# **Electronic Supplementary Material**

Physical activity and incident type 2 diabetes mellitus: A systematic review and doseresponse meta-analysis of prospective cohort studies

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**ESM Table 1** Newcastle-Ottawa quality scores of 28 identified prospective cohort studies retrieved on the association of physical activity and type 2 diabetes

		St	tudy Selection			arability of ohorts		Outcome		Total Score
	Representa tiveness of the exposed cohort	Selection of the non- exposed cohort	Ascertainment of exposure	Demonstration that outcome of interest was not present at start of study	ВМІ	PA intensity	Assessment of outcome	Length of follow up	Adequacy of follow up	
Physicians Health Study (1)	С	A*	С	С	A*	В	С	A*	B*	4
Nurses' Health Study (2)	С	A*	С	С	A*	A*	С	A*	B*	5
British Regional Heart Study (3)	В*	A*	С	B*	A*	A*	В*	A*	A*	8
Women's Health Study (4)	С	A*	С	С	A*	A*	С	A*	B*	5
Eastern and South- Western Finnish adults (5)	A*	A*	С	A*	A*	A*	В*	A*	B*	8
Iowa Women's Health Study (6)	B*	A*	С	С	A*	A*	С	A*	B*	6
University of Pennsylvania Alumni (7)	С	A*	С	С	В	A*	С	A*	С	3
North-Eastern Finnish Adult cohort (8)	A*	A*	С	С	В	A*	С	A*	С	4
MONICA/KORA Augsburg Cohort Study (9)	A*	A*	B*	С	A*	В	B*	A*	B*	8

		Si	tudy Selection			arability of ohorts		Outcome		Total Score
	Representa tiveness of the exposed cohort	Selection of the non- exposed cohort	Ascertainment of exposure	Demonstration that outcome of interest was not present at start of study	ВМІ	PA intensity	Assessment of outcome	Length of follow up	Adequacy of follow up	
Health Professional's Follow-up Study (10)	С	A*	С	С	A*	A*	С	A*	B*	5
Osaka Health Survey (11)	С	A*	С	A*	A*	В*	A*	A*	В*	7
Shanghai Women's Health Study (12)	В*	A*	B*	A*	A*	A*	С	A*	В*	8
Kuopio Ischaemic Heart Disease Risk Factor Study(13)	В*	A*	С	A*	A*	A*	A*	A*	В*	8
The Pitt County Study(14)	С	A*	С	A*	A*	A*	A*	A*	С	6
The Strong Heart Study(15)	С	A*	С	A*	A*	A*	A*	A*	B*	7
English Longitudinal Study of Ageing (16)	B*	A*	С	С	A*	A*	С	A*	В*	6
National Health Insurance Corporation Study(17)	B*	A*	С	A*	A*	В	A*	A*	В*	7
The Multiethnic Cohort (18)	A*	A*	С	B*	A*	A*	B*	A*	B*	8
Chin-Shan community cardiovascular cohort study (CCCC) (19)	A*	A*	С	A*	A*	В	A*/B*	A*	B*	7

		St	tudy Selection			arability of ohorts		Outcome		Total Score
	Representa tiveness of the exposed cohort	Selection of the non- exposed cohort	Ascertainment of exposure	Demonstration that outcome of interest was not present at start of study	ВМІ	PA intensity	Assessment of outcome	Length of follow up	Adequacy of follow up	
Black Women's Health Study (20)	С	A*	С	С	В	A*	B*	A*	С	4
The Australian Diabetes, Obesity and Lifestyle Study (21)	A*	A*	B*	A*	A*	A*	A*	A*	В*	9
The Nord-Trøndelag Health Survey (22)	A*	A*	С	С	A*	В	A*	A*	B*	6
<b>45</b> and Up study (23)	A*	A*	С	С	A*	A*	С	A*	С	5
Honolulu Heart Program (24)	С	A*	С	A*	В	A*	С	A*	В*	5
InterAct Consortium(25)	B*	A*	С	A*	A*	A*	A*	A*	B*	8
China MUCA and China Cardiovascular Health Study (26)	A*	A*	С	A*	A*	A*	<b>A</b> *	A*	В*	8
Japanese Office Workers(27)	С	A*	С	A*	A*	A*	A*	A*	B*	7

### **Key to NOS star allocation**

#### Selection

1) Representativeness of the exposed cohort

A\*= truly representative of the general population

B\* = somewhat representative of the general population

C = selected group of users e.g. nurses, volunteers

2) Selection of the non-exposed cohort

 $A^*$  = drawn from the same community as the exposed cohort

B = drawn from a different source

3) Ascertainment of exposure

B\* = Face-to-face structured interview

C = Self-administered questionnaire

4) Demonstration that outcome of interest was not present at start of study

A\* = Clinical assessment

 $B^*$  = Medical records, medication status of the patient

C = Self-report

D = No demonstration

#### Comparability

1) Comparability of cohorts on the basis of the design or analysis

 $A^* = \text{study adjusts for BMI}$ 

B = study doesn't adjust for BMI

2) Comparability of cohorts on the basis of the design or analysis

 $\overline{A}^*$  = study assesses varying levels of physical activity intensity (not only duration)

B = study doesn't asses varying levels of physical activity intensity

#### Outcome

1) Assessment of outcome

A\* = independent blind clinical assessment

B\* = record linkage

C = self-report

2) Was follow-up long enough for outcomes to occur

 $A^* = Yes (> 2 years)$ 

B = No

3) Adequacy of follow up of cohorts

 $A^*$ = complete follow up (>= 99%)

B \* = subjects lost to follow up unlikely to introduce bias > 80% subjects followed up or description of those lost suggests unlikely to introduce bias

C = follow up rate < 80% and no description of those lost

D = no statement on follow up

**ESM Table 2** Summary of MET h/week and MMET h/week dose assignment calculations for the prospective cohort studies included in the dose-response meta-analysis

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
Physicians Health Study (28)	•0 •1 •2-4 (mid-point: 3) •>5 (mid-point: 6)	45 min/session	MVPA = 4.5 MET	0 3.375 10.125 20.25	0 2.625 7.875 15.75	
Nurses' Health Study (2)		Average minutes/week  None 1–29 (mid-point: 15) 30–59 (mid-point: 45) 60–150 (mid-point: 105) >150 (mid-point: 195)	MVPA = 4.5 MET	0 1.125 3.375 7.875 14.625	0 0.875 2.625 6.125 11.375	
British Regional Heart Study (3)	*"Inactive/occasional" (irregular PA)  *"Light" (more frequent recreational activities or PA <1/week, or regular walking + recreational activity)  *"Moderate" (frequent weekend recreational activities + regular walking, or PA 1x/week)  *"Moderately vigorous" (exercise 1/week or frequent cycling + recreational activities or walking or frequent sporting activities)  *"Vigorous" (very frequent exercise, or frequent exercise + recreational activities)	45 min/session  • "Inactive": 30 min/week • "Light": At least 1/week = 1x/week • "Moderate": frequent recreational weekend activities and sporting activity 1/week = MVPA 2x/week • "Moderately vigorous" = 150 min/week • "Vigorous" = MVPA 5x/week	MVPA = 4.5 MET	2.5 3.375 6.75 11.25 16.875	1.75 2.625 5.25 8.75 13.125	

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
Nurses' Health Study (1980 - 2010) (29)		Average weekly hours of MVPA: • <0.5 h (15 min) •0.5–1.9 h (75 min) •2–3.9 h (180 min) •4–6.9 h (330 min) •>7 h (510 min)	MVPA = 4.5 MET	7.8 13.1 22.4 34.3 68	6.1 10.2 17.4 26.3 53	Updated follow up data and mean MET h durations per exposure category were obtained from the authors to estimate doses for this cohort
Women's Health Study (4)			Kcal EE/week categories for LTPA: • 0–199 (mid-point: 99.5) • 200–599 (mid-point: 299.5) • 600–1499 (mid-point: 749.5) • >1500 (mid-point: 1645.5)	1.37 4.1 10.34 22.69	1.04 3.2 8.04 17.65	Conversion from EE/ week to MET h/ week according to: LTPA EE/week/72.5=LTPA MET h/week 72.5 kg as reported on http://whs.bwh.harvard.edu/participants.ht ml
Eastern and South- Western Finnish adults (5)	<ul> <li>"Low" = light levels of OPA &amp; CPA (&lt;30 minutes) &amp; inactive LTPA</li> <li>"Moderate" = 1 type of LTPA activities/week (= 2 sessions</li> <li>"High" = 2 or 3 types of LTPA/week (Each 2 x per week) = 4 sessions</li> </ul>	45 min/session	MVPA = 4.5 MET	0 6.75 13.5	0 5.25 10.5	
lowa Women's Health Study (6)	<ul> <li>Rare or never</li> <li>1/ week - few times/month (low)</li> <li>2–4/ week (medium)</li> <li>&gt;4 per week (high)</li> </ul>	<ul> <li>45 min/session</li> <li>Rare/ never (= 0 min)</li> <li>Low (= 45 min/week)</li> <li>Medium (= 135 min/week)</li> <li>High (= 225 min/week)</li> </ul>	MVPA = 4.5 MET	0 3.375 10.125 16.9	0 2.625 7.875 13.15	

