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1 **Motivational counseling for reduction of sitting time.**

2 **A community-based randomized controlled trial in sedentary adults**

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38 **ABSTRACT**

39 **Background:** Sedentary behavior is regarded as a distinct risk factor for cardio metabolic morbidity
40 and mortality, but knowledge of the efficacy of interventions targeting reductions in sedentary
41 behavior is limited. **Purpose:** To investigate the effect of an individualized face to face
42 motivational counseling intervention aimed at reducing sitting time. **Design:** A randomized,
43 controlled, observer-blinded community-based trial with two parallel groups using open-end
44 randomization with 1:1 allocation. **Setting/participants:** A total of 166 sedentary adults were
45 consecutively recruited from a population-based study 'The Health2010 Study'. **Intervention:**
46 Participants were randomized to a control (usual lifestyle) or an intervention group with 4
47 individual theory-based counseling sessions. **Main outcome:** Objectively measured overall sitting
48 time (ActivPAL 3TM, 7 days); secondary measures were breaks in sitting time, anthropometric
49 measures and cardio metabolic biomarkers, assessed at baseline and after six months. **Analysis:**
50 Repeated measures multiple regression analyses. **Results:** N=93 were randomized to the
51 intervention group, n=73 to the control group, n=149 completed the study. The intervention group
52 had a mean sitting time decrease of -0.27 h/day, corresponding to 2.9% of baseline sitting time
53 (h/day), the control group increased mean sitting time by 0.06 h/day. The between-group difference
54 in change, -0.32 h/day (95% CI -0.87 to 0.24, p=0.26) was not statistically significant. Significant
55 differences in change in fasting serum insulin of -5.9 pmol/l (95% CI -11.4 to -0.5, p=0.03), in
56 HOMA-IR of -0.28 (95% CI -0.53 to -0.03, p=0.03) and in waist circumference of -1.42 cm (95%
57 CI -2.54 to -0.29, p=0.01) were observed in favor of the intervention group. **Conclusion:** Although
58 the observed decrease in sitting time was not significant, a community-based individually tailored
59 theory-based intervention program aimed at reducing sitting time may be effective for increasing
60 standing and improving cardio metabolic health in sedentary adults. Trial registration at
61 ClinicalTrial.gov (NCT00289237). Data were collected 2010-2012, and analyzed 2013-2014.

62 INTRODUCTION

63 Recent evidence suggests that sedentary behavior may be a distinct risk factor for cardiovascular
64 morbidity and mortality, independent of leisure-time moderate and vigorous physical activity
65 (MVPA).¹⁻⁹ The detrimental health effects of sedentary behaviors, (e.g. during TV-viewing)<sup>1;3-
66 5;10;11</sup> and overall sitting during the day^{2;6-8;12;13} have been shown in observational studies.
67 Additionally, the number of breaks in sitting time may be inversely associated with the cardio
68 metabolic risk profile.¹⁴ It has been proposed that targeting increased participation in MVPA may
69 not be sufficient to offset the health hazards associated with excessive sitting. In a group of healthy
70 middle-aged and elderly women, participation in sustained 10-minute bouts of objectively measured
71 MVPA was unrelated to objectively measured duration of daily sitting time.¹⁵ Recent experimental
72 studies also suggest that neither recommended levels of MVPA¹⁶ or restricted energy intake¹⁷ can
73 fully compensate the negative cardio metabolic health effects of sitting.

74 Sedentary behavior is defined as ‘any waking behavior characterized by an energy expenditure ≤ 1.5
75 Metabolic Equivalents while in a sitting or reclining position’.¹⁸ Adults in western societies spend
76 45-60% of their waking hours in sedentary pursuits, a stark contrast to the small proportion of total
77 time spent in MVPA (5%).¹⁹⁻²² Reduction of sitting time therefore represents a potential target for
78 promotion of health and prevention of chronic disease.

79 Recently, a number of studies have reported on the effects of interventions aimed at reducing and
80 breaking up sedentary time in different settings and populations.²³⁻²⁹ Generally, studies have been
81 small scale feasibility studies with theory-based behavioral interventions,^{26;27} short-term
82 experimental studies^{24;28} or work-site intervention studies conducted in small convenience
83 samples.^{23;25;29} Those studies have shown that reduction of sitting time is possible^{23;25-29} with
84 reductions ranging from 3-4% of total daily sitting time^{26;27} up to approximately 25% of working

85 hours.^{23;25;29} Additionally, positive cardio metabolic effects have been observed in some studies.^{23;24}
86 However, there is a need for randomized trials conducted in population-based study samples in
87 order to confirm that the possible beneficial effects of reducing sitting time are achievable in
88 community settings, and to obtain un-biased estimates of the effects on various health outcomes.
89 Therefore the aim of the present study was to investigate the efficacy of an individually tailored
90 theory-based, motivational counseling intervention on objectively measured sitting time, number of
91 breaks in sitting time, anthropometric measures and cardio metabolic biomarkers, in a population-
92 based sample of sedentary adult men and women. We hypothesized that daily sitting time could be
93 reduced, that number of breaks in sitting time could be increased and that cardio metabolic
94 biomarkers and anthropometric measures could be improved.

