

### Original article

# Weight training and risk of all-cause, cardiovascular disease and cancer mortality among older adults

Prathiyankara Shailendra,<sup>1,2</sup> Katherine L Baldock,<sup>1,2</sup> Lok Sze Katrina Li,<sup>2</sup> Jessica Gorzelitz (1),<sup>3,4</sup> Charles E Matthews (1),<sup>4</sup> Britton Trabert (1),<sup>4,5</sup> Jason A Bennie<sup>6</sup> and Terry Boyle<sup>1,2,\*</sup>

<sup>1</sup>Australian Centre for Precision Health (ACPreH), University of South Australia, Adelaide, SA, Australia, <sup>2</sup>Allied Health and Human Performance, University of South Australia, Adelaide, SA, Australia, <sup>3</sup>Department of Health and Human Physiology, University of Iowa, Iowa City, IA, USA, <sup>4</sup>Metabolic Epidemiology Branch, Division of Cancer Epidemiology and Genetics, National Cancer Institute, Bethesda, MD, USA, <sup>5</sup>Obstetrics and Gynecology, University of Utah, Huntsman Cancer Institute at the University of Utah, Salt Lake City, UT, USA and <sup>6</sup>Population Health Unit, Murrumbidgee Primary Health Network, Wagga Wagga, NSW, Australia

\*Corresponding author. Australian Centre for Precision Health, University of South Australia, Level 8 South, SAHMRI Building, North Terrace, Adelaide, SA 5000, Australia. E-mail: Terry.Boyle@unisa.edu.au

#### Abstract

**Background:** While previous studies indicate muscle-strengthening exercises may reduce mortality risk, further research is needed to increase certainty of the evidence. We investigated overall and dose-response associations between weight training and the risks of all-cause, cardiovascular disease (CVD) and cancer mortality in a large cohort of older adults with long follow-up time and a large number of deaths. We also investigated the joint associations of weight training and aerobic exercise with mortality risk.

**Methods:** Weight training was assessed via self-report in 2004–05 in the National Institutes of Health-American Association of Retired Persons (NIH-AARP) Diet and Health Study (USA; n = 216339), with follow-up to 2019. Cox regression estimated the hazard ratios (HR) and 95% confidence intervals (CI) for the associations between weight training and mortality, after adjusting for confounders including aerobic exercise.

**Results:** Around 25% of participants [mean age = 69.9 years (standard deviation = 5.4), 58% men] reported engaging in weight training over the past year, and there were 79 107 (37%) deaths. Engaging in any weight training (vs none) was associated with lower risks of all-cause (HR = 0.94; 95% CI = 0.93-0.96), CVD (HR = 0.92; 95% CI = 0.90-0.95) and cancer mortality (HR = 0.95; 95% CI = 0.92-0.98). More time spent in weight training was associated with only marginally greater risk reductions. Larger risk reductions were observed among women than men. Performing both aerobic exercise and weight training conferred the greatest mortality risk reduction; weight training was not associated with mortality risk among participants who did no aerobic exercise.

Conclusion: Performing any amount of weight training lowered mortality risk.

Keywords: Mortality, muscle-strengthening activity, older adults, physical activity, strength training, resistance training.

#### **Key Messages**

- Performing any amount of weight training was associated with lower risk of all-cause, CVD and cancer mortality in this cohort of older adults, and those who engaged in both weight training and aerobic exercise were at lowest mortality risk.
- Larger risk reductions were observed among women than men.
- Further emphasis should be placed on the benefits of muscle-strengthening activities for overall health and mortality reduction in public health programmes.

#### Introduction

National and global physical activity guidelines recommend that, in addition to performing moderate-to-vigorous intensity aerobic physical activity (MVPA; e.g. brisk walking, running, cycling), adults should engage in muscle-strengthening exercises involving all major muscle groups on 2 or more days per week.<sup>1–3</sup> Muscle-strengthening exercises, sometimes referred to as resistance training or strength training, involve using weight machines, resistance bands, free weights or body weight to strengthen the musculoskeletal system.<sup>4</sup>

Research indicates engaging in muscle-strengthening exercises has a range of cardiometabolic, musculoskeletal and mental health benefits,<sup>5–9</sup> and that the combination of both muscle-strengthening exercises and aerobic exercise leads to greater improvements in anthropometric outcomes, metabolic syndrome factors and cardiovascular risk markers than

Received: 3 August 2023. Editorial Decision: 25 March 2024. Accepted: 25 May 2024

<sup>©</sup> The Author(s) 2024. Published by Oxford University Press on behalf of the International Epidemiological Association.

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (https://creativecommons.org/licenses/by/4.0/), which permits unrestricted reuse, distribution, and reproduction in any medium, provided the original work is properly cited.

performing either of those activities alone.<sup>10,11</sup> Findings from cohort studies suggest muscle-strengthening exercises may be associated with a lower risk of developing type 2 diabetes, cardiovascular disease (CVD), obesity and some cancers.<sup>12–14</sup> However, although numerous studies suggest aerobic exercise is associated with lower mortality risk among the general population and older adults,<sup>3,15</sup> there are considerably fewer studies on muscle-strengthening exercises and mortality risk.

