

1 **Leisure-time physical activity at moderate and high intensity is associated with**
2 **parameters of body composition, muscle strength and sarcopenia in aged adults with**
3 **obesity and metabolic syndrome** from the PREDIMED-Plus study.

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50 **ABSTRACT**

51 **Aims:** We aimed to examine the associations of leisure-time physical activity (PA) and
52 sedentary behavior (SB) with the prevalence of sarcopenia, body composition and muscle
53 strength among older adults having overweight/obesity and metabolic syndrome, from
54 the PREDIMED-Plus trial.

55 **Methods:** Cross-sectional baseline analysis including 1,539 men and women (65±5y).
56 Sarcopenia was defined as low muscle mass (according to FNIH cut-offs) plus low muscle
57 strength (lowest sex-specific tertile for 30-second chair-stand test). We applied
58 multivariable-adjusted Cox regression with robust variance and constant time (given the
59 cross-sectional design) for the associations of self-reported leisure-time PA and SB with
60 sarcopenia; and multivariable-linear regression for the associations with dual-energy X-
61 ray absorptiometry (DXA)-derived bone mass, fat mass, lean mass and lower-limb
62 muscle strength.

63 **Results:** Inverse associations were observed between sarcopenia and each hourly
64 increment in total [prevalence ratio 0.81 (95% confidence interval, 0.70, 0.93)], moderate
65 [0.80(0.66, 0.97)], vigorous [0.51 (0.32, 0.84)], and moderate-vigorous PA (MVPA)
66 [0.74 (0.62, 0.89)]. Incrementing 1-h/day total-PA and MVPA was inversely associated
67 with body-mass-index, waist circumference (WC), fat mass, and positively associated with
68 bone mass and lower-limb muscle strength (all $P<.05$). One h/day increase in total SB,
69 screen-based SB and TV-viewing was positively associated with body-mass-index, WC
70 and fat mass. Light-PA was not significantly associated with any outcome.

71 **Conclusions:** Total-PA and PA at moderate and high intensities may protect against the
72 prevalence of sarcopenia, have a beneficial role on body composition and prevent loss of
73 muscle strength. SB, particularly TV-viewing, may have detrimental effects on body
74 composition in older adults at high cardiovascular risk.

75 **Keywords:** Sarcopenia, Exercise, Sedentary behavior, Muscle, Epidemiology.

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77 INTRODUCTION

78 Aging-related changes in body composition are characterized by increased fat mass, loss
79 of skeletal muscle mass and declined muscle strength [1]. According to the European
80 Working Group on Sarcopenia in Older People (EWGSOP) [2], the coexistence of low
81 muscle mass and low muscle function (strength or performance) is referred as sarcopenia
82 Although the criteria for sarcopenia remains under debate, a more recent consortium of
83 the FNIH [3] has provided much insight in this topic, and has noted that among different
84 methods measuring body composition and muscle strength and performance, dual-energy
85 X-ray absorptiometry (DXA), handgrip or gait speed are the ones most suitable for
86 measuring muscle mass, muscle strength and performance, respectively. Importantly,
87 despite the differences in definition criteria and methodology used, the presence of
88 sarcopenia and/or these aforementioned changes in body composition have been
89 associated with increased risk of falls and fractures, and loss of independence [4].

90 Physical activity (PA) and sedentary behavior (SB) are major lifestyle factors known to
91 impact metabolic processes involving body composition. The ability of PA to
92 promote/preserve muscle mass and muscle strength while reducing fat mass has been
93 reported cross-sectionally [5–8] and prospectively [1,9,10] across different populations
94 of senior adults. Leisure-time PA consist of any recreational physical activity, also
95 including structured physical exercise, and household activities during spare time which
96 are not associated with regular occupation or transportation activities [11]. Consistent
97 beneficial associations with the aforementioned outcomes have been shown with total
98 leisure-time PA as well as with PA at moderate-vigorous intensities (MVPA), i.e.
99 activities ≥ 4 METs [5–8,10], whereas evidence concerning light-PA (< 4 METs) is
100 currently unclear [6,7].

101 SB refers to activities with prolonged periods of sitting or lying and activities with ≤ 1.5
102 METs of energy expenditure [12]. In the light of the recent Systems of Sedentary
103 behaviors consensus [13], priorities to foster SB research have been focused on home
104 setting factors, such as TV and computer use, among a number of other identified system-
105 based determinants. In fact, also recent studies have emphasized the importance of
106 context-specific SB, including screen-based SB (i.e. TV-viewing and PC use) for body
107 composition and muscular health [14,15]. While some authors have observed detrimental
108 associations with total SB and TV-viewing concerning sarcopenia [15], lean mass [8,15]
109 and muscle strength [6,14], others found no associations [7].
110 Despite these evidences and to the best of our knowledge, such associations of leisure-
111 time PA and SB with body composition, functional performance and sarcopenia have
112 never been explored in older adults with overweight/obesity and metabolic syndrome—a
113 population at higher risk of cardiovascular morbimortality, typically sedentary and
114 physically inactive. This population is likely to benefit from investigations addressing the
115 relationships between PA, SB, body composition, muscle health and sarcopenia.
116 Therefore, we aimed at examining the independent associations with leisure-time PA
117 levels and SB subtypes concerning body composition, muscle strength, and sarcopenia
118 prevalence among community-dwelling older adults with overweight/obesity and
119 metabolic syndrome participating in the PREDIMED-Plus trial.
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122 **METHODS**

