

1 **Association of midlife body composition with old-age health-related**
2 **quality of life, mortality, and reaching 90 years of age: A 32-year**
3 **follow-up of a male cohort**

4

5 **The Helsinki Businessmen study**

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7 Satu K. Jyväkorpi¹, Annele Urtamo¹, Mika Kivimäki¹, Veikko Salomaa², Timo E. Strandberg^{1,3}

8

9 ¹University of Helsinki, Clincum and Helsinki University Central Hospital, Unit of Primary Health
10 Care, Finland

11 ²THL-National Insitute for Health and Welfare, Helsinki, Finland

12 ³University of Helsinki, Clincum, and Helsinki University Hospital, Helsinki, Finland; University
13 of Oulu, Center for Life Course Health Research, Oulu, Finland

14

15 Corresponding author:

16 Satu Jyväkorpi

17 Tukholmankatu 8 B, 00014 University of Helsinki

18 Tel: +358 50 4920970

19 satu.jyvakorpi@gery.fi

20 Orchid ID: 0000-0001-5901-3584

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22 **Data share statement**

23 The data described used in the manuscript will be made available to editors upon request either
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26

27

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34 **Abbreviations:**

35 ANCOVA = Analysis of covariance

36 BF = Body fat

37 BMI = Body mass index

38 CI = Confidence interval

39 HBS = Helsinki Businessmen Study

40 HC = Hip circumference

41 HDL = High density lipoprotein

42 HRQoL = Health Related Quality of Life

43 LDL = Low density lipoprotein

44 OR = Odds ratio

45 SF = Short Form

46 SM = Skeletal muscle

47 WC = Waist circumference

48

49 **Abstract**

50 **Background:** Overweight and obesity increase risk of morbidity and mortality. The relationships
51 between body composition at midlife, health-related quality of life (HRQoL) in old age and
52 longevity are, however, less studied.

53 **Objective:** We examined the association of midlife body composition with successful aging,
54 defined as high HRQoL and reaching 90 years of age during 32-year follow-up.

55 **Design:** Participants were 1354 males from the Helsinki Businessmen Study, born 1919 to 1934. In
56 1985/86 (mean age 60 years) various health measurements were performed. Body fat (BF)% and
57 skeletal muscle mass (SM)% were calculated using validated formulas (including waist and hip
58 circumferences, weight and age) and divided into quartiles. In 2000 and 2007 (mean ages 74 and 80
59 years), HRQoL was assessed using RAND-36/SF-36 scales. Mortality was retrieved from registers
60 through 2018, and longevity determined by calculating the proportion of participants reaching 90
61 years. Logistic regression was used to assess odds ratios (OR) with 95% confidence intervals (CI).

62 **Results:** Higher SM% at midlife in 1985/86 was associated ($P<0.05$) with higher scores in RAND-
63 36 scales Physical functioning, Role limitations caused by physical health problems, Vitality, Social
64 functioning, and General health in old age in 2000. In 2007 only the association with Physical
65 functioning remained statistically significant ($P<0.01$). BF% quartiles in 1985/86 were inversely
66 associated with several RAND-36 scales in 2000 and 2007. During the 32-year follow-up, 982
67 participants died and 281 reached age 90 years of age. Being in the highest SM% quartile at midlife
68 increased (adjusted OR 2.32, 95% CI 1.53, 3.53; lowest SM% quartile as reference) and being in
69 the highest BF% quartile decreased (OR 0.43, 95% CI 0.28, 0.66; lowest BF% quartile as reference)
70 odds of reaching 90 years.

71 **Conclusions:** Desirable body composition in terms of both fat and skeletal muscle mass at midlife
72 was associated with successful aging in males.

73 **Keywords:** Body composition, skeletal muscle, body fat, quality of life, successful aging, longevity

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75

76 Introduction

77 Body composition is key to health and disease, and its derangements are a growing public health
78 problem (1). Body composition is the result of a wide range of factors including physical activity,
79 nutrition, disease, and age-related hormonal changes (2). Excessive body fat (BF) is associated
80 with various chronic diseases from midlife to old age. Skeletal muscle mass (SM), on the other
81 hand, is an important predictor of health in adult life, while severe loss of SM is linked to physical
82 frailty in old age (3). Moreover, SM is an important endocrine organ, affecting eg. glucose
83 metabolism, and low SM may impair glucose tolerance and increase insulin resistance and risk of
84 metabolic complications (4,5).

85 Excessive BF has been shown to be associated with lower muscle quality, and it predicts
86 accelerated loss of lean mass (6). Sarcopenic obesity (i.e. low muscle mass combined with obesity)
87 increases risk of poor functional outcomes compared to either of these conditions alone (7,8). Aging
88 further increases undesirable changes in body composition, such as reduction in lean body mass and
89 total body water and increase in total fat mass, even if the body weight is steady or reduced (9).

90 These changes in body composition occur often simultaneously with decline in physical
91 performance and may increase physical limitations. According to US statistics, 17% of people at
92 age of 50 years have one or more physical limitations, the corresponding figure being 43% at 80
93 years (10). Moreover, physical limitations either self-reported or measured predict onset of
94 disability (11). However, to best of our knowledge there are no studies on how midlife body
95 composition contributes to the health-related quality of life (HRQoL) in old age.

96 Here, we hypothesized that midlife body composition has long-standing consequences for health,
97 physical function and longevity. Therefore, we explored the relationships between body
98 composition (BF% and SM%) at midlife and successful aging, defined as a combination of high
99 HRQoL and reaching 90 years of age among males from the Helsinki Businessmen Study (HBS)
100 cohort during a 32-year follow-up.