The research to date has generally found that musclestrengthening exercises reduce the risk of all-cause mortality.<sup>14,16</sup> A 2022 systematic review and meta-analysis of 16 studies, seven of which examined all-cause mortality, found engaging in muscle-strengthening exercises reduced all-cause mortality risk by 15% compared with doing none.<sup>1</sup> Subsequent studies have observed risk reductions of similar magnitude.<sup>17,18</sup> Engaging in muscle-strengthening exercises has also been associated with a reduced risk of CVD mortality and cancer mortality; however, there are fewer studies and the results are inconsistent.<sup>13,16</sup> Previous studies have generally had information about frequency (e.g. sessions/ week) rather than time spent performing musclestrengthening exercises (e.g. min/week), so the impact of higher volumes of muscle-strengthening exercises on mortality is uncertain.<sup>14,16</sup> Previous studies have also had relatively small sample sizes, small numbers of deaths and/or short follow-up times, which has limited their precision when estimating the association between muscle-strengthening exercises and mortality risk. As such, several areas require further investigation. These include whether the association differs by sex, and whether performing both muscle-strengthening and aerobic exercise (as recommended by physical activity guidelines) confers additional mortality benefits compared with either activity alone.<sup>14,16</sup> Finally, it is unclear if musclestrengthening exercises confer mortality benefits among older adults, as most studies to date have been conducted in study populations with adults of all ages.

The aim of this study was to investigate the dose-response associations between weight training—a specific type of muscle-strengthening exercise—and all-cause mortality, CVD mortality, and cancer mortality overall, stratified by sex in a large cohort of older adults with long follow-up time and a large number of deaths. We hypothesized that weight training would be associated with reduced mortality risk. In addition, we investigated the joint effects of weight training and aerobic exercise on mortality outcomes.

#### Methods

This study used data from the National Institutes of Health-American Association of Retired Persons (NIH-AARP) Diet and Health Study, a cohort of 566 398 AARP members who completed a mailed entry questionnaire regarding demographics, medical history and dietary behaviours in 1995–96.<sup>19</sup> Further information on the cohort is available elsewhere.<sup>19</sup> Participants who completed the entry questionnaire were invited to complete a questionnaire at two subsequent time points: 1996–97 (not used in this analysis) and 2004–05.

The 2004–05 questionnaire collected detailed information on lifestyle and physical activity (including weight training) and was completed by 313791 participants, who ranged in age from 59 to 82 years at the time of completion. These participants form the basis of the current analysis. The following exclusion criteria were then applied (in order): proxy respondents (n = 20054); participants who reported being unable to walk (n = 7418); participants who did not complete the weight training question (n = 13297); and participants with missing data for any confounders included in the analyses (n = 56683). After these exclusions, 216339 participants remained and were included in the current analyses (Supplementary Figure S1, available as Supplementary data at *IJE* online).

## Measurement of weight training and aerobic exercise

In the 2004-05 questionnaire, participants were asked to report the amount of time per week spent performing different physical activities and exercises, including weight training, over the past 12 months. The full definition of weight training provided to participants in the questionnaire was 'weight training or lifting (include free weights and machines)'. Participants could choose from 10 different categories: none, 5 min, 15 min, 30 min, 1 h, 1.5 h, 2-3 h, 4-6 h, 7-10 h and >10 h. Similar questionnaires demonstrate acceptable validity and reliability when measuring weight training and musclestrengthening exercises (intraclass correlation coefficients  $\sim 0.5-0.6$ ).<sup>20,21</sup> To avoid sparsely populated categories, these data were categorized as: none, 5-15 min, 30 min, 1 h, 1.5 h, 2-3 h and 4 h and greater in the primary analyses examining the associations between weight training and mortality risk; and none, 5-30 min, 1-1.5 h and >2 h in the joint effects analyses.

Metabolic equivalents of task-h (MET-h) per week of leisure-time aerobic exercise were calculated based on time spent per week in each of: jogging, tennis, golf, swimming, cycling, walking for exercise and other aerobic activity.<sup>19</sup> To facilitate calculation of MET-h/week of aerobic exercise, the 10 categories described above were converted to a continuous 'h/week' measure, with the mid-point used when the response option was a range, and a MET-value was assigned to each activity. Total MET-h/week of aerobic exercise was categorized as none (0 MET-h/week), with the remaining participants classified into three categories of roughly equal size which were defined as low (0.1 to <10.75), medium (10.75 to <29) and high (>29).

#### Ascertainment of outcome

Mortality data from the Social Security Administration Death Master File and the National Data Index Plus were used to ascertain mortality status. We used Surveillance, Epidemiology, End Result (SEER) codes for underlying and contributing causes of death to identity CVD deaths (codes 50060 to 50110) and cancer deaths (codes 20010 to 37000). Follow-up time began at the 2004–05 questionnaire completion date and ended at death or the time of the last available mortality extract (31 December 2019), whichever occurred first.

#### Statistical analysis

Participant characteristics were categorized using descriptive statistics. Cox proportional hazards regression was used to estimate hazard ratios (HRs) and associated 95% confidence intervals (CIs) for the associations between weight training and all-cause, CVD and cancer mortality, after adjusting for confounders. We used a directed acyclic graph (DAG) to identify confounders based on the determinants of weight training and mortality risk factors. The following variables

were included as confounders in all models: age, sex, education, race, body mass index (BMI), alcohol consumption, self-reported health status (as a proxy for comorbidity and overall health) and healthy eating index,<sup>22</sup> all of which were based on self-reported information from the 1994–95 entry questionnaire; and leisure-time aerobic exercise and cigarette smoking, which were taken from the 2004–05 questionnaire (see Table 1 for further information about these variables). Interaction terms were added to the models and global Wald tests were conducted to determine whether the associations between weight training and mortality risk differed by sex or by concurrent leisure-time aerobic exercise.