123 **Study design and population**

124 This is a cross-sectional analysis of baseline data from the PREDIMED-Plus trial, a 6-
125 year lifestyle intervention for the prevention of cardiovascular morbimortality on 6,874
126 older adults. Detailed study information is available at <http://predimedplus.com/>. The trial
127 was registered at the International Standard Randomized Controlled Trial (ISRCT:
128 <http://www.isrctn.com/ISRCTN89898870>) with registration number 89898870 and date
129 of 24 July 2014.

130 Eligible participants were community-dwelling adults (aged 55-75 in men; 60-75 in
131 women) with body-mass-index (BMI) ≥ 27 and < 40 kg/m², and meeting ≥ 3 metabolic
132 syndrome individual components [16]. Briefly, exclusion criteria included previous
133 history of cardiovascular disease, any chronic medical condition (cancer, inflammatory
134 bowel disease, cirrhosis, etc), having acute infectious processes, any condition
135 compromising physical activity, psychiatric disorders, alcohol and drug abuse,
136 institutionalization, use of specific medications (cytotoxic agents, immune-suppressors,
137 etc) important weight loss in short time and any food allergy to Mediterranean diet foods.
138 Following the PREDIMED-Plus design and protocol, from the 6,874 total participants
139 enrolled in the PREDIMED-Plus trial, only a subsample underwent total dual-energy X-
140 ray absorptiometry (DXA) scans (n=1,564) in 7 out of the 22 PREDIMED-Plus recruiting
141 centers. From those 1,564 participants with DXA measurements, we excluded
142 participants with incomplete DXA data (n=25), resulting in an effective sample size of
143 1,539 for the present study.

144 Study protocol and procedures were approved following the ethical standards of the
145 Declaration of Helsinki by the Institutional Review Boards of the 7 participating centers.
146 All participants provided written informed consent.

147 **Physical activity and sedentary behavior**

148 Leisure-time PA was evaluated with the validated REGICOR questionnaire [17] which
149 included questions to collect information about the type of activity, frequency (number
150 of days) and duration (min/day) performed during a representative month. Time spent in
151 moderate, vigorous and moderate-vigorous PA (min/day) was obtained as described
152 previously [18]. Time spent in total-PA was computed as the sum of the time from all PA
153 intensities.

154 SB were assessed on weekdays and weekends using the validated Nurses' Health Study
155 questionnaire for Sedentary Behaviors [19], which evaluates the time spent over the last
156 year in different sedentary domains according to 12 time-categories. Time spent in total
157 SB and TV-watching (min/day) was further computed as described previously [18].
158 Screen-based SB was additionally estimated as the sum of the time spent watching TV
159 and using PC.

160 **Anthropometry and body composition**

161 Body weight and height were determined and BMI was calculated from weight (kg)
162 divided by height (meters) squared. Waist circumference (WC, cm) was measured with
163 anthropometric tape following the PREDIMED-Plus operations protocol.

164 Dual-energy X-ray absorptiometry scans (DXA, Lunar iDXA and DXA Lunar Prodigy
165 Primo, GEHealthcare) were performed by trained radiology technicians to ascertain body
166 composition. In this analysis, total bone mass (g), fat mass (kg) and lean mass (kg) were
167 evaluated. Appendicular skeletal muscle mass (ASM, kg) was determined as the sum of
168 the muscle mass from the four limbs, divided by 1,000. Skeletal muscle mass index (SMI,
169 kg/m^2) was calculated with the equation $[\text{ASM (kg)}/\text{BMI}]$ [3].

170 **Muscle strength**

171 Lower-limb muscle strength was determined using the validated 30-second chair-stand
172 test [20], based on the times participants stand-and-sit in a chair within 30 seconds,
173 following an established protocol. The 30-second chair-stand test has been also
174 considered a test for measuring performance [21] and its validity has been previously
175 evaluated [20,21], showing good reliability across trials (90.2%) in community-dwelling
176 Spanish seniors.

177 **Sarcopenia**

178 Sarcopenia was defined following the EWGSOP criteria [2] as low muscle mass
179 simultaneously with low muscle strength. Low muscle mass was defined applying the
180 recommended cut-offs according to the FNIH criteria [3]; while low muscle strength was
181 considered as the lowest sex-specific tertile for the number of stands in the 30-second
182 chair-stand test .