101 Subjects and Methods

102 These are secondary analyses of HBS, a Finnish cohort originally consisting of 3490 Caucasian
103 males, born between 1919 and 1934, who have been followed-up since the 1964 (12, 13). All
104 participants in the cohort belong to the highest socio-economic class and they had been mostly
105 business leaders or executives during their working lives. In the present analysis we focused on a
106 representative sample of males in this cohort who had been clinically healthy (no chronic diseases
107 or medications) during the clinic visit in 1974 (n=1815) and who responded to a health survey and
108 underwent laboratory measurements in 1985/86 (n=1399; 81.9% of 1709 eligible) when
109 participants' mean age was 60 years (14).

110 *Measurements*

111 Because participants were living in various parts of Finland during the clinic visit in 1985/1986,
112 measurements were performed locally by trained, registered nurses, who were given written
113 instructions how to measure body mass index (kg/m^2) waist circumference (WC, cm) and hip
114 circumference (HC, cm) according to Larsson et al (15). In addition, questionnaires were used to
115 define their health, medications, and lifestyle, including current smoking (no/yes), weekly alcohol
116 consumption (beer, wine, spirits separately, consumption calculated as grams/week), and regular
117 physical activity (hours /week) (14). Laboratory measurements (blood pressure, fasting serum lipids
118 and blood glucose) were performed with standard methods in certified laboratories.

119 In 1985/86 only 13 males (1%) had body mass index below $20 \text{ kg}/\text{m}^2$. Because their exclusion did
120 not affect main results, all males were analyzed together.

121 In 2000, surviving males were assessed using mailed questionnaires including questions about
122 health, medications, current weight, smoking, alcohol consumption, and physical activity (12,16). In
123 addition, the questionnaire included the RAND-36 HRQoL instrument which is practically identical
124 to Short Form [SF]-36 (17). The instrument has been validated in the Finnish general population
125 (18) and consists of 8 scales: Physical functioning, Role limitations caused by physical health

126 problems, Role limitations caused by emotional health problems, Vitality, Mental health, Social
 127 functioning, Bodily pain, and General health (17). The questionnaire survey with RAND-36 was
 128 repeated in 2007. We calculated the 8 scales using standard procedures (17) and present the scales
 129 separately in the analyses. Using personal identification number, vital status was verified from the
 130 Population Information System of Finland through March 2018 and the proportion of males
 131 reaching 90 years of age was calculated.

132

133 *Calculation of percentages of SM and BF*

134 The BF% and SM% were calculated according to validated anthropometric formulas (19,20) for the
 135 participants who attended clinic visits in 1985/86 as follows: $BF\% = 0.567 \times \text{Waist Circumference}$
 136 $(WC) \text{ in cm} + 0.101 \times \text{age (years)} - 31.8$, and for $SM \text{ (kg)} = 39.5 + 0.665 \times \text{body weight (kg)} - 0.185$
 137 $WC \text{ in cm} - 0.418 \times \text{Hip Circumference (HC) in cm} - 0.08 \times \text{age in years}$. SM% is $SM \text{ (kg)}/\text{body}$
 138 $\text{weight (kg)} \times 100\%$.

139 *Statistical analysis*

140 The SM% and BF% were divided into quartiles. We used descriptive statistics, Armitage test for
 141 trend in proportions, and analysis of covariance (ANCOVA, Bonferroni test for multiple
 142 comparisons) to compare SM% and BF% quartiles. Because dichotomous outcome (yes/no) was
 143 known for all participants, logistic regression was used to compare SM% and BF% quartile groups
 144 in reaching 90 years of age. Other basic assumptions of logistic regression were also met.
 145 Participants were living in various parts of Finland and did not include siblings or family members,
 146 there was linearity in continuous variables used, absence of multicollinearity, and no strongly
 147 influential outliers. Odds ratios (OR) with 95% confidence intervals (CI) were calculated and
 148 adjusted for age and various lifestyle-related variables (current smoking yes/no; alcohol
 149 consumption, grams/week; weekly physical activity, cut-point median time of 3 hours) at baseline.

150 Because the outcome, reaching 90 years of age, was relatively common (>20%) and odds ratios
151 may overestimate associations between risk factors and common outcomes, we repeated data
152 analysis using log-binomial regression instead of logistic regression. This led to very similar
153 estimates, and therefore only logistic regression data are shown.

154 Statistical significance was taken as two-sided P-value <0.05, but because multiple tests were
155 performed in logistic regression, we also report as a sensitivity analysis ORs with P-value <0.001
156 (and 99.9% CIs) to control for type 1 error. Statistical analyses were performed using NCSS
157 statistical software (Kaysville, UT, www.ncss.com, version 8) and SPSS program, version 24
158 (SPSS IBM, Armonk, NY, USA).

159 *Ethics*

160 All participants signed informed consent and the follow-up study was approved by the ethical
161 committee of the Department of Medicine, Helsinki University Central Hospital and the study is
162 registered with ClinicalTrials.gov identifier: NCT02526082.

163 **Results**

164 The flow chart of the study is shown in **Figure 1**. Of the 1399 participants attending the clinic visit
165 in 1985/86, SM% and BF% could be calculated for 1342 and 1351 males, respectively. The quartile
166 cut-offs of SM% were: $Q_1 \leq 34.03339$; $Q_2 > 34.03339$ and ≤ 35.93856 ; $Q_3 > 35.93856$ and
167 ≤ 37.7719 ; $Q_4 > 37.7719$. The corresponding cut-offs of BF% quartiles were: $Q_1 \leq 25.795$; $Q_2 >$
168 25.795 and ≤ 28.7114 ; $Q_3 > 28.7114$ and ≤ 32.10375 ; $Q_4 > 32.10375$.

169 At baseline, higher SM% quartiles were linearly and statistically significantly ($P < 0.05$) associated
170 with various cardiovascular health indicators (**Table 1**), including lower BMI, lower waist and hip
171 circumference, lower systolic and diastolic blood pressure, lower fasting blood glucose levels,
172 higher serum high density lipoprotein (HDL) cholesterol, and lower triglyceride levels. Smoking,

173 coffee or tea drinking were not associated with SM% quartiles and consumption of alcohol was
174 inversely associated with the SM% quartiles ($P = 0.01$). Regular physical activity was more
175 frequent in higher SM% quartiles ($P < 0.01$).

