Supplemental Online Content

del Pozo Cruz B, Ahmadi MN, Lee IM, Stamatakis E. Prospective associations of daily step counts and intensity with cancer and cardiovascular disease incidence and mortality and all-cause mortality. *JAMA Intern Med.* Published online September 12, 2022. doi:10.1001/jamainternmed.2022.4000

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eFigure 27. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD mortality (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).

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eFigure 30. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites incidence outcome (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).

This supplemental material has been provided by the authors to give readers additional information about their work.

	Accelerometer wearing	
	No	Yes
Sample size	398808	103684
Age (years)	56.65	56.05
Sex = male (%)	(8.16) 183714	(7.84) 45401
Sex = Inale (70)	(46.1)	(43.8)
Ethnicity = non-white (%)	49265	9755
	(12.4)	(9.4)
Education = university/college (%)	272178	58031
	(68.2)	(56.0)
Townsend Index of Deprivation (lower scores indicate more affluence)	-1.19	-1.71
	(3.15)	(2.83)
Smoking (%)		
Never	214554	58962
	(54.2)	(57.0)
Previous	135883	37168
	(34.3)	(35.9)
Current	45700	7277
	(11.5)	(7.0)
Alcohol use (%) ^a		
Never	19381	3004
	(4.9)	(2.9)
Previous	15244	2858
Occasional	(3.9) 92832	(2.8) 21029
Occasional	(23.6)	(20.4)
Within guidelines	126547	37598
	(32.1)	(36.5)
Double guidelines	81874	24180
8	(20.8)	(23.4)
Above double guidelines	58008	14465
-	(14.7)	(14.0)
Fruit consumption (servings/day)	3.05	3.21
	(2.64)	(2.50)
Vegetable consumption (servings/day)	4.92	4.91
	(3.41)	(3.17)
Family history of CVD = yes (%)	212025	56790
	(53.2)	(54.8)
Family history of Cancer = yes (%)	100692	26028
Cholesterol medication = yes (%)	(25.2) 75238	(25.1)
(10)	(18.9)	(14.4)
Insulin medication = yes (%)	5534	716
	(1.4)	(0.7)
Hypertension medication = yes (%)	40151	7766
	(10.1)	(7.5)
Body mass index (Kg/m ²)	27.62	26.73
	(4.85)	(4.55)
Self-reported health (0-4)	2.16	1.99
	(0.79)	(0.71)

^aGuidelines for alcohol use in the UK recommend no more than 14 units of alcohol per week for both men and women.

eTable 2. Stepping-based exposure variables definition				
Variable	Definition			
Total steps per day	Average of steps accumulated in a day.			
Walking steps per day	Average of steps accumulated in a day while walking (i.e. excludes running).			
Peak 30-minute cadence ^{1,2}	Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day.			
Incidental steps	Total daily steps at 1-39 steps/minute.			
Purposeful steps	Total daily steps at ≥ 40 steps/minute.			
Light intensity	Total daily steps at <100 steps/minute.			
Moderate intensity	Total daily steps at ≥ 100 and < 130 steps/minute.			
Vigorous intensity	Total daily steps at ≥130 steps/minute.			
Moderate-to-vigorous intensity	Total daily steps at ≥100 steps/minute.			
1.Lee IM, Shiroma EJ, Kamada M, Bassett DR, Matthews CE, Buring JE. Association of Step Volume and Intensity With All-Cause Mortality in Older Women. JAMA Intern Med. Published online May 29, 2019. doi:10.1001/jamainternmed.2019.0899 2.Tudor-Locke C, Ducharme SW, Aguiar EJ, et al. Walking cadence (steps/min) and intensity in 41 to 60-year-old adults: the CADENCE- adults study. Int J Behav Nutr Phys Act. 2020;17(1):137.				

eTable 3. Assessment of CVD and cancer	r incidence (fatal and non-fatal) and definition of		
diseases			
Variable	Definition		
Inpatient hospitalisation	The inpatient hospitalization data were		
	provided by either the Hospital Episode		
	Statistics for England, the Patient Episode		
	Database for Wales, or the Scottish		
	Morbidity Record for Scotland		
Cancer registry	Cancer data linkage was obtained through		
	national cancer registries. For England and		
	Wales, cancer diagnosis data were provided		
	by the Medical Research Information		
	Service, based at the National Health		
	Service Information Centre. For Scotland,		
	cancer diagnosis data were provided by the		
	Information Services Division, which is part		
	of NHS Scotland		
Cardiovascular disease definition	CVD was defined as coronary heart disease,		
	stroke, and heart failure using ICD-10		
	classification: I0, I11, I13, I20-I51, I60-I69.		
Cancer definition	Physical activity-related cancer was defined		
	as 13 site specific cancers (esophageal		
	adenocarcinoma, liver, lung, kidney, gastric		
	cardia, endometrial, myeloid leukemia,		
	myeloma, colon, head and neck, rectal,		
	bladder, and breast) using ICD-10		
	classifications: C0-C3, C5-C9, C40-C42,		
	C45-C49.		

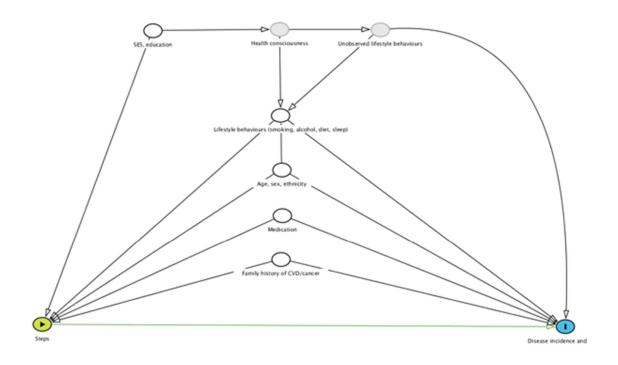
eTable 3 Assessment of CVD and cancer incidence (fatal and non-fatal) and definition of

eTable 4. Tabulation of the number of events per
physical activity-related ³ cancer site considered in
this study

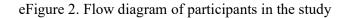
	Fatal	Non-fatal
Esophageal adenocarcinoma	89	65
Liver	47	69
Lung	169	209
Kidney	37	95
Gastric Cardia	9	39
Endometrial	21	130
Myeloid Leukemia	27	29
Myeloma	25	85
Colon	57	416
Head and neck	14	58
Rectal	16	72
Bladder	36	65
Breast	63	756
3. Moore SC, Lee IM, Weiderp	ass E, et a	1.
Association of Leisure-Time P	hysical Ac	tivity With
Risk of 26 Types of Cancer in	1.44 Millio	on Adults.