183 **Other covariates**

184 Participants self-reported data on age, sex, education, marital and employment status,
185 smoking, diabetes and osteoporosis medication. Diabetes was defined as previous diagnosis
186 of diabetes or glycated hemoglobin (HbA1c) $\geq 6.5\%$, use of antidiabetic medication or
187 having fasting glucose >126 mg/dl in the screening visit plus fasting glucose >126 mg/dl at
188 baseline. Dietary factors known to relate with body composition and sarcopenia [22–24]
189 were determined using a semi-quantitative food frequency questionnaire [25]: daily intake
190 of calcium (mg/day), vitamin D ($\mu\text{g/day}$), total protein (g/day), monounsaturated and
191 polyunsaturated fats (mg/day), and consumption of olive oil (g/day) and alcohol (g/day).
192 Nutrients intake was determined using Spanish food composition tables [26,27].
193 Age was classified into ≤ 65 and >65 y. Smoking was categorized as current, former and
194 never smokers. Dichotomous variables were generated for current diabetes and osteoporosis

195 medication. Quintiles of daily intake of nutrients and consumption of olive oil and alcohol
196 were created.

197 **Statistical analyses**

198 Population baseline characteristics were presented as means \pm standard deviations (SD) for
199 quantitative variables, and percentages and numbers for categorical variables. One-way
200 ANOVA and Chi-square tests were used for differences in baseline characteristics across
201 sex categories and quartiles of time in total-PA. Linear regression models were employed to
202 examine the associations with 95% confidence intervals (CI) of 1-hour/day increment in
203 time spent in PA levels and SB subtypes with BMI, WC, body composition and lower-limb
204 muscle strength. Cox regression models with constant time of follow-up ($t=1$) and robust
205 variance, given the cross-sectional design, were fitted to assess the prevalence ratios (PR)
206 for sarcopenia for 1-hour/day increment in time spent in PA levels and SB subtypes.
207 According to methodologists, this method is better suited than logistic regression for cross-
208 sectional studies with frequent prevalent outcomes ($>10\%$ prevalence), such as the present
209 study, since it avoids the overestimation of the prevalence ratios derived from the odds ratios
210 when logistic regression is applied in analysis with frequent outcomes [28,29].

211 Three multivariable-adjusted models were examined. For all study outcomes, model 1 was
212 adjusted for sex and age. Model 2 was further adjusted for diabetes, smoking and alcohol
213 intake. Besides these covariates, and to avoid over-adjustment, model 2 in each outcome
214 was specifically adjusted for factors that may affect these associations. Thus, model 2 for
215 bone mass was further adjusted for osteoporosis medication, vitamin D, calcium, protein,
216 mono and polyunsaturated fats and olive oil, and fat mass and lean mass. Model 2 for body
217 fat mass and lower-limb muscle strength were further adjusted for lean mass. Model 2 for
218 lean mass, ASM and SMI were further adjusted for fat mass. Model 2 for sarcopenia was
219 further adjusted for BMI and consumption of protein. To precisely assess independent

220 associations of a single activity, model 3 was fitted including variables in model 2 plus time
221 spent on the rest of the activities. For instance, when evaluating the associations with total
222 PA, model 3 included variables in model 2 plus time spent in total SB, and *vice versa*. All
223 analyses were stratified by recruiting center.

224 Effect modification by sex, age (≤ 65 and > 65 y) and diabetes on the associations between
225 time spent in PA levels and SB regarding the study outcomes was evaluated with the
226 likelihood ratio test between the fully adjusted model and the same model adding the
227 interaction product-term. Because no effect modification was observed ($P > .05$) all analyses
228 were pooled for the entire study population.

229 To test the robustness of our findings, the following sensitivity analyses were conducted: a)
230 excluding participants engaging in zero minutes of light PA; b) excluding participants
231 engaging in zero minutes of MVPA; and c) evaluating the associations with sarcopenia, body
232 composition compartments and lower-limb muscle strength by fitting quartiles of time spent
233 in total PA, light PA and MVPA.

234 All statistical analyses were conducted using the software Stata14 (StataCorp) and P -values
235 $< .05$ were considered significant.

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242 **RESULTS**

243 On average, study participants were aged 65.3 ± 5.0 y, 52% were men and 14.2% were
244 classified as sarcopenic. Participants spent 74 ± 60 min/day engaging in total leisure-time
245 PA, of which 46 ± 54 min resulted from MVPA, contributing 62% of their total PA time.
246 Mean time of total SB was 5.8 ± 1.8 h/day, of which 3.1 ± 1.6 h/day was for TV-viewing.
247 A complete description of the general characteristics for the entire study population by
248 sex categories is displayed in **Table 1** and across quartiles of total PA in **Supplementary**
249 **table 1**.

