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Exercise interventions for cerebral palsy (Review)

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TABLE OF CONTENTS

SUMMARY OF FINDINGS 4 BACKGROUND 9 DECTWS 11 METHODS 11 Figure 1 15 Figure 2 19 Figure 3 20 Figure 4 22 Figure 5 23 DISCUSSION 29 AUTHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 ACROWLEDGEMENTS 28 Analysis 1.1, Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term. 128 Analysis 1.2, Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.2, Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.4, Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.5, Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term. 129 Analysis 1.6, Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, children and adolescents; short term. 129 Analysis 2.7, Comparison 2 Resistance training versus usual care, Outcome 4 Activity:	ABSTRACT	1
BACKGOUND 9 DBJECTWES 11 DREHODS 11 RESUTS 14 Figure 1. 15 Figure 2. 19 Figure 3. 20 Figure 4. 22 Figure 5. 23 DSCUSSION 32 ACKNOWLEDGEMENTS 33 CHARACTENISTICS OF STUDIES. 44 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: walking endurance; short term. 129 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: walking endurance; short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: walking endurance; short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Activity: dait physical activity: short term. 131 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Activity: dait speed, children and adolescents; short term. 132 Analysis 2.0. Comparison 2 Resistance training versus usual care,	PLAIN LANGUAGE SUMMARY	2
DBJECTIVES 11 METHODS 11 Figure 1. 15 Figure 2. 19 Figure 3. 20 Figure 4. 22 Figure 5. 23 DISCUSSION 29 VITHORS' CONCLUSIONS 29 ADMYDRSY CONCLUSIONS 29 ADAYASING SOF STUDIES 34 ARASCTENTISCS OF STUDIES 49 DATA AND ANALYSES 34 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gait speed, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gait speed, children and adolescents; short term. 131 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: gait speed, children and adolescents; short term.	SUMMARY OF FINDINGS	4
METHODS 11 RESULTS 14 Figure 1 15 Figure 2 19 Figure 3 20 Figure 4 22 Figure 5 23 DSCUSSION 29 DATHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 CKNOW LEDGENENTS 33 REFERENCES 34 Analysis 1.1 Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gaits speed, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Activity: gait speed, children and adolescents; short term. 129 Analysis 1.7. Comparison 1 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; and adolescents; anot function, children and adolescents; intermediate term. 121	BACKGROUND	9
RESULTS 14 Figure 1 15 Figure 2 19 Figure 3 20 Figure 4 22 Figure 5 23 DISCUSSION 22 AUTHORS' CONCLUSIONS 23 ACKNOWLEDGEMENTS 33 REFERENCES 34 Analysis 1.1. Comparison 1 Aerobic exercise versus susual care, Outcome 1 Activity: gross motor function, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus susual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.2. Comparison 1 Aerobic exercise versus susual care, Outcome 3 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus susual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus susual care, Outcome 5 Activity: gait speed, short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus susual care, Outcome 4 Activity: dait physical activity: short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus susual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term. 129 Analysis 2.5. Comparison 1 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 131 Anal	OBJECTIVES	11
Figure 1. 15 Figure 2. 19 Figure 3. 20 Figure 4. 22 Figure 5. 23 DISCUSSION 29 AUTHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 ACKNOW LEDGEMENTS 32 ACKNOW LEDGEMENTS 34 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gross motor function, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gross motor function, children and adolescents; short term. 129 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate	METHODS	11
Figure 2. 19 Figure 3. 20 Figure 4. 22 Figure 5. 23 SIGUSSION 29 AUTHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 ACKNOWLEDGEMENTS 33 REFREENCES 34 CHARACTERISTICS OF STUDIES 49 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, intermediate term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, nitermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic finess; short term. 130 Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic finess; short term. 131 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term. 132	RESULTS	14
Figure 2. 19 Figure 3. 20 Figure 4. 22 Figure 5. 23 SIGUSSION 29 AUTHORS' CONCLUSIONS 29 AUTHORS' CONCLUSIONS 29 ACKNOWLEDGEMENTS 33 REFREENCES 34 CHARACTERISTICS OF STUDIES 49 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, intermediate term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, nitermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic finess; short term. 130 Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic finess; short term. 131 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term. 132	Figure 1	15
Figure 4. 22 Figure 5. 23 DSCUDSION 29 AUTHORS' CONCLUSIONS 32 ACKNOWLEDGEMENTS 33 CHARACTERISTICS OF STUDIES 34 CHARACTERISTICS OF STUDIES 49 DATA AND ANALYSES 128 Analysis 1.1 Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gait speed, short term. 128 Analysis 1.2 Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: gait speed, short term. 129 Analysis 1.4 Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.5 Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term. 129 Analysis 1.5 Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gross motor function, intermediate term. 130 Analysis 2.1 Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, intildren and adolescents; intermediate term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; ahort term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; ahort term. 132 Analysis 2.5. Comparison 2 Resistance	Figure 2.	19
Figure 4. 22 Figure 5. 23 DSCUDSION 29 AUTHORS' CONCLUSIONS 32 ACKNOWLEDGEMENTS 33 CHARACTERISTICS OF STUDIES 34 CHARACTERISTICS OF STUDIES 49 DATA AND ANALYSES 128 Analysis 1.1 Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gait speed, short term. 128 Analysis 1.2 Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: gait speed, short term. 129 Analysis 1.4 Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.5 Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term. 129 Analysis 1.5 Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gross motor function, intermediate term. 130 Analysis 2.1 Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, intildren and adolescents; intermediate term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; ahort term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; ahort term. 132 Analysis 2.5. Comparison 2 Resistance	Figure 3.	20
DISCUSSION 29 AUTHORS' CONCLUSIONS 32 REFERENCES 33 REFERENCES 34 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: garts peed, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gaits geed, short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: gaits geed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gross motor function, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gross motor function, children and adolescents; short term. 130 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; intermediate term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; short term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents; short term. 1	Figure 4.	22
AUTHORS' CONCLUSIONS 32 ACKNOWLEDGEMENTS 33 REFERENCES 34 HARACTERISTICS OF STUDIES 49 DATA AND ANALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gait speed, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: walking endurance; short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, nitermediate term. 129 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term. 130 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; short term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.5. Comparison 2 Resistance training	Figure 5.	23
ACKNOWLEDGEMENTS 33 REFERENCES 34 ADRARCTERISTICS OF STUDIES 49 DATA AND ANALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gait speed, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term. 130 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; short term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term. 132	DISCUSSION	29
REFERENCES 34 CHARACTERISTICS OF STUDIES 49 DATA AND ANNALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, alcivity, short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gait speed, alcivity, short term. 130 Analysis 2.1. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic filtness; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term. 132 Analysis 2.6. Comparison 2 Resistance training versus usual	AUTHORS' CONCLUSIONS	32
REFERENCES 34 CHARACTERISTICS OF STUDIES 49 DATA AND ANNALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 128 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, short term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, alcivity, short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gait speed, alcivity, short term. 130 Analysis 2.1. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic filtness; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term. 132 Analysis 2.6. Comparison 2 Resistance training versus usual	ACKNOWLEDGEMENTS	33
CHARACTERISTICS OF STUDIES 49 DATA AND ANALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gross motor function, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: walking endurance; short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gross motor function, intermediate term. 129 Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: daily physical activity; short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term. 130 Analysis 2.1. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term. 131 adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; short term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; short term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; short term. 132 Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term. 132 Analysis 2.7. Com		34
DATA AND ANALYSES 128 Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term. 128 Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term. 129 Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: gait speed, intermediate term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term. 129 Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term. 130 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, adults; short term. 132 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; 131 Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: gait speed, adults; short term. 132 Analysi		
Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term.128Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term.129Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.129Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term.129Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term.130Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and131adolescents; intermediate term.131adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gait speed, children and adolescents;short term.132Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents;short term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents;132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents;133Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents;133Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 9 Partici		128
Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term.128Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.129Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gait speed, intermediate term.129Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gaits peed, intermediate term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term.130Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; short term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; short term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents; short term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, children and adolescents; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: walking endurance, adults; short term.133Analysis 2.6.		128
Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: walking endurance; short term.129Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: gait speed, intermediate term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity: short term.129Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term.130Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term.131Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: gait speed, adults; short term.132Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short term.133Analysis 2.10. Comparison 2		
Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.129Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: daily physical activity; short term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term.130Analysis 2.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, children and131adolescents; short term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and131adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents;131adolescents; intermediate term.132Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents;132intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents;132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents;133term.133134Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), chi	· · · ·	129
Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term.129Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity: short term.130Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gross motor function, adults; short term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short term.133Analysis 2.10. Comparison 2 Resistance training ve	· · ·	129
Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term. 129 Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term. 130 Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term. 131 Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term. 131 Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; intermediate term. 132 Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, adults; short term. 132 Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term. 132 Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term. 133 Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; intermediate term. 133 Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (child-reported), children and adolescents; short term. 133		
Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term.130Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; short term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133		
Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short132Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short133adolescents; short term.134Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; intermediate term.133 <td></td> <td></td>		
adolescents; short term.131Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.131Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; intermediate term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short133Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; term. <td< td=""><td></td><td></td></td<>		
adolescents; intermediate term.Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; short term.Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: walking endurance, adults; short term.Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; shortAnalysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; shortAnalysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; short term.Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; short term.Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; short term.Analysis 2.13. Compar		
Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; short term.131Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gait speed, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short133term.133Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.134Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; short term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; short term.134Analysis 3.1. Comparison 3 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; short term.134Analysis 3.1. Comparison 3 Resista		131
intermediate term.132Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short133term.134Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents;133intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; short term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, children and adolescents; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 14 Muscle strength, adults; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 14 Auscle strength, adults; short term.134Analysis 3.3. Comparis	Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents;	131
Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.132Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.132Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.133Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; short term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.134Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents;134Intermediate term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: walking endurance; short term. <td< td=""><td></td><td>132</td></td<>		132
Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.132Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.133Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short133term.133Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents;133intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.134Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; short term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136	Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.	132
Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short133Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents;133intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and133adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and133adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short133term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents;134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		132
term	Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.	132
intermediate term.133Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; intermediate term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		133
adolescents; short term.133Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; intermediate term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		133
Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.133Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.133Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; intermediate term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		133
Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short133term.Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents;134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		133
intermediate term.134Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136	Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short	133
Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.134Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.135	Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents;	134
Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.135Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		134
Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.135Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.136		
Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term. 136		
	Analysis 3.4. Comparison 3 Mixed training versus usual care, Outcome 4 Participation; short term.	136

Exercise interventions for cerebral palsy (Review)

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Analysis 3.5. Comparison 3 Mixed training versus usual care, Outcome 5 Participation; intermediate term.	136
Analysis 3.6. Comparison 3 Mixed training versus usual care, Outcome 6 Aerobic fitness; short term.	136
Analysis 3.7. Comparison 3 Mixed training versus usual care, Outcome 7 Muscle strength; short term.	136
Analysis 3.8. Comparison 3 Mixed training versus usual care, Outcome 8 Anaerobic fitness; short term.	137
Analysis 3.9. Comparison 3 Mixed training versus usual care, Outcome 9 Aerobic fitness; intermediate term	137
Analysis 3.10. Comparison 3 Mixed training versus usual care, Outcome 10 Anaerobic fitness; intermediate term	137
Analysis 3.11. Comparison 3 Mixed training versus usual care, Outcome 11 Muscle strength; intermediate term.	137
Analysis 4.1. Comparison 4 Resistance training versus aerobic exercise, Outcome 1 Activity: gross motor function; short term.	138
Analysis 4.2. Comparison 4 Resistance training versus aerobic exercise, Outcome 2 Activity: gait speed; short term	138
Analysis 4.3. Comparison 4 Resistance training versus aerobic exercise, Outcome 3 Activity: gait speed; intermediate term	138
Analysis 4.4. Comparison 4 Resistance training versus aerobic exercise, Outcome 4 Activity: gross motor function; intermediate term.	139
Analysis 4.5. Comparison 4 Resistance training versus aerobic exercise, Outcome 5 Muscle strength; short term.	139
Analysis 4.6. Comparison 4 Resistance training versus aerobic exercise, Outcome 6 Muscle strength; intermediate term	139
Analysis 5.1. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.	140
Analysis 5.2. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 2 Activity: gross motor function, intermediate term.	140
Analysis 5.3. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 3 Activity: gait speed; short term.	140
Analysis 5.4. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 4 Activity: walking endurance; short term.	141
Analysis 5.5. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 5 Aerobic fitness; short term	141
Analysis 6.1. Comparison 6 Resistance training and mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.	142
Analysis 6.2. Comparison 6 Resistance training and mixed training versus usual care, Outcome 2 Activity: gross motor function; intermediate term.	142
Analysis 6.3. Comparison 6 Resistance training and mixed training versus usual care, Outcome 3 Activity: gait speed; short term.	143
Analysis 6.4. Comparison 6 Resistance training and mixed training versus usual care, Outcome 4 Participation; short term	143
Analysis 6.5. Comparison 6 Resistance training and mixed training versus usual care, Outcome 5 Participation; intermediate term.	143
Analysis 6.6. Comparison 6 Resistance training and mixed training versus usual care, Outcome 6 Muscle strength; short term.	144
Analysis 6.7. Comparison 6 Resistance training and mixed training versus usual care, Outcome 7 Muscle strength; intermediate term.	144
ADDITIONAL TABLES	144
APPENDICES	145
CONTRIBUTIONS OF AUTHORS	156
DECLARATIONS OF INTEREST	156
SOURCES OF SUPPORT	157
DIFFERENCES BETWEEN PROTOCOL AND REVIEW	157
INDEX TERMS	157



[Intervention Review]

Exercise interventions for cerebral palsy

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ABSTRACT

Background

Cerebral palsy (CP) is a neurodevelopmental disorder resulting from an injury to the developing brain. It is the most common form of childhood disability with prevalence rates of between 1.5 and 3.8 per 1000 births reported worldwide. The primary impairments associated with CP include reduced muscle strength and reduced cardiorespiratory fitness, resulting in difficulties performing activities such as dressing, walking and negotiating stairs.

Exercise is defined as a planned, structured and repetitive activity that aims to improve fitness, and it is a commonly used intervention for people with CP. Aerobic and resistance training may improve activity (i.e. the ability to execute a task) and participation (i.e. involvement in a life situation) through their impact on the primary impairments of CP. However, to date, there has been no comprehensive review of exercise interventions for people with CP.

Objectives

To assess the effects of exercise interventions in people with CP, primarily in terms of activity, participation and quality of life. Secondary outcomes assessed body functions and body structures. Comparators of interest were no treatment, usual care or an alternative type of exercise intervention.

Search methods

In June 2016 we searched CENTRAL, MEDLINE, Embase, nine other databases and four trials registers.

Selection criteria

We included randomised controlled trials (RCTs) and quasi-RCTs of children, adolescents and adults with CP. We included studies of aerobic exercise, resistance training, and 'mixed training' (a combination of at least two of aerobic exercise, resistance training and anaerobic training).

Data collection and analysis

Two review authors independently screened titles, abstracts and potentially relevant full-text reports for eligibility; extracted all relevant data and conducted 'Risk of bias' and GRADE assessments.

Main results

We included 29 trials (926 participants); 27 included children and adolescents up to the age of 19 years, three included adolescents and young adults (10 to 22 years), and one included adults over 20 years. Males constituted 53% of the sample. Five trials were conducted in

Exercise interventions for cerebral palsy (Review)

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the USA; four in Australia; two in Egypt, Korea, Saudi Arabia, Taiwan, the Netherlands, and the UK; three in Greece; and one apiece in India, Italy, Norway, and South Africa.

Twenty-six trials included people with spastic CP only; three trials included children and adolescents with spastic and other types of CP. Twenty-one trials included people who were able to walk with or without assistive devices, four trials also included people who used wheeled mobility devices in most settings, and one trial included people who used wheeled mobility devices only. Three trials did not report the functional ability of participants. Only two trials reported participants' manual ability. Eight studies compared aerobic exercise to usual care, while 15 compared resistance training and 4 compared mixed training to usual care or no treatment. Two trials compared aerobic exercise to resistance training. We judged all trials to be at high risk of bias overall.

We found low-quality evidence that aerobic exercise improves gross motor function in the short term (standardised mean difference (SMD) 0.53, 95% confidence interval (CI) 0.02 to 1.04, N = 65, 3 studies) and intermediate term (mean difference (MD) 12.96%, 95% CI 0.52% to 25.40%, N = 12, 1 study). Aerobic exercise does not improve gait speed in the short term (MD 0.09 m/s, 95% CI -0.11 m/s to 0.28 m/s, N = 82, 4 studies, very low-quality evidence) or intermediate term (MD -0.17 m/s, 95% CI -0.59 m/s to 0.24 m/s, N = 12, 1 study, low-quality evidence). No trial assessed participation or quality of life following aerobic exercise.

We found low-quality evidence that resistance training does not improve gross motor function (SMD 0.12, 95% CI –0.19 to 0.43, N = 164, 7 studies), gait speed (MD 0.03 m/s, 95% CI –0.02 m/s to 0.07 m/s, N = 185, 8 studies), participation (SMD 0.34, 95% CI –0.01 to 0.70, N = 127, 2 studies) or parent-reported quality of life (MD 12.70, 95% CI –5.63 to 31.03, n = 12, 1 study) in the short term. There is also low-quality evidence that resistance training does not improve gait speed (MD –0.03 m/s, 95% CI –0.17 m/s to 0.11 m/s, N = 84, 3 studies), gross motor function (SMD 0.13, 95% CI –0.30 to 0.55, N = 85, 3 studies) or participation (MD 0.37, 95% CI –6.61 to 7.35, N = 36, 1 study) in the intermediate term.

We found low-quality evidence that mixed training does not improve gross motor function (SMD 0.02, 95% CI -0.29 to 0.33, N = 163, 4 studies) or gait speed (MD 0.10 m/s, -0.07 m/s to 0.27 m/s, N = 58, 1 study) but does improve participation (MD 0.40, 95% CI 0.13 to 0.67, N = 65, 1 study) in the short-term.

There is no difference between resistance training and aerobic exercise in terms of the effect on gross motor function in the short term (SMD 0.02, 95% CI – 0.50 to 0.55, N = 56, 2 studies, low-quality evidence).

Thirteen trials did not report adverse events, seven reported no adverse events, and nine reported non-serious adverse events.

Authors' conclusions

The quality of evidence for all conclusions is low to very low. As included trials have small sample sizes, heterogeneity may be underestimated, resulting in considerable uncertainty relating to effect estimates. For children with CP, there is evidence that aerobic exercise may result in a small improvement in gross motor function, though it does not improve gait speed. There is evidence that resistance training does not improve gait speed, gross motor function, participation or quality of life among children with CP.

Based on the evidence available, exercise appears to be safe for people with CP; only 55% of trials, however, reported adverse events or stated that they monitored adverse events. There is a need for large, high-quality, well-reported RCTs that assess the effectiveness of exercise in terms of activity and participation, before drawing any firm conclusions on the effectiveness of exercise for people with CP. Research is also required to determine if current exercise guidelines for the general population are effective and feasible for people with CP.

PLAIN LANGUAGE SUMMARY

Exercise interventions for improving activity, participation and quality of life in people with cerebral palsy

Review question

Does exercise improve activity, participation in life situations and quality of life in people with cerebral palsy (CP)?

Background

Cerebral palsy (CP) is caused by an injury to an infant's brain that interrupts normal development. People with CP have reduced muscle strength and aerobic fitness, which may impact their ability to perform activities such as standing, walking, running and to participate in everyday life. Exercise is defined as a planned, structured and repetitive activity that aims to improve fitness. Aerobic exercise aims to improve aerobic fitness, while strength training aims to improve muscle strength. Health professionals often prescribe exercise to people with CP, primarily to improve function, but there has been no comprehensive evaluation of the evidence for the effectiveness of these interventions in people with CP.

Study characteristics



In June 2016 we searched for all studies that investigated the effectiveness of exercise for people with CP. We included 29 trials with a total of 926 participants with CP, 53% of whom were male. Five trials were conducted in the USA; four in Australia; two in Egypt, Korea, Saudi Arabia, Taiwan, the Netherlands, and the UK; three in Greece; and one apiece in India, Italy, Norway, South Africa.

One trial included only adults with CP and three trials included adolescents and young adults. Most trials included children with CP who could walk independently, with or without a walking aid. Four trials also included people who used wheeled mobility devices (e.g. wheelchairs) in most settings and one trial included people who used wheeled mobility devices only. Three trials did not clearly report participants' functional ability and only two trials reported participants' manual ability (use of hands when handling objects). Eight trials compared aerobic exercise to usual care (i.e. the care a patient usually receives in practice), 15 trials compared resistance training (a type of exercise to improve muscular strength) to either usual care or no treatment, 4 trials compared mixed training (aerobic exercise and resistance training) to usual care or no treatment, and 2 trials compared aerobic exercise to resistance training.

Key results

Aerobic exercise may improve activity as indicated by motor function but does not appear to improve gait speed, walking endurance, participation or aerobic fitness among children with CP in the short or intermediate term. There is no research regarding the effect of aerobic exercise on participation or quality of life.

Resistance training does not appear to improve motor function, gait speed or participation in the short or intermediate term, or quality of life in the short term, in children and adolescents with CP but may improve muscle strength.

Mixed training does not improve motor function or gait speed but does improve participation in children and adolescents with CP in the short term.

We found no difference between aerobic and resistance training on motor function but a difference in muscle strength in the short term.

Although the evidence suggests that exercise might be safe for people with CP, only 16 trials (55%) included information on adverse events; these trials reported no serious adverse events. All of the studies we found had small numbers of participants, meaning that we cannot be sure the results are accurate.

Quality of the evidence

We judged the quality of evidence for all comparisons to be low or very low. All of the studies had small sample sizes. There were very few trials involving adults with CP or people with CP who could not walk, so our results may not apply to these groups of people. Few trials provided clear detail about the frequency, intensity and duration of exercise prescribed. Further research assessing the effectiveness of exercise for activity and participation is needed. Such research should determine if the amount and intensity of exercise prescribed to people with CP has an impact on its effectiveness, and whether current guidelines on exercise for the general population apply to people with CP.

SUMMARY OF FINDINGS

Summary of findings for the main comparison. Aerobic exercise versus usual care

Aerobic exercise versus usual care

Patient or population: children and adolescents with cerebral palsy Intervention: aerobic exercise

Setting: mixed (community, outpatients, home) Comparison: usual care

Outcomes	Anticipated absolute effects* (95% CI)		Relative effect (95% CI)	Number of par- ticipants	Quality of the evidence	Comments	
	Risk with usual care	Risk with aerobic exercise	(95% CI)	(studies)	(GRADE)		
Activity Gross motor func- tion assessed with the Gross Motor Function Measure (follow-up 0 to 1 month)	The mean gross motor function ranged across control groups from 0.20% to 65.13%	The standardised mean gross motor function in the interven- tion group was 0.53 higher (0.02 higher to 1.04 higher)	_	65 (3)	⊕⊕⊝⊝ Low ^{a,c,d}	Higher score indicates im- proved activity A rule of thumb for inter- preting SMD is that 0.2 represents a small effect, 0.5 a moderate effect, and 0.8 a large effect (Cohen 1988)	
Activity Gait speed assessed with a timed walk test (follow-up 0 to 1 month)	The mean gait speed ranged across control groups from 0.63 m/s to 2.40 m/s	The mean gait speed in the in- tervention groups was 0.09 m/s faster (0.11 m/s slower to 0.28 m/s faster)	_	82 (4)	⊕⊙⊙⊙ Very low ^{a,b,c,d}	Higher speed indicates im- proved activity	

GRADE Working Group grades of evidence

High quality: we are very confident that the true effect lies close to that of the estimate of the effect.

Moderate quality: 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 quality: our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low quality: we have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

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Exercise

interventions for cerebral palsy (Review)

^bHeterogeneity statistically significant: P < 0.1, I² > 40%. ^cNumber of participants < 400. ^dWe did not downgrade on the basis of publication bias, as there can be no direct evidence with so few trials for any given intervention.

Summary of findings 2. Resistance training versus usual care

Resistance training versus usual care

Patient or population: children and adolescents with cerebral palsy **Setting**: mixed (home, physiotherapy clinic, school, community gym) **Intervention**: resistance training **Comparison**: usual care

Outcomes	Anticipated absolute	effects* (95% CI)	Relative effect (95% CI)	Number of par- ticipants	Quality of the evidence	Comments
	Risk with usual care	Risk with resistance train- ing	(5576 Cl)	(studies)	(GRADE)	
Activity Gross motor func- tion assessed with the Gross Motor Function Measure (follow-up 0 to 1 month)	The mean gross motor function ranged across con- trol groups from 60.80% to 81.30%	The standardised mean gross motor function in the intervention groups was 0.12 higher (0.19 lower to 0.43 higher)	_	164 (7)	⊕⊕⊙© Lowa,b,c	A rule of thumb for interpreting the SMD is that 0.2 represents a small effect, 0.5 a moderate ef- fect, and 0.8 a large effect (Co- hen 1988) Higher score indicates im- proved activity
Activity Gross motor func- tion assessed with the Gross Motor Function Measure (follow-up > 1 month to 6 months)	The mean gross motor function ranged across con- trol groups from 61.80% to 74.30%	The standardised mean gross motor function in the intervention groups was 0.13 higher (-0.30 lower to 0.55 higher)	_	85 (3)	⊕⊕⊙⊝ Lowa,b,c	A rule of thumb for interpret- ing SMD is that 0.2 represents a small effect, 0.5 a moderate ef- fect, and 0.8 a large effect (Co- hen 1988) Higher score indicates im- proved activity
Activity Gait speed assessed with a timed walk test (follow-up 0 to 1 month)	The mean gait speed ranged across con- trol groups from 0.30 m/s to 1.17 m/s	The mean gait speed in the intervention groups was 0.03 m/s faster (0.02 m/s slower to 0.07 m/s faster)	_	185 (8)	⊕⊕⊙⊙ Lowa,b,c	Higher speed indicates im- proved activity

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Activity	The mean gait speed ranged across con-	The mean gait speed in the intervention groups was	-	84 (3)	⊕⊕⊝⊝ Lowa,b,c	Higher speed indicates im- proved activity
Gait speed assessed with a timed walk test	trol groups from 0.68 m/s to 1.06 m/s	0.03 m/s slower (0.17 m/s slower to 0.11 m/s faster)				
(follow-up > 1 month to 6 months)						
Participation	The mean participa- tion in the control	The standardised mean par- ticipation in the interven-	_	127	⊕⊕⊝⊝	A rule of thumb for interpret- ing SMD is that 0.2 represents a
Assessed with various neasures	group ranged from 7.40 to 31.14	tion groups was 0.34 higher (0.01 lower to 0.70 higher)		(2)	Low ^{a,b,c}	small effect, 0.5 a moderate ef- fect, and 0.8 a large effect (Co- hen 1988)
follow-up 0 to 1 nonth)						Higher score indicates im- proved participation
The risk in the interv	antion enough (and its OFO)				ative effect of the	intervention (and its 95% CI).
CI: confidence interval; GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confid	SMD: standardised mean o grades of evidence ery confident that the true are moderately confident i dence in the effect estimat	difference. effect lies close to that of the es	stimate of the effect ffect is likely to be o be substantially dif	close to the estimat ferent from the esti	e of the effect, but mate of the effect.	there is a possibility that it is sub-
CI: confidence interval GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confid /ery low quality: we h	SMD: standardised mean o grades of evidence ery confident that the true are moderately confident i dence in the effect estimat ave very little confidence i of bias because it is not po < 400.	difference. effect lies close to that of the es in the effect estimate: the true e e is limited: the true effect may	stimate of the effect iffect is likely to be of be substantially dif iffect is likely to be s rticipants to group a	close to the estimat ferent from the esti substantially differe allocation.	e of the effect, but mate of the effect. nt from the estima	there is a possibility that it is sub-
CI: confidence interval GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confid /ery low quality: we h Ill trials are at high risk Jumber of participants /e did not downgrade	SMD: standardised mean o grades of evidence ery confident that the true are moderately confident i dence in the effect estimat ave very little confidence i of bias because it is not po < 400.	difference. effect lies close to that of the es in the effect estimate: the true e re is limited: the true effect may in the effect estimate: the true e possible to blind personnel or pa bias, as there can be no direct e	stimate of the effect iffect is likely to be of be substantially dif iffect is likely to be s rticipants to group a	close to the estimat ferent from the esti substantially differe allocation.	e of the effect, but mate of the effect. nt from the estima	there is a possibility that it is sub-
CI: confidence interval GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confie Very low quality: we h All trials are at high risk Number of participants Ve did not downgrade of ummary of findings	SMD: standardised mean ogrades of evidence ery confident that the true are moderately confident i dence in the effect estimat ave very little confidence i of bias because it is not po < 400. on the basis of publication 3. Mixed training vers	difference. effect lies close to that of the es in the effect estimate: the true e re is limited: the true effect may in the effect estimate: the true e possible to blind personnel or pa bias, as there can be no direct e	stimate of the effect iffect is likely to be of be substantially dif iffect is likely to be s rticipants to group a	close to the estimat ferent from the esti substantially differe allocation.	e of the effect, but mate of the effect. nt from the estima	there is a possibility that it is sub-
CI: confidence interval GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confid Very low quality: we h All trials are at high risk Jumber of participants Ve did not downgrade of Mixed training versus Patient or population Setting: mixed (school Intervention: mixed tr	SMD: standardised mean ogrades of evidence ery confident that the true are moderately confident i dence in the effect estimat ave very little confidence i of bias because it is not po < 400. on the basis of publication 3. Mixed training vers usual care : children and adolescents , home) aining	difference. effect lies close to that of the es in the effect estimate: the true e e is limited: the true effect may in the effect estimate: the true e ossible to blind personnel or pa bias, as there can be no direct e	stimate of the effect iffect is likely to be of be substantially dif iffect is likely to be s rticipants to group a	close to the estimat ferent from the esti substantially differe allocation.	e of the effect, but mate of the effect. nt from the estima	there is a possibility that it is sub-
CI: confidence interval GRADE Working Group High quality: we are ve Moderate quality: we stantially different. Low quality: our confit Very low quality: we h All trials are at high risk Number of participants We did not downgrade of ummary of findings Mixed training versus	SMD: standardised mean ogrades of evidence ery confident that the true are moderately confident i dence in the effect estimat ave very little confidence i of bias because it is not po < 400. on the basis of publication 3. Mixed training vers usual care : children and adolescents , home) aining	difference. effect lies close to that of the estinate: the effect estimate: the true effect may n the effect estimate: the true effect may n the effect estimate: the true effect stimate: the true effect estimate: the true effect estimate of the effect	stimate of the effect iffect is likely to be of be substantially dif iffect is likely to be s rticipants to group a	close to the estimat ferent from the esti substantially differe allocation.	e of the effect, but mate of the effect. nt from the estima	there is a possibility that it is sub-

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Activity Gross motor function assessed with the Gross Motor Func- tion Measure	function in the controlgrogroups ranged frominte30.76% to 90.11%0.0	standardised mean — ss motor function in the rvention groups was 2 higher (0.29 lower to 8 higher)	163 (4)	⊕⊕⊙⊝ Lowa,b,c	ing SMD is the small effect, (fect, and 0.8 a hen 1988)	nb for interpret- at 0.2 represents a 0.5 a moderate ef- a large effect (Co-
(follow-up 0 to 1 month)					Higher score proved activi	
	r ention group (and its 95% CI) is ; SMD : standardised mean differ	based on the assumed risk in the co rence.	omparison group and the	relative effect of t	he intervention (an	nd its 95% CI).
Moderate quality: we stantially different. Low quality: our confi Very low quality: we h All trials are at high risk Number of participants We did not downgrade ummary of findings	are moderately confident in the dence in the effect estimate is li lave very little confidence in the cof bias because it is not possible < 400.	t lies close to that of the estimate of effect estimate: the true effect is lik mited: the true effect may be substa effect estimate: the true effect is lik e to blind personnel or participants , as there can be no direct evidence of sus aerobic exercise	ely to be close to the esti ntially different from the ely to be substantially dif to group allocation.	estimate of the effe ferent from the estin	ct.	pility that it is sub-
	: children with cerebral palsy					
Setting: home or not r Intervention: resistan Comparison: aerobic e	eported ce training					
Outcomes	Anticipated absolute effects	* (95% CI)	Relative effect (95% CI)	Number of par- ticipants	Quality of the evidence	Comments
	Risk with aerobic exercise	Risk with resistance training	(95% CI)	(studies)	(GRADE)	
Activity Gross motor function assessed with vari- ous measures	The mean gross motor func- tion in the aerobic exercise groups ranged from 44.09% to 63.30%	The standardised mean gross mo function in the intervention grou was 0.02 higher (0.50 lower to 0 higher)	ips	56 (2)	⊕⊕⊙⊙ Lowa,b,c	Higher score indicates im- proved activity

assessed with various measures

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(follow-up 0 to 1 month)

*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). CI: confidence interval; SMD: standardised mean difference.

GRADE Working Group grades of evidence

High quality: we are very confident that the true effect lies close to that of the estimate of the effect.

Moderate quality: 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 quality: our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low quality: we have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

^{*a*}All trials are at high risk of bias because it is not possible to blind participants or personnel to group allocation.

^bNumber of participants < 400.

^cWe did not downgrade on the basis of publication bias, as there can be no direct evidence with so few trials for any given intervention.

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BACKGROUND

Description of the condition

Cerebral palsy (CP) is defined as "a group of permanent disorders of the development of movement and posture, causing activity limitation, that are attributed to non-progressive disturbances that occurred in the developing fetal or infant brain" (Rosenbaum 2007, p 11). Children with CP may also present with cognitive impairments, hearing and visual impairments, communication difficulties and epilepsy (Rosenbaum 2007). Most children with CP are diagnosed at around one to two years of age (Ashwal 2004; Herskind 2015), following a medical history and physical examination that identify a non-progressive motor deficit (Ashwal 2004; Rosenbaum 2007). Neuroimaging techniques, preferably magnetic resonance imaging (MRI), may be used in conjunction with the history and physical examination to establish aetiology and prognosis (Ashwal 2004).

CP is the most common form of childhood disability, with reported prevalence rates of between 1.5 and 3.8 per 1000 live births in different areas of Europe and the USA (SCOPE 2002; Kirby 2011). The prevalence of CP varies not only by geographical location but also by birth weights and gestational age, with higher prevalence rates reported in children born preterm or at low birth weight (Platt 2007; Sellier 2010; Andersen 2011). Other factors associated with CP include multiple births, maternal infection during pregnancy, having a relative with CP, breech position and placental abruption (O'Callaghan 2011; Tollånes 2014; Trønnes 2014). The prevalence of severe CP in Europe, defined by an inability to walk and a severe intellectual disability, is approximately 0.43 per 1000 live births (SCOPE 2002). Children without severe impairments are expected to live well into adulthood (Strauss 1998a; Blair 2001; Brooks 2014). Although less is known about the life expectancy of adults with CP, evidence suggests that adults with CP who maintain a high level of function have a slightly lower life expectancy than the general population (Strauss 1998b; Brooks 2014).

The primary impairments associated with CP include reduced muscle strength (Riad 2012; Nooijen 2014), reduced cardiorespiratory fitness (Verschuren 2010; Nieuwenhuijsen 2011; Nooijen 2014), and poor selective motor control (Østensjø 2004). As a result of these impairments, people with CP may have difficulty performing everyday activities such as eating, dressing, walking, running, jumping and negotiating stairs (Østensjø 2004; Ross 2007; Opheim 2009; Klingels 2012). Intensive rehabilitation is often provided in childhood to improve gross motor function. Indeed, many children who are non-ambulatory at age two to three years will be ambulatory by the time they reach adolescence (Wu 2004). About 54% of five-year-old children in Europe and 56% of eight-year-old children in the USA are independently ambulatory despite having CP (Beckung 2008; Kirby 2011). However, a subsequent decline in gross motor function often occurs in adolescence and young adulthood (Bottos 2001; Sandström 2004; Hanna 2009; Kerr 2011). Up to 50% of adults with CP report experiencing deterioration in walking function from young adulthood (Bottos 2001; Opheim 2009). Adults with CP attribute deterioration in walking function to reduced muscle strength, reduced cardiorespiratory fitness, fatigue and pain (Jahnsen 2004; Opheim 2009). Conversely, adults who experience improvements or no change in walking function over time credit this to improvements in balance, muscle strength and cardiorespiratory fitness (Opheim 2009). Poor gross motor function may also contribute to reduced quality of life and unemployment, which is high among young adults with CP (Soyupek 2010; Verhoef 2014).

Although CP is defined by the presence of motor disorders, the clinical presentation of CP can vary considerably, making it difficult to compare individuals at one point in time or to evaluate changes in an individual's condition over time. Traditionally, CP has been classified according to the type of motor abnormality (for example, spasticity, dystonia, choreoathetosis, ataxia) and anatomical distribution of CP (for example, bilateral, unilateral) (Rosenbaum 2007). More recently, classification systems that allow categorisation of people with CP according to their level of functional impairment have been developed; the Gross Motor Function Classification System (GMFCS) and Manual Ability Classification System (MACS) are two such systems. The GMFCS is a five-point scale that distinguishes between levels of motor function based on functional mobility and the need for assistive technology, particularly mobility aids (Palisano 1997; Palisano 2008). A full description of the GMFCS is presented in Appendix 1. To summarise, from six years of age children in level I of the GMFCS are able to walk indoors and outdoors without assistance and can perform gross motor skills such as running and jumping; children in level II can also walk indoors and outdoors without assistance but have only minimal ability to perform gross motor skills like running and jumping; children in level III require a mobility device to walk indoors and outdoors and may require wheeled mobility for travelling long distances; children in level IV use wheeled mobility in most settings; children in level V are limited in their ability to maintain antigravity head and trunk postures and to control arm and leg movements, and they are transported in a manual wheelchair in all settings. Although developed for children with CP, the GMFCS has been used successfully to classify motor function in adults with CP (Sandström 2004). For its part, the MACS is a five-point scale that classifies how children aged four years or older with CP use their hands when handling objects in daily activities (Eliasson 2006). A full description of the MACS is in Appendix 2. Children in level I of the MACS handle objects easily and successfully. They may have limitations in the ease of performing tasks that require speed and accuracy. Children in level II handle most objects but with reduced quality, speed or both. Children in level III have difficulty handling objects and need help to prepare or modify activities. Children in level IV can only handle a limited selection of easily managed objects in adapted situations and require continuous support and assistance.

Description of the intervention

Exercise is defined as "physical activity that is planned, structured, repetitive, and purposive in the sense that improvement or maintenance of one or more components of physical fitness is an objective" (Caspersen 1985, p 128). The components of physical fitness that exercise may improve include muscle strength, muscle endurance and cardiorespiratory fitness. The focus of this review will be on exercise interventions categorised as resistance training or aerobic training. Resistance training involves the body's muscles working or holding against an applied force. Body weight, free weights, machine weights, and elastic bands are often used to apply force (USDHHS 2008). Current guidelines for resistance training to improve muscle strength for youth suggest that one to three sets of 6 to 15 repetitions of a muscle strengthening exercise should be performed at an intensity of 50% to 85% of one repetition maximum (RM) (i.e. the maximum weight a person can lift with one repetition)

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(Faigenbaum 2009). Alternatively, if people do not perform one RM tests, therapists can establish the intensity by prescribing a repetition range and determining the maximum load that people can lift for the prescribed range. Current guidelines for adults suggest that in order to improve muscle strength, inexperienced people should perform one to three sets of 8 to 12 repetitions of a muscle strengthening exercise at loads corresponding to 60% to 70% of one RM (American College of Sports Medicine 2009). People should engage in resistance training on two to three days per week (American College of Sports Medicine 2009; Faigenbaum 2009), and at least eight weeks of training are required to observe an increase in muscle strength (Faigenbaum 2009). Aerobic training involves moving the body's large muscles in a rhythmic manner for a sustained period of time (USDHHS 2008). Walking, running, cycling and arm ergometry are examples of aerobic exercise. Current guidelines for aerobic exercise to improve cardiorespiratory fitness suggest that people with CP should engage in aerobic exercise two to three times per week at an intensity of 60% to 95% of peak heart rate, between 40% to 80% of heart-rate reserve (HRR) or between 50% and 65% of VO₂ peak (i.e. maximum oxygen consumption), for at least 20 minutes per session (Verschuren 2016). Further, a training programme should continue for at least 8 consecutive weeks when training three times a week or for 16 consecutive weeks when training twice a week (Verschuren 2016). Many exercise programmes target muscle strength, anaerobic fitness, cardiorespiratory fitness or a combination of these components. We will refer to such programmes as 'mixed training'.

How the intervention might work

The goal of treatment for people with CP has shifted from targeting impairments of the motor system to targeting activity limitations and participation restriction, where activity is defined as a person's ability to execute a task, and participation is defined as a person's involvement in a life situation (WHO 2001). Indeed, people with CP have identified improving restricted mobility and poor upper limb function as primary therapeutic goals (Vargus-Adams 2011). However, many experts believe there is an association between motor impairments, activity limitation and participation restriction, so targeting one may well affect another. There is evidence that impairments, particularly muscle strength, are associated with activity in children with CP (Østensjø 2004; Ross 2007; Voorman 2007; Verschuren 2009; Klingels 2012; Park 2013). Although less information is available about the association between cardiorespiratory fitness and activity, aerobic training, resistance training and mixed training have proven efficacy on activity in older adults (Liu 2009; Giné-Garriga 2014), a population who experience similar declines in physical functioning as young adults with CP (Nusselder 2005; Day 2007). Improvements in activity provided by exercise may translate to improved participation in mobility-based behaviours for people with CP (Park 2013; Bjornson 2014).

Exercise may also have benefits in terms of pain relief and quality of life for people with CP. Some adults report using exercise as a treatment for pain and find it moderately effective (Engel 2002; Hirsh 2011), which may positively impact on quality of life. Further, a positive association between physical activity and physical, behavioural, emotional and social quality of life has been reported in children with CP (Bjornson 2008; Maher 2016). As exercise is structured physical activity, the implementation of an exercise programme may result in improvements in quality of life for people with CP.

The aim of this review, to assess the effects of exercise interventions on activity, participation and quality of life in people with CP, reflects the goals of people with CP and their clinicians and therefore is of most interest to users of this review. While the association between physical fitness and activity suggests that improving physical fitness may improve activity, the physiological, biomechanical, and neuromuscular adaptations that may occur as a result of exercise training in people with CP are not understood. It is also possible that the effect of exercise on activity performance may vary according to the person's baseline level of functional ability. For example, improving muscle strength in children in GMFCS level III, who have reduced muscle strength compared to children in GMFCS level I (Eek 2008), may result in greater improvements in activity because of their greater potential for improvement. Conversely, improvements in muscle strength may be small in people with a greater degree of functional impairment because of their inability to exercise at an adequate intensity.

Why it is important to do this review

Although CP begins in childhood, it impacts the individual's whole life course as well as the healthcare system. Identifying appropriate interventions to alleviate disability throughout the life of a person with CP is urgent. Health professionals often recommend exercise for people with CP, partly because of its known importance for improving physical functioning in other populations. This is reflected in the growing number of publications on the topic of exercise in CP.

Ten reviews have investigated the effectiveness of exercise interventions in children with CP (Dodd 2002; Taylor 2005; Anttila 2008; Mockford 2008; Rogers 2008; Verschuren 2008; Scianni 2009; Butler 2010; Novak 2013; Rameckers 2014). Eight of these included articles published up to July 2008 (Dodd 2002; Taylor 2005; Anttila 2008; Mockford 2008; Rogers 2008; Verschuren 2008; Scianni 2009; Butler 2010); one included articles up to December 2012 (Novak 2013), and one included articles published up to August 2014 (Rameckers 2014). Four reviews focused solely on randomised controlled trials (RCTs) (Anttila 2008; Scianni 2009; Butler 2010; Rameckers 2014); the remaining six included experimental or quasi-experimental studies. Nine reviews provided a narrative summary of the evidence (Dodd 2002; Taylor 2005; Anttila 2008; Mockford 2008; Rogers 2008; Verschuren 2008; Butler 2010; Novak 2013; Rameckers 2014). Only one review conducted a meta-analysis of RCTs (Scianni 2009). However, this review specifically examined the effectiveness of muscle strengthening, rather than all exercise interventions, in children with CP.

Two reviews have investigated the effectiveness of exercise interventions in adults with CP (Dodd 2002; Jeglinsky 2010). These reviews included observational studies published up to March 2002 and 2009, respectively. Both reviews conducted descriptive analyses. One meta-analysis specifically investigated the effect of strength training in children and adults with CP (Park 2014b).

An up-to-date and comprehensive assessment of the evidence surrounding exercise interventions in adults and children with CP is required to guide consumers, health professionals and policymakers.

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OBJECTIVES

To assess the effects of exercise interventions in people with CP, primarily in terms of activity, participation and quality of life. Secondary outcomes assessed body functions and body structures. Comparators of interest were no treatment, usual care or an alternative type of exercise intervention.

METHODS

Criteria for considering studies for this review

Types of studies

Randomised controlled trials (RCTs) and quasi-RCTs (where sequence generation is systematically determined but not truly random, for example, based on order of entry or date of birth).

Types of participants

Children, adolescents and adults of any age with a diagnosis of CP, irrespective of level of functional ability (i.e. Gross Motor Function Classification System (GMFCS) levels I to V and the Manual Ability Classification System (MACS) levels I to IV).

Types of interventions

We included studies of exercise that met the definition in Caspersen 1985 (see Description of the intervention). We included studies of aerobic and resistance training and studies that used a combination of exercises, where at least one exercise was categorised as resistance training, aerobic training or anaerobic training (that is, 'mixed training'). We included interventions that targeted both the upper and lower limbs. We did not include studies of stretching interventions. We did not include studies of interventions, such as constraint-induced movement therapy or bimanual therapy, where the intervention did not specifically target one or more components of physical fitness (i.e. muscle strength, muscle endurance and cardiorespiratory fitness).

Comparisons of interest were exercise versus no treatment, usual care or an alternative exercise intervention (e.g. a comparison of resistance training and aerobic exercise).

Types of outcome measures

Primary outcomes

- 1. Activity, defined as a person's ability to execute a task (WHO 2001). Examples of outcome measures for activity include the Gross Motor Function Measure 66- or 88-item (GMFM-66 or GMFM-88; Russell 1989), Assisted Hand Assessment (AHA) (Krumlinde-Sundholm 2003), timed walk tests, Melbourne Assessment of Unilateral Upper Limb Function (MAUULF; Randall 1999), ABILHAND-Kids questionnaire (Arnould 2004), Activities Scale for Kids (ASK; Young 2000), International Physical Activity Questionnaire (IPAQ; Craig 2003), accelerometers, and pedometers. Subdomains of activity are:
 - a. activity capacity (i.e. a person's ability to execute a task in a standardised environment);
 - b. activity capability (i.e. a person's ability to execute a task in his or her daily environment); and
 - c. activity performance (i.e. what a person actually does in his or her environment) (Holsbeeke 2009).

- 2. Participation, defined as a person's involvement in a life situation. This may include participation in domestic life (e.g. acquiring a place to live or managing a household); employment or education; and community, social, and civic life (WHO 2001). Examples of outcome measures for participation include the Paediatric Evaluation of Disability Inventory (PEDI; Haley 1992), the Waisman Activities of Daily Living Scale (W-ADL; Maenner 2013), and Assessment of Life Habits questionnaire (LIFE-H; Fougeyrollas 1998).
- 3. Quality of life, defined as the impact of disease and treatment on physical, psychological and social functioning (Schipper 1996; Solans 2008), as measured by, for example, the Short Form-36 (SF-36) health survey (Ware 1993) and the Child Health Questionnaire (CHQ; Landgraf 1998).
- 4. Incidence and nature of adverse events such as injury, cardiac events, stiffness and delayed onset muscle soreness, where reported.

Secondary outcomes

- 1. Body functions and body structures, defined as changes in physiological systems or in anatomical structures (WHO 2001). These include:
 - a. muscle strength and endurance, as measured by, for example, dynamometry;
 - b. cardiorespiratory fitness, as measured by, for example, the Shuttle Run Test (SRT; Verschuren 2006);
 - c. pain, as measured by, for example, a visual analogue scale (VAS) (McCormack 1988);
 - d. fatigue, as measured by, for example, the Fatigue Severity Scale (Krupp 1989); and
 - e. depression, as measured by, for example, the Center for Epidemiological Studies Depression Scale (CES-D; Radloff 1977).

As studies assessed change in a wide range of body structures and functions following exercise, we limited the included outcomes to those targeted by a specific exercise intervention. For example, for studies of resistance training, we reported the effect on muscle strength; for studies of aerobic exercise we reported the effect on aerobic fitness; for studies of mixed training, we reported the effect on muscle strength and aerobic and anaerobic fitness.

We planned to include studies that used any validated scale that measures these primary and secondary outcomes. However, as trials used a range of outcome measures to assess these outcomes, we included any measure that purported to assess them, regardless of whether or not it was validated specifically in people with CP. See Differences between protocol and review. We collected outcomes for the following time points: short term (zero to one month postintervention), intermediate term (more than one month and up to six months' postintervention), and long term (more than six months' postintervention). We presented all available results for the primary outcomes in 'Summary of findings' tables.

Search methods for identification of studies

Electronic searches

We searched all available years of the following databases in June 2016.

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- 1. Cochrane Central Register of Controlled Trials (CENTRAL; 2016, Issue 5), in the Cochrane Library, which contains the Cochrane Developmental, Psychosocial and Learning Problem Specialised Register (searched 14 June 2016).
- 2. Ovid MEDLINE (1946 to June Week 1 2016).
- 3. Embase Ovid (1980 to 2016 Week 24).
- 4. CINAHL Plus EBSCOhost (Cumulative Index to Nursing and Allied Health Literature; 1937 to 14 June 2016).
- 5. Cochrane Database of Systematic Reviews (CDSR; 2016, Issue 6), part of the Cochrane Library.
- 6. Database of Abstracts of Reviews of Effects (DARE; 2015, Issue 2), part of the Cochrane Library (searched 15 May 2015; DARE was not updated after this date).
- 7. Science Citation Index Web of Science (1970 to 9 June 2016).
- 8. Conference Proceedings Citation Index Science Web of Science (CPCI-S; 1990 to 9 June 2016).
- 9. LILACS (Latin American and Caribbean Health Science Information database; lilacs.bvsalud.org/en; searched 14 June 2016).
- 10.Health Services Research Projects in Progress (HSRPRoj; wwwcf.nlm.nih.gov/hsr_project/home_proj.cfm; searched 23 June 2016).
- 11.OpenGrey (www.opengrey.eu; searched 16 June 2016).
- 12.National Rehabilitation Information Center (www.naric.com; searched 23 June 2016).
- 13.PEDro (Physiotherapy Evidence Database; www.pedro.org.au; searched 23 June 2016).
- 14.UKCRN Study Portfolio (public.ukcrn.org.uk/search; searched 16 June 2016).
- 15.ClinicalTrials.gov (clinicaltrials.gov; searched 16 June 2016).
- 16.World Health Organization International Clinical Trials Registry Platform (WHO ICTRP; www.who.int/ictrp/en; searched 20 June 2016).

We used the search strategy for MEDLINE Ovid, which incorporates the Cochrane highly sensitive search strategy for identifying randomised trials (Lefebvre 2011), and we adapted this strategy, as appropriate, for other sources. We did not limit searches by language, date or publication status. The strategy for each source is reported in Appendix 3. For a detailed record of the searches (including search dates and the number of records found in each source), see Appendix 4.

Searching other resources

We handsearched the reference lists of eligible trials and relevant systematic reviews identified from the search to identify additional studies.