176 BF% quartiles (Table 1) were linearly and statistically significantly ($P < 0.05$) associated with BMI,
177 waist and hip circumference, systolic and diastolic blood pressure, fasting blood glucose levels,
178 serum low density lipoprotein (LDL) cholesterol and triglyceride levels, and inversely associated
179 with HDL cholesterol level. Tea and coffee drinking were not associated with BF% quartiles,
180 whereas smoking and regular physical activity were inversely and consumption of alcohol
181 positively associated with increasing BF% quartiles (Table 1.).

182 The number of participants taking part of the follow-up survey in 2000 was 995 (90.5% of the 1100
183 eligible males). Their BMIs were somewhat lower than at baseline. However, those in SM%
184 quartile Q_1 at baseline had the highest BMI at follow-up, and those in the SM% Q_4 , the lowest BMI
185 (**Table 2**). Neither alcohol consumption nor smoking differed between SM% quartiles in 2000.
186 Regular physical activity at baseline was linearly associated with SM% quartiles at follow-up ($P <$
187 0.01).

188 In 2000, BMI was the highest in BF% Q_4 and declined linearly towards BF% Q_1 ($P < 0.01$). Alcohol
189 consumption was the lowest in the lowest BF% quartile and increased linearly toward the highest
190 BF% Q_4 ($P < 0.01$). Smoking did not differ between the BF% quartiles, whereas physical activity
191 was inversely associated with the BF% quartiles.

192 *Health Related Quality of life*

193 Higher SM% at baseline was linearly associated with higher scores in many of the RAND-36 scales
194 at follow-up in 2000 (**Table 3**). Significant associations were observed in Physical functioning,
195 Role limitations caused by physical health problems, Vitality, Social functioning, and General

196 health. In 2007, the associations with two physical subscales of RAND-36 remained statistically
197 significant: Physical functioning and Role limitations caused by physical health problems (Table 3).
198 BF% quartiles at baseline showed an inverse relationship with several RAND-36 subscales at
199 follow-up in 2000, including Physical functioning, Role limitations caused by physical health
200 problems, Role limitations caused by emotional health problems, Vitality, Social functioning, and
201 General health (Table 3). Several of these inverse relations between BF% quartiles and RAND-36
202 subscales remained at the second follow-up in 2007: Physical functioning, Role limitations caused
203 by physical health problems, Role limitations caused by emotional health problems, Bodily pain,
204 and General health (Table 3).

205 *Mortality and odds for reaching 90 years of age*

206 During 32 years of follow-up through March 2018, 982 (72.7%) participants died (Figure 1). Total
207 mortality was 83.3% (n=280), 74.4% (n=249), 70.2% (n=235), and 62.5% (n=209) with increasing
208 SM% quartiles ($P < 0.001$), respectively, and 64.9% (n=216), 69.8% (n=240), 74.3% (n= 252), and
209 81.1% (n=274) with increasing BF% quartiles ($P < 0.001$), respectively.

210 Through March 2018, 281 (20.8%) participants had reached 90 years of age. Both SM% and BF%
211 were associated with odds of reaching this age (**Table 4**). In the fully adjusted model (age plus
212 lifestyle variables in 1985/86), OR for the highest SM% quartile, as compared to the lowest quartile,
213 was 2.32 (95% CI, 1.53, 3.53). The corresponding OR for the highest BF% quartile, as compared to
214 the lowest quartile, was 0.43 (95% CI, 0.28, 0.66).

215 We also performed a sensitive analysis by setting the threshold for statistical significance at P-
216 value <0.001 , and the main results in the fully adjusted model remained significant. Accordingly,
217 OR for the highest SM% quartile, as compared to the lowest quartile, was 2.32 (99.9% CI 1.14,
218 4.08). The respective OR for the highest BF% quartile, as compared to the lowest quartile, was 0.43
219 (99.9% CI 0.23, 0.93).

220 Discussion

221 In our study among males, desirable body composition -- characterized by higher skeletal muscle
222 mass and lower body fat at midlife -- was associated with lower mortality and increased odds of
223 reaching 90 years of age. In addition, higher skeletal muscle mass in midlife was associated with
224 higher scores in the physical aspects of HRQoL at mean ages of 73 and 80 years. In contrast, higher
225 body fat percentage at midlife was inversely associated with the scores of several aspects of
226 HRQoL at both follow-ups. These results suggest that body composition at midlife has long-
227 standing consequences for longevity and quality of life in old age.

228 As societies are aging, healthy and active aging and good quality of life in old age are increasingly
229 important goals, but there are only few studies on long-term predictors of healthy aging. In a
230 previous report of the Helsinki Businessmen Study, we found that even metabolically healthy
231 overweight and obesity at midlife are related to reduced odds of successful aging (21). To the best
232 of our knowledge, the present study is the first to explore the associations between midlife body
233 composition and components of HRQoL in old age in a longitudinal study design with a very long
234 follow-up time (32 years). The lack of studies may be due to the fact that body composition
235 measuring devices were rarely available for scientific use decades ago. Thus, our study has a novel
236 design for using validated anthropometric formulas based on waist and hip circumference to
237 estimate skeletal muscle mass and body fat at midlife (19,20).