95 **METHODS**

96 **Setting and Participants**

97 Participants for the present randomized controlled trial (RCT), ‘Sedentary Intervention Trial’ (SIT)
98 were consecutively recruited from ‘The Health2010’ study, a population-based epidemiological
99 study carried out at The Research Centre for Prevention and Health (RCPH). A random sample of
100 3,762 men and women between 18 and 69 years of age and living in ten municipalities in the
101 Western part of the Capital Region of Denmark was initially drawn through the Danish Civil
102 Registration Office. The Health 2010 participants (N=1,522, response rate 40.5 %), who self-
103 reported at least 3.5 hours of daily leisure time sedentary behaviors were eligible and approached to
104 participate in the present sedentary behavior trial (‘SIT’). The 3.5 h/day corresponded to the median
105 daily leisure time sitting in a previous population study with similar age and gender distribution.³⁰
106 Additional eligibility criteria included: comprehension of the Danish language, self-report of

107 maximum 8 hours of vigorous physical activity per week³¹, not having a handicap or a functional
108 limitation that prevented reduction of sedentary behavior (e.g. being wheel-chair bound).

109 All eligible participants, approached in the period November 23rd 2010 – August 24th 2011, were
110 given thorough oral and written information about participation in the ‘SIT’ study by one of two
111 research nurses and written informed consent was obtained from all interested participants before
112 formal inclusion in the ‘SIT’. The six month follow-up examinations were concluded on the 1st of
113 March 2012. Ethical approval was obtained from the Ethics Committee of the Capital Region of
114 Denmark (H-1-2010-072) and the study was registered at ClinicalTrial.gov (NCT00289237).

115 **Randomization and blinding**

116 After inclusion, participants were instructed to wear an ActivPAL 3TM® inclinometer^{32;33} for a 7-
117 day period. The ActivPAL data were processed by a ‘blinded’ investigator and the data was not
118 shown to the participant. Participants were then allocated to the control or intervention group by
119 ‘open end’ randomization using computer-generated random numbers (Microsoft® Access,
120 ‘Random’ function), operated by a blinded data manager.

121 Participants returned to the centre after six months to initiate a follow-up 7-day ActivPAL
122 measurement followed by a health examination. Although this was an open trial whereby
123 participants and some of the researchers were aware of randomization group, the research staff
124 conducting the objective measurements and data processing was blinded to the randomization.

125 **Intervention and control group**

126 The behavior change intervention program consisted of four individual theory-based face to face
127 sessions conducted by one of two research nurses. The sessions took place with approximately six-
128 week intervals during a six month period, with the first session taking place on the day the

129 participants were randomized and notified about their allocation status. Each session lasted between
130 30 and 45 minutes. The behavior change intervention program was based on behavioral choice
131 theory,³⁴ incorporating individual behavior goal setting, self-efficacy,³⁵ and motivational
132 interviewing techniques.^{36,37} Participants set specific individual goals for change in sedentary
133 behavior by identifying adequate behavior substitutes or choices and by initiating small changes in
134 availability and access to sedentary behavior in their daily lives³⁴ (e.g. 'I will go to the kitchen for
135 coffee refill when watching TV instead of leaving the coffee pot on in the table in front of me', 'I'll
136 raise my desk to a standing position at the end of a work day, when going to lunch or for a meeting
137 during the day'). At each of the following sessions (sessions 2-4), behavior goals were reviewed
138 and evaluated. Together with the research nurse goals were modified and new goals were set.

139 The intervention program focused on 4 key messages or themes:

- 140 1. Reduce daily TV-viewing
- 141 2. Substitute sitting with standing when possible – at work and/or at home. (No time restrictions)
- 142 3. Break up prolonged sitting – by standing up frequently
- 143 4. Maximum 30 minutes of sitting per episode

144 Written materials containing these key messages, including strategies and suggestions for reduction
145 of sitting time, were handed out to participants at each session in purposefully developed booklets
146 and on postcards and stickers. One key message was introduced at each session along with the
147 corresponding written material. No specific messages were directed at MVPA.

148 Participants randomized to the control group were instructed to maintain their usual lifestyle from
149 randomization to the end of the 6 month follow-up period. After the follow-up assessments were
150 completed at the end of the study, participants in the control group were given a single 45-minute

151 session with individual counseling on reduction of daily sitting time – corresponding to a shortened
152 or compact version of intervention group program.