250 In multivariable-regression analyses (**Table 2** and **Supplementary table 2**), 1-h/day
251 increment in total-PA and MVPA was inversely associated with BMI, WC [and fat mass,
252 and positively associated with bone mass and lower-limb muscle strength. Regarding
253 moderate-PA and vigorous-PA, similar results for these study outcomes were found (all
254 $P<.01$), except for associations between vigorous-PA and bone mass. Only 1-h/day increase
255 in vigorous-PA, but not other PA levels, was significantly associated with greater lean
256 mass and ASM.

257 Associations of time spent across SB subtypes with body composition and lower-limb
258 muscle strength are summarized in **Table 3** and fully displayed in **Supplementary Table**
259 **3**. One h/day increase in SB subtypes was positively associated with BMI, WC and total
260 fat mass (all $P<.01$). Only TV-watching was significantly associated with lower ASM.
261 Otherwise, associations with other body composition compartments and lower-limb
262 muscle strength remained non-significant.

263 No significant associations were found with light-PA and SMI (**Table 2** and **Table 3**).

264 **Figure 1** highlights the multivariable-adjusted PR of sarcopenia concerning 1-h/day
265 increment in PA levels and SB subtypes. Each daily hour increase of total, moderate,
266 vigorous and MVPA was significantly associated with 19% to 49% lower prevalence of

267 sarcopenia—with vigorous-PA exhibiting the lowest prevalence [0.51 (0.32, 0.84)]. No
268 significant associations with any SB subtype were observed.

269 Sensitivity analyses were consistent with the general results. When excluding participants
270 engaging in zero minutes of light-PA (n=1,091), all associations remained significant. Only
271 associations between vigorous-PA and lean mass [β 0.68 kg/m² (-0.24, 1.59)] and ASM [β
272 0.39 kg/m² (-0.09, 0.88)] lost significance in model 3. Results excluding participants
273 engaging in zero minutes of MVPA (n=1,306) were comparable to those from the general
274 analyses. The exceptions where the associations for vigorous-PA with lean mass [β 0.56
275 kg/m² (-0.13, 1.24); $P=0.110$] and ASM [β 0.36 kg/m² (-0.01, 0.72); $P=0.052$] Associations
276 with sarcopenia, body composition compartments and lower-limb muscle strength according
277 to quartiles of time spent in total PA, light PA and MVPA were consistent with the general
278 results (**Supplementary Table 4**).

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292 **DISCUSSION**

293 To the best of our knowledge, this is the first study conducted in Mediterranean old adults
294 at high cardiovascular risk that has examined the associations between PA levels and SB
295 subtypes with the prevalence of sarcopenia, body composition and muscle strength. Main
296 findings revealed that each daily hour increase in total, moderate, vigorous and MVPA
297 was significantly associated with 19%, 20%, 49% and 26%, respectively, lower
298 prevalence of sarcopenia. Further inspection of body composition and muscle strength
299 showed that increasing 1-h/day total-PA or MVPA was inversely associated with BMI,
300 WC and fat mass, but positively associated with bone mass and lower-limb muscle
301 strength.

302 Our findings regarding PA and sarcopenia are according to those from previous cross-
303 sectional [5,6] and prospective studies [10,30] among old individuals. Cross-sectional
304 analyses in 2,264 Korean older adults revealed that individuals engaging moderate- and
305 high-PA vs. light-PA, were 38% and 74%, respectively, less likely to present sarcopenia
306 [5]. Similar findings have been reported in European adults cross-sectionally [6] as well
307 as prospectively, like in the Reykjavik Study [10], where greater MVPA was associated
308 with 36% protection against sarcopenia incidence after 5 years.

309 We did not observe significant associations between sarcopenia risk and SB subtypes,
310 which partially supports some of the findings from previous studies in European and
311 Australian healthy senior adults using accelerometer-derived and self-reported SB
312 subtypes [6,15]. No significant higher risk for sarcopenia for each 30 minutes increase in
313 accelerometer-derived sedentary time was reported among 1,286 old British men [6].
314 Contrary, a 33% increased risk of sarcopenia for 1-h/day increased in self-reported total
315 sitting time, but not with TV-watching was observed in healthy aged adults [15]. Of note,
316 in the present study as well as in the aforementioned studies, these associations were

317 independent from MVPA. These discrepancies may be due to differences in methodology
318 used to quantify SB and to define sarcopenia or in the study population characteristics. In
319 view of these conflicting results, more prospective studies and clinical trials evaluating
320 the effect of time spent in SB on sarcopenia are warranted to clarify these results across
321 different aged populations.