238 There are a number studies with cross-sectional or longitudinal designs with short follow-ups (1 to
239 3 years) on associations between body composition and HRQoL in older people (22-24). In those,
240 especially low muscle mass and higher amount of body fat have been associated with lower HRQoL
241 and mobility limitations (23). Our results suggest that midlife body composition is an important
242 predictor of mobility limitations in old age. There are several plausible explanations for this
243 finding. First, skeletal muscle mass naturally decreases with increasing muscle loss between 40-59

244 years due to hormonal and other lifestyle factors (25). Muscle loss is further accelerated at end of
245 lifespan. Although muscle loss with aging is a normal phenomenon, sarcopenia (also including loss
246 of strength) is a clinical condition that increases risk of falls, functional decline, frailty, and
247 mortality (3, 26,27). Thus, those who already have low percentage of skeletal muscle mass at
248 midlife may be at increased risk of sarcopenia in old age. This would explain the consistent
249 association with the physical component of HRQoL at mean ages of 73 and 80 in our study.
250 Second, the accumulation of body fat and especially the accumulation of visceral fat is associated
251 with harmful consequences of obesity, not only due to the accumulating cardiovascular-metabolic
252 burden, but also due to the increased risk of mobility limitations in old age (1, 24,28). It is therefore
253 plausible that obesity reduces HRQoL. Furthermore, sarcopenic obesity -- where both high body fat
254 and low skeletal muscle mass are present -- is a strong predictor of ill health and poor physical
255 function (29). Although most participants in our cohort were not obese, those in the highest quartile
256 of body fat percentage had consistently lower physical components of HRQoL in old age, and the
257 association remained significant at the last measuring point, where the participants had a mean age
258 of 80 years.

259 In our study midlife smoking was associated with lower body fat percentage at baseline. This is
260 consistent with the finding that tobacco smoking accelerates metabolism and may reduce appetite
261 (30). However, from baseline (1985/86) to the first follow-up (in 2000), the number of current
262 smokers reduced dramatically as some of them died and others ceased smoking. Only 6% reported
263 smoking at the first follow-up in 2000, and there were no differences in the prevalence of smoking
264 between SM% or BF% quartiles. Thus, those who survived and had higher HRQoL were not likely
265 to be smokers. Use of alcohol at baseline, on the other hand, was inversely associated with skeletal
266 muscle mass and linearly associated with body fat, and this association remained significant in body
267 fat quartiles at the latter follow-up in 2007. Alcohol is very energy dense and regular drinking may
268 increase food intake, and thus weight gain. The mean alcohol consumption in the highest BF%

269 quartile was 161 g/week, which equals to moderate drinking in males. However, within the group,
270 there were also those who drank considerably more, which may impair odds for successful aging.
271 Regular physical activity was inversely associated with BF, and linearly associated with SM mass.
272 These findings suggest that unhealthy lifestyle habits may cluster in persons with low SM mass and
273 high BF, contributing to adverse body composition and ultimately reduced HRQoL in old age.

274 **Strengths and limitations**

275 The main strength of our study is the very long follow-up time combined with high participation in
276 two follow-ups 7 years apart and reliable retrieval of information for males who reached 90 years of
277 age from national registers. Using anthropometric formulas instead of a body composition device
278 can be both advantageous and disadvantageous. The formulas we used to estimate percentage of
279 skeletal muscle mass and body fat percentage are well-validated (19,20). On the other hand, use of
280 a golden standard DXA-device to measure body composition would have given additional
281 information on an individual's body composition. This was not possible at the time the baseline
282 measurements were done.

283 Due to the nature of the observational study, no causal relationships can be determined on the basis
284 of this study. In addition, we did not have more detailed information about nutritional factors and
285 dietary intakes at midlife. A further limitation is that the cohort of male survivors in a long-term
286 observational study is obviously selected. The participants were surviving Caucasian males from
287 high socio-economic groups, and their health and characteristics probably differ from those of the
288 general population. Therefore, the results cannot be directly generalized to other populations.
289 However, homogeneousness of the cohort is also a strength through reducing confounding by
290 socioeconomic factors, which may be important in a study related to lifestyle.

291 A limitation of the HRQoL results is that the questionnaire in 1985/86 did not include items about
292 quality of life, and it is unknown whether differences between SM% and BF% already existed at

293 baseline. However, a proxy of HRQoL (and one of RAND-36 items) could be self-rated health,
294 which was asked from the participants using a 5-step scale, the same measure as used in the
295 Whitehall study (31) in 1974, i.e. 12 years earlier than the baseline of the current study (12). In
296 1974, no significant difference in self-rated health was observed between SM% quartiles ($P=0.58$)
297 nor between BF% quartiles ($P=0.24$)(unpublished observations).

298 Furthermore, the associations of SM% and BF% with HRQoL could be affected by mortality and
299 nonresponse, if these occurred differently in quartiles of SM% and BF%. However, longevity was
300 less probable with lower SM% and higher BF%, and nonresponders in 2007 – more frequent with
301 lower SM% and higher BF% (Figure 1) -- had worse HRQoL in 2000 (unpublished observations).
302 This suggests that differences in HRQoL between SM% and BF% quartiles, if anything, would
303 have been even larger, if all participants had responded.

304 Finally, our study sample was not very large, but the long follow-up time combined with the robust
305 results between the body composition and successful aging enhance the significance of this study.

306 Conclusions

307 Components of body composition, both low muscle mass and high fat mass at midlife, appear to
308 adversely affect health-related quality of life in old age and were associated with reduced odds of
309 successful aging.

310

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312 *Conflict of Interest*

313 SKJ, AU, MK, report no disclosures, VS has participated in a conference trip sponsored by Novo
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315 group meeting, TES reports various cooperation (educational, research, consultation) with several
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318 *Author's contribution*

319 SKJ and TES designed the analysis, TES and SKJ carried out the statistical analysis and all the
320 authors wrote the manuscript and approved the final version of it.

