Association of walking pace and handgrip strength with all-cause, cardiovascular and cancer mortality: A UK Biobank observational study

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ABSTRACT

Aims: To quantify the association of self-reported walking pace and handgrip strength with all-cause, cardiovascular, and cancer mortality.

Methods and Results: 230670 women and 190057 men free from prevalent cancer and cardiovascular disease were included from UK Biobank. Usual walking pace was self-defined as slow, steady/average or brisk. Handgrip strength was assessed by dynamometer. Cox-proportional hazard models were adjusted for social deprivation, ethnicity, employment, medications, alcohol use, diet, physical activity, and TV viewing time. Interaction terms investigated whether age, BMI and smoking status modified associations. Over 6.3 years there were 8598 deaths, 1654 from cardiovascular disease and 4850 from cancer. Associations of walking pace with mortality were modified by BMI. In women, the hazard ratio (HR) for all-cause mortality in slow compared to fast walkers were 2.16 (95% CI: 1.68, 2.77) and 1.31 (1.08, 1.60) in the bottom and top BMI tertiles, respectively; corresponding HRs for men were 2.01 (1.68, 2.41) and 1.41 (1.20, 1.66). HRs for cardiovascular mortality remained above 1.7 across all categories of BMI in men and women, with modest heterogeneity in men. Handgrip strength was associated with cardiovascular mortality in men only (HR tertile 1 vs tertile 3 = 1.38; 1.18, 1.62), without differences across BMI categories, while associations with all-cause mortality were only seen in men with low BMI. Associations for walking pace and handgrip strength with cancer mortality were less consistent.

Conclusions: A simple self-reported measure of slow walking pace could aid risk stratification for allcause and cardiovascular mortality within the general population.

Keywords: Cardiorespiratory fitness, handgrip strength, mortality, walking pace

INTRODUCTION

Physical fitness, composed of cardiorespiratory fitness and muscle strength, is an important predictor of mortality and an established target for cardiovascular disease prevention.¹⁻³ Cardiorespiratory fitness, in particular, has shown a strong graded association with the risk of allcause and cardiovascular mortality and is advocated as a clinical vital sign with importance for risk classification.^{1,4} Therefore, pragmatic methods of harnessing cardiorespiratory fitness for risk stratification are warranted. Studies have found an association between objectively assessed walking pace and mortality, mostly in older adult,⁵⁻⁷ with other studies suggesting self-reported walking pace may act as an important marker of cardiovascular health.^{8,9} Thus, self-reported walking pace could function as a pragmatic measure of physical fitness and a simple index of mortality risk. However, in order to have clinical and public health relevance, research is needed to examine the extent to which pragmatic self-reported measures of walking pace are associated with cardiorespiratory fitness and whether the associations with all-cause and cause-specific mortality are maintained after adjustment for potential confounding variables or vary across potential effect modifiers such as age, smoking status or body mass index (BMI).

Muscle strength is also an important and reproducible measure of physical fitness and has been associated with all-cause and cardiovascular mortality.² Handgrip strength is correlated with overall muscle strength and is associated with functional and health outcomes including frailty, falls, nutritional status and mortality.¹⁰⁻¹⁵ Investigating whether associations persist after adjustment for other pragmatic measures of physical fitness, such as walking pace, and maintained across potential effect modifiers will further help elucidate the potential clinical importance of handgrip strength within the general population.

The aim of this analysis was to quantify the associations of self-reported walking pace and handgrip strength with all-cause, cardiovascular and cancer mortality in men and women from UK Biobank

and to investigate whether associations with all-cause mortality are maintained across categories of age, BMI and smoking status.

METHODS

UK Biobank

This analysis used data from 502,639 participants within UK Biobank, the methods and aim of which have been reported elsewhere¹⁶. In brief, UK Biobank is a large prospective cohort of middle-aged adults designed to support biomedical analysis focused on improving the prevention, diagnosis and treatment of chronic disease. Between March 2006 and July 2010, individuals living within 25 miles of one of the 22 study assessment centres located throughout England, Scotland and Wales were recruited and provided comprehensive data on a broad range of demographic, clinical, lifestyle, and social outcomes. All participants provided written informed consent and the study was approved by the NHS National Research Ethics Service (Ref: 11/NW/0382).