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
University of Pennsylvania Alumni (7)			Weekly total EE for LTPA in 500 kcal increments:  •<500 (mid-point: 250)  •500 – 999 (mid-point: 750)  •1000-1499 (mid-point: 1250)  •1500 -1999 (mid-point: 1750)  •2000 -2499 (mid-point: 2250)  •2500 -2999 (mid-point: 2750)  •3000 – 3499 (mid-point: 3250)  •≥ 3500 (mid-point: 3750)	3.28 9.84 16.4 22.97 29.53 36.09 42.65 49.21	2.55 7.65 12.76 17.87 22.97 28.07 33.17 38.27	Conversion from EE/ week to MET h/ week according to the following formula: LTPA EE/week/76.2=LTPA MET h/week  Average weight of males in the general population: 168 lbs (= 76.2 kg) ((30))
North-Eastern Finnish Adult cohort (8) (I)			Weekly LTPA EE (kcal) categories for men defined as: •Low: 0–1100 (mid-point: 550) •Moderate: 1101–1900 (mid-point: 1500.5) •High: >1900 (mid-point: 2299)	6.8 18.5 28.3	5.3 14.4 22	LTPA EE/week/81.3=LTPA MET h/week  BMI: 25.1 Height: 180 cm Mean weight: 81.3 kg  Height info: http://www.pisa2006.helsinki.fi/finland/starstics/statistics.htm
North-Eastern Finnish Adult cohort (8) (II)			Weekly LTPA EE (kcal) categories for women:  • Low: 0–900 (mid-point: 450)  • Moderate: 901–1500 (mid-point: 1200.5)  • High >1500 (mid-point: 1799.5)	6.9 18.3 25.7	5.4 14.2 20	Weight information source: ((31)) LTPA EE/week/65.6=LTPA MET h/week  BMI: 23.8 Height: 166 cm Mean weight: 65.6 kg  Height info: http://www.pisa2006.helsinki.fi/finland/statstics/statistics.htm  Weight information source ((31))

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
MONICA/KORA Augsburg Cohort Study (9)		<1 h/week = Mean of 30 min/week  4 LTPA categories:  • "No activity" = No sports in summer or winter  • "Low activity" = Irregular, <1 h/week in <1 season  • "Moderate" = Regular 1 h/week in <1 season  • "High" = regular >2 h/week in all seasons	MVPA = 4.5 ME I	0 2.25 4.5 11.25	0 1.75 3.5 8.75	
Health Professional's Follow-up Study (10)		seasons		MET h/week LTPA : • 0–5.9 (median=2.7) • 6 –13.7 (median=9.6) • 13.8–24.2 (median=18.6) • 24.3–40.8 (median = 31.6) • ≥40.9 (median=48.2)	2.1 7.47 14.47 24.56 37.45	
The Pitt County Study (14)	4 categories of LTPA  "Inactive" = No strenuous PA, walking, home maintenance or gardening in a week.  "Low" = > 15 min of home maintenance or gardening  "Moderate" = Strenuous PA but not >3 times/week at 20 min/session  "Strenuous" = Strenuous exercise/work >3/week and >20 min at a time	0 min 15 min of MVPA 60 min of VPA 105 min VPA	MVPA = 4.5 MET  VPA = 8 MET	0 1.125 8 14	0 0.875 7 12.25	

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
The Strong Heart Study (15)			LIPA MET h/week:  No activity < 8 MET h 8 – 24 MET h > 24 MET h	0 3.47 15.3 64.2	0 2.625 11.9 49.93	Mean ME I h/ exposure category values were obtained upon correspondence with the authors
Shanghai Women's Health Study (12)		0 48 min 150 min 300 min	MET h/day/year	0 3.6 11.25 27	0 2.8 8.75 17.5	Additional data received from lead author
English Longitudinal Study of Ageing (16)	3 category index:	45 min/session	Light intensity PA = 3 MET	0 4.5	0 2.5	
	<ul> <li>Physical inactivity</li> <li>Low-intensity but not MVPA at least 1/week (at least 1/week = 2/week)</li> <li>MVPA at least 1/week (at least 1/week (at least 1/week = 2/week)</li> </ul>		MVPA = 4.5 MET	6.75	4.75	
National Health Insurance Corporation Study (17)		• "Inactive" (0 minutes/week) • "Low" (1–149 min/week = 75 min) • "Medium" (150–299 min/week) = 225 min) • "High" (≥300 min/week) = 375 min	MVPA = 4.5 MET	0 5.625 16.875 28.125	0 4.375 13.125 21.875	

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
China Multicenter Collaborative Study of Cardiovascular Epidemiology (ChinaMUCA) and China CardiovascularHealth Study (26)			• Sedentary (PAL 1.00-1.39) =1.2 • Low active (PAL 1.40-1.59) = 1.5 • Active (PAL 1.60-1.89) = 1.75 • Very active (PAL >1.89) = 2.04	136.22 173.04 202.86 238.56	24.22 61.04 90.86 126.56	Males:
Chin-Shan community cardiovascularcohort study (CCCC)] (19)	Frequency of sports exercise:  • Never • Rarely (30 min/week) • Sometimes (2/week) • Often (5/week)	• Never • Rarely = (30 min/week) • Sometimes = (2/week) • Often = (5/week)	MVPA = 4.5 MET	0 2.25 6.75 16.875	0 1.75 4.75 13.125	
Black Women's Health Study (20)		• 0 h/week • <1 h/week = 0.5 h • 1–2 h/week = 1.5 h • 3–4 h/week = 3.5 h • 5–6 h/week = 5.5 h • ≥7 h/week = 7.5 h	MVPA = 4.5 MET	0 2.25 6.75 15.75 20.25 33.75	0 1.75 5.25 12.25 15.75 26.25	

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
Kuopio Ischaemic Heart Disease Risk Factor Study (13)	Frequency/month	1 h 8.8 h = 528 min 7.3 h = 438 min	4.1 ME I/h 4.1 MET/h 6.4 MET/h	4.1 36.08 46.72	3.1 27.28 39.42	
EPIC-InterAct (25)		4 category index which incorporates OPA and LTPA:  • Inactive" = Sedentary job and no LTPA • "Moderately inactive" = Sedentary job with 0.5 h LTPA/day or standing job with no LTPA • "Moderately active" = Sedentary job with 0.5 -1 h LTPA/day or standing job with 0.5 h LTPA/day or physical job with no LTPA • "Active" = Sedentary job with >1 h LTPA or standing job with .0.5 h LTPA or physical job with .0.5 h LTPA or physical job with some LTPA or heavy manual job		0 9.966 20.04 33.4	0 8.216 14.79 24.65	Doses were assigned with the aid of findings from a validation study (32)

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
Honolulu Heart Program(24)	Hours/week in each of 5 activity levels with 8 h of sleep per day excluded from the daily total	24 h PA index with 18 h of wake time	MVPA = 4.5 MET Activity levels were multiplied by a weight based on mean oxygen consumption required to perform the activities of the category	129.85 153.3 168 187.25 300.3	17.85 41.3 56 75.25 188.2	Excl. 8 h per day as sleeping time  For MMET = Subtract 7x16 to account for 1 MET/h less per wake time
Japanese male office worker cohort (27)			Daily EE/kg for total PA:  •<33.1 (mid-point: 16.55)  •33.1–36.7 (mid-point: 34.9)  •36.8–40.3 (mid-point: 38.55)  •≥40.4 (mid-point: 42.15)	119.13 244.3 269.85 295.05	N/A	Mean Japanese weight:59.017 kg  Adjustment for male only cohort ~ +5 kg to the mean adult japanese weight. = Mean of 64 kg assumed for a Japanese office worker (33)  Level 1: 16.55*64=(1059.2 total PA EE/person/day)*7=7624.4 total PA EE/week  Level 2: 34.9*64=(2233.6 total PA EE/person/day)*7=15635.2 total PA EE/week  Level 3: 38.55*64=(2467.2 total PA EE/person/day)*7=17270.4 total PA EE/person/day)*7=17270.4 total PA EE/week  Level 4: 42.15*64=(2697.6 total PA EE/person/day)*7=18883.2 total PA EE/week  Conversion from EE/ week to MET h/ week:
The Multiethnic Cohort (I) (18)	PA frequency for strenuous sport was collapsed into 4 categories: •Never •0.5 – 1 h/week •2 – 3 h/week •>4h/week	•Never (= 0 min) •0.5 – 1 h/week (45 min/week) •2 – 3 h/week (150 min/week) •>4h/week (=270 min/week)	MVPA = 4.5 MET	0 3.375 11.25 20.25	0 2.625 8.75 15.75	PA EE/week/64=total PA MET h/week

Study	Frequency	Duration	Intensity	Assigned MET h/week	MMET h/week	Additional information
The Multiethnic Cohort	PA trequency for	•Never (= 0 min)	MVPA = 4.5 ME I	0	0	
<b>(II)</b> (18)	strenuous sport was	•0.5 – 1 h/week (45		3.375	2.625	
	collapsed into 4	min/week)		11.25	8.75	
	categories: •Never •0.5 – 1 h/week •2 – 3 h/week	•2 – 3 h/week (150 min/week) •>4h/week (=270 min/week)		20.25	15.75	
Osaka Health Survey (11)	<ul><li>&gt;4h/week</li><li>3 categories of LTPA</li></ul>	45 min/session	MVPA = 4.5 MET	0	0	
	frequency on both			5.0625	3.9375	
	weekdays and	0 min		15.1875	11.8125	
	weekends:	67.5 min				
	•0	202.5 min				
	•1-2 (1.5 sessions)					
	• ≥3 (4.5 sessions)					

MET h per week doses were assigned from descriptions identified within the individual studies or from correspondence with study authors.