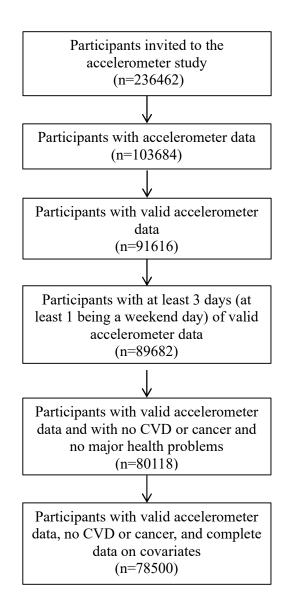
JAMA Intern Med. 2016;176(6):816-825.

eFigure 1. A priori Defined Directed Acyclic Graph



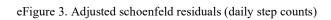
SES, socioeconomical status. The directed acyclic graph above was generated via an opensource application (Johannes Textor, Benito van der Zander, Mark K. Gilthorpe, Maciej Liskiewicz, George T.H. Ellison. Robust causal inference using directed acyclic graphs: the R package 'dagitty'. *International Journal of Epidemiology* 45(6):1887-1894, 2016. <u>https://doi.org/10.1093/ije/dyw341</u>).

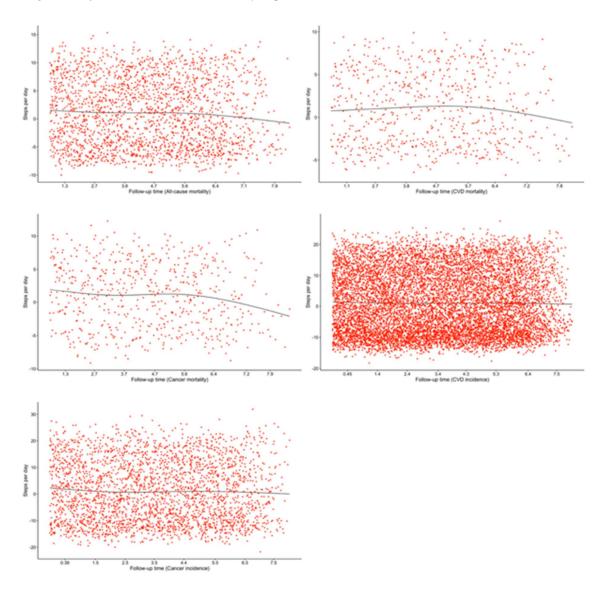


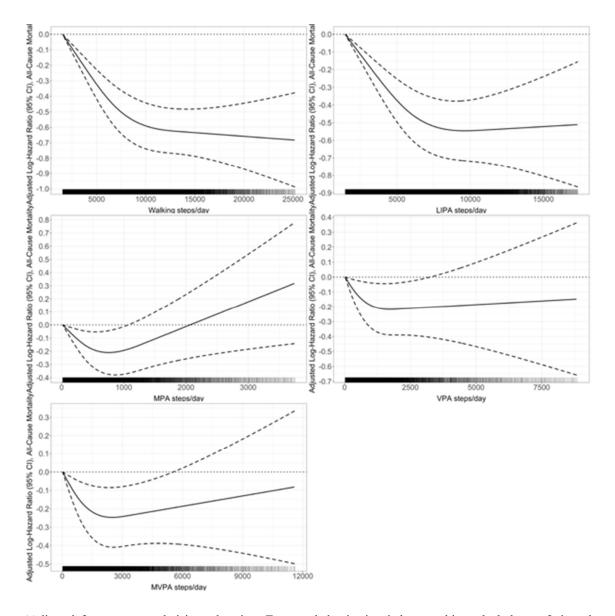


eTable 5. Characteristics of accelerometer sample excluded from analysis	5*
Sample size	25184
Age (years)	61.51 (7.92)
Sex = male (%)	10264
	(40.8)
Ethnicity = non-white (%)	2669 (10.6)
Education = university/college (%)	
Townsend Index of Deprivation (lower scores indicate more affluence)	-1.6 (2.9)
Smoking (%)	
Never	13794 (55.4)
Previous	9259 (37.2)
Current	1854 (7.4)
Alcohol use (%) ^a	
Never	808 (3.3)
Previous	766 (3.1)
Occasional	5169 (21.0)
Within guidelines	8854 (35.9)
Double guidelines	5661 (23.0)
Above double guidelines	3376 (13.7)
Fruit consumption (servings/day)	3.2 (2.6)
Vegetable consumption (servings/day)	5.0 (3.3)
Family history of CVD = yes (%)	13,777 (54.7)
Family history of Cancer = yes (%)	6431 (25.5)
Cholesterol medication = yes (%)	3968 (15.8)
Insulin medication = yes (%)	187 (0.7)
Hypertension medication = yes (%)	2173 (8.6)
Sleep (accelerometer-measured, min/day)	408.8
	(100.4)
Accelerometer wear days	6.6 (1.0)
Total steps per day	6915.06 (4,645.52)
Walking steps per day	5817.73
	(3,389.56)
Incidental steps per day	3185.80
Purposeful steps per day	(1,324.19) 2631.93
i uiposeiui steps pei uay	(2,328.58)
LIPA steps per day	6256.94
	(3,429.85)
MPA steps per day	556.96 (742.07)
VPA steps per day	727.38
	(1,598.02)
MVPA steps per day	1284.34
Deals 20 min and an an (stans/min)	(2124.28)
Peak 30-min cadence (steps/min)	36.3 (15.6)

*Due to missing data, values may not equal total sample size entered in the main analyses