For each variable reported in this analysis, we provide the UK Biobank Data-Field (DF) number. Each DF is linked to detailed information on measurement procedures through the UK Biobank website.¹⁷

Measures of physical function

The primary exposures of interest were self-reported walking pace and objectively assessed handgrip strength. Walking pace was assessed by the following single item question: "How would you describe your usual walking pace? 1) Slow pace, 2) Steady/ average pace, 3) Brisk pace" (DF 924). Handgrip strength was assessed through the use of a hydraulic hand dynamometer (Jamar J00105) whilst sitting (DF 46 and 47). The elbow of the arm holding the dynamometer was placed against the side of the body and bent to a 90° angle with the forearm placed on an armrest. Participants were instructed to squeeze the handle of the dynamometer as hard as they could for 3

seconds. Both left and right hand strengths were measured. The values for the right and left hand were summed and divided by two to estimate an average.

Cardiorespiratory fitness

A subsample of UK Biobank undertook a 6 minute graded submaximal fitness test on a cycle ergometer (DF 6025). We utilised data from this subsample to inform the main analyses by investigating the association of cardiorespiratory fitness with walking pace and handgrip strength. The start and end workload of the graded fitness test was set according to age, height, weight, resting heart rate and sex to ensure a similar relative intensity across the population. Heart rate was monitored before, during, and after the exercise test via a 4-lead ECG. Maximal cardiorespiratory fitness was estimated by: 1) Fitting a linear regression line between heart rate and power output for each stage of the test; 2) The regression line was extrapolated to the age-predicted maximal heart rate with the formula $208 - 0.7 \times age;^{18}$ 3) Maximal oxygen uptake was estimated from the regression equation for the relationship between work rate (power) and oxygen uptake (oxygen uptake [ml.kg⁻¹.min⁻¹] = 7 + (10.8 x work rate [Watts])/body mass [kg]).¹⁹ Individuals without cancer and cardiovascular disease who completed at least one stage of the fitness test were included for this sub-analysis (n = 51282).

Data on potential confounders, mediators and effect modifiers

This study utilised the following covariates: anthropometric (BMI); demographic (age, sex, ethnicity, social deprivation [Townsend index], and employment status); health (number of medications); and lifestyle (smoking [never, past, current], alcohol, diet, TV viewing time and self-reported physical activity). Diet was assessed through a food frequency questionnaire; alcohol (DF 1558), fresh fruit (DF 1309), cooked vegetable (DF 1289) and processed meat (DF 1349) were included. Physical activity was assessed as weekly frequency of walking (DF 864), moderate (DF 884) or vigorous (DF 904) intensity physical activity lasting at least 10 minutes in duration. Further details for each variable are available on the UK Biobank Website²⁰.

Mortality status follow-up

UK Biobank undertook comprehensive data linkage for mortality status. Date and cause of death were obtained from the National Health Service (NHS) Information Centre for participants from England and Wales, and from the NHS Central Register, Scotland for participants from Scotland. Detailed information about the linkage procedure is available online²⁰. Linkage captured all deaths occurring until 31st January 2016 for England and 30th November 2015 for Scotland. We defined cancer and cardiovascular disease mortality using the International Classification of Diseases edition 10 (ICD-10): C00-C97 for cancer death and I00-I79 for cardiovascular death.

Participants included

From the initial sample of 502,639 participants, those with prevalent cancer (number of selfreported cancers [DF 134]; n = 41706) or prevalent cardiovascular disease (defined as myocardial infarction, stroke, or angina [DF 6150]; n = 27573) were excluded from the analysis. From the remaining sample, only subjects with information available for all covariates were included (missing data were less than 2% for all included covariates), leaving 420,727 individuals who at baseline were free from cancer and cardiovascular disease and had complete walking pace, handgrip strength (both left and right measures) and covariate data (eFigure 1 in the Supplement).