321

322 **References**

- 323 1. Arnlov J, Ingelsson E, Sundstrom J, Lind L. Impact of body mass index and the metabolic
324 syndrome on the risk of cardiovascular disease and death in middle-aged men. *Circulation*.
325 2010;121:230–236.
- 326 2. Guo SS, Zeller C, Chumlea WC, Siervogel RM. Aging, body composition, and lifestyle: the Fels
327 Longitudinal Study. *Am J Clin Nutr* 1999;3:405-411.
- 328 3. Morley JE, Argiles JM, Evans WJ, Bhasin S, Cella D, Deutz NE, Doehner W, Fearon KC,
329 Ferrucci L, Hellerstein MK et al. Nutritional recommendations for the management of sarcopenia. *J*
330 *Am Med Dir Assoc* 2010;11:391-396. Doi:10.1016/j.jamda.2010.04.014
- 331 4. Scott D, de Courten B, Ebeling PR. Sarcopenia: a potential cause and consequence of type 2
332 diabetes in Australia's ageing population? *Med J Aust* 2016;205:329-33. Doi:10.5694/mja16.00446.
- 333 5. Stefanaki C, Pervanidou P, Boschiero D, Chrousos GP. Chronic stress and body composition
334 disorders: implications for health and disease. *Hormones* 2018;17:33-43. Doi:10.1007/s42000-018-
335 0023-7.
- 336 6. Zamboni M, Mazzali G, Fantin F, Rossi A, Di Francesco V. Sarcopenic obesity: a new category
337 of obesity in the elderly. *Nutr Metab Cardiovasc Dis* 2008;18:388–95.
338 Doi:10.1016/j.numecd.2007.10.002.
- 339 7. Baumgartner RN. Body composition in healthy aging. *Ann NY Acad Sci* 2000;904:437–48.
- 340 8. Waters DL, Baumgartner RN. Sarcopenia and obesity. *Clin Geriatr Med* 2011;27:401–21.
- 341 9. Ritz P, Vol S, Berrut G, Tack I, Arnaud MJ, Tichet J. Influence of gender and body composition
342 on hydration and body water spaces. *Clin Nutr* 2008;27:740-746.
343 Doi.org/10.1016/j.clnu.2008.07.010.

- 344 10. Holmes J, Powel-Giner E, Lethbridge-Cejku M, Heyman. Aging differently: physical
345 limitations among adults aged 50 years and over: United States, 2001–2007. NCHS Data Brief
346 2009; No. 20.
- 347 11. Seidel D, Brayne C, Jagger C. Limitations in physical functioning among older people as a
348 predictor of subsequent disability in instrumental activities of daily living, *Age & Ageing*, 2011;4:
349 463–469. doi.org/10.1093/ageing/afr054.
- 350 12. Strandberg TE, Salomaa V, Strandberg AY, Vanhanen H, Sarna S, Pitkälä K, Rantanen K,
351 Savela S, Pienimäki T, Huohvanainen E, et al. Cohort Profile: The Helsinki Businessmen Study
352 (HBS). *Int J Epidemiol*. 2016;45:1074-1074h. Doi.org/10.1093/ije/dyv310.
- 353 13. Huohvanainen E, Strandberg AY, Stenholm S, Pitkälä KH, Tilvis RS, Strandberg TE.
354 Association of self-rated health in midlife with mortality and old age frailty: a 26-year follow-up of
355 initially healthy men. *J Gerontol A Biol Sci Med Sci*. 2016;71:923-8. Doi: 10.1093/gerona/glv311.
- 356 14. Salomaa V. Long-term effect of primary prevention measures of coronary heart disease on the
357 risk factor levels in middle-aged men. In Finnish with English summary, University of Helsinki,
358 Helsinki 1988
- 359 15. Larsson B, Svärsudd K, Welin L, Wilhelmsen L, Björntorp P, Tibblin G. Abdominal adipose
360 tissue distribution, obesity and risk of cardiovascular disease and death: 13 year follow up of
361 participants in the study of men born in 1913. *Br Med J* 1984;288:1401-1404
- 362 16. Strandberg A, Strandberg TE, Salomaa VV, Pitkälä K, Häppölä O, Miettinen TA. A follow-up
363 study found that cardiovascular risk in middle age predicted mortality and quality of life in old age.
364 *J Clin Epidemiol*. 2004;57:415-21. Doi: 10.1016/j.jclinepi.2003.09.013
- 365 17. Hays RD, Morales LS. The RAND-36 measure of health-related quality of life. *Ann Med*.
366 2001;33:350–357.

- 367 18. Aalto AM, Aro AR, Teperi J. RAND-36 as a measure of health-related quality of life.
368 Reliability, construct validity and reference values in the Finnish general population. Helsinki,
369 Finland: Stakes, Research Reports; No. 101, 1999,
- 370 19. Lean ME, Han TS, Deurenberg P. Predicting body composition by densitometry from simple
371 anthropometric measurements. *Am J Clin Nutr* 1996;63:4-14.
- 372 20. Al-Gindan YY, Hankey C, Govan L, Gallagher D, Heymsfield SB, Lean ME. Derivation and
373 validation of simple equations to predict total muscle mass from simple anthropometric and
374 demographic data. *Am J Clin Nutr* 2014;100:1041-1051. Doi:10.3945/ajcn.113.070466
- 375 21. Jyväkorpi SK, Urtamo A, Strandberg AY, von Bonsdorff M, Salomaa V, Kivimäki M, Luotola
376 K, Strandberg TE. Associations of overweight and metabolic health with successful aging: 32-year
377 follow-up of the Helsinki Businessmen Study. *Clin Nutr*. 2019;S0261-5614:30266-3. Doi:
378 10.1016/j.clnu.2019.06.011
- 379 22. Kim S, Leng XI, Kritchevsky SB. Body composition and physical function in older adults with
380 various comorbidities, *Inno Aging*, 2017;1:igx008, <https://doi.org/10.1093/geroni/igx008>
- 381 23. Verlaan S, Aspray TJ, Bauer JM, Cederholm T, Hemsworth J, Hill TR, McPhee JS, Piasecki M,
382 Seal C, Sieber CC et al. Nutritional status, body composition, and quality of life in community-
383 dwelling sarcopenic and non-sarcopenic older adults: A case-control study. *Clin Nutr* 2017;1:267 –
384 274. doi: 10.1007/s11136-018-1904-6.
- 385 24. Wang L, Crawford JD, Reppermund S, Trollor J, Campbell L, Baune BT, Sachdev P, Brodaty
386 H, Samaras K, Smith E. Body mass index and waist circumference predict health-related quality of
387 life, but not satisfaction with life, in the elderly. *Qual Life Res* 2018;27:2653-2665.
388 Doi:10.1007/s11136-018-1904-6.