Two sensitivity analyses were performed to evaluate risk of bias associated with reverse causation and potential confounding by chronic conditions: (i) excluding participants who self-reported being diagnosed with cancer, angina, heart attack, coronary disease or stroke prior to follow-up questionnaire completion; and (ii) censoring deaths within the first 2 years of follow-up. Further sensitivity analyses were conducted to assess the impact of selection bias induced by: (i) a large proportion of the 1994-95 entry cohort not completing the 2004-05 questionnaire; and (ii) participants being excluded from the analytical sample due to missing data. We first compared the characteristics of the analytical sample with participants who did not complete the follow-up questionnaire and participants who were excluded due to missing data. Second, we predicted the likelihood of being in the analytical sample using the variables reported in Supplementary Table S1 (available as Supplementary data at IJE online), then used these values as weights in an inverse probability weighting-based analysis to account for the potential impact of exclusions.

We investigated the joint associations of weight training and aerobic exercise by creating joint categories of weight training and aerobic exercise consisting of all combinations of four categories of weight training and four categories of leisure-time aerobic exercise.

Proportional hazards assumptions were tested using the Schoenfeld residuals technique; no meaningful violations were observed. All statistical analyses were conducted using Stata v15.1 (StataCorp, College Station, TX). Reporting follows the STROBE guidelines.<sup>23</sup>

#### Results

The mean age of participants was 69.9 years (standard deviation = 5.4 years) at the time of the 2004–05 follow-up questionnaire completion. There was a higher proportion of men (58%) and White participants (94%), and 46% were college graduates. There were 79107 (37%) deaths during the follow-up period (median follow-up time = 15 years), of which 56% and 31% were due to CVD and cancer, respectively.

Approximately 25% of participants reported doing weight training (Table 1). Weight training was more prevalent among younger participants, men, participants with a BMI in the 18.5–<25 kg/m<sup>2</sup> range, participants with higher levels of aerobic exercise, participants with higher education levels and participants who self-reported their health as excellent.

#### Weight training and mortality risk

Compared with participants who did not perform any weight training, participants who performed any weight training had with only marginally greater risk reductions. The associations between weight training and the risks of all-cause and CVD mortality, but not cancer mortality, were stronger in women compared with men ( $P_{interaction} < 0.001$  for all-cause and CVD mortality in all analyses,  $P_{interaction}$  for cancer mortality = 0.339 and 0.141 in None/Some and categorical weight training analyses, respectively; Table 2). Among women, performing any weight training was associated with 12%, 16% and 9% decreased risks of all-cause, CVD and cancer mortality, respectively, compared with doing none. The corresponding risk reductions among men were 3%, 4% and 4%.

The results of sensitivity analyses investigating the potential impact of reverse causation, confounding by chronic conditions and selection bias, were not meaningfully different from the original analyses (Supplementary Table S2, available as Supplementary data at *IJE* online).

#### Joint effects of weight training and aerobic exercise on mortality risk

The association between weight training and mortality risk appeared to vary by level of aerobic leisure-time exercise, although the interaction term for weight training and aerobic exercise was statistically significant for CVD mortality ( $P_{Interaction} = 0.014$ ) but not all-cause mortality ( $P_{Interaction} = 0.264$ ) or cancer mortality ( $P_{interaction} = 0.586$ ).

There was no evidence that weight training was associated with all-cause or cancer mortality among participants who did not report performing any aerobic leisure-time exercise, and only weak evidence of a potential reduction in CVD mortality was observed (Figure 1; Supplementary Table S3, available as Supplementary data at *IJE* online). Similarly, engaging in weight training appeared to confer little mortality benefit among participants who reported the highest levels of aerobic exercise. For example, HRs for all-cause mortality among participants in the highest category aerobic exercise were 0.78 (95% CI = 0.76–0.80) for no weight training and 0.73 (95% CI = 0.70–0.76) for 2+ h/week of weight training, compared with participants who performed neither weight training nor aerobic exercise.

Increasing min/week of weight training did appear to provide additional mortality benefits to those provided by aerobic exercise among participants who performed low-to-intermediate levels of aerobic activity. For all-cause mortality, among those who performed 0.1-<10.75 MET-h/week of aerobic exercise, the HRs decreased from 0.95 (95% CI = 0.93-0.98) for no weight training to 0.80 (95% CI = 0.70-0.93) for the highest level of weight training, and the corresponding HRs among participants who performed 10.75-<29 75 MET-h/week were 0.84 (95% CI = 0.82-0.86) and 0.74 (95% CI = 0.69-0.79). Larger risk reductions were observed for CVD mortality than for cancer mortality.

Increasing levels of aerobic exercise were associated with increasingly greater risk reductions for all three mortality outcomes, across all levels of weight training.