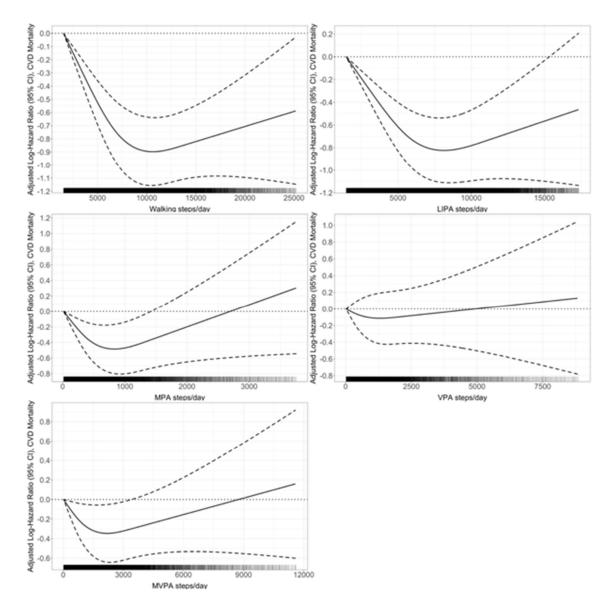






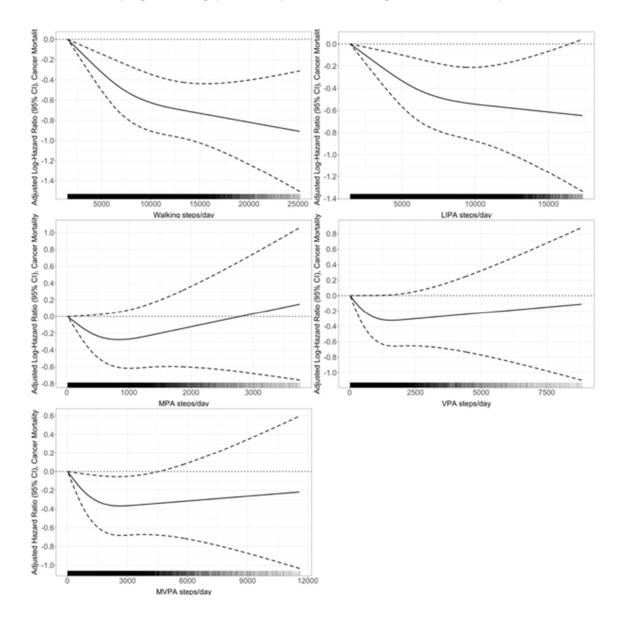
eFigure 4. Dose–response association (Adjusted^{*a*} hazard ratios and associated 95% confidence interval band) between secondary exposures and all-cause mortality.

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For LIPA steps, models were further adjusted for MPA and VPA steps (and vice versa). For MVPA steps, models were adjusted for LIPA steps. CVD, Cardiovascular disease; LIPA, Light intensity physical activity; MPA, Moderate intensity physical activity; VPA, Vigorous intensity physical activity; MVPA, Moderate-to-vigorous physical activity; LIPA steps/day, Total daily steps at <100 steps/minute; MPA steps/day, Total daily steps at \geq 100 and < 130 steps/minute. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



eFigure 5. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between secondary exposures and CVD mortality.

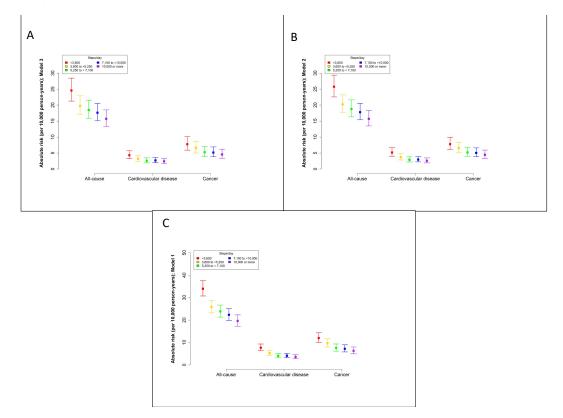
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For LIPA steps, models were further adjusted for MPA and VPA steps (and vice versa). For MVPA steps, models were adjusted for LIPA steps. CVD, Cardiovascular disease; LIPA, Light intensity physical activity; MPA, Moderate intensity physical activity; VPA, Vigorous intensity physical activity; MVPA, Moderate-to-vigorous physical activity; LIPA steps/day, Total daily steps at <100 steps/minute; MPA steps/day, Total daily steps at \geq 100 and < 130 steps/minute. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



eFigure 6. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between secondary exposures and physical activity-related cancer composite 13 sites mortality outcome.

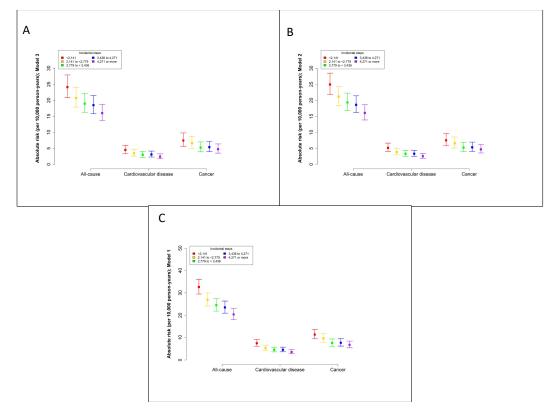
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For LIPA steps, models were further adjusted for MPA and VPA steps (and vice versa). For MVPA steps, models were adjusted for LIPA steps. CVD, Cardiovascular disease; LIPA, Light intensity physical activity; MPA, Moderate intensity physical activity; VPA, Vigorous intensity physical activity; MVPA, Moderate-to-vigorous physical activity; LIPA steps/day, Total daily steps at <100 steps/minute; MPA steps/day, Total daily steps at ≥100 and < 130 steps/minute. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 7. Sequentially adjusted dose–response quintile-based absolute risk describing the associations between daily steps and mortality outcomes (events rate per 10000 person-years and associated 95% confidence intervals).