(I)/(II) indicate sub-cohorts with independently reported risk estimates for diabetes type 2

CPA: Commuting physical activity; EE: Energy expenditure (in kcal); LTPA: Leisure time physical activity; MET: Metabolic Equivalent of Task; MMET: Marginal Metabolic Equivalent of Task, MET unit accounting for the individual basal metabolic rate; MVPA: Moderate to vigorous physical activity; OPA:

Occupational physical activity; PA: Physical activity

Black font: Reported PA duration, frequency or intensity

Red: Assumed Green: Calculated

Blue: Additional PA duration, frequency or intensity data received upon correspondence with the authors

**ESM Table 3** Summary of MET h/week and MMET h/week sensitivity analysis dose assignment calculations for the prospective cohort studies included in the dose-response meta-analysis

#### Scenario A:

i. **Session duration:** 45 min

ii. Intensity: MVPA = 4.5 MET/h; LPA= 3 MET/h; VPA= 8 MET/h

## Scenario B:

i. **Session duration:** 30 min

ii. Intensity: MVPA = 4.5 MET/h; LPA= 3 MET/h; VPA= 8 MET/h

#### Scenario C:

i. **Session duration**: 45 min

ii. Intensity: MVPA = 3.5 MET/h; LPA= 2 MET/h; VPA= 7 MET/h

#### Scenario D:

i. **Session duration**: 30 min

ii. Intensity: MVPA = 3.5 MET/h; LPA= 2 MET/h; VPA= 7 MET/h

Study	Scenar Assig MET h/	ned	Scenario A: MMET h/week	Scenari Assigi MET h/v	ned	Scenario B: MMET h/week	Scenar Assigi MET h/v	ned	Scenario C: MMET h/week	Scenar Assigned h/we	d MET	Scenario D: MMET h/week
Physicians Health	0	ID	0	0	D	0	0		0	0	ID	0
Study (1)	1.6875		1.3125	1.125		0.875	1.3125		0.9375	0.875		0.625
,	3.375		2.625	2.25		1.75	2.625		1.875	1.75		1.25
	10.125		7.875	6.75		5.25	7.875		5.625	3.5		3.75
	20.25		15.75	13.5		10.5	15.75		11.25	7		7.5
Nurses' Health	0	ı	0	0		0	0	1	0	0	ı	0
Study (2) (I)/(II)	1.125		0.875	1.125		0.875	0.875		0.625	0.875		0.625
	3.375		2.625	3.375		2.625	2.625		1.875	2.625		1.875
	7.875		6.125	7.875		6.125	6.125		4.375	6.125		4.375
	14.625		11.375	14.625		11.375	11.375		8.125	11.375		8.125

Study	Scenari Assign MET h/w	ned	Scenario A: MMET h/week	Scenari Assigr MET h/v	ned	Scenario B: MMET h/week	Scenari Assigr MET h/v	ned	Scenario C: MMET h/week	Scenar Assigne h/we	d MET	Scenario D: MMET h/week
British Regional Heart Study (3)	2.5 3.375	ID	1.75 2.625	1.125 2.25	D	0.875 1.75	1.75 2.625		1.25 1.875	0.875 1.75	ID	0.625 1.25
neart Study (3)	6.75		5.25	2.23 5		3.5	5.25		3.75	3.5		2.5
	6.75 11.25											
			8.75	7.25		5.25	8.75		6.25	5.25		3.75
	16.875		13.125	11.25		8.75	13.125		9.375	8.75		6.25
Women's Health	1.37		1.04	1.37		1.04	1.37		1.04	1.37		1.04
Study (4)	4.1		3.2	4.1		3.2	4.1		3.2	4.1		3.2
	10.34		8.04	10.34		8.04	10.34		8.04	10.34		8.04
	22.69		17.65	22.69		17.65	22.69		17.65	22.69		17.65
Eastern and South-	0		0	0		0	0		0	0		0
Western Finnish	6.75		5.25	4.5		3.5	5.25		3.75	3.5		2.5
adults (5)	13.5		10.5	9		7	10.5		7.5	7		5
lowa Women's	0	ID	0	0	D	0	0	I	0	0	ID	0
Health Study (6)	3.375		2.625	2.25		1.75	2.625		1.875	1.75		1.25
	10.125		7.875	6.75		5.25	7.875		5.625	5.25		3.75
	16.9		13.15	11.25		8.75	13.15		9.375	8.75		6.25
University of	3.28		2.55	3.28		2.55	3.28		2.55	3.28		2.55
Pennsylvania	9.84		7.65	9.84		7.65	9.84		7.65	9.84		7.65
Alumni (7)	16.4		12.76	16.4		12.76	16.4		12.76	16.4		12.76
( )	22.97		17.87	22.97		17.87	22.97		17.87	22.97		17.87
	29.53		22.97	29.53		22.97	29.53		22.97	29.53		22.97
	36.09		28.07	36.09		28.07	36.09		28.07	36.09		28.07
	42.65		33.17	42.65		33.17	42.65		33.17	42.65		33.17
	49.21		38.27	49.21		38.27	49.21		38.27	49.21		38.27
North-Eastern	6.8		5.3	6.8		5.3	6.8		5.3	6.8		5.3
Finnish Adult	18.5		14.4	18.5		14.4	18.5		14.4	18.5		14.4
cohort (8) (I)	28.3		22	28.3		22	28.3		22	28.3		22

Study	Scenar Assig MET h/	ned	Scenario A: MMET h/week	Scenario Assign MET h/w	ed	Scenario B: MMET h/week	Scenar Assig MET h/	ned	Scenario C: MMET h/week	Scenario D: Assigned MET h/week	Scenario D: MMET h/week
North-Eastern Finnish Adult cohort (8) (II)	6.9 18.3 25.7		5.4 14.2 20	6.9 18.3 25.7		5.4 14.2 20	6.9 18.3 25.7		5.4 14.2 20	6.9 18.3 25.7	5.4 14.2 20
MONICA/KORA Augsburg Cohort Study (9) (I)	0 2.25 4.5 11.25	ID	0 1.75 3.5 8.75	0 2.25 4.5 11.25	D	0 1.75 3.5 8.75	0 1.75 3.5 8.75	I	0 1.25 2.5 6.25	0 ID 1.75 3.5 8.75	0 1.25 2.5 6.25
MONICA/KORA Augsburg Cohort Study (9) (II)	0 2.25 4.5 11.25	ID	0 1.75 3.5 8.75	0 2.25 4.5 11.25	D	0 1.75 3.5 8.75	0 1.75 3.5 8.75	I	0 1.25 2.5 6.25	0 ID 1.75 3.5 8.75	0 1.25 2.5 6.25
Health Professional's Follow-up Study (34)	0 2.025 7.275 27		0 1.575 5.658 21	0 2.025 7.275 27		0 1.575 5.658 21	0 1.575 5.658 21		0 1.125 4.04 15	0 1.575 5.658 21	0 1.125 4.04 15
The Pitt County Study (14)	0 1.125 8 14	ID	0 0.875 7 12.25	0 1.125 8 14	D	0 0.875 7 12.25	0 0.875 7 12.25	I	0 0.625 6 10.5	0 ID 0.875 7 12.25	0 0.625 6 10.5
The Strong Heart Study (15)	0 3.47 15.3 64.2		0 2.625 11.9 49.93	0 3.47 15.3 64.2		0 2.625 11.9 49.93	0 3.47 15.3 64.2		0 2.625 11.9 49.93	0 3.47 15.3 64.2	0 2.625 11.9 49.93
Shanghai Women's Health Study (12)	0 3.6 11.25 27		0 2.8 8.75 17.5	0 3.6 11.25 27		0 2.8 8.75 17.5	0 3.6 11.25 27		0 2.8 8.75 17.5	0 3.6 11.25 27	0 2.8 8.75 17.5