153 **Primary outcome**

154 The outcomes were assessed by identical methods in all participants at baseline and after six
155 months. The primary outcome, total volume of daily sitting, and the secondary outcome, number of
156 breaks in sitting time, were assessed by objective measurement over 7 days after inclusion in the
157 study, the *baseline period*, and again the last 7 days before the follow-up health examination, the
158 *follow-up period*. Measurements were obtained using an ActivPAL 3TM® Activity Monitor (PAL
159 Technologies, Glasgow, Scotland, UK). This is a small (2.0 x 1.4 x 0.3 inches) and light (20.1
160 grams) triaxial accelerometer-based device which is worn on the anterior upper right thigh and kept
161 in place by an adhesive pad, a Palstickie®, supplied with the monitor. The monitor uses
162 accelerometer-derived information about thigh position to estimate time spent in different body
163 positions (i.e. sitting/lying, standing and stepping). Data were processed using the ActivPAL
164 software (version 6.4.1) as 0.05 second events.³⁸ The ActivPAL monitor has previously been
165 validated against direct observation^{32;33} and is currently considered the best choice for objective
166 measurement of sitting/lying. The ActivPAL has also been found to be sensitive to change in sitting
167 time.³²

168 During the ActivPAL measurement period, participants kept a log sheet, reporting when they went
169 to bed at night, when they awoke in the morning, and any other time they were lying down sleeping
170 or resting in bed during the day. The ActivPAL was to be removed for showering, bathing, or
171 swimming, which was also reported.

172 **Secondary outcomes**

173 Self-reported sitting time at work and during leisure on an average weekday was measured by a
174 questionnaire, the Physical Activity Scale 2.1.,³¹ a modified version of the Physical Activity
175 Scale.^{39;40} Construct validity of PAS 2.1. has previously been established by cognitive
176 interviewing.³¹ Information on weekly MVPA, sociodemographic characteristics, and other lifestyle
177 variables was also reported by questionnaire.

178 Anthropometric measures: Height was measured without shoes to the nearest centimeter; weight
179 was measured in light clothing without shoes to the nearest 0.1 kg along with body fat % by
180 bioelectrical impedance (Tanita BC-420MA), and BMI was calculated as kg/m². Waist
181 circumference was measured midway between the lower rib margin and the iliac crest to the nearest
182 0.5 centimeter with a non-elastic tape measure.

183 Cardio metabolic biomarkers: Venous blood samples were drawn after an overnight fast (≥ 8
184 hours). Fasting serum insulin levels were analyzed using fluoro-immunoassay (Auto-Delfia; Perkin
185 Elmer, Waltham, MA, USA) and fasting plasma glucose levels were enzymatically analyzed by the
186 hexokinase/glucose-6-phosphate dehydrogenase (G6PDH) method (Hitachi 912; Roche diagnostics,
187 Indianapolis, IN, USA). Using the homeostasis model assessment (HOMA), hepatic insulin
188 resistance (HOMA-IR) and basal insulin secretion (HOMA %B) were estimated from fasting
189 glucose and insulin levels.⁴¹ The proportion of HbA_{1c} was assessed using high-pressure liquid
190 chromatography (HPLC, Tosoh G7, Roche Diagnostics). Total cholesterol, high-density
191 lipoprotein cholesterol (HDL), and triglycerides were measured by enzymatic procedures
192 (BoeringerMannheim, Germany). Low-density lipoprotein cholesterol (LDL) was calculated by
193 Friedewald's formula.

194 **Statistical analysis**

195 Data are presented as mean (standard deviation) or as frequencies (%). General linear **mixed** models
196 for repeated measures ANCOVA (SAS proc mixed) were performed to determine the effect of the
197 intervention on the primary and secondary outcomes by including an interaction term between
198 randomization group (intervention vs. control) and time from baseline to 6 months follow-up.
199 Selection of co-variates was based on being differently distributed between intervention and control
200 group at baseline and results are presented in an unadjusted model and a model adjusted for sex and
201 work status. **Loss to follow-up was assumed to be missing at random.**

202 All data processing and analyses were performed in SAS (version 9.3) and mixed model analyses
203 were performed using SAS Proc Mixed procedure. A significance level of 0.05 was chosen.

204

205 **Sample size**

206 Power calculations were performed before the initiation of the study. In a previous population-based
207 study of a similar age and gender distribution,³⁰ participants with ≥ 3.5 hours of self-reported leisure
208 time sitting per day had a mean self-reported leisure time sitting duration of 4.96 (1.64) hours per
209 day. We expected an intervention effect of approximately 45 minutes of reduction in sitting time per
210 day, based on findings in a feasibility study by Kozey-Keadle et al.²⁷ Assuming an alpha and beta of
211 0.05 and 0.2, respectively, we calculated that 150 participants should be included in the study.
212 Moreover, we expected approximately 10% attrition, and hence, included 166 participants for
213 randomization in the present study.

214

215 **RESULTS**

216 **Participant flow and baseline characteristics**

217 During the inclusion period, 689 Health2010 participants were assessed for eligibility. Among
218 these, 299 (43% of participants assessed for eligibility) met eligibility criteria (Figure 1), and 171
219 (57% of eligible participants) initially consented to participate and started the 7-day baseline
220 ActivPAL monitoring period. Five participants withdrew after initially accepting participation, but
221 before randomization; one due to skin reaction to the adhesive ActivPAL 'Palstickie', whereas four
222 withdrew for undisclosed reasons. Recruitment and inclusion was terminated when 166 participants
223 were consecutively randomized after completion of the pre-randomisation ActivPAL
224 measurements. Randomization resulted in 73 participants being assigned to the control group and
225 93 to the intervention group. Between randomization and the six month follow-up, five participants
226 (6.8%) withdrew from the control group and 12 participants (12.9%) withdrew from the
227 intervention group. The most frequently reported reason for withdrawal was lack of time (Figure 1).
228 There was a similar gender distribution (53% women) in participants (n=166) and non-participants
229 from the H2010 Study sample (n=523), whereas mean age in participants was slightly higher (51.3
230 (14.2)) than in non-participants (47.3 (13.7)).