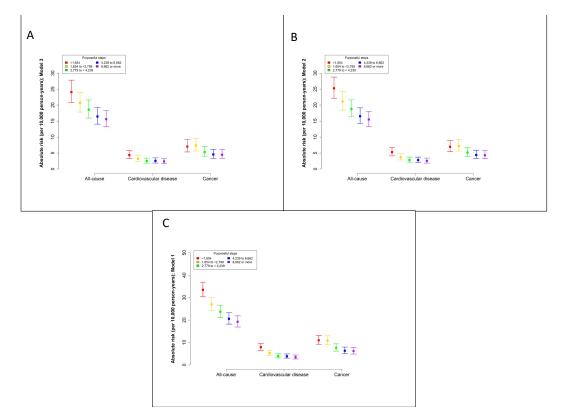
- A. Model 1 adjusted for age and sex
- B. Model 2 adjusted for model 1 + ethnicity, education, ses, smoking, alcohol, fruit and vegetable intake, and sleep.
- C. Model 3 adjusted for model 2 + family history of CVD and cancer and medication use (cholesterol, insulin, and hypertension)

eFigure 8. Sequentially adjusted dose–response quintile-based absolute risk describing the associations between incidental steps and mortality outcomes (events rate per 10000 person-years and associated 95% confidence intervals)



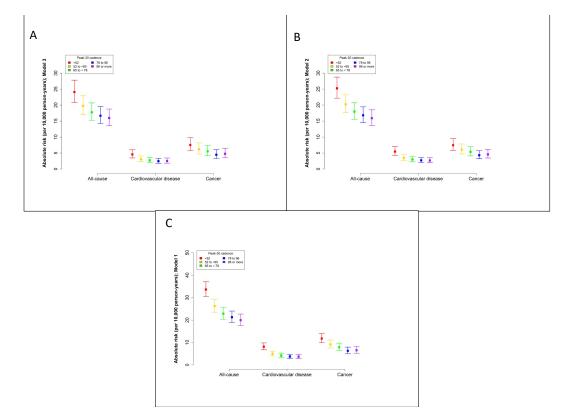
- A. Model 1 adjusted for age and sex
- B. Model 2 adjusted for model 1 + ethnicity, education, ses, smoking, alcohol, fruit and vegetable intake, and sleep.
- C. Model 3 adjusted for model 2 + family history of CVD and cancer and medication use (cholesterol, insulin, and hypertension)

eFigure 9. Sequentially adjusted dose–response quintile-based absolute risk describing the associations between purposeful steps and mortality outcomes (events rate per 10000 person-years and associated 95% confidence intervals).

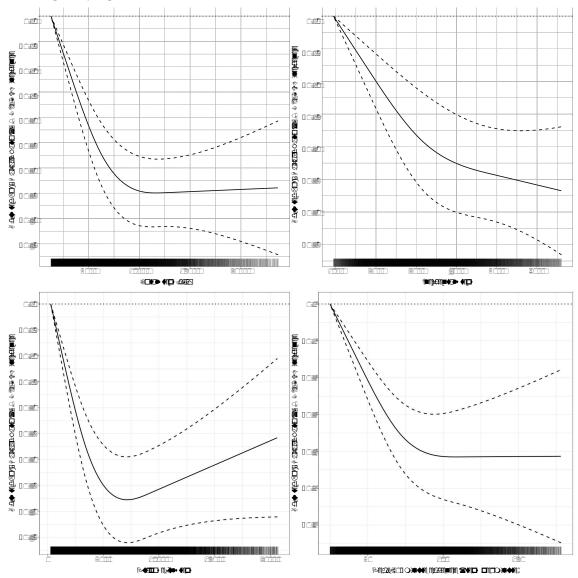


- A. Model 1 adjusted for age and sex
- B. Model 2 adjusted for model 1 + ethnicity, education, ses, smoking, alcohol, fruit and vegetable intake, and sleep.
- C. Model 3 adjusted for model 2 + family history of CVD and cancer and medication use (cholesterol, insulin, and hypertension)

eFigure 10. Sequentially adjusted dose–response quintile-based absolute risk describing the associations between peak-30 cadence and mortality outcomes (events rate per 10000 person-years and associated 95% confidence intervals)

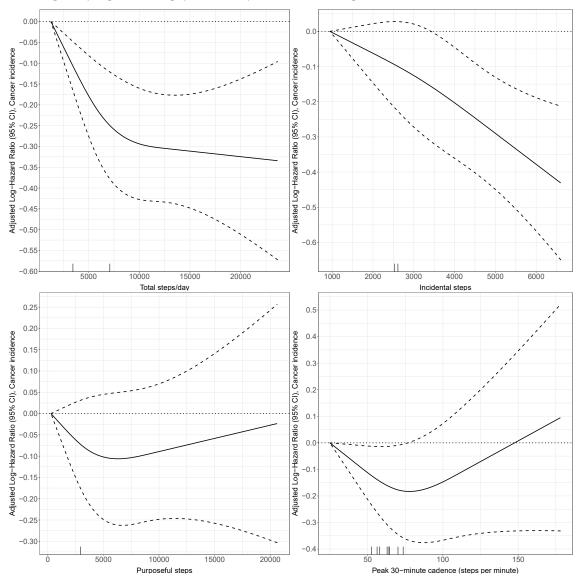


- A. Model 1 adjusted for age and sex
- B. Model 2 adjusted for model 1 + ethnicity, education, ses, smoking, alcohol, fruit and vegetable intake, and sleep.
- C. Model 3 adjusted for model 2 + family history of CVD and cancer and medication use (cholesterol, insulin, and hypertension)



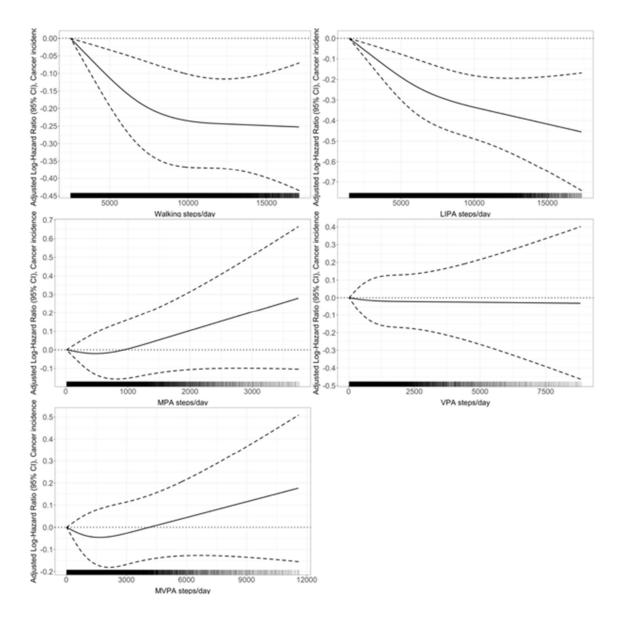
eFigure 11. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD incidence.