Table 1	<ul> <li>Characteristics of</li> </ul>	of the participants in t	he NIH-AARP Die	t and Health stuc	ly who returned	d the follow-up	questionnaire,	by levels	of weight
training	(n = 216339)								

	Weight training/week							
Characteristic	None	5–15 min	30 min	1 h	1.5 h	2–3 h	4+ h	
<i>n</i> (%)	162 246 (75.0)	9800 (4.5)	11 223 (5.2)	11 687 (5.4)	8188 (3.8)	8792 (4.1)	4403 (2.0)	
Age (years), 2004–05, mean (SD)	70.1 (5.4)	69.4 (5.5)	69.5 (5.4)	69.4 (5.4)	69.1 (5.2)	68.9 (5.3)	68.8 (5.2)	
Sex (%)								
Men	57.0	62.2	59.9	61.4	62.3	64.3	69.7	
Women	43.0	37.8	40.1	38.6	37.7	35.7	30.3	
Race (%)								
White	94.0	93.5	94.3	94.7	95.2	94.9	92.6	
Black	3.1	3.1	2.7	2.3	1.9	1.9	3.7	
Hispanic	1.5	1.4	1.5	1.5	1.5	1.7	2.4	
Other	1.4	2.0	1.6	1.4	1.3	1.5	1.3	
Smoking status, 2004–05 (%)								
Never smoker	40.3	41.9	41.2	39.9	38.5	38.6	38.1	
Former smoker, quit 10+ years ago	42.5	46.3	47.0	48.1	49.3	49.5	49.1	
Former smoker, quit 5–9 year ago	3.4	2.9	2.7	3.0	3.2	2.8	2.5	
Former smoker, quit <5 years ago	2.8	2.1	2.0	2.0	2.2	1.9	2.3	
Former smoker, time since quitting unknown	4.3	3.7	4.1	4.1	4.0	4.3	4.8	
Current smoker	6.7	3.1	3.1	2.8	2.6	2.9	3.2	
Education level, 1995–96 (%)								
<12 years	4.0	1.8	1.6	1.5	1.7	1.4	2.7	
12 years or completed high school	19.5	10.9	11.2	10.0	9.7	9.6	12.8	
Post high-school training	10.4	7.7	7.4	7.9	6.6	7.1	7.5	
Some college	24.2	20.7	22.0	21.8	21.8	21.1	23.4	
College graduate	41.9	58.9	57.8	58.8	60.2	60.8	53.6	
Self-reported health, 1995–96 (%)								
Excellent	16.9	22.0	24.6	27.1	30.1	32.5	33.7	
Very good	37.9	40.0	41.7	41.3	41.6	41.1	39.4	
Good	35.8	31.0	28.1	27.0	24.3	22.6	22.4	
Fair	8.6	6.4	5.3	4.2	3.6	3.6	4.2	
Poor	0.8	0.6	0.4	0.4	0.3	0.3	0.3	
Body mass index $(kg/m^2)$ 1995–96 (%)	0.0	0.0	0.1	0.1	0.0	0.0	0.5	
<18.5	0.9	11	1.0	1.0	12	11	1.0	
18 5-<25	34.0	42.0	42.9	44.8	45.4	45.8	43.0	
25-<30	43.0	40.9	41.8	40.6	40.2	40.3	42.8	
30 - < 35	16.0	12.1	11.0	10.8	10.5	10.0	10.2	
35+	6.1	4.0	3 3	2.8	2.6	2.8	2.9	
Body mass index $(kg/m^2)$ 1995–96 mean (SD)	27 1 (4 9)	26 2 (4 6)	26.0(4.6)	259(42)	25.8(4.1)	258(41)	259(48)	
Alcohol intake $(g/day)$ 1995–96 mean $(SD)$	12.8(34.7)	12.6(29.6)	12 1 (28 0)	123(27.8)	12.9(27.0)	12.9(28.8)	146(353)	
Healthy Fating Index score 1995–96 mean (SD)	67 5 (9 5)	697(89)	704(88)	70.8 (8.6)	71 3 (8 5)	71 1 (8 6)	70.4 (8.8)	
Aerobic leisure-time evercise (MET-h/week)	20.2(26.6)	250(250)	31.8(27.2)	38.2(31.0)	40.6(31.7)	497(383)	97.0 (84.5)	
2004–05, mean (SD)	20.2 (20.0)	23.0 (23.0)	51.0 (27.2)	56.2 (51.0)	40.0 (31.7)	49.7 (30.3)	J7.0 (04.3)	
Aerobic leisure-time exercise (MET-h/week), 2004–05 (%)								
0	13.6	1.7	1.5	1.3	1.3	1.4	1.9	
0.1-<10.75	31.9	31.0	16.6	9.9	6.5	5.1	3.9	
10.75-<29	30.3	36.3	39.2	36.5	35.0	24.9	11.8	
29+	24.2	30.9	42.6	52.4	57.2	68.6	82.4	

MET, metabolic equivalent of task; NIH-AARP, National Institutes of Health-American Association of Retired Persons; SD, standard deviation.