231 Despite the rather uneven number of participants, the two randomized groups appeared relatively
232 well balanced (table 1) with the exception of an uneven gender distribution with 63 % women in the
233 intervention group vs. 49 % in the control group. Likewise, 60 % of participants in the intervention
234 group were not working, as opposed to only 47% in the control group. Median sitting time was 9.49
235 h/day (interquartile range: 8.19 – 10.50) in the intervention group and 9.64 h/day (interquartile
236 range 8.61 – 10.69) in the control group.

237 **Data quality and attendance**

238 The minimum number of valid days accepted for inclusion of ActivPAL data were 2 full days of
239 measurement. Measurements were only considered valid, if the monitor had been removed less than

240 2 hours per day for showering, bathing or swimming. At baseline, three participants had missing
241 ActivPAL data because of defective recording with the ActivPAL monitors or corrupt files. Two
242 participants had two full days of measurement only, 5 participants had 4 days of measurement, and
243 all others (n=156) had 5-7 days of measurement. At the six month follow-up, two participants had
244 incomplete and missing data due to monitor problems, two participants had 3 full days of
245 measurement only, 4 participants had 4 days of measurement, whereas all others (n=141) had 5-7
246 days of measurement. One of the participants with missing ActivPal data at baseline later dropped
247 out of the study leaving ActivPal data from n=79 and n= 66 participants in the intervention and the
248 control group, respectively.

249 Attendance to the counseling sessions in the intervention group was nearly complete. Only two
250 participants were unable to meet at the Centre for counseling session number four, but the
251 intervention material was sent by mail instead.

252 **Effectiveness of the intervention**

253 Objectively measured daily sitting time decreased in the intervention group and increased in the
254 control group, however, the difference in change between groups in favor of the intervention group
255 was not statistically significant -0.32 h/day (95% CI -0.87 to 0.24, p=0.26) (Table 2) With further
256 adjustment for sex and work status, the significance level was unchanged (table 2). In the
257 intervention group, the reduction in objectively measured sitting time was -0.27 h/day,
258 corresponding to a 2.9% reduction in daily sitting time compared to baseline. In the intervention
259 group the median change in sitting time was -0.14 h/day (interquartile range -1.09 to 0.72). In the
260 control group median change in sitting time was -0.07 h/day (interquartile range -0.7 to 0.68). In
261 the intervention group, 32% (n=25) reduced their daily sitting time at least 0.75 h/day, whereas a
262 similar reduction in sitting time was found in 24% (n=16) of the control group.

263 For objectively measured standing time, there was a significant difference in change in favor of the
264 intervention group of 0.44 h/day (95% CI 0.08 to 0.80, $p=0.02$). Results in favor of the intervention
265 group were also seen for objectively measured stepping time, however this was not statistically
266 significant ($p=0.11$). No significant differences in change in number of breaks were found (Table
267 2).

268 Change in self-reported leisure time sitting was significantly more pronounced in the intervention
269 than in the control group: mean difference in change from baseline to 6 months was -0.81 h/day
270 (95% CI -1.36;-0.27, $p=0.004$). In contrast, there were no significant differences in change in self-
271 reported sitting at work, (Table 2). For the other secondary outcomes, anthropometric and cardio
272 metabolic biomarkers, there was a mean difference in change in waist circumference from baseline
273 to 6 months of -1.42 cm (95% CI -2.54 to -0.29, $p=0.01$) in favor of the intervention group (Table
274 3). When adjusted for sex and work status, this finding remained statistically significant ($p=0.01$).
275 For weight and body fat, there were small differences in change in favor of the intervention group,
276 but differences were not statistically significant (Table 3). Finally, the difference in change in serum
277 insulin was -5.9 pmol/l (95% CI -11.4 to -0.50, $p=0.03$), and -0.28 (95% CI -0.53 to -0.03, $p=0.03$) in
278 insulin resistance, estimated by HOMA-IR, in favor of the intervention group. There was no
279 statistically significant effect of the intervention on serum lipids, fasting glucose, HOMA-%B or
280 HbA_{1c} (Table 3).

281 Analyses were also performed with sitting time expressed as percentage of waking hours (non-sleep
282 wear-time) and with break-rate (breaks per hour of sitting time), as suggested by Lyden et al.⁴²
283 However, results were essentially unchanged (data not shown).

284 **DISCUSSION**

285 In this population-based study sample of community-dwelling adults, a six month individually
286 tailored theory-based intervention program aimed at reducing daily sitting time was effective for
287 improving cardio-metabolic biomarkers. The overall sitting time was reduced by 2.9% in the
288 intervention group, but this reduction was not statistically different when compared to the control
289 group. However, standing time significantly increased in the intervention group and fasting serum
290 insulin level, HOMA-IR and waist circumference were reduced. To our knowledge this is the first
291 randomized trial to investigate the effects of a behavioral intervention targeted at reducing overall
292 sedentary behavior in a population-based sample of sedentary adults.

