	3	during	24
Koh 2009 (B)	aerobic	walking	-	moderate	unsupervised	45	3	not report- ed	24
Konstanti- nidou 2002 (A)	combined	Calisthenics, steps, flexibili- ty, stretching and resistance exercises	ergometer	moderate	Sports physician, physical educa- tion instructor	40	3	On non- HD days	26
Konstanti- nidou 2002 (B)	combined	Stationary cycling and low- er limbs exercises	ergometer, resistance bands and weights	moderate to vigorous	sports physician, physical educa- tion instructor	20 aero- bic + resis- tance	3	during	26

Cochrane

(Continued)									
Konstanti- nidou 2002 (C)	combined	stationary cycling	ergometer	moderate	not supervised	30 aero- bic+resis- tance	5	not report- ed	26
Kopple 2007 (A)	aerobic	stationary cycling	ergometer	moderate	investigator	70	3	during	20
Kopple 2007 (B)	resistance	lower limbs exercises	leg exten- sion/leg curl and leg press/ calf extension apparatus	moderate	investigator	not report- ed	3	just before	20
Kopple 2007 (C)	combined	50% of (A) and 50% of (B)	A + B	moderate	investigator	not report- ed	3	just before and during	20
Koufaki 2003	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	40	3	during	12
Kouidi 1997	aerobic	stationary cycling, walking or jogging, callisthenics, aerobics, swimming and/or game sports	not reported	moderate	physician, exer- cise physiologist, trainer	90	3-4	on non-HD days	26
Kouidi 2003	aerobic	stationary cycling	not reported	not reported	supervised not further defined	not report- ed	3	during	52
Kouidi 2004a	aerobic	stationary cycling	not reported	not reported	supervised not further defined	not report- ed	3	during	26
Kouidi 2005	aerobic	stationary cycling	not reported	not reported	supervised not further defined	not report- ed	3	during	43.4
Kouidi 2008	combined	stationary cycling and ab- dominal and lower limbs ex- ercises	ergometer, weights and elastic bands	moderate to vigorous	exercise trainers, physician	110	3	during	43.4
Kouidi 2010	combined	stationary cycling and lower limbs exercises	ergometer, free weights and re- sistance bands	not reported	exercise trainers, physician	100	3	during	52
Lee 2001	aerobic	stationary cycling and walk- ing	ergometer, treadmill	moderate	not reported	40	2-4	just prior	12

(Continued)									
Liao 2016	aerobic	stationary cycling	ergometer	moderate to vigorous	physician and nurse	30	3	during	12
Ma 2018	combined	aerobics, resistance, and flexibility training not fur- ther defined	not reported	not reported	not reported	20	3	during	104
Makhlough 2012	range of movement	rotating the wrist, wrist up and down, ankle twisting motion	-	light to moderate	not reported	15	3	during	8
Marchesan 2016	combined	stationary cycling and up- per and lower limbs, thorax and abdominal exercises	ergometer and step	moderate	not reported	45+resis- tance	3	during	24
Marinho 2016	resistance	lower limbs exercises	resistance bands and an- kle cuffs	moderate	physical educa- tion teacher	varied	3	during	8
Martin-Ale- many 2016	resistance	upper and lower limbs exer- cises	ankle weights and resistance springs	moderate	not reported	40	2	during	12
Martins do Valle 2020	resistance	lower and upper limbs exer- cises	ankle weights and dumbbells	moderate to vigorous	supervised not further defined	varied	3	during	12
Matsumoto 2007	aerobic	stationary cycling	ergometer	moderate	research assis- tants	20	3	just before	52
McAdams-De- Marco 2018	aerobic	stationary cycling	ergometer	not reported	research assis- tants	varied	3	during	12
McGregor 2018	aerobic	stationary cycling	ergometer	moderate	exercise physiol- ogists	70	3	during	10
Mitsiou 2015	not report- ed	not reported	not reported	not reported	not reported	not report- ed	not report- ed	during	26
Miura 2015	aerobic	stationary cycling	ergometer	light to moderate	not reported	60	3	during	12
Molsted 2004	combined	step and circuit training and stationary cycling	ergometer, step	moderate to vigorous	physiotherapist	70	2	not report- ed	21.7
		stationary cycling		vigorous				ed	