#### Discussion

In this large cohort study of older adults, engaging in any amount of weight training was associated with a 6% lower risk of all-cause mortality, 8% lower risk of CVD mortality and 5% lower risk of cancer mortality, compared with those who did none. Larger risk reductions were observed for women than men, particularly for all-cause mortality (3% in men vs 12% in women) and CVD mortality (4% in men vs 16% in women). Engaging in high levels of both weight training and aerobic leisure-time exercise was associated with greater mortality benefits compared with performing either of these activities alone, particularly among participants who performed low-to-intermediate levels of aerobic exercise. Our results are consistent with previous studies on the association between muscle-strengthening exercises and mortality risk and provide strong evidence for this association among older adults. Consistent with our study, recent research indicates engaging in muscle-strengthening exercises reduces the risk of all-cause mortality by 10–15% compared with doing none, and also reduces the risk of both CVD and cancer mortality.<sup>14,16,17,24</sup> Given fewer than 20% of adults in the USA, Europe and Australia meet current muscle-strengthening activity guidelines,<sup>25–27</sup> and that this figure is even lower among older adults,<sup>4</sup> our study supports a greater emphasis on increasing the prevalence of muscle-strengthening exercises among older adults.

Table 2. Associations between weight training and all-cause, cardiovascular disease and cancer mortality overall and among men and women (n = 216 339)

			All-cause mortality		CVD mortality	Cancer mortality		
Weight training	Person-years	Deaths	HR (95% CI), <i>P</i> <sup>a</sup>	Deaths	HR (95% CI), P <sup>a</sup>	Deaths	HR (95% CI), <i>P</i> <sup>a</sup>	
All participants								
No weight training	2 046 616	62 5 50	1.00 (Reference)	35 217	1.00 (Reference)	19 314	1.00 (Reference)	
Some weight training	715 998	16 557	0.94 (0.93-0.96), <0.001	8953	0.92 (0.90–0.95), <0.001	5459	0.95 (0.92-0.98), 0.001	
Categories (none is reference)								
5–15 min/week	127 589	3199	0.95 (0.91-0.98), 0.003	1746	0.92 (0.88-0.97), 0.001	997	0.93 (0.87-0.99), 0.030	
30 min/week	147 705	3564	0.95 (0.92–0.98), 0.003	1954	0.94 (0.89–0.98), 0.005	1173	0.97 (0.91-1.03), 0.292	
1 h/week	154 615	3653	0.97 (0.93–1.00), 0.045	1964	0.94 (0.90-0.98), 0.008	1178	0.95 (0.89-1.01), 0.081	
1.5 h/week	109 111	2421	0.94 (0.90-0.98), 0.002	1309	0.93 (0.87-0.98), 0.007	801	0.93 (0.86-0.99), 0.036	
2–3 h/week	118 202	2474	0.91 (0.88–0.95), <0.001	1297	0.88 (0.83–0.93), <0.001	883	0.96 (0.90-1.03), 0.248	
4+ h/week	58 776	1246	0.92 (0.87–0.97), 0.004	683	0.92 (0.85–0.99), 0.027	427	0.91 (0.83-1.01), 0.066	
Men								
No weight training	1 136 981	39 247	1.00 (Reference)	22 586	1.00 (Reference)	12 405	1.00 (Reference)	
Some weight training	437 390	11 851	0.97 (0.95–0.99), 0.012	6599	0.96 (0.93–0.99), 0.005	3874	0.96 (0.92-1.00), 0.032	
Categories (none is reference)								
5–15 min/week	77 216	2277	0.97 (0.93-1.01), 0.137	1269	0.94(0.89 - 1.00), 0.041	684	0.92 (0.85-0.99), 0.025	
30 min/week	86 204	2460	0.98 (0.94–1.02), 0.344	1.372	0.96 (0.91-1.02), 0.161	829	1.01 (0.94-1.09), 0.692	
1 h/week	92 664	2598	1.01 (0.97–1.05), 0.781	1442	0.99 (0.94–1.04), 0.696	844	0.99 (0.92-1.06), 0.696	
1.5 h/week	66 505	1729	0.96 (0.91–1.01), 0.102	979	0.97 (0.91–1.04), 0.373	559	0.92 (0.85-1.01), 0.067	
2–3 h/week	74 479	1821	0.95 (0.90–0.99), 0.030	991	0.93 (0.87–0.99), 0.024	632	0.96 (0.89-1.05), 0.380	
4+ h/week	40 321	969	0.94 (0.88–1.01), 0.075	546	0.96 (0.88–1.04), 0.334	326	0.91 (0.82-1.02), 0.112	
Women								
No weight training	906 635	23 303	1.00 (Reference)	12 631	1.00 (Reference)	6909	1.00 (Reference)	
Some weight training	278 608	4703	0.88 (0.85-0.91), <0.001	2354	0.84 (0.80–0.88), <0.001	1585	0.91 (0.86-0.97), 0.002	
Categories (none is reference)								
5–15 min/week	50 374	922	0.90 (0.84–0.96), 0.002	477	0.88 (0.80-0.96), 0.006	313	0.97 (0.86-1.08), 0.542	
30 min/week	61 500	1104	0.89 (0.83–0.94), <0.001	582	0.88 (0.81–0.96), 0.003	344	0.87 (0.78-0.97), 0.013	
1 h/week	61 950	1055	0.88 (0.82–0.93), <0.001	522	0.83 (0.75–0.90), <0.001	334	0.86 (0.77-0.96), 0.009	
1.5 h/week	42 605	692	0.89 (0.82–0.96), 0.002	330	0.81 (0.73–0.91), <0.001	242	0.93 (0.82-1.06), 0.287	
2–3 h/week	43 723	653	0.84 (0.77–0.90), <0.001	306	0.76 (0.67–0.85), <0.001	251	0.95 (0.84-1.09), 0.484	
4+ h/week	18 455	277	0.86 (0.76–0.97), 0.012	137	0.81 (0.68–0.96), 0.013	101	0.93 (0.76-1.13), 0.446	

CI, confidence interval; CVD, cardiovascular disease; HR, hazard ratio; kg, kilogram; m, metre.