Data collection and analysis

Selection of studies

Two review authors (JMR and EEC) independently checked the titles and abstracts of the search results and excluded studies that did not meet the inclusion criteria outlined above (Criteria for considering studies for this review). In cases that appeared to meet the inclusion criteria, or where there was any doubt as to whether we should have excluded the report, we retrieved the full text of the report. Two review authors (JMR and EEC) independently reviewed these papers against the inclusion criteria (Criteria for considering

studies for this review), resolving any disagreements regarding the exclusion of a report at any stage through discussion, and where necessary, through consultation with a third review author (SN). We recorded our decisions in a PRISMA diagram (Moher 2009).

Data extraction and management

Two review authors (JMR and EEC) independently extracted data using a standardised form developed for the purpose. We resolved disagreements regarding the extraction of data by discussion. If we could not reach a resolution, we consulted a third review author (NEO'C). The form included the following information, where available.

- 1. Country of origin.
- 2. Study design.
- 3. Sample size: treatment and control groups.
- 4. Study population (treatment and control groups): sex, age, ethnicity, distribution of CP, type of motor abnormality and gross motor function. Where sufficient information was provided, we classified children and adults according to GMFCS level and MACS level, as these scales provide a comprehensive indication of functional ability above that provided by classifying individuals according to type of motor abnormality and anatomical distribution of CP. Although we proposed to classify general gross motor function as unaided walking, walking with aids or unable to walk (Beckung 2008), most studies reported the GMFCS level of participants. Therefore, we reported the GMFCS level where available and use of mobility aids when the GMFCS level was not available.
- 5. Intervention: aim of the intervention, type of exercise programme (e.g. aerobic exercise), mode of delivery (e.g. home programme), type(s) of location(s) where the intervention occurred (including any necessary infrastructure or relevant features), supervised or unsupervised programme, exercise mode (e.g. cycle ergometry, treadmill), exercise dose (i.e. duration, intensity, and frequency of exercise), tailoring of intervention to individual, modification of intervention (what, why, when, how), duration of programme. Following data extraction, we combined information on modification of the intervention to individual to create one category.
- 6. Intervention provider: profession, expertise, background, specific training received.
- 7. Fidelity or adherence to programme: how or by whom this was assessed.
- 8. Outcome measures (Types of outcome measures).
- 9. Results: short-term (zero to one month postintervention), intermediate-term (greater than one month to six months' postintervention), and long-term (more than six months' postintervention) follow-up.
- 10. Measures of adherence to the exercise programme.
- 11.Adverse effects.
- 12.Conflicts of interest.
- 13.Declarations of conflicts of interest.

14.Sources of funding.

Assessment of risk of bias in included studies

Two review authors (JMR and EEC) independently assessed risk of bias using Cochrane's tool for assessing risk of bias

Exercise interventions for cerebral palsy (Review)

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(Higgins 2011a). A third review author (SN) resolved any persistent disagreements between them. We assessed the following domains: random sequence generation, allocation concealment, blinding of participants and personnel, blinding of outcome assessment, incomplete outcome data, selective reporting and other sources of bias. We scored all domains as being at low, high or unclear risk of bias. We present operational definitions for making judgements on each domain in Appendix 5.

We assigned included studies an overall rating of high, low or unclear risk of bias. Where we rated one or more domains at high risk of bias, we rated the study at high risk of bias overall. Where we did not rate a study at high risk of bias for any domain but rated it at unclear risk of bias for one or more domains, we rated that study at unclear risk of bias overall. We rated a study at low risk of bias overall if we rated it as low risk of bias for all domains.

Measures of treatment effect

Dichotomous data

No study used dichotomous outcomes. Table 1 outlines our plans for dealing with such data and other methodological decisions that were not possible or appropriate to deploy, should it be necessary to use these methods in future updates of this review. Please also see our protocol (Ryan 2015).

Continuous data

Where pooled studies used the same scale on a continuous outcome measure, we presented the effect size as a mean difference (MD) with 95% confidence intervals (CI). Where studies used different scales to measure the same construct or used different versions of an outcome measure that scored the outcome differently, we presented the standardised mean difference (SMD) with 95% CI. We used a rule of thumb to interpret the magnitude of effect for the SMD: 0.2 represents a small effect, 0.5 a moderate effect, and 0.8 a large effect (Cohen 1988).

Clinically important differences

Clinically important differences have been developed for a number of outcome measures (for example, GMFM, weeFIM, one-minute walk test) for ambulatory children and adolescents (GMFCS levels I to III) aged 4 to 19 years old (Oeffinger 2008; Hassani 2014). However, there are no well-established and accepted thresholds for clinically important differences across the range of possible outcome measures and possible participants. Where possible, our discussion of the results considered the size of effects for our primary outcomes in light of contemporary research literature on clinically important differences.

Unit of analysis issues

Cluster-randomised trials

We did not include any cluster-randomised trials in this review. See our protocol, Ryan 2015, and Table 1 for details of methods archived for use in future updates of this review.

Cross-over trials

We identified only one cross-over trial from which we were unable to include any data. See Ryan 2015 and Table 1 for details of methods archived for use in future updates of this review.

Studies with multiple treatment groups

Where studies included multiple treatment groups, we combined results across all eligible intervention groups and compared them with the combined results across all eligible control groups, making single pairwise comparisons.

Dealing with missing data

Where the report of an included study presented insufficient data to enter into the meta-analysis, we requested access to missing data from the authors with two reminder requests sent at monthly intervals in the event of non-response. We specifically requested data relating to the effect of the intervention (e.g. means and standard deviations (SDs)) on any of the outcomes of interest (e.g. adverse events) and details of dropouts. We did not routinely request other methodological details or information relating to the 'Risk of bias' assessments. See Ryan 2015 and Table 1 for details of methods archived for use in future updates of this review. We conducted analyses using only the available data; we did not impute missing data.

Assessment of heterogeneity

We assessed clinical variation across studies by comparing the distribution of important factors among trials (for example, participant age, sex, and functional ability (GMFCS level), characteristics of the interventions). We assessed statistical heterogeneity and its impact using the Chi² test and the I² statistic (Deeks 2011). We used the Chi² test to determine whether differences in effects across studies are compatible with chance alone and the I² statistic to describe the percentage of the variability in effect estimates that was due to heterogeneity rather than sampling error (chance).

Assessment of reporting biases

We considered the possible influence of publication and smallstudy biases on review findings. Where we identified sufficient data (equal to or greater than 10 studies in a meta-analysis), we examined funnel plots and used the test proposed by Egger 1997 to test for funnel plot asymmetry.

Data synthesis

We pooled the results from included studies using Review Manager 5 (RevMan 5) software (Review Manager 2014). Comparisons of interest were exercise versus no treatment or usual care, and comparisons of one type of exercise intervention versus another. We did not pool data from these two comparisons together in a single meta-analysis. We believe that the effect sizes for each of these comparisons are likely to vary considerably and that it is not theoretically justifiable to include exercise and usual care in one comparison group. Where studies compared two types of exercise interventions, we interpreted and discussed the results in the context of the evidence, or lack of evidence, of the effectiveness of each exercise intervention compared to usual care or no treatment.

We attempted to deal with clinical heterogeneity by performing separate meta-analyses for each category of exercise intervention (i.e. resistance training, aerobic training and mixed training). We believe that the type of exercise performed could impact the effect size and that combining these interventions could mask the true effect of each individual intervention. We performed separate meta-analyses for studies in children and adolescents

Exercise interventions for cerebral palsy (Review)

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versus adults. We defined children as aged 19 years and below, adolescents as individuals aged 10 to 19 years inclusive, and adults as aged 20 years and older (WHO 2013). We report the results for adults and for children and adolescents as a group as the majority of studies included children and adolescents, as opposed to children or adolescents only.

We used a random-effects model to combine studies since we expected studies to vary somewhat in terms of the interventions, comparisons and populations. We considered separate metaanalyses for different types of exercise intervention and for shortterm (zero to one month postintervention), intermediate-term (more than one month to six months' postintervention), and long-term (more than six months' postintervention) outcomes. Where meta-analyses were not possible, we conducted a narrative synthesis of the data. We also considered contextual data in our interpretation of the evidence.

There was large variation between studies in terms of the muscle groups targeted by interventions and the muscle groups whose strength investigators assessed as an outcome measure. We therefore applied the following rules for extracting outcome data to be included in the pooled analysis. Where resistance training targeted one muscle group in the lower limbs, we extracted data on a continuous scale for the targeted muscle group. Where resistance training interventions targeted multiple muscle groups in the lower limbs and assessed the strength of multiple muscle groups, we extracted data on a continuous scale for the knee extensors. We chose to extract data on the knee extensors because the knee extensors were the most commonly trained and assessed muscle group. Where exercises targeted right and left limbs and presented data on each limb individually, we took the data from the right limb. Where concentric and eccentric muscle strength was trained and assessed, we took data on concentric muscle strength, as this was more common. Where data on muscle strength at multiple speeds was presented, we took data for muscle strength assessed at 60°/ second as this was the most consistently assessed speed across trials. Where trials of lower limb resistance training, aerobic training or mixed training (that included lower limb resistance training) assessed activity using the GMFM-88 or GMFM-66, we extracted the combined score (%) for dimensions D and E (i.e. standing, walking, running and jumping), as studies often only assessed activities in dimensions D and E. If this was not available, we extracted the individual score (%) for dimension E (i.e. walking, running and jumping), and if this was not available, we extracted the total score (%) for the GMFM-88 or GMFM-66.

Subgroup analysis and investigation of heterogeneity

Due to the small number of trials that could be included in each meta-analysis, we did not conduct subgroup analysis. See Ryan 2015 and Table 1 for our published strategy for subgroup analysis and investigation of heterogeneity.

Sensitivity analysis

We assessed the influence of our analysis model by reanalysing data using a fixed-effect model instead of a random-effects model. Other planned sensitivity analyses were not possible. Please see Ryan 2015 and Table 1 for our published strategy for exploring the impact of studies at high risk of bias due to missing data, and the influence of using imputed correlation coefficients in meta-analyses including cross-over and cluster trials.

Summary of findings table

Two authors (JMR, EEC) used the GRADE approach to assess the quality of the body of evidence (Guyatt 2008). To ensure consistency of GRADE judgements, we applied the criteria below to each domain equally for all key comparisons.

- Limitations of studies: downgrade once if less than 75% of included studies are at low risk of bias across all 'Risk of bias' domains.
- 2. Inconsistency: downgrade once if heterogeneity is statistically significant (P < 0.10) and $I^2 > 40\%$.
- 3. Indirectness: downgrade once if more than 50% of the participants are outside the target group.
- 4. Imprecision: downgrade once if fewer than 400 participants for continuous data and fewer than 300 events for dichotomous data (Guyatt 2011).
- 5. Publication bias: downgrade where there is direct evidence of publication bias.

We presented the GRADE judgements for all outcomes for comparisons of aerobic exercise versus usual care, resistance training versus usual care, mixed training versus usual care, and aerobic exercise versus resistance training, in the Effects of interventions section. We also presented GRADE ratings for outcomes where there were sufficient data to conduct metaanalyses for comparisons of aerobic exercise versus usual care, resistance training versus usual care, mixed training versus usual care, and aerobic exercise versus resistance training in 'Summary of findings' tables, which we constructed using GRADEpro GDT 2014.

See Summary of findings for the main comparison; Summary of findings 2; Summary of findings 3; Summary of findings 4.

RESULTS

Description of studies

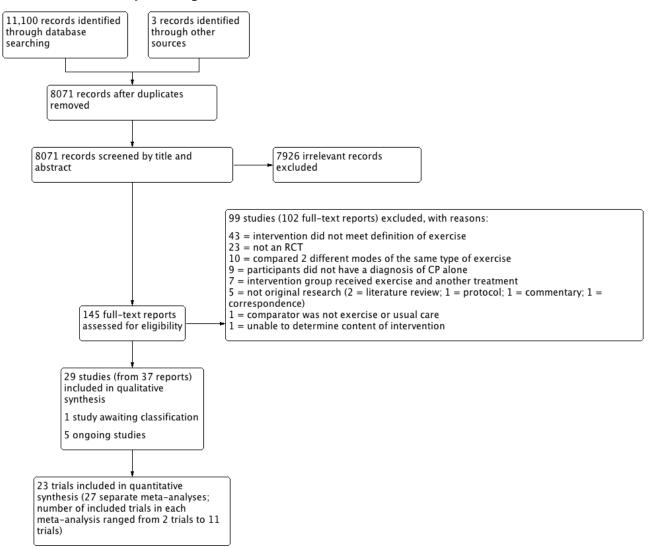
Results of the search

We ran our searches in May 2015 and updated them in June 2016. Our searches yielded 11,100 original research studies, reviews and abstracts from databases and a further three reports from additional sources (Seniorou 2007; Nsenga Leunkeu 2013; Mitchell 2016). After removing duplicates, we screened the titles and abstracts of the remaining 8071 records against our inclusion criteria (Criteria for considering studies for this review) and retrieved 145 full-text reports for further assessment. We used translators to evaluate two reports published in languages other than English. We excluded 99 studies (102 reports) that did not meet the inclusion criteria (see Excluded studies; Characteristics of excluded studies tables) and included 29 studies (from 37 reports) in the review (see Included studies; Characteristics of included studies tables). One additional trial, Carlon 2014, was only available as a conference abstract and is described in the Characteristics of studies awaiting classification tables. We also identified five ongoing trials (Gillett 2015; ISRCTN90378161; NCT02754128; NCT02766491; RBR-5rh6cg), which we describe in Characteristics of ongoing studies tables. Figure 1 presents a summary flow diagram.

Exercise interventions for cerebral palsy (Review)

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Figure 1. 8071-7926-Study flow diagram.



Included studies

This review includes 29 studies from 37 reports (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Verschuren 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Bryant 2013; Chen 2012; Chrysagis 2012; Mattern-Baxter 2013; Taylor 2013; Tedla 2014; Emara 2015; Lee 2015; Mitchell 2016). We provided a detailed description of each included study in the Characteristics of included studies tables.

Design

All included trials were RCTs. One trial used a randomised crossover design (Reid 2010). One trial allowed participants to continue or cross-over into the exercise training programme after nine months of training (Van den Berg-Emons 1998); we analysed this trial up to the point at which the participants were allowed to cross over. Two trials were quasi-randomised (Unnithan 2007; Mattern-Baxter 2013). Twenty-six trials contained two arms (Van den Berg-Emons 1998; Dodd 2003; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Verschuren 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Mattern-Baxter 2013; Taylor 2013; Tedla 2014; Emara 2015; Lee 2015; Mitchell 2016), two trials contained three arms (McCubbin 1985; Bryant 2013), and one trial contained four arms (Engsberg 2006).

Participants

The 29 trials involved a total of 926 participants. The number of participants per trial ranged from 12 in Chrysagis 2009 to 102 in Mitchell 2016. We judged 24 trials to be small (N < 50) (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Reid 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013; Taylor 2013; Emara 2015; Lee 2015), four trials to be medium sized (between 50 and 100 participants) (Verschuren 2007; Fowler 2010; Scholtes 2010; Tedla 2014), and one trial to be large (≥ 100 participants) (Mitchell 2016). The exact number of participants in Olama 2011 was unclear, as authors provided contradictory information. All

Exercise interventions for cerebral palsy (Review)

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participants had a diagnosis of CP. Two trials included children only (aged less than 10 years) (Mattern-Baxter 2013; Emara 2015), six trials included adolescents only (aged 10 to 19 years) (Unger 2006; Unnithan 2007; Gharib 2011; Olama 2011; Smania 2011; Chrysagis 2012), 17 trials included children and adolescents up to the age of 20 years (Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Fowler 2010; Reid 2010; Scholtes 2010; Johnston 2011; Pandey 2011; Chen 2012; Bryant 2013; Tedla 2014; Lee 2015; Mitchell 2016; Verschuren 2007), three trials included adolescents and young adults from the age of 14 to 22 years (McCubbin 1985; Chrysagis 2009; Taylor 2013), and one trial included adults over the age of 20 years (Maeland 2009). All trials included males and females. The mean percentage of males and females in the included studies was 53% (SD 10%; range 31% to 67%) and 47% (SD 10%; range 33% to 69%), respectively.

Twenty-six trials included people with spastic CP (Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Verschuren 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Taylor 2013; Tedla 2014; Emara 2015; Lee 2015; Mitchell 2016). Of these, 10 specifically included people with spastic diplegia (Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Maeland 2009; Fowler 2010; Taylor 2013; Tedla 2014; Emara 2015), one included children with unilateral CP only (Mitchell 2016), two did not state if participants had unilateral or bilateral CP (Pandey 2011; Lee 2015), and the remainder included participants with unilateral and bilateral spastic CP (Van den Berg-Emons 1998; Unger 2006; Verschuren 2007; Lee 2008; Chrysagis 2009; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Smania 2011; Chen 2012; Chrysagis 2012). One trial included children with dyskinetic and spastic (unilateral and bilateral) CP (Bryant 2013), one trial included children with spastic and hypotonic CP (Mattern-Baxter 2013), and one trial included children with spastic, athetoid, ataxic and mixed CP (McCubbin 1985).

Eight trials included people classified in GMFCS levels I, II and III (Dodd 2003; Engsberg 2006; Seniorou 2007; Chrysagis 2009; Fowler 2010; Scholtes 2010; Chrysagis 2012; Lee 2015). Six trials included people classified in GMFCS levels I and II (Liao 2007; Verschuren 2007; Lee 2008; Chen 2012; Mattern-Baxter 2013; Mitchell 2016). Two trials included people in GMFCS levels I, II, III and IV (Smania 2011; Tedla 2014). One trial included people in GMFCS level II only (Gharib 2011), two included people in GMFCS levels II and III (Maeland 2009; Taylor 2013), one included people in GMFCS levels II, III and IV (Johnston 2011), and one included people in GMFCS levels IV and V (Bryant 2013). Eight trials did not state participants' GMFCS level (McCubbin 1985; Van den Berg-Emons 1998; Unger 2006; Unnithan 2007; Olama 2011; Pandey 2011; Emara 2015; Reid 2010). Of these, three trials stated that participants were able to walk with or without aids or that they occasionally used a wheelchair (Unger 2006; Unnithan 2007; Pandey 2011), one trial stated that participants were able to walk independently (Olama 2011), and one trial reported including people who were both ambulant and wheelchair bound (Van den Berg-Emons 1998). Two trials reported the number of participants classified in each MACS level; participants were in MACS levels I, II and III (Reid 2010; Mitchell 2016).

Only one trial reported the ethnicity of participants, which included African Americans, whites, Asians and others (Fowler 2010).

Settings

Trials took place in a number of geographical locations: Australia (Dodd 2003; Reid 2010; Taylor 2013; Mitchell 2016), Egypt (Gharib 2011; Olama 2011), Greece (Unnithan 2007; Chrysagis 2009; Chrysagis 2012), India (Pandey 2011), Italy (Smania 2011), Korea (Lee 2008; Lee 2015), Norway (Maeland 2009), Saudi Arabia (Tedla 2014; Emara 2015), South Africa (Unger 2006), Taiwan (Liao 2007; Chen 2012), the Netherlands (Van den Berg-Emons 1998; Verschuren 2007; Scholtes 2010), the UK (Seniorou 2007; Bryant 2013), and the USA (McCubbin 1985; Engsberg 2006; Fowler 2010; Johnston 2011; Mattern-Baxter 2013).

Interventions

Aerobic exercise

Six trials compared aerobic exercise to usual care (Van den Berg-Emons 1998; Chrysagis 2009; Gharib 2011; Smania 2011; Chrysagis 2012; Emara 2015). One trial compared aerobic exercise to a physical therapy session, the content of which was not clear (Mattern-Baxter 2013). One trial with three arms compared aerobic exercise on a static bike, aerobic exercise on a treadmill and usual care (Bryant 2013). Modes of aerobic exercise included swimming; cycling on a stationary bike; wheelchair driving; negotiating stairs; and walking or running on a gait trainer, treadmill or overground.

Resistance training

Ten trials compared lower limb resistance training to usual care, active movements without resistance or no physiotherapy (Dodd 2003; Liao 2007; Seniorou 2007; Lee 2008; Maeland 2009; Scholtes 2010; Pandey 2011; Taylor 2013; Lee 2015; Mitchell 2016). Two trials compared upper limb, lower limb, and trunk resistance training to usual care (Unger 2006; Tedla 2014).

One trial with four arms compared resistance training of the dorsiflexors, plantarflexors, plantar and dorsiflexors verus no resistance training (Engsberg 2006). Two trials compared upper limb strength training to normal activity or active movements without resistance (McCubbin 1985; Reid 2010).

Mixed training

Four trials compared mixed training to usual care (Unnithan 2007; Verschuren 2007; Fowler 2010; Chen 2012). Mixed training consisted of aerobic and lower limb resistance training (Fowler 2010; Chen 2012); upper and lower limb and trunk resistance training and aerobic exercise (Unnithan 2007); and aerobic, anaerobic and general strength training (Verschuren 2007).

Two trials compared aerobic exercise to resistance training, either of the trunk, upper and lower limbs (Johnston 2011), or of the lower limbs only (Olama 2011).

Duration, frequency and intensity of interventions

The duration of aerobic exercise programmes ranged from two weeks (Smania 2011) to nine months (Van den Berg-Emons 1998). The duration of resistance training programmes ranged from 4 weeks (Pandey 2011) to 20 weeks (Mitchell 2016). The duration of mixed training programmes ranged from 12 weeks (Unnithan 2007; Fowler 2010; Chen 2012) to 8 months (Verschuren 2007). The

Exercise interventions for cerebral palsy (Review)

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duration of interventions in one trial comparing aerobic exercise to lower limb resistance training was 12 weeks (Johnston 2011). It was not clear how long the intervention was in a second trial comparing aerobic exercise to resistance training, as authors provided contradictory information (Olama 2011).

Aerobic exercise

For aerobic interventions, participants were prescribed the intervention for two days per week (Verschuren 2007; Chrysagis 2009), three days per week (Unnithan 2007; Fowler 2010; Gharib 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Emara 2015), four days per week (Van den Berg-Emons 1998), five days per week (Johnston 2011; Smania 2011), six days per week (Mattern-Baxter 2013), or seven days per week (Olama 2011). The prescribed duration of exercise per session ranged from 12 min to 60 min. One trial did not report duration of exercise (Olama 2011). The intensity of aerobic exercise was unclear or not reported in eight trials (Verschuren 2007; Chrysagis 2009; Gharib 2011; Johnston 2011; Olama 2011; Chen 2012; Bryant 2013; Mattern-Baxter 2013). One trial reported that participants were prescribed walking "at a comfortable speed" (Chrysagis 2012), and one trial reported that participants were prescribed walking at 75% of their comfortable walking speed (Emara 2015). Two trials did not report the intensity prescribed but reported that walking speed gradually increased throughout the programme (Smania 2011), and that the mean time spent at > 70% of heart rate reserve was 49% (SD 17%) when heart rate was randomly measured in participants during the intervention (Van den Berg-Emons 1998). Two trials prescribed an intensity of 65% to 75% of heart rate maximum and 70% to 80% of heart rate reserve, respectively (Unnithan 2007; Fowler 2010).

Seven trials did not report fidelity to the intervention (Unnithan 2007; Chrysagis 2009; Gharib 2011; Olama 2011; Smania 2011; Chen 2012; Emara 2015), although one reported excluding participants from the analysis if they missed more than three of the prescribed sessions (Gharib 2011). Six trials reported attendance at sessions (Van den Berg-Emons 1998; Verschuren 2007; Fowler 2010; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013); the average adherence to the prescribed number of sessions across trials ranged from 77.0% to 93.0%. One trial also reported the average time walked per session (Mattern-Baxter 2013), which was 28.2 min a day (range 9.6 min a day to 39.3 min a day) out of a prescribed 20.0 to 40.0 min a day. Only one trial reported fidelity to the intensity of the intervention, reporting that the mean percentage of heart-rate maximum attained during sessions was 52.2% (SD 12.2%; range 8% to 77%) (Fowler 2010).

Resistance training

For resistance training interventions, participants were prescribed the intervention for one to three days a week (Unger 2006), two days per week (Verschuren 2007; Pandey 2011; Taylor 2013), three days a week (McCubbin 1985; Dodd 2003; Liao 2007; Seniorou 2007; Unnithan 2007; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Olama 2011; Chen 2012; Tedla 2014; Lee 2015), five days a week (Johnston 2011), or six days a week (Mitchell 2016). Therapists prescribed the following sets, repetitions and intensity: two sets of 10 repetitions to fatigue (although only 0.25 kg, 0.45 kg or 0.90 kg weights were used) (Lee 2008); two sets of 10 repetitions at 75% of one RM (Chen 2012); three sets of eight repetitions at an intensity that was not reported, 50% to 70% of maximum torque, or 10 RM, respectively (McCubbin 1985; Seniorou 2007; Reid 2010); **Cochrane** Database of Systematic Reviews

three sets of 8 to 10 repetitions at 8 to 12 RM (Dodd 2003); three sets of 10 to 12 repetitions at 60% to 80% of one RM (Taylor 2013); three sets of 6 to 10 repetitions at 80% of one RM (Tedla 2014); four sets of 12 to 15 repetitions at 60% to 75% of one RM and four to six repetitions at 85% of one RM (Maeland 2009); one to three sets of 6 to 12 repetitions to fatigue (Unger 2006); six sets of five repetitions at 80% or more of one RM (Engsberg 2006); 7 to 11 activities with 5 to 10 repetitions or 20 repetitions, depending on the exercise, at 75% of one RM (Mitchell 2016); two sets of 10 repetitions at 20% of one RM and one set of as many repetitions as possible at 50% of one RM (Liao 2007); one set of 10 repetitions at 50% of 10 RM gradually increased to 10 repetitions at 100% of 10 RM (Olama 2011); three to five sets of 8 to 15 repetitions depending on the exercise (intensity not reported) (Unnithan 2007). Three trials did not report the number of sets, repetitions or intensity prescribed (Verschuren 2007; Johnston 2011; Pandey 2011). Two trials did not report the sets or repetitions prescribed but reported that exercises were performed at eight RM and 10 RM in Lee 2015 and Fowler 2010, respectively.

Eleven studies did not provide information on fidelity to the intervention (McCubbin 1985; Engsberg 2006; Unger 2006; Seniorou 2007; Unnithan 2007; Lee 2008; Olama 2011; Pandey 2011; Chen 2012; Tedla 2014; Lee 2015). Of the studies that reported adherence to the prescribed number of sessions (Dodd 2003; Verschuren 2007; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Taylor 2013), the average adherence ranged from 88.9% (Reid 2010) to 93.3% (Dodd 2003). One study reported that participants completed, on average, 32.4 hours of potential 60 hours of training (Mitchell 2016). One trial reported that the mean sets performed were 147.7 (SD 23.4) out of a possible 162 (Dodd 2003). Only two trials provided information on fidelity to the intensity of the intervention. One trial reported that the mean rating of exertion at the end of each session was 6.9 (SD 1.1) out of 10, and that participants increased their training load from session 3 to 24 by a mean of 183% (SD 23%) (Taylor 2013). The second trial reported that participants increased their training load by a mean of 17.5 kg (SD 11.7 kg; range 0 kg to 40.8 kg) (Fowler 2010).

Comparator

Nineteen trials reported that the control group received usual physiotherapy. Eleven trials reported the general content of usual physiotherapy (Liao 2007; Unnithan 2007; Lee 2008; Maeland 2009; Gharib 2011; Smania 2011; Chrysagis 2012; Bryant 2013; Tedla 2014; Emara 2015; Lee 2015), although this varied across individuals, and it was difficult to determine what usual care each participant received. Ten trials stated the dose of usual physiotherapy that was prescribed (Van den Berg-Emons 1998; Liao 2007; Unnithan 2007; Scholtes 2010; Gharib 2011; Smania 2011; Chrysagis 2012; Tedla 2014; Emara 2015; Lee 2015). Four trials indicated that participants in the control group continued with usual activities, but not specifically usual physiotherapy (Chrysagis 2009; Fowler 2010; Reid 2010; Chen 2012). For two trials, it was unclear what the control group did during the intervention period; one trial reported that the control group received no strengthening (Engsberg 2006) and the authors of the second trial provided no information about the comparator (Unger 2006). Two trials investigating the effect of resistance training prescribed active movements without resistance to the control group (McCubbin 1985; Seniorou 2007). Pandey 2011 was the only trial to specifically state that all participants were not allowed to receive usual physiotherapy for the duration of the trial. Two trials tracked physiotherapy received

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by all participants and reported that physiotherapy did not differ between the control and intervention group (Verschuren 2007; Mitchell 2016), although neither trial reported the content of physiotherapy, and one trial did not report the dose (Verschuren 2007).

Ongoing studies

All ongoing studies were RCTs conducted in Australia (Gillett 2015), Canada (NCT02754128), the UK (ISRCTN90378161; NCT02766491), and Brazil (RBR-5rh6cg). The number of participants that studies recruited ranged from 22 in NCT02754128 to 60 in ISRCTN90378161. Participants were aged 15 to 30 years (Gillett 2015), 7 to 17 years (NCT02754128), 7 to 14 years (NCT02766491), 4 to 11 years (RBR-5rh6cg), and 10 to 19 years (ISRCTN90378161). Three studies specified that they included children with spastic CP (RBR-5rh6cg; Gillett 2015; NCT02766491). Two studies specified that participants had hemiplegia and diplegia (Gillett 2015; NCT02754128), one specified that participants had diplegia (RBR-5rh6cg), and two did not specify the anatomical distribution of CP participants (ISRCTN90378161; NCT02766491). Two studies included participants in GMFCS level I and II (Gillett 2015; NCT02754128), two studies included participants in GMFCS levels I, II and III (ISRCTN90378161; NCT02766491), and one study included participants in GMFCS level IV (RBR-5rh6cg). Three studies were investigating the effect of resistance training compared to usual care (ISRCTN90378161), no training (Gillett 2015), or conventional stretching and upper limb exercises (NCT02766491). One study was comparing resistance training to aerobic exercise (NCT02754128), and one study was comparing trunk control exercises to conventional aquatic therapy (RBR-5rh6cg).

Studies awaiting classification

The single study awaiting classification was a RCT comparing aerobic training delivered three times a week for nine weeks, to an arts programme of the same duration, delivered in a school in Australia (Carlon 2014). Participants were 19 children with a mean age of three years classified in GMFCS levels I, II and III. We contacted the authors when we identified the study to determine if the results had been published as a full report and were informed that a full report had not been published at the time. We were unable to determine from the abstract if the intervention met our definition of aerobic exercise.

Excluded studies

We excluded 99 studies (102 reports) following full-text screening and provide details in the Characteristics of excluded studies table. The main reasons for exclusion were as follows: in 43 studies the intervention did not meet the definition of aerobic or resistance training, as defined in the Description of the intervention section; 23 studies were not RCTs; 10 studies compared two different modes of the same type of exercise (Stackhouse 2007; Willoughby 2010; Kim 2012; Olama 2012; Grecco 2013a; Grecco 2013b; Moreau 2013; Su 2013; Hussein 2014; Swe 2015); nine studies did not involve participants with a diagnosis of CP alone (Katz-Leurer 2009; Speyer 2010; Salem 2012; Angulo-Barroso 2013; Ayhan 2014; Hammond 2014; Williams 2014; Lowe 2015; Hsieh 2016); in seven studies exercise was provided as intervention in conjunction with another treatment, and it was not possible to determine the independent effect of exercise on the outcome (Patikas 2006; Bandholm 2012; Williams 2013; Slaman 2014; Van Wely 2014; Preston 2015; Sherief AEAA 2015); five studies were not original research (Hornyak 2008; Taylor 2009; Boyd 2010; Roberti 2011; Verschuren 2014); one study did not use exercise or usual care as the comparator (Ahlborg 2006); and in one study it was not possible to determine the content of the intervention (Kumar 2013).

Risk of bias in included studies

Figure 2 and Figure 3 summarise the 'Risk of bias' assessments for all included trials. We judged all trials as being at high risk of bias overall because we rated at least one domain as being at high risk of bias.



Figure 2. Risk of bias summary: review authors' judgements about each risk of bias item for each included study.

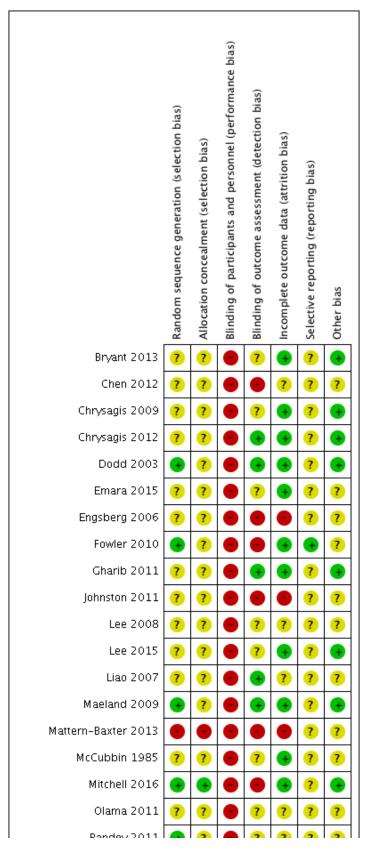




Figure 2. (Continued)

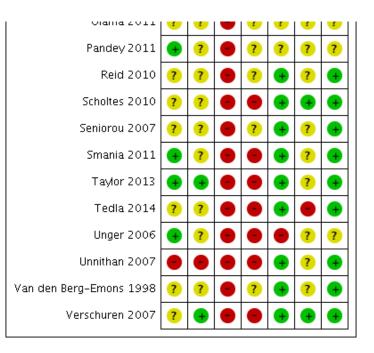
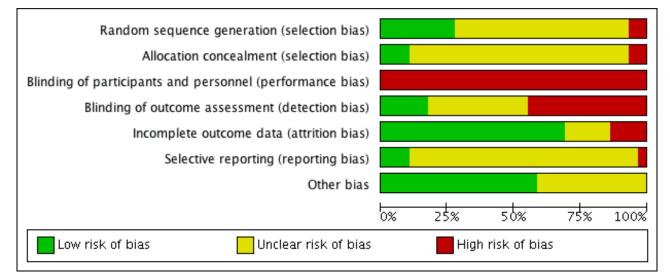


Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.



Allocation

Random sequence generation

We judged eight trials to be at low risk of selection bias, as two used adequate methods of sequence generation (Taylor 2013; Mitchell 2016), and six provided sufficient information to determine the sequence generation was random (Dodd 2003; Unger 2006; Maeland 2009; Fowler 2010; Pandey 2011; Smania 2011).

We judged 19 trials to be at unclear risk of selection bias: 18 because they failed to adequately report the methods used to generate a random sequence (McCubbin 1985; Van den Berg-Emons 1998; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Chrysagis 2009; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama

Exercise interventions for cerebral palsy (Review)

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2011; Chen 2012; Chrysagis 2012; Bryant 2013; Tedla 2014; Emara 2015; Lee 2015), and one because, although the authors stated that participants were randomly assigned to two groups using a four-block randomisation process, they did not describe the randomisation process (Verschuren 2007).

We rated two trials at high risk of selection bias as they used a quasirandomisation method (Unnithan 2007; Mattern-Baxter 2013).

Allocation concealment

We judged three trials to be at low risk of selection bias as they used adequate methods of allocation concealment (Verschuren 2007; Taylor 2013; Mitchell 2016).



We judged 24 trials that failed to adequately report the methods used to conceal allocation at unclear risk of selection bias (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Tedla 2014; Emara 2015; Lee 2015).

We judged two trials at high risk of selection bias as they used a quasi-randomisation method (Unnithan 2007; Mattern-Baxter 2013).

Blinding

Blinding of participants and personnel (performance bias)

We judged all 29 trials to be at high risk of performance bias as participants or personnel were not blinded to group allocation (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Verschuren 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013; Taylor 2013; Tedla 2014; Emara 2015; Lee 2015; Mitchell 2016).

Blinding of outcome assessment (detection bias)

We rated 13 trials at high risk of detection bias (Engsberg 2006; Unger 2006; Unnithan 2007; Verschuren 2007; Fowler 2010; Scholtes 2010; Johnston 2011; Smania 2011; Chen 2012; Mattern-Baxter 2013; Taylor 2013; Tedla 2014; Mitchell 2016). Objective outcomes were assessed by assessors blinded to group allocation in six of these trials, but participants or their parents were responsible for self-report measures, and they were not blinded to group allocation, suggesting that a lack of blinding was likely to have affected a number of trial outcomes (Unger 2006; Verschuren 2007; Fowler 2010; Scholtes 2010; Smania 2011; Taylor 2013). The remaining studies stated that at least some of the assessments were conducted by a person who was not blind to group allocation (Engsberg 2006; Unnithan 2007; Johnston 2011; Chen 2012; Mattern-Baxter 2013; Tedla 2014; Mitchell 2016).

Eleven trials did not report adequate information to determine if outcome assessors were blinded to group allocation, so we judged these trials to be at unclear risk of detection bias (McCubbin 1985; Van den Berg-Emons 1998; Seniorou 2007; Lee 2008; Chrysagis 2009; Reid 2010; Olama 2011; Pandey 2011; Bryant 2013; Emara 2015; Lee 2015).

We rated five trials at low risk of detection bias because assessors were blinded to group allocation (Dodd 2003; Liao 2007; Maeland 2009; Gharib 2011; Chrysagis 2012).

Incomplete outcome data

We considered 20 trials to be at low risk of attrition bias, as they had no missing data, low rates of missing data (10% or less) that were evenly distributed across groups, or the authors performed an intention-to-treat analysis (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Seniorou 2007; Unnithan 2007; Verschuren 2007; Chrysagis 2009; Maeland 2009; Fowler 2010; Reid 2010; Scholtes 2010; Gharib 2011; Smania 2011; Chrysagis 2012; Bryant 2013; Taylor 2013; Tedla 2014; Emara 2015; Lee 2015; Mitchell 2016). We judged five trials, which did not provide sufficient information on the number of participants who withdrew from the study or the reasons for withdrawal, at unclear risk of bias (Liao 2007; Lee 2008; Olama 2011; Pandey 2011; Chen 2012).

Four trials had high rates of missing data (three had 20% or more, and one had more than 30%), missing data were not evenly distributed across groups, or reasons for missing data were likely to be related to the trial outcome and were therefore judged as being at high risk of bias (Engsberg 2006; Unger 2006; Johnston 2011; Mattern-Baxter 2013).

Selective reporting

We considered three trials to be at low risk of reporting bias, as they adequately reported outcome data described in the trial protocol (Verschuren 2007; Fowler 2010; Scholtes 2010).

We judged 25 trials to be at unclear risk of bias: one because a trial protocol was available and outcomes were reported across two reports, but some outcomes were not reported to date (Mitchell 2016), and 24 because a trial protocol was not available and therefore it was not possible to determine if all expected outcomes were reported (McCubbin 1985; Van den Berg-Emons 1998; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Lee 2008; Chrysagis 2009; Maeland 2009; Reid 2010; Gharib 2011; Johnston 2011; Olama 2011; Pandey 2011; Smania 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013; Taylor 2013; Emara 2015; Lee 2015).

We rated one trial at high risk of bias because data on strength and gross motor function were reported incompletely and could not be entered into a meta-analysis (Tedla 2014).

Other potential sources of bias

We considered 12 trials to be at unclear risk of other potential bias: 2 for not providing any demographic data (McCubbin 1985; Olama 2011), 3 for providing demographic and baseline outcome data for those who completed the study only (Liao 2007; Chen 2012; Mattern-Baxter 2013), 3 for being unclear about whether all demographic or baseline data were reported (Lee 2008; Pandey 2011; Emara 2015); 1 for providing outcome data only for those who completed the study (Unger 2006), 1 for providing demographic and baseline data only for those who completed and for not reporting all outcome data for those who completed the study (Johnston 2011), 1 for reporting demographic data only for those participants who improved (Engsberg 2006), and 1 for unaccounted missing data (Fowler 2010).

We considered 17 trials to be at low risk of bias for other potential bias (Van den Berg-Emons 1998; Dodd 2003; Seniorou 2007; Unnithan 2007; Verschuren 2007; Chrysagis 2009; Maeland 2009; Reid 2010; Scholtes 2010; Gharib 2011; Smania 2011; Chrysagis 2012; Bryant 2013; Taylor 2013; Tedla 2014; Lee 2015; Mitchell 2016).

Effects of interventions

See: Summary of findings for the main comparison Aerobic exercise versus usual care; Summary of findings 2 Resistance training versus usual care; Summary of findings 3 Mixed training versus usual care; Summary of findings 4 Resistance training versus aerobic exercise

Exercise interventions for cerebral palsy (Review)

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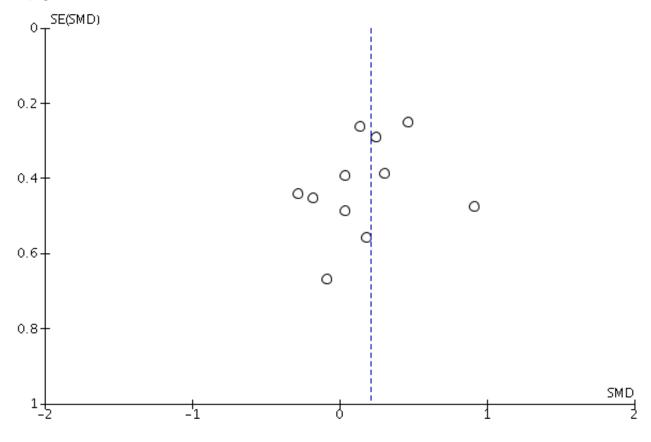


All 29 trials reported outcomes at short-term time points (i.e. zero to one month postintervention). Nine trials reported outcomes at intermediate time points (i.e. more than one month to six months' postintervention) (Dodd 2003; Seniorou 2007; Verschuren 2007; Lee 2008; Scholtes 2010; Johnston 2011; Bryant 2013; Mattern-Baxter 2013; Taylor 2013). One trial reported that assessments were made preintervention, postintervention and at 'follow-up' but did not state when the follow-up assessment occurred (Pandey 2011). We

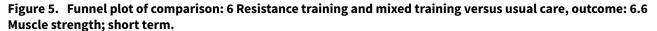
therefore did not report this follow-up time point. No trials reported outcomes at time points beyond six months' postintervention.

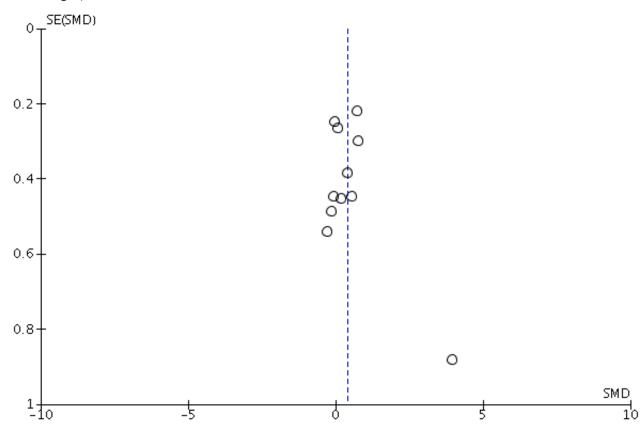
As described under Data synthesis, we performed separate metaanalyses for studies in adults versus children and adolescents. We reported the results for three trials involing adolescents and adults (aged 10 to 22 years) (McCubbin 1985; Chrysagis 2009; Taylor 2013). We identified sufficient data (at least 10 studies in a metaanalysis) to examine funnel plots for two meta-analyses and found no indication of publication bias (see Figure 4; Figure 5).

Figure 4. Funnel plot of comparison: 6 Resistance training and mixed training versus usual care, outcome: 6.1 Activity: gross motor function; short term.









Comparison 1: aerobic exercise versus usual care

Seven trials comparing aerobic exercise to usual care reported at least one primary or secondary outcome of interest (Van den Berg-Emons 1998; Chrysagis 2009; Gharib 2011; Smania 2011; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013). One trial did not report any outcomes of interest (Emara 2015), so we do not report its results.

Activity

Pooled results

Three trials assessed gross motor function in children and adolescents following aerobic exercise (Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013). For one three-armed trial, we combined data from two aerobic exercise groups (bike and treadmill exercise) to compare all modes of aerobic exercise to usual care (Bryant 2013).

The pooled analysis of three trials in 65 children and adolescents indicated that aerobic training improved gross motor function compared to usual care in the short term (SMD 0.53, 95% CI 0.02 to 1.04, P = 0.04, $I^2 = 0$ %, Analysis 1.1; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013).

Pooled analysis of four trials with high heterogeneity involving 82 children and adolescents (GMFCS levels I to IV), demonstrated that aerobic exercise did not result in an improvement in gait speed in the short term (MD 0.09 m/s, 95% CI –0.11 m/s to 0.28 m/s, P = 0.38,

Exercise interventions for cerebral palsy (Review)

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I² = 78%, Analysis 1.2; Gharib 2011; Smania 2011; Chrysagis 2012; Mattern-Baxter 2013).

Single study results

One trial of 12 adolescents and young adults (GMFCS levels I to III; Chrysagis 2009) reported that there was no between-group difference in gross motor function over time (P = 0.11). Although the authors reported that the mean score on dimension E of the GMFM increased following aerobic exercise from 59.02% to 65.04% in the short-term, and that there was little short-term change in the comparison group (mean increased from 59.02% to 59.95%), they did not provide information to allow us to calculate the mean difference between groups.

One trial, Smania 2011 (18 children and adolescents, GMFCS levels I to IV), reported a greater short-term improvement in walking endurance, measured as the distance walked (meters) at self-selected walking speed for six minutes, following aerobic exercise (Cohen's d = 0.44, P = 0.02). Using the postintervention means and SDs provided for each group, we calculated an MD of 41.00 m per 6 minutes walked (95% CI -46.98 m to 128.98 m, Analysis 1.3).

Only one trial, Mattern-Baxter 2013 (12 children), compared aerobic exercise versus usual care for gait speed in the intermediate term, reporting no between-group difference in gait speed (MD –0.17 m/ s, 95% CI –0.59 m/s to 0.24 m/s, Analysis 1.4).



Although gross motor function was measured in children and adolescents with CP at an intermediate time point in two trials, the authors of one trial did not provide a P value to support this or data to allow us to calculate it (Bryant 2013). The second trial, Mattern-Baxter 2013, found that gross motor function improved in the intermediate term (MD 12.96%, 95% CI 0.52% to 25.40%, 12 children, Analysis 1.5).

One trial, Van den Berg-Emons 1998 (20 children and adolescents, ambulatory and non-ambulatory), reported that there was no evidence of a difference in the change in daily physical activity (calculated as the ratio of total energy expenditure to sleeping or resting metabolic rate between participants in the intervention group versus usual care after nine months of aerobic exercise (P value not reported). Using the postintervention means and SDs provided, we calculated an MD of 0.21 (95% CI 0.04 to 0.38, Analysis 1.6) in daily physical activity.

Participation

No trial assessed the effect of aerobic exercise in comparison to usual care on participation.

Quality of life

No trial comparing aerobic exercise to usual care assessed quality of life.

Adverse events

Pooled results

No meta-analysis was possible.

Single study results

Four out of eight trials did not report monitoring adverse events (Van den Berg-Emons 1998; Chrysagis 2009; Gharib 2011; Emara 2015). Four trials reported that participants did not experience any adverse events (Smania 2011; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013).

Body functions and body structures

Pooled results

No meta-analysis was possible.

Single study results

Only one trial in 20 ambulatory and non-ambulatory children and adolescents assessed aerobic fitness following an aerobic exercise intervention (Van den Berg-Emons 1998). This trial reported that after nine months of aerobic exercise there was a greater increase in aerobic fitness in participants with CP in the intervention group compared to the usual care group in the short term (P < 0.05; effect size and exact P value not reported). Our analysis of postintervention means and SDs provided an MD of 0.06 W/kg FFM (95% CI –0.71 W/kg FFM to 0.83 W/kg FFM, Analysis 1.7).

Sensitivity analysis

To assess the influence of our analysis model on the results, we repeated the pooled analyses using a fixed-effect model instead of a random-effects model. Using a fixed-effect model, there was some evidence of an improvement in gait speed following aerobic exercise compared to usual care in 82 children and adolescents in the short term, with no change in heterogeneity (MD 0.08 m/s,

Exercise interventions for cerebral palsy (Review)

95% CI 0.02 m/s to 0.14 m/s, P = 0.01, I^2 = 78%, 4 studies; Gharib 2011; Smania 2011; Chrysagis 2012; Mattern-Baxter 2013; analysis not shown). Using a fixed-effect model instead of a random-effects model had no impact on gross motor function in children and adolescents in the short term (analysis not shown).

Quality of evidence

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that aerobic exercise improves gross motor function and physical activity but does not improve walking endurance or aerobic fitness in children and adolescents with CP in the short term. There is very low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision, once for inconsistency) that aerobic exercise does not improve gait speed in children and adolescents with CP in the short term.

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that aerobic exercise improves gross motor function but does not improve gait speed in children and adolescents with CP in the intermediate term.

See Summary of findings for the main comparison.

Comparison 2: resistance training versus usual care

Fourteen trials comparing resistance training to usual care reported outcomes of interest; 11 included children and adolescents (Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Reid 2010; Scholtes 2010; Pandey 2011; Tedla 2014; Lee 2015; Mitchell 2016), one included adolescents only (Unger 2006), one included adolescents and young adults aged 14 to 22 years (Taylor 2013), and one included adults (Maeland 2009). One trial investigating the effect of resistance training in adolescents and young adults with CP did not report any outcomes of interest, so we do not report the results of this trial (McCubbin 1985).

Activity

Pooled results

Eight trials assessed gross motor function in children and adolescents using the GMFM (Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Scholtes 2010; Tedla 2014; Lee 2015). Tedla 2014 did not report between-group differences, so we could not extract data. For the remaining seven studies, resistance training did not improve short-term gross motor function more than usual care (SMD 0.12, 95% CI –0.19 to 0.43, P = 0.45, $l^2 = 0\%$, 164 children and adolescents, Analysis 2.1; Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Scholtes 2010; Lee 2015). Gross motor function also did not improve following resistance training in 85 children and adolescents (GMFCS levels I to III) in the intermediate term (SMD 0.13, 95% CI –0.30 to 0.55, P = 0.57, $l^2 = 0\%$, 3 studies, Analysis 2.2; Dodd 2003; Lee 2008; Scholtes 2010).

Short-term gait speed did not improve in 185 children and adolescents with CP (GMFCS levels I to III) following resistance training compared to usual care (MD 0.03 m/s, 95% CI –0.02 m/s to 0.07 m/s, P = 0.20, I² = 0%, 8 studies, Analysis 2.3; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Lee 2008; Scholtes 2010; Pandey 2011). In addition, resistance training did not improve intermediate-term gait speed in comparison to usual care (MD –0.03 m/s, 95% CI –0.17 m/s to 0.11 m/s, P = 0.65, I² = 0%, 84

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children and adolescents, 3 studies, Analysis 2.4; Dodd 2003; Lee 2008; Scholtes 2010).

Single study results

Maeland 2009 reported that resistance training did not improve short-term gait speed in 12 adults in GMFCS levels II and III (MD 0.30 m/s, 95% CI –0.28 m/s to 0.88 m/s, Analysis 2.5); short-term gross motor function, assessed with a timed stair climb (MD 10.00 stairs, 95% CI –13.55 stairs to 33.55 stairs, Analysis 2.6); or short-term walking endurance (MD 127.00 m, 95% CI –95.08 m to 349.08 m, Analysis 2.7).