293 Recent non-worksite intervention studies have employed strategies aimed at reducing sedentary
294 behavior in older adults²⁶ or in overweight and obese adults.^{27;28} Only one of these studies had a
295 randomized control group and applied a TV-lockout intervention (usual daily TV-viewing time was
296 reduced to half by using a TV-lockout device),²⁸ whereas two studies used behavioral
297 interventions.^{26;27} Generally, these studies found reductions in objectively measured sitting time
298 from 3.2% to 4.3%.²⁶⁻²⁸ In comparison, the reduction in objectively measured sitting time in the
299 intervention group in our study was 2.9%.

300 Previous intervention studies have targeted the reduction in sitting time at the work site, either
301 through point-of-choice prompts to reduce sitting time at work,²⁵ installation of Sit-Stand
302 workstations,²³ or a multicomponent intervention comprising organizational, environmental and
303 individual elements.²⁹ One study found no effect on total sitting time, but a significant effect on the
304 number of and duration of sitting events.²⁵ Another study found a decrease in overall daily
305 ActivPAL measured sitting time of 97 minutes and an increase in HDL cholesterol of 0.26 mmol/l
306 in the intervention group, whereas no other significant biomarker differences were found.²³ A recent
307 study found a workplace sitting time reduction of 125 minutes per day following a four-week
308 intervention period, but no statistically significant effects on anthropometric or cardio-metabolic

309 health outcomes.²⁹ In comparison, in the present study a mean decrease of 16 minutes per day was
310 found in the intervention group, but no effect of the intervention on lipid parameters. However, the
311 present study differed markedly from the work site studies in several ways. The worksite studies
312 included - presumably rather homogeneous – samples of employees from academic institutions or
313 government agencies, whereas participants in this study were a relatively heterogeneous group of
314 adults recruited from a population-based study sample. The interventions in the occupational studies
315 included specific work site installations, thus the physical environment was altered to be more
316 conducive to standing during the work day, whereas the intervention in this study was behavioral,
317 individualized, and addressed all domains of daily life. Furthermore, the present study had a longer
318 intervention period.

319 The finding of significant differences in fasting serum insulin of 5.9 pmol/l and in HOMA-IR of
320 -0.28 in favor of the intervention group is consistent with results from the recent experimental study
321 by Stephens et al.¹⁷ The acute response of 1 day of prolonged sitting was a sizeable reduction (18%)
322 in insulin action over 24 hours compared to a condition where sitting was minimized. Likewise, in
323 another experimental crossover trial in overweight or obese adults,²⁴ Dunstan and colleagues found
324 an acute lowering effect of breaking up prolonged sitting time by light or moderate intensity
325 physical activity breaks on postprandial glucose and insulin responses, when compared to a day of
326 prolonged sitting. A study by Duvivier and colleagues found a beneficial effect of adding periods of
327 light intensity activity or MVPA to a 14 hour a day sitting regime on insulin sensitivity and **fasting**
328 **triglycerides and non-HDL cholesterol**.¹⁶ Another recent randomized crossover study by Peddie et
329 al found a beneficial effect on insulin and glucose levels of interrupting prolonged sitting by regular
330 short activity breaks (**vigorous treadmill exercise (7.4 METs)**), but no effect on triglyceride level.⁴³

331 The intervention effects in the present study appear modest from a clinical or individual
332 perspective. From a public health perspective, however, the magnitude of the improved fasting

333 serum insulin and the reduction in waist circumference, if sustained in the longer term, could
334 potentially have a significant impact at population level.⁴⁴

335 The present study has some strengths and limitations. Strengths include the objective measurement
336 of sedentary behavior and cardio-metabolic biomarkers, the relatively large population-based study
337 population, and the fact that the study was undertaken in a community setting under free-living
338 conditions. The latter two makes generalizability relatively high and findings can most likely be
339 generalized to other adult, populations with moderate to high sitting time. A limitation is that
340 outcome assessors were not fully blinded. The two nurses who conducted the motivational
341 counseling sessions also performed the six month follow-up examination. However, the primary
342 outcome, objectively measured sitting time, and cardio metabolic biomarkers were processed by
343 blinded investigators and are hence unlikely to have been affected by the nurses' knowledge of
344 participants' allocation status. Ideally, sitting time should have been measured continuously
345 throughout the intervention period, as the 7-day baseline and follow-up measurement periods are
346 not necessarily representative of sedentary behavior in the remaining 23 weeks. Consequently
347 adherence to the intervention program cannot be evaluated by objective measures. Another possible
348 limitation of the study is the uneven number of participants allocated to intervention and control
349 group. This risk exists when using 'open ended' randomization, which is, however, also considered
350 the optimal form of randomization, as it renders an equal chance of group allocation to every
351 participant. Finally, no detailed measurement of dietary habits was included in the present study. It
352 therefore cannot be ruled out that intervention group participants were generally made conscious of
353 lifestyle health issues and improved their dietary habits as a result of the intervention. This possible
354 spill-over effect of the intervention may potentially have contributed to the positive changes, for
355 example the reduction in waist circumference in the intervention group. Likewise, it cannot be fully
356 ascertained whether the observed changes in cardio metabolic markers have in fact occurred as a

357 result of reduction in sitting time or whether increases in MVPA have contributed. Although results
358 indicate that standing time increased significantly in the intervention group compared to the control
359 group, the intervention may have affected cardio metabolic biomarkers via various pathways.