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering



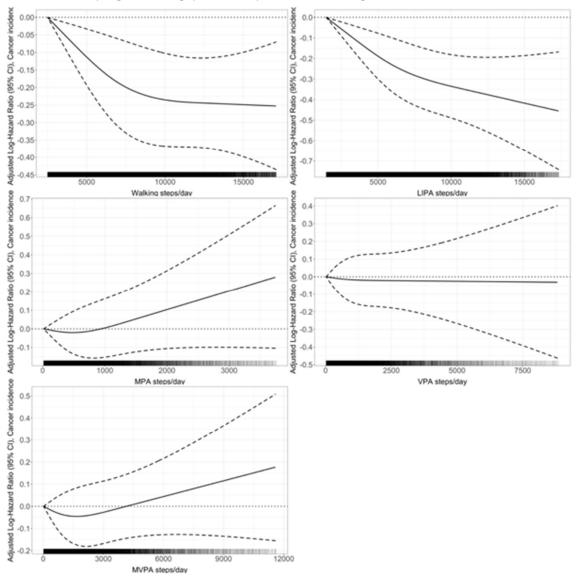
eFigure 12. Dose-response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites incidence outcome.

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering



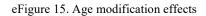
eFigure 13. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between secondary exposures and CVD incidence

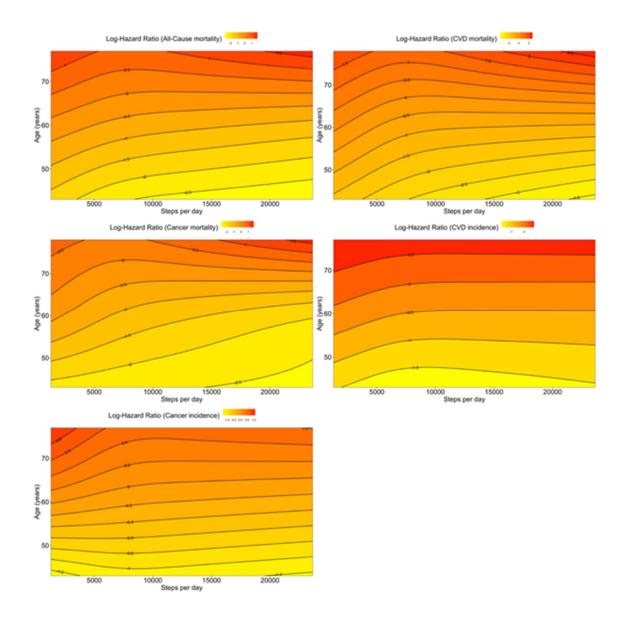
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For LIPA steps, models were further adjusted for MPA and VPA steps (and vice versa). For MVPA steps, models were adjusted for LIPA steps. CVD, Cardiovascular disease; LIPA, Light intensity physical activity; MPA, Moderate intensity physical activity; VPA, Vigorous intensity physical activity; MVPA, Moderate-to-vigorous physical activity; LIPA steps/day, Total daily steps at <100 steps/minute; MPA steps/day, Total daily steps at ≥100 and < 130 steps/minute; MVPA steps/day, Total daily steps at ≥100 steps/minute. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



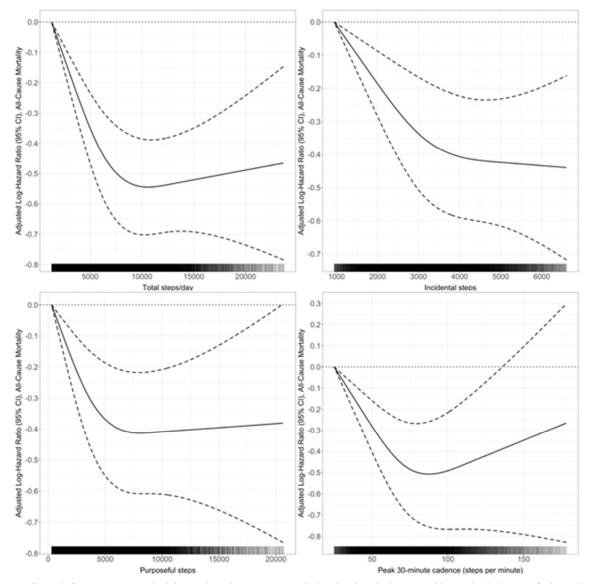
eFigure 14. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between secondary exposures and physical activity-related cancer composite 13 sites incidence outcome.