- 389 25. Janssen I Heymsfield SB, Ross R. Low relative skeletal muscle mass (sarcopenia) in older
390 persons is associated with functional impairment and physical disability. *J Am Geriatr Soc*
391 2002;50(5):889-96. Doi: 10.1046/j.1532-5415.2002.50216.x
- 392 26. Schaap LA, van Schoor NM, Lips P, Visser M. Associations of sarcopenia definitions, and their
393 components, with the incidence of recurrent falling and fractures: the longitudinal aging study
394 Amsterdam. *J Gerontol A Biol Sci Med Sci* 2018;73: 1199–204. Doi:10.1093/gerona/glx245.
- 395 27. Cruz-Jentoft AJ, Bahat G, Bauer J, Boirie Y, Bruyère O, Cederholm T, Cooper C, Landi F,
396 Rolland Y, Sayer AA, Schneider SM et al. Sarcopenia: revised European consensus on definition
397 and diagnosis. *Age Ageing* 2019; 48:16–31. doi: 10.1093/ageing/afy16
- 398 28. Meadows R, Bower JK. Associations of anthropometric measures of obesity with physical
399 limitations in older adults. *Disabil Rehabil* 2018;1–6. doi:10.1080/09638288.2018.1516815.
- 400 29. Hirani V, Naganathan V, Blyth F, Le Couter DG, Seibel MJ, Waite LM, Handelsman DJ,
401 Cumming RG. Longitudinal associations between body composition, sarcopenic obesity and
402 outcomes of frailty, disability, institutionalisation and mortality in community-dwelling older men:
403 The Concord Health and Ageing in Men Project. *Age Ageing* 2017;46:413–420.
404 Doi.org/10.1093/ageing/afw214.
- 405 30. Perkins KA. Metabolic effects of cigarette smoking. *J Appl Physiol* 1992;2:401-9.
- 406 31. Marmot MG, Shipley MJ. Do socioeconomic differences in mortality persist after retirement?
407 25 year follow up of civil servants from the first Whitehall study. *Br Med J* 1996;313: 1177–1180
- 408
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TABLE 1. Age-adjusted characteristics of participants of the Helsinki Businessmen Study (HBS) according to skeletal muscle % and body fat % quartiles at baseline in 1985/86

SM% quartiles (Q) ¹	Q ₁ n = 336	Q ₂ n = 336	Q ₃ n = 335	Q ₄ n = 335	P-value
Characteristics					
Age, mean, years (SE)	61.0 (0.2)	60.5 (0.2)	59.3 (0.2)	58.4 (0.2)	< 0.01
BMI, kg/m ² (SE)	27.2 (0.2)	26.3 (0.2)	26.0 (0.2)	24.8 (0.2)	< 0.01
Waist circumference, cm (SE)	102 (0.4)	98 (0.4)	96 (0.4)	90 (0.4)	< 0.01
Hip circumference, cm (SE)	106 (0.3)	101 (0.3)	99 (0.3)	93 (0.3)	< 0.01
Use of alcohol, g/week (SE)	138 (7.7)	121 (7.7)	109 (7.7)	104 (7.7)	0.01
Current smoking, %	16	18	15	19	0.29
Regular physical activity, %	74	75	78	83	< 0.01
Regular physical activity, h/week (SE)	3.4 (0.2)	3.6 (0.2)	3.8 (0.2)	4.3 (0.2)	0.036
Coffee, cups/d (SE)	3.5 (0.1)	3.7 (0.1)	3.9 (0.1)	3.5 (0.1)	0.10
Tea, cups/d (SE)	1.1 (0.1)	1.2 (0.1)	1.1 (0.1)	1.2 (0.1)	0.45
Systolic BP, mm Hg (SE)	142 (0.9)	140 (0.9)	139 (0.9)	138 (0.9)	< 0.01
Diastolic BP, mm Hg (SE)	88 (0.5)	89 (0.5)	88 (0.5)	86 (0.5)	0.013
Fasting blood glucose, mmol/L (SE)	5.1 (0.1)	4.9 (0.1)	4.8 (0.1)	4.9 (0.1)	< 0.01
Cholesterol, mmol/L (SE)	6.5 (0.1)	6.5 (0.1)	6.4 (0.1)	6.5 (0.1)	0.84
HDL cholesterol, mmol/L (SE)	1.3 (0.0)	1.4 (0.0)	1.4 (0.0)	1.5 (0.0)	< 0.01
LDL cholesterol, mmol/L (SE)	4.4 (0.1)	4.4 (0.1)	4.4 (0.1)	4.5 (0.1)	0.70
Triglycerides, mmol/L (SE)	1.7 (0.1)	1.5 (0.0)	1.4 (0.0)	1.3 (0.0)	< 0.01
BF% quartiles (Q) ²	Q ₁ n = 333	Q ₂ n = 343	Q ₃ n = 339	Q ₄ n = 336	P-value
Characteristics					
Age, mean, years (SE)	59.2 (0.2)	59.8 (0.2)	60.1 (0.2)	60.0 (0.2)	0.015
BMI kg/m ² (SE)	23.2 (0.1)	25.1 (0.1)	26.5 (0.1)	29.3 (0.1)	< 0.01
Waist circumference, cm (SE)	86 (0.2)	94 (0.2)	99 (0.2)	108 (0.2)	< 0.01
Hip circumference, cm (SE)	92 (0.3)	98 (0.3)	101 (0.3)	107 (0.3)	< 0.01
Use of alcohol, g/week (SE)	93 (7.6)	110 (7.5)	106 (7.5)	161 (7.6)	< 0.01
Current smoking, %	21	16	17	14	< 0.01
Regular physical activity, %	86	80	78	66	< 0.01
Regular physical activity, h/week (SE)	4.4 (0.2)	3.8 (0.2)	3.9 (0.2)	3.0 (0.2)	< 0.001
Coffee, cups/d (SE)	3.4 (0.1)	3.6 (0.1)	3.7 (0.1)	3.8 (0.1)	0.26
Tea, cups/d (SE)	1.2 (0.1)	1.1 (0.1)	1.1 (0.1)	1.0 (0.1)	0.11
Systolic BP, mm Hg (SE)	134 (0.9)	140 (0.9)	142 (0.9)	144 (0.9)	< 0.01
Diastolic BP, mm Hg (SE)	84 (0.5)	87 (0.5)	89 (0.5)	90 (0.5)	< 0.01
Fasting blood glucose, mmol/L (SE)	4.7 (0.1)	4.8 (0.1)	5 (0.1)	5.2 (0.1)	< 0.01
Cholesterol, mmol/L (SE)	6.4 (0.1)	6.6 (0.1)	6.5 (0.1)	6.5 (0.1)	0.16
HDL cholesterol, mmol/L (SE)	1.5 (0.0)	1.4 (0.0)	1.3 (0.0)	1.3 (0.0)	< 0.01
LDL cholesterol, mmol/L (SE)	4.3 (0.1)	4.6 (0.1)	4.4 (0.1)	4.3 (0.1)	0.03
Triglycerides, mmol/L (SE)	1.2 (0.0)	1.4 (0.0)	1.6 (0.0)	1.9 (0.0)	< 0.01