Study	Scenari Assigr MET h/v	ned	Scenario A: MMET h/week	Scenario Ba Assigned MET h/weel		Scenario B: MMET h/week	Scenario Assign MET h/w	ed	Scenario C: MMET h/week	Scenari Assigned h/wee	IMET	Scenario D: MMET h/week
English Longitudinal Study	0 3.375	ID	0 2.25	0 [ 3.375	)	0 2.625	0 2.25	ı	0	0 1.5	ID	0
of Ageing (16)	7.03125		5.90625	4.6875		3.9375	5.90625		4.71	3.9375		3.1875
National Health	0	I	0	0		0	0	I	0	0	I	0
nsurance	5.625		4.375	5.625		4.375	4.375		3.125	4.375		3.125
Corporation Study	16.875		13.125	16.875		13.125	13.125		9.375	13.125		9.375
(17)	28.125		21.875	28.125		21.875	21.875		15.625	21.875		15.625
Chin-Shan	0	ID	0	0 [	)	0	0	I	0	0	I	0
community	2.25		1.75	2.25		1.75	1.75		1.25	1.75	D	1.25
cardiovascular	6.75		4.75	4.5		3.5	4.75		3.75	3.5		2.5
cohort study (CCCC)(19)	16.875		13.125	11.25		8.75	13.125		9.375	8.75		6.25
Black Women's	0	ı	0	0		0	0		0	0		0
Health Study (20)	2.25		1.75	2.25		1.75	1.75		1.25	1.75		1.25
- , ,	6.75		5.25	6.75		5.25	5.25		3.75	5.25		3.75
	15.75		12.25	15.75		12.25	12.25		8.75	12.25		8.75
	20.25		15.75	20.25		15.75	15.75		13.75	15.75		13.75
	33.75		26.25	33.75		26.25	26.25		18.75	26.25		18.75
Kuopio Ischaemic	4.1		3.1	4.1		3.1	4.1		3.1	4.1		3.1
Heart Disease Risk	36.08		27.28	36.08		27.28	36.08		27.28	36.08		27.28
Factor Study (13)	46.72		39.42	46.72		39.42	46.72		39.42	46.72		39.42
The Multiethnic	0	ı	0	0		0	0	I	0	0	I	0
Cohort (I) (18)	3.375		2.625	3.375		2.625	2.625		1.875	2.625		1.875
	11.25		8.75	11.25		8.75	8.75		6.25	8.75		6.25
	20.25		15.75	20.25		15.75	15.75		11.25	15.75		11.25
The Multiethnic	0	ı	0	0		0	0	ı	0	0	ı	0
Cohort (II) (18)	3.375		2.625	3.375		2.625	2.625		1.875	2.625		1.875
• • • • • • • • • • • • • • • • • • • •	11.25		8.75	11.25		8.75	8.75		6.25	8.75		6.25
	20.25		15.75	20.25		15.75	15.75		11.25	15.75		11.25

Study	Scenari Assigr MET h/v	ned	Scenario A: MMET h/week	Scenar Assig MET h/v	ned	Scenario B: MMET h/week	Scenario Assign MET h/w	ed	Scenario C: MMET h/week	Scenari Assigned h/wed	MET	Scenario D: MMET h/week
Osaka Health Survey(11)	0 5.0625 15.1875	ID	0 3.9375 11.8125	0 3.375 10.125	D	0 2.625 7.875	0 3.9375 11.8125	I	0 2.8125 8.4375	0 2.625 7.875	ID	0 1.875 5.625
The Australian Diabetes, Obesity and Lifestyle Study (21)	0 5.625 14.625	I	0 4.375 11.375	0 5.625 14.625		0 4.375 11.375	0 4.375 11.375	I	0 3.125 8.125	0 4.375 11.375	I	0 3.125 8.125
The Nord- Trøndelag Health Survey (22)	0 1.6875 3.375 8.4375 23.625	ID	0 1.3125 2.625 6.5625 18.375	0 1.125 2.25 5.625 15.75	D	0 0.875 1.75 4.375 12.25	0 1.3125 2.625 6.5625 18.375	I	0 0.9375 1.875 4.6875 13.125	0 0.875 1.75 4.375 12.25	ID	0 0.625 1.25 3.125 8.75
<b>45 and Up study</b> (23)	5.625 16.875 28.125	I	4.375 13.125 21.875	5.625 16.875 28.125	I	4.375 13.125 221.875	4.375 13.125 21.875	I	3.125 9.375 15.625	4.375 13.125 21.875	I	3.125 9.375 15.625
Nr of studies to which assumptions had to be applied for:	15/27			9/27ª			15/27²			15/27ª		
Intensity Duration	15/2 9/27			0/2 9/2			15/27 0/27			15/2 9/27		

<sup>&</sup>lt;sup>a</sup> Number of studies for which assumptions had to be adjusted relative to baseline assumptions made for LTPA dose harmonization in Scenario A MET h/week doses were assigned from descriptions identified within the individual studies or from correspondence with study authors.

(I)/(II) indicate sub-cohorts with independently reported risk estimates for diabetes type 2.

Black font: Reported PA duration, frequency or intensity

Red: Assumed Green: Calculated

Blue: Additional PA duration, frequency or intensity data received upon correspondence with the authors

LPA: Low intensity physical activity; MET: Metabolic Equivalent of Task; MMET: Marginal Metabolic Equivalent of Task MET; PA: Physical activity; MVPA: Moderate to vigorous physical activity; VPA: Vigorous physical activity

**ESM Table 4** Summary of the characteristics of 28 prospective cohort studies that investigate the association between levels of PA and incident type 2 diabetes, identified in the systematic literature search

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week*)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Helmrich et al, 1991 (7)	USA; University of Pennsylvani a Alumni	5990	М	39–68	14	3.4 (202/5990)	Paffenberger Physical Activity Index/College Alumnus Questionnaire (PAI-CAQ) <sup>b</sup>	Weekly EE for LTPA	Weekly total EE for LTPA in 500 kcal (2092 kJ) increments:	RR for T2D relative to most inactive group: 1.00 0.94 (0.9, 0.98) 0.79 0.78 0.68 0.90 0.86 0.52		Age
Burchfiel et al, 1995 (24)	USA; Honolulu Heart Program <sup>c</sup>	6815	M	45–68	6	5.7 (391/6815)	Questionnaire	H/week in each of five activity levels (multiplied by a weight based on mean oxygen consumption required to perform the activities of the category)	Composite score based on 24 h PA dose calculated by summing the hours spent in each activity intensity level and multiplying by a respective weight factor and categorised into quintiles:  • 24.1–29.0 (129.9)  • 29.1–30.7 (153.3)  • 30.8–33.2 (168.0)  • 33.3–36.2 (187.3)  • 36.3–65.5 (300.3)	OR for T2D relative to most inactive group: 1.00 0.86 (0.64, 1.16) 0.81 (0.60, 1.09) 0.72 (0.53, 1.03) 0.47 (0.33, 0.67)	Cohort of middle-aged Japanese-American men	Age
Lynch et al, 1996 (13)	Finland; Kuopio Ischaemic Heart Disease Risk Factor Study <sup>c</sup>	2682	М	42–60	18	23.9 (640/2682)	Minnesota LTPA Questionnaire <sup>b</sup> (modified)	Frequency/mont h; intensity was estimated on a scale of 1 (lowest) –3 (highest)	1 year retrospective leisure time physical activity assessment of 15 common PA types:	PA (>5.5 MET intensity) >40 min/week had an OR of 0.83 (0.63, 1.10) for T2D compared with participants reporting less duration/intensity of PA.  The OR for T2D observed for PA below an intensity of 5.5 MET but >2 h/week was 0.83 (0.66, 1.03)		Age, fasting baseline glucose, serum triacylglycerol, BP, parental history of diabetes, alcohol consumption, BMI
Haapanen et al, 1997 (8)	Finland; North- Eastern Finnish Adult cohort (I)	1340	М	35–63 <sup>d</sup>	10	4.8 (64/1340)	Self- administered questionnaire	LTPA EE/week Frequency of vigorous PA/week	Weekly LTPA EE (kcal) categories for men:  • low: 0–1100 (6.8) • moderate: 1101–1900 (18.5) • high: >1900 (28.3)  Participants were asked to report average frequency of vigorous activity (≥6 MET) as: • ≥1/week • <1/week	RR for T2D relative to most inactive group: 1.00 1.54 (0.83, 2.84) 1.63 (0.92, 2.88)	Risk estimates were reversed to set the lowest level of LTPA as the referent category using the Hamling method (35)	Age