322 The ability of moderate- and vigorous-PA to improve age-related changes in body
323 composition, have been cross-sectionally [7,8] and prospectively reported [1,9,10],
324 among different populations of lean [6,9] and overweight old adults [8,10] using both
325 self-reported and accelerometry-derived PA data. For instance, in a cohort of Australian
326 women, total PA and PA at moderate and high intensities were beneficially associated
327 with total lean mass, ASM and muscle strength [7]. In addition, a dose-response
328 relationship between PA intensity and lean mass and lower-limb muscle strength have
329 been reported among Tasmanian old adults [8]. Other Europeans and American cohorts
330 have prospectively confirmed these associations regarding muscle mass and muscle
331 strength with physical performance [9,10]. We also found support for this, yet only
332 vigorous-PA was significantly associated with lean mass and ASM. In line with this,
333 moderate-intensity exercise interventions have improved muscle strength/functionality,
334 but not the reversion of age-related muscle loss [31,32] –suggesting that higher PA
335 intensities may be necessary to achieve benefits in muscle mass among seniors [33,34].

336 In agreement with our findings regarding bone mass, prospective analyses in American
337 [35] and European cohorts [32,36], and meta-analyses of randomized control trials [37]
338 have also reported positive associations between PA and bone mineral density in seniors.
339 Light-PA was not significantly associated with any study outcome, which agrees with
340 most of previous studies finding marginal [6] or no significant associations [36,38,39]. It

341 is therefore plausible that an intensity threshold exists for PA to benefit body composition,
342 muscle strength and eventually, reducing sarcopenia risk.

343 Similar to our study, a number of previous investigations across different cohorts have
344 reported positive associations between SB subtypes and BMI, WC and fat mass [15,40],
345 and inverse associations with lean mass or ASM [8,15], using self-reported [15] and
346 accelerometer-derived data [8]. Although in the English Longitudinal Study of Ageing of
347 6,228 old adults, Hamer et al. reported significantly lower grip strength in those participants
348 watching TV ≥ 6 h/day had, compared to those watching TV < 2 h/day [14], the associations
349 with lower-limb performance/strength in this and other studies were consistently non-
350 significant [7,8,15], regardless of the methodology used. This suggests that detrimental
351 associations between SB and muscle strength may be site-specific, with particular
352 vulnerability at the upper extremities.

353 A number of limitations in this study should be acknowledged. First, the cross-sectional
354 design does not allow to make any causal inference of the observed associations, and
355 therefore reverse causation cannot be excluded. Second, despite using validated
356 questionnaires [17,19], the use of self-reported PA and SB data may be more subject to bias
357 in comparison to more objective methods or due to the fact that the questionnaires used in
358 our study captured different timeframes of habitual engagement of PA and SB. Third, the
359 sole use of 30-second chair-stand test to evaluate lower-limb muscle strength represents a
360 major limitation for the definition of sarcopenia in our study, since this test was not included
361 as recommended methodology to assess low muscle strength by the last FNIH report 2014
362 [3]. However, this test has previously shown to reflect also the individual's physical
363 performance, as it involves muscle mobilization requiring strength and further physical
364 demands, such as balance. Along with this, the 30-second chair-stand test showed good
365 validity/reliability in previous studies [20,21]. Fourth, directly-measured data on muscle

366 performance was not determined in our study, thus we could not assess the prevalence of
367 severe sarcopenia. Five, given the observational nature of the study, residual confounding
368 may remain despite adjusting for several potential confounders; and finally, our findings
369 are limited to Mediterranean old adults with overweight/obesity and metabolic syndrome,
370 thus they cannot be extrapolated to other populations. Our study also has strengths: the large
371 sample size of men and women, the control for potential confounders and the inclusion of
372 sensitivity analyses; the use of low muscle mass and low strength to define sarcopenia as
373 they are well recognized components for sarcopenia diagnosis according to EWGSOP [2],
374 and finally, we included DXA-derived body composition, which is preferred over other
375 methods due to its reliability.

376 **CONCLUSION**

377 The findings from the current study support the idea that the increment in daily time spent
378 in total-PA and PA at moderate and high intensities may protect against sarcopenia, improve
379 body composition and prevent muscle strength decline in Mediterranean older adults with
380 overweight/obesity and metabolic syndrome. Contrary, increasing SB, particularly TV-
381 watching, may exert detrimental effects on body composition. This illustrates the importance
382 of addressing sarcopenia prevention with focus on MVPA in older adults at high
383 cardiovascular risk.

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390 **STATEMENT OF AUTHORSHIP**

391 FA, AB, DC, MD-R, VM-S, RE, MF, EG-G, JW, JL, JL-M, AM, MAM-G, JMO, XP, DR,
392 PM-M, LS-M, CV, FT, JT, JV, JV-L and JSS collected all the data from the PREDIMED-
393 PLUS trial. NR-E and J.S.S designed the study; NR-E, AD-L and J.S.S performed the
394 analysis; NR.-E and J.S.S wrote the first draft of the manuscript and all authors contributed
395 to the editing of the manuscript. All authors approved the final version of the manuscript

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397 **CONFLICT OF INTEREST STATEMENTS**

398 The authors have declared that no competing interest exist.

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Table 1. Baseline characteristics of study population in the whole study population and according to sex categories.