Statistical analysis

In the subsample with cardiorespiratory fitness data, the association between walking pace, handgrip strength and cardiorespiratory fitness was tested using analysis of variance models adjusted for age and stratified by sex.

In the full cohort, Cox-proportional hazard models were used to investigate the association of walking pace and handgrip strength with mortality using age as time scale. Handgrip strength was categorised into sex-specific tertiles and separate models were undertaken for all-cause mortality, cardiovascular and cancer mortality. The proportional hazards assumption was verified for all variables by inspection of the plots of the Schoenfeld residual for covariates. Models were arranged

a priori to investigate the impact of incremental adjustment. Model 1 was unadjusted. Model 2 adjusted for ethnicity, social deprivation, employment status, and number of prescribed medication. Model 3 additionally adjusted for smoking, alcohol consumption, and fruit, vegetable, and processed meat intake. Model 4 additionally adjusted for BMI, TV viewing, and frequency of walking, moderate-intensity, and vigorous-intensity physical activity. Lastly, model 5 was additionally mutually adjusted for walking pace and handgrip strength. For the regression models including all covariates in Model 4 plus walking pace or handgrip strength, the C-indexes were calculated and calibration plots displayed.

Interaction terms were fitted to Model 5 to assess whether BMI (as tertiles), age (tertiles) and smoking (never, past, current) modified associations with mortality; in addition the interaction between walking pace and grip strength was also tested. A sensitivity analysis was also conducted to investigate the effect of removing all deaths occurring within the first 24 months of follow-up (women = 424, men = 797) to reduce the possible impact of reverse causation.

All analyses were performed with Stata V14.1 (Stata Corporation, College Station, Texas, USA). Results stratified by sex are reported with 95% CI; a two-sided p-value <0.05 was considered statistically significant.

RESULTS

The mean (range) age of included participants was 56.4 (38.9 - 73.7) years. The distribution of the categorical covariates for the 230,670 women and 190,057 men included in the analyses are displayed in Figure 1. Table 1 shows BMI, age and mean handgrip strength values across categories of walking pace and handgrip strength. Categories of walking pace were similarly distributed between men and women. However, mean values of handgrip strength for sex-specific tertiles showed little overlap between men and women; those in the highest tertile for women had a mean (SD) value of 30.4 (3.3) kg whereas men in the lowest tertile had a mean value of 30.6 (4.9) kg (Table 1).

In the subsample who undertook the fitness test, self-reported walking pace showed a strong graded association with cardiorespiratory fitness in men and women (Table 2). Slow walkers achieved an average fitness level of 28.1 (27.7, 28.6) ml·kg⁻¹·min⁻¹ and 35.9 (35.3, 36.5) ml·kg⁻¹·min⁻¹ in women and men, respectively; whereas brisk walkers achieved levels of 34.0 (33.8, 34.1) ml·kg⁻¹·min⁻¹ and 42.0 (41.8, 42.2) in men and women, respectively. Associations between handgrip strength and cardiorespiratory fitness were less pronounced.

Table 3 reports the number of deaths for men and women by walking pace group and tertile of handgrip strength. During a median (IQR) follow-up of 6.3 [7.0-7.6] years [2,926,917 person-years], 8598 (2.0%) all-cause deaths, 4850 (1.2%) cancer deaths, and 1654 (0.4%) cardiovascular deaths occurred. The number of all-cause, cancer and cardiovascular deaths were greater in men (5184, 2682, 1188 respectively) than women (3414, 2168, and 466 respectively).

The hazard ratios (HRs) for walking pace and handgrip strength are shown graphically in Figure 2 with corresponding estimates presented in eTable 1 and eTable 2. Walking pace was associated with all-cause and cardiovascular mortality, but not cancer mortality, in women and men across all models. Handgrip strength was associated with cardiovascular mortality in men and with all-cause mortality in women and men, although results for women were attenuated after removing deaths occurring within the first 24 months. Handgrip strength was not associated with cancer mortality in women, whereas in men a weaker handgrip appeared to be associated with a lower risk of cancer, particularly after removing deaths occurring within the first 24 months occurring within the first 24 months.