<sup>a</sup> Adjusted for the following confounders: age, sex, education, race, body mass index, leisure-time aerobic exercise, cigarette smoking, alcohol

consumption, Healthy Eating Index, self-reported health status.

Our study provides additional insights into whether the association between muscle-strengthening exercises and mortality differs by sex, with stronger associations for all-cause mortality and CVD mortality observed among women than men. Whileeas one previous study also found weight training was associated with larger mortality risk reductions among women than men,<sup>17</sup> most studies have reported no effect modification by sex<sup>18,28–30</sup> and a potential biologically plausible mechanism is not clear. Future research should further investigate whether the association between musclestrengthening exercises and mortality differs by sex and potential biological mechanisms. Regardless, our study provides robust evidence that weight training is associated with lower mortality risk among older women, a group among whom the prevalence of participation in muscle-strengthening exercises is particularly low.<sup>4</sup>

The large number of participant deaths in this cohort allowed us to investigate the joint effects of weight training and aerobic exercise with greater depth and precision than previous studies. Consistent with previous research, engaging in both exercise types provided the greatest mortality benefits,<sup>30–33</sup> supporting the recommendations of physical activity guidelines to perform muscle-strengthening exercises in addition to aerobic MVPA.<sup>1</sup> Weight training was associated with mortality benefits additional to those conferred by aerobic exercise among older adults who engaged in low-tointermediate levels of aerobic activity. However, it was not associated with all-cause or cancer mortality among participants who performed no aerobic exercise, and only weakly associated with CVD mortality. Weight training was not associated with additional mortality risk reductions among participants with the highest levels of aerobic exercise, perhaps due to this group already being at lower mortality risk. One possible explanation for these findings is that musclestrengthening exercises, when performed in combination with aerobic exercise, may enhance the health benefits conveyed by aerobic exercise. This is supported by evidence from clinical trials which indicates that, for a range of anthropometric and cardiometabolic markers, the benefits gained by performing a combination of muscle-strengthening exercise and aerobic exercise are greater than those seen for aerobic exercise alone, even when the benefits of muscle-strengthening exercises alone may be modest.<sup>10,11,34</sup> Similar findings have been observed for mortality due to a range of causes.<sup>35</sup> There are also likely to be cross-over effects between aerobic and muscle-strengthening exercises, e.g. weight-bearing aerobic exercises (e.g. running) may improve muscle and bone strength, and some muscle-strengthening exercises may improve cardiovascular fitness.

Our study is one of the few large cohorts to have information on min/week of muscle-strengthening exercise, so provides novel and important insights into the dose-response



**Figure 1.** Associations between joint categories of weight training and moderate to leisure-time aerobic exercise and the risks of (A) all-cause mortality, (B) cardiovascular disease mortality and (C) cancer mortality. MET-h, metabolic equivalent of task-h

relationship. We found increasing levels of weight training were associated with only marginally greater risk reductions. A recent dose-response meta-analysis of four studies found a 'U-shaped' relationship between resistance training volume and all-cause mortality risk, with the largest reduction (33%) observed at 60 min/week and greater volumes associated with smaller or no risk reductions.<sup>16</sup> Although somewhat in contrast, both studies suggest large volumes of muscle-strengthening exercises are not needed to receive the optimal mortality benefits.

Strengths of this study include its prospective design, large sample size and long follow-up period, recording a large number of deaths. Having information about min/week spent performing weight training meant we could look at the doseresponse relationship with mortality in more detail than most previous studies, which have only measured frequency of muscle-strengthening exercises (e.g. number of sessions/week).

Limitations included the use of self-reported weight training data, which are susceptible to measurement error. Few studies have investigated the validity and reliability of selfreported weight training,36 and participants may have reported time spent at the gym rather than time engaged in weight training activity specifically. We lacked data on the weight, number of repetitions and intensity of weight training, as well as information about other forms of musclestrengthening exercises (including those performed in other domains, such as occupational settings). Lacking information about the frequency of weight training limited direct evaluation of the current physical activity guidelines, which recommend two or more sessions/week rather than a specific number of minutes. We also only had information about weight training at a single point in time. Future cohort studies with more detailed measures of muscle-strengthening exercises, assessed longitudinally and with valid and reliable tools, would provide stronger evidence about the association with mortality risk. Finally, participants were predominantly older, White adults who were more likely to have higher education levels and better health than the general population, and it is not known if the observed results are generalizable to populations with different sociodemographic characteristics.

Our results indicate engaging in muscle-strengthening exercises such as weight training has mortality benefits for older adults, even at low levels. Given the low prevalence in the general population, and older adults in particular,<sup>4,25</sup> further emphasis should be placed on the benefits of musclestrengthening activities for mortality risk reduction in public health programmes.

#### **Ethics approval**

The NIH-AARP Diet and Health Study was approved by the Special Studies Institutional Review Board of the National Cancer Institute. All participants gave informed consent.