General characteristics	Whole population N=1,539	Men n=803	Women n=736	P-value
Age, y	65.3±5.0	63.9±5.4	66.7±4.1	<.001
BMI, kg/m ²	32.5±3.3	32.3±3.2	32.8±3.5	.005
Waist circumference, cm	107.3±9.3	110.8±8.6	103.5±8.6	<.001
Obesity, n (%)	1,147 (74)	586(73)	561(76)	.144
Central obesity, n (%)	1,440 (94)	714(89)	726(98)	<.001
Type 2 diabetes, n (%)	467 (30)	246(31)	221(30)	.796
Hypertriglyceridemia, n (%)	643 (42)	359(45)	284(39)	.015
High blood pressure, n (%)	1,448 (94)	769(96)	679(92)	.004
Hyperglycemia, n (%)	1,037 (67)	543(67)	494(67)	.834
Low HDL-c, n (%)	764 (49)	372(46)	392(53)	.007
Depression, n (%)	396(26)	125(16)	271(37)	<.001
Sarcopenia, n (%)	219 (14.2)	131(16)	88(12)	.014
Body composition compartments				
Total bone mass, kg	2.6±0.5	3.0±0.4	2.1±3.1	<.001
Total fat mass, kg	34.1±7.3	3.2±7.2	3.6±6.9	<.001
Total lean mass, kg	48.3±9.8	56.2±6.3	39.8±4.3	<.001
ASM, kg	21.5±4.9	25.4±3.3	17.3±2.2	<.001
SMI, kg/m ²	8.0±1.2	8.8±0.9	7.1±0.7	<.001
30-second chair-stand test, n	13.1±5.3	15.0±5.3	13.4±5.2	<.001
Smoking habit, n (%)				
Never	655(43)	169(21)	486(66)	<.001
Former	692(45)	495(62)	197(27)	
Current	192(12)	139(17)	53(7)	
Education status, n (%)				

Primary education	745(48)	310(39)	435(59)	<.001
Secondary education	466(31)	271(34)	195(27)	
Academic/graduate	328(21)	222(27)	106(14)	
Marital status, n (%)				
Single/divorced	210(14)	97(12)	113(15)	<.001
Married	1170(76)	679(85)	491(67)	
Widower	159(10)	27(3)	132(18)	
Employment status, n (%)				
Working	324(21)	234(29)	90(12)	<.001
Non-working	305(20)	67(8)	238(32)	
Retired	910(59)	502(63)	408(56)	
Osteoporosis medication, n (%)	9 (0.6)	1(0.1)	8(1.1)	.013
Dietary intake				
Alcohol, g/day	11.7±15.6	18.1±17.5	4.8±8.2	<.001
Olive oil, g/day	43.2±15.9	43.3±15.6	43.1±16.2	.831
Total protein, g/day	97.2±22.6	98.3±22.7	96.1±22.5	.053
Monounsaturated fats, mg/day	56.1±15.8	58.1±15.2	53.9±16.1	<.001
Polyunsaturated fats, mg/day	17.0±6.7	17.1±6.7	16.2±6.6	<.001
Calcium, mg/day	1019.3±353.5	1005.7±350.9	1034.1±356.0	.116
Vitamin D, µg/day	5.9±3.2	5.8±3.2	5.9±3.3	.314
Total leisure-time physical activity,				
min/day	73.6±60.3	84.7±68.7	61.5±46.6	<.001
Light PA, min/day	27.9±33.4	29.5±35.6	26.2±30.8	.054
Moderate PA, min/day	32.7±47.3	40.3±55.3	24.5±34.9	<.001
Vigorous PA, min/day	12.9±22.2	14.9±25.4	10.8±17.8	.003
MVPA, min/day	45.7±53.7	55.3±61.9	35.3±40.7	<.001
Sedentary time, h/day	5.8±1.8	6.0±1.9	5.6±1.8	.004

Screen-based sedentary time¹, h/day	4.1±1.9	4.2±2.0	3.9±1.9	.032
TV-viewing time, h/day	3.1±1.6	2.8±1.5	3.3±1.7	<.001

Data presented as mean ± SD unless otherwise indicated. BMI, body-mass-index; HDL-c, high-density lipoprotein-cholesterol; PA, physical activity; MVPA, moderate-vigorous physical activity; SB, sedentary behavior; ASM, appendicular skeletal muscle; SMI, skeletal muscle index.

¹Screen-based sedentary time defined as the sum of time spent watching TV plus using computer.

Pearson's chi-square test for categorical and one-factor ANOVA for continuous variables.

P-value for differences between sex categories.