BMI was found to modify associations for walking pace in men and women and handgrip strength in men. Interaction tests and stratified results are displayed in Figure 3. For women, the hazard ratio (HR) for all-cause mortality in slow walkers compared to fast walkers ranged from 2.16 (1.68, 2.77) to 1.31 (1.08, 1.60) for participants in the bottom and top BMI tertiles, respectively. Corresponding HRs for men were 2.01 (1.68, 2.41) and 1.41 (1.20, 1.66). HRs for cardiovascular mortality for slow walkers compared to fast walkers remained above 1.7 across all categories of BMI in men and

women, with a modest heterogeneity in men. A slow walking pace was also associated with cancer mortality in women and men with low BMI. Weak handgrip strength was only associated with a higher risk of all-cause mortality in men with low BMI (1.29; 1.13, 1.48) whereas associations between handgrip strength and cardiovascular mortality were not affected by BMI status.

Interaction analysis revealed the association of walking pace with all-cause (p= 0.572 for women; p = 0.626 for men) and cardiovascular mortality (p= 0.068 for women; p = 0.634 for men) were not modified by handgrip strength or vice-versa. In addition, the association of walking pace and handgrip strength with all-cause or cardiovascular morality were not modified by age or smoking status.

The C-indexes for the most adjusted models containing walking pace or handgrip strength demonstrated moderate discrimination (ranging from 0.63 to 0.71), with good calibration (eFigure 2).

DISCUSSION

Within a large national sample of UK adults free from cancer and cardiovascular disease, a simple measure of self-perceived walking pace was associated with a higher risk of all-cause and cardiovascular mortality; the strength of association persisted after adjustment for handgrip strength and was modified by levels of BMI. For those with low BMI, slow walkers had over twice the risk of cardiovascular and all-cause mortality compared to fast walkers in men and women. Associations were weaker at higher levels of BMI, yet slow walkers still had over 1.7 times the risk of cardiovascular mortality compared to fast walkers. Handgrip strength was not consistently associated with mortality in women, whereas in men a weak handgrip was associated with a higher risk of all-cause mortality within the lowest tertile of BMI and with cardiovascular mortality across all categories of BMI. There was no clear pattern of association of walking pace or handgrip strength with cancer mortality, although a slow walking pace was associated with a higher risk of cancer mortality in women and men with low BMI.

The strength of association with all-cause mortality reported in the present study is consistent with previous research in smaller cohorts using objective measures of walking pace, primarily in older adults. A recent meta-analysis including a total of 12,901 subjects across 9 studies reported a higher risk of all-cause mortality for the slowest walkers compared to the fastest walkers (HR = 1.89 [1.46, 2.46]) when measured objectively⁶. The Whitehall II study reported similar finding in middle aged adults⁵. Other studies have also investigated the association between self-reported walking pace and cardiovascular events in occupational cohorts.^{8,9,21} For example, in the Women's Health Study of health care professionals in the United States and Puerto Rico, those reporting a fast walking pace had around half the risk of a cardiovascular event compared to slow walkers⁹. Similar results were seen for men in the Health Professionals' Follow-up Study.⁸ Self-reported measures of physical fitness have also been associated with frailty and health outcomes in community dwelling cohorts or older adults.^{22,23}

Our findings extend previous evidence by reporting the novel finding that a simple semi-quantitative measure of walking pace that asks individuals to rate their walking pace as slow, steady/average or brisk is associated with objectively measured cardiorespiratory fitness and by evidencing that this simple measure is associated with mortality within the general population. A further novel finding revealed the strength of the association of walking pace with all-cause (men and women), cardiovascular (men) and cancer (men) mortality was modified by BMI, with stronger associations seen in the lowest tertile of BMI. Low levels of BMI are associated with malnutrition and higher levels of sarcopenia,^{24,25} therefore our results suggest the potential importance of self-reported walking pace in differentiating between healthy and unhealthy forms of low BMI.