One trial in 48 adolescents and young adults in GMFCS levels II and III reported that, in the short term, resistance training did not improve gait speed (MD 0.01 m/s, 95% CI –0.06 m/s to 0.07 m/s), gross motor function (MD 0.9%, 95% CI –3.0% to 4.7%), walking endurance (MD 0.1 m, 95% CI –20.6 m to 20.9 m), or steps per day (MD –28 steps/day, 95% CI –1373 steps/day to 1317 steps/day) (Taylor 2013). The same trial reported that resistance training did not improve these outcomes in the intermediate term: gait speed (MD – 0.05 m/s, 95% CI –0.11 m/s to 0.02 m/s), gross motor function (MD 1.0%, 95% CI –2.6% to 4.5%), walking endurance (MD –12.3 m, 95% CI –34.8 m to 10.2 m) and steps per day (MD –1093 steps/day, 95% CI –2316 steps/day to 130 steps/day).

Only one trial, Mitchell 2016, assessed walking endurance in 101 children and adolescents (GMFCS level I and II), reporting that walking endurance improved following resistance training in comparison to usual care (MD 38.9 m, 95% CI 12.3 m to 64.5 m). Mitchell 2016 also reported that resistance training did not result in an improvement in steps per day among children in the short term (MD 563.7 steps/day, 95% CI –706.5 steps/day to 1833.8 steps/day).

Participation

Pooled results

Pooled analysis of two trials, Mitchell 2016 and Scholtes 2010, suggested that short-term participation did not improve in participants in GMFCS levels I to III receiving resistance training versus usual care (SMD 0.34, 95% CI –0.01 to 0.70, P = 0.06, $I^2 = 0\%$, 127 children and adolescents, Analysis 2.8).

Single study results

One trial, Scholtes 2010, also reported that participation did not improve in the intermediate term (P = 0.12). We calculated an MD of 0.37 (95% CI –6.61 to 7.35, 36 children and adolescents, Analysis 2.9) based on the data provided.

Quality of life

Pooled results

No meta-analysis was possible.

Single study results

One trial, Engsberg 2006, reported that parent-reported quality of life improved in 12 children and adolescents with CP following resistance training in the short term, but participant-reported quality of life did not. However, there was no evidence of a between-group difference for parent- or child-reported quality of life based on the data provided (parent reported: MD 12.70, 95% CI –5.63 to 31.03, Analysis 2.10; child reported: MD 11.70, 95% CI –8.32 to 31.72, Analysis 2.11).

Exercise interventions for cerebral palsy (Review)

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Adverse events Pooled results

No meta-analysis was possible.

Single study results

Seven trials did not state whether they recorded any adverse events (McCubbin 1985; Unger 2006; Seniorou 2007; Lee 2008; Reid 2010; Pandey 2011; Lee 2015). Two trials stated that neither participants nor physiotherapists reported or registered any adverse events (Engsberg 2006; Maeland 2009). One trial stated that there were no adverse events that led to missing training sessions (Dodd 2003). However, one participant reported pressure on the shoulders from a loaded backpack, and two participants reported mild foot and ankle discomfort during heel raises (Dodd 2003). Exercises were modified and enabled participants to continue without incident (Dodd 2003). Four trials stated that participants reported muscle soreness, which subsided with rest (Scholtes 2010; Taylor 2013; Tedla 2014; Mitchell 2016). One trial reported that a participant in the intervention group had seizures during the study period, but this was deemed unrelated to the training (Mitchell 2016). In addition, two participants in one trial reported minor calf strain and discomfort to the plantar fascia, respectively, resulting in the programme being adjusted but no missed sessions (Taylor 2013). One trial reported that participants experienced discomfort due to wearing a weighted vest (Liao 2007).

Body structure and function

Twelve trials assessed muscle strength following resistance training (Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Lee 2008; Maeland 2009; Reid 2010; Scholtes 2010; Pandey 2011; Taylor 2013; Tedla 2014; Mitchell 2016). It was not possible to extract data from Tedla 2014, which did not present the necessary numerical data. We did not report muscle strength from a second trial because the authors only included data on participants whose muscle strength improved in the analysis (Engsberg 2006).

Pooled results

Muscle strength improved more with resistance training compared to usual care in children and adolescents in GMFCS levels I to III, both in the short term (SMD 0.53, 95% CI 0.00 to 1.06, P = 0.05, $I^2 = 70\%$, 247 children and adolescents, 8 studies, Analysis 2.12; Dodd 2003; Liao 2007; Seniorou 2007; Lee 2008; Reid 2010; Scholtes 2010; Pandey 2011; Mitchell 2016) and in the intermediate term (SMD 0.50, 95% CI 0.06 to 0.94, P = 0.03, $I^2 = 1\%$, 84 children and adolescents, 3 studies, Analysis 2.13; Dodd 2003; Lee 2008; Scholtes 2010).

Single study results

The results of a single trial, Taylor 2013, suggested that resistance training resulted in a short-term improvement in muscle strength in 48 adolescents and young adults in GMFCS levels II and III (MD 26.7% increase in strength, 95% CI 7.9% to 45.5%) but not in the intermediate term (MD 21.7%, 95% CI –17.3% to 61.7%).

One trial, Maeland 2009, which included 12 adults in GMFCS levels II and III, reported that resistance training did not result in short-term improvements in muscle strength compared to usual care (P = 0.78). We calculated an MD of -7.00 Nm/s (95% CI -58.74 Nm/s to 44.74 Nm/s, Analysis 2.14) using data provided.



Sensitivity analysis

To assess the influence of our analysis model on the results, we repeated the pooled analyses using a fixed-effect model instead of a random-effects model. Using a fixed-effect model resulted in a change in the effect size for the effect of short-term muscle strength in 247 children and adolescents, with no change in heterogeneity (SMD 0.55, 95% CI 0.28 to 0.81, P < 0.001, I² = 70%, 8 studies; Dodd 2003; Liao 2007; Seniorou 2007; Lee 2008; Reid 2010; Scholtes 2010; Pandey 2011; Mitchell 2016; analysis not shown). Using a fixed-effect model instead of a random-effects model had no impact on intermediate-term muscle strength in children and adolescents, or on gait speed, gross motor function, or participation in children and adolescents at any time point.

Quality of evidence

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that resistance training improves walking endurance but does not improve short-term gross motor function, gait speed, physical activity, participation or quality of life in children and adolescents with CP. There is very low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision, once for consistency) that resistance training does improve short-term muscle strength in children and adolescents with CP. There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that resistance training improves muscle strength but does not improve short-term gait speed, gross motor function, walking endurance or physical activity in adolescents and young adults with CP. There is also low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that resistance training does not improve gait speed, gross motor function, walking endurance or muscle strength in adults with CP in the short term.

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that resistance training does not improve gait speed, gross motor function, walking endurance, physical activity or muscle strength in adolescents and young adults with CP in the intermediate term. There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that resistance training improves intermediate-term muscle strength but not gross motor function, gait speed or participation in children and adolescents with CP.

See Summary of findings 2.

Comparison 3: mixed training versus usual care

Four trials compared mixed training to usual care in children and adolescents with CP (Verschuren 2007; Fowler 2010; Chen 2012) or adolescents only (Unnithan 2007).

Activity

Pooled results

There was no statistically significant between-group difference in short-term gross motor function in 163 participants in GMFCS levels I to III receiving mixed training versus usual care (SMD 0.02, 95% CI -0.29 to 0.33, P = 0.90, I² = 0%, 4 studies, Analysis 3.1; Unnithan 2007; Verschuren 2007; Fowler 2010; Chen 2012).

Single study results

One trial, Fowler 2010, reported that mixed training did not improve self-selected short-term gait speed over 30 seconds (MD 0.10 m/s, 95% CI -0.07 m/s to 0.27 m/s, 58 children and adolescents, Analysis 3.2) or walking endurance, measured as the speed per distance completed on the 600-yard (548.6 m) walk test (MD 6.50 m/min, 95% CI -14.91 m/min to 27.91 m/min, 55 children and adolescents, Analysis 3.3) compared to usual care, in participants in GMFCS levels I to III.

One study, Verschuren 2007, reported that mixed training improved gross motor function, as measured by dimension D but not dimension E of the GMFM, in 65 children and adolescents (GMFCS levels I and II) in the intermediate term. When we calculated effect sizes using postintervention means and SDs, the effect size for dimension D was MD –2.62% (95% CI –6.26% to 1.03%) and dimension E was MD –4.90% (95% CI –11.58% to 1.78%). No trials reported intermediate changes in gait speed or walking endurance.

Participation

Pooled results

No meta-analysis was possible.

Single study results

One trial, Verschuren 2007, reported that short-term participation (as indicated by 'overall activities' scores for participation on the CAPE; King 2004) improved in 65 children and adolescents (GMFCS levels I and II) following mixed training compared to usual care (P = 0.002). We calculated an MD of 0.40 (95% CI 0.13 to 0.67, Analysis 3.4) using the data provided. There was no between-group difference in participation in the intermediate term (MD 0.15, 95% CI -0.25 to 0.54, Analysis 3.5).

Quality of life

Pooled results

No meta-analysis was possible.

Single study results

Two trials, Fowler 2010 and Verschuren 2007, reported quality of life following mixed training. One trial, Verschuren 2007, recorded quality of life in children and adolescents using the TNO-AZL Questionnaire for Children's Health-Related Quality of Life Parent Form (TACQOL-PF; Vogels 2000). Scale scores are obtained by adding items scores (between three and seven per scale) within scales and transforming crude scale scores to a 0to-100 scale. Quality of life was scored on seven scales (pain and symptoms, basic motor functioning, autonomy, cognitive functioning, social functioning, global positive emotions, and global negative emotions). The authors reported a statistically significant between-group difference in scores on the 'basic motor functioning' (P = 0.001) and 'cognitive functioning' (P = 0.04) subscales in the short term but did not report effect sizes. We calculated short-term effect sizes for 'basic motor functioning' (MD 3.80, 95% CI 1.71 to 5.89) and 'cognitive functioning' (MD 1.10, 95% CI -1.02 to 3.22); analyses not shown. The authors reported that there were no between-group differences on any scale in the intermediate term (Verschuren 2007).

Another trial, Fowler 2010, assessed quality of life in 58 children and adolescents (GMFCS levels I to III) using participant responses for

Exercise interventions for cerebral palsy (Review)

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the Pediatric Quality of Life Inventory SF15 (PedsQL; Chan 2005) and parent responses for the Pediatric Outcomes Data Collection Instrument (PODCI; Daltroy 1998). We extracted the total score from the PedsQL (scale 0 to 100, higher score indicates more positive quality of life) and the score from each of the four sections in the PODCI (global functioning and symptoms, happiness, treatment expectations, and satisfaction with symptoms). There was no between-group difference in participant-reported quality of life in the short term (MD 3.5, 95% CI –2.0 to 8.8). There was a between-group difference in score on one of five subscales (treatment expectations) on the PODCI in the short-term (MD 17.7, 95% CI 5.2 to 30.0).

Adverse events

Pooled results

No meta-analysis was possible.

Single study results

One trial, Unnithan 2007, did not report adverse events. One trial, Chen 2012, reported observing no adverse effects during the intervention for either group. One trial, Verschuren 2007, reported that one child fell and fractured her radius during an exercise session. One trial, Fowler 2010, reported observing 28 mild events (in 18 participants) potentially related to the study: 6 observed falls; 17 complaints of soreness, muscle cramping or mild pain; 4 reports of fatigue; and 1 skin rash related to wearing a heart-rate monitor. Fowler 2010 also noted 30 events unrelated to study procedures: illness, tooth loss, headache, stomachache, tonsillectomy and skin irritation from orthotic use.

Body structure and function

Pooled results

Mixed training did not result in improved short-term aerobic fitness in 78 children and adolescents (GMFCS levels I, II and II), compared to usual care (SMD 0.05, 95% CI –0.39 to 0.50, P = 0.81, $I^2 = 0\%$, 2 studies, Analysis 3.6; Unnithan 2007; Verschuren 2007).

Mixed training also did not improve short-term muscle strength in 150 children and adolescents with CP compared to usual care (SMD 0.08, 95% CI –0.24 to 0.40, P = 0.63, $I^2 = 0\%$, 3 studies, Analysis 3.7; Verschuren 2007; Fowler 2010; Chen 2012).

Single study results

One trial, Verschuren 2007 (65 children and adolescents), reported that anaerobic fitness improved in the short term (MD 25.20 W, 95% CI 8.89 W to 41.51 W, Analysis 3.8). The same trial reported that mixed training did not result in improvements in intermediate-term aerobic fitness (MD –0.13 min achieved on shuttle run/walk test, 95% CI –2.10 min to 1.84 min, Analysis 3.9) or anaerobic fitness (MD –31.28 W, 95% CI –71.89 W to 9.33 W, Analysis 3.10), but it did result in improved muscle strength. We calculated an effect size using postintervention means and SDs: MD –1.04 repetitions (95% CI –10.33 repetitions to 8.25 repetitions, Analysis 3.11).

Sensitivity analysis

Using a fixed-effect model rather than a random-effects model had no impact on the effect size for muscle strength or gross motor function among children (Verschuren 2007; Fowler 2010; Chen 2012). Analyses not shown. **Quality of evidence**

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that mixed training does not improve gross motor function, gait speed, walking endurance, aerobic fitness or muscle strength in children and adolescents with CP in the short term. There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that mixed training improves short-term participation and anaerobic fitness in children and adolescents with CP.

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that mixed training does not improve intermediate-term gross motor function, participation, aerobic fitness, anaerobic fitness or muscle strength in children and adolescents with CP.

See Summary of findings 3.

Comparison 4: resistance training versus aerobic exercise

Two trials, Johnston 2011 and Olama 2011, compared resistance training to aerobic exercise.

Activity

Pooled results

In two studies, the pooled results indicated there was no statistically significant between-group change in short-term gross motor function among 56 children and adolecents (SMD 0.02, 95% CI –0.50 to 0.55, P = 0.94, I^2 = 0%; Johnston 2011; Olama 2011).

Single study results

In one study in 26 children and adolescents (GMFCS levels II to IV), there was no statistically significant between-group change in short-term gait speed (MD 0.12 m/s, 95% CI -0.15 m/s to 0.39 m/s, Analysis 4.2; Johnston 2011), intermediate-term gait speed (MD 0.19 m/s, 95% CI -0.05 m/s to 0.43 m/s, Analysis 4.3) or intermediate-term gross motor function (MD 4.70%, 95% CI -12.70% to 22.10%, Analysis 4.4).

Participation

Pooled results

No meta-analysis was possible.

Single study results

One study in 26 children and adolescents, Johnston 2011, reported a between-group difference in participation as measured by the 'with whom' subscale of the CAPE (King 2004) in the short term (P = 0.05). As the authors did not provide an effect size, we calculated this using the data provided and found a between-group difference for the 'diversity' (MD 6.60, 95% CI 0.35 to 12.85) and 'with whom' (MD 0.48, 95% CI 0.08 to 0.88) subscales of the CAPE, but not for the subscales of 'intensity' (MD 0.48, 95% CI -0.22 to 1.18), 'where' (MD 0.28, 95% CI -0.07 to 0.63) or 'enjoyment' (MD -0.06, 95% CI -0.48 to 0.36). The authors did not report an effect size for between-group differences in participation in the intermediate term (Johnston 2011). Using the data provided, we calculated effect sizes for the following subscales: 'diversity' (MD 5.20, 95% CI -2.53 to 12.93), 'intensity' (MD 0.59, 95% CI -0.34 to 1.52), 'with whom' (MD 0.18, 95% CI -0.35 to 0.71), 'where' (MD 0.45, 95% CI -0.35 to 0.71), 'where' (MD 0.4

Exercise interventions for cerebral palsy (Review)

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-0.01 to 0.91) and 'enjoyment' (MD -0.02, 95% CI -0.55 to 0.51). Analyses not shown.

Quality of life

Pooled results

No meta-analysis was possible.

Single study results

One study of 26 children and adolescents, Johnston 2011, reported a group-by-time interaction for child-reported quality of life (P = 0.02) but did not state at what time point between-group differences occurred. Our analysis of the reported data indicated that there was no between-group difference in child-reported quality of life or adult-reported quality of life in the short term (child reported: MD 9.50, 95% CI –4.59 to 23.59; adult reported: MD 6.00, 95% CI –11.29 to 23.29) or intermediate term (child reported: MD –1.00, 95% CI –14.61 to 12.61; adult reported: MD 5.10, 95% CI –8.63 to 18.83). Analyses not shown.

Adverse events

Pooled results

No meta-analysis was possible.

Single study results

One trial did not state that adverse events were recorded (Olama 2011). Some children reported experiencing leg pain in one trial (Johnston 2011).

Body structure and function

Pooled results

There was no statistically significant between-group change in muscle strength in the short term (SMD -0.11, 95% CI -0.64 to 0.41, P = 0.67, I² = 0%, 56 children and adolescents, 2 studies, Analysis 4.5; Johnston 2011; Olama 2011).

No trial assessed aerobic fitness following the intervention.

Single study results

There was no statistically significant between-group change in muscle strength in the intermediate term (MD -0.03 N/kg, 95% CI -2.71 to 2.65 N/kg, 26 children and adolescents, 1 study, Analysis 4.6; Johnston 2011).

Sensitivity analysis

Repeating the analysis using a fixed-effect model instead of a random-effects model had no impact on the effect size or statistical significance of any outcome (analyses not shown).

Quality of evidence

There is low-quality evidence (RCT evidence: high, downgraded once for limitations, once for imprecision) that there is no difference between the effect of aerobic exercise and resistance training on gait speed, gross motor function, quality of life, or muscle strength in children and adolescents with CP in the short or intermediate term.

See Summary of findings 4.

Post hoc analyses

Folllowing identification of included studies, we noted a large overlap between the content of aerobic exercise interventions and mixed training interventions, and a large overlap between the content of resistance training interventions and mixed training interventions, respectively. We therefore conducted the following post hoc analyses: pooled analyses of aerobic exercise and mixed training versus usual care, and pooled analyses of resistance training and mixed training versus usual care. These analyses only include trials in children and adolescents, as no mixed training trials included adults with CP. Given the post hoc nature of these analyses, readers should interpret results with caution.

Comparison 1: aerobic exercise (incorporating mixed training interventions) versus usual care

Pooled results

In a meta-analysis of seven studies, aerobic exercise improved short-term gross motor function more than usual care in 228 children and adolescents (SMD 0.36, 95% CI 0.09 to 0.62, P = 0.008, $I^2 = 0\%$, Analysis 5.1; Unnithan 2007; Verschuren 2007; Fowler 2010; Chen 2012; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013). However, a meta-analysis of two studies (77 children and adolescents) showed that aerobic exercise did not improve gross motor function in the intermediate term (SMD 0.25, 95% CI –1.15 to 1.64, P = 0.73, $I^2 = 77\%$, Analysis 5.2; Verschuren 2007; Mattern-Baxter 2013).

In a meta-analysis of five studies (140 children and adolescents), aerobic exercise did not result in an improvement in gait speed (MD 0.10 m/s, 95% Cl –0.05 m/s to 0.24 m/s, P = 0.18, $l^2 = 70\%$, Analysis 5.3; Fowler 2010; Gharib 2011; Smania 2011; Chrysagis 2012; Mattern-Baxter 2013). In addition, aerobic exercise did not improve short-term walking endurance in two studies of 73 children and adolescents (MD 8.43 m, 95% Cl –12.38 m to 29.23 m, P = 0.43, $l^2 = 0\%$, Analysis 5.4; Fowler 2010; Smania 2011). There was also no between-group difference in short-term aerobic fitness following aerobic exercise in three studies of 98 children and adolescents (SMD 0.06, 95% Cl –0.34 to 0.45, P = 0.78, $l^2 = 0\%$, Analysis 5.5; Van den Berg-Emons 1998; Unnithan 2007; Verschuren 2007).

Sensitivity analysis

Using a fixed-effect model rather than a random-effects model had no impact on the short-term results for aerobic fitness, gross motor function or walking endurance. There was, however, a change in the short-term effect size for gait speed (MD 0.08 m/s, 95% CI 0.02 m/ s to 0.14 m/s, P = 0.006, I² = 70%, 140 participants). Using a fixedeffect model also had a small impact on the intermediate effect size for gross motor function (SMD –0.16, 95% CI –0.62 to 0.29, P = 0.49, I² = 77%, 228 participants). Analyses not shown.

Comparison 2: resistance training (incorporating mixed training interventions) versus usual care

Pooled results

A meta-analysis found that resistance training did not improve gross motor function in the short term (SMD 0.21, 95% CI –0.01 to 0.43, P = 0.06, I^2 = 0%, 327 children and adolescents, 11 studies, Analysis 6.1; Dodd 2003; Engsberg 2006; Liao 2007; Seniorou 2007; Unnithan 2007; Verschuren 2007; Lee 2008; Fowler 2010; Scholtes 2010; Chen 2012; Lee 2015) nor in the intermediate term (SMD

Exercise interventions for cerebral palsy (Review)

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-0.08, 95% CI -0.41 to 0.24, P = 0.62, I² = 1%, 150 children and adolescents, 4 studies Analysis 6.2; Dodd 2003; Verschuren 2007; Lee 2008; Scholtes 2010).

Similarly, a meta-analysis of nine studies involving 243 children and adolescents found that resistance training did not improve short-term gait speed (MD 0.03 m/s, 95% CI –0.01 m/s to 0.08 m/s, P = 0.13, $I^2 = 0\%$, Analysis 6.3; Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Seniorou 2007; Lee 2008; Fowler 2010; Scholtes 2010; Pandey 2011).

Resistance training improved participation in the short term (SMD 0.35, 95% CI 0.07 to 0.64, P = 0.02, I^2 = 0%, 192 children and adolescents, 3 studies, Analysis 6.4; Verschuren 2007; Scholtes 2010; Mitchell 2016) but not the intermediate term (MD 0.15, 95% CI –0.24 to 0.54, P = 0.46, I^2 = 0%, 101 children and adolescents, 2 studies, Analysis 6.5; Verschuren 2007; Scholtes 2010).

Resistance training improved muscle strength more than usual care in the short term (SMD 0.38, 95% CI 0.01 to 0.76, P = 0.04, I² = 66%, 397 children and adolescents, 11 studies, Analysis 6.6; Dodd 2003; Liao 2007; Seniorou 2007; Verschuren 2007; Lee 2008; Fowler 2010; Reid 2010; Scholtes 2010; Pandey 2011; Chen 2012; Mitchell 2016) but not in the intermediate term (SMD 0.28, 95% CI –0.16 to 0.71, P = 0.21, I² = 37%, 149 children and adolescents, 4 studies, Analysis 6.7; Dodd 2003; Verschuren 2007; Lee 2008; Scholtes 2010).

Sensitivity analysis

Using a fixed-effect model rather than a random-effects model impacted the effect sizes for strength in the short term (SMD 0.36, 95% CI 0.16 to 0.56, P = 0.001, I^2 = 66%, 397 participants) and intermediate term (SMD 0.25, 95% CI –0.08 to 0.58, P = 0.13, I^2 = 37%, 149 participants). Analyses not shown.

Summary of post hoc analyses

In agreement with the results of the analysis of aerobic exercise only compared to usual care, the pooled analysis showed some evidence that in children and adolescents with CP, aerobic exercise improves gross motor function in the short – but not intermediate – term, and no evidence that it improves short-term gait speed, walking endurance or aerobic fitness.

The pooled analysis of resistance training and mixed training compared to usual care showed similar results to the analysis of resistance training only in terms of some evidence of an improvement in strength in the short term and no improvement in gross motor function or gait speed in the short term or strength in the intermediate term. The pooled analysis, however, found that participation improved following resistance training in the short term, which differed from findings in the primary analysis.

DISCUSSION

Summary of main results

Overall, we are unable to provide firm conclusions regarding the effectiveness of exercise interventions for people with CP due to the poor quality of the available evidence, which was exacerbated by small sample sizes and, in turn, probably led to an underestimation of the extent of heterogeneity. As a result, there is substantial uncertainty related to the effect estimates presented in this review.

The results of the included trials suggest that aerobic exercise may result in a small improvement in gross motor function in the short and intermediate term and may improve physical activity. There is low- to very low-quality evidence that aerobic exercise does not improve gait speed, walking endurance, or aerobic fitness, although these findings are mostly based on single studies. Generally, the results suggest that resistance training does not improve any aspect of activity or participation in people with CP but may improve muscle strength in children, adolescents and young adults in the short term and in children and adolescents in the intermediate term. In addition, a trial of 101 participants reported that resistance training improves walking endurance in children with CP in the short term (Mitchell 2016). Mixed training may improve participation, but at present, there is low-quality evidence that it does not improve activity, muscle strength or aerobic fitness in children with CP. These conclusions, however, are largely based on the results of a single study (Verschuren 2007). Although only two trials compared resistance training and aerobic exercise in children and adolescents with CP (Johnston 2011; Olama 2011), the results indicate that there is no difference in terms of gait speed, gross motor function, quality of life or muscle strength between types of exercise in the short or intermediate term. Based on the data provided, exercise appears to be safe for people with CP, with no serious adverse events reported. The evidence for safety, however, is also incomplete because only 16 trials reported that they collected data on adverse events; it is unclear whether or not this reflects an absence of adverse events or a failure to report them. A small number of participants reported experiencing events potentially related to the intervention, including muscle soreness, mild knee, foot and ankle pain, shoulder pain from wearing weighted vests, fatigue, minor calf strain and falls (one of which resulted in a fracture).

Overall completeness and applicability of evidence

The evidence base for the use of exercise interventions for people with CP is incomplete. There is a lack of trials examining the effectiveness of both aerobic and resistance exercise in adults with CP. There is also a lack of trials investigating the intermediateand long-term effectiveness of exercise for people with CP; the longest follow-up time point was four months postintervention (Verschuren 2007). Given the decline in mobility among young adults with CP (Bottos 2001; Day 2007; Opheim 2009), the effectiveness of exercise for slowing or preventing decline in activity and participation in the long term is a real concern (Shortland 2009). Few trials investigated the effectiveness of exercise for nonambulatory people with CP. Only 5 out of 23 trials that included an intervention that targeted the lower limbs included children and adolescents in GMFCS levels IV or V (Van den Berg-Emons 1998; Johnston 2011; Smania 2011; Bryant 2013; Tedla 2014), and only one trial included children and adolescents in GMFCS levels IV and V only (Bryant 2013).

The evidence for the effectiveness of interventions at the level of participation on the *International Classification of Functioning*, *Disability and Health* (ICF) is incomplete (WHO 2001), with only 4 out of 29 included trials assessing participation (Verschuren 2007; Scholtes 2010; Johnston 2011; Mitchell 2016). Although three of these trials assessed participation using the CAPE, one trial reported the individual scores for each subscale of the CAPE, rather than the total score, making it difficult to pool results. Similarly, the evidence for the effect of exercise interventions on quality

Exercise interventions for cerebral palsy (Review)

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of life is incomplete. Of the four trials that assessed quality of life in children and adolescents, three assessed both participantand parent-reported quality of life, often with contradictory results (Engsberg 2006; Fowler 2010; Johnston 2011). Further, two trials reported the individual results for several subscales of the outcome measure and not an overall score, making it difficult to draw any firm conclusions regarding the impact of exercise on quality of life (Verschuren 2007; Fowler 2010).

Most trials assessed activity capacity, activity performance or both, with most assessing capacity rather than performance. The most commonly used measures of activity capacity were the GMFM and self-selected gait speed. There was variation, however, in the component of the GMFM that trials assessed and reported (e.g. dimension E only, dimension D and E combined, total score) and in the methods they used to assess self-selected gait speed (e.g. walking on a treadmill or walking over ground). Further, as a number of trials used the GMFM to assess motor capacity in adolescents outside of the age range for which the tool was developed (i.e. five months to 16 years) (Dodd 2003; Unnithan 2007; Verschuren 2007; Chrysagis 2009; Chrysagis 2012; Bryant 2013; Taylor 2013), the score may not accurately reflect participants' activity capacity. Perhaps unexpectedly, a number of trials did not assess physical fitness, particularly aerobic fitness; 3 of 14 trials assessed aerobic fitness following an aerobic exercise intervention, and 15 of 21 trials assessed muscle strength following a resistance training intervention.

There was consistently incomplete reporting of the comparator across trials. Although most trials reported that the control group received usual physiotherapy, reporting of the content and dose of usual care was poor. Reported physiotherapy varied considerably across individuals, and it was not clear from many trials if participants received the dose and content of physiotherapy prescribed.

We were unable to locate or extract data from three trials (Chrysagis 2009; Bryant 2013; Tedla 2014), which may have had an impact on the results of our meta-analysis. In particular, the inclusion of data on muscle strength and gross motor function from Tedla 2014 would have increased the power of the analysis through the addition of a further 62 participants. Data from Bryant 2013 were only missing for the intermediate time point, but data on gross motor function from 31 participants in this trial may have impacted the outcome of the meta-analysis.

The most effective dose of exercise for people with CP is currently unknown, making it difficult to prescribe exercise in this population. This is reflected in the large variation in the frequency, duration and intensity of exercise prescribed. There is a body of evidence supporting the effectiveness of exercise in general to improve aerobic fitness and muscle strength, which has contributed to the development of exercise prescription guidelines for the general population (American College of Sports Medicine 2009; Faigenbaum 2009; Garber 2011), and more recently, for people with CP (Verschuren 2016). While many included trials failed to report the volume and intensity of exercise prescribed, those that did report the dose of exercise prescribed did not meet current guidelines for exercise prescription for people with CP (Verschuren 2016). For example, of the 11 trials included in the pooled analysis of resistance training and mixed training in comparison to usual care, six were prescribed for a shorter duration than that suggested in guidelines (Dodd 2003; Liao 2007;

Seniorou 2007; Lee 2008; Reid 2010; Pandey 2011), four trials either did not report the number of sets and repetitions performed or the number of sets and repetitions were lower than guideline recommendations (Verschuren 2007; Fowler 2010; Pandey 2011; Mitchell 2016), and five trials either did not report the intensity of the exercises or the intensity was lower than current guideline recommendations (Liao 2007; Verschuren 2007; Lee 2008; Scholtes 2010; Pandey 2011). Similarly, of the 14 trials that included aerobic exercise in the intervention, 12 did not state the intensity of exercise prescribed (Van den Berg-Emons 1998; Verschuren 2007; Chrysagis 2009; Gharib 2011; Johnston 2011; Olama 2011; Smania 2011; Chen 2012; Chrysagis 2012; Bryant 2013; Mattern-Baxter 2013; Emara 2015). It is possible that inadequate prescription of exercise volume and intensity across trials contributed to the limited or null improvement in physical fitness. At present, however, there is insufficient evidence to suggest that higher doses of exercise, in line with the current guidelines for people with and without CP (American College of Sports Medicine 2009; Faigenbaum 2009; Garber 2011, Verschuren 2016), result in improvements in physical fitness, activity or participation in people with CP. Further, the feasibility of delivering these higher doses of exercise to people with CP is not clear. There was variation in fidelity to the intervention among trials that reported this measure, perhaps because it was not feasible to deliver the dose prescribed in this population. This would suggest that failure to implement exercise interventions as planned contributed to the findings of this review. The findings of this review highlight the need for clearer reporting of the frequency, intensity and duration of exercise prescribed in future trials and further investigation into the effect and feasibility of following generic exercise guidelines for people with CP.

Quality of the evidence

Using the GRADE criteria (Guyatt 2008), we rated the quality of evidence for all comparisons as low or very low, mainly as a result of an overall high risk of bias for all trials and issues of imprecision.

We judged all trials as being at high risk of bias for blinding of participants. Unfortunately, while this is unavoidable when examining the effect of exercise in a trial, it does not negate the risk of introducing performance bias by not blinding participants and personnel. We also judged a large number of trials as being at high or unclear risk of bias for blinding of outcome assessment. It was recently reported that less than a quarter of physical therapy trials were adequately blinded and that blinding was poorly reported in trials (Armijo-Olivo 2017). Our decision to classify studies that used a self-report outcome as being at high risk of bias inflated the number of trials at high risk for blinding of outcome assessment. We believe that, although conservative, classifying these trials as being at high risk of bias was the correct decision given that lack of, or unclear, double blinding is associated with an average 22% exaggeration of intervention effects when assessing self-report outcomes (Savović 2012). However, lack of blinding does not appear to exaggerate the intervention effect when assessing objective outcomes (Wood 2008; Savović 2012). Readers should therefore interpret the results for objective outcomes in these trials (judged at high risk of bias due to the inclusion of subjective outcomes) as being at low risk of detection bias. We judged most trials as being at unclear risk of bias for random sequence generation and allocation concealment; a recent study reported that only 39.7% and 11.5% of physical therapy trials achieved adequate random sequence generation and allocation

Exercise interventions for cerebral palsy (Review)

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concealment, respectively (Armijo-Olivo 2015). Intervention effect estimates may be exaggerated by 11% in trials with inadequate or unclear sequence allocation and by 15% for trials with unclear or inadequate allocation concealment (Savović 2012). Effect sizes are exaggerated even further for self-report outcomes whereas there is little evidence that inadequate or unclear sequence generation or allocation concealment exaggerates the intervention effect in trials with objective outcomes (Wood 2008; Savović 2012). A number of trials reported using sealed envelopes to generate a random sequence, which in itself appears to adequately prevent bias when compared to central randomisation (Herbison 2011). However, several of the trials in this review did not report enhanced security, such as use of opaque envelopes; intervention effects are exaggerated by, on average, 13% for trials that state that they use envelopes but do not give details of enhanced security (Herbison 2011).

Only four trials published a trial protocol prior to the publication of results. In addition, some of these trials were not the most recently published trials, suggesting that this issue is not improving with time. Chan 2004 found that statistically significant outcomes were 2.4 times more likely to be fully reported compared to non-statistically significant outcomes. Although we judged most included trials as being at low risk of bias for incomplete outcome data, because of low levels of missing data, very few trials stated that they performed an intention-to-treat analysis. Excluding participants from the analysis can result in biased estimates of treatment effects (Nüesch 2009; Abraha 2015), although the direction of bias is unpredictable (Nüesch 2009).

Finally, 24 of the 29 included trials had small sample sizes (fewer than 50 participants), and as a result, data from a relatively small number of participants (65 to 397 participants) were included in pooled analyses. The small number of participants included in analyses may have impacted our findings in one of two ways, either inflating effect sizes or resulting in statistically non-significant findings (Pereira 2012; Button 2013; Dechartres 2013). Effect sizes calculated on samples of fewer than 50 participants are, on average, 23% larger than estimates from trials with sample sizes of more than 50 participants (Dechartres 2013). However, the relatively small number of participants included in the analysis may have resulted in analyses that lacked statistical power. In addition, the small number of participants led to large within-study variations and subsequently low between-study variations; in many analyses heterogeneity was estimated as zero. Therefore, although the small sample sizes in trials may have resulted in analyses that lacked statistical power, our analyses likely underestimated the degree of heterogeneity, so we may have observed erroneously significant results.

Potential biases in the review process

We conducted an extensive search of all available literature, including grey literature. We included trials regardless of publication date or language and are therefore confident that this review includes all published evidence on the topic to date.

A broad and varied number of interventions may be described as exercise. The inclusion of certain interventions such as those described as 'gait training' or 'treadmill training' may be contentious. In order to ensure consistency when deciding if an intervention was classified as aerobic or resistance training, we referred to the definitions of exercise published by Caspersen

1985 and the US Department of Health and Human Services (USDHHS 2008). Repetitive movement of the lower limbs during walking on a treadmill or over ground meets the description of aerobic exercise, regardless of the support required to perform this repetitive movement. Similarly, we included any mode of exercise that involved repetitive movement of large muscle groups over a sustained period of time. For example, we included a swimming intervention where it was clear that participants swam continuously for a sustained period. We only included studies of resistance training if it was clear that the body's muscles were working or holding against an applied force, such as body weight, free weights, machine weights, or elastic bands (USDHHS 2008). We did not include studies that provided an intervention in addition to exercise, such as botulinum toxin, motivational interviewing or ankle foot orthoses, as it was not possible to distinguish the independent effect of exercise on the outcomes assessed.

We included postintervention means and SDs in our meta-analysis if they were presented in the published report. We also calculated effect sizes for single studies where the effect size was not provided using postintervention means and SDs. This may have resulted in conservative estimates of the effect size as there was significant heterogeneity of the baseline scores in many studies. It may also explain the difference between the P values and CIs for effect estimates reported in some trials and those that we calculated.

Three authors are chartered physiotherapists and lecturers in physiotherapy. As professionals who might be involved in the delivery of exercise interventions, it is plausible that they might be perceived as having a bias favouring the effectiveness of exercise.

Agreements and disagreements with other studies or reviews

The results of this review are generally inconsistent with the results of previous reviews primarily because of differences in the number and type of included trials and because most previous reviews used narrative syntheses when drawing conclusions. Most authors accepted the statistical significance of comparisons reported in individual trials as indication of a positive effect. Rogers 2008 and Butler 2010 reviewed the literature on the effects of aerobic exercise for children with CP. Rogers 2008 presented the evidence from three trials, one RCT that was included in this review (Van den Berg-Emons 1998), one cohort study and one RCT that we did not include in this review because the only difference between the intervention and control group was that the intervention group received active encouragement. Using a narrative synthesis of the literature, the authors concluded that aerobic exercise improved aerobic fitness and that no study assessed activity (Rogers 2008). Similarly, using a narrative synthesis of three trials included in this review (Van den Berg-Emons 1998; Unnithan 2007; Verschuren 2007), Butler 2010 concluded that aerobic exercise may increase aerobic fitness but there was not enough evidence to indicate that it improved activity.

Two meta-analyses assessed the effectiveness of resistance training for children and children and adults with CP (Scianni 2009; Park 2014b). Scianni 2009 reported that there was no short- or intermediate-term improvement in strength, gross motor function or gait following resistance training in children with CP. Two of the six included studies are not included in the current review because they used electrical stimulation (Van der Linden 2003; Kerr 2006). Park 2014b reported that there was strong evidence for a large effect of strengthening interventions in people with

Exercise interventions for cerebral palsy (Review)

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CP, based on the results of 12 trials described in 13 reports. We included 11 of these trials in our review (Dodd 2003; Engsberg 2006; Unger 2006; Liao 2007; Unnithan 2007; Lee 2008; Maeland 2009; Fowler 2010; Scholtes 2010 (two reports); Chen 2012). However, similar to Scianni 2009, Park 2014b also included two trials that assessed the effect of electrical stimulation (Van der Linden 2003; Kerr 2006). Although Park 2014b did not report a clear plan for the meta-analysis, it appears that the conclusion was based on a standardised effect size calculated by pooling all outcomes from all trials; this may explain the discrepancy between their results and ours. Further, although subgroup analyses were conducted according to outcome, it is not clear which trials contributed to each subgroup analysis, and it appears that more than one outcome measure from each trial was included in a single meta-analysis. Park 2014b also combined the results of trials involving children and adults with CP, respectively.

A third systematic review investigated the effect of upper limb resistance training on all levels of the ICF Framework in children with CP (Rameckers 2014). Using a narrative synthesis, the authors concluded that strength training had a small to large effect on strength and that no study assessed the effect on activity or participation. These conclusions were informed by two trials included in the current review (McCubbin 1985; Reid 2010). However, we did not believe McCubbin 1985 assessed muscle strength and therefore did not include it in the meta-analysis.

In contrast to our findings, a review of all physiotherapy interventions for adults with CP concluded that there was moderate-quality evidence that strength training improves gait in adolescents and adults with CP (Jeglinsky 2010). However, this conclusion was based on the results of one RCT, Unger 2006, and therefore does not represent a thorough review of the literature as presented here.

Only one review to date has summarised the evidence for aerobic, anaerobic and resistance training for children with CP (Verschuren 2008). In contrast with our findings, the authors concluded that exercise may improve strength and aerobic capacity but that more trials are needed to determine the effects on activity, participation and quality of life. Authors drew these conclusions using a narrative synthesis of the results from both randomised and non-randomised trials that took place prior to September 2006 and therefore do not represent the most recent or best quality evidence.

AUTHORS' CONCLUSIONS

Implications for practice

This review shows that there is low- to very low-quality evidence that aerobic exercise results in a small improvement in gross motor function but not aerobic fitness, and that resistance training results in a small improvement in muscle strength but not activity or participation in people with CP. Exercise appears to be safe for people with CP from the limited evidence available.

Historically, rehabilitation efforts for children with disabilities have focused on improving impairments in body structures and function in order to improve activity (Rosenbaum 2012). The results of this review do not support the hypothesis that improving impairments will improve activity, as we found no correlation between improvements in physical fitness and improvements in activity following an exercise intervention. The ICF framework highlights the complex interplay between body structures and functions, activity, participation, and environmental and personal factors (WHO 2001), and personal and environmental factors are likely to be key explanatory factors in this divergence between the effect on impairments and activity limitations.

Aerobic exercise may be important to improve activity capacity in terms of gross motor function, but for now it is not clear whether it improves activity performance. A gap exists between activity capacity and performance (Holsbeeke 2009), which raises an interesting question regarding the focus of exercise (i.e. to improve what a person can do in a controlled environment or to improve what a person actually does in their environment). Promoting participation in physical activity and aerobic exercise in general may be more important than prescribing an exercise programme for a fixed duration, not only for improving activity capacity, as demonstrated by the results of this review, but also for improving activity performance. Also, achieving a sustained impact on activity capacity through aerobic exercise is likely to require frequent participation and hence increased activity performance.

The studies included in this review show that exercise can be delivered through a range of modes, in a range of settings and by a range of providers. It is important not to think of exercise solely as therapy. While we were unable to examine the impact of different modes of delivery of exercise, such as group sessions, on outcomes in the current review, people, and particularly children, may be more likely to engage in exercise over their lifespan if they enjoy it, if they believe it has health benefits, and if they perceive it to be an opportunity for social interaction (Verschuren 2012). The studies included in this review used a wide variety of modes of aerobic exercise, such as cycling on a static bike at home, cycling on a static adapted bike in school, supported and unsupported treadmill walking, swimming and wheelchair driving, indicating that the type of exercise prescribed can be adapted according to the person's ability and preference. For resistance training, free weights or weight machines may increase the intensity of exercise, especially in adolescents and adults. This will likely require identifying opportunities to access this type of equipment in the community.

Implications for research

The current evidence for exercise in people with CP is comprised of mostly small studies at high or unclear risk of bias. There is an urgent need for larger, more rigorous and more completely reported RCTs. Further, a number of key issues need to be addressed by future trials before any firm conclusions can be made regarding the effectiveness of exercise interventions for people with CP.

With regard to trial reporting, exercise interventions need to be prescribed, assessed, and adapted in a systematic fashion to enable development of an evidence-based guideline specific to people with CP. Reporting fidelity to the intervention, in terms of the content as well as the dose of exercise, should be a prerequisite when publishing trials on the effectiveness of exercise interventions for people with CP in order to differentiate between intervention failure and implementation failure. We recommend that all authors of trials investigating exercise interventions in people with CP use the recently developed *Template for Intervention Description and Replication Checklist* (TIDieR) to ensure that they describe interventions in sufficient detail to allow

Exercise interventions for cerebral palsy (Review)

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replication (Hoffmann 2014). Adherence to this standard will also aid the implementation process of interventions that show clinical benefits. The research community should also reach a consensus on the content of the comparator when evaluating the effectiveness of exercise interventions for people with CP. If comparing the intervention to usual care, investigators should track and report the content and dose for both the intervention and comparison group. We acknowledge that usual care, particularly physiotherapy, can vary widely between countries, medical centres, and even health professionals in the same centre, and therefore it may not be possible to control its content and dose. However, clear guidance on how to record and report usual care will greatly improve the transparency of future trials.

With regard to trial design, future trials need to assess the effect of exercise interventions on quality of life and adverse events in people with CP on all levels of the ICF (WHO 2001). Guidance is urgently required regarding choice of outcome measures and reporting of data to allow for comparison across trials. Amongst the few trials that assessed participation and quality of life, there was inconsistency regarding the outcome measures used and the data reported, preventing pooled analysis of individual trials. Participation is an important outcome of any intervention for people with CP, and currently the impact of exercise on participation is not well understood because studies have not used outcome measures that specifically measure participation. Further, we propose that all trials assess activity capacity and activity performance. These are potentially different constructs (Holsbeeke 2009), but both are important to people with CP, their families and health professionals. While there were relatively few (and minor) adverse events that were potentially related to the intervention, the literature does not confirm that no serious adverse events occurred as a result of exercise, as approximately half of the included studies did not report monitoring this outcome. Implementation of a standardised method of recording and reporting adverse events would ensure more consistent and deliberative reporting. Further, trials need to include measures of body structures and functions in order to determine the likely process of change in activity, participation and quality of life. At present, exercise is believed to produce similar neuromuscular and physiological adaptations in people with and without CP. This assumption merits further examination in order to develop evidence-based exercise prescription guidelines for people with CP.The results of this review are only representative of a subgroup of people with CP, as most included studies in children and adolescents fall into GMFCS levels I to III. We identified a stark dearth of studies involving adults with CP and people with moderate-to-severe CP. There are evident difficulties entailed in recruiting adults with CP into clinical trials, and people with moderate-to-severe CP may be not even be able to complete an exercise intervention as prescribed, so a feasibility study may be necessary to determine the appropriateness of trials on exercise interventions in these populations. It is essential, however, that the research community makes efforts towards identifying and overcoming these associated difficulties to prevent discrimination against this group of people.

With regard to future priorities, studies should investigate the longterm effects of exercise on function and health in people with CP. A higher prevalence of a number of conditions such as pain, fatigue, depression, cardiovascular disease, type 2 diabetes mellitus and hypertension has been reported in adults with CP in comparison to the general population (Jahnsen 2004; Opheim 2009; Van Der Slot 2012; Peterson 2015). They also report experiencing a decline in muscle strength, aerobic fitness and mobility in young adulthood (Opheim 2009). The benefits of exercise for people with CP may be in the maintenance of physical fitness, activity and participation, and in the prevention of chronic disease, rather than the improvement of function alone. Future trials need to include long-term follow-up in order to thoroughly examine the effects of exercise throughout the lifespan of a person with CP.

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Exercise interventions for cerebral palsy (Review)



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CHARACTERISTICS OF STUDIES

Characteristics of included studies [author-defined order]

McCubbin 1985

McCubbin 1985	
Methods	Design: randomised controlled trial
	Country of origin: USA
	Intervention(s): resistance training
	Unit of allocation: individual
Participants	Number of participants : 30 participants; 20 participants randomised to a resistance training interven- tion or repetitive movement exercise without resistance; 10 participants selected for matched control group not reported in this review
	Resistance exercise group n = 10; no resistance exercise group n = 10
	Age: 10-20 years. Mean age not reported.
	Sex: not reported
	Ethnicity: not reported
	GMFCS level: not reported
	MACS level: not reported
	Type of motor abnormality : spastic (n = 15), athetoid-ataxic (n = 1), spastic-athetoid (n = 1), mixed (n = 2), and athetoid (n = 1); resistance exercise group: spastic (n = 7), athetoid-ataxic (n = 1), spas-tic-athetoid (n = 1), mixed (n = 1); no resistance exercise group: spastic (n = 8), athetoid (n = 1), mixed (n = 1)
	Anatomical distribution of CP: not reported
	Inclusion criteria: not reported

Young 2000

Østensjø 2004

Ryan 2015

j.1469-8749.2004.tb01021.x]

10.1002/14651858.CD011660]

* Indicates the major publication for the study

Exclusion criteria: not reported

Exercise interventions for cerebral palsy (Review)

McCubbin 1985 (Continued)	
Interventions	Aim of the intervention: not reported
	Type of exercise programme: resistance training
	Exercise mode: elbow extension exercise using Super Mini-Gym Model 180, Mini Gym, Inc.
	Comparator : identical protocol of elbow extension exercise without resistance. All participants participated in regularly scheduled school activities, which included physical education, physical therapy, oc- cupational therapy, speech therapy, and typical school classes. Subjects using medication prior to the experiment continued to use the medication recommended by their physician.
	Setting: not reported
	Intervention provider: primary researcher or registered physical therapist
	Duration of programme: 6 weeks
	Exercise dose: 3 sets of 10 repetitions, 3 days per week. Intensity not reported
	Tailoring of intervention to individual: not reported
	Fidelity to prescribed intervention: not reported
	Monitoring of adverse events: not reported
Outcomes	Assessment time points: baseline (week 0), interim (week 3), postintervention (week 6)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	 Movement time measured using a movement time analyser to measure time of hand movement dur- ing 90 degrees of elbow extension. 10 scores were recorded and the mean score (in milliseconds) was used in analysis
	2. The rate of torque development of elbow extension was evaluated using the Cybex II Isokinetic Sys- tem. 3 maximal trials were performed with the highest torque recorded used for the analysis
Notes	Source of funding: not reported
	Potential conflicts of interest: not reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : the participants were "matched according to type and severity of cere- bral palsy according to the classification system employed by the Nation- al Association of Sport for Cerebral Palsy Following the classification the matched experimental subjects were randomly selected to either experimen- tal group".
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : the participants were "matched according to type and severity of cerebral palsy according to the classification system employed by the National Association of Sport for Cerebral Palsy Following the classification the matched experimental subjects were randomly selected to either experimental group".
		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement

Exercise interventions for cerebral palsy (Review)

McCubbin 1985 (Continued)

Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Comment : did not provide information to indicate if the assessor was blinded to group allocation or not
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : no missing data
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment: no demographic data provided

Methods	Design: randomised controlled trial	
	Country of origin: Netherlands	
	Intervention(s): aerobic exercise	
	Unit of allocation: individual	
Participants	Number of participants : 20 randomised; exercise group: n = 10; control group: n = 10	
	Age : mean (SD) 9.2 (1.4) years; exercise group: mean (SD) = 9.5 (1.6) years; control group: mean (SD)= 8.8 (1.1) years	
	Sex (male/female): 11/9	
	Ethnicity: authors reported 19 white participants, 1 participant born in Sri Lanka	
	GMFCS level : ambulant (i.e. GMFCS levels I, II or III; n = 10) and 'wheelchair bound' (i.e. GMFCS levels IV or V; n = 10)	
	Type of motor abnormality: spastic CP, 2 participants additionally had ataxia	
	Anatomical distribution of CP : diplegia (n = 16) and tetraplegia (n = 4)	
	Inclusion criteria: age 7-13 years, diagnosis of spastic CP	
	Exclusion criteria: not stated	
Interventions	Aim of the intervention : to increase daily physical activity, aerobic power, anaerobic power and mus- cle strength	
	Type of exercise programme: aerobic	
	Exercise mode : aerobic exercises such as cycling, wheelchair driving, running, swimming, training on a "flying saucer" and mat exercises	
	Comparator : all participants participated in a normal school and therapy programme. School pro- gramme included 2, 45-min gymnastic lessons per week. Therapy was based on personal needs and varied from no therapy to more than 2.5 h per week	

Exercise interventions for cerebral palsy (Review)

Van den Berg-Emons 1998 (Co	ntinued) Setting: school		
	Intervention provider: not stated		
	Duration of programme : 18 months. After 9 months the participants in both groups were given the opportunity to participate in the next training programme of 2 times per week (45 min per session). 8 participants in exercise group and all participants in control group chose to participate. Data only extracted for first 9 months as after this participants were not randomised to a group		
	Exercise dose : 4 sessions per week, 45 min per session. Heart rate (HR) was measured randomly to get an indication of training intensity. Mean (SD) HR during training sessions in first 2 months was mean (SD) 135 (10) bpm; mean (SD) percentage of time spent at ≥ 70% of HR reserve during training was 49 (17) %		
	Fidelity to prescribed intervention: attendance was 84% (range 78 to 88%) in first 2 months		
	Monitoring of adverse events: not reported		
Outcomes	Assessment time points: baseline (week 0) and postintervention (9 months)		
	Primary outcome: no primary outcome measure stated		
	Outcomes:		
	1. Physical activity was calculated as the ratio of total energy expenditure to sleeping or resting meta- bolic rate. Sleeping metabolic rate was measured in participants who stayed overnight in a respiration chamber. Resting metabolic rate was measured using a ventilated hood. Total energy expenditure was calculated from heart rate using the heart rate flex principle. Participants wore a heart rate mon- itor continuously from 9 am until they went to bed on one day.		
	2. Fat mass was determined using the thickness of 4 skinfolds (biceps, triceps, subscapular and suprail- iac) using a callipers. Reported in kg.		
	3. Aerobic fitness was determined by peak aerobic power (reported in W per kg fat free mass) measuring during a maximal graded exercise test on a cycle or arm crank ergometer. Protocol not described.		
	4. Anaerobic fitness was determined by peak anaerobic power and mean anaerobic power measured during an exercise test on a cycle or arm crank ergometer. Protocol not described. Reported in W per kg fat free mass.		
	5. Muscle strength of the knee extensors and flexors was measured using an isokinetic dynamometer at 30°/s. Reported in Nm.		
Notes	Source of funding: not stated		
	Potential conflicts of interest: not stated; none perceived		

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[a]t the beginning of the first year (in September, after the summer holidays), the children were matched pairwise for physical ability, mental function, and if possible, for age, gender and body composition. After matching, the children of each pair were randomly assigned to an experimental group (EXP, n=10) or control group (CON, n=10)".
		Comment : insufficient information regarding the method of sequence alloca tion was provided to make a judgement.
Allocation concealment (selection bias)	Unclear risk	Quote : "[a]t the beginning of the first year (in September, after the summer holidays), the children were matched pairwise for physical ability, mental function, and if possible, for age, gender and body composition. After matching, the children of each pair were randomly assigned to an experimental group (EXP, n=10) or control group (CON, n=10)".