Table 2. Multivariable-adjusted β -coefficients (95% CI) for body composition and lower-limb muscle strength¹ per 60 min/day increase PA time.

	n	Total-PA	<i>P</i> -value	Light-PA	<i>P</i> -value	Moderate-PA	<i>P</i> -value	Vigorous-PA	<i>P</i> -value	MVPA	<i>P</i> -value
BMI, kg/m²	1539	-0.47 (-0.64, -0.30)	<.001	-0.20 (-0.49, 0.09)	.188	-0.50 (-0.71, -0.29)	<.001	-0.83 (-1.27, -0.38)	<.001	-0.57 (-0.75, -0.38)	<.001
WC, cm	1539	-1.23 (-1.66, -0.80)	<.001	-0.47 (-1.24, 0.29)	.233	-1.28 (-1.83, -0.73)	<.001	-2.43 (-3.58, -1.27)	<.001	-1.50 (-1.99, -1.02)	<.001
Bone mass, g	1539	23.1 (8.1, 38.1)	.003	20.7 (-5.6, 47.1)	.124	24.6 (5.47, 43.6)	.012	21.5 (-18.5, 61.5)	.292	23.9 (6.98, 40.9)	.006
Fat mass, kg	1539	-0.90 (-1.22, -0.58)	<.001	-0.18 (-0.74, 0.39)	.548	-0.98 (-1.39, -0.57)	<.001	-1.88 (-2.73, -1.02)	<.001	-1.16 (-1.52, -0.79)	<.001
Lean mass, kg	1539	0.02 (-0.23, 0.27)	.865	-0.39 (-0.83, 0.04)	.079	0.05 (-0.26, 0.37)	.739	0.66 (0.02, 1.33)	.048	0.17 (-0.11, 0.45)	.228
ASM, kg	1537	0.03 (-0.10, 0.16)	.646	-0.19 (-0.42, 0.04)	.108	0.04 (-0.13, 0.21)	.634	0.41 (0.05, 0.76)	.024	0.11 (-0.04, 0.26)	.141
SMI, kg/m²	1537	0.01 (-0.03, 0.05)	.773	-0.04 (-0.11, 0.03)	.272	0.01 (-0.04, 0.06)	.766	0.08 (-0.02, 0.19)	.127	0.02 (-0.02, 0.07)	.330
Lower-limb muscle strength, No. counts¹	1539	0.56 (0.30, 0.82)	<.001	-0.04 (-0.51, 0.43)	.868	0.51 (0.17, 0.83)	.003	1.87(1.17, 2.57)	<.001	0.77 (0.48, 1.07)	<.001

BMI, body-mass-index; CI, confidence interval; PA, physical activity; MVPA, moderate-vigorous PA; ASM, appendicular skeletal muscle; SMI, skeletal muscle index; WC, waist circumference. Adjusted for sex, age, study center, smoking, diabetes and quintiles of alcohol intake and time spend in other self-reported physical activities or sedentary behavior, as appropriate, for all study outcomes. For bone mass, the model was additionally adjusted for fat mass and lean mass, osteoporosis medication, and quintiles of intake of olive oil, protein, calcium, vitamin D and mono and polyunsaturated fats. For lean mass, ASM and SMI it was additionally adjusted for fat mass. For fat mass and 30-second chair-stand test, it was additionally adjusted for lean mass.

¹Lower-limb muscle strength determined with 30-second chair-stand test based on the number of counts participants stand-and-sit within 30s.

Table 3. Multivariable-adjusted β -coefficients (95% CI) for body composition and lower-limb muscle strength¹ per 60 min/day increase SB time.

	n	Total SB	<i>P</i> -value	Screen-based SB	<i>P</i> -value	TV-viewing	<i>P</i> -value
BMI, kg/m²	1539	0.25 (0.16, 0.35)	<.001	0.12 (0.04, 0.21)	.004	0.20 (0.10, 0.31)	<.001
WC, cm	1539	0.60 (0.35, 0.83)	<.001	0.41 (0.19, 0.63)	<.001	0.43 (0.17, 0.70)	.001
Bone mass, g	1539	0.02 (-8.21, 8.24)	.993	-2.88 (-10.5, 4.72)	.460	-4.64 (-13.7, 4.45)	.315
Fat mass, kg	1539	0.47 (0.30, 0.65)	<.001	0.33 (0.17, 0.49)	<.001	0.26 (0.07, 0.46)	.008
Lean mass, kg	1539	-0.09 (-0.15, 0.13)	.900	-0.06 (-0.19, 0.06)	.325	-0.13 (-0.28, 0.02)	.088
ASM, kg	1537	-0.01 (-0.08, 0.05)	.825	-0.01 (-0.08, 0.05)	.683	-0.09 (-0.17, -0.01)	.026
SMI, kg/m²	1537	0.00 (-0.02, 0.02)	.866	-0.01 (-0.03, 0.01)	.408	0.00 (-0.02, 0.03)	.882
Lower-limb muscle strength, No. counts¹	1539	-0.13 (-0.27, 0.02)	.083	0.01 (-0.12, 0.15)	.877	-0.14 (-0.30, 0.03)	.098