Associations between handgrip strength and mortality are somewhat weaker than those reported in smaller studies which did not stratify by sex ^{13,26,27}. As this study demonstrated, there is very little overlap in the distribution of handgrip strength between men and women. Therefore, conventional methods of adjustment for sex may not fully account for this difference.

This study did not find a clear pattern of association for walking pace or handgrip strength with the risk of cancer mortality in men or women. However, unexpectedly a weaker handgrip appeared to be associated with a slightly lower risk of cancer in men, a finding that was strengthened after removing deaths occurring within the first 24 months. To date, the literature in this area has been discrepant. Some studies have reported an association between higher levels of physical fitness and lower cancer mortality risk,^{2,28-30} while others have reported no association despite finding clear associations with cardiovascular mortality.^{14,15,31} Of note, another large population study also reported the finding that weaker handgrip strength was associated with an lower risk of cancer mortality.¹³ The equivocal nature of the evidence may reflect factors like age and the predominant types of cancer within the measured population; however, based on this study and the literature to date it is likely that simple indices of physical performance, such as walking pace and handgrip strength, are more reflective of cardiovascular health within the general population. The lower risk of cancer mortality with lower muscular strength reported for men in this study and in men and women in another large study warrants further investigation.¹³

Our results are supported by interventional research. Physical fitness, which was strongly associated with walking pace in the present study, is modifiable with exercise training. Although inter-individual responses to exercise training are heterogeneous, the majority gain some improvement. For example, in a population of healthy, pre-frail and frail older adults (>65 years), all individual increased their functional capacity following an exercise intervention.³² Other studies have shown that the majority of adults respond with increased cardiorespiratory fitness following aerobic exercise training.^{33,34} Importantly, exercise training has also been shown to reduce the risk of chronic disease and improve cardiometabolic risk factors,³⁵ including type 2 diabetes,³⁶ blood pressure,^{37,38} and cholesterol.³⁹ The fact that physical fitness is modifiable through exercise intervention and that exercise interventions in turn improve health status provide a robust mechanism that supports the association between measures of physical fitness and cardiovascular mortality reported in this study.

The present study has important strengths, including the large sample size and number of deaths which allowed stratification by sex. The extensively phenotyped population also allowed a comprehensive investigation into possible confounding or mediating influences on the association between walking pace, handgrip strength and mortality. However, important limitations remain. Although participants with cancer, myocardial infarction, stroke, or angina were excluded from the analysis, other potentially relevant conditions, such as a history of coronary artery bypass grafting or revascularization were not captured and reverse causality remains a possible explanation of our findings, including the interactions with BMI. Moreover, although we adjusted for a wide range of demographic, clinical and lifestyle factors, many covariates were self-reported, thus residual confounding due limited measurement precision also remains a possibility. This analysis was also limited to mortality end-points.

Conclusions

In conclusion, this study found that a slow self-reported walking pace was associated with a higher risk of all-cause and cardiovascular mortality in women and men with associations remaining robust across multiple layers of adjustment. Association were stronger in those with low BMI. Self-reported walking pace could therefore represent an important measure of physical fitness with potential applicability for risk stratification within the general population, particularly those presenting with low BMI. Future research is needed to investigate the extent to which walking pace can be used to improve the prognostic performance of common cardiovascular risk scores across different categories of BMI. Compared to self-reported walking pace, handgrip strength appeared to be a less generalizable marker of risk within the general population.