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For LIPA steps, models were further adjusted for MPA and VPA steps (and vice versa). For MVPA steps, models were adjusted for LIPA steps. CVD, Cardiovascular disease; LIPA, Light intensity physical activity; MPA, Moderate intensity physical activity; VPA, Vigorous intensity physical activity; MVPA, Moderate-to-vigorous physical activity; LIPA steps/day, Total daily steps at <100 steps/minute; MPA steps/day, Total daily steps at \geq 100 and < 130 steps/minute. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



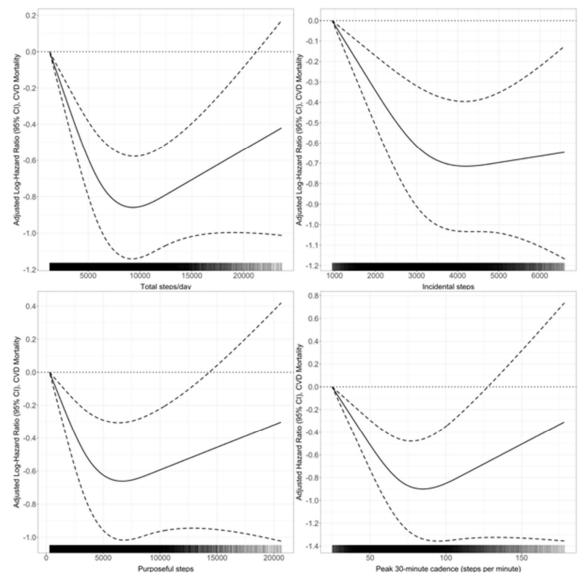


Paralel lines represent lack of interaction between age and steps per day. Darker areas represent higher risks. The lack of interaction between age and daily steps is evident by the somewhat parallel contour lines in all plots.



eFigure 16. Dose–response association (Adjusted^{*a*} hazard ratios and associated 95% confidence interval band) between primary exposures and all-cause mortality (Sensitivity analysis: left-truncation at 2 years follow-up).

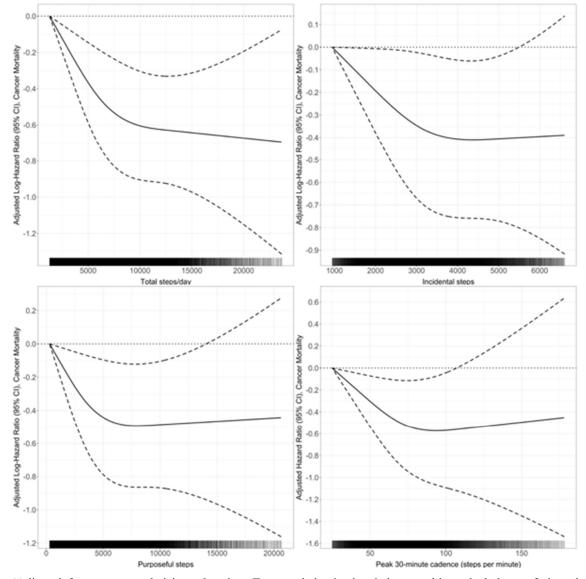
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



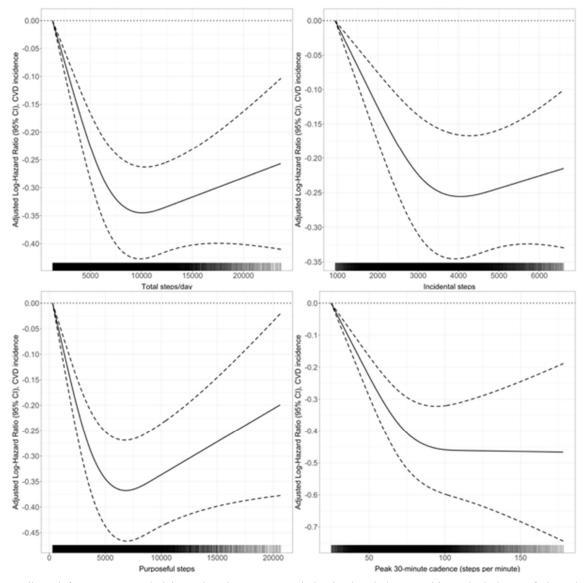
eFigure 17. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD mortality (Sensitivity analysis: left-truncation at 2 years follow-up).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 18. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites mortality outcome (Sensitivity analysis: left-truncation at 2 years follow-up).

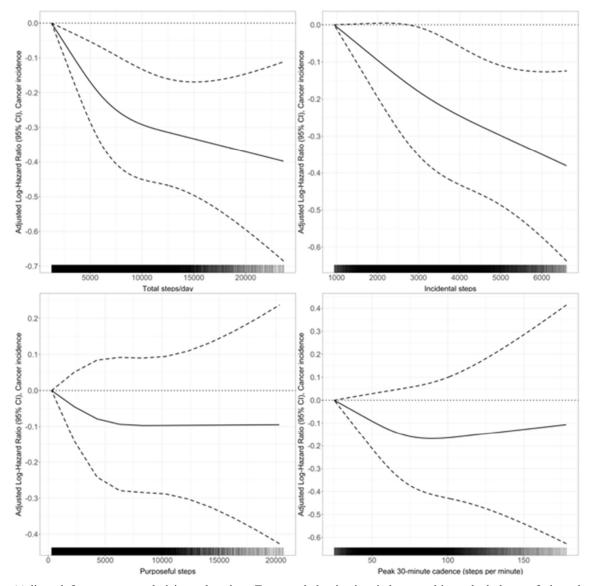


^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



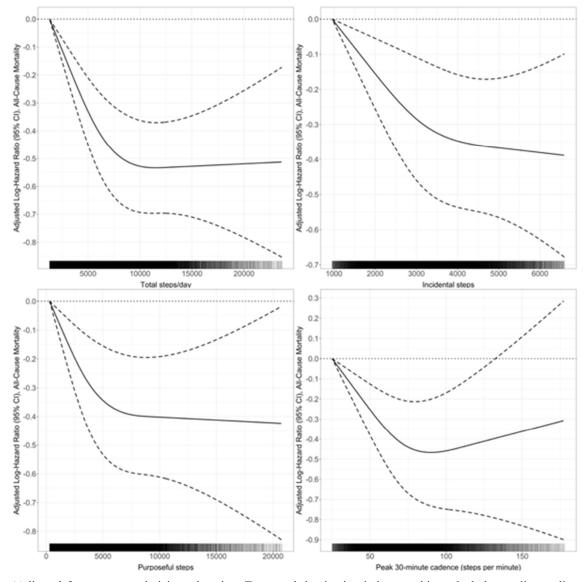
eFigure 19. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD incidence (Sensitivity analysis: left-truncation at 2 years follow-up).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