BMI = body mass index; BF = body fat; ; Q = quartile; SE = standard error; SM = skeletal muscle. Armitage test for trend in proportions and analysis of covariance (ANCOVA) were used to compare the quartiles. ¹ The cut-offs for SM% are: $Q_1 \leq 34.03339$; $Q_2 > 34.03339$ and ≤ 35.93856 ; $Q_3 > 35.93856$ and ≤ 37.7719 ; $Q_4 > 37.7719$. ² The cut-offs for BF% are: $Q_1 \leq 25.795$; $Q_2 > 25.795$ and ≤ 28.7114 ; $Q_3 > 28.7114$ and ≤ 32.10375 ; $Q_4 > 32.10375$.

Table 2. Age-adjusted characteristics of participants of the Helsinki Businessmen Study (HBS) during follow-up in 2000 according to skeletal muscle % and body fat % quartiles at baseline

Baseline SM% quartiles (Q) ¹	Q ₁ n = 229	Q ₂ n = 244	Q ₃ n = 253	Q ₄ n = 269	P-value
Characteristics in 2000					
Age, mean, years (SE)	75 (0.2)	75 (0.2)	73 (0.2)	72 (0.2)	<0.01
BMI, kg/m ² (SE)	26.6 (0.2)	25.7 (0.2)	25.9 (0.2)	24.6 (0.2)	< 0.01
Use of alcohol, g/week (SE)	123 (9.1)	120 (8.8)	122 (8.7)	109 (8.4)	0.67
Current smoking, %	5.7	5.3	6.7	6.3	0.62
Regular physical activity, %	76	82	86	85	< 0.01
Regular physical activity, h/week (SE)	5.0 (0.4)	5.6 (0.4)	5.7 (0.3)	5.3 (0.3)	0.46
Baseline BF% quartiles (Q) ²	Q ₁ n = 264	Q ₂ n = 263	Q ₃ n = 240	Q ₄ n = 235	P-value
Characteristics in 2000					
Age, mean, years (SE)	73 (0.2)	74 (0.2)	74 (0.2)	74 (0.2)	0.01
BMI kg/m ² (SE)	23.4 (0.2)	24.9 (0.2)	26.4 (0.2)	28.4 (0.2)	< 0.01
Use of alcohol, g/week (SE)	103 (8.3)	110 (8.4)	116 (8.9)	146 (9.0)	< 0.01
Current smoking, %	6.8	5.7	6.3	5.1	0.50
Regular physical activity, %	90	84	85	72	< 0.01
Regular physical activity, h/week (SE)	5.9 (0.3)	5.5 (0.3)	5.3 (0.4)	4.7 (0.4)	0.13

BMI = body mass index; BF = body fat; ; Q = quartile; SE = standard error; SM = skeletal muscle. Armitage test for trend in proportions and analysis of covariance (ANCOVA) were used to compare the quartiles. ¹ The cut-offs for SM% are: Q₁ ≤ 34.03339; Q₂ > 34.03339 and ≤ 35.93856; Q₃ > 35.93856 and ≤ 37.7719; Q₄ > 37.7719. ² The cut-offs for BF% are: Q₁ ≤ 25.795; Q₂ > 25.795 and ≤ 28.7114; Q₃ > 28.7114 and ≤ 32.10375; Q₄ > 32.10375.