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Conflicts of Interest: None

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Data source: This research has been conducted using the UK Biobank Resource under Application

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Figure 1: Distribution in participant characteristics for men and women across tertiles of handgrip strength and categories of walking pace

Figure 2: Association of walking pace and handgrip strength with all-cause, cancer, and cardiovascular mortality

Figure 3: Association of walking pace and handgrip strength with all-cause, cardiovascular and cancer mortality across categories of BMI

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Table 1: Age and BMI characteristics across categories of walking pace and handgrip strength

		Women		Men				
	Grip Third							
	First	Second	Third	First	Second	Third		
Age (years)	59.1 (7.3)	56.7 (7.8)	53.1 (7.7)	58.8 (7.9)	56.8 (8.0)	53.8 (7.9)		
Body mass index (kg/m ²)	27.3 (5.2)	26.8 (5.0)	26.8 (5.1)	27.6 (4.4)	27.6 (4.1)	28.0 (4.0)		
Grip strength (kg)	17.2 (3.6)	23.9 (1.4)	30.4 (3.3)	30.6 (4.9)	39.9 (2.0)	49.3 (4.8)		
Walking pace								
Slow	9850 (11.8%)	3935 (5.4%)	2636 (3.5%)	6370 (9.9%)	2956 (4.8%)	2055 (3.2%)		
Average	47557 (57.2%)	38744 (53.6%)	35980 (47.9%)	36250 (56.1%)	32244 (52.5%)	31227 (48.8%)		
Brisk	25759 (31.0%)	29648 (41.0%)	36561 (48.6%)	22014 (34.1%)	26180 (42.7%)	30761 (48.0%)		
	Walking Pace							
	Slow	Average	Brisk	Slow	Average	Brisk		
Age (years)	58.2 (7.7)	57.0 (8.0)	55.2 (8.0)	58.8 (7.8)	57.1 (8.2)	55.4 (8.2)		
Body mass index (kg/m ²)	31.9 (7.1)	27.7 (5.0)	25.1 (3.8)	30.3 (5.8)	28.2 (4.2)	26.7 (3.5)		
Grip strength (kg)	19.6 (6.9)	23.2 (6.0)	24.9 (5.9)	34.7 (10.0)	39.4 (8.6)	41.2 (8.4)		

Numbers are mean (SD) or number (%).

Table 2: Age adjusted cardiorespiratory fitness across walking speed and handgrip categories

Women (n = 26593)								
Walking speed	Slow	Steady	Brisk	Р				
Mean fitness (ml/kg.min)	28.1 (27.7, 28.6)	30.9 (30.7, 31.0)	34.0 (33.8, 34.1)	< 0.001				
Hand grip strength	Tertile 1	Tertile 2	Tertile 2					
Mean fitness (ml/kg.min)	31.4 (31.2, 31.6)	32.2 (32.0, 32.4)	32.9 (32.7, 33.0)	< 0.001				
	Me	en (n = 24689)						
Walking speed	Slow	Steady	Brisk	Р				
Mean fitness (ml/kg.min)	35.9 (35.3, 36.5)	39.3 (39.1, 39.5)	42.0 (41.8, 42.2)	< 0.001				
Hand grip strength	Tertile 1	Tertile 2	Tertile 2					

40.6 (40.4, 40.8)

Data as marginal mean (adjusted for age) (95% CI)

39.5 (39.3, 39.7)

Mean fitness (ml/kg.min)

41.1 (40.9, 41.3)

<0.001

Table 3: Number of mortalit	v events across catego	ries of walking pace	and handgrip stren	gth in women and men
				0

	Women					Men				
		N	Mortality events			Mortality events				
Exposure	Participants	All-cause	Cancer	Cardiovascular	Participants	All-cause	Cancer	Cardiovascular		
Grip strength										
First third	83166	1644 (2.0%)	967 (1.2%)	244 (0.3%)	64634	2465 (3.8%)	1151 (1.8%)	591 (0.9%)		
Second third	72327	1032 (1.4%)	680 (0.9%)	128 (0.2%)	61380	1535 (2.5%)	825 (1.3%)	363 (0.6%)		
Third third	75177	738 (1.0%)	521 (0.7%)	94 (0.1%)	64043	1184 (1.8%)	706 (1.1%)	234 (0.4%)		
Walking pace										
Slow	16421	563 (3.4%)	239 (1.5%)	108 (0.7%)	11381	853 (7.5%)	287 (2.5%)	245 (2.2%)		
Average	122281	1866 (1.5%)	1237 (1.0%)	247 (0.2%)	99721	2898 (2.9%)	1553 (1.6%)	660 (0.7%)		
Brisk	91968	985 (1.1%)	692 (0.8%)	111 (0.1%)	78955	1433 (1.8%)	842 (1.1%)	283 (0.4%)		
Total	230670	3414 (1.5%)	2168 (0.9%)	466 (0.2%)	190057	5184 (2.7%)	2682 (1.4%)	1188 (0.6%)		
Excluding first 2 years	230246	2990 (1.3%)	1936 (0.8%)	382 (0.2%)	189260	4387 (2.3%)	2338 (1.2%)	960 (0.5%)		