#### **Data availability**

Data for the NIH-AARP Diet and Health Study are maintained by the National Cancer Institute, Division of Cancer Epidemiology and Genetics. Data described in the manuscript, a code book and analytical code will be made available upon request pending study approval from the NIH-AARP Diet and Health Study Steering Committee. Further details are provided at [https://www.nihaarpstars.com/].

#### Supplementary data

Supplementary data are available at IJE online.

#### **Author contributions**

P.S. and T.B. designed and conducted the statistical analyses, with significant intellectual input from all other authors. All authors contributed to interpretation of the results. The manuscript was drafted by P.S. and T.B. and critically revised by all other authors. All authors read and approved the final manuscript. C.E.M. is a member of the NIH-AARP cohort study team. T.B. is responsible for the overall content as guarantor.

#### Funding

This research was supported [in part] by the Intramural Research Program of the NIH, National Cancer Institute.

#### Acknowledgements

We are indebted to the participants in the NIH-AARP Diet and Health Study for their outstanding cooperation. We wish to acknowledge Dr Arthur Schatzkin who was instrumental in conceiving and establishing the NIH-AARP Diet and Health Study. We also thank former and current study leaders at the National Cancer Institute and AARP, including Louise A Brinton, Laurence S Freedman, Albert R Hollenbeck, Victor Kipnis, Michael F Leitzmann, Linda M Liao, Charles E Matthews, Yikyung Park, Rashmi Sinha, Amy F Subar and Mary H Ward. Additionally, we are thankful to Sigurd Hermansen and Kerry Grace Morrissey from Westat for study outcomes ascertainment and management and to Leslie Carroll and her team at Information Management Services for data support and analysis.

#### **Conflict of interest**

None declared.

#### References

- Bull FC, Al-Ansari SS, Biddle S *et al.* World Health Organization 2020 guidelines on physical activity and sedentary behaviour. *Br J Sports Med* 2020;54:1451–62.
- Department of Health and Social Care. UK Chief Medical Officers' Physical Activity Guidelines. London: Department of Health and Social Care, 2019. https://assets.publishing.service. gov.uk/media/5d839543ed915d52428dc134/uk-chief-medicalofficers-physical-activity-guidelines.pdf (16 April 2024, date last accessed).
- King AC, Whitt-Glover MC, Marquez DX et al.; 2018 PHYSICAL ACTIVITY GUIDELINES ADVISORY COMMITTEE. Physical activity promotion: highlights from the 2018 physical activity guidelines advisory committee systematic review. Med Sci Sports Exerc 2019;51:1340–53.
- Bennie JA, Shakespear-Druery J, De Cocker K. Muscle-strengthening exercise epidemiology: a new frontier in chronic disease prevention. Sports Med Open 2020;6:40.
- Zhao R, Zhao M, Xu Z. The effects of differing resistance training modes on the preservation of bone mineral density in postmenopausal women: a meta-analysis. Osteoporos Int 2015; 26:1605–18.
- Gordon BR, McDowell CP, Lyons M *et al*. The effects of resistance exercise training on anxiety: a meta-analysis and meta-regression analysis of randomized controlled trials. *Sports Med* 2017; 47:2521–32.
- MacDonald HV, Johnson BT, Huedo-Medina TB et al. Dynamic resistance training as stand-alone antihypertensive lifestyle therapy: a meta-analysis. J Am Heart Assoc 2016;5:e003231.
- Gordon BR, McDowell CP, Hallgren M *et al.* Association of efficacy of resistance exercise training with depressive symptoms: meta-analysis and meta-regression analysis of randomized clinical trials. *JAMA Psychiatry* 2018;75:566–76.