(Continued)									
Momeni 2014	aerobic	stationary cycling	mini bike	not reported	not reported	30	3	during	12
Mortazavi 2013	aerobic	stationary cycling	ergometer	light to moderate	not reported	30	3	during	16

Momeni 2014	aerobic	stationary cycling	mini bike	not reported	not reported	30	3	during	12
Mortazavi 2013	aerobic	stationary cycling	ergometer	light to moderate	not reported	30	3	during	16
Olvera-Soto 2016	resistance	upper and lower limbs exer- cises	weights and resistance bands	not reported	supervised not further defined	50	2	during	12
Ouzouni 2009	combined	stationary cycling and ab- dominal and lower limbs ex- ercises	ergometer, weights and re- sistance bands	not reported	physician and ex- ercise physiolo- gist	20 aero- bic + resis- tance	3	during	43.4
Painter 2002a	aerobic	stationary cycling	ergometer	moderate to vigorous	research assis- tants	40	3	during	21.7
Paluchamy 2018	aerobic	stationary cycling	ergometer	not reported	not reported	20	3	during	12
Parsons 2004	aerobic	stationary cycling	ergometer	light to moderate	not reported	45	3	during	12
PEAK 2006	resistance	upper and lower limbs exer- cises	ankle and free- weights dumb- bells	moderate to vigorous	exercise physiol- ogist	45	3	prior and during	12
Pellizzaro 2013	resistance	knee extensions	free leg weights	moderate	not reported	varied	3	during	10
Rahimi- moghadam 2017	resistance	modified pilates	not reported	not reported	pilates profes- sional	45	3	on non-HD days	8
Reboredo 2010	aerobic	stationary cycling	horizontal er- gometer	moderate	supervised not further defined	50	3	during	12
Rezaei 2015	range of movement	joints warming actions, stretching, lower back and abdominal exercises, and deep breathing exercises.	not reported	light to moderate	unsupervised	35	3	not dur- ing, home- based	10
Rosa 2018	resistance	upper and lower limbs exercises	weights and resistance bands	not reported	exercise physiol- ogist	50	3	prior and during	12

Cochrane

(Continued)									
Rouchon 2016	combined	stationary cycling and up- per and lower limbs exercis- es	ergometer, weights	not reported	not reported	35	2	not applic- able,PD patients only	12
Samara 2016	aerobic	swimming	pool, foam tubes, buoyan- cy belts, pad- dles	moderate	trainer	60	3	on non-HD days	16
Segura-Orti 2009	resistance	lower limbs exercises	ankle weights	moderate	physiotherapist	35	3	during	24
Sheshadri 2020	aerobic	walking and weekly activity goals	pedometer	not applica- ble	unsupervised	not applic- able	not applic- able	not during	12
Soliman 2015	range of movement	range of motion exercises	not reported	light to moderate	not reported	15	3	during	8
Song 2012a	resistance	upper and lower limbs exer- cises	ankle weights and resistance bands	moderate	investigator and research assis- tant	30	3	not during	12
Suhardjono 2019 (A)	aerobic	stationary cycling	ergometer	moderate	physician	30	2	during	12
Suhardjono 2019 (B)	combined	stationary cycling and lower limbs exercises	ankle weights	light to moderate	physicians	not report- ed	2	during	12
Toussaint 2008	aerobic	stationary cycling	ergometer	not reported	unsupervised	30	3	during	12
Tsuyuki 2003	aerobic	cycling, walking and jogging	ergometer	moderate	physician	30	2-3	on non-HD days	20
Uchiyama 2019	combined	walking, upper and lower limbs exercises	resistance bands	moderate	unsupervised	30	3	not applic- able, PD patients only	12
van Vilsteren 2005	combined	stationary cycling, callis- thenics, steps and flexibility and resistance exercises	multi-trainer	moderate	not reported	60	3	during and prior to HD	12