Exercise interventions for cerebral palsy (Review)



Van den Berg-Emons 1998 (Continued)

		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Comment : no information is provided regarding who conducted assessments
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[t]he project lasted 2 years and included two training periods of 9 months".
		Comment : only data from the first 9 month training period is reported as par- ticipants were not randomly allocated to a training group or control group for the second 9 month training period. No missing data for first 9 month training period
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes.
Other bias	Low risk	Comment : no other sources of bias identified

Dodd 2003

Juu 2005			
Methods	Design: randomised controlled trial		
	Country of origin: Australia		
	Intervention(s) : resistance training programme of the ankle plantarflexors, knee extensors, and hip extensors		
	Unit of allocation: individual		
Participants	Number of participants : 21 randomised; exercise group: n = 11; control group: n = 10		
	Group characteristics reported for all randomised participants.		
	Age : 8-18 years; exercise group: mean (SD) = 12.7 (2.8) years; control group: mean (SD) = 13.5 (3.4) years		
	Sex (male/female): 10/11; exercise group: 4/7; control group: 6/4		
	Ethnicity: not stated		
	GMFCS level : level I (n = 7), level II (n = 5), level III (n = 9); exercise group: level I (n = 2), level II (n = 2), level II (n = 7); control group: level I (n = 5), level II (n = 3), level III (n = 2)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP: diplegia		
	Inclusion criteria : aged 8-18 years, spastic diplegia, able to walk independently with or without a walk- ing aid (GMFCS level I-III), able to follow commands		
	Exclusion criteria : fixed flexion deformity at the knee, hip greater than 25 degrees or fixed equinus of more than 10 degrees, current participation in other management strategies such as serial casting, bot-		

Exercise interventions for cerebral palsy (Review)



Dodd 2003 (Continued)	ulinum toxin or orthopaedic surgery in the previous 12 months, participation in a strength-training pro- gramme within the previous 3 months
Interventions	Aim of the intervention: to increase muscle strength, physical activity and walking ability
	Type of exercise programme : resistance training. Participants also instructed to continue their nor- mal activities including school and sport and attend usual physiotherapy provided the programme didn't include progressive resistance training (usual physiotherapy is a 45-min consultation once or twice a month).
	Exercise mode : exercises to increase strength of ankle plantarflexors, knee extensors, and hip extensors i.e. bilateral heel raises, bilateral half squats, step-ups
	Comparator : participants in control group: instructed to continue their normal activities including school and sport. Also attended usual physiotherapy provided the programme didn't include progressive resistance training
	Setting: participant's home. Unsupervised individual session
	Intervention provider: physiotherapist
	Duration of programme: 6 weeks
	Exercise dose : 3 sets of 8-10 repetitions of each exercise (20-30 min sessions). 3 times per week. Load was adjusted by adding free weights to a backpack worn by participant to ensure participants could complete 8-12 repetitions to fatigue. At the end of 2nd and 4th week the physiotherapist visited participants and adjusted the load to ensure they can only complete between 8-12 repetitions of each exercise.
	Tailoring of intervention to individual : at the first session the training load was adjusted to ensure that each participant obtained optimal strength training
	Fidelity to prescribed intervention : adherence monitored with a self-report exercise diary. Mean (SD) sessions 16.8 (2.4) out of 18. Mean (SD) sets 147.7 (23.4) out of 162
	Monitoring of adverse events : no adverse events reported that led to missing training sessions. 1 par- ticipant reported pressure on the shoulders from the loaded backpack. 2 participants reported mild foot and ankle discomfort during heel raises. Exercise was modified and enabled participants to contin- ue without incident. Not stated how adverse events were monitored.
Outcomes	Assessment time points : baseline (week 0), postintervention (week 6), 12 weeks postintervention (week 18)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	1. Muscle strength of ankle plantarflexors, knee extensors, and hip extensors measured using hand-held dynamometry (recorded in kg).
	2. Gross motor function measured using dimensions D and E of the Gross Motor Function Measure. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. Scores presented as percentage of total possible score (higher score indicates better gross motor function). Separate scores were calculated for dimensions D and E and a total score was calculated by combining dimensions D and E.
	 Self-selected walking speed over 10 m (recorded in m/min). Timed stair test. Participants walk up and down 3 steps of standard size (17.5 cm) as quickly as possi-
	ble, using rails if required (recorded in s).
	 A physiotherapist experience in assessing movement disorders assessed all outcomes. Self-concept was measured with the Self-Perception Profile for Children. The scale assess children's perceptions of themselves across the domains of scholastic competence, physical appearance, and behavioural conduct, as well as a global perception of their worth or esteem as a person. A higher score indicates a more positive self-concept. This outcome was only assessed on children and adolescents

Exercise interventions for cerebral palsy (Review)



Dodd 2003 (Continued)

aged 8 to 16 years (n = 17; exercise group, n = 10; control group, n = 7) as it is only suitable for this age group.

Notes

Source of funding: not stated

Potential conflicts of interest: not stated; none perceived

Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "[p]articipants were allocated randomly to either the strength train- ing or control group using a concealed method. Twenty-two identical pieces of paper were placed in an opaque container, 11 with the words 'experimental group', 11 with the words 'control group' written on them. In another opaque container, the name of each participant was written on 21 separate pieces of paper. Allocation was achieved by drawing a piece of paper from each contain er. This process continued until all participants were allocated to a group".
Allocation concealment (selection bias)	Unclear risk	Quote : "[p]articipants were allocated randomly to either the strength train- ing or control group using a concealed method. Twenty-two identical pieces of paper were placed in an opaque container, 11 with the words 'experimental group', 11 with the words 'control group' written on them. In another opaque container, the name of each participant was written on 21 separate pieces of paper. Allocation was achieved by drawing a piece of paper from each contain er. This process continued until all participants were allocated to a group".
		Comment : unclear if an independent person did the allocation. Implementa- tion of the sequence could be open to bias
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote : "[a] physiotherapist who was blind to group allocation and experi- enced in assessing movement disorders took all outcome measures"
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : low rate of missing data. 1 participant (5%) withdrew from control group, reason given
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment : no other sources of bias identified

Engsberg 2006

Methods

Design: randomised controlled trial

Country of origin: USA

Intervention(s): strength training



Engsberg 2006 (Continued)	Unit of allocation: individual			
Participants	Number of participants : 15 participants randomised. Data reported on 12 participants. Dorsiflexor exercise group: n = 3; plantarflexor group: n = 4; dorsi- and plantarflexor group: n = 2; control group: n = 3			
	Age : mean (SD) = 9.9 (3.5) years; dorsiflexor group: mean (SD) = 12.5 (5.4) years; plantarflexor group: mean (SD) = 8.6 (2.3) years; dorsi- and plantarflexor group: mean = 7.6 (unable to calculate SD); control group: mean (SD) 10.7 (2.2) years			
	Sex (male/female): 5/10 (randomised) 3/9 (reported); dorsiflexor exercise group: 0/3; plantarflexor group: 2/2; dorsi- and plantarflexor group: 0/2; control group: 1/2			
	Ethnicity: not stated			
	GMFCS level : level I (n = 5), level II (n = 5) and level III (n = 2); dorsiflexor exercise group: level I (n = 1), level II (n = 1), and level III (n = 1); plantarflexor group: level I (n = 2), level II (n = 1) and level III (n = 1); dorsi- and plantarflexor group: level I (n = 1) and level II (n = 1); control group: level I (n = 1) and level II (n = 2)			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of CP: diplegia			
	Inclusion criteria : diagnosis of spastic diplegia, GMFCS level I, II or III, cognitive skills to actively partic- ipate (follow simple commands), ability to perform 6-8 repetitions of walking 9 m, passive dorsiflexion range of motion to at least -5 degrees with knee extended, ability to actively dorsi- and plantarflex the foot. Unclear whether hypertonicity of the plantar flexors as measured by the Ashworth scale was an in- clusion or exclusion criterion			
	Exclusion criteria : surgical intervention in last 12 months, casting procedures or botulinum toxin injection in last 6 months, selective dorsal rhizotomy, intrathecal baclofen, motor deficits secondary to neurological injury/illness beginning after the first month of life, or children with moderate-to-severe dystonia, athetosis, ataxia.			
Interventions	Aim of the intervention: to determine if an increase in ankle muscle strength results in an increase in function			
	Type of exercise programme: resistance training			
	Exercise mode : isokinetic dynamometer. Active assisted protocol in the passive mode was used. Par- ticipants were instructed to contribute as much force to the moving lever arm as possible.			
	Comparator : comparison of dorsiflexor strength training, plantarflexor strength training, dorsi- and plantarflexor strength training, and control group undergoing no strength training (no more detail provided)			
	Setting: physiotherapy clinic			
	Intervention provider: physiotherapist			
	Duration of programme: 12 weeks			
	Exercise dose : all strength training groups (1, 2 and 3) performed 3 sessions each week. In each session they performed 3 sets of 5 repetitions at 30°/s concentrically and eccentrically and 3 sets of 5 repetitions at 90°/s concentrically and eccentrically. Total number of repetitions was 30 for a muscle group per session. Load was equal to or greater than 80% of maximum load determined at start of session			
	Tailoring of intervention to individual : at the start of each session the participant had 3 attempts to meet or exceed previous maximum			
	Fidelity to prescribed intervention: not stated			



Engsberg 2006 (Continued)

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Engsberg 2006 (Continued)		e events : spasticity and end range dorsiflexion was monitored once a week. Pase and dorsiflexion range of motion did not reduce. No other adverse effects
Outcomes	Assessment time poin	its : baseline (week 0) and postintervention (week 12)
	Primary outcome: no	primary outcome measure stated
	Outcomes:	
	at 30°/s and 90°/s u their best performa 2. Spasticity of the pla	he plantar- and dorsiflexors. Concentric and eccentric muscle strength measured sing a dynamometer. 3-5 trials were permitted to allow the participants achieve nce. The maximum torque value normalised to participants' mass was reported. Intar- and dorsiflexors was measured using a dynamometer at 10, 30, 60, 90 and ons were conducted to identify a stable trial. Torque-angle data were processed /s].
	 Gait speed (in cm/s), stride length (in cm), cadence (in steps/min), ankle dorsiflexion at initial contact, ankle dorsiflexion maximum during stance, ankle dorsiflexion maximum during swing, and knee flexion minimum (all reported in degrees) were assessed as the participant walked barefoot at a self-selected pace along a 9-m walkway, using motion analysis. At least 6 trials of data were collected from each participant. Gross motor function was assessed using the GMFM-88. The Gross Motor Abilities Estimate was reported (higher score indicate better function). The score from dimension E was also reported. Possible range of scores 0 to 72 for dimension E. Scores presented as percentage of total possible score (higher score indicates better gross motor function). 	
	23 items, with each	sessed using the PedsQL 4.0 Core Scales child report and parent report. It includes item scored using a 5-point scale (higher score indicates better quality of life). ion was assessed using a goniometer (reported in degrees).
Notes	Source of funding : National Institute of Neurological Disorders and Stroke at the National Institut Health (ROI NS 046434)	
	Potential conflicts of interest: not stated; none perceived	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote (from abstract) : "Data were obtained from 12 children with spastic diplegia who were assigned randomly to a dorsiflexor group, a plantarflexor group, a dorsi- and plantarflexor group, or a control group."
		Quote : "A key factor for participation was the parents agreeing to permit ran- dom assignment to 1 of 4 groups: (1) dorsiflexion strength training (DF group), (2) plantarflexion strength training (PF group), (3) dorsi- and plantarflexion

(2) plantarflexion strength training (PF group), (3) dorsi- and plantarflexion strength training (DF&PF group), and (4) control group undergoing no strength group), (3) dorsi- and plantarflexion training program (control group)".

Comment: insufficient information regarding the method of sequence allocation was provided to make a judgement

Allocation concealment (selection bias)	Unclear risk	Comment : insufficient information regarding allocation concealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation

Exercise interventions for cerebral palsy (Review)

Cochrane

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Engsberg 2006 (Continued)		
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[i]t should also be noted that the investigators were not blinded to the training regimen of the subjects, since the study was too small to permit such a separation of tasks" Quote : "[t]he same study therapist collected dorsiflexion end-range outcome values for all subjects before and after the intervention"
		Comment : no indication who assessed outcomes other than "dorsiflexion end-range outcome values". Both self-reported and objective outcome mea- sures were used. As participants and their parents were not blinded to group allocation, at least 1 outcome is at high risk of bias.
Incomplete outcome data	High risk	Quote: "[h]owever, the final analysis included data from 12 of the 15 subjects"
(attrition bias) All outcomes		Quote : "[t]hree of the 15 subjects were eliminated from the investigation they did not demonstrate any gains in ankle strength (i.e. any increase above base- line values for a muscle being trained). The reasons for a lack of strength in- creases included: our inability to recognize the limited understanding of maxi- mum effort for one subject, use of an older Biodex that did not permit an 80% target line, and a therapist who did not follow the protocol. As previously dis- cussed the purpose of the investigation was to focus only on those subjects that increased in strength since we wanted to examine any relationship that might exist between strength gain and change in function".
		Comment : although the authors justify their exclusion of 3 participants, this represents a missing data rate of 20%. Further, reason for missing data is likely to be related to the true outcome.
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes.
Other bias	Unclear risk	Comment : demographic data provided only for those who completed the tri- al. Outcome data only provided for those who improved on measured out- comes

Unger 2006	
Methods	Design: randomised controlled trial
	Country of origin: South Africa
	Intervention(s): strength training; specifically targeted at disadvantaged students
	Unit of allocation: individual
Participants	Number of participants : 37 randomised; exercise group: n = 24; control group: n = 13. Data presented on 31 participants; exercise group: n = 21; control group: n = 10
	Age : exercise group: mean (range) = 15.86 (13.5 to 18.92) years; control group: mean (range) = 16.28 (14.0 to 18.33) years
	Sex (male/female): 19/12; exercise group: 13/8; control group: 6/4
	Ethnicity: not stated
	GMFCS level : no assistive devices (i.e. GMFCS level I or II; n = 29), and crutches or occasional use of wheelchair (i.e. GMFCS level III; n = 2); exercise group: no assistive devices (n = 19) and crutches or occasional use of wheelchair (n = 2); control group: no assistive devices (n = 10)

Exercise interventions for cerebral palsy (Review)

gia (n = 14), and triplegia (n = 1); exercise n = 1); control group:: hemiplegia (n = 8),			
without a walking aid, be in good health, able			
uch as a baclofen pump or selective dorsal onths, botulinum toxin injection in last 6 al level			
Type of exercise programme: resistance training			
argeting upper and lower limbs and trunk. 8 bells, ankle and wrist cuffs and bar with disc Ills were used for support or to provide an un-			
ructions on performance criteria by the re- vision of the exercise programmes			
ions progressed to 3 sets of 12 repetitions			
was set to allow at least 1 set of 6 to 10 repe- resistance was increased and repetitions re-			
tial programme was recorded on a participa- nd recording their own programme.			
Monitoring of adverse events: not reported			
tion (week 8)			
hase (reported in degrees), knee angle at heel i), stride length (reported in mm), and cadence ional gait analysis while participants were in- l without orthotics, down an 11 m walkway. A to ensure that at least 3 trials captured a com-			
reported questionnaire consisting of 6 state- scores 0-30. Higher score indicates better per-			
ing a self-reported questionnaire consisting of ge of scores 0-25. Higher score indicates better			

Exercise interventions for cerebral palsy (Review)



Unger 2006 (Continued)

Potential conflicts of interest: not stated. None perceived.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "Following pre-testing subjects were systematically randomised in- to either the experimental group or the control group with every third name drawn from the hat being allocated to the control group".
		Comment : description suggests sequence generation based on a non-random component.
Allocation concealment (selection bias)	Unclear risk	Quote : "Following pre-testing subjects were systematically randomised in- to either the experimental group or the control group with every third name drawn from the hat being allocated to the control group".
		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement. Did not indicate who conducted sequence genera- tion and how the allocation was concealed following sequence generation.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation.
Blinding of outcome as- sessment (detection bias)	High risk	Quote : "The research assistants for both outcome measures were blinded to group allocation for both pretesting and at eight-week testing".
All outcomes		Quote : "A short, self-administered questionnaires shown in the Appendix were used to assess perceptions of body image and functional competence. The themes relating to body image were identified from the physical appear- ance and attributes sub scale of the Piers Harris Children's Self-Concept Scale. Themes for section B were decided on in consultation with the school thera- pists and included activities required by the child for successful functioning in his or her environment."
		Quote: "Subjects selected the most applicable phrase."
		Comment : both self-reported and objective outcome measures were used. As participants were not blinded to group allocation at least 1 outcome is at high risk of bias.
Incomplete outcome data (attrition bias) All outcomes	High risk	Comment : 6 participants (16%) were withdrawn in total. 3 participants (13%) withdrew from the intervention group; 2 were "withdrawn due to absenteeism" and 1 was withdrawn "sport participation". 3 participants (23%) were withdrawn from the control group due to "sport participation", "incorrect diagnosis", and "participated in PRE".
		Indicates a high rate of missing data that's not evenly distributed across groups. Reasons for withdrawal aren't clear.
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes.
Other bias	Unclear risk	Comment : data analysis only included those who completed the trial not all those who enrolled.

Exercise interventions for cerebral palsy (Review)



iao 2007	
Methods	Design: randomised controlled trial
	Country of origin: Taiwan
	Intervention(s): resistance training (loaded sit-to-stand exercise)
	Unit of allocation: individual
Participants	Number of participants : 24 randomised, 20 analysed. Exercise group: n = 10; control group: n = 10;
	Baseline characteristics provided for n = 20 analysed participants
	Age : exercise group: mean (SD) = 85.6 (20.8) months; control group: mean (SD) = 91.3 (17.5) months.
	Sex (male/female): 12/8; exercise group: 7/3; control group: 5/5
	Ethnicity: not stated
	GMFCS level : level I (n = 10), level II (n = 10); exercise group: level I (n = 4) and level II (n = 6); control group: level I (n = 6) and level II (n = 4)
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: diplegia
	Inclusion criteria : aged 5-12 years, spastic diplegia, GMFCS level I or II, able to stand up from a chair ir dependently and maintain standing for more than 5 seconds, able to follow verbal instructions, without obvious limitation in passive range of motion of the lower limbs, able to attend physiotherapy at least once a week before and during the study, not received strength training in last 3 months, parenta commitment to allow participation without altering current therapy or activity
	Exclusion criteria : orthopaedic intervention, selective dorsal rhizotomy or botulinum toxin injection t the lower extremities within 6 months, or orthopaedic problems or medical conditions that prevented participants from participating in the exercises
Interventions	Aim of the intervention: to increase muscle strength
	Type of exercise programme: resistance training
	Exercise mode: loaded sit-to-stand exercise with weighted vest
	Comparator : all participants in both groups performed their regular physiotherapy programme which included passive range-of-motion exercises, positioning, balance training, functional training, and neurodevelopment training. At the start of the study both groups had 2 participants per group who had physiotherapy twice a week and 8 participants per group who received physiotherapy once a week. During a SARS epidemic participants in both groups decreased or stopped physiotherapy. In the control group, 1 participant received physiotherapy 2 days per week, 5 participants received physiotherapt 1 day per week, 1 participant received physiotherapy once every 2 weeks, and 3 participants didn't receive any physiotherapy. In the exercise group, 4 participants received physiotherapy 1 day per week, participants received physiotherapy once every 2 weeks, and 4 participants discontinued physiotherapy py
	Setting: home-based programme
	Intervention provider : trainer taught participants and their caregivers how to perform the exercise and modify the exercise during a visit to the home or study site, every other week. Exercises at home supervised by caregiver
	Duration of programme: 6 weeks
	Exercise dose : 3 sets of sit-to-stand exercises; 2 sets of 10 repetitions at 20% of 1 repetition maximum (RM), 1 set performing as many repetitions as possible at 50% 1 RM until fatigue. 3 times per week



iao 2007 (Continued)	Tailoring of intervention to individual : resistance applied was progressively increased to ensure the participant was performing exercises at 50% of 1 RM every 2 weeks			
	Fidelity to prescribed intervention : trainer insured the compliance of exercises during the training period via telephone interview. An exercise diary was provided to the caregiver to document the partic- ipant's exercise date, weight and number of repetitions in each exercise session. Participants in the ex- perimental group performed the loaded sit-to-stand exercise mean (SD) 18.0 (3.2) times (range 12 to 21 times) during the 6-week period. All participants performed exercises at least twice a week and 3 partic- ipants performed exercises more than 3 times. Participants' average maximum repetitions of 50% of 1 RM sit-to-stand varied from 20 to 100 each session.			
	Monitoring of adverse events : most participants in the exercise group reported pressure on the shoul- ders from the body vest during the loaded sit-to-stand. No pain or injury due to training was reported.			
Outcomes	Assessment time points: baseline (week 0) and postintervention (week 6)			
	Primary outcome: no primary outcome measure stated			
	Outcomes:			
	 Gross motor function was assessed using dimensions D and E of the GMFM-88. The scores on the 2 dimensions, presented as a percentage of total possible points, were averaged. Possible range 0-100. Higher score indicates better gross motor function 			
	2. Self-selected gait speed was calculated using the time it took participants to walk 10 m. The average of 3 trials was used in analysis. Reported in m/min.			
	3. Muscle strength was assessed using the 1 RM of the loaded sit-to-stand, defined as the maximal load the participant can carry while standing up one time. Reported in kg.			
	4. Knee extensor muscle strength was measured using a hand-held dynamometer. Each leg was as- sessed 3 times and the average torque of 3 separate trials of both legs was used in data analysis. Re- ported in kg.			
	5. Gait efficiency was measured using the physiologic cost index, calculated as the difference between the resting heart rate and walking heart rate, divided by walking speed. The higher the physiologic cost index the higher the energy consumption during walking.			
Notes	Source of funding: grant from the National Science Council, Taiwan			
	Potential conflicts of interest: authors report no conflict of interest; none perceived			
Risk of bias				

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote: "[r]andomised block design"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Comment : insufficient information regarding allocation concealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote : "[o]ne blinded tester (Y-CL) who is a physical therapist with pediatric assessment experience (including GMFM-88, gait speed) for 6 years, conducted the outcome measures and demographic data collection."

Exercise interventions for cerebral palsy (Review)

Liao 2007 (Continued)		
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Quote : "[o]f 24 children, 4 children (2 in experimental group, 2 in control group) withdrew before the study's completion because parents were concerned about SARS and did not want their children to come to the laboratory, which was located inside a hospital, for a follow-up test"
		Quote : "Some of the demographic data of these children who withdrew differed from the participant children. Compared with the participant children, the children who withdrew were statistically significantly older (109.8±6.4mo), heavier (26.1±4.3kg), and taller (127.0±10.9cm). However, their outcome measure data for the pre-assessment were similar to the participant children in this study."
		Comment : attrition accounted for but high level of missing data (17%). Miss- ing data evenly distributed across groups and no reported differences in base- line data between groups.
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment : baseline outcome and demographic data only includes those who completed the trial

Seniorou 2007

Methods	Design: randomised controlled trial		
	Country of origin: UK		
	Intervention(s): resistance training		
	Unit of allocation: individual		
Participants	Number of participants : 21 participants randomised. 20 participants completed study and included in analysis. Resistance exercise group: n = 11; no resistance exercise group: n = 9		
	Age : mean (SD) = 12.5 (SD 2.5) years (range 7.9 to 16.0 years)		
	Sex (male/female): 10/10		
	Ethnicity: not stated		
	GMFCS level : level I ($n = 3$); level II ($n = 13$) and level III ($n = 4$)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP: diplegia		
	Inclusion criteria: ambulant children and adolescents with spastic diplegic CP, surgery indicated		
	Exclusion criteria: botulinum toxin or orthopaedic surgery in previous year		
Interventions	Aim of the intervention: to increase muscle strength		
	Type of exercise programme: resistance training		
	Exercise mode : exercises for the hip flexors, hip extensors, hip abductors, knee flexors and knee extensors bilaterally		
	Comparator : identical programme performed with no weights (against gravity only).		

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Seniorou 2007 (Continued)	Sotting not constant
	Setting: not reported
	Intervention provider: clinician.
	Duration of programme: 6 weeks.
	Exercise dose : 3 times per week. 3 sets of 10 repetitions for each muscle group using free weights. The weight was determined using a 10RM (repetition maximum) for each muscle group.
	Tailoring of intervention to individual : re-assessment and incremental weight increase were dictated by the participant's progress.
	Fidelity to prescribed intervention: not reported
	Monitoring of adverse events: not reported
Outcomes	Assessment time points : baseline (week 0), postintervention (week 6), 20 weeks postintervention (week 26)
	Primary outcomes: primary outcome not stated.
	 Isometric muscle strength was assessed using a combination of fixed and hand-held dynamometry. 5 proximal muscle groups were tested: hip flexors, hip extensors and hip abductors, knee flexors and knee extensors at both 90 and 30 degrees of flexion. The highest force of 3 efforts for each participant was used in the data analysis.
	 3-dimensional gait analysis was performed using a Vicon 612 motion analysis system (Vicon, Oxford UK) with 15 retro-reflective surface markers on the pelvis and lower limbs. Each participant walked along a 10 metre walkway at their self-selected walking speed and a minimum of 4 walking trials was collected. The following parameters were reported: a. Normalised walk speed
	b. Range pelvic rotation (°)
	c. Max knee extensors stance (°)
	d. Knee flexion initial contact (°)
	e. Max thigh rotation (°) 3. Gross motor function was assessed using the gross motor function measure (GMFM-88). Data present-
	ed for dimension E (%) and total score (%), where a higher score indicates better function.
Notes	Source of funding: Oxfordshire Health Services Research Committee and the Wishbone Trust
	Potential conflicts of interest: authors report no conflict of interest.
Risk of bias	

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "Six months post-operatively, each child was allocated to one of two strengthening groups—AE or RS."
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : "Six months post-operatively, each child was allocated to one of two strengthening groups—AE or RS."
		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement.
Blinding of participants and personnel (perfor- mance bias)	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation.

Exercise interventions for cerebral palsy (Review)

Seniorou 2007 (Continued) All outcomes

Cochrane

Library

Unclear risk	Comment : did not provide information to indicate if the assessor was blinded to group allocation or not.
Low risk	Quote : "One child failed to complete the study for reasons unrelated to treat- ment. Therefore, 20 patients were included in the further analysis. 9 complet- ed programme AE and 11 programme RS. One child failed to complete the 12- month assessment". Comment : low rate of missing data (<10%).
Unclear risk	A protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes.
Low risk	Comment : no other sources of bias identified.
	Low risk Unclear risk

Methods	Design: quasi-randomised controlled trial Country of origin: Greece		
	Intervention(s): resistance and aerobic interval training		
	Unit of allocation: individual		
Participants	Number of participants : 13 allocated to groups; exercise group: n = 7; control group: n = 6. Data reported for all allocated participants.		
	Age : exercise group: mean (SD) = 15.9 (1.5) years; control group: mean (SD) = 15.7 (1.2) years		
	Sex (male/female): 4/9; exercise group: 2/5; control group: 2/4		
	Ethnicity: not stated		
	GMFCS level : no walking aids (i.e. GMFCS level I or II) (n = 4) and anterior walker (i.e. GMFCS level III) (n = 9); exercise group: no walking aids (n = 2) and anterior walker (n = 5); control group: no walking aids (n = 2) and anterior walker (n = 4)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP: diplegia		
	Inclusion criteria : diagnosis of spastic diplegia, able to walk with or without aids, aged 14-18 years, no been subjected to any orthopaedic surgical operation and had not received botulinum toxin injections in the preceding year, attended a similar physical therapy programme that did not include any form of systematic exercise		
	Exclusion criteria: not stated		
Interventions	Aim of the intervention: to increase strength and aerobic capacity		
	Type of exercise programme: mixed training (resistance training and aerobic interval training)		
	Exercise mode:		
	1. Resistance training protocol to strengthen elbow flexors and extensors, shoulder abductors, flexors extensors, internal and external rotators, forearm pronators and supinators, quadriceps, hamstrings		

Exercise interventions for cerebral palsy (Review)

Unnithan 2007 (Continued)

gastrocnemius, trunk and abdominal muscle groups. Exercises performed were biceps curls, triceps extensions, side arm lifts, knee and hip extensions, hip abductions, heel rises, push ups, and sit ups.

- 2. Game for general strengthening consisting of lateral trunk rotations using a medicine ball and a variety of drills that involved passing the ball and push ups and sit ups using the ball.
- 3. Aerobic interval training consisting of 3, 60 m outdoor uphill walking repetitions (gradient 5%, with or without an anterior walker).

Comparator: all participants maintained normal physical therapy, which consisted of individualised physical therapy on 2 days per week for 45 min, carried out by a physical therapist, based on Bobath treatment

Setting: not stated. Group, supervised sessions

Intervention provider: not stated

Duration of programme: 12 weeks

Exercise dose: 3 sessions per week. Each session lasting approximately 70 min (10 min warm-up, 20 min strength training protocol, 10 min game for general strengthening, 20-22 min aerobic interval training, 7 min breathing and passive stretching)

- 1. Resistance training protocol: initially 3 sets of 20 repetitions were performed for all upper-body exercises using hand weights of 2-3 kg. 4 sets of 10 repetitions of hip and knee extensions performed using ankle weights of between 0.5 and 1.0 kg. Heel rises performed using ankle weights of between 0.5 and 1.0 kg (sets and repetitions not stated). 3 sets of 15 repetitions of hip abductions. 3 sets of 8 repetitions of push-ups. 5 sets of 10 repetitions of sit-ups.
- 2. Game for general strengthening: 1 kg medicine ball used. Sets and repetitions not stated.
- 3. Aerobic interval training: the initial intensity was approximately 65% of the age-predicted maximal heart-rate. The average time taken to cover the 60 m distance was 100 s (range 90 s to 112 s). Active recovery (walking) between each repetition (work:rest ratio of 1:3). As the individuals improved their physical fitness the number of repetitions were increased, with the work:rest ratio maintained at 1:3. The intensity of the training sessions had increased to 75% of age-predicted maximal heart rate by the end of the 12th week.

Tailoring of intervention to individual: intensity was progressively increased every 3 weeks on an individualised basis by increasing the number of sets by 1 and repetitions by 5 while the weight remained stable.

Fidelity to prescribed intervention: not reported

Monitoring of adverse events: not reported

Outcomes

Assessment time points: baseline (week 0), postintervention (week 12)

Primary outcome: no primary outcome measure stated

Outcomes:

- 1. Gross motor function was assessed using dimensions D and E of the GMFM-88. It appears the score for D and E respectively were summed to provide an overall score. Possible range 0-100. Higher score indicates better performance.
- 2. Submaximal oxygen uptake (VO₂) reported in ml.kg⁻¹.min⁻¹, ventilation (V_E) reported in L.min⁻¹, and respiratory exchange ratio (RER) were measured during 4 min of arm cranking on an arm-crank ergometre at a submaximal workload (power output of 2.5 W at 50 rpm). V_E/VO₂ and VO₂ as a percentage of VO₂ peak were calculated.
- 3. Peak oxygen uptake (VO_{2peak}), V_E, HR, RER, heart-rate (in bpm) and blood lactate (in mM) were measured during a maximal exercise test on an arm-crank ergometer. The initial workload of 2.5 W was maintained for 4 min and the workload was increased at a rate of 2.5 W per min until volitional exhaustion (cadence of 50 rpm maintained throughout).

Notes Source of funding: not stated

Exercise interventions for cerebral palsy (Review)



Unnithan 2007 (Continued)

Potential conflicts of interest: not stated

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	High risk	Quote : "[t]he subjects were recruited on a staggered basis during a 6-month period and, therefore, were randomly allocated into either the training group (n=7) or control group (n=6) on the basis of timing of their recruitment into the study".
		Comment : description of method of sequence generation suggests it was based on a non-random component
Allocation concealment (selection bias)	High risk	Quote : "[t]he subjects were recruited on a staggered basis during a 6-month period and, therefore, were randomly allocated into either the training group (n=7) or control group (n=6) on the basis of timing of their recruitment into the study".
		Comment : description of method of allocation suggests that it was not con- cealed
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[t]his researcher was aware of which group each CP participant be- longed to"
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[a]Il subjects (training and control) participated in all the week 0 and week 12 tests, and none of the subject data were excluded from the subsequent data analyses".
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes.
Other bias	Low risk	Comment: no other sources of bias identified

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versenuren 2001	
Methods	Design: randomised controlled trial
	Country of origin: Netherlands
	Intervention(s): mixed training; resistance training, aerobic and anaerobic training
	Unit of allocation: individual
Participants	Number of participants : 68 randomised; exercise group: n = 34; control group: n = 34
	Group characteristics reported for all randomised participants
	Age : exercise group: mean (SD) = 11.6 (2.5) year; control group: mean (SD) = 12.7 (2.7) year
	Sex (male/female): 44/24; exercise group: 20/14; control group: 24/10

Exercise interventions for cerebral palsy (Review)

Verschuren 2007 (Continued)			
	Ethnicity: not stated		
	GMFCS level : level I (n = 47) and level II (n = 21); exercise group: level I (n = 24) and level II (n = 10); control group: level I (n = 23) and II (n = 11)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP : unilateral (n = 45) and bilateral (n = 23); exercise group: unilateral (n = 23) and bilateral (n = 11); control group: unilateral (n = 22) and bilateral (n = 12)		
	Inclusion criteria : aged 7-20 years, diagnosed with spastic CP, GMFCS level I or II, able to follow simple verbal commands, receiving rehabilitation services at the time of the study.		
	Exclusion criteria : orthopaedic surgery or neurosurgery and/or botulinum toxin injection in previous 6 months, cardiac or respiratory conditions that could be negatively affected by exercise		
Interventions	Aim of the intervention : to increase aerobic fitness (for first 4 months) and anaerobic fitness (for sec- ond 4 months)		
	Type of exercise programme: mixed; aerobic, anaerobic, and muscle strengthening		
	Exercise mode : 8 standardised aerobic exercises lasting 3-6 min. 8 standardised anaerobic exercises lasting 20-30 seconds. The task-specific exercises, such as running and changing direction of the body abruptly, step-ups, and negotiating stairs, were repeated throughout the programme and aimed to improve daily functioning. Participants also continued with usual care.		
	Comparator : all participants received usual care, which ranged from no treatment to various therapeu- tic approaches. There was no difference in usual care between groups as tracked from medical records		
	Setting: school; supervised groups of 4-6 participants		
	Intervention provider : led by 2 local paediatric physiotherapists who received standardised fitness programme training prior to the start of the programme		
	Duration of programme : 8 months; aimed to increase aerobic fitness in the first 4 months and in- crease anaerobic fitness in second 4 months,		
	Exercise dose : 2 days per week. Each session lasted 45 min (5 min warm up, 25 to 35 min functional aerobic, anaerobic, and muscle strengthening exercises performed in a circuit, 5 min cool down).		
	Tailoring of intervention to individual: not stated		
	Fidelity to prescribed intervention : median attendance was 56 out of a possible 60 sessions (93%). All participants attended at least 85% of the training sessions. Unclear how fidelity was monitored.		
	Monitoring of adverse events : during a training session 1 participant fell and fractured her radius. She missed 4 training sessions because she was wearing a cast.		
Outcomes	Assessment time points : baseline (week 0), postintervention (month 8), 4 months postintervention (month 12)		
	Primary outcomes: primary outcomes were aerobic and anaerobic capacity		
	1. Aerobic capacity was measured in terms of the level achieved on a 10 m shuttle run test. Reported in minutes.		
	2. Anaerobic capacity was measured using mean power (reported in watts) derived from the Muscle Power Sprint Test.		
	Secondary outcomes : agility, muscle strength, body mass index, self-perception, gross motor func- tion, participation, and health-related quality of life		
	 Agility was measured using the 10 X 5 m sprint test (unit of measurement unclear) Muscle strength of the lower limbs was assessed using the 30 second repetition maximum (unit of measurement unclear). 		

Exercise interventions for cerebral palsy (Review)



Verschuren 2007 (Continued)

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3. Body mass index was calculated as weight in kilograms divided by height in metres squared

- 4. Self-perception was measured using the Self-Perception Profile for Children. The domains of athletic competence, physical appearance, and global perception of their worth or esteem as a person were assessed. Score for each domain presented as a percentage.
- 5. Gross motor function was assessed using Dimensions D and E of the GMFM. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. It is not clear if scores are presented as the sum of the scores for each item or as a percentage of total possible score. Higher score indicates better gross motor function.
- 6. Participation was measured with the Children's Asssessment of Participation and Enjoyment (CAPE). The intensity scores of all types of activity (i.e. recreational, active physical, social, skill-based, and self-improvement activities), was assessed to reflect the average amount of time that a participant spent participating in different activities.
- 7. The TNO-AZL Questionnaire for Children's Health-Related Quality of Life Parent Form (TACQOL-PF) was used to assess health-related quality of life. Parents provided a single score for each pair of items (functional item and corresponding emotional item) in 5 scales (pain and symptoms, basic motor functioning, autonomy, cognitive functioning, social functioning). The sum scores may range from 0 to 32 for these scales. Parents provide a single score for each single item in a global positive emotional functioning scale. The scores may range from 0 to 16 for these scales. For all scales, a low score indicates a lower quality of life.

Source of funding: Dr WM Phelps foundation

Potential conflicts of interest: authors declare no conflicts of interest; none perceived

Notes

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[p]articipants were randomly assigned to 2 groups using a 4-block randomization protocol. Each block represented all participants from 1 school. The groups within each block consisted of children at level I or II on the GMFCS. From each block and group every participant was randomly allocated to the training group or the control group".
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Low risk	Quote : "[a]n independent off-site researcher not involved in the assessments used a concealed method for allocation."
		Comment: probably done
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[t]o reduce bias 8 assessors who were not the treating therapist and who were blinded for the treatment modality undertook the testing without review of previous scores"
		Comment : participation and quality of life were assessed using self-report outcome measures. As participants and their parents were not blinded to group allocation, at least 1 outcome is at high risk of bias.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : all data analyses were carried out according to a pre-established analysis plan and were performed according to the intention-to-treat principle". 4% attrition during baseline measures, evenly distributed, and accounted for.

Exercise interventions for cerebral palsy (Review)



Lee 2008

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Verschuren 2007 (Continued)

Selective reporting (re- porting bias)	Low risk	Comment : a summary of the study protocol is available and pre-specified out- comes reported
Other bias	Low risk	Comment : no other sources of bias identified

Methods Design: randomised controlled trial Country of origin: Korea Intervention(s): resistance training Unit of allocation: individual Participants Number of participants: 17 randomised; exercise group: n = 9; control group: n = 8 Age: exercise group: mean (SD) = 6.3 (2.1) years; control group: mean (SD) = 6.3 (2.9) years Sex (male/female): 10/7; exercise group: 4/5; control group: 6/2 Ethnicity: not stated GMFCS level: II or III, distribution per group not stated Type of motor abnormality: spastic CP **Anatomical distribution of CP**: diplegia (n = 9) and hemiplegia (n = 8); exercise group: diplegia (n = 4)and hemiplegia (n = 5); control group: diplegia (n = 5) and hemiplegia (n = 3)Inclusion criteria: aged 4-12 years, spastic diplegia or hemiplegia CP, GMFCS level II or III Exclusion criteria: unable to follow commands, fixed contracture at hip or knee joint of more than 25°, medical or orthopedic diseases that prevented them from exercising, received orthopedic surgery of the lower limb or injection of antispastic drug (e.g. botulinum toxin injection) Aim of the intervention: to increase muscle strength in the lower limbs Interventions Type of exercise programme: resistance training Exercise mode: squat-to-stand, lateral step-up, stair walk up and down, isotonic exercise of lower limb muscles, isokinetic exercises using a bike. Comparator: conventional physiotherapy including neurodevelopmental therapy, range of movement exercises and gait training. Dose not reported Setting: unclear Intervention provider: physiotherapist

Duration of programme: 5 weeks

Exercise dose: a 60 min session was delivered 3 times per week. For isotonic exercise 1 of 3 weights (0.25 kg, 0.45 kg or 0.9 kg) was used to provide resistance to voluntary muscle contraction during the exercise. Participants completed 2 sets of 10 repetitions in each muscle group.

Tailoring of intervention to individual: selected weight depended on the ability of the participant to complete 2 sets of 10 repetitions.

Fidelity to prescribed intervention: not reported



Lee 2008 (Continued) Monitoring of adverse events: not reported

Outcomes	Assessment time points : baseline (week 0), postintervention (week 5), 6 weeks postintervention (week 11)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	 Muscle strength was assessed using manual muscle testing (possible score of 0 to 5, higher score in- dicates better muscle strength) and number of lateral step-ups and squats to stand performed during 30 seconds.
	 Gross motor function was assessed using the GMFM-88. Scores for dimensions D and E and total scores were presented as a percentage of total possible score (higher score indicates better gross motor func- tion).
	3. Muscle tone was assessed using the Modified Ashworth Scale (MAS). Possible score of 0-5, higher score indicates greater spasticity.
	 The following parameters at self-selected speed using assistive devices if necessary were assessed using motion analysis: a. Gait speed reported in cm/s
	b. Stride length reported in cm
	c. Cadence metric not reported
	d. Time in single support reported as percentage of gait cycle
	e. Time in double support reported as percentage of gait cycle
	f. Maximal and minimal angle of hip flexion
	g. Maximal and minimal angle of knee flexion
	h. Maximal and minimal angle of ankle plantarflexion
	i. Maximal and minimal angle of pelvic anterior tilt
	j. Maximal and minimal moment of hip flexion
	k. Maximal and minimal moment of knee flexion
	l. Maximal and minimal moment of ankle plantarflexion
	m. Maximal and minimal hip power
	n. Maximal and minimal knee power
	o. Maximal and minimal ankle power
Notes	Source of funding: not stated

Potential conflicts of interest: not reported; none perceived

Risk of bias

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Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[p]articipants were allocated randomly to either the experimental group or control group using concealed methods"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : "[p]articipants were allocated randomly to either the experimental group or control group using concealed methods"
		Comment : insufficient information regarding the method of allocation con- cealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias)	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation

Exercise interventions for cerebral palsy (Review)



Lee 2008 (Continued) All outcomes

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Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Comment : did not provide information to indicate if the assessor was blinded to group allocation or not
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Comment : does not state if anyone withdrew or number of participants in- cluded in the analysis
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment : unclear whether all demographic or outcome data for all participants are reported

Methods	Design: randomised controlled trial		
	Country of origin: Greece		
	Intervention(s): aquatic programme		
	Unit of allocation: individual		
Participants	Number of participants : 12 randomised; exercise group: n = 6; control group: n = 6		
	Group characteristics were reported for all randomised participants		
	Age : 13-20 years; exercise group: mean (SD) = 16 (2.89) years; control group: mean (SD) = 16.66 (2.65) years.		
	Sex (male/female): 7/5; exercise group: 4/2; control group: 3/3		
	Ethnicity: not stated		
	GMFCS level : according to inclusion criteria levels I, II and III (i.e. able to walk with or without aids). Level of lower-limb gross motor function of included participants is not stated		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of motor abnormality : according to inclusion criteria tetraplegia and diple gia; anatomical distribution of included participants is not stated		
	Inclusion criteria : diagnosis of spastic tetraplegia or diplegia, able to walk with or without aids, able follow simple commands		
	Exclusion criteria: undergone surgery in last 12 months or receive medication for spasticity		
nterventions	Aim of the intervention : to improve gross motor function and range of movement and to reduce spar ticity		
	Type of exercise programme: aerobic		
	Exercise mode : swimming. Participants worked on the basic backstroke and crawl swimming styles		
	Comparator : participants in the control group continued their normal activities and physiotherapy sessions provided from the school staff. Dose not reported.		

Exercise interventions for cerebral palsy (Review)



Chrysagis 2009 (Continued)	Setting : 25 m swimmir	ng pool. Supervised sessions. Unclear if individual or group session.
	Intervention provider	r: 2 physical educators trained in swimming skills for children with CP from the onsible for the swimming programme
	Duration of programm	ne : 10 weeks
		pants in the exercise group received 45 min of exercise (10 min warm up of of training), 2 days per week
		ion to individual : training was individualised according to each participant's s were used if necessary
	Fidelity to prescribed	intervention: not reported
	Monitoring of adverse	e events: not reported
Outcomes	Assessment time poin	nts: baseline (week 0) and 4 weeks postintervention (week 14)
	Primary outcome: no	primary outcome measure stated
	Outcomes:	
	velopmental and Ph (GMFM). It is not cle known. However, ra indicates better gro 2. Spasticity of the rig Ashworth Scale (scc	In was measured by 2 members of the Laboratory of Adapted Physical Activity/De- hysical Disabilities using dimensions D and E of the Gross Motor Function Measure ear if the GMFM-66 or GMFM-88 was used and therefore the range of scores is un- aw scores were converted to a percentage of total possible points. A higher score ss motor function. ht and left hip adductors and knee flexors was evaluated according the modified ored as 0, 1, 1 ⁺ , 2, 3, 4); higher score indicates greater spasticity the right and left shoulder, hip and knee was measured using a goniometer
Notes	Source of funding: no	ne stated
	Potential conflicts of	interest: none stated; no conflicts of interest perceived
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[p]articipants were randomly allocated (sealed envelopes) to experi- mental and the control group"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : "[p]articipants were randomly allocated (sealed envelopes) to experi- mental and the control group".
		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Quote : "[t]wo members of the Laboratory of Adapted Physical Activity/Devel- opmental and Physical Disabilities carried out the above measurements in the school gym at the beginning and at the end of the intervention program"

Exercise interventions for cerebral palsy (Review)



Chrysagis 2009 (Continued)		Comment : insufficient information regarding assessment procedures was pro- vided to make a judgement
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[t]here was no drop out during the study" Comment : no missing data
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment: no other sources of bias identified

Methods	Design: randomised controlled trial			
	Country of origin: Norway			
	Intervention(s): resistance training using a seated leg press			
	Unit of allocation: individual			
Participants	Number of participants : 12 randomised; exercise group: n = 6; control group: n = 6			
	Baseline characteristics reported on all randomised participants			
	Age : exercise group: mean 41 years (range 32 to 69 years); control group: mean 45 years (range 27 to 69 years) years)			
	Sex (male/female): 4/8; exercise group: 2/4; control group: 2/4			
	Ethnicity: not stated			
	GMFCS level : level II (n = 7) and level III (n = 5); exercise group: level II (n = 4) and level III (n =2); contro group: level II (n = 3) and level III (n = 3)			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of CP: diplegia			
	Inclusion criteria : age 18 years and older, spastic diplegia, GMFCS level II or III, experiencing difficulti walking but able to walk for 6 min with or without minimal support from another person, motivated to participate in progressive resistance training, able to understand and perform progressive resistance exercise training under supervision			
	Exclusion criteria : participation in strength training for the lower limbs in the past 12 months, severe cognitive disorders			
Interventions	Aim of the intervention: to improve muscle strength			
	Type of exercise programme: resistance training			
	Exercise mode: seated leg press			
	Comparator : participants in the control group continued their pre-study individual ongoing sympton relief treatment or training regime. This consisted of mainly passive physiotherapy treatment such as stretching, mobilisation of joints and massage			
	Setting: physiotherapy clinic; supervised individual session			

Exercise interventions for cerebral palsy (Review)

Maeland 2009 (Continued)	
	Intervention provider: physiotherapist
	Duration of programme: 8 weeks
	Exercise dose : 4 sets of 12-15 repetitions at 60%-75% of 1 repetition maximum (1 RM), 3 days per week for first 2 weeks. 4 sets of 4-6 repetitions to fatigue (i.e. 4 RM to 6 RM, 85% of 1 RM), 3 days per week for 6 weeks
	Tailoring of intervention to individual : when the participants managed to complete 15 repetitions or 6 repetitions (depending on the week) in all 4 sets resistance was increased by 5 kg to 10 kg
	Fidelity to prescribed intervention: 90% of all sessions were completed
	Monitoring of adverse events : no adverse events or effects were reported or registered by the participants or physiotherapists
Outcomes	Assessment time points: baseline (week 0), 2 weeks postintervention (week 10)
	Primary outcome:
	1. Walking capacity was measured on the 6-min walk test. Distance completed was reported in m.
	Seconday outcomes:
	 Self-selected comfortable gait speed and self-selected maximum gait speed were measured during the 10-metre walk test. Reported in m/s.
	2. Muscle strength was measured with the timed stands test. The time required to complete 10 full stands from sitting was recorded in s.
	3. Gross motor function was assess using the stair climb test. The time required to walk up and down 9 steps, as quickly as possible was recorded in s.
	4. Isokinetic muscle power was measured for concentric action of the knee extensors at 60°/s. Reported in Nm/s.
	5. Perceived exertion was measured using the Borg 20 grades rating scale of perceived exertion, a nu- merical rating scale from 6 (no perceived exertion) to 20 (maximal exertion).
Notes	Source of funding: Helse_Ost RHF (Eastern Norway Regional Health Authority)
	Potential conflicts of interest: authors report no conflict of interest; none perceived

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "[t]he randomisation was done by computerised generating of 12 Bernoulli trials (0s and 1s) using SPSS"
Allocation concealment (selection bias)	Unclear risk	Quote : "[s]ealed numbered envelopes with information about the treatment were prepared by the statistician"
		Comment : did not state if envelopes were opaque or if the statistician was in- dependent to the research team
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote : "[a]ll functional outcome measures were carried out by two experi- enced senior physiotherapists who were blinded as to which group the partici- pants belonged"

Exercise interventions for cerebral palsy (Review)

Maeland 2009 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment: no missing data
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment : no other sources of bias identified

Fowler 2010

Methods	Design: randomised controlled trial
	Country of origin: USA
	Intervention(s): stationary cycling programme
	Unit of allocation: individual
Participants	Number of participants : 64 randomised; exercise group: n = 33; control group: n = 31; Baseline mea- sures n = 62; exercise group: n = 31; control group: n = 31
	Group characteristics reported for n = 62
	Age : exercise group: mean (95% CI) = 11.1 (9.9 to 12.3) years; control group: mean (95% CI) = 11.6 (10.6 to 12.6) years
	Sex (male/female): 29/33; exercise group: 18/13; control group: 11/20.
	Ethnicity : African American (n = 8), white (n = 33), Asian (n = 6), other (not specified; n = 15); exercise group: African American (n = 5), white (n = 18), Asian (n = 1), other (n = 7); control group: African American (n = 3), white (n = 15), Asian (n = 5), other (n = 8)
	GMFCS level : levels I (n = 19), level II (n = 14), level III (29); exercise group: level I (n = 11), level II (n = 8), level III (n = 12). control group: level I (n = 8), level II (n = 6), level III (n = 17)
	Type of motor abnormality: spastic CP
	Anatomical distribution of motor abnormality: diplegia
	Inclusion criteria : diagnosis of spastic diplegia, aged 7-18 years, ability to comply with simple verbal directions, GMFCS levels I-III, selective motor control rating of good or fair
	Exclusion criteria : neurological surgery, orthopaedic surgery or implantation of a baclofen pump with in the 12 months preceding enrolment; botulinum toxin injections within the preceding 3 months; se- rial casting or new orthotic devices within the preceding 3 months; initiation of oral medications that affect the neuromuscular system (e.g. baclofen) within the preceding 3 months; initiation of physical therapy, exercises, sports activity, or change in assistive devices for walking within the preceding 3 months; inability or unwillingness to maintain age-appropriate behaviour; serious medical conditions such as diabetes, cardiovascular disease, or uncontrolled seizures; current participation in a fitness programme that included a minimum of once-weekly cardiorespiratory endurance exercise; significan hip, knee, or ankle joint contractures preventing passive movement of the lower limbs through pedal- ing cycle; poor bilateral voluntary selective motor control
Interventions	Aim of the intervention: to improve muscle strength and walking and running endurance
	Type of exercise programme: mixed (resistance and aerobic)

Type of exercise programme: mixed (resistance and aerobic)



Fowler 2010 (Continued)

Exercise mode: stationary cycling. In order to perform lower limb strengthening the bike seat was unlocked and allowed to slide backward. Up to 10 tension cords each providing 4.5 kg (10 lb) of force acted to pull the seat forward. Participant extended lower limbs to prevent the seat being pulled forward

Comparator: no cycling. All participants completed physical activity diaries to assess levels of physical activity throughout duration of the study

Setting: physiotherapy clinic

Intervention provider: physical therapist

Duration of programme: 12 weeks

Exercise dose: 30, 60-min sessions, 3 times a week. During cardiorespiratory phase participants exercised at 70%-80% heart rate max (HRmax) (calculated using the Karvonen formula); aimed for 15-30 min at this intensity. Mean (SD) typical exercise heart rate across all sessions was 147.2 (14.4) bpm (range 117-176 bpm) representing a mean (SD) percentage of HRmax of 52.2 (12.2) (range = 8%-77%). For lower limb strengthening the starting resistance was the attachment of 1 tensioning cord. Resistance was progressed to next cord when 10 revolutions were performed while keeping seat in desired zone. If a participant could not cycle with the seat unlocked or if the maximum resistance was reached a 'constant power' resistance mode was used. Number of revolutions prescribed not stated. The mean (SD) maximum load was 12.2 (12.1) kg (26.9 (26.6) lb) over the first 3 days of the intervention and 29.7 (15.5) kg (65.5 (34.2) lb) by the end of the intervention, representing a mean (SD) gain of 17.5 (11.7) kg (38.6 (25.7) lb) (range = 0-40.1 kg (0-90 lb)).