360 The rate of drop-out was relatively low (10.2%), but it was still twice as high in the intervention
361 group relative to the control group. This indicates that the time commitment for the intervention
362 group participants may have been an obstacle to some participants.

363 The study population in the present study was rather heterogeneous with a large variation in
364 baseline sitting time, baseline MVPA level and change in sitting time. Future studies should
365 therefore aim to identify subgroups with high/low responsiveness. Maintenance of reduction in
366 sitting time and improvement of cardio metabolic biomarkers is another aspect that should be
367 further investigated in future studies with longer follow-up time.

368 In conclusion, our findings suggest that a six month individually tailored theory-based intervention
369 program may increase standing time, improve fasting serum insulin and reduce waist circumference
370 in sedentary men and women recruited from a population-based study sample. Future studies
371 should investigate different types of interventions targeted at reduction and break up of sitting time
372 in order to provide further directions for public health action against the detrimental health effects
373 of sedentary behavior.

374

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380

381 The authors declare that there was no racial or gender bias in the selection of participants.

382 **Contributorship**

383 TJ, DRW, DWD, AL and MA were responsible for the conception and design of the study. TJ, AL
384 MA, TR and SR were involved in the design of the study and development of the intervention. MA
385 analysed data and wrote first draft of the manuscript. DRW, DWD, TJ, AL and MA discussed data
386 analyses and interpretation and contributed to subsequent versions of the manuscript. DRW, DWD,
387 TJ, AL, TM, SR and MA critically revised the manuscript and approved the final version of the
388 manuscript.

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534 **Figure Legend**

535 **Figure 1:** Consolidated Standards of Reporting Trials (CONSORT) diagram of participants' flow
536 through the trial.

Figure 1: Consolidated Standards of Reporting Trials (CONSORT) diagram of participant's flow through the trial.