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Haapanen et al, 1997 (8)	Finland; North- Eastern Finnish Adult cohort (II)	1500	F	35–63 <sup>d</sup>	10	3.6 (54/1500)	Self- administered questionnaire	LTPA EE/week Frequency of vigorous PA/week	Weekly LTPA EE (kcal) categories for women:  • low: 0–900 (6.9) • moderate: 901–1500 (18.3) • high: >1500 (25.7)  Participants were asked to report average frequency of vigorous activity (≥6 MET) as: • ≥1/week • <1/week	RR for T2D relative to most inactive group: 1.00 2.64 (1.28, 5.44) 2.23 (0.95, 5.23)	Risk estimates were reversed to set the lowest level of LTPA as the referent category using the Hamling method (35)	Age
James et al, 1998 (14)	USA; Pitt County Study	916	M/F	25–55	5	8.5 (78/916)	Questionnaire	Physical activity index based on frequency of physical activity enough to work up a sweat and result in heavy breathing Frequency of strenuous work/exercise >20 min at a time	Four categories of LTPA level defined as:  • 'inactive' = individuals who did not report any strenuous work/exercise, walking, home maintenance or gardening in a week. (0)  • 'low' = some home maintenance work (>15 min) or gardening during an average week (1.1)  • 'moderate' = some strenuous work/exercise but not >3 times/week at 20 min/session (4.5)  • 'strenuous' = strenuous exercise/work >3/week and >20 min at a time (7.9)	OR for T2D relative to most inactive group: 1.00 0.51 (0.20, 1.28) 0.35 (0.12, 0.98) 0.65 (0.26, 1.63)	Cohort exclusively consisted of individuals of African- American ethnicity	Age, sex, education, WHR, BMI
Folsom et al, 2000 (6)	USA; Iowa Women's Health Study	34,257	F	55-69	12	5.8 (1997/34,257)	Mailed questionnaire Standard questions from the Gallup poll with the addition of questions specific to LTPA	Frequency/week	Initial assessment of any habitual PA (Y/N) Participants reporting regular PA needed to specify frequency of moderate and vigorous PA (>6 MET) which was categorised into quartiles:  • rare or never (0)  • 1/week-few times/month (3.4)  • 2-4/week (10.1)  • >4/week (16.9)	RR for T2D relative to most inactive group: 1.00 0.80 (0.71, 0.90) 0.65 (0.58, 0.74) 0.51 (0.43, 0.59)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.90 (0.79, 1.01) 0.86 (0.76, 0.98) 0.73 (0.62, 0.85)	Cohort consisted exclusively of postmenopausal women	Age, education, smoking, alcohol intake, oestrogen replacement, energy intake, wholegrain intake, dietary score, family history of diabetes (+ BMI and WHR in further adjusted model)

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Okada et al, 2000 (11)	Japan; Osaka Health Survey	6013	М	35–60	10	7.4 (444/6013)	Questionnaire	Min/week during the week or weekend	Three categories of weekly LTPA:  'sedentary': no regular exercise (0)  'moderate': 1 h/week (5.1)  'vigorous': ≥1 h/week exercise 'enough to work up a sweat' (15.2)	RR for T2D relative to most inactive group: 1.00 0.65 (0.45, 0.95) 0.52 (0.35, 0.79)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.80 (0.71, 0.99) 0.55 (0.34, 0.87)		Age, daily alcohol consumption, smoking habits, BP levels, parental history of T2D (+ BMI in the BMI- adjusted model)
Wannamethee et al, 2000 (3)	UK; British Regional Heart Study	7735	M	40–59	16.8	2.5 (196/7735)	Minnesota LTPA Questionnaire in a nurse- administered face to face interview <sup>b</sup>	Weekly frequency of three intensity categories (combined to an overall PA score): (1) regular walking and cycling (2) recreational activity e.g. gardening or pleasure walking (3) sporting activity e.g. running, golf, swimming or tennis	A PA score was calculated depending on dose/type of regular exercise. Scores were categorised into five groups:  • 'inactive/occasional' (irregular walking or recreational activity) (2.5)  • 'light' (more frequent recreational activities or exercise <1/week, or regular walking + recreational activity) (3.4)  • 'moderate' (frequent weekend recreational activities + regular walking, or sporting activity 1/week) (6.8)  • 'moderately vigorous' (exercise 1/week or frequent cycling + recreational activities or walking or frequent sporting activities) (11.3)  • 'vigorous' (very frequent exercise, or frequent exercise + recreational activities) (16.9)	RR for T2D relative to most inactive group: 1.00 0.65 (0.42, 1.00) 0.60 (0.38, 0.95) 0.42 (0.24, 0.72) 0.36 (0.21, 0.62)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.66 (0.42, 1.02) 0.65 (0.41, 1.03) 0.48 (0.28, 0.83) 0.46 (0.27, 0.79)		Age, smoking, alcohol, social class, pre-existing CHD (+ BMI in the BMI-adjusted model)
Hu et al, 2004 (5)	Finland; Eastern and South- Western Finnish adults	4369	M/F	45–64	9.4	2.8 (120/4369)	Mailed questionnaire	Min/week	A simplified index for LTPA scores was derived and reported in three categories:  • 'low' = light levels of occupational, commuting (<30 min) and inactive LTPA (0)  • 'moderate' = 1 type of LTPA activity/week (3.4)  • 'high' = 2 or 3 types of LTPA/week (8.4)	RR for T2D relative to most inactive group: 1.00 0.71 (0.46, 1.12) 0.32 (0.19, 0.56) BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.85 (0.54, 1.34) 0.43 (0.25, 0.74)		Age, study year, sex, systolic BP, smoking, education (+ BMI in the BMI- adjusted model)

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Nakanishi et al, 2004 (27)	Japan; Japanese male office worker cohort	2924	М	35–59	7	5.8 (168/2924)	Self- administered 1 day activity record	Daily EE for total PA	Quartiles of daily EE/kg for 20 activities:	RR for T2D relative to most inactive group: 1.00 0.65 (0.45, 0.95) 0.52 (0.35, 0.79) 0.27 (0.16, 0.45)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.76 (0.52, 1.11) 0.70 (0.46, 1.06) 0.41 (0.24, 0.70)		Age, family history of diabetes, alcohol consumption, cigarette smoking, weekly EE on PA, systolic BP, HDL-cholesterol and triacylglycerol at baseline (+ BMI in the BMI-adjusted model)
Weinstein et al, 2004 (4)	USA; Women's Health Study	37,878	F	55	6.9	3.6 (1361/37,878)	The NHS II Physical Activity & Inactivity Questionnaire	LTPA EE/week and min/week walking	EE/week (kcal) for LTPA in categories of:  • 0–199 (1.4) • 200–599 (4.1) • 600–1499 (10.3) • >1500 (22.7)	HR for T2D relative to most inactive group: 1.00 0.78 (0.68, 0.90) 0.69 (0.59, 0.80) 0.74 (0.63, 0.88) BMI-adjusted HR for T2D relative to most inactive group: 1.00 0.91 (0.79, 1.06) 0.86 (0.74, 1.01) 0.82 (0.70, 0.97)		Age, family history of diabetes, smoking, alcohol, hormone therapy, hypertension, high cholesterol, dietary factors, randomised treatment group within the Women's Health Study (+ BMI in the BMI-adjusted model)
Hsia et al, 2005 (36)	USA; Women's Health Initiative <sup>e</sup>	87,907	F	63.8	5.1	2.6 (2271/87,907)	Women's Health Initiative Questionnaire	MET h/week	Categories of weekly MET h for total physical activity:	RR for T2D relative to most inactive group: 1.00 0.91 (0.80, 1.03) 0.80 (0.70, 0.91) 0.86 (0.75, 0.99) 0.78 (0.67, 0.91)	Ethnically diverse population of white, African-American, Hispanic, American Indian and Asian postmenopausal women	Age, BMI, alcohol, education, smoking, hypertension, hypercholesterolae mia, dietary fibre intake, percent energy from carbohydrate
Meisinger et al, 2005 (9)	Germany; MONICA/K ORA Augsburg Cohort Study (I)	4069	М	24–75 <sup>d</sup>	7.4	3.6 (145/4069)	Face-to-face interview	H/week Frequency/seas on (summer/winter)	Four categories of LTPA defined as:  • 'no activity' = no sports in summer or winter (0)  • 'low activity' = irregular, <1 h/week in at least one season (2.3)  • 'moderate' = regular 1 h/week in at least one season (4.5)  • 'high' = regular > 2 h/week in both seasons (11.3)	HR for T2D relative to most inactive group: 1.00 0.86 (0.57, 1.29) 0.73 (0.45, 1.20) 0.73 (0.45, 1.20)		Age, survey, actual hypertension, dyslipidaemia, parental history of diabetes, regular smoking, alcohol intake, education, BMI