Tailoring of intervention to individual: if participant could not cycle independently manual assistance was provided until independence was achieved. Resistance and pedaling rate were adjusted based on HR and Children's Effort Rating Table (scale 1-10)

Fidelity to prescribed intervention: attendance at cycling sessions was 89.6%.

Monitoring of adverse events: did not state how adverse events were assessed. 28 mild events (for 18 participants) potentially related to the study: 6 observed falls, 17 complaints of soreness, muscle cramping or mild pain, 4 reports of fatigue, and 1 skin rash related to HR monitor. 30 events unrelated to study procedures: illness, tooth loss, headache, stomach ache, tonsillectomy, and skin irritation from orthotic use.

Outcomes

Assessment time points: baseline (week 0) and postintervention (week 12)

Primary outcomes

- 1. Walking and running endurance measured using the 600 yard walk/run (548.6 m) test where participants were directed to walk or run as fast as they could. Speed per distance completed was outcome recorded (m/min).
- 2. Self-selected walking speed was assessed using the 30-second walk test and reported in m/min.
- 3. Gross motor function was assessed using dimensions D and E of the GMFM-66. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. Scores presented as percentage of total possible score (higher score indicates better gross motor function). Separate scores were calculated for dimensions D and E and a total score was calculated by combining dimensions D and E.
- 4. Muscle strength was assessed by measuring left and right peak knee extensor and flexor moments at 0, 30, 60 and 120°/s using an isokinetic dynamometer.

Secondary outcomes

 Health-related quality of life was assessed using the Pediatric Quality of Life Inventory 4.0 Generic Core Scales SF15 (PedsQL) and the Pediatric Outcomes Data Collection Instrument (PODCI). Participants reported quality of life using the PedsQL, which assesses 4 dimensions of function. Scores in 3 of the 4 dimensions (emotional, social, and school) were combined to provide a psychosocial health summary. A higher score indicates a more positive health-related quality of life. Parents reported quality of life using a parent proxy version of the PODCI. The PODCI contains 4 sections: global functioning and symptoms, happiness, treatment expectations, and satisfaction with symptoms. A higher score represents a more positive health-related quality of life.



Fowler 2010 (Continued)	Assessments were conducted by an evaluator who was blinded to participant group assignment and had to pass a rigorous standardization procedure for each outcome measurement protocol by demon- strating at least 90% competency.
Notes	Source of funding: Foundation for Physical Therapy
	Potential conflicts of interest : not stated. Authors acknowledged other corporate funders, donors and discounts; none perceived

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "[t]he participants were randomly assigned to a control (no interven- tion) or an intervention (cycling) group. Randomization was blocked by age (7-11y, 12-18y) and lower extremity selective voluntary motor control ability (good, fair) to minimize the effects of physical impairment and maturation.
		Quote : "[g]roup assignment was determined using a computerized random number generator."
Allocation concealment (selection bias)	Unclear risk	Quote (from the protocol) : "[a]n enrolment form containing the subject's age and selective motor control ability will be submitted to the PTClinResNet Da- ta Management Centre for subject randomisation. Families will be notified of their child's assignment to the control or intervention group following baseline evaluation".
		Comment : although it appears randomisation was conducted by a third party independent of the study, insufficient information is provided to make a clear judgement regarding allocation concealment.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation.
Blinding of outcome as-	High risk	Quote : "[e]valuators were blinded to group assignment"
sessment (detection bias) All outcomes		Quote : "[b]oth questionnaires were administered following a standardized protocol outlined in the study's manual of procedures".
		Quote : "[t]he youngest children in the study (7y) used the PedsQL young child version; older children (8-12y) completed the child version and adolescents (13-18y) used the teen version. The PODCI parent proxy versions for parents or guardians of children (2-10y) or adolescents (11-18y) were used."
		Quote : "[e]ach question and all possible answers were read to the participant and the evaluator recorded the selected answer".
		Comment : both self-reported and objective outcome measures were used. As participants and their parents were not blinded to group allocation, at least 1 outcome is at high risk of bias.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : attrition and withdrawal accounted for. Low rate of missing data (~ 10%) and evenly distributed across groups
Selective reporting (re- porting bias)	Low risk	Comment: all prespecified outcomes reported

Exercise interventions for cerebral palsy (Review)



Fowler 2010 (Continued)

Other bias

Unclear risk

Methods	Design: cross-over randomised controlled trial
	Country of origin: Australia
	Intervention(s): resistance training
	Unit of allocation: individual
Participants	Number of participants : 14 randomised; exercise group: n = 7; control group: n = 7
	Age: mean (SD) 11 (2) years (range 9 to 15 years)
	Sex (male/female): 6/8
	Ethnicity: not stated
	GMFCS level: not stated
	MACS level : level I ($n = 4$), level II ($n = 8$) and level III ($n = 2$)
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: hemiplegia ($n = 13$) and triplegia ($n = 1$)
	Inclusion criteria : elbow flexor spasticity, the ability to follow 2-step instructions, no previous up- per-limb surgery, and no upper-limb strength training or pharmacological treatment for spasticity (bot ulinum toxin A) in the past 12 months
	Exclusion criteria: as stated in inclusion criteria
Interventions	Aim of the intervention : predicted that eccentric strength training would improve peak torque and work, torque-angle relationship, and electromyographic activation
	Type of exercise programme: resistance training
	Exercise mode : upper limb eccentric exercises using training rig that provided loaded assistance to draw the elbow into extension in a gravity-eliminated position. The arm was returned to 110° elbow flexion at the end of each repetition by a partner so the participant did not perform a concentric muscl action
	Comparator: normal activity
	Setting: home-based programme
	Intervention provider: training partner assisted (not stated who was training partner)
	Duration of programme: 6 weeks
	Exercise dose : 3 sessions per week, 3 sets of 10 repetitions, starting at 50% maximum eccentric torque and progressing to 70% maximum eccentric torque at by the last week in increments of 5%
	Tailoring of intervention to individual : each participant was provided with an individually adapted eccentric training rig that permitted eccentric extension of the elbow flexors only

Exercise interventions for cerebral palsy (Review)



Reid 2010 (Continued)

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(Continued)	Monitoring of adverse quence of eccentric tra	e events: not reported: states no child reported muscle soreness as a conse- ining		
Outcomes	Assessment time points: baseline (week 0) and postintervention (week 6) Primary outcome: not stated			
	Outcomes:			
	 Concentric muscle strength of the elbow flexors at 30°/s, 60°/s and 90°/s was assessed during 3 isoki netic trials using a dynamometer. The best performance was used in analysis. Peak torque and wor normalised to body mass were reported in Nm/kg. Eccentric muscle strength of the elbow flexors at 30°/s was assessed during 3 isokinetic trials using a dynamometer. The best performance was used in analysis. Peak torque and work normalised to body mass were reported in Nm/kg. Activation of the biceps brachii and brachioradialis during the isokinetic assessments was collected using surface electromyography. 			
Notes	Source of funding: not	t stated		
	Potential conflicts of	interest: not stated; none perceived		
Risk of bias				
Bias	Authors' judgement	Support for judgement		
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[p]articipants with CP were randomised into two groups and completed the same eccentric training programme: group I (n=7) trained in the first 6 weeks of the study, while group II (n=7) acted as a control group, maintaining their normal activity for 6 weeks, and then completed the 6-week eccentric training programme".		
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement		
Allocation concealment (selection bias)	Unclear risk	Quote : "[p]articipants with CP were randomised into two groups and completed the same eccentric training programme: group I (n=7) trained in the first 6 weeks of the study, while group II (n=7) acted as a control group, maintaining their normal activity for 6 weeks, and then completed the 6-week eccentric training programme".		
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement		
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation		
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Comment : did not provide information to indicate if the assessor was blinded to group allocation or not		
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment: no missing data		
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes		

to indicate that published report includes all expected outcomes.

Exercise interventions for cerebral palsy (Review)



Cochrane Database of Systematic Reviews

Reid 2010 (Continued)

Other bias

Low risk

Methods	Design: randomised controlled trial		
	Country of origin: Netherlands		
	Intervention(s): resistance training		
	Unit of allocation: individual		
Participants	Number of participants : 51 participants randomised. Exercise group: n = 26; control group: n = 25 (49 participants included in analysis, some demographic data presented for 49 participants only)		
	Age : mean (SD) = 10 years 5 months (1 year 10 months) (range 6 years 0 months to 13 years 10 months) exercise group: mean (SD) = 10 years 4 months (1 year 10 months); control group: mean (SD) = 10 years 3 months (2 years 3 months)		
	Sex (male/female): 29/22; exercise group: 16/8; control group: 13/12		
	Ethnicity: not stated		
	GMFCS level : level I (n = 25); level II (n = 17) and level III (n = 7); exercise group: level I (n = 13), level II (n = 8) and level III (n = 3); control group: level I (n = 12), level II (n = 9) and level III (n = 4)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP : unilateral (n = 17) and bilateral (n = 32); exercise group: unilateral (n = 7) and bilateral (n = 17); control group: unilateral (n = 10) and bilateral (n = 15)		
	Inclusion criteria : aged 6-13 years, able to accept and follow verbal instructions, able to walk indepen dently indoors with or without walking aids (GMFCS level I, II or III), able to participate in a group training programme		
	Exclusion criteria : unstable seizures, any treatment of spasticity or surgical procedures in last 3 months (for botox) or 6 months (for surgery), any change in medication expected during the study period and suffering from any other diseases that interfered with physical activity		
Interventions	Aim of the intervention: to increase muscle strength		
	Type of exercise programme: resistance training		
	Exercise mode : 2 loaded exercises: leg press and sit-to-stand. Unloaded or low loaded exercises: later- al step-up, forward step-up, half knee-rise. Each participant completed 4 different exercises on a circui of 5 stations. The intervention replaced usual care.		
	Comparator : conventional physiotherapy programme, 1-3 sessions per week, dose and content not recorded		
	Setting: special school		
	Intervention provider: physiotherapist		
	Duration of programme: 12 weeks (33 sessions in total as 3 sessions cancelled)		
	Exercise dose : each session was 45-60 min. 3 sessions per week. Each session consisted of 4 different exercises performed as a station in a circuit. The exercises were a leg press exercise on a child-adapted leg-press, sit-to-stand, lateral step-up and half knee-rise. 3 sets of 8 repetitions of each exercise was performed. The stations were a combination of a high load exercise, a low load exercise and an unloaded exercise. The high load exercise (bilateral leg-press) was performed at 100% 8 RM, the loaded		

Exercise interventions for cerebral palsy (Review)

Scholtes 2010 (Continued)	exercise (bilateral sit-to-stand) was performed at 75% 8 RM, the loaded game (unilateral half-knee rise, lateral step-up or forward step-up) was performed at 25% 8 RM, and the unloaded game (unilateral half-knee rise, lateral step-up or forward step-up) was performed with no resistance. The first 6 weeks were intended to slowly build up training to these loads. The final 6 weeks were performed at these loads
	Tailoring of intervention to individual : training loads were adjusted to new individual levels of strength, if necessary, as determined by the 8 RM test
	Fidelity to prescribed intervention : mean compliance was 92.3% (71% to 100%). A mean of 32 out of 36 sessions (range 30 to 33) were attended. Reasons for absence were illness (41.4%), medical appointment (8.6%), vacation (6.9%), other/unknown (43.1%).
	Monitoring of adverse events : each week 1-6 participants reported mild-to-moderate muscle sore- ness, recorded on a Likert scale (no, mild, moderate, severe, extremely severe)
Outcomes	Assessment time points : baseline (week 0), postintervention (week 12), 6 weeks postintervention (week 18)
	Primary outcomes: gross motor function, functional muscle strength, and walking ability
	1. Gross motor function was measured with the GMFM-66. Possible range 0 to 100 (higher score indicates better gross motor function).
	2. Muscle strength was measured using the 30-s lateral step-up test and the 30-s sit-to-stand test. The number of step-ups the participant could perform in 30 seconds and the number of sit-to-stands the participant could perform in 30 seconds were recorded respectively.
	3. Walking ability was assessed by the time (in seconds) and number of footsteps needed to walk 10 m. The participant was instructed to walk at a self-selected comfortable speed.
	4. Walking ability was assessed by the distance (in m) walked during 1 min. The participant was instruct- ed to walk at his/her fastest attainable speed.
	5. Walking ability was assessed by the time (in seconds) needed to climb a set of 4- or 5-step stairs. The participant was instructed to walk as fast as possible.
	Secondary outcomes: muscle strength, anaerobic power, mobility, and participation
	 Muscle strength was assessed using the 6 RM test on a leg-press. Reported as a percentage of body weight.
	2. Isometric muscle strength of the unilateral hip flexors and abductors, knee flexors and extensors, and ankle plantarflexor muscles were measured with a handheld dynamometer. The mean peak force of 3 tests were reported in n.
	3. Anaerobic power was evaluated during a 20-s full out cycle-test on a child-adapted cycle ergometer. Mean power over 20 seconds was reported in W/kg.
	 Mobility was assessed using the 28 item Mobility Questionnaire. The response options are given on a 5 point scale. Possible range 0-100. Higher score indicates better mobility.
	5. Participation was measured using the Children's Asessment of Participation and Enjoyment question- naire. It was completed by parents with the child. 17 items were chosen from the sub-scales of physi- cal, recreational, and skill based activities to represent frequency in sports participation.
	 Range of motion of the hamstrings, adductors, rectus femoris, soleus and gastrocnemius muscles was assessed with goniometry during the 3rd of 3 slow passive stretches (> 3 seconds). Reported in de- grees.
	7. Spasticity in the hamstrings, adductors, rectus femoris, soleus and gastrocnemius muscles was measured using goniometry by assessing the joint angle at which a 'catch' (defined as a sudden increase in muscle tone, blocking further movement) occurred in a fast passive stretch (< 1 second). Reported on a scale of 0-5. Not clear how the grade was determined.
Notes	Source of funding : the Johanna Kinder-Fonds (2005/0123-357), the Adriaanstichting, and the Phelps Stichting (2006016)
	Potential conflicts of interest : not stated. In the published protocol the authors stated no competing interests; no conflicts of interest perceived.

Exercise interventions for cerebral palsy (Review)

Scholtes 2010 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[r]andomization was performed for each school separately by one in- dependent physiatrist. The children were pre-stratified according to 3 stratifi- cation variables: sex, GMFCS level (I, II-III), age (youngest: 6-9y; oldest 10-13y), and subsequently randomised to one of two groups using sealed envelopes".
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : "[r]andomization was performed for each school separately by one in- dependent physiatrist. The children were pre-stratified according to 3 stratifi- cation variables: sex, GMFCS level (I, II-III), age (youngest: 6-9y; oldest 10-13y), and subsequently randomised to one of two groups using sealed envelopes"
		Comment: did not state if envelopes were opaque
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[t]wo independent research assistants performed all assessments and data entry"
		Comment : both self-reported and objective outcome measures were used. As participants and their parents were not blinded to group allocation, at least 1 outcome is at high risk of bias.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[o]f the 51 included children, one dropped out before T0 (GMFCS III, girl, 13 years 1 month, intervention group) due to a hip-injury which made test- ing and training impossible, and one was lost to follow-up at T1 (GMFCS II, girl, 12 years 1 month, intervention group) due to an unexpected long term stay abroad. Consequently, data on 49 children were included in the analyses (intervention group: n = 24, control group: n = 25)".
		Quote : "[n]ot all children could be motivated to complete all tests on all occasions"
		Comment : missing data rate of approximately 4%. There is some (minimal) variation in the n reported for each outcome at each time point. Reasons for dropout accounted for
Selective reporting (re- porting bias)	Low risk	Comment: all prespecified outcomes reported
Other bias	Low risk	Comment : no other sources of bias identified

Gharib 2011

Methods

Design: randomised controlled trial

Country of origin: Egypt

Intervention(s): Gait Trainer assisted walking exercises

Exercise interventions for cerebral palsy (Review)



Gharib 2011 (Continued)	Unit of allocation: individual
Participants	Number of participants : 30 randomised; exercise group: n = 15; control group: n = 15
	Group characteristics reported for all randomised participants
	Age : mean (SD) = 11.55 (1.11) years; exercise group: mean (SD) = 11.87 (1.06) years; control group: mean (SD) = 11.23 (1.11) years
	Sex (male/female): 16/14; exercise group: 10/5; control group: 6/9
	Ethnicity: not stated
	GMFCS level: level II (n = 30)
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: hemiplegia
	Inclusion criteria : unclear if the following is stated as inclusion criteria or a description of participants included: age 10-13 years, hemiparetic CP with a mild degree of spasticity in affected lower limbs (Mod-ified Ashworth Scale score < 2), classified as Gross Motor Function Classification Scale level II
	Exclusion criteria : fracture, sprain or strain injury of the lower extremities in the past 6 months, neuro- logical or orthopaedic surgery in the last 12 months, botulinum toxin application for at least 6 months before the study, exercise-induced asthma, a congenital heart defect with cardiac compromise, ag- gressive or self-harming behaviours, cognitive impairment (not being able to follow simple verbal com- mands and instructions during tests and training), uncontrolled seizure disorder
Interventions	Aim of the intervention: to improve walking parameters
	Type of exercise programme: aerobic
	Exercise mode : Biodex Gait Trainer 2 (no body weight support)
	Comparator : all participants in both groups received a physical therapy exercise session of 30 min, 3 times per week. This included stretching, strengthening, practicing of activities of daily living, balance and gait exercises
	Setting: physiotherapy clinic
	Intervention provider: physiotherapist
	Duration of programme: 3 months
	Exercise dose: participants encouraged to walk continuously for 15 min, 3 times per week
	Tailoring of intervention to individual : a rest break of 1-3 min was given to a participant if they re- quested it. The gait trainer belt speed was gradually increased for each participant.
	Fidelity to prescribed intervention : not reported; if a participant missed > 3 sessions because of med- ical reasons or an inability to participate the participant was considered as dropped from the study. No participant was excluded
	Monitoring of adverse events: not reported
Outcomes	Assessment time points: baseline (week 0) and postintervention (3 months)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	The following procedure was used to measure all gait parameters. The speed of the belt was slowly in- creased to a comfortable pace for the participant at which point the participant walked continuously

Exercise interventions for cerebral palsy (Review)



 Walking speed was reported in m/s Step length was reported in m Ambulation index, a composite score based on foot-to-foot time distribution and average step was reported on a scale of 0-100, where 100 indicates better gait parameters. Time on each foot was recorded as a percentage of gait cycle. 	
1. Walking speed was reported in m/s 2. Step length was reported in m 3. Ambulation index, a composite score based on foot-to-foot time distribution and average step	
 Walking speed was reported in m/s 	p cycle,
·	
Gharib 2011 (Continued) for 3 min while data was collected. This was repeated 3 times and the average was taken for each parameter.	h gait

Potential conflicts of interest: not stated; none perceived

Risk of bias	Risk of bias		
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[c]hildren were randomly assigned to one of two groups"	
tion (selection bias)		Quote : "[r]andom assignment of children was conducted into two stages. Stage one involved instructing two physical therapists who were working in the paediatric physical therapy outpatient clinic to report all children who ful- filled the inclusion criteria of the study (registration diagnosis, age, level of Gross Motor Function Classification System scale), and had no exclusion crite- ria. The second stage involved randomly assigning the children to either the experimental group or the control group by using sealed envelopes"	
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement	
Allocation concealment (selection bias)	Unclear risk	Quote : "[t]he second stage involved randomly assigning the children to either the experimental group or the control group by using sealed envelopes. The randomisation process was carried out by a registration clerk who was not involved in any part of the study"	
		Comment : insufficient information provided to determine if a person involved in the study could predict allocation	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation	
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote : "[a]ll children (in both groups) were evaluated prior to the commence- ment of baseline training and at the end of the three-month training period (post-treatment) by the same examiner who was blinded to which group each child was assigned"	
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment: no missing data	
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes	

Comment: no other sources of bias identified

Exercise interventions for cerebral palsy (Review)

Other bias

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Low risk

Methods	Design: randomised controlled trial
	Country of origin: USA
	Intervention(s): a comparison of supported treadmill training and strength training
	Unit of allocation: individual
Participants	Number of participants : 34 randomised. Outcome and demographic data only presented for participants who completed the study (n = 26); treadmil exercise group: n =14; strengthening exercise group: n = 12
	Age : mean (SD) = 9 years 6 months (2 years 2 months); treadmill exercise group: mean (SD) = 9 years 7 months (2 years 2 months); strengthening exercise group: mean (SD) = 9 years 6 months (2 years 4 months)
	Sex (male/female): 14/12; treadmill exercise group: 7/7; strengthening exercise group: 7/5
	Ethnicity: not stated
	GMFCS level : level II (n = 2), level III (n = 15), level IV (n = 9); treadmill exercise group: level II (n = 1), level III (n = 9), level IV (n = 4); strengthening exercise group: level II (n = 1), level III (n = 6), level IV (n = 5)
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP : diplegia (n = 12), triplegia (n = 2) and quadriplegia (n = 12); treadmill exercise group: diplegia (n = 8) and quadriplegia (n = 6); strengthening exercise group: diplegia (n = 4), triplegia (n = 2), and quadriplegia (n = 6)
	Inclusion criteria : spastic CP, marginal ambulatory function (defined as decreased gait velocity < 80% of age-expected value regardless of GMFCS level, or GMFCS level III or IV), ability to take 8 steps inde- pendently with or without assistive devices, able to complete 3D gait analysis, body weight < 68 kg (150 lb), age 6-13 years, ability to follow multiple-step commands
	Exclusion criteria : medical condition that would be negatively affected by exercise; lower extremity orthopaedic surgery in the past year, botulinum toxin A in the past 6 months, dorsal rhizomotomy in the past 2 years; flexion contractures > 30° at the hip or > 20° at the knee or plantarflexion contractures > 15°, intrathecal baclofen, dystonia, athetoid or mixed types of CP
Interventions	Aim of the intervention : to increase strength, motor control, gait, gross motor skills, and physical function, and improve quality of life, self-concept, satisfaction and participation (not sure if needed)
	Type of exercise programme: aerobic exercise versus resistance training
	Exercise mode : treadmill exercise group: completed a programme of partial body-weight-supported treadmill training
	Comparator : strengthening exercise group completed a programme of step-ups, squats, upper and lower limb progressive resistance exercise, and core strengthening
	Setting : home-based programme. First 2 weeks were delivered at study site or in home. Final 10 weeks were delivered at home
	Intervention provider : physiotherapist delivered programme for 2 weeks and parents delivered it for 10 weeks
	Duration of programme: 12 weeks
	Exercise dose : all participants were prescribed 2 sessions of 30 min, 5 days/week for first 2 weeks and session of 30 min, 5 days/week for final 10 weeks
	Tailoring of intervention to individual:

Exercise interventions for cerebral palsy (Review)

Johnston 2011 (Continued)	Treadmill exercise group: each participant wore a harness and the parent or physiotherapist guided
	their leg if necessary, and ankle-foot orthoses were used where necessary. Initial training speed based on participant's baseline gait speed and adjusted as needed based on participant's response. Aimed to decrease body-weight support to < 30% and increase speed to normal values during the 2-week induc tion period, based on the participant's ability to maintain normal gait pattern
	Strengthening exercise group: assistance was provided as needed, and assistive devices were allowed for weight bearing. The exercises were advanced on an individual basis by increasing the number of repetitions and then adding resistance by cuff weights.
	Fidelity to prescribed intervention : parents of participants kept weekly logs. All participants who completed the study attained an adherence rate of at least 80%
	Monitoring of adverse events : did not state how adverse events were assessed. 2 participants in the treadmill group complained of leg/knee discomfort which resolved without intervention. 1 participant developed a blister from the AFO during the induction period, unclear group allocation
Outcomes	Assessment time points : baseline (week 0), postintervention (week 12), 4 weeks postintervention (week 16)
	Primary outcome:
	 Gait speed (m/s) was measured using 3-dimensional motion analysis. Participants walked at their sel selected speed using their commonly used assistive device and ankle orthosis.
	Outcomes:
	 Spasticity of the plantarflexors and knee flexors was assessed with the participant in a semi-supin position on a dynamometer reported in J/°/s.
	 Muscle strength of the plantar- and dorsiflexors, and knee flexors and extensors was measured usin a dynamometer. The highest value from 3 trials at 10°/s was defined as the maximal strength an reported in n/kg.
	3. Motor control of the quadriceps was tested using a dynamometer 10°/s and reported in N.
	 Cadence was measured using 3-dimensional motion analysis and reported in steps/min. Participan walked at their self-selected speed using their commonly used assistive device and ankle-foot orth sis.
	Stride length was measured using 3-dimensional motion analysis and reported in m. Participan walked at their self-selected speed using their commonly used assistive device and ankle orthosis.
	 Gross motor function was measured using dimensions A to E of the GMFM. Possible range 0 to 10 (higher score indicates better gross motor function).
	7. Gross motor function was measured using the parent report of the Pediatric Outcomes Data Colle tion Instrument (PODCI). The parent report for the PODCI global function score and the transfers ar mobility scale were used as the primary measures of function. Although not stated it appears the sta dardised score was reported, the value of which can range from 0 to 100, where a higher score inc cates better function.
	8. The Children's Assessment of Participation and Enjoyment (CAPE) questionnaire was used to me sure participation. It was completed by the participant with the therapist's assistance. Participation scores were provided on 5 subscales: intensity (higher score indicates more time spent participating 'with whom' (higher score indicates more social activities), diversity (higher score indicates more or verse participation), 'where' (higher score indicates more community-based activities) and enjoyme (higher score indicates more pleasure experienced from participating).
	 Quality of life was measured using the Pediatric Quality of Life Inventory-CP Module (PedsQLCP). Bo the primary caregiver and the participant completed the PedsQLCP. The summary score is on a 0-10 scale with a higher score reflecting better quality of life.
	10.Self-concept was measured with the child-reported Piers-Harris Children's Self-Concept Scale. A hig er total self-concept score demonstrates greater self-concept.
	11.The Canadian Occupation Performance Measure (COPM) was used to set individualised goals and a sess the degree of satisfaction with goals attained. Administered as a semi-structured interview b tween therapist and participant. Higher scores indicate improvement.

Exercise interventions for cerebral palsy (Review)



Johnston 2011 (Continued)

Notes

Source of funding: Shriners Hospitals for Children (grant no. 9147)

Potential conflicts of interest: authors report no financial conflict of interest; none perceived

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[c]hildren were randomly assigned to the SSTTEP or exercise group using a block randomization schedule at each site using blocks of eight group assignments (four of each intervention per block)."
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Comment : insufficient information regarding allocation concealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[t]wo sites were able to blind the evaluators to group assignment (representing 16 children) but the third site was unable to because of personnel issues. All evaluators were blinded to results and trained on each measure for which written study specific protocols were available at each site"
		Comment : 26 participants completed the study. Therefore 10 participants were assessed by evaluators not blinded to group allocation. Both self-report and objective outcome measures were used. As participants and their parents were not blinded to group allocation, at least 1 outcome is at high risk of bias.
Incomplete outcome data	High risk	Quote : "[d]id not receive allocated intervention (n=2)"
(attrition bias) All outcomes		Quote : "Lost to follow-up (n=2). Discontinued participation part way through intervention period".
		Quote : "[s]ix participants were lost to follow-up at the 12-week point because of personal and family reasons not related to the intervention. Two participants (in the SSTTEP group) did not participate in data collection after the washout period."
		Quote : "[c]hildren were required to complete at least 40 out of 50 full sessions (80% adherence) to participate in data collection.
		Comment : 34 participants were randomised and only 26 participants were included in the analysis (24% missing data). Missing data were evenly distributed across groups (22% and 25%, respectively). All reasons for withdrawals not specified. As-treated analysis undertaken with substantial departure of the intervention received from that assigned at randomisation
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment : baseline and demographic data is only reported for those who completed the trial; not all completers' data is reported for all outcomes, no reason given

Exercise interventions for cerebral palsy (Review)



Olama 2011

Methods	Design: randomised controlled trial			
	Country of origin: Egypt			
	Intervention(s): a comparison of a treadmill training programme to a resistance training programme			
	Unit of allocation: individual			
Participants	Number of participants : unclear. States 30 participants but also states that there were 12 males and 8 females, and 12 participants with left hemiplegia and 8 participants with right hemiplegia; treadmill ex ercise group: n = 15; Resistance exercise group: n =15			
	Age : mean (SD) = 13.73 (0.85) years			
	Sex (male/female): 12/8			
	Ethnicity: not stated			
	GMFCS level: not stated			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of CP: hemiplegia			
	Inclusion criteria : does not state inclusion criteria but describes sample as: age 12-15 years, able to un derstand any command, with an IQ level within normal range, free from any associated disorders othe than spasticity, spasticity in the range of 1+ and 2 on the Modified Ashworth Scale, free from structural changes of the joints of the lower limbs, able to walk independently			
	Exclusion criteria: not stated			
Interventions	Aim of the intervention: to increase muscle strength			
	Type of exercise programme : treadmill exercise group: aerobic; resistance exercise group: resistance training			
	Exercise mode : treadmill exercise group: treadmill; resistance exercise group: resistance training of quadriceps and hamstrings			
	Comparator : both groups received an exercise programme consisting of neurodevelopmental ther- apy, proprioceptive training, facilitation of righting reactions, stretching, and gait training, daily for 6 months			
	Setting: not stated			
	Intervention provider: not stated			
	Duration of programme : unclear. States participants received 6 months of treatment but also states that the exercise procedure for the resistance training group was repeated 3 times per week for 3 months.			
	Exercise dose : treadmill exercise group: daily (no indication of duration of treadmill training); resis- tance exercise group: 3 times per week			
	Tailoring of intervention to individual:			
	Treadmill exercise group: participants can hold on with 2 hands until they feel confident to walk with- out support.			
	Resistance exercise group: the resistance added to the exercise was gradually increased starting at 10 repetitions at 50% of 10 RM, 10 repetitions at 75% of 10 RM and 10 repetitions at 100% of 10 RM			
	Fidelity to prescribed intervention: not reported			

Exercise interventions for cerebral palsy (Review)

Olama 2011 (Continued)			
	Monitoring of adverse events: not reported		
Outcomes	Assessment time points: baseline (week 0) and postintervention (6 months)		
	Primary outcome: no primary outcome measure stated.		
	Outcomes:		
	 Concentric muscle strength of the knee flexors and knee extensors at 60°/s and 180°/s were measured using a dynamometer. The mean ratio of peak torque to body weight of 3 tests was used in analysis. 		
	2. Gross motor function was measured with the Bruininks-Oseretsity test for motor proficiency. Higher score indicates better gross motor function.		

Notes

Source of funding: not stated

Potential conflicts of interest: not stated; none perceived

Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote: "[t]he study sample was divided into two groups of equal size"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment	Unclear risk	Quote: "[t]he study sample was divided into two groups of equal size"
(selection bias)		Comment : insufficient information regarding allocation concealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Quote : "[d]ouble-blind evaluation was conducted for each child individually before and after six months of treatment".
		Comment : insufficient information regarding who was blinded to make judge- ment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Comment : no indication of how many participants completed the study; no indication of how many participants were included in the analysis.
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment: no demographic data provided

Pandey 2011

Methods

Design: randomised controlled trial

Country of origin: India

Intervention(s): resistance training

Exercise interventions for cerebral palsy (Review)



Pandey 2011 (Continued)	Unit of allocation: individual		
Participants	Number of participants: 18 randomised; exercise group: n = 9; control group: n = 9		
	Age: not stated		
	Sex (male/female): 11/7		
	Ethnicity: not stated		
	GMFCS level: not stated		
	Type of motor abnormality: spastic		
	Anatomical distribution of CP: diplegia		
	Inclusion criteria : diagnosis of spastic CP; aged 5-10 years, able to walk with or without aid, able to extend knee from 90° to 45° or more in sitting position with full passive range of motion in supine, no known cognitive impairment, able to flex knee to 90° in prone position, spasticity grade 2 or less than 2 on Modified Ashworth Scale (for hip adductors and abductors, knee flexors, and ankle plantarflexors)		
	Exclusion criteria : known cognitive impairment, orthopaedic or medical condition that prevents exer- cising, cerebellar symptoms, known visual, speech, hearing disorders, systemic medical problem which prevents exercising, lower limb surgery within 12 months, on anticonvulsant, antispastic medication, non-ambulatory		
Interventions	Aim of the intervention: to increase lower limb strength, improve segmental control of lower limbs, and improve balance		
	Type of exercise programme: resistance training		
	Exercise mode : bilateral heel raises, sit-to-stand exercises, standing balance exercises, step-ups, vestibular ball supported half squat		
	Comparator : not clearly stated. States "none were allowed to attend physiotherapy other than inter- vention protocol"		
	Setting: not stated		
	Intervention provider: physiotherapist		
	Duration of programme: 4 weeks		
	Exercise dose: 1 h session, twice per week. Number of sets, repetitions and resistance used not stated		
	Tailoring of intervention to individual: not stated		
	Fidelity to prescribed intervention: not reported		
	Monitoring of adverse events: not reported		
Outcomes	Assessment time points : baseline (week 0), postintervention (week 4), follow-up (not stated how long after intervention)		
	Primary outcome: no primary outcome measure stated.		
	Outcomes:		
	 Muscle strength was assessed by the number of lateral step-ups completed in 15 seconds. Gross motor function was assessed by assessing the minimum height of a chair (reported in cm) that participants could stand up from for 3 successive repetitions. 		
	3. Gross motor function was assessed by the motor assessment scale, sit-to-stand sub-test. It is not clear how this was conducted or the outcome reported or measured.		

Exercise interventions for cerebral palsy (Review)



Pandey 2011 (Continued) 4. Gross motor function was assessed by the stride length (reported in m) and cadence (reported in steps/min) calculated from the 10-m walk test. It's not stated if this was performed at maximum speed or comfortable speed.

5. It is stated that walking speed was assessed during the 10-m walk test and the 2-min walk test. However, only 1 walking speed (in m/s) is presented and it's not clear if it was recorded during the 10-m walk test or 2-min walk test. Further, it's not clear if it was self-selected or maximum gait speed

Source of funding: not stated

Potential conflicts of interest: not stated; none perceived

Risk of bias

Notes

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "[a]ll eligible subjects were randomly assigned to intervention group and control group, through use of random number generator with sealed envelopes"
Allocation concealment (selection bias)	Unclear risk	Quote : "[a]Il eligible subjects were randomly assigned to intervention group and control group, through use of random number generator with sealed envelopes"
		Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Quote : "[t]he study employed a randomized single blind controlled trial de- sign consisting of two groups and three measurements, training was conduct- ed in one-hour sessions twice a week for four weeks"
		Comment : unclear who was blinded to group allocation
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Comment : no indication of how many participants completed the study; no indication of how many participants were included in the analysis
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment : there is no clear statement of numbers in each group or how many were analysed. Unclear whether baseline data is just for those who completed the trial

Smania 2011

Methods

Design: randomised controlled trial

Country of origin: Italy

Intervention(s): repetitive locomotor training

Exercise interventions for cerebral palsy (Review)



Smania 2011 (Continued)	Unit of allocation: individual			
Participants	Number of participants : 18 randomised; exercise group: n = 9; control group: n = 9			
	Group characteristics reported for all randomised participants			
	Age : mean (SD) = 13.30 (2.91) years; exercise group: mean (SD) = 13.87 (2.79) years; control group: mean (SD) = 12.72 (3.08) years			
	Sex (male/female): 10/8; exercise group: 4/5; control group: 6/3			
	Ethnicity: not stated			
	GMFCS level : level I (n = 6), level II (n = 2), level III (n = 3), and level IV (n = 7); exercise group: level I (n = 3), level II (n = 2) and level IV (n = 4); control group: level I (n = 3), level III (n = 3) and level IV (n = 3)			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of CP : tetraplegia (n = 7) and diplegia (n = 11); exercise group: tetraplegia (n = 4) and diplegia (n = 5); control group: tetraplegia (n = 3) and diplegia (n = 6)			
	Inclusion criteria : bilateral lower limb (diplegia or tetraplegia) CP, aged 10-18 years, GMFCS levels II, III or IV, ability to walk independently with or without an aid for at least 10 m, maintain a sitting position without assistance, follow instructions, participate in the programme			
	Exclusion criteria : lower limb spasticity ≥ 2 on the Modified Ashworth Scale, severe lower limb con- tractures, cardiovascular diseases, orthopaedic surgery or neurosurgery in the past 12 months or botu- linum toxin injections in past 6 months			
Interventions	Aim of the intervention: to increase walking speed and endurance			
	Type of exercise programme: aerobic			
	Exercise mode : gait trainer (Gait Trainer GT I). 30 min of repetitive locomotor therapy on the gait train- er and 10 min of passive joint mobilisation and stretching			
	Comparator : usual physiotherapy: passive joint mobilisation and stretching of lower limbs, strength- ening exercises including leg-press and sit-to-stand exercises with resistance adapted for participant's ability, balance and gait exercises. Each type of exercise lasted 10, 15 and 15 min, respectively			
	Setting: rehabilitation gym in medical centre			
	Intervention provider : a physiotherapist delivered the exercise programme to both the exercise group and the control group			
	Duration of programme: 2 weeks			
	Exercise dose : both groups received 10 sessions of 40 min, 5 days per week.			
	Tailoring of intervention to individual : gait speed and step length were individually set according to the gait parameters recorded at baseline. Walking speed was gradually increased over the course of the 2 weeks if the participant completed the last training session without discomfort or fatigue. Partial body weight support was decreased from 30% to 0% over the duration of the sessions. The criterion for reduction was the participants' ability to avoid their knees collapsing into flexion during the stance phase because of increased load of body weight.			
	Fidelity to prescribed intervention: not reported			
	Monitoring of adverse events : no adverse events that led to a missed session, no joint pain or muscle spasms were reported during or after the intervention. Unclear how adverse events were monitored.			
Outcomes	Assessment time points : baseline (week 0), postintervention (week 2), 4 weeks postintervention (weel 6)			
	Primary outcome: gait speed and walking endurance			

Exercise interventions for cerebral palsy (Review)



Smania 2011 (Continued)

Trusted evidence. Informed decisions. Better health.

m/s

ing speed. Distance was reported in metres

Secondary outcomes: gait parameters

	analysis. The follow a. Joint angles of th swing; unclear w b. Speed; unclear v c. Cadence; unclea	re measured while participants walked at their self-selected speed using motion ing parameters were measured: he hip, knee and ankle at initial contact, middle stance, and initial swing and middle that unit joint angle was reported in. what unit speed was reported in. r what unit cadence was reported in. lear what unit step length was reported in.	
Notes	Source of funding: the CariVerona Fondation (PACIS) Potential conflicts of interest: declares no conflicts of interest; none perceived		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Low risk	Quote : "[b]efore the start of the study participants were allocated to the experimental group or the control group via computerised randomisation. The randomisation sequence was generated by a research assistant not involved with the study and the group allocation was concealed using sealed numbered envelopes"	
Allocation concealment (selection bias)	Unclear risk	Quote : "[t]he randomisation sequence was generated by a research assistant not involved with the study and the group allocation was concealed using sealed numbered envelopes. The randomisation list was locked in a desk drawer accessible only to the principal investigator".	
		Comment : PI has access to the randomised list, unclear whether envelopes were opaque.	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation.	
Blinding of outcome as- sessment (detection bias)	High risk	Quote : "[t]he participants were evaluated by the same examiner who was un- aware of treatment allocation".	
All outcomes		Comment : both self-reported and objective outcome measures were used. Al- though it does not state who completes the self-report outcome measure, giv- en its nature, we assume participants and/or their parents were not blinded to group allocation.	
Incomplete outcome data	Low risk	Quote: "[n]o children withdrew from the study"	
(attrition bias) All outcomes		Comment: no missing data	
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes	
Other bias	Low risk	Comment : no other sources of bias identified	

1. Self-selected gait speed was measured (in seconds) while the participant walked 10 m. Reported in

2. Walking endurance was measured while the participant walked for 6 min at his/her self-selected walk-

Exercise interventions for cerebral palsy (Review)



Chen 2012

Methods	Design: randomised controlled trial			
	Country of origin: Taiwan			
	Intervention(s): home-based virtual cycling training programme			
	Unit of allocation: individual			
Participants	Number of participants : 30 randomised; cycling group: n = 14; control group: n = 16.			
	Age, sex, GMFCS level and anatomical distribution reported for 27 participants (cycling group, n = 13, and control group, n = 14) who completed the study.			
	Age : 6-12 years; cycling group: mean (SD) = 8.7 (2.1) years; control group: mean (SD) = 8.5 (2.2) years			
	Sex (male/female): 18/9; cycling group: 9/4; control group: 9/5			
	Ethnicity: not reported			
	GMFCS level : levels I and II; cycling group: level I (n = 10) and level II (n = 3); control group: level I (n = 11) and level II (n = 3)			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of motor abnormality : diplegia (n = 19) and hemiplegia (n = 8); cycling group: diplegia (n = 10) and hemiplegia (n = 3); control group: diplegia (n = 9) and hemiplegia (n = 5)			
	Inclusion criteria : aged 6-12 years, diagnosis of spastic CP with GMFCS levels I-II, in prepubertal stage, ability to walk independently, ability to undergo motor function and isokinetic muscle test, ability to comprehend commands and cooperate during an examination			
	Exclusion criteria : children and adolescents with recognised chromosomal abnormalities, a progress sive neurological disorder or severe concurrent illness or disease that is not typically associated with CP, active medical conditions such as pneumonia, any major surgery or nerve block in the preceding months, hormonal disturbance, poor tolerance for performing the isokinetic test or a poor ability to operate during assessment			
Interventions	Aim of the intervention: to improve muscle strength, motor function and bone density			
	Type of exercise programme: mixed (aerobic and resistance training)			
	Exercise mode : cycling group: cycled on an ergometer and sit-to-stand exercises (n = 14). Virtual cy- cling system; CD-ROM allowed participant to cycle in virtual world and guided participant through ex cises			
	Comparator : encouraged to perform general physical activity at home which involved walking, run- ning, jogging or sports or recreational exercises at school or at home for 30-40 min/day, 3 days/week. Supervised by parent			
	Setting: home-based programme; individual session			
	Intervention provider: virtual cycling system			
	Duration of programme: 12 week programme			
	Exercise dose : cycling group: received 40 min of exercise (20 min of cycling and 2 sets of 10 repetitions of loaded sit-to-stand exercises), 3 days per week			
	Tailoring of intervention to individual : the resistance of the bike was progressively increased depending on the ability of the participant. The loaded sit-to-stand exercises were adjusted by adding weight bags to a back-pack worn by the participant, and the weight of the bags ranged from 0.5 to 3 kg. Participants trained with loads of 75% of 1 RM.			