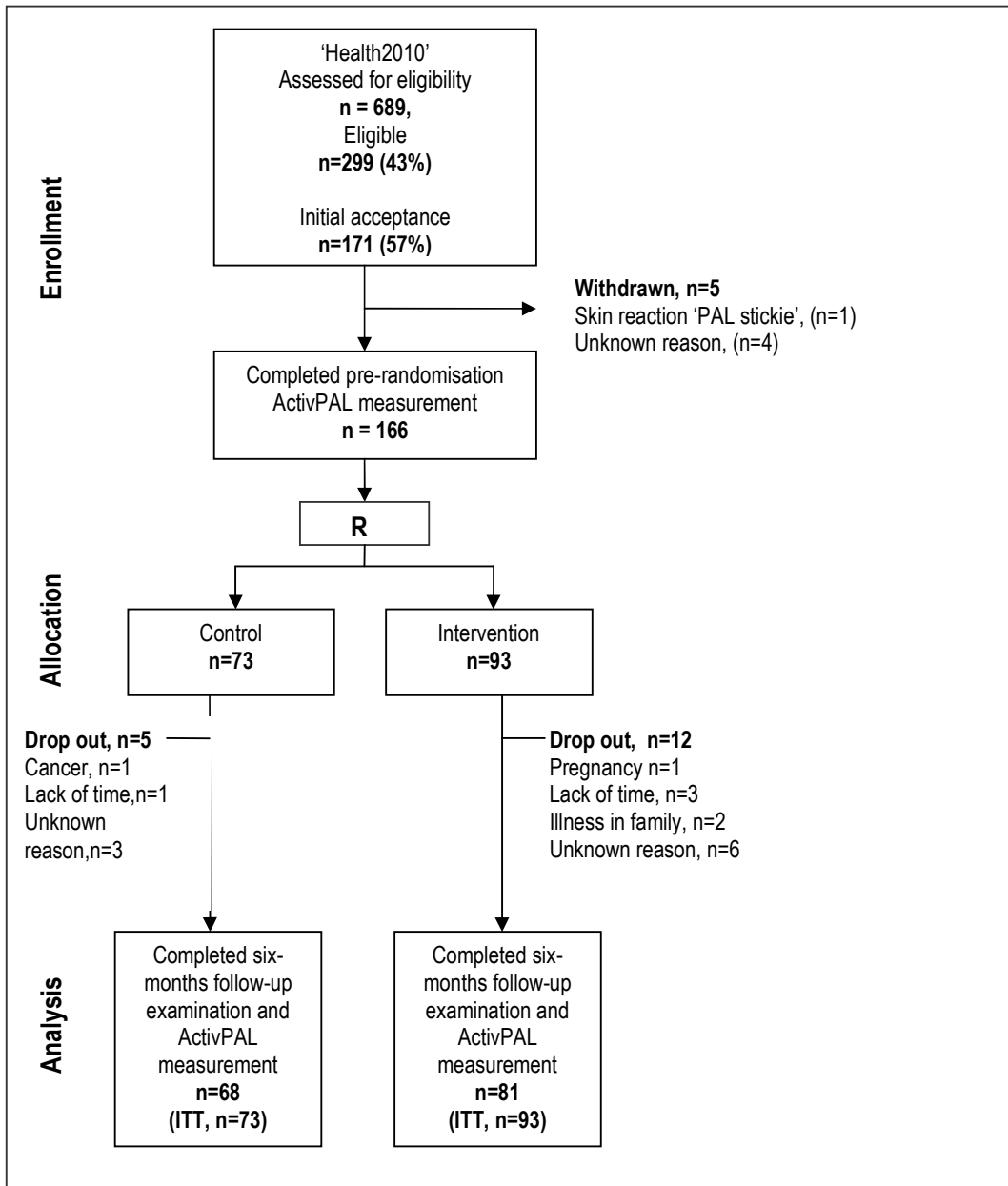


Table 1. Baseline characteristics of participants by randomisation group.
Data are presented as mean (SD), unless otherwise stated.

Characteristic	Intervention N=93	Control N=73
Women (%)	63%	49%
Age (years)	52.2 (13.8)	51.8 (14.3)
Seasonal variation, (included winter, (%))	51%	53%
Daily smokers (%)	16%	15%
Employment status (not working (%))	60%	47%
Sitting time (ActivPAL) ^a h/day	9.29 (2.0)	9.78 (1.8)
Standing time (ActivPAL) ^a h/day	4.15 (1.1)	4.10 (1.2)
Stepping time (ActivPAL) ^a h/day	1.78 (0.6)	1.74 (0.7)
Number of breaks (ActivPAL) ^a per day	60.2 (17.8)	59.6 (19.2)
Total non-sleep wear time (ActivPAL) ^a h/day	15.3 (1.2)	15.7 (1.0)
Self-reported vigorous physical activity h/week ^b	0.5 (0.0-2.0)	0.0 (0.0-1.0)
Self-reported moderate physical activity h/week ^b	2 (0.5-5.0)	2 (1-4.0)
BMI, kg/m ²	27.1 (5.1)	27.5 (4.9)
Weight, kg	79.0 (15.7)	82.1 (17.0)
Waist circumference, cm	92.7 (12.9)	95.2 (14.3)
Body fat (%)	32.8 (9.0)	32.1 (9.7)
Cholesterol, mmol/l	5.30 (0.9)	5.31 (1.1)
HDL cholesterol, mmol/l	1.57 (1.5)	1.51 (1.4)
LDL cholesterol, mmol/l	3.18 (3.0)	3.20 (3.0)
Triglycerides, mmol/l	1.17 (1.0)	1.32 (1.2)
Fasting plasma glucose, mmol/l	5.48 (0.9)	5.71 (1.1)
Fasting serum insulin, pmol/l	50.26 (43.4)	54.55 (46.9)
Fasting HbA _{1c} , %	5.57 (0.5)	5.59 (0.6)
Fasting HbA _{1c} , mmol/mol	37.5 (5.6)	37.6 (6.7)
HOMA-IR (insulin resistance) ^{b, c}	1.41 (0.9-2.1)	1.63 (1.1-2.6)
HOMA-%B (basal insulin secretion) ^{b, c}	68.3 (44.8-93.4)	73.0 (44.9-97.9)

^a ActivPAL baseline data for n=91 intervention group participants and n=72 control group participants only.

^b Median and interquartile range.

^c Homeostatis Model Assessment

Table 2: Change in sitting time, physical activity & wear time from baseline to 6 month follow-up.

Outcome (<i>intervention n; control n</i>)	Intervention group mean (SD)			Control group mean (SD)			Intervention vs. control group Mixed models [#]		
	Baseline	6 months	Within particip. change	Baseline	6 months	Within particip. change	Difference in change, between groups mean (95% CI)	p	p [§]
Objective measures by ActivPAL									
Sitting, ActivPAL, h/day (79;66)	9.3 (1.8)	9.0 (1.7)	-0.27 (1.7)	9.8 (1.8)	9.9 (1.8)	0.06 (1.7)	-0.32 (-0.87;0.24)	0.26	0.31
Breaks (sitting), ActivPAL, no/day (79/66)	59.9 (15)	60.3 (15)	0.5 (14.8)	59.2 (19)	59.7 (18)	0.4 (12.2)	-0.74 (-5.8; 4.4)	0.77	0.69
Standing, ActivPAL, h/day (79;66)	4.2 (1.1)	4.4 (1.3)	0.21 (1.0)	4.1 (1.2)	3.9 (1.3)	-0.22 (1.2)	0.44 (0.8; 0.80)	0.02	0.02
Stepping, ActivPAL, h/day (79;66)	1.8 (0.6)	1.9 (0.6)	0.1 (0.5)	1.7 (0.7)	1.7 (0.7)	-0.04 (0.6)	0.15(-0.04;0.33)	0.11	0.13
Non-sleep wear time, h/day (79/66)	15.2 (1.3)	15.3 (0.9)	0.04 (1.0)	15.6 (0.9)	15.4 (1.1)	-0.21 (0.9)	0.27 (-0.05;0.60)	0.09	0.09
Self-report measures									
Self-report sitting, (leisure) ,h/day (76;65)	5.3 (1.8)	4.4 (1.7)	-0.93 (1.6)	5.0 (1.7)	4.9 (2.2)	-0.03 (1.7)	-0.81 (-1.36;-0.27)	0.004	0.004
Self-report sitting (work), h/day (33;34)	4.4 (2.4)	4.0 (2.4)	-0.41 (1.3)	4.4 (2.4)	4.3 (2.4)	-0.05 (1.2)	-0.47 (-1.06;0.12)	0.12	0.11
Self-report vigorous PA, h/week (76;65)	1.3 (1.6)	1.3 (2.2)	0.07 (1.5)	0.8 (1.4)	0.8 (2.1)	0.05 (2.0)	0.29 (-0.20;0.78)	0.25	0.23