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Meisinger et al, 2005 (9)	Germany; MONICA/K ORA Augsburg Cohort Study (II)	4034	F	24–75 <sup>d</sup>	7.4	2 (82/4034)	Face-to-face interview	H/week Frequency/seas on (summer/winter)	Four categories of LTPA defined as:  • 'no activity' = no sports in summer or winter (0)  • 'low activity' = irregular, <1 h/week in at least one season (2.3)  • 'moderate' = regular 1 h/week in at least one season (4.5)  • 'high' = regular, >2 h/week in both seasons (11.3)	HR for T2D relative to most inactive group: 1.00 0.85 (0.51, 1.41) 0.59 (0.31, 1.11) 0.21 (0.05, 0.86)		Age, survey, actual hypertension, dyslipidaemia, parental history of diabetes, regular smoking, alcohol intake, education, BMI
Villegas et al, 2006 (12)	China; Shanghai Women's Health Study	70,658	F	40-70	4.6	2.8 (1973/70,658)	Face-to-face interview at baseline and the Shanghai Women's Health Study Physical Activity Questionnaire (SWHS- PAQ) <sup>b</sup>	MET h/day/year	EE for retrospective regular LTPA during previous 5 years in MET h/day/year, DPA (including walking), CPA (bus or vehicle, walking or cycling <30 min/day or 30+ min/day) and EE in OPA (high/medium/low)  LTPA h/day:  0 (0)  <0.8 (3.6)  1.89 (31.3)  >1.99 (27.0)	RR for T2D relative to most inactive group: 1.00 0.89 (0.76, 1.03) 0.99 (0.85, 1.15) 0.83 (0.70, 0.97)		Age, daily calories, education level, income level, occupation, smoking, alcohol, hypertension, chronic diseases
Carlsson et al, 2007 (22)	Sweden; Nord- Trøndelag Health Survey	38,800	M/F	≥20	11	1.9 (738/38,800)	Questionnaire	Exercise frequency ranging from 'never' to 'every day'	Frequency of weekly LTPA:  never (0)  <1x/week (1.7)  1x/week (3.4)  2-3x/week (8.4)  every day (23.6)	RR for T2D relative to most inactive group: 1.00 0.79 (0.64, 0.99) 0.61 (0.48, 0.77) 0.60 (0.48, 0.73) 0.49 (0.37, 0.66)	Risk estimates were reversed to set the lowest level of LTPA as the referent category using the Hamling method (35)	Sex, smoking, BMI
Magliano et al, 2008 (21)	Australia; The Australian Diabetes, Obesity and Lifestyle Study	5842	M/F	50.9	5	3.8 (224/5842)	Interviewer- administered Active Australia questionnaire	Total LTPA time derived from sum of the time spent performing MVPA + double the time spent performing VPA in the previous week	Categories of weekly LTPA min/week:  • inactive (0 min/week)  • insufficient (1–49 min/week) (5.6)  • sufficient (≥150 min/week) (14.6)	OR for T2D relative to most inactive group: 1.00 0.97 (0.58, 1.63) 0.64 (0.46, 0.89)  BMI-adjusted OR for T2D relative to most inactive group: 1.00 0.86 (0.58, 1.27) 0.50 (0.35, 0.72)	Risk estimates were reversed to set the lowest level of LTPA as the referent category using the Hamling method (35)	Age, sex, waist circumference, smoking, education, hypertension, family history of diabetes, log FPG, hypertriacylglycerol aemia, low HDL-cholesterol and cholesterol

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Chien et al, 2009 (19)	Taiwan; Chin-Shan community cardiovascu lar cohort study (CCCC)	1639	M/F	>35	9.02	19 (312/1639)	Baecke Questionnaire of Habitual Physical Activity <sup>b</sup>	Sports, occupational and leisure PA frequency was rated on a 5- point Likert scale	Frequency of sports exercise was reported in quartiles corresponding to:  never (0) rarely (2.3) sometimes (6.8) often (16.9)	RR for T2D relative to most inactive group: 1.00 0.83 (0.62, 1.12) 0.70 (0.52, 0.94) 0.74 (0.54, 1.03)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.82 (0.60, 1.12) 0.65 (0.47, 0.89) 0.68 (0.49, 0.95)		Age, sex, the metabolic syndrome, smoking, current alcohol drinking, marital status, education level, occupation, hypertension status, HDL-cholesterol, triacylglycerols, glucose levels, family history of diabetes, BMI
Fretts et al, 2009 (15)	USA; The Strong Heart Study	1651	M/F	45-74	10	27.5 (454/1651)	Questionnaire	LTPA MET h/week Total PA MET h/week	LTPA MET h/week:  no activity  <8 MET h/week (3.5) 8-24 MET h/week (15.3) >24 MET h/week (64.2)  Total PA MET h/week:  no activity <30 MET h/week 30-106 MET h/week >106 MET h/week	OR for T2D relative to most inactive group: 1.00 1.04 (0.74, 1.47) 0.76 (0.55, 1.07) 0.68 (0.49, 0.95)  BMI-adjusted OR for T2D relative to most inactive group: 1.00 1.09 (0.76, 1.56) 0.80 (0.56, 1.15) 0.75 (0.53, 1.00)	Cohort consisted exclusively of individuals of American Indian ethnicity	Age, study site, sex, education, cigarette smoking, alcohol use, family history of diabetes, systolic BP, diastolic BP, HDL-cholesterol, LDL-cholesterol, plasma fibrinogen, BMI
Krishnan et al, 2009 (20)	USA; Black Women's Health Study	45,668	F	21–69	10	6.4 (2928/45,668)	Questionnaire b	H/week spent on VPA (e.g. running, swimming), walking for exercise and walking to and from work	MVPA was reported in categories of:  • 0 h/week (0)  • <1 h/week (2.3)  • 1–2 h/week (6.8)  • 3–4 h/week (15.8)  • 5–6 h/week (20.3)  • ≥7 h/week (33.8)	RR for T2D relative to most inactive group: 1.00 0.90 (0.82, 0.99) 0.77 (0.69, 0.85) 0.53 (0.45, 0.63) 0.49 (0.38, 0.64) 0.43 (0.31, 0.59)		Age, time period, family history of diabetes, years of education, family income, marital status, cigarette use, alcohol use, energy intake, coffee consumption, television watching, walking

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Siegel et al, 2009 (1)	USA; Physicians Health Study	20,757	M	40–84	23.1	8.8 (1836/20,757)	Paffenberger Physical Activity Index/College Alumnus Questionnaire (PAI-CAQ) <sup>b</sup>	Weekly frequency of vigorous exercise 'enough to work up a sweat'	Weekly vigorous exercise in number of times/week:  • rarely/never (0)  • 1–3/month (1.7)  • once/week (3.4)  • 2–4/week (10.1)  • ≥5 times/week (20.3)	RR for T2D relative to most inactive group: 1.00 0.84 (0.72, 0.98) 0.78 (0.68, 0.91) 0.63 (0.55, 0.73) 0.49 (0.41, 0.59)		Age, alcohol use, smoking, history of high cholesterol, history of hypertension (+ BMI in the BMI-adjusted model)
										BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.84 (0.72, 0.98) 0.81 (0.70, 0.93) 0.69 (0.61, 0.79) 0.58 (0.48, 0.69)		
Demakakos et al, 2010 (16)	UK; English Longitudinal Study of Ageing (ELSA)	7466	M/F	62.9-68.3	3.8	3.5 (258/7466)	Questionnaire	Frequency/week	Frequency of each vigorous, moderate and low intensity PA:	HR for T2D relative to most inactive group: 1.00 0.76 (0.51, 1.13) 0.49 (0.33, 0.71)		Age, age-squared, sex, marital status, educational attainment, total household wealth (+ BMI in the BMI- adjusted model)
									Combined to a derived summary three-category index:  • physical inactivity (0)  • low-intensity but not vigorous/moderate-intensity physical activity at least once a week (3.4)  • MVPA or VPA at least once a week (7.0)	BMI-adjusted HR for T2D relative to most inactive group: 1.00 0.83 (0.56, 1.23) 0.57 (0.39, 0.84)		
Ekelund et al, 2012 (25)	Denmark, France, Germany, Italy, Spain, Sweden, UK and the Netherlands ; EPIC- InterAct (I)	EPIC total cohort 340,234; InterAct subcoho rt 15,934; men 6009	MF; MF; M	51.4–55.4 (mean)	12.3	3.6 (12,403/340,234); 4.9 (778/15,934); 6.5 (391/6009)	EPIC Physical Activity Questionnaire including questions specific to LTPA	Physical activity index (including OPA)	Four category index which incorporates OPA and LTPA:  • 'inactive' = sedentary job and no LTPA (0)  • 'moderately inactive' = sedentary job with 0.5 h LTPA/day or standing job with no LTPA (10.0)  • 'moderately active' = sedentary job with 0.5–1 h LTPA/day or standing job with 0.5 h LTPA/day or standing job with 0.5 h LTPA/day or	HR for T2D relative to most inactive group: 1.00 0.89 (0.78, 1.01) 0.73 (0.64, 0.85) 0.69 (0.60, 0.80)		Education, smoking status, alcohol consumption, energy intake, BMI

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
									physical job with no LTPA (20.0)  • 'active' = sedentary job with >1 h LTPA or standing job with 0.5 h LTPA or physical job with some LTPA or heavy manual job (33.4)			
Ekelund et al, 2012 (25)	Denmark, France, Germany, Italy, Spain, Sweden, UK and the Netherlands ; EPIC- InterAct (II)	EPIC total cohort 340,234; InterAct subcoho rt 15,934; women 9925	M/F; M/F; F	51.4–55.4 (mean)	12.3	3.6 (12,403/340,234); 4.9(778/15,934); 4(397/9925)	EPIC Physical Activity Questionnaire including questions specific to LTPA	Physical activity index (including OPA)	Four category index which incorporates OPA and LTPA:  • 'inactive' = sedentary job and no LTPA (0)  • 'moderately inactive' = sedentary job with 0.5 h LTPA/day or standing job with no LTPA (10.0)  • 'moderately active' = sedentary job with 0.5–1 h LTPA/day or standing job with 0.5 h LTPA/day or physical job with no LTPA (20.0)  • 'active' = sedentary job with >1 h LTPA or standing job with 0.5 h LTPA or physical job with 5.5 h LTPA or physical job with some LTPA or heavy manual job (33.4)	HR for T2D relative to most inactive group: 1.00 0.93 (0.89, 0.98) 0.89 (0.78, 1.01) 0.79 (0.68, 0.91)		Education, smoking status, alcohol consumption, energy intake, BMI
Grøntved et al, 2012 (34)	USA; Health Professional s Follow-up Study	32,002	М	44–79	18	7.1 (2278/32,002)	Health Professionals Follow-Up Study Physical Activity Questionnaire & a biannual follow-up questionnaire <sup>b</sup>	Aerobic exercise min/week	Total time spent on aerobic exercise of at least moderate intensity (≥3 METs); participants grouped into four categories:  • none (0)  • 1–59 min (2.0)  • 60–149 min (7.3)  • ≥150 min (27.0)	RR for T2D relative to most inactive group: 1.00 0.93 (0.81, 1.06) 0.61 (0.60, 0.80) 0.55 (0.42, 0.55)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 1.00 (0.88, 1.15) 0.80 (0.69, 0.92) 0.61 (0.53, 0.70)		Age, smoking, alcohol consumption, coffee intake, race, family history of diabetes, total energy, trans fat, polyunsaturated fat to saturated fat ratio, cereal fibre, wholegrain, and glycaemic load, weight, physical activity of at least moderate intensity, TV viewing (+ BMI in the BMI-adjusted model)