Table 3. Age-adjusted subscales of health-related quality of life (HRQoL) during follow-up according to different skeletal muscle % and body fat % quartiles at baseline

Baseline SM% quartiles (Q) ¹	Q ₁ n = 203 (2000) n = 115 (2007)	Q ₂ n = 228 (2000) n = 139 (2007)	Q ₃ n = 235 (2000) n = 167 (2007)	Q ₄ n = 257 (2000) n = 175 (2007)	P-value
RAND-36 scales in 2000 and 2007, points					
Physical Functioning (SE)					
2000	73 (1.5)	77 (1.4)	79 (1.4)	82 (1.3)	< 0.01
2007	68 (2.0)	73 (1.9)	75 (1.7)	77 (1.7)	< 0.01
Role limitations caused by physical health problems (SE)					
2000	60 (2.5)	69 (2.4)	70 (2.3)	74 (2.3)	< 0.01
2007	57 (1.4)	68 (3.1)	73 (2.9)	70 (2.8)	< 0.01
Role limitations caused by emotional health problems (SE)					
2000	73 (2.3)	77 (2.2)	79 (2.2)	79 (2.1)	0.23
2007	73 (3.1)	81 (2.7)	79 (2.5)	79 (2.5)	0.23
Vitality (SE)					
2000	66 (1.4)	69 (1.3)	70 (1.3)	71 (1.2)	0.039
2007	69 (1.7)	70 (1.5)	72 (1.4)	69 (1.4)	0.34
Mental Health (SE)					
2000	81 (1.6)	81 (1.1)	81 (1.1)	82 (1.1)	0.90
2007	80 (1.4)	81 (1.2)	83 (1.1)	80 (1.1)	0.40
Social Functioning (SE)					
2000	83 (1.4)	85 (1.4)	85 (1.3)	89 (1.3)	0.018
2007	82 (1.6)	83 (1.4)	85 (1.3)	84 (1.3)	0.25
Bodily Pain (SE)					
2000	76 (1.5)	79 (1.4)	78 (1.4)	81 (1.4)	0.059
2007	76 (1.9)	82 (1.7)	78 (1.6)	81 (1.6)	0.17
General Health (SE)					
2000	56 (1.2)	61 (1.2)	61 (1.2)	61 (1.1)	0.013
2007	55 (1.6)	61 (1.4)	60 (1.3)	61 (1.3)	0.75

Continues...

TABLE 3 continued

Baseline BF % quartiles(Q) ²	Q ₁ n = 244 (2000) n = 170 (2007)	Q ₂ n = 243 (2000) n = 174 (2007)	Q ₃ n = 224 (2000) n = 129 (2007)	Q ₄ n = 219 (2000) n = 129 (2007)	P-value
RAND-36 scales in 2000 and 2007, points					
Physical Functioning (SE)					
2000	82 (1.3)	80 (1.3)	75 (1.4)	74 (1.4)	< 0.01
2007	79 (1.7)	74 (1.7)	71 (1.9)	69 (1.9)	< 0.01
Role limitations caused by physical health problems (SE)					
2000	77 (2.3)	73 (2.3)	62 (2.4)	61 (2.4)	< 0.01
2007	76 (2.9)	68 (2.8)	67 (3.3)	56 (3.3)	< 0.01
Role limitations caused by emotional health problems (SE)					
2000	83(2.1)	77 (2.1)81	75 (2.2)	72 (2.3)	< 0.01
2007	80 (2.5)	(2.4)	76 (2.9)	70 (2.9)	0.03
Vitality (SE)					
2000	72 (1.3)	69 (1.3)	68 (1.3)	66 (1.3)	< 0.01
2007	72 (1.4)	70 (1.4)	69 (1.6)	67 (1.6)	0.14
Mental Health (SE)					
2000	83 (1.1)	81 (1.1)	81 (1.1)	80 (1.1)	0.20
2007	83 (1.1)	82 (1.1)	80 (1.3)	78 (1.3)	0.08
Social Functioning (SE)					
2000	88 (1.3)	87 (1.3)	84 (1.4)	83 (1.4)	0.027
2007	85 (1.3)	85 (1.3)	84 (1.5)	81 (1.5)	0.25
Bodily Pain (SE)					
2000	81 (1.4)	79 (1.4)	77 (1.4)	77 (1.4)	0.13
2007	82 (1.6)	77 (1.6)	81 (1.8)	76 (1.9)	0.02
General Health (SE)					
2000	63 (1.1)	61 (1.1)	58 (1.2)	58 (1.2)	< 0.01
2007	63 (1.3)	60 (1.3)	58 (1.5)	56 (1.5)	< 0.01

¹ The cut-offs for SM% are: Q₁ ≤ 34.03339; Q₂ > 34.03339 and ≤ 35.93856; Q₃ > 35.93856 and ≤ 37.77179; Q₄ > 37.77179. ² The cut-offs for BF% are: Q₁ ≤ 25.795; Q₂ > 25.795 and ≤ 28.7114; Q₃ > 28.7114 and ≤ 32.10375; Q₄ > 32.10375.

TABLE 4. Odds ratios of reaching 90 years of age during the 32-year follow-up of the Helsinki Businessmen Study according to skeletal muscle % and body fat % quartiles at baseline

	OR of reaching 90 years of age ¹			
	SM% Q ₁	SM% Q ₂	SM% Q ₃	SM% Q ₄
Model 1 (95%, CI)	1.0 (ref)	1.18 (0.79, 1.76)	1.90 (1.27, 2.83)	2.36 (1.56, 3.56)
Model 2 (95%, CI)	1.0 (ref)	1.19 (0.79, 1.79)	1.80 (1.19, 2.70)	2.32 (1.53, 3.53)
	OR of reaching 90 years of age ¹			
	BF% Q ₁	BF% Q ₂	BF% Q ₃	BF% Q ₄
Model 1 (95%, CI)	1.0 (ref)	0.72 (0.49-1.05)	0.78 (0.53-1.13)	0.41 (0.27, 0.62)
Model 2 (95%, CI)	1.0 (ref)	0.74 (0.50, 1.09)	0.77 (0.52, 1.13)	0.43 (0.28, 0.66)

BF = body fat; CI = confidence interval; OR = odds ratio; Q = quartile; SM = skeletal muscle. ¹OR was calculated using logistic regression with lowest quartile as reference (OR=1.0). Model 1 adjusted for age at baseline in 1985/86; Model 2 adjusted for age, smoking (yes/no), alcohol consumption (grams/week), and regular exercise (cut-point 3 hours/week) at baseline in 1985/86.

Legend for the figure:

Figure 1. Flowchart of the study. BF = body fat; SM = skeletal muscle.