(Continued) Wilund 2010	aerobic	stationary cycling	ergometer	moderate	research assis- tants	45	3	during	16
Wu 2014d	aerobic	stationary cycling	ergometer	moderate to vigorous	not reported	20	3	during	12
Yurtkuran 2007	yoga	modified yoga exercise	not reported	not reported	not reported	30	2	not report- ed	12
Zhao 2017	aerobic	road cycling	bicycle, road	not reported	not reported	70	6	after	18



Appendix 4. 2011 search strategies

The search strategies below were created by the authors of the initial review. These strategies have not been updated and were not used for the 2021 update.

DATABASE	Search terms
CINAHL	1. exertion/
	2. therapeutic exercise/
	3. exercise test/
	4. physical fitness/
	5. or/1-4
	6. exercise.tw.
	7. (resistance training or resistance program\$).tw.
	8. (physical fitness or physical rehabilitation).tw.
	9. (strength\$ and (muscle\$ or program\$ or training)).tw.
	10.or/6-9
	11.or/5,10
	12.uremia/
	13.ur?emi\$.tw.
	14.12 or 13
	15.(hemodialysis or haemodialysis).tw.
	16.dialysis.tw.
	17.renal replacement therapy/
	18.kidney failure chronic/
	·
	19. (kidney failure or renal failure or kidney disease or renal disease).tw.
	20.(CKD or CKF or CRD or CRF or ESKD or ESKF or ESRD or ESRF).tw.
	21.or/15-20
	22.or/14,21
	23.and/11,22
Webscience (Science citation	1. (exertion OR exercise therapy OR physical education and training OR physical fitness OR exercise
index and Social science citation index)	program* OR exercise training) AND (uremia OR ur?emia OR hemodialysis OR haemodialysis OR peritoneal dialysis OR renal* OR kidney*)
	2. (excertion OR exercise* OR motion therapy* OR physical educ* OR physical train* OR physical fit-
	ness*) AND (uremia OR ur?emia OR hemodialysis OR haemodialysis OR peritoneal dialysis OR re-
	nal* OR kidney*) AND (controlled clinical trial* OR CCT OR clinical trial* OR CT OR Randomized
	controlled trial* OR RCT)
BIOSIS	1. exertion.mp.
	2. exercise therapy.mp.
	3. exercise test.mp.
	4. (physical education and training).mp. [mp=title, book title (english), original language book title
	(non-english), abstract, concept codes, biosystematic codes, chemicals & biochemicals, diseases,
	major concepts, methods & equipment, organisms, parts, structures & systems of organisms, se-
	quence data, super taxa, taxa notes, time, geopolitical locations, gene name, miscellaneous de- scriptors]
	5. physical fitness.mp.
	6. 1 or 2 or 3 or 4 or 5
	7. exercise program\$.mp.
	8. exercise training.mp.
	9. 7 or 8
	10.6 or 9