BMI, body-mass-index; CI, confidence interval; ASM, appendicular skeletal muscle; SMI, skeletal muscle index; WC, waist circumference; SB, sedentary behavior. Adjusted for sex, age, study center, smoking, diabetes and quintiles of alcohol intake and time spend in other self-reported physical activities or sedentary behavior, as appropriate, for all study outcomes. For bone mass, the model was additionally adjusted for fat mass and lean mass, osteoporosis medication, and quintiles of intake of olive oil, protein, calcium, vitamin D and mono and

polyunsaturated fats. For lean mass, ASM and SMI it was additionally adjusted for fat mass. For fat mass and 30-second chair-stand test, it was additionally adjusted for lean mass.

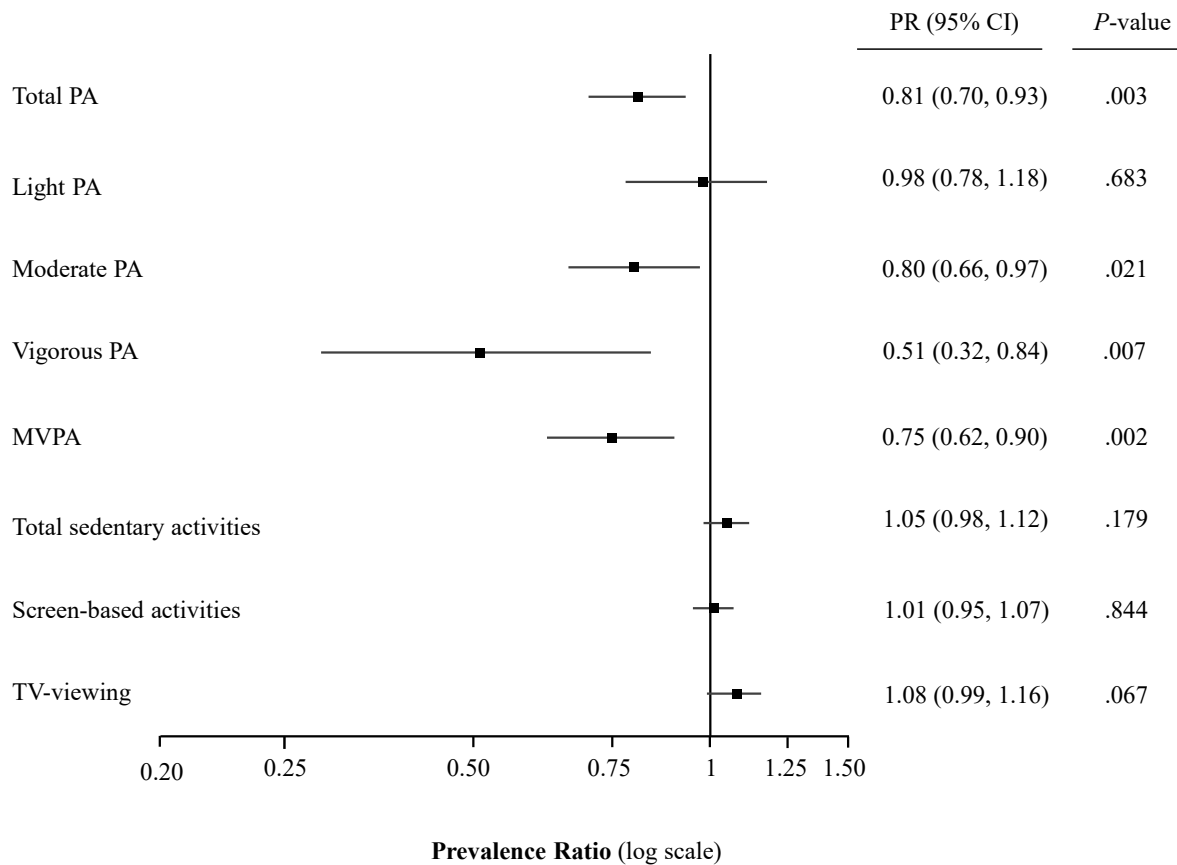


Figure 1. Multivariable-adjusted PR (95%CI) for sarcopenia per 60 min/day increase time in PA and SB (n= 1,537).

CI, confidence interval; PA, physical activity; MVPA, moderate-vigorous PA; PR, prevalence ratios. The model corresponds to the fully adjusted model, including sex, age, study center, smoking, diabetes, body-mass-index, quintiles of alcohol and protein intake and the corresponding time spent in other self-reported physical activities or sedentary behavior, as appropriate.