Exercise interventions for cerebral palsy (Review)



Chen 2012 (Continued)	Fidelity to prescribed intervention : compliance recorded for both groups by a research assistant. Parents and participants were interviewed about the implementation of the programmes by a research assistant via telephone every 1-2 weeks. The participants and caregivers were followed up at the reha- bilitation unit every month. Compliance data not reported.			
	Monitoring of adverse events : no adverse effects observed during intervention for either group; not stated how adverse events were monitored			
Outcomes	Assessment time points : baseline (week 0) and postintervention (week 12) Primary outcome : no primary outcome measure stated			
	Outcomes:			
	 Gross motor function assessed with the Bruininks-Oseretsky Test of Motor Proficiency (BOTMP) by a physiotherapist. A raw score is obtained for 8 norm-referenced subtests and converted to point scores. By comparing with norm, the point score is converted into a standard score for each subtest; a higher standard score indicates a better subtest performance. The sum of the standard scores for 4 sub-tests (running speed and agility, balance, bilateral coordination, and strength) contributed to the standard score of gross motor composite; a higher score indicates better gross motor function. Gross motor function was also assessed using the GMFM-66. Possible range 0-100 (higher score indi- 			
	 cates better gross motor function) 3. Muscle strength was assessed by a physiotherapist with an isokinetic dynamometer. Knee extension and flexion torque was measured in the more-affected lower limb during repeated extension-flexion. Testing angular velocity was set to 60°/s, 90°/s and 120°/s, and range of motion was set to 70° starting with the knee flexed at 80° and ending in an extension at -10°. The isokinetic peak torque of the knee extensor and knee flexor was normalised by body weight (Nm/kg). The strength change index (%) was calculated as percentage of (post-treatment isokinetic peak torque - pre-treatment isokinetic peak torque. 			
	4. Muscle strength of the trunk was assessed using curl-ups. The maximum number of curl-ups per- formed correctly in 1 min was recorded.			
	 Areal Bone Mineral Density (aBMD) (g/cm²) was measured at the lumbar spine (L1 to L4) and the distal femur of the more affected limb using Dual X-ray absorptiometry. 			
Notes	Source of funding: National Science Council of the Republic of China, Taiwan			
	Potential conflicts of interest : authors state they have no conflicts of interest; no conflicts of interest perceived			
Risk of bias				

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[p]articipants were randomly assigned to the hVCT group or to the control group"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Comment : no information provided to indicate if allocation was concealed or not
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[a] physical therapist, who was not blinded to group allocation was trained to use an isokinetic dynamometer and the GMFM as a precondition of study participation"

Exercise interventions for cerebral palsy (Review)

Chen 2012 (Continued)		
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Quote : "[o]ne child in the control group and one child in the experimental groups dropped out due to lack of time to complete the study"
		Comment : in addition to the participants who dropped out of the study due to lack of time, 1 participant in the control group's data was excluded from the analysis; reasons not provided The reasons for missing data are not described in enough detail to make a judgement regarding the effect of attrition bias on results
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported
Other bias	Unclear risk	Comment : demographic and baseline data provided for those who completed the study only

Methods	Design: randomised controlled trial			
	Country of origin: Greece			
	Intervention(s): treadmill training programme			
	Unit of allocation: individual			
Participants	Number of participants : 22 randomised; exercise group: n = 11; control group: n = 11			
	Group characteristics were reported for all randomised participants			
	Age : 13-19 years; exercise group: mean (SD) = 15.9 (1.97) years; control group: mean (SD) = 16.09 (1.51) years.			
	Sex (male/female): 13/9; exercise group: 6/5; control group: 7/4			
	Ethnicity: not stated			
	GMFCS level : level I (n = 5), level II (n = 9), level III (n = 8); exercise group: level I (n = 3), level II (n = 4), level III (n = 4); control group: level I (n = 2), level II (n = 5), level III (n = 4)			
	Type of motor abnormality: spastic CP			
	Anatomical distribution of motor abnormality : diplegia (n = 19) and tetraplegia (n = 3); exercise group: diplegia (n = 9), tetraplegia (n = 2); control group: diplegia (n = 10), tetraplegia (n = 1)			
	Inclusion criteria : a diagnosis of spastic CP (tetraplegia or diplegia), GMFCS levels I-III, ability to follow simple commands			
	Exclusion criteria : undergone surgery in previous 12 months, received botulinum toxin injections in the previous 6 months, had cardiovascular disease or uncontrolled epilepsy			
Interventions	Aim of the intervention: to improve gross motor function and self-selected walking speed			
	Type of exercise programme : aerobic training. Parents and guardians were instructed to continue the daily activities of their participants, and not to initiate any additional interventions or increase the usual daily physical activities of their participants throughout the duration of the programme			
	Exercise mode : treadmill training (no bodyweight support).			
	Comparator : conventional physiotherapy delivered by a physical therapist. 3 sessions of 45 min each consisting of 3 sets of exercises with mat activities, balance, and gait training and functional gross mo tor activities; adherence not reported			

Exercise interventions for cerebral palsy (Review)

Chrysagis 2012 (Continued)	Setting: special school; individual supervised session
	Intervention provider: 2 physical therapists
	Duration of programme: 12 weeks
	Exercise dose : participants in the exercise group underwent treadmill training on 3 days per week for a total of 34 sessions (2 sessions cancelled). Each session consisted of a 10 min warm-up, walking for a maximum of 30 min at a comfortable speed, and a 5 min cool down. Participants completed between 12.00 and 20.00 min (mean (SD) 16.43 (2.59) min) per session at the start, and between 23.90 and 29.80 min (mean (SD) 27.54 (2.12) min) per session, at the end of the training. Mean (SD) treadmill speed increased from 1.76 (0.41) km/h at the beginning to 3.00 (0.60) km/h at the end of the programme.
	Tailoring of intervention to individual : training was customised to each participant's ability and the physiotherapist's supervision of the participants' safety and facilitation of the walking pattern, without elicitation of abnormal movements. Adolescents held handrails when needed.
	Fidelity to prescribed intervention : therapists kept logbooks during each session for the exercise group. Adherence ranged from 26 to 34 sessions (mean (SD) 29.45 (2.84) sessions).
	Monitoring of adverse events : no change in spasticity as measured by Modified Ashworth Scale. No participant experienced a fall or complained of fatigue, soreness or pain as recorded by the therapist in a logbook.
Outcomes	Assessment time points: baseline (week 0) and postintervention (week 12)
	Primary outcomes: gross motor function and self-selected walking speed
	 Gross motor function was measured by 2 trained assessors (experienced physiotherapists) using di- mensions D and E of the Gross Motor Function Measure. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. Scores presented as percentage of total possible score (higher score indicates better gross motor function).
	2. Self-selected walking speed was evaluated with the 10-m walk test by 2 trained assessors (experienced physiotherapists). Reported as m/min
	Secondary outcomes:
	 Spasticity of the knee flexors, knee extensors, and foot plantar flexors were evaluated according to the modified Ashworth Scale (scored as 0, 1, 1⁺, 2, 3, 4); a higher score indicates greater spasticity
Notes	Source of funding: none stated
	Potential conflicts of interest: authors report no conflict of interest; none perceived
Risk of bias	

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[t]he participants were stratified according to GMFCS level and sex and then randomly allocated to the experimental and control groups. The process was led by a PhD student working in the laboratory who was not in- volved with the participants or the experimental treatment. The student used sealed envelopes to assign the participants to the experimental and control conditions".
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Quote : "[t]he process was led by a PhD student working in the laboratory who was not involved with the participants or the experimental treatment. The student used sealed envelopes to assign the participants to the experimental and control conditions"

Exercise interventions for cerebral palsy (Review)



Chrysagis 2012 (Continued)

Jong Dorr (continued)		Comment : did not state if envelopes were opaque
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote : "[t]wo trained assessors who were blinded to the treatment alloca- tions carried out the measurements at the school"
Incomplete outcome data	Low risk	Quote : "[t]here were no dropouts"
(attrition bias) All outcomes		Comment : no missing data reported
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported. No convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment: no other sources of bias identified

Bryant 2013

Methods	Design: randomised controlled trial Country of origin: UK			
	Intervention(s): aerobic exercise intervention using a static bike or treadmill			
	Unit of allocation: individual			
Participants	Number of participants : 35 randomised; cycling group: n = 11; treadmill group: n = 12; control group: r = 12			
	Group characteristics were reported for all randomised participants			
	Age : 8-17 years; mean (SD) = 13 years 9 months (2 years 3 months); cycling group: mean (SD) = 14.3 (1.9) years; treadmill group: mean (SD) = 13.5 (2.6) years; control group: mean (SD) = 13.8 (2.3) years			
	Sex (male/female): 14/21; cycling group: 6/5; treadmill group: 3/9; control group: 5/7			
	Ethnicity: not reported			
	GMFCS level : level IV (n = 23) and V (n = 12); cycling group: level IV (n = 8) and V (n = 3); treadmill group: level IV (n = 8) and V (n = 4); control group: level IV (n = 7) and V (n = 5).			
	Type of motor abnormality : dyskinetic (n = 14) and spastic (n = 21); cycling group: dyskinetic (n = 4) and spastic (n = 7); treadmill group: dyskinetic (n = 3) and spastic (n = 9); control group: dyskinetic (n = 7) and spastic (n = 5)			
	Anatomical distribution of motor abnormality: bilateral			
	Inclusion criteria : aged 8-17 years, CP at GMFCS level IV and V, able to pedal on adapted static bike and walk with partial bodyweight support on a treadmill			
	Exclusion criteria : undergone orthopaedic surgery to the spine or lower limbs within the last year, cog nitive or behavioural impairment preventing understanding or compliance with instructions			
Interventions	Aim of the intervention: to improve gross motor function			

Exercise interventions for cerebral palsy (Review)

Bryant 2013 (Continued)	Type of exercise prog	ramme: aerobic training	
	Exercise mode : cycling group cycled on an adapted static bike (n = 11); treadmill group walked supported with a hoist on a treadmill, minimal operating speed 0.5km/h (n = 12)		
	Comparator : control group received usual physiotherapy such as stretching and exercise on a mat, use of a standing frame and swimming, dose not reported (n = 12)		
	Setting: 4 special schools; unclear if supervised individual or group session		
	Intervention provider: not stated		
	Duration of programme: all groups received a 6-week programme		
	Exercise dose : cycling group and treadmill group received 30 min of exercise (including transfers), 3 days per week. An exercise test was used to determine starting level of exercise and to monitor progress in ability. However, the exercise intensity is unclear.		
	-	ion to individual : bikes were adapted to provide extra postural support. Partici- p were supported during walking on treadmill with a hoist.	
	Fidelity to prescribed intervention : the number of training sessions attended were recorded but it's not stated how attendance was recorded. Mean (SD) number of sessions attended was 14.6 (3.1) for cycling group and 13.8 (4.2) for treadmill group		
	Monitoring of adverse events : reports no adverse events. In the treadmill group 1 participant with- drew due to hospitalisation because of gastric problems and 1 participant withdrew due to a reoccur- rence of long-standing hip pain; not stated how adverse events were monitored		
Outcomes	Assessment time points : baseline (week 0), postintervention (week 6), 6 weeks postintervention (week 12), 12 weeks postintervention (week 18)		
	Primary outcome: no primary outcome measure stated		
	Outcomes:		
	 Gross motor function measured with the Gross Motor Function Measure (GMFM-66). Possible range 0-100 (higher score indicates better gross motor function). 		
	2. Gross motor function measure with dimensions D and E of the GMFM-88. Possible range 0-39 for di- mension D and 0-72 for dimension E (higher score indicates better gross motor function).		
	3. GMFM-66 and dimensions D and E of the GMFM-88 were assessed at all time points and 1 researcher		
	acted as an assessor for all but 3 of the study assessments 4 Speed (kpb) and duration of evercise (min) were evaluated for those in cycling group and treadmill		
	4. Speed (kph) and duration of exercise (min) were evaluated for those in cycling group and treadmill group at week 0 and week 6		
Notes	Source of funding: Nat	tional Institute for Health Research (NIHR)	
	Potential conflicts of i	interest: did not declare conflicts of interest; no perceived conflicts of interest.	
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[t]he study administrator provided nine sealed envelopes (three for each group) to each site which were randomly placed by a third party in the participant files and were opened after the baseline assessment"	
		Comment : insufficient information regarding the method of sequence alloca-	

Exercise interventions for cerebral palsy (Review)

Bryant 2013 (Continued)		
Allocation concealment (selection bias)	Unclear risk	Quote : "[t]he study administrator provided nine sealed envelopes (three for each group) to each site which were randomly placed by a third party in the participant files and were opened after the baseline assessment"
		Comment : unclear if the study administrator was independent to the study or if envelopes were opaque
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Quote : "[o]ne researcher acted as assessor for all but three of the study assess- ments and was blinded as to which arm of the study each participant was allo- cated"
		Comment : assessor may not have been blinded as to which group all of the participants were allocated to
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : low rate of missing data (<10%). Missing data evenly distributed across groups. Missing data accounted for
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment : no other sources of bias identified

lattern-Baxter 2013			
Methods	Design: quasi-randomised controlled trial		
	Country of origin: USA		
	Intervention(s): locomotor treadmill training		
	Unit of allocation: individual		
Participants	Number of participants : 15 randomised. Reported data on 12; exercise group: n = 6; control group: n 6		
	Age: exercise group: mean (SD) = 21.76 (6.50) months; control group: mean (SD) = 21.25 (6.07) months		
	Sex (male/female): 8/4; exercise group: 3/3; control group: 5/1		
	Ethnicity : African-American (n = 2), Asian (n = 2), Hispanic (n = 2), white (n = 6); exercise group: Asian (n = 1), Hispanic (n = 1), white (n = 4); control group: African-American (n = 2), Asian (n = 1), Hispanic (n = 1) white (n = 2)		
	GMFCS level : level I (n = 4) and level II (n = 8); exercise group: level I (n = 2) and level II (n = 4); control group: level I (n = 2) and level II (n = 4)		
	Type of motor abnormality : spastic (n = 7) and hypotonic (n = 5); exercise group: spastic (n = 4) and hypotonic (n = 2); control group: spastic (n = 3) and hypotonic (n = 3)		
	Anatomical distribution of CP : participants with spastic CP had hemiplegia and diplegia; not stated how many in each group.		

Exercise interventions for cerebral palsy (Review)

Mattern-Baxter 2013	
	Inclusion criteria : diagnosis of CP in GMFCS level I or II, aged 9-36 months, signs of walking readiness indicated by both the ability to sit for at least 30 seconds unsupported in ring sitting or W sitting, the ability to take 10 consecutive steps when held on hands or torso
	Exclusion criteria : diagnosis of a genetic syndrome, independent ambulation without an assistive device, previous or current use of treadmill intervention during physiotherapy, use of medication to control spasticity in the past 6 months
Interventions	Aim of the intervention: to improve gross motor skills
	Type of exercise programme: aerobic
	Exercise mode: treadmill
	Comparator : all participants received their weekly scheduled physiotherapy sessions in their homes or in the clinic excluding treadmill training. Content not described, dose not recorded
	Setting: home-based programme
	Intervention provider : parent assisted the participant. The parent was supervised once a week by a physiotherapist
	Duration of programme: 6 weeks
	Exercise dose : 2 sessions per day on 6 days per week. Each session lasted 10-20 min
	Tailoring of intervention to individual : participants used their orthotics and bilateral hand rails to hold on while walking. Parents assisted the child in leg advancement if needed and provided as little manual support as needed at the pelvis. The starting treadmill speed was determined during the initial training session and was increased as quickly as possible throughout the sessions. The speed was determined as the fastest possible speed during which a participant could move his feet independently without dragging them for more than 5 seconds. A range of treadmill speeds was determined at each weekly visit and maintained throughout that week. Participants were encouraged to self-correct before parent intervened
	Fidelity to prescribed intervention : mean completion rate of 87.5% for the 12 weekly training ses- sions. Participants walked an average 28.2 min/d (range 9.6 to 39.3 min/d)
	Monitoring of adverse events : parents were asked about adverse events weekly; no adverse events re- ported.
Outcomes	Assessment time points : baseline (week 0), postintervention (week 6), 4 weeks postintervention (week 10), 12 weeks postintervention (week 18)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	1. Gross motor function was measured using dimensions D and E of the GMFM-66. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. It is not clear if scores are presented as the sum of the scores for each item or as a percentage of total possible score. Higher score indicates better gross motor function.
	 Gross motor function was measured using the locomotion sub scale of the Peabody Developmental Motor Scales-2 (PDMS-2). A higher score indicates better gross motor function.
	3. Walking function was measured with the time taken to complete the 10-metre walk test reported in
	s. It is not stated if this is maximum walking speed or self-selected walking speed.4. Walking function was measured with the Functional Mobility Scale, scale not described, units of measured with the Functional Mobility Scale, scale not described.
	surement not reported 5. Walking function was measured with the number of alternating steps in 10 seconds, scale not de- scribed
	 Gross motor function was measured with the mobility sub scale of the Pediatric Evaluation of Disabil- ity Inventory (PEDI) administered via parent interview. A higher score indicates more independence.

Exercise interventions for cerebral palsy (Review)

Mattern-Baxter 2013 (Continued)

Notes

Source of funding: supported by a research grant from the paediatric section of the American Physical Therapy Association

Potential conflicts of interest: authors report no conflicts of interest; none perceived

Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	High risk	Quote : "[q]uasi-randomization was conducted to achieve matched groups by age and GMFCS levels"	
		Quote : "[t]he children were quasi-randomized by the principal investigators and matched by GMFCS levels and age"	
		Comment : description of method of sequence generation suggests it was based on a non-random component	
Allocation concealment (selection bias)	High risk	Quote : "[t]he children were quasi-randomized by the principal investigators and matched by GMFCS levels and age"	
		Comment : description suggests the principal investigators were aware of group allocation	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation	
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[b]linding for the GMFM-66 and PDMS-2 was achieved by videotaping the children's gross motor skills at their homes. The videotapes were subsequently reviewed by a physical therapist who was blinded to group allocation"	
		Quote : "[t]he timed 10-m walk test (10MWT), the Functional Moblity Scale (FMS), and the number of alternating steps in 10 seconds were used as measures of walking function. They were scored by clinical observation by a non blinded assessor."	
		Quote : "In order to gain the parents' perspective on their child's motor abili- ties, the mobility sub scale of the PEDI was administered via parent interview."	
		Comment : although 1 outcome measure was assessed by an assessor blinded to group allocation, the other measure was not assessed by a blinded assessor. A self-report measure was also used. As parents and assessors were not blinded to group allocation, at least 1 outcome is at high risk of bias.	
Incomplete outcome data (attrition bias)	High risk	Quote : "[a]ttrition (n=3). Illness (n=1). Family reasons (n=1). Change of diagnosis to genetic syndrome (n=1).	
All outcomes		Comment: missing data rate of 20%. All missing data in intervention group	
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes	
Other bias	Unclear risk	Comment : demographic and baseline outcome data only provided for those who completed the trial	

Exercise interventions for cerebral palsy (Review)



Methods	Design: randomised controlled trial				
	Country of origin: Australia				
	Intervention(s): resistance training				
	Unit of allocation: individual				
Participants	Number of participants : 49 randomised; exercise group: n = 24; control group: n = 25 (demographic data reported on n = 48 as 1 person withdrew from the exercise group after allocation but before the start of training)				
	Age : mean (SD) 18 years 1 month (1 year 11 months); e xercise group: mean (SD) = 18 years 2 months (1 year 11 months); control group: mean (SD) = 18 years 7 months (2 years 11 months)				
	Sex (male/female): 26/22; exercise group: 13/10; control group: 13/12				
	Ethnicity: not stated				
	GMFCS level : level II (n = 29) and level III (n = 19); exercise group: level II (n = 13) and level III (n = 10); control group: level II (n = 16) and level III (n = 9)				
	Type of motor abnormality: spastic CP				
	Anatomical distribution of CP: diplegia				
	Inclusion criteria : diagnosis of spastic diplegia, aged 14-22 years, GMFCS level II or III, able to follow simple instructions				
	Exclusion criteria : participated in strength training in previous 6 months, single event multi-level surgery in last 2 years, contractures > 20 degrees at hips and knees				
Interventions	Aim of the intervention: to improve mobility				
	Type of exercise programme: resistance training				
	Exercise mode : weights machine; 4-6 individualised exercises to target deficits identified during gait analysis. Targeted muscles were the knee extensors (7 people), the plantarflexors (4 people), the hip extensors (3 people), the hip abductors (2 people) and generalised extensors represented by the leg press (7 people)				
	Comparator : usual recreation and physiotherapy provided it did not include progressive resistance training				
	Setting: community gym; individually or in pairs				
	Intervention provider: physiotherapist				
	Duration of programme: 12 weeks (24 sessions)				
	Exercise dose : 2 sessions per week, 3 sets of 10-12 repetitions to fatigue (i.e. 60%-80% 1 RM), at least 5 on Borg Rating of Percieved Exertion scale				
	Tailoring of intervention to individual : when the participant was able to complete 3 sets of 12 repeti- tions of an exercise the weight to be lifted was increased				
	Fidelity to prescribed intervention : participants kept logbooks detailing exercise, weight lifted, num ber of repetitions, sets and details of injuries. Mean (SD) sessions performed 21.9 (2.4). The mean (SD) rating of perceived exertion at the end of each session was 6.9 (1.1). Participants increased their training load of exercises for targeted muscles from session 3 to session 24 by a mean (SD) of 183% (23%).				
	Monitoring of adverse events : short-term muscle soreness reported by most participants but resolve in a few days. 1 participant reported minor calf strain, and 1 participant reported minor discomfort to plantar fascia; the programme was adjusted but participants did not miss sessions.				



Taylor 2013 (Continued)

Outcomes

Assessment time points: baseline (week 0), postintervention (week 12), 12 weeks postintervention (week 24)

Primary outcome: mobility

1. Mobility was measured with the 6-min walk test. Participants were instructed to walk as far as they could in 6 min and the distance (in m) was recorded

Secondary outcomes: mobility-related function, gross motor function, muscle performance

- 1. Mobility-related function was assessed with self-selected walking speed over 10 m (reported in m/s)
- 2. Mobility-related function was assessed with a timed stairs test. The time participants took to walk up and down 3 stairs was recorded in seconds
- 3. Gross motor function was assessed with dimensions D and E of the GMFM-66. Possible range of scores 0 to 39 for dimension D and 0 to 72 for dimension E. Scores are presented as a percentage of total possible score for each domain. Higher score indicates better gross motor function.
- 4. Gait was assessed using the Gait Profile Score, measured using motion analysis, which provides an overall measure of gait kinematic deviation from normal in degrees.
- 5. Participant-rated mobility was measured using the Functional Mobility Scale, which describes the level of assistance that participants require to cover different distances and environments. The score at each distance (5 m, 50 m, and 500 m) was reported on a scale of 1-6. Higher score indicates better mobility.
- 6. Participant-rated mobility was measured using the Functional Assessment Questionnaire, a 10-level report of the level that best describes typical walking ability. Higher score indicates better mobility.
- 7. Muscle strength was measured using 1 RM of a leg press and a reverse leg press (reported in kg).
- 8. Isometric muscle strength of the targeted muscles for each participant was measured using a handheld dynamometer. The percentage increase in muscle strength was reported.
- 9. Physical activity was assessed using the ActivPAL activity monitor. Participants were instructed to wear the monitor for 7 consecutive days; number of steps per day and time spent in sitting and lying (reported in hours per day) were reported.

Source of funding: National Health and Medical Research Council of Australia

Potential conflicts of interest: the authors report no conflict of interest; none perceived

Risk of bias

Notes

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote : "[a] separate randomisation procedure was prepared for each stratum (GMFCS levels II and III) using permuted blocks. An independent researcher generated a block allocation sequence for each stratum by drawing pieces of paper from a sealed container and then sealing assignments in sequentially numbered opaque envelopes. The research coordinator allocated participants after enrolment and baseline testing".
Allocation concealment (selection bias)	Low risk	Quote : "[a] separate randomisation procedure was prepared for each stratum (GMFCS levels II and III) using permuted blocks. An independent researcher generated a block allocation sequence for each stratum by drawing pieces of paper from a sealed container and then sealing assignments in sequentially numbered opaque envelopes. The research coordinator allocated participants after enrolment and baseline testing".
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation

Exercise interventions for cerebral palsy (Review)



Taylor 2013 (Continued)		
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[a]ssessments were completed in a hospital gait laboratory by an as- sessor blinded to group allocation".
		Quote : "[i]n addition, two participant-rated mobility outcomes were assessed: the Functional Mobility Scale, which describes the level of assistance that chil- dren with CP required to cover different distances and environments; and the Functional Assessment Questionnaire, a 10-level report of the level that best describes typical walking ability".
		Comment : both self-reported and objective outcome measures were used. As participants were not blinded to group allocation at least 1 outcome is at high risk of bias.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[t]he intention-to-treat principle was applied with available data of all participants who were allocated and commenced their programme included in analyses".
		Quote : "[o]ne participant withdrew from the intervention group after alloca- tion but before the start of training because surgery was scheduled unexpect- edly".
		Comment : low rate of missing data (2%) and intention-to-treat analysis under- taken
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes.
Other bias	Low risk	Comment : no other sources of bias identified

Fedla 2014	
Methods	Design: randomised controlled trial
	Country of origin: Saudi Arabia
	Intervention(s): strength training of trunk and lower extremity muscles
	Unit of allocation: individual
Participants	Number of participants : 62 randomised; exercise group: n = 31, control group: n = 31. Data presented on 60 participants; exercise group: n = 30; control group: n = 30
	Age : mean (SD) = 8.94 (2.46) year; exercise group: mean (SD) = 9.14 (2.46) years; control group: mean (SD) = 8.74 (2.49) years
	Sex (male/female): 40/20
	Ethnicity: not stated
	GMFCS level: presented as bar chart; unable to determine number of participants in each level
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: diplegia
	Inclusion criteria : diagnosis of spastic diplegia, 5-14 years, able to sit for 10 seconds with back unsup ported and feet supported, minimum required score on the mini-mental state examination, GMFCS levels I, II, III or IV, able to move the affected lower limbs at least in gravity eliminated position

Exercise interventions for cerebral palsy (Review)



Tedla 2014 (Continued)	Exclusion criteria : orthopaedic surgery in past year, botulinum toxin injection in past 6 months, history of selective dorsal rhizotomy, receiving medication that alters muscle tone or strength			
Interventions	Aim of the intervention: to increase muscle strength			
	Type of exercise programme: resistance training			
	Exercise mode : muscles with > 50% weakness from the normal were targeted. Exercise group completed a circuit consisting of the following exercises: quadruped position upper extremity and lower extremity lifts for trunk extensors, curl-ups and leg lifts for trunk flexors, trunk and lower extremity combined in wall squats and sit-to-stand with weighted vest, hip flexors, knee extensors and dorsiflexors with sand bags in high sitting position, hip extensors and knee flexors with sand bags in prone positions, hip adductors with sand bags in side lying position, heel rises on a step for plantar flexors with weighted vest			
	Comparator : 3-5 sessions a week of conventional physiotherapy consisting of range of motion exercises and stretching, positioning or adaptive equipment prescription, movement transitions and mobility training, functional activities and gait training. Average duration per session was 60-90 min			
	Setting: not stated; supervised programme			
	Intervention provider: the investigator			
	Duration of programme: 6 weeks			
	Exercise dose : 3 sessions per week for a total of 18 sessions. All exercises performed at 80% of 1 repetition maximum (RM). The number of repetitions varied between exercises and individuals but ranging between 6 and 10 for 3 sets in the initial periods. Conventional physiotherapy continued as usual 1-2 days/week Tailoring of intervention to individual : when the participant reached 3 sets of 12 repetitions without any difficulty or according to the new 80% of 1 RM at each week, the participant progressed to higher weights. With changed weight, the exercise repetitions were started again with 6-10 in 3 sets			
	Fidelity to prescribed intervention : parent/caregiver help was taken to maintain a log book. This recorded the type of exercise, the number of repetitions, the amount of weight lifted and any addition-al information; adherence not reported			
	Monitoring of adverse events : muscle soreness was reported in a few subjects but subsided with rest and no additional treatment was required			
Outcomes	Assessment time points: baseline (week 0), postintervention (week 6)			
	Primary outcome: no primary outcome measure stated			
	Outcomes:			
	 Muscle strength of the trunk flexors and extensors, hip flexors, extensors, abductors and adduc- tors, knee flexors and extensors, and dorsiflexors and plantarflexors was measured by handheld dy- namometry. The best of 3 trials was used in analysis. Reported in lb 			
	 Gross motor function was assessed using the GMFM. Score for each dimension reported separately. Possible range on each dimension 0 to 100 (higher score indicates better gross motor function) Balance was assessed using the Pediatric Balance Scale. Unclear what units the score reported in 			
Notes	Source of funding: not stated			
	Potential conflicts of interest: not stated; none perceived			
Risk of bias				
Bias	Authors' judgement Support for judgement			

Exercise interventions for cerebral palsy (Review)

Tedla 2014 (Continued)

Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[i]n this randomized controlled trial subjects were randomly allocated either to 6 wk strengthening experimental group or to a conventional intervention control group".
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment (selection bias)	Unclear risk	Comment : insufficient information regarding allocation concealment was pro- vided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "[b]efore and after the training the assessments were conducted by the same investigator".
		Quote : "[i]n addition, the investigator did not review or have access to the pre- training values at the post-training assessments."
		Quote : "[i]deally the person conducting the assessment should have been blinded to the training schedule. Because limited manpower and long duration there was difficulty in blinding the tester."
		Comment : although not stated, text suggests assessor was not blinded to group allocation.
Incomplete outcome data	Low risk	Quote: "[d]rop out due to absence (No-1)"
(attrition bias) All outcomes		Quote : "[d]rop out due to surgery (No-1)"
		Comment : figure 2 indicates that 1 person dropped out from each group. This represents a low rate of missing data (3%) that is evenly distributed across groups.
Selective reporting (re- porting bias)	High risk	Comment : data on strength and gross motor function measure are reported incompletely and cannot be entered into a meta-analysis.
Other bias	Low risk	Comment: no other sources of bias identified

Methods	Design: randomised controlled trial
	Country of origin: Saudi Arabia
	Intervention(s): gait training programme using antigravity treadmill
	Unit of allocation: individual
Participants	Number of participants : 36 randomised; exercise group: n = 17; control group: n = 17
	30 completed the trial (exercise group: n = 15; control group: n = 15). Unclear if baseline data provided on 34 participants or 30 participants
	Age : 6-8 years; exercise group: mean (SD) = 6.799 (0.77) years; control group: mean (SD) = 6.402 (0.68) years.

Exercise interventions for cerebral palsy (Review)

Emara 2015 (Continued)	Sex (male/female): 18/12; exercise group: 9/6; control group: 9/6
	Ethnicity: not stated
	GMFCS level: not stated
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: diplegia
	Inclusion criteria : spasticity grades ranged from 1 to 1+ according to the modified Ashworth scale, no hearing defects, no fixed deformity of both lower limbs, absence of cognitive or visual impairment that could compromise the performance of the tasks
	Exclusion criteria: not stated
Interventions	Aim of the intervention: to improve balance
	Type of exercise programme: aerobic exercise
	Exercise mode : gait training using the Alter G anti-gravity treadmill in addition to the same physical therapy programme provided to the control group
	Comparator : physical therapy programme of 1 hour session, 3 times per week, based on neurodevel- opmental approach, including facilitation, strengthening exercises, gait training in parallel bars, and negotiating obstacles
	Setting : participant's home. Unsupervised individual session. Antigravity treadmill training was pro- vided in an outpatient department
	Intervention provider: not stated
	Duration of programme: 12 weeks
	Exercise dose : 20 min of walking on anti-gravity treadmill, 3 times per week, at a comfortable walking speed
	Tailoring of intervention to individual : a comfortable treadmill speed was selected for all partici- pants as 75% of their comfortable speed during over-ground walking
	Fidelity to prescribed intervention: not reported
	Monitoring of adverse events: not reported
Outcomes	Assessment time points: baseline (week 0), postintervention (week 12)
	Primary outcome: no primary outcome measure stated
	Outcomes:
	1. Spasticity was assessed using the Modified Ashworth Scale. The degree of spasticity was evaluated by passive movement for both limbs while the child was completely relaxed and lying supine on a mat with the head in mid position. The mean score of 3 tests was used in analysis
	2. Dynamic postural control was assessed by calculating an overall stability index, anteroposterior sta- bility index, and mediolateral stability index using the Biodex
Notes	Source of funding: not stated
	Potential conflicts of interest: authors reported no potential conflicts of interest
Risk of bias	
Bias	Authors' judgement Support for judgement

Exercise interventions for cerebral palsy (Review)

Emara 2015 (Continued)		
Random sequence genera- tion (selection bias)	Unclear risk	Quote : "[t]hey were divided randomly into two groups"
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement
Allocation concealment	Unclear risk	Quote : "[t]hey were divided randomly into two groups".
(selection bias)		Comment : insufficient information regarding the method of allocation con- cealment was provided to make a judgement
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Comment : did not provide information to indicate if the assessor was blinded to group allocation or not
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : attrition and withdrawal accounted for. Low rate of missing data (< 10%) and evenly distributed across groups
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Unclear risk	Comment : unclear whether all demographic or outcome data for all participants are reported

Lee 2015	
Methods	Design: randomised controlled trial
	Country of origin: Korea
	Intervention(s): resistance training programme
	Unit of allocation: individual
Participants	Number of participants : 26 randomised; exercise group: n = 13; control group: n = 13
	Group characteristics reported for all randomised participants
	Age : 5-10 years; exercise group: mean (SD) = 6.1 (2.7) years; control group: mean (SD) = 6.9 (2.5) years
	Sex (male/female): 13/13; exercise group: 8/5; control group: 5/8
	Ethnicity: not stated
	GMFCS level: levels I to III
	Type of motor abnormality: spastic CP
	Anatomical distribution of CP: not stated
	Inclusion criteria : aged 5-10 years, the ability to follow verbal instructions, able to walk independently indoors with or without a walking aid (GMFCS level I-III)

Exercise interventions for cerebral palsy (Review)



Lee 2015 (Continued)		stable seizures, any treatment for spasticity or surgical procedures in the last in medication expected during the study period, any other disease that would nysical activity	
Interventions	Aim of the intervention	on : to change lower limb muscle architecture and motor function	
	Type of exercise programme: resistance training		
	Exercise mode : 3 functional training items: loaded sit-to-stand for 5 min, loaded lateral step-up and half knee-rise for 10 min, unloaded lateral step-up and half knee-rise for 10 min		
	Comparator : all partic sions a week for 6 weel	ipants received 30 min of general neurodevelopmental treatment (NDT), 3 ses- <s< td=""></s<>	
	Setting: not stated		
	Intervention provider	r: not stated	
	Duration of programm	ne: 6 weeks	
	Exercise dose: 3 session	ons per week	
		ion to individual : the training load was initially set to 5% of participants' body ely increased base on repeated estimation of the 8 RM	
	Fidelity to prescribed intervention: not reported		
	Monitoring of adverse events: not reported		
Outcomes	Assessment time points: baseline (week 0), postintervention (week 6)		
	Primary outcome: no primary outcome measure stated		
	Outcomes:		
	 The right side muscle thickness of the quadriceps femoris was assessed using ultrasonography (US); the average of 3 measures was used in analysis. 		
	2. The cross-sectional area of rectus femoris was assessed using US; the average of 3 measures was used in analysis.		
	3. The muscle thickness had pennation angle of gastrocnemius was assessed using US; the average of 3 measures was used in analysis.		
	4. Gross motor function measured using the Gross Motor Function Measures-88; possible range of scores		
	0 to 100 (higher score indicates better gross motor function). 5. Mobility limitations were assessed using the parent-reported questionnaire, the MobQue.		
Notes	Source of funding: not stated		
	Potential conflicts of interest: not stated; none perceived		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Quote: "[t]his study had a single-blind randomized controlled design"	
		Comment : insufficient information regarding the method of sequence alloca- tion was provided to make a judgement	
Allocation concealment	Unclear risk	Quote : "[t]his study had a single-blind randomized controlled design"	
(selection bias)		Comment : insufficient information regarding the method of allocation con- cealment was provided to make a judgement	

Exercise interventions for cerebral palsy (Review)

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Lee 2015 (Continued)

Library

Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Quote : "[t]his study had a single-blind randomized controlled design" Comment : did not provide information to indicate if the assessor was blinded to group allocation or not
Incomplete outcome data (attrition bias) All outcomes	Low risk	Comment : no missing outcome data
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is not available for this study and therefore unable to determine if all expected outcomes are reported; no convincing text provided to indicate that published report includes all expected outcomes
Other bias	Low risk	Comment: no other sources of bias identified

Mitchell 2016

Methods	Design: randomised controlled trial		
	Country of origin: Australia		
	Intervention(s) : web-based multimodal therapy programme that included upper limb, cognitive, visu- al perceptual, and physical activity training		
	Unit of allocation: individual		
Participants	Number of participants : 102 participants randomised. Baseline assessments conducted on 101 partic- ipants. Exercise group: n = 51; control group: n = 50		
	Group characteristics reported for 101 participants		
	Age : mean (SD) 11 years 9 months (2 years 4 months), range 8 to 17 years; exercise group: mean (SD) = 11 years 3 months (2 years 4 months); control group: mean (SD) = 11 years 4 months (2 years 6 months)		
	Sex (male/female): 52/49; exercise group: 26/25; control group: 26/24		
	Ethnicity: not stated		
	GMFCS level : level I (n = 45), level II (n = 56); exercise group: level I (n = 20), level II (n = 31); control group: level I (n = 25), level II (n = 25)		
	MACS level : level I (n = 24), level II (n = 76), level III (n = 1); exercise group: level I (n = 11), level II (n = 39), level III (n = 1); control group: level I (n = 13), level II (n = 37), level III (n = 0)		
	Type of motor abnormality: spastic CP		
	Anatomical distribution of CP: unilateral		
	Inclusion criteria : aged 8-18 years, unilateral CP, classified in GMFCS levels I and II, classified in MACS levels I to III, sufficient cooperation and cognitive understanding to perform required tasks. Computer and internet access were provided at home if participants did not have access on entry to the study		



Trusted evidence. Informed decisions. Better health.

Mitchell 2016 (Continued)	Exclusion criteria : unstable epilepsy or medical conditions that precluded participation in training. Entry was delayed to trial if the participant had undergone upper-limb botulinum neurotoxin A injec- tions or surgery in the previous 2 months or 6 months, respectively			
Interventions	Aim of the intervention : to improve activity capacity and performance, reduce mobility limitations, improve recreational participation, improve occupational performance, upper limb function, and visual perception			
	Type of exercise programme: resistance training			
	Exercise mode : physical activity games were interspersed with upper-limb and visual-perceptual games. Gross-motor exercises comprised of 40% of the overall programme and included sequences of repetitive multi-joint bodyweight functional exercises (e.g. sit-to-stand, alternate lunging, step-ups, side step-ups onto a block, squatting, balancing on balance foam)			
	Comparator : participants in the wait list control group continued care as usual for the duration of the programme			
	12 (23%) participants in the exercise group received physiotherapy totaling on average 0.47 hours over the 20 weeks. 15 (30%) participants in the control group received physiotherapy totaling on average 3.9 hours over the 20 weeks			
	Setting : participant's home. Unsupervised individual session. Therapists were available to participants and their families via email, telephone or videoconferencing to provide encouragement and technical support.			
	Intervention provider: web-based programme			
	Duration of programme: 20 weeks			
	Exercise dose : 30-min programme, 6 days per week (total potential dose of 60 hours)			
	Tailoring of intervention to individual : the intensity of the lower-limb strength exercises for week 1 were determined by setting tasks for approximately 75% of repetition maximum. These were incre- mented weekly by the physiotherapists remotely, by increasing the repetitions, speed, step height, and balance challenge in response to individual performance and feedback from the participants and their parents or caregivers.On average, week 1 started with 7 activities of 5-10 repetitions, lasting approximately 60 seconds per activity, and progressed to 11 games of up to 20 repetitions lasting approximately 90 seconds with the addition of step blocks and balance foam.			
	Fidelity to prescribed intervention : on average participants completed mean (SD) 32.4 (17.2) hours of training over the 20-week period, logging in for 24.2 (5.5) min on 77.7 (35.7) days. The total dose of therapy ranged from 3.7 to 74.7 hours per participant.			
	Monitoring of adverse events : 2 instances of minor musculoskeletal pain were recorded during the in- tervention; participants were instructed to modify the movement and continued with the programme. 1 participant in the intervention group had seizures but these were deemed unrelated to training			
Outcomes	Assessment time points: baseline (week 0), postintervention (week 22)			
	Primary outcomes:			
	 Activity capacity was assessed by recording the maximal repetitions of sit-to-stand, lateral step-up using a 20 cm step, and half-kneel to standing for the dominant and non-dominant legs over a 30 s period. A composite score was created by summing repetitions from each task. 			
	2. Walking endurance was measured using the 6-min walk test. The distance participants walked for 6 min along a flat straight 10 m corridor, was recorded.			
	 Activity performance was assessed using an ActiGraph GT3X+ tri-axial accelerometer. Time in seden- tary, light, moderate, and vigorous activity was reported. 			
	 ADL motor and processing skills were assessed using the Assessment of Motor and Process Skills (AM- PS). 			
	5. Impaired hand use in bimanual tasks was assessed using the Assisting Hand Assessment (AHA).			

Exercise interventions for cerebral palsy (Review)



Mitchell 2016 (Continued)	
	6. Upper limb unimanual speed and dexterity were assessed using the Jebsen-Taylor Test of Hand Func- tion (JTTHF).
	7. Quality of reach, grasp, release, and manipulation of the impaired upper limb was assessed using the Melbourne Assessment of Unilateral Upper Limb Function (MUUL).
	Secondary outcomes:
	1. Mobility limitations were assessed using a parent-reported questionnaire (28-item Mobility Question- naire). Possible range 0 to 100.
	2. Participation was assessed using the Assessment of Life Habits (Life-H). Possible range 0 to 10 (higher score indicates higher participation).
	3. Self-perceived occupational performance was assessed using the Canadian Occupational Perfor- mance Measure (COPM).
	4. Visual perception was assessed using the Test of Visual Perceptual Skill (non-motor) 3rd edition (TVPS-3).
Notes	Source of funding : financial support was obtained from Queensland Government Co-Investment Pro- gram Grant "EBrain", a Financial Markets Foundation for Children research grant, an Australian Post- graduate Award, a National Health and Medical Research Council Career Development Fellowship, and a Smart State Fellowship.
	Potential conflicts of interest : the authors stated that they had no interests that might be perceived as posing a conflict or bias.
Risk of bias	

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Quote (from trial protocol) : "[c]hildren will be matched in pairs according to age, gender and level of functional ability based on MACS level The randomisation process will involve randomly allocating a number '1' or '2' to each member of the pair. As each pair is entered, they will be allocated to the next consecutive envelope, which will be opened by the non-study personnel who will read and record the treatment allocation from the paper inside the envelope. Treatment allocation will be recorded on a piece of folded paper inside each envelope, in random order".
Allocation concealment (selection bias)	Low risk	Quote (from trial protocol) : "[c]hildren will be matched in pairs according to age, gender and level of functional ability based on MACS level The ran- domisation process will involve randomly allocating a number '1' or '2' to each member of the pair. As each pair is entered, they will be allocated to the next consecutive envelope, which will be opened by the non-study personnel who will read and record the treatment allocation from the paper inside the envelope. Treatment allocation will be recorded on a piece of folded paper inside each envelope, in random order".
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Comment : given the nature of the intervention participants were not blinded to treatment allocation
Blinding of outcome as- sessment (detection bias) All outcomes	High risk	Quote : "it should be noted that while the study's primary outcome (the Assessment of Motor and Process Skills) was blinded, study personnel were not able to be blinded to group allocation for other outcomes measures".
		Comment : the outcomes included in the results of this review were assessed by an unblinded assessor

Exercise interventions for cerebral palsy (Review)

Mitchell 2016 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Low risk	Quote : "[i]n the intervention group, 47 started training and were assessed at 20 weeks (92% retention); in the waitlist control group, 44 were assessed at 20 weeks (86% retention)" Quote : "[d]ata were analysed according to the intention-to-treat principle"
Selective reporting (re- porting bias)	Unclear risk	Comment : a protocol is available for the trial. The majority of outcomes are reported in the 2 included reports. However, some outcomes are not reported to date e.g. quality of life, participation assessed with the Participation and Environment Measure for Children and Youth.
Other bias	Low risk	Comment: no other sources of bias identified

bpm: beats per minute; **CP**: cerebral palsy; **HR**: heart rate; **GMFCS**: Gross Motor Function Classification System; **GMFM**: Gross Motor Function Measure; **IQ**: intelligence quotient; **RER**: respiratory exchange ratio; **rpm**: rotations per minute; **RM**: repetition maximum; **SD**: standard deviation; **US**: ultrasound; **VO**₂ : volume of oxygen.

Characteristics of excluded studies [author-defined order]

Study	Reason for exclusion
Bar-or 1976	Not a randomised controlled trial
Hutzler 1998	Not a randomised controlled trial
Jones 2001	Not a randomised controlled trial
Ketelaar 2001	Intervention did not meet pre-stated definition of exercise
Benda 2003	Intervention did not meet the pre-stated definition of exercise
Blundell 2003	Not a randomised controlled trial
Cherng 2004	Intervention did not meet pre-stated definition of exercise
Ahlborg 2006	Comparison was whole body vibration therapy
Kandrali 2006	Intervention did not meet the pre-stated definition of exercise
Patikas 2006	Intervention was provided following surgery; unable to determine independent effects of exercise
Reid 2006	Intervention did not meet the pre-stated definition of exercise
Dodd 2007	Not a randomised controlled trial
Gordon 2007	Intervention did not meet the pre-stated definition of exercise
Ozer 2007	Intervention did not meet the pre-stated definition of exercise
Stackhouse 2007	Compared strengthening with neuromuscular electrical stimulation to strengthening without neu- romuscular electrical stimulation
Weindling 2007	Intervention did not meet the pre-stated definition of exercise
Bar-Haim 2008	Intervention did not meet the pre-stated definition of exercise

Exercise interventions for cerebral palsy (Review)



Study	Reason for exclusion
Fernandes 2008	Intervention did not meet the pre-stated definition of exercise
Hornyak 2008	This is a commentary
Jannink 2008	Intervention did not meet the pre-stated definition of exercise
Davis 2009	Intervention did not meet pre-stated definition of exercise
Katz-Leurer 2009	Included children with cerebral palsy and traumatic brain injury and presented data on combined group
McGibbon 2009	Not a randomised controlled trial
Taylor 2009	This is a correspondence not a study
Salem 2009	Intervention did not meet the pre-stated definition of exercise
Batista 2010	Not a randomised controlled trial
Boyd 2010	Protocol of constraint-induced movement therapy compared to bimanual therapy (do not meet pre-stated definition of exercise)
Brown 2010	Not a randomised controlled trial
Harbourne 2010	Compared different interventions (not exercise interventions) and different means of delivery.
Herrero 2010	Intervention did not meet the pre-stated definition of exercise
Kumar 2010	Not a randomised controlled trial
Maher 2010	Intervention did not meet the pre-stated definition of exercise
Mehta 2010	Compared Bobath therapy to conventional therapy. Neither intervention met the pre-stated defini- tion of exercise.
Sorsdahl 2010	Not a randomised controlled trial
Speyer 2010	Participants did not have a diagnosis of cerebral palsy
Willoughby 2010	Compared two modes of aerobic exercise
Yonetsu 2010	Intervention did not meet the pre-stated definition of exercise
Batra 2011	Intervention did not meet the pre-stated definition of exercise
Choi 2011	Intervention did not meet pre-stated definition of exercise
Hung 2011	Intervention did not meet the pre-stated definition of exercise
Roberti 2011	This is a literature review not a study
Sakzewski 2011a	Intervention did not meet the pre-stated definition of exercise
Sakzewski 2011b	Intervention did not meet the pre-stated definition of exercise. It compared constraint-induced movement therapy to bimanual training.

Exercise interventions for cerebral palsy (Review)



Study	Reason for exclusion
Silva e Borges 2011	Intervention did not meet the pre-stated definition of exercise
Yabunaka 2011	Intervention did not meet the pre-stated definition of exercise
Bandholm 2012	Compared resistance training and usual care following botulinum toxin treatment; unable to differ- entiate effects of exercise and botulinum toxin treatment
Dimitrijević 2012	Intervention did not meet pre-stated definition of exercise
Herrero 2012	Intervention did not meet the pre-stated definition of exercise
Kang 2012	Intervention did not meet the pre-stated definition of exercise
Kim 2012	Compared 2 modes of resistance training
Nsenga Leunkeu 2012	Participants not randomly allocated into groups
Olama 2012	Compared resistance training to resistance training and myofeedback
Sakzewski 2012	Intervention did not meet the pre-stated definition of exercise
Salem 2012	Participants did not have a diagnosis of cerebral palsy
Angulo-Barroso 2013	Participants did not have a diagnosis of cerebral palsy
Chang 2013	Not a randomised controlled trial
Fedrizzi 2013	Intervention included constraint-induced movement therapy
Grecco 2013a	Compared two modes of aerobic exercise
Grecco 2013b	Compared two modes of aerobic exercise.
Green 2013	Not a randomised controlled trial
Hutzler 2013	Not a randomised controlled trial
Jeng 2013	Not a randomised controlled trial
Kumar 2013	Unable to determine the content of the intervention from the published report
Moreau 2013	Compared 2 types of strength training (fast and slow training)
Nsenga Leunkeu 2013	Participants not randomly allocated into groups
Rimmer 2013	Intervention did not meet the pre-stated definition of exercise
Su 2013	Compared two modes of aerobic exercise
Williams 2013	The participants in the intervention group received botulinum toxin type A treatment before or af- ter strength training (this was not consistent for participants). Unable to determine independent effects of exercise
Abd El-Kafy 2014	Intervention did not meet the pre-stated definition of exercise

Exercise interventions for cerebral palsy (Review)



Study	Reason for exclusion
Ayhan 2014	Participants did not have a diagnosis of cerebral palsy
Chiu 2014	Intervention did not meet pre-stated definition of exercise
Franki 2014	Intervention did not meet the pre-stated definition of exercise
Hammond 2014	Participants did not have a diagnosis of cerebral palsy
Hussein 2014	Compared 2 modes of aerobic exercise
Lee 2014a	Intervention did not meet the pre-stated definition of exercise
Lee 2014b	Not a randomised controlled trial
Park 2014a	Intervention did not meet pre-stated definition of exercise
Schroeder 2014	Not a randomised controlled trial
Slaman 2014	Intervention group received 3 interventions; only 1 was exercise. Unable to determine independent effects of exercise
Van Wely 2014	Intervention group did not receive exercise alone; unable to determine independent effects of exer- cise
Verschuren 2014	This is a literature review not a study
Walsh 2014	Not a randomised controlled trial but a case study
Williams 2014	Participants did not have a diagnosis of cerebral palsy
AlSaif 2015	Intervention did not meet the pre-stated definition of exercise
Bohm 2015	Intervention did not meet the pre-stated definition of exercise
Capio 2015	Not a randomised controlled trial
Chen 2015	Not a randomised controlled trial
El-Basatiny 2015	Intervention did not meet pre-stated definition of exercise
Gillaux 2015	Intervention did not meet the pre-stated definition of exercise
Hamah 2015	Intervention did not meet the pre-stated definition of exercise
Kim 2015	Not a randomised controlled trial
Lai 2015	Not a randomised controlled trial
Lowe 2015	Participants do not have a diagnosis of cerebral palsy
Preston 2015	Participants in intervention group received botulinum toxin and exercise; unable to determine in- dependent effects of exercise
Sherief AEAA 2015	Participants in intervention group received treadmill training and dynamic ankle-foot orthoses; un- able to determine independent effects of exercise

Exercise interventions for cerebral palsy (Review)



Study	Reason for exclusion
Swe 2015	Compared 2 modes of aerobic exercise
Temcharoensuk 2015	Intervention did not meet the pre-stated definition of exercise
Declerck 2016	Intervention did not meet the pre-stated definition of exercise
Hsieh 2016	Participants did not have a diagnosis of cerebral palsy

Characteristics of studies awaiting assessment [author-defined order]

Carlon 2014

Methods	Design: randomised controlled trial
	Country of origin: Australia
	Intervention(s): aerobic training programme
	Unit of allocation: individual
Participants	Number of participants: 19
	Age: mean 13 years 10 months
	Sex: not stated
	Ethnicity: not stated
	GMFCS level : level I ($n = 4$), level II ($n = 9$) and level III ($n = 6$)
	Type of motor abnormality: not stated
	Anatomical distribution of CP: not stated
	Inclusion criteria : aged 8-18 years, diagnosis of CP, classified in GMFCS level I, II or III, attending 1 of 3 specialist schools, reliable yes/no response
	Exclusion criteria : surgery or botulinum toxin A to the lower-limbs, or aerobic training, in the pre- vious 6 months
Interventions	Aim of the intervention: not stated
	Type of exercise programme: aerobic training
	Exercise mode: individualised programme that considered student activity preference
	Comparator: arts programme of same duration
	Setting: school; supervised in ratio of 1:1 or 1:2
	Intervention provider: not stated
	Duration of programme: 9 weeks
	Exercise dose : 3 weekly sessions of 30 min each
	Tailoring of intervention to individual : individualised programme that considered student activi- ty preference

Exercise interventions for cerebral palsy (Review)



Carlon 2014 (Continued)	Fidelity to prescribed intervention : participants attended, on average, 79% of sessions. Across the intervention there was 80% adherence to target heart rate (target heart rate not stated)
	Monitoring of adverse events : 3 non-serious adverse events in 2 participants which were expected and related to the intervention. 1 participant hit her leg on a bike and 1 participant tripped on 2 separate instances with no adverse consequences.
Outcomes	Assessment time points: unclear; baseline (week 0) and week 9
	Primary outcomes: feasibility and safety
	1. Feasibility was assessed by programme attendance and adherence to training heart rate targets measured by heart rate monitors.
	2. Safety was measured by incidence of adverse events, recorded as serious or non-serious, expect- ed or unexpected, and related or unrelated.
	Secondary outcomes: cardiovascular function and physical activity
	1. Cardiovascular function was measured using the 6-min walk test and the Muscle Power Sprint Test.
	2. Physical activity was measured using RT3 activity monitors.
Notes	Source of funding: Dr WM Phelps Foundation
	Potential conflicts of interest: authors declare no conflicts of interest.

CP: cerebral palsy; **GMFCS**: Gross Motor Function Classification System.

Characteristics of ongoing studies [author-defined order]

Gillett 2015

Trial name or title	FAST CP
Methods	Design: randomised controlled trial
	Country of origin: Australia
	Intervention(s) : mixed training (lower limb resistance exercises and functional anaerobic exercises)
	Unit of allocation: individual
Participants	Number of participants: 40
	Inclusion criteria : aged 15 to 30 years, confirmed diagnosis of spastic hemiplegia or diplegia-type CP, able to walk independently, GMFCS level I or II, maximum passive ankle dorsiflexion range of motion of < 5° (knee fully extended)
	Exclusion criteria : lower limb surgery in the past 2 years and/or botulinum toxin-A injections to the lower extremities within the past 6 months, unable to provide sufficient cooperation and cognitive understanding to participate in the intervention, participated in lower limb resistance training within the past 6 months
Interventions	Aim of the intervention: to alter skeletal muscle properties and improve muscle function
	Type of exercise programme : mixed training (lower limb resistance exercises and functional anaerobic exercises).
	Exercise mode : 5 resistance exercises: seated knee calf raise, leg press, seated straight knee calf press, tibialis anterior raise, standing calf raise. Functional anaerobic exercises including step-ups,

Exercise interventions for cerebral palsy (Review)



Gillett 2015 (Continued)

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	beanbag run, lateral step-ups, 5 m sprint, obstacle course, shuttle sprint, up and down stairs, agili- ty run, 4 cone run
	Comparator: no training; allowed to continue with usual activities
	Setting: tertiary institution gymnasium
	Intervention provider: 2 experienced trainers with tertiary qualifications in the field of exercise science
	Duration of programme: 12 weeks
	Exercise dose : 3 weekly sessions of 75 min each (total dosage 45 hours)
	Tailoring of intervention to individual : initial 12 RM load will be determined in the first week of training for each participant. Load will be added if the participant can complete more than the required number of repetitions in all sets of that exercise (i.e. not exercising to fatigue)
	Fidelity to prescribed intervention and monitoring of adverse events : each participant will complete a training diary to document their progress report injuries and fatigue levels. Any injuries or adverse outcomes attributable to the intervention or testing protocol will be documented and reported. Participants will report any leg pain, lower leg stiffness and level of leg fatigue prior to each training session using a visual analogue scale (0-100 mm).
Outcomes	Assessment time points : baseline (week 0), immediately postintervention (week 12), 12 weeks postintervention (week 24)
	Primary outcomes:
	 Muscle volume of the medial gastrocnemius, lateral gastrocnemius, soleus, and tibialis anterior muscles assessed using MRI
	2. Neuromuscular properties (i.e. passive mechanical, active mechanical, and neural properties) of the medial gastrocnemius muscle
	Secondary outcomes:
	1. Intramuscular fat content of the medial gastrocnemius, lateral gastrocnemius, soleus, and tibialis anterior muscles assessed using MRI
	2. Isometric muscle strength of the plantarflexor and dorsiflexor muscles assessed using a isokinetic dynamometer
	3. Anaerobic power assessed using the Muscle Power Sprint Test
	 Agility assessed using the time to complete 10 × 5 m sprints Functional strength assessed using the 30 s RM test i.e. the sum of the total number of repetitions
	completed in 30 s for a lateral step-up, sit-to-stand, and stand from half kneel
	6. Walking ability assessed using the maximum distance completed during the 6-min walk test
	 In vivo muscle mechanics during walking (i.e. muscle fascicle behaviour during walking, muscle activation and timing of the medial gastrocnemius, lateral gastrocnemius, soleus, and tibialis an- terior)
	8. Participation assessed using the Assessments of Life Habits (Life-H) questionnaire
Starting date	23 January 2015
Contact information	Jarred Gillett; email: j.gillett1@uq.edu.au
Notes	Source of funding : the National Health and Medical Research Council of Australia Postgraduate Scholarship and Australian Rotary Health/Rotary Club of St Ives Funding Partner Scholarship
	Potential conflicts of interest: authors declare no conflicts of interest.