[#] Analyses performed by linear mixed models with a treatment x time interaction term characterizing the intervention effect of interest.

[§] p values adjusted for sex and work status.

Table 2: Change in anthropometric measures and biomarkers from baseline to 6 month follow-up.

Outcome (intervention n; control n)	Intervention group mean (SD)			Control group mean (SD)			Intervention vs. control group Mixed models #		
	Baseline	6 months	Within particip. change	Baseline	6 months	Within Particip change	Difference in change, between groups mean (95%CI)	p	p [§]
Anthropometrics									
Waist circumference cm (81;68)	93.5 (12.9)	92.3 (12.9)	-1.18 (4.0)	95.5 (14.5)	95.7 (15.0)	0.24 (2.7)	-1.42 (-2.54;-0.29)	0.01	0.01
Weight kg (81;68)	79.7 (15.7)	78.8 (15.2)	-0.84 (3.1)	82.1 (17.6)	82.1 (17.7)	0.007 (2.2)	-0.83 (-1.73;0.06)	0.07	0.07
Body Fat % (81;67)	32.9(8.9)	32.4 (9.2)	-0.5 (2.6)	31.4 (9.7)	31.6 (9.6)	0.2 (2.3)	-0.74 (-1.55;0.07)	0.07	0.08
Biomarkers									
Fasting plasma glucose, mmol/l (81;68)	5.5 (0.9)	5.3 (0.7)	-0.2 (0.5)	5.8 (1.1)	5.6 (0.9)	-0.2 (0.7)	0.003 (-0.20;0.19)	0.98	0.98
Fasting serum insulin, pmol/l (80;68)	49.4 (31.3)	43.2 (28.0)	-6.2 (16.1)	54.9 (33.2)	54.5 (35.2)	-0.4 (17.3)	-5.9 (-11.4;-0.50)	0.03	0.03
HbA _{1c} , % (81;66)	5.6 (0.5)	5.7 (0.5)	0.08 (0.3)	5.6 (0.6)	5.7 (0.6)	0.07 (0.2)	0.01 (-0.07;0.09)	0.73	0.73
HbA _{1c} , mmol/mol (81;66)	37.6 (5.8)	38.6 (5.5)	1.0 (3.3)	37.8 (6.9)	38.6 (6.2)	0.8 (2.3)	0.16 (-0.79;1.12)	0.74	0.74
HOMA-IR (80;68)	1.8 (1.4)	1.5 (1.1)	-0.3 (0.7)	2.1 (1.5)	2.1 (1.6)	-0.03 (0.8)	-0.28 (-0.53;-0.03)	0.03	0.03
HOMA-%B (80;68)	72.5 (42.1)	67.4 (36.9)	-5.1 (28.8)	77.1 (45.9)	80.2 (42.6)	3.0 (33.6)	-9.33 (-19.50;0.83)	0.07	0.08
Total cholesterol mmol/l (81;68)	5.3 (0.9)	5.1 (0.8)	-0.26 (0.7)	5.3 (1.1)	5.3 (0.9)	-0.08 (0.6)	-0.18 (-0.39;0.31)	0.09	0.12
HDL cholesterol mmol/l (81;68)	1.6 (0.4)	1.6 (0.4)	-0.03 (0.2)	1.5 (0.4)	1.5 (0.4)	-0.02 (0.2)	-0.0004 (-.077;.076)	0.99	0.89
LDL cholesterol mmol/l (81;68)	3.2 (0.9)	3.0 (0.8)	-0.21 (0.6)	3.2 (1.0)	3.2 (0.9)	-0.06 (0.5)	-0.15 (-0.33;0.04)	0.11	0.15
Triglycerides mmol/l (81;68)	1.13 (0.5)	1.07 (0.6)	-0.06 (0.4)	1.34 (0.7)	1.32 (-0.7)	-0.02 (0.6)	-0.06 (-0.23;0.10)	0.45	0.43

Analyses performed by linear mixed models with a treatment x time interaction term characterizing the intervention effect of interest.

§ p values adjusted for sex and work status.