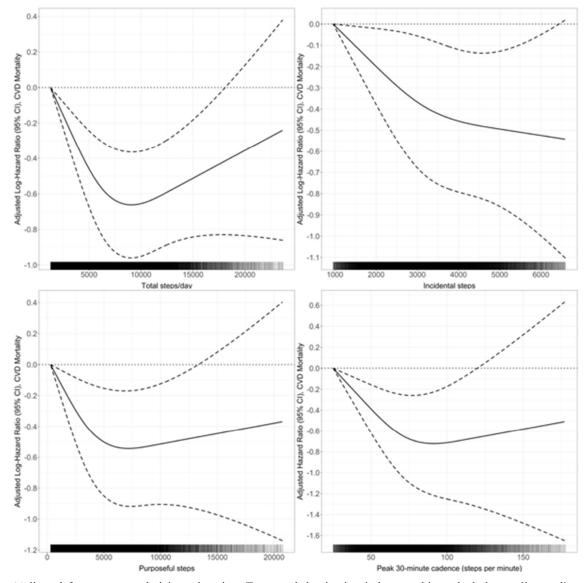
eFigure 20. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites incidence outcome (Sensitivity analysis: left-truncation at 2 years follow-up).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



eFigure 21. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and all-cause mortality (Sensitivity analysis: addition of body mass index, and sleep and dietary score).

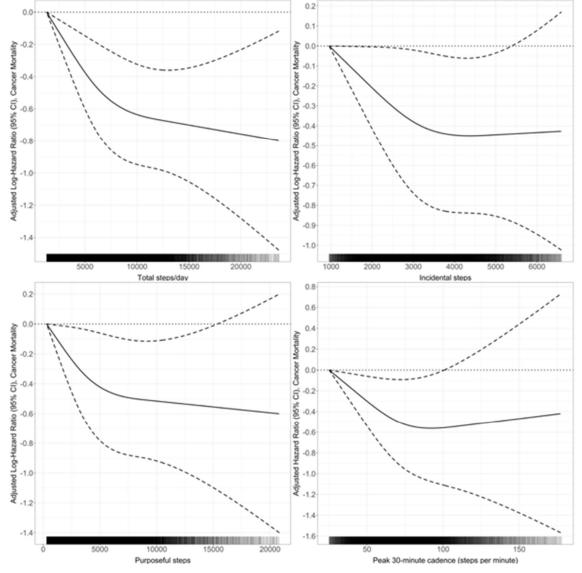
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, diet quality, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), sleep quality, body mass index, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at ≥40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



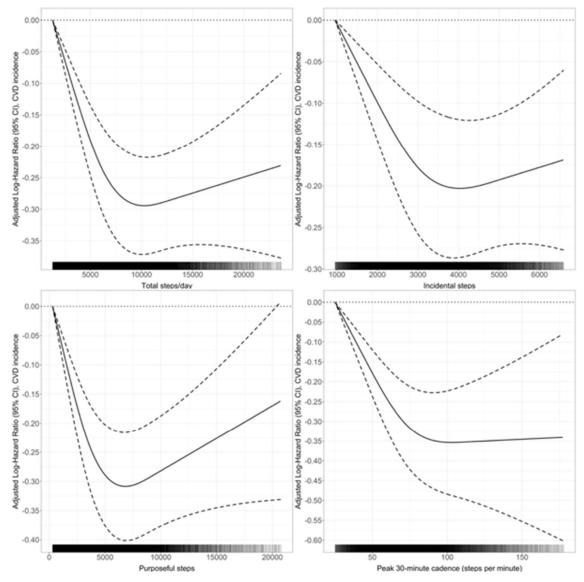
eFigure 22. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD mortality (Sensitivity analysis: addition of body mass index, and sleep and dietary score).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, diet quality, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), sleep quality, body mass index, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at ≥40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 23. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites mortality outcome (Sensitivity analysis: addition of body mass index, and sleep and dietary score).



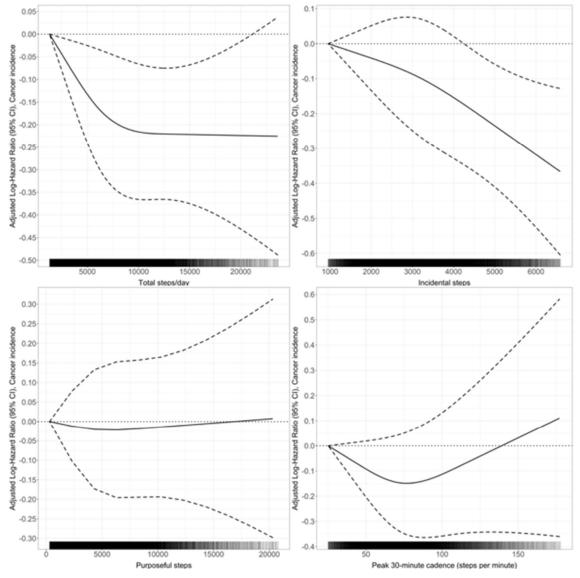
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, diet quality, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), sleep quality, body mass index, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at ≥40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



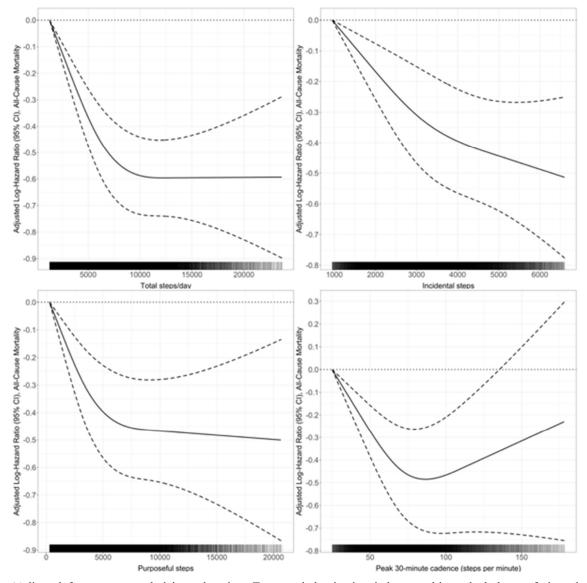
eFigure 24. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD incidence (Sensitivity analysis: addition of body mass index, and sleep and dietary score).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, diet quality, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), sleep quality, body mass index, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at ≥40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 25. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites incidence outcome (Sensitivity analysis: addition of body mass index, and sleep and dietary score).