Figure 1: Distribution in participant characteristics for men and women across tertiles of handgrip strength and categories of walking pace

Ethnic group		White	Non-White								
Grip third		1st	2nd	3rd							
Walking pace		Slow	Average	Brisk							
Smoking status		Never	Previous	Current	Not answered						
Alcohol use (frequency)		Never	1-3/month	1-2/week	3-4/week	Daily	Special occasions	Not answered			
Medications (n)		0	<mark>_</mark> 1	2	3	4	5	6	7+		
Processed meat intake (f	requency)	Never	<1/week	1/week	2-4/week	5-6/week	1+/daily	Unknown	Not answered		
Watching TV (hr/day)		0	1	2	3	4+	Unknown	<1	Not answered		
Cooked vegetables (spor	on/day)	0	1	2	3	4	5+	Unknown	_ <1	Not answered	
Current employment		Unpaid	Student	Paid/self-employe	d 📕 Family care	Other	Not answered	Retired	Unable	Unemployed	
Fresh fruit intake (piece/o	day)	0	1	2	3	4	5+	Unknown	<1	Not answered	
Townsend score (tenth)		1st	2nd	3rd	4th	5th	6th	7th	8th	= 9th	10th
Discrete all and the	Walking	0	1	2	3	4	5	6	7	Unknown	Not answered
(day/wook 10+ mine)	Moderate	0	1	2	3	4	5	6	7	Unknown	Not answered
(day week 10+ mins)	Vigorous	0	1	2	3	4	5	6	7	Unknown	Not answered



Figure 2: Association of walking pace and handgrip strength with all-cause, cancer, and cardiovascular mortality

Model adjustment

Model 1: Unadjusted

Model 2: Townsend score, ethnicity, employment, number of medications

▲ Model 3: Model 2 + smoking, alcohol use, diet (fresh fruit, cooked vegatable, and processed meat intake)

Model 4: Model 3 + time TV viewing, physical activity (walking, moderate, vigourous), body mass index

Model 5: Model 4 + walking pace (grip strength analysis) or grip strength (walking pace analysis)
Model 5 + exclusion of deaths occurred in the first two years

References (hazard ratio 1, dotted lines) are the third teriles for grip strength (highest) and "brisk" pace for walking pace. For grip strength, the first estimate refers to the second tertile and the second estimate to the first tertile (lowest grip strength). For walking pace, the first estimate refers to "average" pace and the second to "slow" pace. Spikes indicate 95%Cl.

Figure 3: Association of walking pace and handgrip strength with all-cause, cardiovascular and cancer mortality across categories of BMI



Mean (SD) of body mass index was: 22.2 (1.6), 26.2 (1.1), and 32.8 (4.3) Kg/m² in the bottom, middle, and top tertile in women; 23.7 (1.6), 27.3 (0.9), and 32.3 (3.4) Kg/m² in the corresponding tertiles in men. Hazard ratios adjusted for variables included in Model 5: Townsend score, ethnicity, employment, number of medications, smoking, alcohol use, diet (fresh fruit, cooked vegatable, processed meat intake), time TV viewing, physical activity (walking, moderate, vigoorusus), body mass index (tertiles), walking pace earloss the three levels of body mass index.