- Ashton RE, Tew GA, Aning JJ et al. Effects of short-term, medium-term and long-term resistance exercise training on cardiometabolic health outcomes in adults: systematic review with metaanalysis. Br J Sports Med 2020;54:341–48.
- Liang M, Pan Y, Zhong T *et al.* Effects of aerobic, resistance, and combined exercise on metabolic syndrome parameters and cardiovascular risk factors: a systematic review and network meta-analysis. *Rev Cardiovasc Med* 2021;22:1523–33.
- Morze J, Rücker G, Danielewicz A *et al*. Impact of different training modalities on anthropometric outcomes in patients with obesity: A systematic review and network meta-analysis. *Obes Rev* 2021;22:e13218.
- 12. Giovannucci EL, Rezende LFM, Lee DH. Muscle-strengthening activities and risk of cardiovascular disease, type 2 diabetes, cancer and mortality: a review of prospective cohort studies. *J Intern Med* 2021;**290**:789–805.
- 13. Nascimento W, Ferrari G, Martins CB *et al.* Muscle-strengthening activities and cancer incidence and mortality: a systematic review and meta-analysis of observational studies. *Int J Behav Nutr Phys Act* 2021;18:69.
- Momma H, Kawakami R, Honda T *et al.* Muscle-strengthening activities are associated with lower risk and mortality in major non-communicable diseases: a systematic review and metaanalysis of cohort studies. *Br J Sports Med* 2022;56:755–63.
- 15. Hupin D, Roche F, Gremeaux V *et al*. Even a low-dose of moderate-to-vigorous physical activity reduces mortality by 22% in adults aged  $\geq 60$  years: a systematic review and meta-analysis. *Br J Sports Med* 2015;49:1262–67.
- Shailendra P, Baldock KL, Li LSK *et al.* Resistance training and mortality risk: a systematic review and meta-analysis. *Am J Prev Med* 2022;63:277–85.
- Gorzelitz J, Trabert B, Katki HA *et al.* Independent and joint associations of weightlifting and aerobic activity with all-cause, cardio-vascular disease and cancer mortality in the Prostate, Lung, Colorectal and Ovarian Cancer Screening Trial. *Br J Sports Med* 2022;56:1277–83.
- Coleman CJ, McDonough DJ, Pope ZC *et al*. Dose–response association of aerobic and muscle-strengthening physical activity with mortality: a national cohort study of 416 420 US adults. *Br J Sports Med* 2022;56:1218–23. doi: 10.1136/bjsports-2022-105519.
- Mazzilli KM, Matthews CE, Salerno EA *et al*. Weight training and risk of 10 common types of cancer. *Med Sci Sports Exerc* 2019; 51:1845–51.
- Chasan-Taber S, Rimm EB, Stampfer MJ *et al.* Reproducibility and validity of a self-administered physical activity questionnaire for male health professionals. *Epidemiology* 1996;7:81–86.
- 21. Shakespear-Druery J, De Cocker K, Biddle SJH *et al.* Muscle-Strengthening Exercise Questionnaire (MSEQ): an assessment of concurrent validity and test–retest reliability. *BMJ Open Sport Exerc Med* 2022;8:e001225.
- 22. Arem H, Reedy J, Sampson J *et al.* The Healthy Eating Index 2005 and risk for pancreatic cancer in the NIH-AARP Study. *J Natl Cancer Inst* 2013;105:1298–305.
- 23. von Elm E, Altman DG, Egger M, STROBE Initiative *et al.* Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) statement: guidelines for reporting observational studies. *Bmj* 2007;**335**:806–808.
- Marino KR, Vishnubala D, Oja P. Muscle-strengthening activities to improve health outcomes: what the evidence supports. *Br J Sports Med* 2022;56:831–32.
- Bennie JA, De Cocker K, Teychenne MJ *et al.* The epidemiology of aerobic physical activity and muscle-strengthening activity guideline adherence among 383,928 U.S. adults. *Int J Behav Nutr Phys Act* 2019;16:34.
- Bennie JA, De Cocker K, Smith JJ *et al.* The epidemiology of muscle-strengthening exercise in Europe: a 28-country comparison including 280,605 adults. *PLoS One* 2020;15:e0242220.
- 27. Bennie JA, Pedisic Z, van Uffelen JGZ et al. The descriptive epidemiology of total physical activity, muscle-strengthening exercises and sedentary behaviour among Australian adults—results from

the National Nutrition and Physical Activity Survey. *BMC Public Health* 2016;16:73.

- 28. Nie J, Haberstroh M, Acosta T *et al.* Independent and joint associations between leisure time physical activity and strength activities with mortality outcomes in older adults at least 65 years of age: a prospective cohort study. *J Gerontol A Biol Sci Med Sci* 2021; 76:2122–31.
- Patel AV, Hodge JM, Rees-Punia E *et al*. Relationship between muscle-strengthening activity and cause-specific mortality in a large US cohort. *Prev Chronic Dis* 2020;17:E78–78.
- Siahpush M, Farazi PA, Wang H et al. Muscle-strengthening physical activity is associated with cancer mortality: results from the 1998–2011 National Health Interview Surveys, National Death Index record linkage. *Cancer Causes Control* 2019;30:663–70.
- 31. Stamatakis E, Lee IM, Bennie J *et al.* Does strength-promoting exercise confer unique health benefits? a pooled analysis of data on 11 population cohorts with all-cause, cancer, and cardiovascular mortality endpoints. *Am J Epidemiol* 2018;187:1102–12.

- Schoenborn C, Stommel MP. Adherence to the 2008 adult physical activity guidelines and mortality risk. *Am J Prev Med* 2011; 40:514–21.
- Zhao G, Li C, Ford ES *et al.* Leisure-time aerobic physical activity, muscle-strengthening activity and mortality risks among US adults: the NHANES linked mortality study. *Br J Sports Med* 2014;48:244–49.
- 34. Mann S, Beedie C, Jimenez A. Differential effects of aerobic exercise, resistance training and combined exercise modalities on cholesterol and the lipid profile: review, synthesis and recommendations. *Sports Med* 2014;44:211–21.
- 35. Zhao M, Veeranki SP, Magnussen CG *et al.* Recommended physical activity and all cause and cause specific mortality in US adults: prospective cohort study. *BMJ* 2020;370:m2031.
- Shakespear-Druery J, De Cocker K, Biddle SJH et al. Assessment of muscle-strengthening exercise in public health surveillance for adults: A systematic review. Prev Med 2021; 148:106566.

© The Author(s) 2024. Published by Oxford University Press on behalf of the International Epidemiological Association.

This is an Open Access article distributed under the terms of the Creative Commons Attribution License (https://creativecommons.org/licenses/by/4.0/), which permits unrestricted reuse, distribution, and reproduction in any medium, provided the original work is properly cited. International Journal of Epidemiology, 2024, 53, 1–8 https://doi.org/10.1093/ije/dyae074 Original article