(Continued)	11.uremia.mp. 12.ur?emia.mp. 13.11 or 12 14.renal replacement therapy.mp. 15.haemodialysis.mp. 16.hemodialysis.mp. 17.renal transplant\$.mp. 18.peritoneal dialysis.mp. 19.14 or 15 or 16 or 17 or 18 20.kidney failure chronic.mp. 21.chronic kidney failure.mp. 22.chronic renal failure.mp. 23.20 or 21 or 22
	24.13 or 19 or 23 25.10 and 24
PEDRO	 abstract & Title: renal Therapy: fitness training
AMED	 exp Exertion/ exercise therapy.mp. or Treatment group Exercise therapy/ exp Exercise testing/ or exercise test.mp. (physical education and training).mp. exp Physical fitness/ 1 or 2 or 3 or 4 or 5 exercise program?.mp. exercise training.mp. 7 or 8 6 or 9 uremia.mp. ur?emia.mp. haemodialysis.mp. haemodialysis.mp. replacement therapy.mp. peritoneal dialysis.mp. exendialysis.mp. or exp Hemodialysis/ oxp Kidney failure chronic/ chronic kidney failure.mp. chronic kidney failure.mp. or 20 or 21 or 22 and 24
PsycINFO	 exp EXERCISE/ or exercise.mp. exp Dialysis/ or Kidney Diseases/ or Organ Transplantation/ or Kidneys/ 1 AND 2 limit 3 to human
AGELINE	 Exercise OR Exertion OR Fitness OR Training uremia OR renal OR kidney OR hemodialysis OR peritoneal dialysis



(Continued)	3. Combine with AND4. Limit to Research/Academic and Professional/Provider	
KoreaMed	 exercise [ALL] AND nephrol [ALL] exercise [ALL] AND kidney [ALL] 	

HISTORY

Date	Event	Description
22 May 2017	Amended	Updated search strategies for MEDLINE, EMBASE & CENTRAL

CONTRIBUTIONS OF AUTHORS

- Amelie Bernier-Jean designed the new features of the systematic review and meta-analyses from the previous version, coordinated the review process, screened all the search results, assessed all the studies for quality, extracted data for all studies, analysed data, conducted the meta-analysis, and had the primary role in writing the manuscript.
- Nadim Berudi screened search results, assessed studies for quality, extracted data, conducted the independent double verification of all results provided in the manuscript and reviewed the final manuscript.
- Nicola Bondonno screened search results, assessed studies for quality, extracted data and reviewed the final manuscript.
- Gabrielle Williams screened search results, assessed studies for quality, extracted data and reviewed the final manuscript.
- Jonathan Craig contributed to the design of the new features of the systematic review and meta-analyses from the previous version and reviewed the final manuscript.
- Germaine Wong contributed to the design of the new features of the systematic review and meta-analyses from the previous version and reviewed the final manuscript.

DECLARATIONS OF INTEREST

- Amelie Bernier-Jean has declared that they have no conflict of interest
- · Nadim A Beruni has declared that they have no conflict of interest
- Nicola Patricia P Bondonno has declared that they have no conflict of interest
- Gabrielle Williams has declared that they have no conflict of interest
- Armando Teixeira-Pinto has declared that they have no conflict of interest
- Jonathan C Craig has declared that they have no conflict of interest
- · Germaine Wong has declared that they have no conflict of interest

SOURCES OF SUPPORT

Internal sources

• No sources of support provided

External sources

• NHMRC, Australia

Postgraduate Scholarship Scheme (GNT1151246)

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

Types of participants

The protocol and the original review published in Heiwe 2011 included adults at all stages of CKD, including those not undergoing dialysis and kidney transplant recipients. The current reviews included only adults undergoing maintenance dialysis. Adults with CKD not undergoing dialysis and kidney transplant recipients will be the subject of separate reviews.



Types of outcome measures

All the outcomes selected in the protocol and in Heiwe 2011 were updated. However, only the outcomes that are important to patients, their caregivers and health professionals were reported in the text of the review. Whenever appropriate, we conducted exploratory meta-analyses of the remaining outcomes.

Statistical assessment

All meta-analyses were random-effects meta-analyses. As in Heiwe 2011, in this update we did not adjust for the degree of anaemia, the degree of glomerular filtration rate, the duration of uraemia and dialysis adequacy as the protocol initially intended it.

INDEX TERMS

Medical Subject Headings (MeSH)

Exercise; Fatigue [etiology]; Quality of Life; *Renal Dialysis; *Resistance Training

MeSH check words

Adult; Humans