Exercise interventions for cerebral palsy (Review)

ISRCTN90378161

Trial name or title	Feasibility and efficacy of resistance training in cerebral palsy (CP)
Methods	Design: randomised controlled trial
	Country of origin: UK
	Intervention(s): resistance training compared to usual care
	Unit of allocation: individual
Participants	Number of participants: 60
	Inclusion criteria : adolescents with CP aged 10-19 years (amended from 12-19 years on 12/07/2016), the ability to walk independently with or without a mobility aid, the ability to activate the ankle plantarflexors
	Exclusion criteria : lower limb orthopaedic surgery in the past year, botulinum toxin type A (Botox) injections or serial casting in the past 6 months, receiving intrathecal baclofen, unable to comply with the protocol
Interventions	Aim of the intervention: not stated
	Type of exercise programme: progressive resistance training
	Exercise mode: single-joint plantarflexor exercises
	Comparator : all participants will be instructed to continue their usual physiotherapy programme and usual activities
	Setting: local physiotherapy department or gym and participants' homes
	Intervention provider: physiotherapist
	Duration of programme: 10 weeks
	Exercise dose : 30 sessions; 10 sessions in a class and 20 sessions at home. 4 sets of 8-12 repetitions to fatigue per session.
	Tailoring of intervention to individual : the intensity of the exercise will be based on individual strength capacity to ensure that the participant can perform 8 to 12 repetitions to fatigue
	Fidelity to prescribed intervention and monitoring of adverse events: not reported
Outcomes	Assessment time points : baseline (week 0), immediately postintervention (week 10), and 12 weeks postintervention (week 22)
	Primary outcomes:
	1. Gait efficiency measured using indirect calorimetry
	Secondary outcomes:
	 Physical activity measured using accelerometry Participation measured using Assessment of Life Habits questionnaire (Life-H). Gross Motor Function measured using components D and E of Gross Motor Function Measure (GMFM) and gait speed during 10-m walking trial Muscle strength measured using isokinetic dynamometry. Muscle activity during dynamometry and treadmill walking measured using EMG Muscle and tendon force measured using isokinetic dynamometry and ultrasonography Muscle and tendon length measured using ultrasonography and motion analysis Muscle and tendon stiffness measured using ultrasonography and motion analysis

Exercise interventions for cerebral palsy (Review)



ISRCTN90378161 (Continued)

9. Muscle, tendon and fascicle strain measured using ultrasonography
10.Muscle and tendon cross-sectional area measured using ultrasonography
11.Quality of life measured using EQ-5D-Y and CHU 9D
August 2015

Starting date	August 2015
Contact information	Jennifer.Ryan@brunel.ac.uk
Notes	Source of funding : Action Medical Research and Chartered Society of Physiotherapy Charitable Trust
	Potential conflicts of interest: not reported

RBR-5rh6cg

Trial name or title	Aquatic physical therapy in the trunk control in children with cerebral palsy: randomized clinical trial							
Methods	Design: randomised controlled trial							
	Country of origin: Brazil							
	Intervention(s): resistance training versus aerobic							
	Unit of allocation: individual							
Participants	Number of participants: 24							
	Inclusion criteria : clinical diagnosis of spastic diplegic CP; level IV Gross Motor Function Classifica- tion System (GMFCS); aged 4 years to 10 years and 11 months							
	Exclusion criteria : uncooperative patients; unable to understand the proposed activities; under- going orthopedic surgery or peripheral blocks less than 6 months previously							
Interventions	Aim of the intervention: not stated							
	Type of exercise programme : aquatic exercises emphasising trunk control versus conventional aquatic therapy							
	Exercise mode: trunk control exercises							
	Comparator: conventional aquatic therapy							
	Setting: not reported							
	Intervention provider: not reported							
	Duration of programme: 8 weeks							
	Exercise dose : 16 sessions; 35 min twice a week							
	Tailoring of intervention to individual: not reported							
	Fidelity to prescribed intervention and monitoring of adverse events: not reported							
Outcomes	Assessment time points: pre- and postintervention measures							
	Primary outcomes:							
	 Trunk control measurement scale; improved control considered as at least 5% variation between pre- and postintervention measures 							

Exercise interventions for cerebral palsy (Review)

RBR-5rh6cg (Continued)

Secondary outcomes:

	1. Gross Motor Function Measure (GMFM-88)
	2. Surface electromyography
	3. Visual Analogue Scale (VAS) trunk
	4. Flexometer Wells
	5. Child Health Questionnaire (CHQ, PF-50)
Starting date	November 2015
Contact information	Mirna Sayuri Kanashiro email: mitie_kakihata@hotmail.com
Notes	Source of funding: institution: Associação de Assistência à Criança Deficiente
	Potential conflicts of interest: not reported

NCT02766491							
Trial name or title	Improving Stretching Interventions for Children with CP						
Methods	Design: randomised controlled trial						
	Country of origin: UK						
	Intervention(s): resistance training (stretching and strengthening exercises for the calf muscles)						
	Unit of allocation: individual						
Participants	Number of participants: 30						
	Inclusion criteria : diagnosis of spastic CP, GMFCS level I-III, able to perform at least 1 bilateral heel raise, aged 7-14 years						
	Exclusion criteria : orthopaedic or neural surgery to the lower limb 2 years prior to or planned dur- ing the intervention, botulinum toxin A injections 6 months prior to or planned during the interven- tion, a learning or behaviour impairment that prevents full participation in the intervention						
Interventions	Aim of the intervention: to stiffen the tendon and increase the amount of stretch in the muscle						
	Type of exercise programme : resistance training (calf muscle resistance exercises) and stretching exercises						
	Exercise mode: heel raises and stretching exercises						
	Comparator: conventional stretching to the calf muscle, resisted upper limb bicep curls						
	Setting: not reported						
	Intervention provider: not reported						
	Duration of programme: 10 weeks						
	Exercise dose : strengthening exercises 4 times a week for 10 weeks, stretching exercises for the final 6 weeks of the intervention						
	Tailoring of intervention to individual : exercise load can be reduced by changing to bilateral heel raises, giving external support, reducing the range of motion or performing the heel raises while seated. Exercise load will be progressively increased by adding weight in the form of water bottles to a rucksack worn on the participant's back. Extra load will be added to biceps curls by using water bottles held in the hand						

Exercise interventions for cerebral palsy (Review)



NCT02766491 (Continued)

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	Fidelity to prescribed intervention and monitoring of adverse events: not reported							
Outcomes	Assessment time points: baseline (week 0), immediately postintervention (week 10)							
	Primary outcomes:							
	 Change in muscle fascicle length of the gastrocnemius (mm) using B-mode ultrasound images Change in gastrocnemius muscle length (mm) using B-mode ultra sound images 							
	Secondary outcomes:							
	1. Change in ankle range of motion (degree, goniometer)							
	2. Change in Achilles tendon stiffness (Nm) quantified as the change in tendon length per change in tendon force, using B-mode ultrasound images (also measured at 4 weeks after baseline)							
	3. Change in maximal dorsiflexion angle during gait (degree) quantified from kinematic data ob- tained during gait analysis							
	4. Change in lengthening properties of the muscle fascicles (mm) using B-mode ultrasound images							
	 Changes in step length during gait (m) quantified from kinematic data obtained during gait analy- sis 							
	6. Changes in ankle power at push off during gait quantified from kinematic and kinetic data ob- tained during gait analysis							
Starting date	June 2016							
Contact information	Email: B.M.Kalkman@2014.ljmu.ac.uk							
Notes	Source of funding: not reported							
	Potential conflicts of interest: not reported							

NCT02754128							
Trial name or title	BeFast or BeStrong: linking neuroplasticity with the outcomes of walking-based interventions: a feasibility trial comparing a motor learning versus a strength-based program in children with CP						
Methods	Design: randomised controlled trial						
	Country of origin: Canada						
	Intervention(s) : motor learning-based gait-related training intervention versus a functional lower limb strength training intervention						
	Unit of allocation: individual						
Participants	Number of participants: 22						
	Inclusion criteria : aged 7-17 years, diagnosis of hemiplegic or diplegic CP, GMFCS level I or II, able to follow testing and motor imagery instructions, able to actively participate in a minimum of 45 min of physical activity, show evidence of independent dorsiflexion of both ankles, able to commit to attendance of sessions 2-3 times weekly for 6 weeks						
	Exclusion criteria : orthopaedic surgery within the last 9 months (muscle) or 12 months (bone), botulinum toxin-A (BTX-A) injections to lower limb in the last 4 months, inability to put BTX-A on hold during trial, severe spasticity (may be a contraindication for neuroimaging procedures), seizure disorder (if not fully controlled by medication for 12 months), not prepared or unable to discontinue any formal lower limb therapy intervention or physical activity programme during the trial, involved in another intervention study, standard MRI contraindications (e.g. magnetic implants, inability to lay still, claustrophobia etc.)						

Exercise interventions for cerebral palsy (Review)



NCT02754128 (Continued)

Interventions

Aim of the intervention: to compare a motor learning-based gait-related training intervention to a functional lower limb strength training intervention, and to evaluate functional, neural and participation outcomes for children and young people with CP

Type of exercise programme: resistance training versus aerobic training

Exercise mode: motor learning (ML)-based gait-related training programme, and a 3-5 min mental motor imagery script to practice on days when there are no active training sessions. Exercises are designed to improve advanced gross motor skills and athleticism

Comparator: functional strength training programme designed to improve gait-related skills and a 3-5 min home programme of strength exercises to practice on days when there are no active training sessions

Setting: not reported

Intervention provider: not reported

Duration of programme: 6 weeks (a maximum of 7 weeks will be permitted)

Exercise dose: 45-min training sessions 2-3 times a week, over 6 weeks for a total of 16 active sessions (training can extend to a 7th week if necessary), plus home based training for non-active training days. Total training will be 5 times per week.

Tailoring of intervention to individual: not reported

Fidelity to prescribed intervention and monitoring of adverse events:

Feasibility process indicators: retention rate, perceived intervention benefit (using a combination of participant and parent ratings) retention rate (number enrolled compared to number completed).

Feasibility resource indicators: adherence rate (number of completed sessions), data collection time (projected versus actual), data collection completion (% missing data)

Feasiblity management indicators: intervention fidelity (within session effort scores); intervention fidelity/contamination (video sessions every 2 weeks to assess session content: STRONG group: 1 RM lower limb strength progression, FAST group: session content via Motor Learning Strategy Rating Instrument (MLSRI); intervention fidelity (completion of diaries for motor imagery/strength home practice); intervention fidelity/contamination (log books for PA-participation tracking/management in active intervention and 4 month follow-up); treatment administration (PTA/RKin session summary form data); acceptability of intervention (aggregate score of: parent/staff satisfaction scale, child physical activity enjoyment scale (PACES), child intervention satisfaction score)

Adverse events recorded as part of the feasibility science indicators through to study completion

Outcomes

Assessment time points: 7 days pre/7 days post/4-months post-training intervention unless otherwise stated

Primary outcomes:

1. Change from baseline in advanced motor skills on the Challenge Module, a measure of advanced motor skills

Secondary outcomes:

- 1. Change from baseline in functional activity in lower-limb related cortical areas, assessed using functional MRI (fMRI) measured at 7 days pre/7 days post
- 2. Change from baseline in resting state activity, assessed using resting state fMRImeasured at 7 days pre/7 days post
- 3. Change from baseline in microstructure of brain, assessed using diffusion tensor imaging (DTI) measured at 7 days pre/7 days post

Exercise interventions for cerebral palsy (Review)

NCT02754128 (Continued)

- 4. Change from baseline in Physical Activity self-efficacy, self-report measure of task efficacy and barrier efficacy for physical activity
- 5. Change from baseline in walking activity, assessed using an Actigraph accelerometer
- 6. Change from baseline in gait kinematics as measured using an electronic walkway for time/distance parameters of footsteps via GAITRite system
- 7. Change from baseline in physical activity participation, as measured using the Participation and Environment Measure for Children/Youth, a parent report measure of participation
- 8. Change from baseline in walk speed on the 6-min walk test
- 9. Change from baseline in targeted goal abilities and satisfaction with performance as measured by the Canadian Occupational Performance Measure via 3-5 individualized walking-based activity/participation tasks set at baseline with assessor and child/parent
- 10. Change from baseline in targeted goal abilities as measured by Goal Attainment Scaling (GAS) using 3-5 individualised walking-based activity/participation set at baseline with assessor and child/ parent

Other outcome measures

- 1. Motor learning as evaluated using retention and transfer tests at weeks 2, 4, 6; sessions 5, 10 and 15 designated training transfer sessions
- 2. Motor learning content of interventions, as assessed using the Motor Learning Strategy Rating Instrument (week 2, 4, 6) to determine the extent to which motor learning strategies are used in an intervention session
- 3. Intervention programme enjoyment, assessed using a modified version of the Physical Activity Enjoyment Scale (PACES) at 7 days post-training
- 4. Intervention session enjoyment using a study specific questionnaire 16 times over 6 weeks at the end of each intervention.
- 5. Rating of exertion, assessed using Pictorial Children's Effort Rating Table at the mid-point of each session and 2 min from the end of each session
- 6. Heart rate (beats per minute) via the radial pulse, at 4 time points per session: 1 min before start of session, at 22 min (mid-point), at 43 min (2 min before end), and at 45 min (end) of each session
- 7. Body pain, using the FACES pain scale and body diagrams to show areas of pain at 2 min before and 2 min after each intervention session
- 8. Parent satisfaction scale at 7 days post-training intervention using a study-specific questionnaire
- 9. Staff satisfaction scale at 7 days post-training intervention using a study-specific questionnaire
- 10.Lower limb strength using a 30-s bilateral lateral step-up test
- 11. Ankle range of motion and dorsiflexion force, passive and active range using a goniometer
- 12. Dorsiflexion force using surface electromyography (EMG)
- 13.Lower limb joint-sense position using a semi-goniometer
- 14.Mental chronometry; timed while walking a 10-m distance, and then timed while imagining walking the same distance

15.Motor Imagery Questionnaire for Children (MIQ-C)

Starting date	June 2016
Contact information	Alicia J Hilderley email: ahilderley@hollandbloorview.ca
Notes	Source of funding: not reported
	Potential conflicts of interest: not reported

CP: cerebral palsy; EMG: electromyography; GMFCS: Gross Motor Function Classification System; GMFM: Gross Motor Function Measure; PA: physical activity; PTA/RKin: Physiotherapist Assistant/Registered Kinesiologist.



DATA AND ANALYSES

Comparison 1. Aerobic exercise versus usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Activity: gross motor function, short term	3	65	Std. Mean Difference (IV, Ran- dom, 95% CI)	0.53 [0.02, 1.04]
2 Activity: gait speed, short term	4	82	Mean Difference (IV, Random, 95% CI)	0.09 [-0.11, 0.28]
3 Activity: walking endurance; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
4 Activity: gait speed, intermediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
5 Activity: gross motor function, in- termediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
6 Activity: daily physical activity; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
7 Aerobic fitness; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only

Analysis 1.1. Comparison 1 Aerobic exercise versus usual care, Outcome 1 Activity: gross motor function, short term.

Study or subgroup	Aerob	oic exercise	Usual care		Std. Mean Difference		Weight	Std. Mean Difference		
	N	Mean(SD)	n(SD) N Mean(SD) Ran			ndom, 95% Cl		Random, 95% CI		
Bryant 2013	20	1.9 (3.3)	11	0.2 (1.8)				45.7%	0.56[-0.19,1.31]	
Chrysagis 2012	11	71.7 (17.7)	11	65.1 (16.5)				36.15%	0.37[-0.48,1.21]	
Mattern-Baxter 2013	6	16.9 (4.8)	6	13.9 (1.8)			+	18.15%	0.76[-0.43,1.95]	
Total ***	37		28				•	100%	0.53[0.02,1.04]	
Heterogeneity: Tau ² =0; Chi ² =0	0.3, df=2(P=0.86); I ² =0%								
Test for overall effect: Z=2.04(P=0.04)									
			Favo	urs usual care	-4	-2	0 2	4 Favours a	erobic	

Analysis 1.2. Comparison 1 Aerobic exercise versus usual care, Outcome 2 Activity: gait speed, short term.

Study or subgroup	Aerob	Aerobic exercise		Usual care		Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ran	dom, 95	% CI			Random, 95% Cl
Chrysagis 2012	11 1 (0.1) 11 0.8 (0.2)			34.26%	0.22[0.08,0.35]						
Gharib 2011	15	0.7 (0.1)	15	0.6 (0.1)						38.95%	0.04[-0.03,0.11]
Mattern-Baxter 2013	6	0.7 (0.5)	6	2.4 (1.5)			_			2.31%	-1.7[-2.94,-0.46]
Smania 2011	9	1 (0.3)	9	0.8 (0.2)			+			24.48%	0.15[-0.1,0.4]
			Fa	vours aerobic	-5	-2.5	0	2.5	5	- Favours usua	l care

Exercise interventions for cerebral palsy (Review)



Study or subgroup	Aerol	bic exercise	Usual care		Mean Difference				Weight	Mean Difference	
	N	Mean(SD)	N	Mean(SD)		Ran	dom, 95	% CI			Random, 95% Cl
Total ***	41		41				•			100%	0.09[-0.11,0.28]
Heterogeneity: Tau ² =0.02; Ch	i ² =13.46, df=3(F	P=0); I ² =77.72%									
Test for overall effect: Z=0.88	(P=0.38)										
			Fav	ours aerobic	-5	-2.5	0	2.5	5	- Favours usua	Il care

Analysis 1.3. Comparison 1 Aerobic exercise versus usual care, Outcome 3 Activity: walking endurance; short term.

Study or subgroup	Aerob	ic exercise	Us	ual care	l care Mean Difference		Weight	Mean Difference			
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI					Random, 95% CI	
Smania 2011	9	360 (128.7)	9	319 (39.6)				-+		0%	41[-46.98,128.98]
			Favo	urs usual care	-100	-50	0	50	100	Favours aerol	bic

Analysis 1.4. Comparison 1 Aerobic exercise versus usual care, Outcome 4 Activity: gait speed, intermediate term.

Study or subgroup	Aerob	Aerobic exercise Usual care		ual care		Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Random, 95% CI			Random, 95% CI		
Mattern-Baxter 2013	6	0.4 (0.3)	6	0.6 (0.4)			+			0%	-0.17[-0.59,0.24]
			Favo	urs usual care	-1	-0.5	0	0.5	1	Favours aerobi	c

Analysis 1.5. Comparison 1 Aerobic exercise versus usual care, Outcome 5 Activity: gross motor function, intermediate term.

Study or subgroup	Aerob	ic exercise	Usual care			Mean Difference				Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% CI					Random, 95% CI	
Mattern-Baxter 2013	6	34.5 (14.9)	6	21.5 (4.4)					0%	12.96[0.52,25.4]	
			Favoi	ırs usual care	-50	-25	0	25	50	Favours aerobio	2

Analysis 1.6. Comparison 1 Aerobic exercise versus usual care, Outcome 6 Activity: daily physical activity; short term.

Study or subgroup	Aerob	ic exercise	Usual care Mean Difference				Weight	Mean Difference			
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% CI				Random, 95% CI		
Van den Berg-Emons 1998	10	1.6 (0.2)	10	1.3 (0.2)		1			-	0%	0.21[0.04,0.38]
			Favo	urs usual care	-0.5	-0.25	0	0.25	0.5	Favours aerobi	c

Analysis 1.7. Comparison 1 Aerobic exercise versus usual care, Outcome 7 Aerobic fitness; short term.

Study or subgroup	Aerob	Aerobic exercise Usu		Usual care		Mean Difference				Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% CI				Random, 95% Cl		
Van den Berg-Emons 1998	10	1.2 (0.8)	10	1.2 (0.9)						0%	0.06[-0.71,0.83]
			Favo	urs usual care	-2	-1	0	1	2	Favours aerobi	c

Comparison 2. Resistance training versus usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Activity: gross motor function, children and adolescents; short term	7	164	Std. Mean Difference (IV, Random, 95% CI)	0.12 [-0.19, 0.43]
2 Activity: gross motor function, children and adolescents; intermediate term	3	85	Std. Mean Difference (IV, Random, 95% CI)	0.13 [-0.30, 0.55]
3 Activity: gait speed, children and ado- lescents; short term	8	185	Mean Difference (IV, Ran- dom, 95% CI)	0.03 [-0.02, 0.07]
4 Activity: gait speed, children and ado- lescents; intermediate term	3	84	Mean Difference (IV, Ran- dom, 95% CI)	-0.03 [-0.17, 0.11]
5 Activity: gait speed, adults; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
6 Activity: gross motor function, adults; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
7 Activity: walking endurance, adults; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
8 Participation, children and adoles- cents; short term	2	127	Std. Mean Difference (IV, Random, 95% CI)	0.34 [-0.01, 0.70]
9 Participation, children and adoles- cents; intermediate term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
10 Quality of life (parent-reported), chil- dren and adolescents; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
11 Quality of life (child-reported), chil- dren and adolescents; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
12 Muscle strength, children and adoles- cents; short term	8	247	Std. Mean Difference (IV, Random, 95% CI)	0.53 [0.00, 1.06]
13 Muscle strength, children and adoles- cents; intermediate term	3	84	Std. Mean Difference (IV, Random, 95% CI)	0.50 [0.06, 0.94]
14 Muscle strength, adults; short term	1		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only

Exercise interventions for cerebral palsy (Review)



Analysis 2.1. Comparison 2 Resistance training versus usual care, Outcome 1 Activity: gross motor function, children and adolescents; short term.

Study or subgroup	Resista	nce training	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% CI
Dodd 2003	11	69 (21.4)	10	75.3 (21.3)	+	13.13%	-0.28[-1.14,0.58]
Engsberg 2006	9	69 (28.4)	3	71.4 (10)		5.7%	-0.09[-1.39,1.22]
Liao 2007	10	82.7 (2.2)	10	80.6 (2.2)	+	11.24%	0.91[-0.02,1.84]
Seniorou 2007	11	55.6 (28)	9	60.8 (26.5)	+	12.5%	-0.18[-1.07,0.7]
Lee 2008	9	62.7 (34.1)	8	61.4 (33.9)		10.74%	0.04[-0.92,0.99]
Scholtes 2010	24	76.1 (11.8)	24	73.1 (12.4)		30.2%	0.24[-0.32,0.81]
Lee 2015	13	81.9 (16.1)	13	81.3 (14.3)		16.49%	0.04[-0.73,0.81]
Total ***	87		77		•	100%	0.12[-0.19,0.43]
Heterogeneity: Tau ² =0; Chi ² =	=4.4, df=6(P=0.62)); I ² =0%					
Test for overall effect: Z=0.76	6(P=0.45)						
-			Favo	urs usual care	-2 -1 0 1 2	Favours re	sistance

Favours usual care

avours resistance

Analysis 2.2. Comparison 2 Resistance training versus usual care, Outcome 2 Activity: gross motor function, children and adolescents; intermediate term.

Study or subgroup	Resista	ance training	aining Usual care Std. Mean Difference		Weight	Std. Mean Difference				
	Ν	N Mean(SD)		N Mean(SD)		Rai	ndom, 95% CI		Random, 95% Cl	
Dodd 2003	11	69.6 (21.4)	9	74.3 (21.4)				23.4%	-0.21[-1.09,0.67]	
Lee 2008	9	63 (34.4)	8	61.8 (34)				20.16%	0.03[-0.92,0.99]	
Scholtes 2010	24	76.6 (13)	24	72.7 (12.8)				56.44%	0.3[-0.27,0.87]	
Total ***	44		41				-	100%	0.13[-0.3,0.55]	
Heterogeneity: Tau ² =0; Chi ² =	0.94, df=2(P=0.6	2); I ² =0%								
Test for overall effect: Z=0.57	(P=0.57)									
			Favo	urs usual care	-2	-1	0 1	² Favours re	sistance	

Analysis 2.3. Comparison 2 Resistance training versus usual care, Outcome 3 Activity: gait speed, children and adolescents; short term.

Study or subgroup	Resista	nce training	Us	ual care	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% Cl
Dodd 2003	11	0.8 (0.4)	10	0.8 (0.3)		2.17%	-0.04[-0.34,0.26]
Unger 2006	21	1.1 (0.2)	10	1.2 (0.1)	-+	11.04%	-0.05[-0.18,0.08]
Engsberg 2006	9	0.9 (0.3)	3	0.8 (0.3)		1.1%	0.12[-0.3,0.54]
Liao 2007	10	1 (0.1)	10	1 (0.1)		31.96%	0.04[-0.04,0.12]
Seniorou 2007	11	0.3 (0.1)	9	0.3 (0.1)		25.07%	0[-0.09,0.09]
Lee 2008	9	0.7 (0.4)	8	0.7 (0.4)		1.28%	0.06[-0.33,0.45]
Scholtes 2010	23	1 (0.3)	23	1.1 (0.4)		4.6%	-0.04[-0.25,0.17]
Pandey 2011	9	0.7 (0.1)	9	0.6 (0.1)		22.79%	0.1[0.01,0.19]
Total ***	103		82		•	100%	0.03[-0.02,0.07]
Heterogeneity: Tau ² =0; Chi ² =	=5.06, df=7(P=0.6	5); I ² =0%					
			Favo	urs usual care ⁻¹	-0.5 0 0.5	¹ Favours res	istance

Exercise interventions for cerebral palsy (Review)



Study or subgroup	Resistance training Usual care			Mean Difference				Weight	Mean Difference		
	Ν	Mean(SD)	Ν	Mean(SD)		Random, 95% CI				Random, 95% Cl	
Test for overall effect: Z=1.29(P=0.2)					I					
			Favo	ours usual care	-1 -0.5 0 0.5		1	Favours resis	stance		

Analysis 2.4. Comparison 2 Resistance training versus usual care, Outcome 4 Activity: gait speed, children and adolescents; intermediate term.

Study or subgroup	Resista	Resistance training		ual care	Меа	Mean Difference		Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Ran	dom, 95% CI		Random, 95% CI
Dodd 2003	11	0.8 (0.4)	9	0.9 (0.3)			23.23%	-0.05[-0.34,0.24]
Lee 2008	9	0.8 (0.4)	8	0.7 (0.4)	-		14.89%	0.1[-0.26,0.47]
Scholtes 2010	24	1 (0.3)	23	1.1 (0.3)	-		61.88%	-0.06[-0.24,0.12]
Total ***	44		40			•	100%	-0.03[-0.17,0.11]
Heterogeneity: Tau ² =0; Chi ² =	0.64, df=2(P=0.7	3); I ² =0%						
Test for overall effect: Z=0.45	(P=0.65)							
			Favo	urs usual care ⁻¹	-0.5	0 0.5	¹ Favours res	stance

Analysis 2.5. Comparison 2 Resistance training versus usual care, Outcome 5 Activity: gait speed, adults; short term.

Study or subgroup	Resista	nce training	Us	ual care		Mear	n Diffe	rence		Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rand	lom, 9	5% CI			Random, 95% Cl
Maeland 2009	6	1.1 (0.4)	6	0.8 (0.6)		-		•		0%	0.3[-0.28,0.88]
			Favoi	urs usual care	-1	-0.5	0	0.5	1	Favours resist	ance

Analysis 2.6. Comparison 2 Resistance training versus usual care, Outcome 6 Activity: gross motor function, adults; short term.

Study or subgroup	Resista	nce training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	6 CI			Random, 95% Cl
Maeland 2009	6	38 (29)	6	28 (5)		-			1	0%	10[-13.55,33.55]
			Favo	urs usual care	-50	-25	0	25	50	Favours resist	ance

Analysis 2.7. Comparison 2 Resistance training versus usual care, Outcome 7 Activity: walking endurance, adults; short term.

Study or subgroup	Resista	nce training	g Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rai	ndom, 95%	6 CI			Random, 95% CI
Maeland 2009	6	425 (186)	6	298 (206)				+		0%	127[-95.08,349.08]
			Favou	urs usual care	-500	-250	0	250	500	Favours resis	stance

Exercise interventions for cerebral palsy (Review)

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Analysis 2.8. Comparison 2 Resistance training versus usual care, Outcome 8 Participation, children and adolescents; short term.

Study or subgroup	Resista	nce training	training Usual care Std. Mean Difference			Weight	Std. Mean Difference			
	N	Mean(SD)	Ν	Mean(SD)		Rai	ndom, 95% CI			Random, 95% Cl
Scholtes 2010	22	39 (13.3)	17	31.1 (13.6)					29.64%	0.57[-0.07,1.22]
Mitchell 2016	46	8 (2.3)	42	7.4 (2.5)					70.36%	0.25[-0.17,0.67]
Total ***	68		59				•		100%	0.34[-0.01,0.7]
Heterogeneity: Tau ² =0; Chi ² =	0.69, df=1(P=0.4	1); I ² =0%								
Test for overall effect: Z=1.92	(P=0.06)									
			Favo	urs usual care	-2	-1	0 1	2	Favours res	istance

Analysis 2.9. Comparison 2 Resistance training versus usual care, Outcome 9 Participation, children and adolescents; intermediate term.

Study or subgroup	Resista	nce training	Usual care			Mean Difference				Weight I	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	% CI		I	Random, 95% Cl
Scholtes 2010	19	32.2 (9.3)	17	31.8 (11.8)				1		0%	0.37[-6.61,7.35]
			Favo	urs usual care	-10	-5	0	5	10	Favours resistan	ce

Analysis 2.10. Comparison 2 Resistance training versus usual care, Outcome 10 Quality of life (parent-reported), children and adolescents; short term.

Study or subgroup	Resista	nce training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	6 CI			Random, 95% Cl
Engsberg 2006	9	66.3 (13.8)	3	53.6 (14.1)				+		0%	12.7[-5.63,31.03]
			Favo	urs usual care	-50	-25	0	25	50	Favours resist	ance

Analysis 2.11. Comparison 2 Resistance training versus usual care, Outcome 11 Quality of life (child-reported), children and adolescents; short term.

Study or subgroup	Resista	nce training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	% CI			Random, 95% CI
Engsberg 2006	9	70.8 (11.9)	3	59.1 (16.3)	1	1		·		0%	11.7[-8.32,31.72]
			Favo	urs usual care	-50	-25	0	25	50	Favours resis	stance

Analysis 2.12. Comparison 2 Resistance training versus usual care, Outcome 12 Muscle strength, children and adolescents; short term.

Study or subgroup	Resista	nce training	Usual care			Std. Mean Difference				Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	CI			Random, 95% Cl
Dodd 2003	11	33.1 (15.8)	10	25.5 (9.9)			+-			12.67%	0.55[-0.33,1.42]
Liao 2007	10	6.1 (1.3)	10	6.2 (1.3)			-			12.66%	-0.08[-0.95,0.8]
			Favo	urs usual care	-10	-5	0	5	10	Favours resi	stance

Exercise interventions for cerebral palsy (Review)



Study or subgroup	Resista	nce training	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% CI
Seniorou 2007	11	1.3 (0.5)	9	1.2 (0.5)	- - -	12.6%	0.19[-0.69,1.07]
Lee 2008	9	13.2 (5.4)	8	14.1 (5.8)	_ + _	11.89%	-0.15[-1.11,0.8]
Scholtes 2010	24	5.4 (1.1)	24	4.5 (1.2)	+	15.68%	0.77[0.19,1.36]
Reid 2010	7	184.7 (15.3)	7	211.8 (116.7)	_+	10.94%	-0.3[-1.36,0.75]
Pandey 2011	9	6.3 (1.1)	9	2.7 (0.5)		6.31%	3.95[2.22,5.67]
Mitchell 2016	46	63.5 (26)	43	46.8 (18.3)	-+-	17.25%	0.73[0.3,1.16]
Total ***	127		120		•	100%	0.53[0,1.06]
Heterogeneity: Tau ² =0.37; Ch	ni²=23.3, df=7(P=	0); I ² =69.95%					
Test for overall effect: Z=1.97	r(P=0.05)						
			Favo	urs usual care -10	-5 0 5	¹⁰ Favours re	sistance

Analysis 2.13. Comparison 2 Resistance training versus usual care, Outcome 13 Muscle strength, children and adolescents; intermediate term.

Study or subgroup	Resista	nce training	Usual care		Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% CI
Dodd 2003	11	32.5 (11.4)	9	25.2 (7.8)		23.17%	0.7[-0.21,1.62]
Lee 2008	9	13.7 (5.9)	8	14.4 (5.9)		21.29%	-0.11[-1.07,0.84]
Scholtes 2010	23	5.2 (1)	24	4.5 (1.2)		55.54%	0.65[0.06,1.23]
Total ***	43		41		•	100%	0.5[0.06,0.94]
Heterogeneity: Tau ² =0; Chi ² =	=2.01, df=2(P=0.3	7); I ² =0.67%					
Test for overall effect: Z=2.2	1(P=0.03)						
			Favo	urs usual care	-2 -1 0 1 2	Favours re	sistance

Analysis 2.14. Comparison 2 Resistance training versus usual care, Outcome 14 Muscle strength, adults; short term.

Study or subgroup	Resista	nce training	Us	ual care		Ме	an Differe	nce		Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	6 CI			Random, 95% Cl
Maeland 2009	6	76 (50)	6	83 (41)						0%	-7[-58.74,44.74]
			Favoi	urs usual care	-100	-50	0	50	100	Favours resis	tance

Comparison 3. Mixed training versus usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Activity: gross motor function; short term	4	163	Std. Mean Difference (IV, Random, 95% CI)	0.02 [-0.29, 0.33]
2 Activity: gait speed; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only

Exercise interventions for cerebral palsy (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
3 Activity: walking endurance; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
4 Participation; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
5 Participation; intermediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
6 Aerobic fitness; short term	2	78	Std. Mean Difference (IV, Random, 95% CI)	0.05 [-0.39, 0.50]
7 Muscle strength; short term	3	150	Std. Mean Difference (IV, Random, 95% CI)	0.08 [-0.24, 0.40]
8 Anaerobic fitness; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
9 Aerobic fitness; intermediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
10 Anaerobic fitness; intermedi- ate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
11 Muscle strength; intermedi- ate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only

Analysis 3.1. Comparison 3 Mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.

Study or subgroup	Mixe	d training	Us	ual care		Std. M	lean Difference		Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ran	idom, 95% CI			Random, 95% Cl
Verschuren 2007	32	87.2 (12.8)	33	90.1 (11.1)					39.86%	-0.24[-0.73,0.25]
Unnithan 2007	7	33.9 (17.9)	6	30.8 (12.5)					7.94%	0.18[-0.91,1.28]
Fowler 2010	29	70.8 (11)	29	69.3 (10.3)					35.75%	0.14[-0.38,0.65]
Chen 2012	13	84.2 (11.7)	14	81 (8.8)					16.45%	0.3[-0.46,1.06]
Total ***	81		82				•		100%	0.02[-0.29,0.33]
Heterogeneity: Tau ² =0; Chi ² =1	L.89, df=3(P=0.6	; I ² =0%								
Test for overall effect: Z=0.12(P=0.9)									
			Favo	urs usual care	-2	-1	0 1	2	Favours mixe	d

Favours usual care

Analysis 3.2. Comparison 3 Mixed training versus usual care, Outcome 2 Activity: gait speed; short term.

Study or subgroup	Mixe	d training	Us	ual care		Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	% CI			Random, 95% CI
Fowler 2010	29	1.1 (0.3)	29	1 (0.3)	1	I				0%	0.1[-0.07,0.27]
			Favo	urs usual care	-0.5	-0.25	0	0.25	0.5	Favours mixed	

Exercise interventions for cerebral palsy (Review)

Study or subgroup	Mixe	d training	Us	ual care		Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	1dom, 95%	CI			Random, 95% Cl
Fowler 2010	27	90.6 (38.4)	28	84.1 (42.6)		_				0%	6.5[-14.91,27.91]
			Favoi	urs usual care	-50	-25	0	25	50	Favours mixed	

Analysis 3.3. Comparison 3 Mixed training versus usual care, Outcome 3 Activity: walking endurance; short term.

Analysis 3.4. Comparison 3 Mixed training versus usual care, Outcome 4 Participation; short term.

Study or subgroup	Mixe	d training	Us	ual care		Mean Difference			Weight	Mean Difference	
	N	Mean(SD)	Ν	Mean(SD)		Rar	ndom, 95%	6 CI			Random, 95% CI
Verschuren 2007	32	0 (0.5)	33	-0.4 (0.6)		1	_			0%	0.4[0.13,0.67]
			Favoi	urs usual care	-1	-0.5	0	0.5	1	Favours mixed	

Analysis 3.5. Comparison 3 Mixed training versus usual care, Outcome 5 Participation; intermediate term.

Study or subgroup	Mixe	d training	Usual care			Mea	an Differe	nce		Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI					Random, 95% CI	
Verschuren 2007	32	1.8 (0.9)	33	1.7 (0.7)						0%	0.15[-0.25,0.54]
			Favo	urs usual care	-1 -0.5 0 0.5 1		1	Favours mixe	ed training		

Analysis 3.6. Comparison 3 Mixed training versus usual care, Outcome 6 Aerobic fitness; short term.

Study or subgroup	Mixe	d training	Us	ual care	Std. Mean Difference				Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95% Cl			Random, 95% Cl
Unnithan 2007	7	20.8 (5.9)	6	18 (4.1)		-	+		15.99%	0.5[-0.61,1.62]
Verschuren 2007	32	7.7 (4.1)	33	7.8 (4)					84.01%	-0.03[-0.52,0.45]
Total ***	39		39				-		100%	0.05[-0.39,0.5]
Heterogeneity: Tau ² =0; Chi ² =0	0.74, df=1(P=0.39	9); I ² =0%								
Test for overall effect: Z=0.24((P=0.81)									
			Favo	urs usual care	-2	-1	0	1 2	Favours mixe	d

Analysis 3.7. Comparison 3 Mixed training versus usual care, Outcome 7 Muscle strength; short term.

Study or subgroup	Mixe	d training	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% Cl
Verschuren 2007	32	37.4 (18.8)	33	38.5 (19.4)		43.57%	-0.05[-0.54,0.43]
Fowler 2010	28	0.9 (0.3)	29	0.9 (0.4)		38.19%	0.07[-0.44,0.59]
Chen 2012	13	1.6 (0.8)	15	1.4 (0.6)	+	18.24%	0.41[-0.34,1.16]
Total ***	73		77		•	100%	0.08[-0.24,0.4]
			Favo	urs usual care -2	-1 0 1	² Favours m	ixed training

Exercise interventions for cerebral palsy (Review)



Study or subgroup	Mixe	Mixed training Usual care		ual care		Std. I	Mean Diffe	rence		Weight S	td. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	CI			Random, 95% Cl
Heterogeneity: Tau ² =0; Chi ² =1.	.02, df=2(P=0.6); I ² =0%									
Test for overall effect: Z=0.49(F	P=0.63)										
			Favou	Irs usual care	-2	-1	0	1	2	Favours mixed	training

Analysis 3.8. Comparison 3 Mixed training versus usual care, Outcome 8 Anaerobic fitness; short term.

Study or subgroup	Mixe	d training	Usual care			Mea	an Differe	nce		Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	ndom, 95%	% CI			Random, 95% Cl
Verschuren 2007	32	20.4 (38)	33	-4.8 (28.2)		I	-		_	0%	25.2[8.89,41.51]
			Favoi	urs usual care	-50	-25	0	25	50	Favours mixed	

Analysis 3.9. Comparison 3 Mixed training versus usual care, Outcome 9 Aerobic fitness; intermediate term.

Study or subgroup	Mixe	d training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95ª	% CI			Random, 95% Cl
Verschuren 2007	32	7.7 (4.1)	33	7.8 (4)						0%	-0.13[-2.1,1.84]
			Favoi	urs usual care	-5	-2.5	0	2.5	5	Favours mixed	

Analysis 3.10. Comparison 3 Mixed training versus usual care, Outcome 10 Anaerobic fitness; intermediate term.

Study or subgroup	Mixe	d training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ran	1dom, 95%	6 CI			Random, 95% CI
Verschuren 2007	32	94.5 (75.7)	33	125.8 (90.9)					0%	-31.28[-71.89,9.33]	
			Favo	ours usual care	-100	-50	0	50	100	Favours mixed	

Analysis 3.11. Comparison 3 Mixed training versus usual care, Outcome 11 Muscle strength; intermediate term.

Study or subgroup	Mixe	d training	Usual care			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rai	ndom, 95%	6 CI			Random, 95% CI
Verschuren 2007	32	37.4 (18.8)	33	38.5 (19.4)						0%	-1.04[-10.33,8.25]
			Favoi	urs usual care	-20	-10	0	10	20	Favours mixed	

Comparison 4. Resistance training versus aerobic exercise

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Activity: gross motor function; short term	2	56	Std. Mean Difference (IV, Random, 95% CI)	0.02 [-0.50, 0.55]

Exercise interventions for cerebral palsy (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2 Activity: gait speed; short term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
3 Activity: gait speed; intermediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
4 Activity: gross motor function; in- termediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only
5 Muscle strength; short term	2	56	Std. Mean Difference (IV, Random, 95% CI)	-0.11 [-0.64, 0.41]
6 Muscle strength; intermediate term	1		Mean Difference (IV, Random, 95% CI)	Subtotals only

Analysis 4.1. Comparison 4 Resistance training versus aerobic exercise, Outcome 1 Activity: gross motor function; short term.

Study or subgroup	Aerob	Aerobic exercise		nce training		Std. M	Aean Differend	e	Weight		Std. Mean Difference	
	N	Mean(SD)	Ν	Mean(SD)		Rai	ndom, 95% Cl				Random, 95% CI	
Olama 2011	15	44.1 (40.1)	15	46.7 (4.3)						53.77%	-0.09[-0.8,0.63]	
Johnston 2011	14	63.3 (16.2)	12	60.1 (25.1)		-		-		46.23%	0.15[-0.62,0.92]	
Total ***	29		27							100%	0.02[-0.5,0.55]	
Heterogeneity: Tau ² =0; Chi ² =0	0.2, df=1(P=0.66); I ² =0%										
Test for overall effect: Z=0.08(P=0.94)											
			Favo	urs resistance	-2	-1	0	1	2	Favours aerob	ic	

Analysis 4.2. Comparison 4 Resistance training versus aerobic exercise, Outcome 2 Activity: gait speed; short term.

Study or subgroup	Aerob	oic exercise	Resistance training			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ra	ndom, 95%	6 CI			Random, 95% CI
Johnston 2011	14	0.6 (0.3)	12	0.5 (0.4)						0%	0.12[-0.15,0.39]
			Favo	urs resistance	-1	-0.5	0	0.5	1	Favours aerobio	2

Analysis 4.3. Comparison 4 Resistance training versus aerobic exercise, Outcome 3 Activity: gait speed; intermediate term.

Study or subgroup	Aerob	ic exercise	Resistance training			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	1dom, 95%	% CI			Random, 95% CI
Johnston 2011	14	0.6 (0.3)	12	0.4 (0.3)	· · · · · · · · · · · · · · · · · · ·				0%	0.19[-0.05,0.43]	
			Favo	urs resistance	-1	-0.5	0	0.5	1	Favours aerobio	2

Exercise interventions for cerebral palsy (Review)

Analysis 4.4. Comparison 4 Resistance training versus aerobic exercise, Outcome 4 Activity: gross motor function; intermediate term.

Study or subgroup	Aerob	ic exercise	Resistance training			Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rai	ndom, 95%	6 CI			Random, 95% CI
Johnston 2011	14	65.3 (16.5)	12	60.6 (26.7)						0%	4.7[-12.7,22.1]
			Favou	urs resistance	-50	-25	0	25	50	Favours aerobio	2

Analysis 4.5. Comparison 4 Resistance training versus aerobic exercise, Outcome 5 Muscle strength; short term.

udy or subgroup Aerobic e		oic exercise	ercise Resistance training		Std. Mean Difference				Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Ran	dom, 95% CI			Random, 95% CI
Johnston 2011	14	3.6 (2.8)	12	3.8 (4.2)			-		46.36%	-0.06[-0.83,0.71]
Olama 2011	15	29.5 (4)	15	30.1 (3.9)					53.64%	-0.16[-0.88,0.56]
Total ***	29		27						100%	-0.11[-0.64,0.41]
Heterogeneity: Tau ² =0; Chi ² =0	0.03, df=1(P=0.8	5); I ² =0%								
Test for overall effect: Z=0.42(P=0.67)									
			Favo	urs resistance	-2	-1	0 1	2	Favours aero	bic

Analysis 4.6. Comparison 4 Resistance training versus aerobic exercise, Outcome 6 Muscle strength; intermediate term.

Study or subgroup	Aerob	Aerobic exercise		Resistance training		Mean Difference				Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	ndom, 95%	6 CI			Random, 95% CI
Johnston 2011	14	3.7 (3.3)	12	3.7 (3.7)						0%	-0.03[-2.71,2.65]
			Favo	urs resistance	-5	-2.5	0	2.5	5	Favours aerobio	2

Comparison 5. Aerobic exercise and mixed training versus usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Activity: gross motor function; short term	7	228	Std. Mean Difference (IV, Ran- dom, 95% CI)	0.36 [0.09, 0.62]
2 Activity: gross motor function, in- termediate term	2	77	Std. Mean Difference (IV, Ran- dom, 95% CI)	0.25 [-1.15, 1.64]
3 Activity: gait speed; short term	5	140	Mean Difference (IV, Random, 95% CI)	0.10 [-0.05, 0.24]
4 Activity: walking endurance; short term	2	73	Mean Difference (IV, Random, 95% CI)	8.43 [-12.38, 29.23]
5 Aerobic fitness; short term	3	98	Std. Mean Difference (IV, Ran- dom, 95% CI)	0.06 [-0.34, 0.45]

Exercise interventions for cerebral palsy (Review)



Analysis 5.1. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.

Study or subgroup	Aerob	ic + mixed Usual care Std. Mean Difference		Weight	Std. Mean Difference		
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% Cl
Unnithan 2007	7	33.9 (17.9)	6	30.8 (12.5)		5.83%	0.18[-0.91,1.28]
Verschuren 2007	32	1.5 (6.4)	33	-0.9 (3.5)		28.69%	0.46[-0.03,0.95]
Fowler 2010	29	70.8 (11)	29	69.3 (10.3)		26.26%	0.14[-0.38,0.65]
Bryant 2013	20	1.9 (3.6)	11	0.2 (1.8)	++	12.44%	0.53[-0.22,1.28]
Chrysagis 2012	11	71.7 (17.7)	11	65.1 (16.5)		9.79%	0.37[-0.48,1.21]
Chen 2012	13	84.2 (11.7)	14	81 (8.8)		12.08%	0.3[-0.46,1.06]
Mattern-Baxter 2013	6	16.9 (4.8)	6	13.9 (1.8)	+	4.92%	0.76[-0.43,1.95]
Total ***	118		110		•	100%	0.36[0.09,0.62]
Heterogeneity: Tau ² =0; Chi ² =1.62	, df=6(P=0.9	5); I ² =0%					
Test for overall effect: Z=2.63(P=0	0.01)						

Favours usual care -4

4 -2

4 Favours aerobic + mixed

Analysis 5.2. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 2 Activity: gross motor function, intermediate term.

Study or subgroup	Aerol	pic + mixed	Us	ual care	St	d. Mean Difference	Weight	Std. Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		Random, 95% Cl		Random, 95% Cl
Verschuren 2007	32	85.5 (16.2)	33	90.4 (10.7)			58.27%	-0.35[-0.85,0.14]
Mattern-Baxter 2013	6	34.5 (14.9)	6	21.5 (4.4)			41.73%	1.09[-0.16,2.34]
Total ***	38		39				100%	0.25[-1.15,1.64]
Heterogeneity: Tau ² =0.81; Chi ²	=4.44, df=1(P=	0.04); l ² =77.46%						
Test for overall effect: Z=0.35(P	=0.73)							
			Favo	urs usual care	-2	-1 0 1 2	Favours a	erobic + mixed

Analysis 5.3. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 3 Activity: gait speed; short term.

Study or subgroup	Aerob	oic + mixed	Us	ual care	Mean Differe	nce	Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95%	6 CI		Random, 95% CI
Fowler 2010	29	1.1 (0.3)	29	1 (0.3)	+		23.1%	0.1[-0.07,0.27]
Gharib 2011	15	0.7 (0.1)	15	0.6 (0.1)	•		32.12%	0.04[-0.03,0.11]
Smania 2011	9	1 (0.3)	9	0.8 (0.2)	+		16.92%	0.15[-0.1,0.4]
Chrysagis 2012	11	1 (0.1)	11	0.8 (0.2)	-		26.59%	0.22[0.08,0.35]
Mattern-Baxter 2013	6	0.7 (0.5)	6	2.4 (1.5)	—+—		1.28%	-1.7[-2.94,-0.46]
Total ***	70		70		•		100%	0.1[-0.05,0.24]
Heterogeneity: Tau ² =0.02; Chi	² =13.51, df=4(P	=0.01); I ² =70.4%						
Test for overall effect: Z=1.33(P=0.18)							
			Favo	urs usual care	-5 -2.5 0	2.5 5	Favours aer	obic + mixed

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Analysis 5.4. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 4 Activity: walking endurance; short term.

Study or subgroup	Aerob	Aerobic + mixed		ual care	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% CI
Fowler 2010	27	90.6 (38.4)	28	84.1 (42.6)		94.41%	6.5[-14.91,27.91]
Smania 2011	9	360 (128.7)	9	319 (39.6)		- 5.59%	41[-46.98,128.98]
Total ***	36		37		•	100%	8.43[-12.38,29.23]
Heterogeneity: Tau ² =0; Chi ² =0	0.56, df=1(P=0.46	6); I²=0%					
Test for overall effect: Z=0.79((P=0.43)						
			Favo	urs usual care	-100 -50 0 50 100	Favours aer	obic + mixed

Analysis 5.5. Comparison 5 Aerobic exercise and mixed training versus usual care, Outcome 5 Aerobic fitness; short term.