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Lee et al, 2012 (17)	South Korea; National Health Insurance Corporation Study	675,496	М	39.4	7.5	7.9 (52,995/675,496)	Questionnaire	Frequency and duration of LTPA that 'causes sweating'	Physical activity volume was calculated and participants were classified into four categories:  • 'inactive' (0 min/week) (0)  • 'low' (1–149 min/week) (5.6)  • 'medium' (150–299 min/week) (16.9)  • 'high' (≥300 min/week) (28.1)	HR for T2D relative to most inactive group: 1.00 0.98 (0.96, 0.99) 0.94 (0.91, 0.96) 0.94 (0.91, 0.97)  BMI-adjusted HR for T2D relative to most inactive group: 1.00 0.95 (0.93, 0.97) 0.90 (0.87, 0.93) 0.91 (0.89, 0.94)		Age, smoking status, alcohol intake, hypertension, parental diabetes, baseline glucose (+ BMI in the BMI- adjusted model)
Steinbrecher et al, 2012 (18)	USA; The Multiethnic Cohort (I)	35,976 (men)	М	45–75	14	12.6 (4527/35,927)	Questionnaire	H/week of strenuous sport, vigorous work or moderate activity	Physical activity frequency for strenuous sport was collapsed into four categories:  • never (0)  • 0.5–1 h/week (3.4)  • 2–3 h/week (11.3)  • >4 h/week (20.3)	HR for T2D relative to most inactive group: 1.00 0.94 (0.87, 1.02) 0.85 (0.77, 0.94) 0.80 (0.72, 0.88)		Age, ethnicity, education, BMI
Steinbrecher et al, 2012 (18)	USA; The Multiethnic Cohort (II)	38,937 (women)	F	45–75	14	10.4 (4034/38,937)	Questionnaire	H/week of strenuous sport, vigorous work or moderate activity	Physical activity frequency for strenuous sport was collapsed into four categories:  never (0)  0.5–1 h/week (3.4)  2–3 h/week (11.3)  >4 h/week (20.3)	HR for T2D relative to most inactive group: 1.00 1.00 (0.91, 1.09) 0.85 (0.75, 0.96) 0.67 (0.57, 0.79)		Age, ethnicity, education, BMI
Shi et al, 2013 (37)	China; Shanghai Men's Health Study <sup>e</sup>	51,464	М	54.1	5.4	2.5 (1304/51,464)	The Shanghai Men's Health Study Physical Activity Questionnaire	Appraisal of LTPA, DPA and CPA Participants had to indicate whether they had undertaken any LTPA ≥1/week over the preceding 5 years	LTPA volume was reported as four categories of MET h/week/year:  • none (0) • low (<1.2) (4.2) • medium (1.2–3) (14.7) • high (≥3) (27.3)	HR for T2D relative to most inactive group: 1.00 0.79 (0.65, 0.96) 0.87 (0.72, 1.04) 0.87 (0.75, 1.07) Fully-adjusted HR for T2D relative to most inactive group: 1.00 0.80 (0.65, 0.97) 0.89 (0.74, 1.07) 0.91 (0.76, 1.08)		Age, energy intake, smoking, alcohol consumption, education level, occupation, income level, hypertension, family history of diabetes (+ BMI and WHR in fullyadjusted model)

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
Fan et al, 2015 (26)	China; China Multicenter Collaborativ e Study of Cardiovasc ular Epidemiolog y (China MUCA) and China Cardiovasc ular Health Study	6348	M/F	49.2	7.9	7.5 (478/6348)	Questionnaire	Physical activity level (PAL) = method to estimate total daily energy expenditure (38)	Average h/day spent in vigorous activity (e.g. jogging), moderate activity (e.g. yard work), light activity (e.g. office work), sedentary activity (e.g. TV) and periods of reclining during the previous 12 months  Four PAL categories:  • sedentary (PAL 1.00–1.39) (136.2)  • low active (PAL 1.40–1.59) (173.0)  • active (PAL 1.60–1.89) (202.9)  • very active (PAL >1.89) (238.6)	HR for T2D relative to most inactive group: 1.00 0.92 (0.69, 1.22) 0.70 (0.52, 0.93) 0.55 (0.42, 0.73)  BMI-adjusted HR for T2D relative to most inactive group: 1.00 0.82 (0.62, 1.09) 0.63 (0.47, 0.83) 0.47 (0.36, 0.61)	Urban residents only included in the analysis as the PAL questionnaire was not considered valid for rural participants	Age, sex, geographic region, educational level, cigarette smoking, alcohol consumption, family history of diabetes (+ BMI in the BMI adjusted model)
Grøntved et al, 2014 (2)	USA; Nurses' Health Study (2000– 2008) (I)	51,642	F	53-81	8	4.2 (2158/51,642)	The NHS Physical Activity & Inactivity questionnaire with biannual postal questionnaire s <sup>b</sup>	MVPA min/week	MVPA defined as brisk walking, jogging, running, bicycling, tennis, swimming, other aerobic exercise, other vigorous exercise and stair climbing (>3 METs) and categorised into quintiles according to average min/week:  • none (0)  • 1–29 (1.1)  • 30–59 (3.4)  • 60–150 (7.9)  • >150 (14.6)	RR for T2D relative to most inactive group: 1.00 0.84 (0.73, 0.97) 0.76 (0.66, 0.88) 0.68 (0.60, 0.77) 0.48 (0.42, 0.54)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.94 (0.81, 1.09) 0.88 (0.76, 1.02) 0.85 (0.74, 0.96) 0.66 (0.58, 0.75)		Race, alcohol, weight training, coffee intake, smoking, postmenopausal hormone use, oral contraceptive use, menopausal status, family history of diabetes, total calorie intake, saturated to polyunsaturated fat ratio, trans fat, cereal fibre, wholegrains, glycaemic load (+ BMI in the BMI-adjusted model)
Grøntved et al, 2014 (2)	USA; Nurses' Health Study II (2001– 2009) (II)	47,674	F	36-55	8	2.8 (1333/47,674)	The NHS II Physical Activity & Inactivity Questionnaire with biannual postal questionnaire s <sup>b</sup>	MVPA min/week	MVPA defined as brisk walking, jogging, running, bicycling, tennis, swimming, other aerobic exercise, other vigorous exercise, and stair climbing (-3 METs) and categorised into quintiles according to average min/week:  • none (0)  • 1–29 (1.1)  • 30–59 (3.4)  • 60–50 (7.9)  • >150 (14.6)	RR for T2D relative to most inactive group: 1.00 0.80 (0.67, 0.95) 0.68 (0.57, 0.82) 0.63 (0.54, 0.74) 0.42 (0.36, 0.50)  BMI-adjusted RR for T2D relative to most inactive group: 1.00 0.94 (0.79, 1.13) 0.83 (0.69, 1.00) 0.86 (0.73, 1.01)		Race, alcohol, weight training, coffee, smoking, post-menopausal hormone use, oral contraceptive use, menopausal status, family history of diabetes, total calorie intake, saturated to polyunsaturated fat ratio, trans fat, cereal fibre, wholegrains,

Study	Country; study name	Cohort size	Sex	Age at baseline (years)	Follow- up (years)	% Cumulative incidence (cases/cohort)	Method of PA assessment	PA unit	PA assessment (PA dose in MET h/week²)	Reported OR/RR/HR (95% CI)	Comments	Adjustments
										0.70 (0.59, 0.83)		glycaemic load (+ BMI in the BMI- adjusted model)
Ding et al, 2015 (23)	Australia; 45 and Up study	54,997	M/F	≥45	3.4	1.6 (888/54,997)	Active Australia survey	PA calculated as the sum of time spent in walking, MVPA and VPA (weighted by a factor of two) in the previous week	Total min MVPA/week:	OR for T2D relative to most inactive group: 1.00 0.72 (0.56, 0.94) 0.71 (0.85, 0.97)	Risk estimates were reversed to set the lowest level of LTPA as the referent category using the Hamling method (35)	Age, sex, BMI, SES, health status, BP, blood cholesterol, weight, family history of T2D/heart disease, smoking, alcohol, sitting time, sleep, fruit and vegetable intake, psychological distress