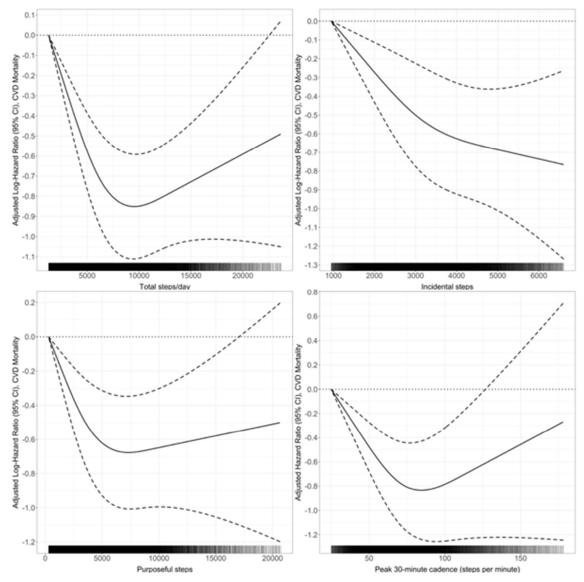


^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, diet quality, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), sleep quality, body mass index, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at ≥40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



eFigure 26. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and all-cause mortality (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).

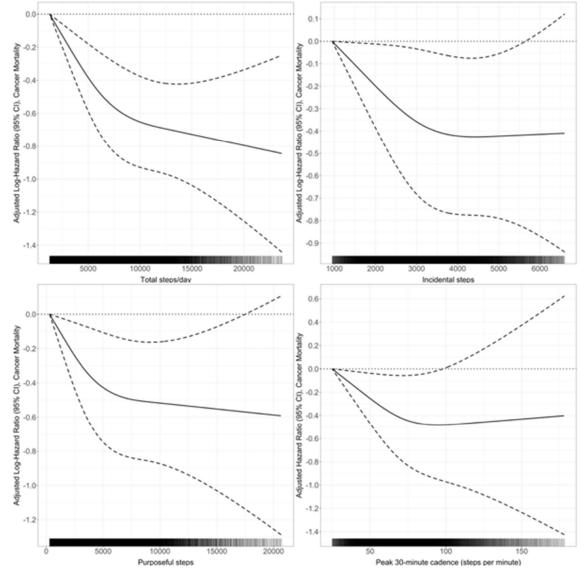
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



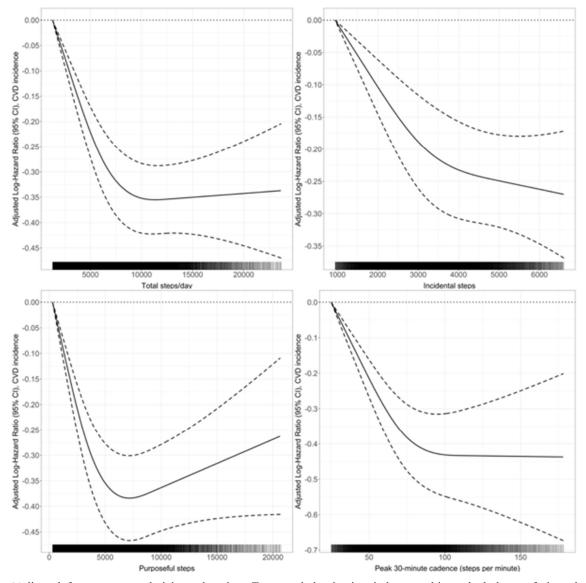
eFigure 27. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD mortality (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at 240 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 28. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites mortality outcome (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).



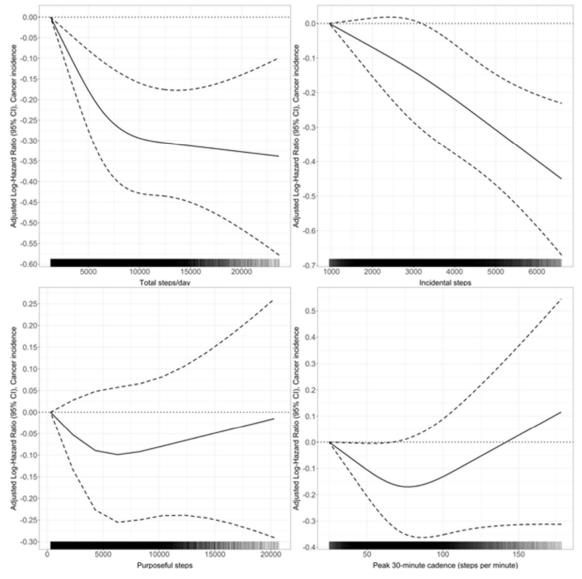
^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.



eFigure 29. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and CVD incidence (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).

^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.

eFigure 30. Dose–response association (Adjusted^a hazard ratios and associated 95% confidence interval band) between primary exposures and physical activity-related cancer composite 13 sites incidence outcome (Sensitivity analysis: restricted to participants with valid wear time days of 20h or more).



^aAdjusted for age, sex, ethnicity, education, Townsend deprivation index, smoking, alcohol use, fruit and vegetable consumption, family history of CVD and Cancer, medication use (cholesterol, insulin, and hypertension), accelerometer-measured sleep, and wear accelerometer days. For incidental steps, models were further adjusted for purposeful steps (and vice versa). CVD, Cardiovascular disease; Total steps/day, Average of steps accumulated in a day; Incidental steps, Total daily steps at 1-39 steps/minute; Purposeful steps, Total daily steps at \geq 40 steps/minute; Peak 30-min cadence, Average steps/minute recorded for the 30 highest, but not necessarily consecutive, minutes in a day. Dose-response associations were assessed with restricted cubic splines with knots at 10th, 50th, and 90th centiles of the distribution of the exposure of interest. Darker colors in the lower bars represent a higher sample clustering.