Study or subgroup	Aerol	oic + mixed	Us	ual care		Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Random, 95% CI		Random, 95% Cl
Van den Berg-Emons 1998	10	1.2 (0.8)	10	1.2 (0.9)			20.54%	0.07[-0.81,0.94]
Unnithan 2007	7	20.8 (5.9)	6	18 (4.1)		+	12.7%	0.5[-0.61,1.62]
Verschuren 2007	32	7.7 (4.1)	33	7.8 (4)			66.76%	-0.03[-0.52,0.45]
Total ***	49		49			•	100%	0.06[-0.34,0.45]
Heterogeneity: Tau ² =0; Chi ² =0.75	5, df=2(P=0.6	9); I ² =0%						
Test for overall effect: Z=0.28(P=0	0.78)							
			Favo	urs usual care	-2	-1 0 1 2	Favours a	erobic + mixed

Comparison 6. Resistance training and mixed training versus usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size	
1 Activity: gross motor function; short term	11	327	Std. Mean Difference (IV, Random, 95% CI)	0.21 [-0.01, 0.43]	
2 Activity: gross motor function; intermediate term	4	150	Std. Mean Difference (IV, Random, 95% CI)	-0.08 [-0.41, 0.24]	
3 Activity: gait speed; short term	9	243	Mean Difference (IV, Random, 95% CI)	0.03 [-0.01, 0.08]	
4 Participation; short term	3	192	Std. Mean Difference (IV, Random, 95% Cl)	0.35 [0.07, 0.64]	
5 Participation; intermediate term	2	101	Mean Difference (IV, Random, 95% CI)	0.15 [-0.24, 0.54]	
6 Muscle strength; short term 11		397	Std. Mean Difference (IV, Random, 95% Cl)	0.38 [0.01, 0.76]	

Exercise interventions for cerebral palsy (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
7 Muscle strength; intermediate term	4	149	Std. Mean Difference (IV, Random, 95% CI)	0.28 [-0.16, 0.71]

Analysis 6.1. Comparison 6 Resistance training and mixed training versus usual care, Outcome 1 Activity: gross motor function; short term.

Study or subgroup	Resista	nce + mixed	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% Cl
Dodd 2003	11	69 (21.4)	10	75.3 (21.3)		6.51%	-0.28[-1.14,0.58]
Engsberg 2006	9	69 (28.4)	3	71.4 (10)		2.83%	-0.09[-1.39,1.22]
Unnithan 2007	7	33.9 (17.9)	6	30.8 (12.5)		4.04%	0.18[-0.91,1.28]
Liao 2007	10	82.7 (2.2)	10	80.6 (2.2)	+	5.57%	0.91[-0.02,1.84]
Verschuren 2007	32	1.5 (6.4)	33	-0.9 (3.5)		19.86%	0.46[-0.03,0.95]
Seniorou 2007	11	55.6 (28)	9	60.8 (26.5)	+	6.19%	-0.18[-1.07,0.7]
Lee 2008	9	62.7 (34.1)	8	61.4 (33.9)	+	5.32%	0.04[-0.92,0.99]
Fowler 2010	29	70.8 (11)	29	69.3 (10.3)	+	18.18%	0.14[-0.38,0.65]
Scholtes 2010	24	76.1 (11.8)	24	73.1 (12.4)		14.97%	0.24[-0.32,0.81]
Chen 2012	13	84.2 (11.7)	14	81 (8.8)		8.36%	0.3[-0.46,1.06]
Lee 2015	13	81.9 (16.1)	13	81.3 (14.3)		8.17%	0.04[-0.73,0.81]
Total ***	168		159		•	100%	0.21[-0.01,0.43]
Heterogeneity: Tau ² =0; Chi ² =	5.84, df=10(P=0.8	83); I ² =0%					
Test for overall effect: Z=1.87	(P=0.06)						
			Favo	urs usual care -2	-1 0 1	² Favours re	sistance + mixed

Analysis 6.2. Comparison 6 Resistance training and mixed training versus usual care, Outcome 2 Activity: gross motor function; intermediate term.

Study or subgroup	Resista	nce + mixed	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% Cl
Dodd 2003	11	69.6 (21.4)	9	74.3 (21.4)	+	13.41%	-0.21[-1.09,0.67]
Verschuren 2007	32	85.5 (16.2)	33	90.4 (10.7)		42.97%	-0.35[-0.85,0.14]
Lee 2008	9	63 (34.4)	8	61.8 (34)		11.56%	0.03[-0.92,0.99]
Scholtes 2010	24	76.6 (13)	24	72.7 (12.8)		32.06%	0.3[-0.27,0.87]
Total ***	76		74		•	100%	-0.08[-0.41,0.24]
Heterogeneity: Tau ² =0; Chi ² =	3.03, df=3(P=0.3	9); I ² =1.07%					
Test for overall effect: Z=0.49	(P=0.62)						
			Favoi	urs usual care -2	-1 0 1	² Favours re	sistance + mixed



Analysis 6.3. Comparison 6 Resistance training and mixed training versus usual care, Outcome 3 Activity: gait speed; short term.

Study or subgroup	Resista	ance + mixed	Us	ual care	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% CI
Dodd 2003	11	0.8 (0.4)	10	0.8 (0.3)		2.03%	-0.04[-0.34,0.26]
Engsberg 2006	9	0.9 (0.3)	3	0.8 (0.3)		1.03%	0.12[-0.3,0.54]
Unger 2006	21	1.1 (0.2)	10	1.2 (0.1)	+	10.36%	-0.05[-0.18,0.08]
Liao 2007	10	1 (0.1)	10	1 (0.1)	-	29.99%	0.04[-0.04,0.12]
Seniorou 2007	11	0.3 (0.1)	9	0.3 (0.1)	-	23.52%	0[-0.09,0.09]
Lee 2008	9	0.7 (0.4)	8	0.7 (0.4)		1.2%	0.06[-0.33,0.45]
Fowler 2010	29	1.1 (0.3)	29	1 (0.3)	+	6.18%	0.1[-0.07,0.27]
Scholtes 2010	23	1 (0.3)	23	1.1 (0.4)		4.31%	-0.04[-0.25,0.17]
Pandey 2011	9	0.7 (0.1)	9	0.6 (0.1)	-+-	21.38%	0.1[0.01,0.19]
Total ***	132		111		◆	100%	0.03[-0.01,0.08]
Heterogeneity: Tau ² =0; Chi ² =	=5.64, df=8(P=0.6	9); I ² =0%					
Test for overall effect: Z=1.52	2(P=0.13)			1			
			Favo	urs usual care -1	-0.5 0 0.5	¹ Favours res	istance + mixed

Favours resistance + mixed

Analysis 6.4. Comparison 6 Resistance training and mixed training versus usual care, Outcome 4 Participation; short term.

Study or subgroup	Resista	nce + mixed	Us	ual care	Sto	l. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Random, 95% Cl		Random, 95% CI
Verschuren 2007	32	1.9 (0.7)	33	1.7 (0.6)			34.02%	0.37[-0.12,0.86]
Scholtes 2010	22	39 (13.3)	17	31.1 (13.6)			19.56%	0.57[-0.07,1.22]
Mitchell 2016	46	8 (2.3)	42	7.4 (2.5)			46.43%	0.25[-0.17,0.67]
Total ***	100		92			•	100%	0.35[0.07,0.64]
Heterogeneity: Tau ² =0; Chi ² =	0.69, df=2(P=0.7	1); I ² =0%						
Test for overall effect: Z=2.42	(P=0.02)						1	
			Favo	urs usual care -2	-1	0 1	² Favours re	sistance + mixed

Analysis 6.5. Comparison 6 Resistance training and mixed training versus usual care, Outcome 5 Participation; intermediate term.

Study or subgroup	Resista	nce + mixed	Us	ual care		Меа	an Difference		Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	ndom, 95% CI			Random, 95% Cl
Verschuren 2007	32	1.8 (0.9)	33	1.7 (0.7)			+		99.68%	0.15[-0.25,0.54]
Scholtes 2010	19	32.2 (9.3)	17	31.8 (11.8)					0.32%	0.37[-6.61,7.35]
Total ***	51		50				•		100%	0.15[-0.24,0.54]
Heterogeneity: Tau ² =0; Chi ² =	Heterogeneity: Tau ² =0; Chi ² =0, df=1(P=0.95); I ² =0%									
Test for overall effect: Z=0.74	(P=0.46)									
			Favo	urs usual care	-10	-5	0	5 10	Favours res	istance + mixed



Analysis 6.6. Comparison 6 Resistance training and mixed training versus usual care, Outcome 6 Muscle strength; short term.

Study or subgroup	Resista	nce + mixed	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% Cl
Dodd 2003	11	33.1 (15.8)	10	25.5 (9.9)		8.3%	0.55[-0.33,1.42]
Liao 2007	10	6.1 (1.3)	10	6.2 (1.3)		8.29%	-0.08[-0.95,0.8]
Verschuren 2007	32	37.4 (18.8)	33	38.5 (19.4)	+	12.11%	-0.05[-0.54,0.43]
Seniorou 2007	11	1.3 (0.5)	9	1.2 (0.5)	-+-	8.23%	0.19[-0.69,1.07]
Lee 2008	9	13.2 (5.4)	8	14.1 (5.8)	-+-	7.65%	-0.15[-1.11,0.8]
Reid 2010	7	184.7 (15.3)	7	211.8 (116.7)	_• <u> </u> -	6.88%	-0.3[-1.36,0.75]
Scholtes 2010	24	5.4 (1.1)	24	4.5 (1.2)	+	11.06%	0.77[0.19,1.36]
Fowler 2010	28	0.9 (0.3)	29	0.9 (0.4)	+	11.77%	0.07[-0.44,0.59]
Pandey 2011	9	6.3 (1.1)	9	2.7 (0.5)	— — • —	3.59%	3.95[2.22,5.67]
Chen 2012	13	1.6 (0.8)	15	1.4 (0.6)		9.43%	0.41[-0.34,1.16]
Mitchell 2016	46	63.5 (26)	43	46.8 (18.3)	+	12.68%	0.73[0.3,1.16]
Total ***	200		197		•	100%	0.38[0.01,0.76]
Heterogeneity: Tau ² =0.24; Chi	i²=29.2, df=10(P	=0); I ² =65.75%					
Test for overall effect: Z=2.01(P=0.04)						
			Favo	urs usual care ⁻¹⁰	-5 0 5	¹⁰ Favours re	sistance + mixed

Analysis 6.7. Comparison 6 Resistance training and mixed training versus usual care, Outcome 7 Muscle strength; intermediate term.

Study or subgroup	Resista	nce + mixed	Us	ual care	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% CI
Dodd 2003	11	32.5 (11.4)	9	25.2 (7.8)	++	16.96%	0.7[-0.21,1.62]
Verschuren 2007	32	37.4 (18.8)	33	38.5 (19.4)		36.81%	-0.05[-0.54,0.43]
Lee 2008	9	13.7 (5.9)	8	14.4 (5.9)		15.9%	-0.11[-1.07,0.84]
Scholtes 2010	23	5.2 (1)	24	4.5 (1.2)		30.33%	0.65[0.06,1.23]
Total ***	75		74		•	100%	0.28[-0.16,0.71]
Heterogeneity: Tau ² =0.07; Cl	hi²=4.74, df=3(P=	0.19); I ² =36.76%					
Test for overall effect: Z=1.25	5(P=0.21)						
			Favo	urs usual care	-2 -1 0 1 2	Favours re	sistance + mixed

ADDITIONAL TABLES

Table 1. Additional methods table

Binary data	We planned to present the relative risk (or risk ratio) with a 95% confidence interval, and calculate the number needed to treat for an additional beneficial outcome as an absolute measure of treat- ment effect. We will report the odds ratio (OR) with a 95% confidence interval in future updates of this review, as most studies with a dichotomous outcome report the OR.
Cluster trials	We planned to seek direct estimates of the effect from an analysis that accounted for cluster de- sign. Where the analysis in a cluster trial did not account for the cluster design, we planned to use the approximately correct analysis approach, presented in the <i>Cochrane Handbook for Systematic</i> <i>Reviews of Interventions</i> (Higgins 2011c).

Exercise interventions for cerebral palsy (Review)

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Table 1. Additional methods table (Continued)

Crossover trials	Where studies presented repeated measurements over time, we planned to only include data from 1 time point from an individual study in any single meta-analysis. If inadequate data were available to conduct this analysis, we planned to only include data from the first phase of the cross-over tri- al, as if it were from a parallel trial design. We planned to combine the results of cross-over studies with those of parallel studies by imputing the post-treatment, between-condition correlation coef- ficient from an included study that presents individual participant data, and use this to calculate the standard error of the SMD, using the generic inverse-variance method.
Assessment of reporting bi- ases	Where we identified evidence of publication bias, we planned to consider its likely influence on the observed effect sizes in our interpretation of the results. However, as common tests of publication bias lack sensitivity, we planned to consider the possible influence that a dominance of small trials might have on pooled effect sizes in our interpretation.
Subgroup analysis and iden- tification of heterogeneity	We planned to further explore possible clinical heterogeneity through preplanned subgroup analy- sis based on important clinical features. We predicted that some trials would include ambulatory participants only (i.e. people who could walk with or without a mobility aid; GMFCS level I, II, and III), and some studies would include participants who could walk without a mobility aid only (i.e. GMFCS level I and II). Where adequate data allowed, we planned to undertake 2 subgroup analyses for studies that included ambulatory people only (i.e. GMFCS level I, II, and III), and for studies that included ambulatory people who walk without a mobility aid only (i.e. GMFCS level I and II).
Sensitivity analysis	We planned to explore the impact of studies at high risk of bias by reanalysis after excluding stud- ies rated at overall high risk of bias. We also planned to explore the impact of excluding studies at high risk of bias for missing data through reanalysis. We planned to explore the influence of using imputed correlation coefficients in our approach to including cross-over and cluster trials by re- analysing these data with adjusted (higher and lower) coefficient values.

GMFCS: Gross Motor Function Classification System; SMD: standardised mean difference.

APPENDICES

Appendix 1. Gross Motor Function Classification System

Level I: walks without restrictions; limitations in more advanced gross motor skills

Before 2nd birthday: infants move in and out of sitting and floor sit with both hands free to manipulate objects. Infants crawl on hands and knees, pull to stand, and take steps holding onto furniture. Infants walk between 18 months and 2 years of age without the need for any assistive mobility device.

From age 2 to 4th birthday: children floor sit with both hands free to manipulate objects. Children perform movements in and out of floor sitting and standing without adult assistance. Children walk as the preferred method of mobility without the need for any assistive mobility device.

From age 4 to 6th birthday: children get into and out of and sit in a chair without the need for hand support. Children move from floor and chair sitting to standing without the need for objects for support. Children walk indoors and outdoors and climb stairs. Emerging ability to run and jump.

From age 6 to 12th birthday: children walk at home, school, outdoors and in the community. Children are able to walk up and down curbs without physical assistance and stairs without the use of a railing. Children perform gross motor skills, such as running and jumping, but speed, balance, and coordination are limited. Children may participate in physical activities and sports depending on personal choices and environmental factors.

From age 12: youth walks at home, school, outdoors and in the community. Youth is able to walk up and down curbs without physical assistance and stairs without the use of a railing. Youth performs gross motor skills, such as running and jumping, but speed, balance, and coordination are limited. Youth may participate in physical activities and sports depending on personal choices and environmental factors.

Level II: walks without assistive devices; limitations walking outdoors and in the community

Before 2nd birthday: infants maintain floor sitting but may need to use their hands for support to maintain balance. Infants creep on their stomach or crawl on hands and knees. Infants may pull to stand and take steps holding onto furniture.

From age 2 to 4th birthday: children floor sit but may have difficulty with balance when both hands are free to manipulate objects. Children perform movements in and out of sitting without adult assistance. Children pull to stand on a stable surface. Children crawl on hands and knees with a reciprocal pattern, cruise holding onto furniture and walk using an assistive mobility device as preferred methods of mobility.

From age 4 to 6th birthday: children sit in a chair with both hands free to manipulate objects. Children move from the floor to standing and from chair sitting to standing but often require a stable surface to push or pull up on with their arms. Children walk without needing any assistive mobility device indoors and for short distances on level surfaces outdoors. Children climb stairs holding onto a railing but are unable to run or jump.

From age 6 to 12th birthday: Children walk in most settings. Children may experience difficulty walking long distances and balancing on uneven terrain, on inclines, in crowded areas, in confined spaces, or when carrying objects. Children walk up and down stairs holding onto a railing or with physical assistance if there is no railing. Outdoors and in the community, children may walk with physical assistance or a hand-held mobility device or use wheeled mobility when travelling long distances. Children have at best only minimal ability to perform gross motor skills such as running and jumping. Limitations in performance of gross motor skills may necessitate adaptations to enable participation in physical activities and sports.

From age 12: youth walk in most settings. Environmental factors (such as uneven terrain, inclines, long distances, time demands, weather, and peer acceptability) and personal preference influence mobility choices. At school or work, youth may walk using a hand-held mobility device for safety. Outdoors and in the community, youth may use wheeled mobility when travelling long distances. Youth walk up and down stairs holding a railing or with physical assistance if there is no railing. Limitations in performance of gross motor skills may necessitate adaptations to enable participation in physical activities and sports.

Distinctions between levels I and II

Compared with children at level I, children at level II have limitations in the ease of performing movement transitions; walking outdoors and in the community; the need for assistive mobility devices when beginning to walk; quality of movement; and the ability to perform gross motor skills such as running and jumping.

Level III: walks with assistive mobility devices; limitations walking outdoors and in the community

Before 2nd birthday: infants maintain floor sitting when the low back is supported. Infants roll and creep forward on their stomachs.

From age 2 to 4th birthday: children maintain floor sitting often by 'W-sitting' (sitting between flexed and internally rotated hips and knees) and may require adult assistance to assume sitting. Children creep on their stomach or crawl on hands and knees (often without reciprocal leg movements) as their primary methods of self-mobility. Children may pull to stand on a stable surface and cruise short distances. Children may walk short distances indoors using an assistive mobility device and adult assistance for steering and turning.

From age 4 to 6th birthday: children sit on a regular chair but may require pelvic or trunk support to maximise hand function. Children move in and out of chair sitting using a stable surface to push on or pull up with their arms. Children walk with an assistive mobility device on level surfaces and climb stairs with adult assistance. Children are frequently transported when travelling for long distances or outdoors on uneven terrain.

From age 6 to 12th birthday: children walk using a hand-held mobility device in most indoor settings. When seated, children may require a seat belt for pelvic alignment and balance. Sit-to-stand and floor-to-stand transfers require physical assistance of a person or support surface. When travelling long distances, children use some form of wheeled mobility. Children may walk up and down stairs holding onto a railing with supervision or physical assistance. Limitations in walking may necessitate adaptations to enable participation in physical activities and sports, including a self-propelling manual wheelchair or powered mobility.

From age 12: youth is capable of walking using a hand-held mobility device. In comparison with individuals at other levels, young people at level III demonstrate more variability in methods of mobility depending on physical ability and environmental and personal factors. When seated, youth may require a seat belt for pelvic alignment and balance. Sit-to-stand and floor-to-stand transfers require physical assistance from a person or support surface. At school, youth may self-propel a manual wheelchair or use powered mobility. Outdoors and in the community, young people are transported in a wheelchair or use powered mobility. Youth may walk up and down stairs holding onto a railing with supervision or physical assistance. Limitations in walking may necessitate adaptations to enable participation in physical activities and sports, including self-propelling a manual wheelchair or powered mobility.

Distinctions between levels II and III

Differences are seen in the degree of achievement of functional mobility. Children at level III need assistive mobility devices and frequently orthoses to walk, while children at level II do not require assistive mobility devices after age 4.

Exercise interventions for cerebral palsy (Review)

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Level IV: self-mobility with limitations; children are transported or use power mobility outdoors and in the community

Before 2nd birthday: infants have head control but require trunk support for floor sitting. Infants can roll to supine and may roll to prone.

From age 2 to 4th birthday: children floor sit when placed but are unable to maintain alignment and balance without using their hands for support. Children frequently require adaptive equipment for sitting and standing. Children achieve self-mobility for short distances (within a room) through rolling, creeping on stomach, or crawling on hands and knees without reciprocal leg movement.

From age 4 to 6th birthday: children sit on a chair but need adaptive seating for trunk control and to maximise hand function. Children move in and out of chair sitting with assistance from an adult or a stable surface to push or pull up on with their arms. Children may at best walk short distances with a walker and adult supervision but have difficulty turning and maintaining balance on uneven surfaces. Children are transported in the community. Children may achieve self-mobility using a power wheelchair.

From age 6 to 12th birthday: children use methods of mobility that require physical assistance or powered mobility in most settings. Children require adaptive seating for trunk and pelvic control and physical assistance for most transfers. At home, children use floor mobility (roll, creep, or crawl), walk short distances with physical assistance, or use powered mobility. When positioned, children may use a body support walker at home or school. At school, outdoors, and in the community, children are transported in a manual wheelchair or use powered mobility. Limitations in mobility necessitate adaptations to enable participation in physical activities and sports, including physical assistance, powered mobility or both.

From age 12: youth uses wheeled mobility in most settings. Youth requires adaptive seating for pelvic and trunk control. Youth requires physical assistance from one or two people for transfers. Youth may support weight with their legs to assist with standing transfers. Indoors, youth may walk short distances with physical assistance, use wheeled mobility, or, when positioned, use a body support walker. Youth are physically capable of operating a powered wheelchair. When a powered wheelchair is not feasible or available, youth are transported in a manual wheelchair. Limitations in mobility necessitate adaptations to enable participation in physical activities and sports, including physical assistance or powered mobility, or both.

Distinctions between levels III and IV

Differences in sitting ability and mobility exist, even allowing for extensive use of assistive technology. Children at level III sit independently, have independent floor mobility and walk with assistive mobility devices. Children at level IV function in sitting (usually supported), but independent mobility is very limited. Children at level IV are more likely to be transported or to use power mobility.

Level V: self-mobility is severely limited even with the use of assistive technology

Before 2nd birthday: physical impairments limit voluntary control of movement. Infants are unable to maintain antigravity head and trunk postures in prone and sitting. Infants require adult assistance to roll.

From age 2 to 12th birthday: Children are transported in a manual wheelchair in all settings. Children are limited in their ability to maintain antigravity head and trunk postures and control arm and leg movements. Assistive technology is used to improve head alignment, seating, standing, and/or mobility, but limitations are not fully compensated for by equipment. Transfers require complete physical assistance of an adult. At home, children may move short distances on the floor or may be carried by an adult. Children may achieve self-mobility using powered mobility with extensive adaptations for seating and control access. Limitations in mobility necessitate adaptations to enable participation in physical activities and sports, including physical assistance and using powered mobility.

From age 12: youth are transported in a manual wheelchair in all settings. Youth are limited in their ability to maintain antigravity head and trunk postures and control arm and leg movements. Assistive technology is used to improve head alignment, seating, standing, and mobility, but limitations are not fully compensated for by equipment. Transfers require physical assistance from one or two people or a mechanical lift. Youth may achieve self-mobility using powered mobility with extensive adaptations for seating and control access. Limitations in mobility necessitate adaptations to enable participation in physical activities and sports, including physical assistance and using powered mobility.

Distinctions between levels IV and V

Children at level V lack independence even in basic antigravity postural control. Child achieves self-mobility only if he or she can learn how to operate an electrically powered wheelchair.

Appendix 2. Manual Ability Classification System

Level I: handles objects easily and successfully. At most, limited in the ease of performing manual tasks requiring speed and accuracy. However, any limitations in manual abilities do not restrict independence in daily activities.

Level II: handles most objects but with somewhat reduced quality or speed of achievement, or both. May avoid or achieve with some difficulty certain activities; might use alternative ways of performance, but manual abilities do not usually restrict independence in daily activities.

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Exercise interventions for cerebral palsy (Review)



Level III: handles objects with difficulty; needs help to prepare or modify activities, or both. The performance is slow and achieved with limited success regarding quality and quantity. Performs activities independently if they have been set up or adapted.

Level IV: handles a limited selection of easily managed objects in adapted situations. Performs parts of activities with effort and with limited success. Requires continuous support and assistance, adapted equipment or both, for even partial achievement of the activity.

Level V: does not handle objects and has severely limited ability to perform even simple actions. Requires total assistance.

Distinctions between levels I and II. Children at level I may have limitations in handling very small, heavy or fragile objects, which demand detailed fine motor control or efficient coordination between hands. Limitations may also involve performance in new and unfamiliar situations. Children at level II perform almost the same activities as children at level I, but the quality of performance is decreased or the performance is slower. Functional differences between hands can limit effectiveness of performance. Children at level II commonly try to simplify handling of objects, for example by using a surface for support instead of handling objects with both hands.

Distinctions between Levels II and III. Children at level II handle most objects, although slowly or with reduced quality of performance. Children at level III commonly need help to prepare the activity or require that adjustments be made to the environment, or both, since their ability to reach or handle objects is limited. They cannot perform certain activities and their degree of independence is related to the supportiveness of the environmental context.

Distinctions between Levels III and IV. Children at level III can perform select activities if the situation is prearranged and if they receive supervision and plenty of time. Children at level IV need continuous help during the activity and can at best participate meaningfully in only parts of an activity.

Distinctions between Levels IV and V. Children at level IV perform part of an activity with continuous help. Children at level V might at best participate with a simple movement in special situations, for example by pushing a button.

Appendix 3. Search strategies

Cochrane Central Register of Controlled Trials (CENTRAL) in the Cochrane Library, which contains the Cochrane Developmental, Psychosocial and Learning Problems Specialised Register

#1[mh "cerebral palsy"] #2cerebral next pals* #3((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/5 spastic*) #4((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/3 ataxi*) #5"Little* disease" #6{or #1-#5} #7[mh exercise] #8[mh "Exercise Movement Techniques"] #9[mh "Exercise Therapy"] #10[mh ^"Physical Education and Training"] #11[mh "Physical Endurance"] #12[mh ^"Physical Fitness"] #13[mh Sports] #14[mh Hydrotherapy] #15[mh "Equine-Assisted Therapy"] #16aerobic' #17(cycle or cycling) #18[mh ergometry] #19ergometry #20(treadmill or tread next mill) #21(exercise* or strength* or fitness) #22(flexibility or stretching) #23((weight* near/1 lift*) or weight next training) #24(hippotherapy or equine* therapy or equine assist* or horse*) #25sport* #26(walking or running) #27(aquatic* or swim*) #28[mh "Physical Therapy Modalities"] #29(physiotherapy or physical next therap*) #30(resistance or resisted) #31physical next activit* #32{or #7-#31} #33#6 and #32 in Trials



Ovid MEDLINE(R)

This strategy uses the Cochrane highly sensitive search strategy for identifying randomised trials in Lines 33 to 43 (Lefebvre 2011).

- 1 cerebral palsy/ 2 cerebral pals\$.tw. 3 ((Hemiplegi\$ or diplegi\$ or quadriplegi\$ or unilateral\$) adj5 spastic\$).tw. 4 ((Hemiplegi\$ or diplegi\$ or quadriplegi\$ or unilateral\$) adj3 ataxi\$).tw. 5 Little\$ disease.tw. 6 or/1-5 7 exp Exercise/ 8 exp Exercise Movement Techniques/ 9 exp Exercise Therapy/ 10 exp "Physical Education and Training"/ 11 Physical Endurance/ 12 Physical Fitness/ 13 exp Sports/ 14 Hydrotherapy/ 15 Equine-Assisted Therapy/ 16 aerobic\$.tw. 17 (cycle or cycling).tw. 18 ergometry.tw. 19 (treadmill or tread-mill).tw. 20 ergometry/ 21 (exercise\$ or strength\$ or fitness).tw. 22 (flexibility or stretching).tw. 23 ((weight\$ adj1 lift\$) or weight training).tw. 24 (hippotherapy or equine\$ therapy or equine assist\$ or horse\$).tw. 25 sport\$.tw. 26 (walking or running).tw. 27 (aquatic\$ or swim\$).tw. 28 Physical Therapy Modalities/ 29 (physiotherapy or physical therap\$).tw. 30 (resistance or resisted).tw. 31 physical activit\$.tw. 32 or/7-31 33 randomized controlled trial.pt. 34 controlled clinical trial.pt. 35 randomi#ed.ab. 36 placebo\$.ab. 37 drug therapy.fs. 38 randomly.ab. 39 trial.ab. 40 groups.ab. 41 or/33-40 42 exp animals/ not humans.sh. 43 41 not 42 44 6 and 32 and 43 **Embase Ovid**
- 1 cerebral palsy/ 2 cerebral pals\$.tw. 3 ((Hemiplegi\$ or diplegi\$ or quadriplegi\$ or unilateral\$) adj5 spastic\$).tw. 4 ((Hemiplegi\$ or diplegi\$ or quadriplegi\$ or unilateral\$) adj3 ataxi\$).tw. 5 Little\$ disease.tw. 6 or/1-5 7 exp exercise/ 8 exp kinesiotherapy/ 9 physical education/ 10 endurance/ 11 fitness/ 12 exp sport/



50 32 and 49

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13 hydrotherapy/ 14 hippotherapy/ 15 aerobic\$.tw. 16 (cycle or cycling).tw. 17 (treadmill or tread-mill).tw. 18 ergometry/ 19 ergometry.tw. 20 (exercise\$ or strength\$ or fitness).tw. 21 (flexibility or stretching).tw. 22 ((weight\$ adj1 lift\$) or weight training).tw. 23 (hippotherapy or equine\$ therapy or equine assist\$ or horse\$).tw. 24 sport\$.tw. 25 (walking or running).tw. 26 (aquatic\$ or swim\$).tw. 27 exp physiotherapy/ 28 (physiotherapy or physical therap\$).tw. 29 physical activit\$.tw. 30 (resistance or resisted).tw. 31 or/7-30 32 6 and 31 33 Randomized controlled trial/ 34 controlled clinical trial/ 35 Single blind procedure/ 36 Double blind procedure/ 37 triple blind procedure/ 38 Crossover procedure/ 39 (crossover or cross-over).tw. 40 ((singl\$ or doubl\$ or tripl\$ or trebl\$) adj1 (blind\$ or mask\$)).tw. 41 Placebo/ 42 placebo.tw. 43 prospective.tw. 44 factorial\$.tw. 45 random\$.tw. 46 assign\$.ab. 47 allocat\$.tw. 48 volunteer\$.ab. 49 or/33-48

CINAHL Plus EBSCOhost (Cumulative Index to Nursing and Allied Health Literature)

S48 S32 AND S47 S47 S33 OR S34 OR S35 OR S36 OR S37 OR S38 OR S39 OR S40 OR S41 OR S42 OR S43 OR S44 OR S45 OR S46 S46 (MH "Treatment Outcomes") S45 (MH "Program Evaluation") S44 TI ("prospective study" or "prospective research") or AB("prospective study" or "prospective research") S43 TI ("follow-up study" or "follow-up research") or AB ("follow-up study" or "follow-up research") S42 AB((trebl* N1 mask*) or (trebl* N1 blind*)) S41 AB("cross over" or crossover) S40 (MH "Crossover Design") S39 AB((tripl* N1 mask*) or (tripl* N1 blind*)) S38 AB ((doubl* N1 mask*) or (doubl* N1 blind*)) S37 AB ((singl* N1 mask*) or(singl* N1 blind*)) S36 AB(trial) S35 AB(random*) S34 (MH "Random Assignment") S33 (MH "Clinical Trials+") S32 S6 AND S31 S31 S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14 OR S15 OR S16 OR S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24 OR S25 OR S26 OR S27 OR S28 OR S29 OR S30 S30 physical activit* S29 (resistance or resisted)

S28 (physiotherapy or physical therap*) S27 (MH "Physical Therapy+") S26 (aquatic* or swim*) S25 (walking or running) S24 sport* S23 (hippotherapy or equine* therapy or equine assist* or horse*) S22 (weight* N1 lift*) or weight training S21 (flexibility or stretching) S20 (exercise* or strength* or fitness) S19 (treadmill or tread-mill) S18 (cycle or cycling) S17 ergometry S16 (MH "Ergometry") S15 (MH "Sports+") S14 (MH "Physical Activity") S13 (MH "Physical Fitness+") S12 (MH "Physical Endurance+") S11 (MH "Physical Education and Training+") S10 (MH "Sports+") S9 (MH "Physical Fitness+") S8 (MH "Leisure Activities+") S7 (MH "Exercise+") S6 S1 OR S2 OR S3 OR S4 OR S5 S5 "Little* disease" S4 ((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) N3 ataxi*) S3 ((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) N5 spastic*) S2 cerebral pals* S1 (MH "Cerebral Palsy")

Cochrane Database of Systematic Reviews (CDSR), part of the Cochrane Library

#1[mh "cerebral palsy"] #2(cerebral next pals*):ti,ab #3((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/5 spastic*):ti,ab 251 #4((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/3 ataxi*):ti,ab #5("Little* disease"):ti,ab #6{or #1-#5} #7[mh exercise] #8[mh "Exercise Movement Techniques"] #9[mh "Exercise Therapy"] #10[mh ^"Physical Education and Training"] #11[mh "Physical Endurance"] #12[mh ^"Physical Fitness"] #13[mh Sports] #14[mh Hydrotherapy] #15[mh "Equine-Assisted Therapy"] #16aerobic*:ti,ab #17(cycle or cycling):ti,ab #18[mh ergometry] #19ergometry:ti,ab #20(treadmill or tread next mill):ti,ab #21(exercise* or strength* or fitness):ti,ab #22(flexibility or stretching):ti,ab #23((weight* near/1 lift*) or weight next training):ti,ab #24(hippotherapy or equine* next therapy or equine next assist* or horse*):ti,ab #25sport*:ti,ab #26(walking or running):ti,ab #27(aquatic* or swim*):ti,ab #28[mh "Physical Therapy Modalities"] #29(physiotherapy or physical next therap*):ti,ab #30(resistance or resisted):ti,ab #31(physical next activit*):ti,ab



#32{or #7-#31} #33#6 and #32 in Cochrane Reviews (Reviews and Protocols)

Database of Abstracts of Reviews of Effects (DARE), part of the Cochrane Library

#1[mh "cerebral palsy"] #2(cerebral next pals*):ti,ab #3((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/5 spastic*):ti,ab #4((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) near/3 ataxi*):ti,ab #5("Little* disease"):ti,ab #6{or #1-#5} #7[mh exercise] #8[mh "Exercise Movement Techniques"] #9[mh "Exercise Therapy"] #10[mh ^"Physical Education and Training"] #11[mh "Physical Endurance"] #12[mh ^"Physical Fitness"] #13[mh Sports] #14[mh Hydrotherapy] #15[mh "Equine-Assisted Therapy"] #16aerobic*:ti,ab #17(cycle or cycling):ti,ab #18[mh ergometry] #19ergometry:ti,ab #20(treadmill or tread next mill):ti,ab #21(exercise* or strength* or fitness):ti,ab #22(flexibility or stretching):ti,ab #23((weight* near/1 lift*) or weight next training):ti,ab #24(hippotherapy or equine* next therapy or equine next assist* or horse*):ti,ab #25sport*:ti,ab #26(walking or running):ti,ab #27(aquatic* or swim*):ti,ab #28[mh "Physical Therapy Modalities"] #29(physiotherapy or physical next therap*):ti,ab #30(resistance or resisted):ti,ab #31(physical next activit*):ti,ab #32{or #7-#31} #33#6 and #32 in Other Reviews

Science Citation Index Web of Science

#14 #13 AND #12 # 13 TS=(random* or control NEAR/1 group* or assign* or allocat*) # 12 #11 AND #5 # 11 #10 OR #9 OR #8 OR #7 OR #6 # 10 TS= ("weight* lift*" or "weight training") #9 TS= (physiotherapy or "physical therap*" or "physical education") #8 TS=(aerobic* or aquatic* or cycle or cycling or ergometry or swim* or running or walking or treadmill or "tread mill" or sport* or horse* or equine* or hippotherapy) # 7 TS=(flexibility or stretching) #6 Ts=(exercise* or strength* or fitness) # 5 #4 OR #3 OR #2 OR #1 # 4 TS= "littles disease" # 3 TS=((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) NEAR/3 ataxi*) # 2 TS=((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) NEAR/5 spastic*) #1TS=(cerebral pals*)

Conference Proceedings Citation Index - Science Web of Science

#14 #13 AND #12 # 13 TS=(random* or control NEAR/1 group* or assign* or allocat*) # 12 #11 AND #5 # 11 #10 OR #9 OR #8 OR #7 OR #6 # 10 TS= ("weight* lift*" or "weight training")



#9 TS= (physiotherapy or "physical therap*" or "physical education")

- #8 TS=(aerobic* or aquatic* or cycle or cycling or ergometry or swim* or running or walking or treadmill or "tread mill" or sport* or horse* or equine* or hippotherapy)
- # 7 TS=(flexibility or stretching)
- # 6 Ts=(exercise* or strength* or fitness)
- # 5 #4 OR #3 OR #2 OR #1
- # 4 TS= "littles disease"
- # 3 TS=((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) NEAR/3 ataxi*)
- # 2 TS=((Hemiplegi* or diplegi* or quadriplegi* or unilateral*) NEAR/5 spastic*)
- # 1 TS=(cerebral pals*)

LILACS (Latin American and Caribbean Health Science Information database)

(lilacs.bvsalud.org/en)

(tw:(cerebral pals*) OR mh:("Cerebral Palsy") OR tw:(littles disease)) AND (instance: "regional") AND (db:("LILACS") AND type_of_study: ("clinical_trials"))

Health Services Research Projects in Progress (HSRPRoj)

(wwwcf.nlm.nih.gov/hsr_project/home_proj.cfm)

We searched for the following words and phrases individually and then manually de-duplicated the records:

cerebral pals*; ataxi*; hemiplegi*; diplegi*; quadriplegi*; unilateral*; Little's disease; spastic*

OpenGrey

(www.opengrey.eu)

cerebral pals* OR hemiplegi* OR diplegi* OR quadriplegi*

National Rehabilitation Information Center

(www.naric.com)

We searched for the following words and phrases individually and then manually de-duplicated the records:

#1 Cerebral palsy #2 hemiplegia OR diplegia OR quadriplegia OR unilateral #3 spastic #4 ataxic

PEDro (Physiotherapy Evidence Database)

www.pedro.org.au

cerebral pals*

UKCRN Study Portfolio

(public.ukcrn.org.uk/search)

The following searches were conducted individually and the records were de-duplicated manually: #1 cerebral palsy #2 cerebral pals* #3 cerebral pals\$ #4 cerebral palsy OR hemiplegia #5 hemiplegia #6 diplegia #7 quadriplegia #8 ataxic #9 spastic #10 little disease

ClinicalTrials.gov

(clinicaltrials.gov)

cerebral palsy

World Health Organization International Clinical Trials Registry Platform (WHO ICTRP)

(www.who.int/ictrp/en)

We searched for the following words and phrases individually and then manually de-duplicated the records:

#1 cerebral palsy (in the condition field)
#2 cerebral palsy (in the title field)
#3 hemiplegia OR diplegia OR quadriplegia OR unilateral OR ataxia OR spastic (in the condition)
#4 hemiplegia OR diplegia OR quariplegia OR unilateral OR ataxia OR spastic (in the title)

Exercise interventions for cerebral palsy (Review)

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#5 Little disease (in the title field) #6 Little disease (in the condition)

Appendix 4. Record of searches

Database	Search date	Database date range/is- sue/volume	Number of records	Limits
Cochrane Central Register of Controlled	7 May 2015	2015 Issue 4	76	_
Trials (CENTRAL), in the Cochrane Library	14 June 2016	2016, Issue 5	0	Publication year from 2015 to 2016
MEDLINE Ovid	7 May 2015	1946 to May Week 1 2015	954	_
	14 June 2016	1946 to June Week 1 2016	103	ed=20150501-201606
Embase Ovid	7 May 2015	1980 to 2015 Week 18	1095	_
	14 June 2016	1980 to 2016 Week 24	172	em=201519-201624
CINAHL Plus EBSCOhost (Cumulative In-	7 May 2015	1937 to 7 May 2015	819	_
dex to Nursing and Allied Health Litera- ture)	14 June 2016	1937 to 14 June 2016	65	EM = 201504-
Cochrane Database of Systematic Reviews	7 May 2015	2015 Issue 5	2	_
(CDSR), part of the Cochrane Library	14 June 2016	2016 Issue 6	0	_
Database of Abstracts of Reviews of Ef- fects (DARE), part of the Cochrane Library	7 May 2015	2015 Issue 2	11	_
Tects (DARE), part of the Cochrane Library	Not searched	No new content after 2015 Issue 2	Not searched	Not searched
Science Citation Index Web of Science	7 May 2015	1970 to 6 May 2015	730	_
	14 June 2016	1970 to 9 June 2016	160	2015-2016
Conference Proceedings Citation Index - Science Web of Science	7 May 2015	1990 to 6 May 2015	36	_
	14 June 2016	1990 to 9 June 2016	5	2015-2016
LILACS (Latin American and Caribbean Health Science Information database;	7 May 2015	All available years	14	_
lilacs.bvsalud.org/en)	14 June 2016	All available years	4	Deduplicated with previous records
Health Services Research Projects in	13 May 2015	All available years	103	_
Progress (HSRPRoj; wwwcf.nlm.ni- h.gov/hsr_project/home_proj.cfm)	23 June 2016	All available years	137	_
OpenGrey (www.opengrey.eu)	6 May 2015	All available years	260	_

Exercise interventions for cerebral palsy (Review)

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(Continued)

(continued)	16 June 2016	All available years	0	Deduplicated with previous records
National Rehabilitation Information Cen- ter (www.naric.com)	13 May 2015	All available years	1564	_
(www.nanc.com)	23 June 2016	All available years	417	_
PEDro (Physiotherapy Evidence Data-	15 May 2015	All available years	320	_
base; www.pedro.org.au)	17 June 2016	All available years	59	_
UKCRN Study Portfolio (pub- lic.ukcrn.org.uk/search)	6 May 2015	All available years	41	_
ic.ukciii.org.uk/searchj	16 May 2016	All available years	21	_
ClinicalTrials.gov (www.clinicaltrials.gov)	6 May 2015	All available years	890	_
	16 June 2016	All available years	112	_
World Health Organization International Clinical Trials Registry Platform (WHO IC-	10 June 2015	All available years	1998	_
TRP; www.who.int/ictrp/en)	20 June 2016	All available years	883	_
Total records			11,100	

Appendix 5. Risk of bias criteria: operational definitions

Adequate sequence generation?

- 1. Low risk of bias: based on a random component judged to be both appropriate and sufficiently well described.
- 2. High risk of bias: based on any non-random component.
- 3. Unclear risk of bias: insufficient information regarding sequence generation process to permit judgement of low or high risk of bias.

Adequate allocation concealment?

- 1. Low risk of bias: method of concealment allocation employed prohibited foresight of participant assignment.
- 2. High risk of bias: method of concealment allocation employed permitted possible foresight of participant assignment.
- 3. Unclear risk of bias: method of concealment allocation not described or described in insufficient detail to permit judgement of low or high risk of bias.

Blinding of outcome assessment?

- 1. Low risk of bias: outcome assessor (including participants with respect to self-reported outcomes) blinded to participants' allocated intervention, and unlikely that blinding broken; OR no or incomplete blinding but judged that a given outcome unlikely to be influenced by lack of blinding.
- 2. High risk of bias: outcome assessor (including participants with respect to self-report outcomes) unblinded to participants' allocated intervention; OR outcome assessor blinded to allocated intervention but likely that blinding may have been broken (and a given outcome is likely to be influenced by lack of blinding).
- 3. Unclear risk of bias: insufficient information to permit judgement of low or high risk of bias.

Blinding of participants?

- 1. Low risk of bias: participants blinded to allocated intervention and unlikely that blinding broken; OR no or incomplete blinding but judged that a given outcome unlikely to be influenced by lack of blinding.
- 2. High risk of bias: participants not blinded to allocated intervention; OR participants blinded to allocated intervention but likely that blinding may have been broken (and a given outcome is likely to be influenced by lack of blinding).
- 3. Unclear risk of bias: insufficient information to permit judgement of low or high risk of bias.

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Blinding of care provider?

- 1. Low risk of bias: care provider blinded to allocated intervention and unlikely that blinding broken; OR no or incomplete blinding but judged that a given outcome unlikely to be influenced by lack of blinding.
- 2. High risk of bias: care provider not blinded to allocated intervention; OR care provider blinded to allocated intervention but likely that blinding may have been broken (and a given outcome is likely to be influenced by lack of blinding).
- 3. Unclear risk of bias: insufficient information to permit judgement of low or high risk of bias.

Incomplete outcome data addressed?

- 1. Low risk of bias: no missing outcome data. Reasons for missing data unlikely to be related to the true outcome. Missing outcome data balanced across intervention groups with similar reasons for omissions. Dichotomous outcomes: proportion of missing outcomes compared with observed event risk not enough to have a clinically relevant impact on the intervention effect estimate. Continuous outcomes: difference in means or standardised mean difference among missing outcomes not enough to have a clinically relevant impact on observed effect size. Missing data imputed using appropriate methods. Intention-to-treat analysis undertaken. Less than or equal to 10% dropout rate.
- 2. High risk of bias: reason for missing outcome data likely to be related to the true outcome. Dichotomous outcomes: proportion of missing outcomes compared with observed event risk enough to induce a clinically relevant bias in intervention effect estimate. Continuous outcomes: difference in means or standardised mean difference among missing outcomes enough to induce a clinically relevant bias on observed effect size. As-treated analysis undertaken with substantial departure of the intervention received from that assigned at randomisation. Equal to or greater than 30% dropout rate.
- 3. Unclear risk of bias: insufficient reporting of attrition or exclusions to permit judgement of low or high risk of bias. Greater than 10% and less than 30% dropout rate.

Selective outcome reporting?

- 1. Low risk of bias: all primary outcomes of interest adequately reported with point estimates and measures of variance for all time points.
- 2. High risk of bias: incomplete reporting of prespecified outcomes. One or more primary outcomes is reported using measurements, analysis methods, or subsets of data that were not prespecified. One or more reported primary outcomes were not prespecified. One or more outcomes of interest reported incompletely and cannot be entered into a meta-analysis. Results for a key outcome expected to have been reported, excluded.
- 3. Unclear risk of bias: insufficient information to permit judgement of low or high risk of bias.

Free of other biases

- 1. Low risk of bias: appears free of other sources of bias.
- 2. High risk of bias: results may have been confounded by at least one important risk of bias (design-specific, fraudulent, other).
- 3. Unclear risk of bias: other sources of bias may be present but there is either insufficient information to assess whether an important risk of bias exists; OR insufficient rationale or evidence regarding whether an identified problem will introduce bias.

CONTRIBUTIONS OF AUTHORS

Jennifer M Ryan: conceived and designed the review protocol; implemented the search strategy, applied eligibility criteria, assessed studies, extracted and analysed data; assessed the quality of the evidence using the GRADE approach; led the write-up of the review; and has overall responsibility for the review.

Elizabeth E Cassidy: informed the protocol design; applied eligibility criteria, assessed studies, extracted data; assessed the quality of the evidence using the GRADE approach; and assisted with the write-up of the review.

Stephen G Noorduyn: informed the protocol design; acted as the third reviewer; and assisted with the write-up of the review.

Neil E O'Connell: informed the protocol design; oversaw the data synthesis; acted as the third reviewer; and assisted with the write-up of the review

DECLARATIONS OF INTEREST

Jennifer M Ryan, Elizabeth E Cassidy, and Neil E O'Connell are chartered physiotherapists and lecturers in physiotherapy. As professionals who might be involved in the delivery of exercise interventions, it is plausible that they might be perceived as having a bias favouring the effectiveness of exercise.

Jennifer M Ryan is receiving funding from Action Medical Research and the Chartered Society of Physiotherapy Charitable Trust, to evaluate the feasibility, acceptability and efficacy of resistance training for adolescents with CP.

Elizabeth E Cassidy: none known.

Exercise interventions for cerebral palsy (Review)

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Stephen G Noorduyn: Stephen was lead author on Noorduyn 2011, which was screened by JR and EC.

Neil E O'Connell: none known.

SOURCES OF SUPPORT

Internal sources

• Brunel University London, UK.

Provided JMR and NEO'C with the time required to undertake this review

External sources

• None, Other.

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

- 1. The title has been changed from 'Exercise interventions for adults and children with cerebral palsy' to 'Exercise interventions for cerebral palsy', in accordance with guidance from the Cochrane Handbook (Higgins 2011b).
- 2. We planned to include studies that used any validated scale that measured the predefined primary and secondary outcomes. However, as trials used a range of outcome measures for these outcomes, we included any measure that purported to measure them, regardless of whether or not it was validated specifically in people with CP.
- 3. Although we proposed to classify general gross motor function as unaided walking, walking with aids, or unable to walk (Beckung 2008), most studies reported the GMFCS level of participants. Therefore, we reported the GMFCS level when available and the use of mobility aids when the GMFCS level was not available.
- 4. We were not able to use the following methods in the review, which we archived for use in future updates (see Table 1).
 - a. We planned to present the relative risk (or risk ratio (RR)) with 95% CI and calculate the number needed to treat for an additional beneficial outcome as an absolute measure of treatment effect, where studies used dichotomous outcomes (see Measures of treatment effect and Ryan 2015). We did not, however, identify any studies that reported dichotomous outcomes. Further, we will report the odds ratio (OR) with 95% CI in updates of this review as most studies report the OR rather than the RR if the outcome is dichotomous.
 - b. We planned to combine the results of cross-over studies with those of parallel studies by imputing the post-treatment betweencondition correlation coefficient from an included study that presented individual participant data, and using this to calculate the standard error of the SMD, using the generic inverse-variance method (see Unit of analysis issues and Ryan 2015).
 - c. We planned to further explore possible clinical heterogeneity through preplanned subgroup analysis based on important clinical features. We predicted that some trials would include ambulatory participants only (i.e. people who could walk with or without a mobility aid; GMFCS level I, II, and III), and some studies would include participants who could walk without a mobility aid only (i.e. GMFCS level I and II). Where adequate data allowed, we planned to undertake two subgroup analyses for studies that include ambulatory people only (i.e. GMFCS level I, II, and III) and for studies that include ambulatory people who walk without a mobility aid only (i.e. GMFCS level I and II). However, due to the small number of trials amenable to meta-analyses, we did not conduct subgroup analysis.
 - d. We planned to explore the impact of studies at high risk of bias by reanalysis with studies rated at overall high risk of bias excluded. However, we could not do this as all studies were rated at high risk of bias.
 - e. As stated in the 'Dealing with missing data' section, we planned to include all studies in the main analysis and exclude studies that were at high risk of bias for incomplete outcome data as a sensitivity analysis (Ryan 2015). However, this was not possible because of the small number of trials in each meta-analysis.
 - f. We planned to explore the influence of using imputed correlation coefficients in our approach to including cross-over and cluster trials (see Unit of analysis issues) by reanalysing these data with adjusted (higher and lower) coefficient values (Ryan 2015). However, we did not identify any cluster trials or include data from cross-over trials in any pooled analyses.
- 5. We stated in the protocol that Dr Brian Timmons would validate the final list of studies (Ryan 2015). Dr Timmons did not validate the final list of studies, so we deleted this sentence from the 'Selection of studies' section in the review.
- 6. Following identification of included studies, we noted a large overlap between the content of aerobic exercise interventions and mixed training interventions and between the content of resistance training interventions and mixed training interventions, respectively. We therefore decided to conduct a post hoc pooled analysis of aerobic exercise and mixed training versus usual care and resistance training and mixed training versus usual care.

INDEX TERMS

Medical Subject Headings (MeSH)

*Exercise; *Motor Skills; *Walking Speed; Cerebral Palsy [*rehabilitation]; Muscle Spasticity [rehabilitation]; Publication Bias; Randomized Controlled Trials as Topic; Resistance Training [*methods]



MeSH check words

Adolescent; Adult; Child; Female; Humans; Male; Young Adult