<sup>&</sup>lt;sup>a</sup>Doses were assigned from descriptions identified within the individual studies or from correspondence with study authors. Full details of MET h dose assignment are listed in ESM Table 2, together with the MMET h/week calculations (see ESM)

(I)/(II) indicate subcohorts with independently reported risk estimates for T2D within the same publication

CCCC: Chin-Shan community cardiovascular cohort study; China MUCA: China Multicenter Collaborative Study of Cardiovascular Epidemiology; CPA, commuting physical activity; CVD, cardiovascular disease; DPA: daily living physical activity; EE, energy expenditure; ELSA: English longitudinal study of ageing; EPIC-InterAct: European Prospective Investigation into Cancer and Nutrition-InterAct; F, Female; FPG, fasting plasma glucose; M, Male; MEC, Multiethnic cohort; MONICA/KORA: Monitoring Trends and Determinants on Cardiovascular Diseases/Cooperative Research in the Region of Augsburg Cohort Study; NHS, Nurses' Health Study; OPA, occupational physical activity; SES, socioeconomic status

<sup>&</sup>lt;sup>b</sup>Denotes PA assessment questionnaires with published validation and/or reproducibility studies

<sup>°</sup>Studies updated with further follow-up data obtained from the authors

dTotal cohort

eStudies included in the sensitivity analysis using variance-weighted least squares regression analysis

## **ESM Figure 1** Systematic literature review search terms and strategy

**#1** (activity, physical[MeSH Terms]) OR exercise[MeSH Terms]) OR aerobic exercise[MeSH Terms]) OR motor activity[MeSH Terms]) OR walking[MeSH Terms]) OR oR sport[MeSH Terms]) OR physical fitness[MeSH Terms])) OR physical activity[Title/Abstract]) OR exercise[Title/Abstract]

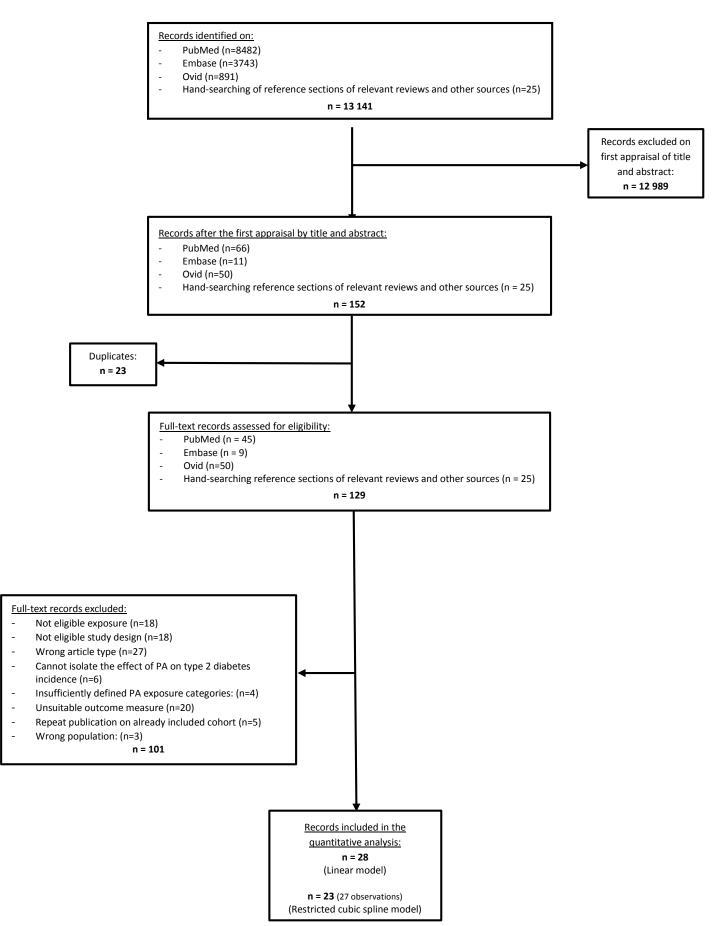
**#2** (Type 2 diabetes mellitus[MeSH Terms]) OR niddm[MeSH Terms]) OR diabetes[MeSH Terms]) OR diabetes mellitus, type 2[MeSH Terms]) OR diabetes mellitus, type ii[MeSH Terms]) OR disorders, glucose metabolism[MeSH Terms]) OR impaired glucose[Title/Abstract]) OR type 2 diabetes[Title/Abstract]) OR insulin resistance[Title/Abstract]) OR glucose tolerance[Title/Abstract]

**#3** (inciden\*[Title/Abstract]) OR cohort[Title/Abstract]) OR prospective[Title/Abstract]) OR longitudinal[Title/Abstract]) OR prospective studies[MeSH Terms]) OR cohort study[MeSH Terms]) OR cohort study[MeSH Terms])

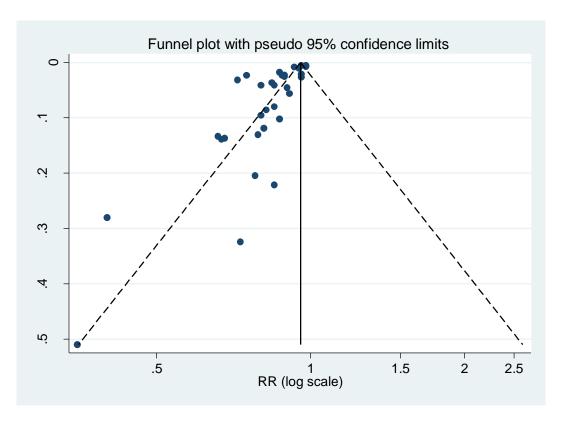
#### #1 and #2 and #3

Search terms for the retrieval of studies in Embase were modified and consisted of combinations of: 'physical activity', 'diabetes mellitus type 2', 'NIDDM', 'noninsulin dependent diabetes', 'cohort study ', 'observational study' or 'longitudinal study'.

## ESM Figure 2 PRISMA flow chart

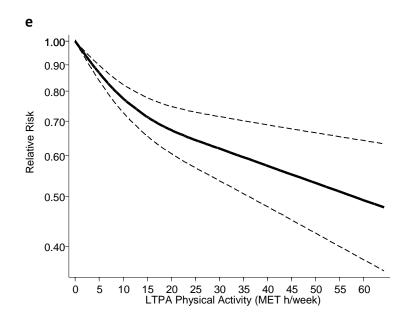


**ESM Figure 3** Funnel plot for the linear association of the RR for type 2 diabetes against the standard error of the RR per 10 MET h/week increment of physical activity



Dashed lines indicate the 95% confidence interval. Each data point represents one of the 32 observations extracted from 28 cohorts. Egger's test for publication bias: p=0.000

**ESM Figure 4** Comparison of the dose-response association and predicted RR point-estimates between LTPA and incidence of diabetes type 2 between the model using restricted cubic splines and the restricted cubic spline model including linear trend estimates of studies not eligible for flexible modelling



	II.	а	e <sup>1</sup>		
LTPA MET h/week	RR	95% CI		RR	95% CI
2.25	0.93	(0.92, 0.95)		0.94	(0.92, 0.95)
4.50	0.87	(0.84, 0.90)		0.88	(0.85, 0.91)
10.00	0.76	(0.71, 0.81)		0.77	(0.73, 0.82)
11.25	0.74	(0.69, 0.80)		0.76	(0.71, 0.81)
22.50	0.64	(0.56, 0.73)		0.66	(0.58, 0.74)
30.00	0.60	(0.51, 0.70)		0.62	(0.54, 0.72)
60.00	0.47	(0.34, 0.65)		0.50	(0.38, 0.65)

LTPA converted to MET h per week with results pooled in a two stage random effects model. RRs were derived from a common lowest physical activity category within each study. Listed exposure levels were chosen to represent meaningful and easy to interpret PA volumes equivalent to the following: 30 min of MVPA; 1 hour of MVPA; Rounded value to allow for comparison with GLS PA exposure increment; 150 minutes of PA/current recommended guidelines; double the recommended guidelines and two high PA exposure levels investigating the risk reductions at the higher end of the LTPA spectrum. The bold line indicates the pooled restricted cubic spline model and the black dashed line indicates the 95% confidence intervals of the pooled curve.

Duration assumption was necessary in 9 out of 27 observations and 9 out of 29 observations (applied as 45 min/session in (a) and (e) respectively). Intensity assumption was necessary in 15 out of 27 observations and 17 out of 29 observations (applied as LPA=3 MET, MVPA=4.5 MET, and VPA=8 MET in Scenario (a) and (e) respectively).

<sup>&</sup>lt;sup>1</sup> Model (e) includes additional observations from two independent cohorts that did not report sufficient data to include them in the full model. Sensitivity analysis using variance-weighted least squares (vwls) regression to estimate a linear-within study trend was applied to publications from the Women's Health Initiative (36) and the Shanghai Men's Health Study (37).

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