

1 **Acute effects of breaking up prolonged sitting with standing and light-intensity**
2 **physical activity on cardiometabolic risk factors**

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17 **Breaking up prolonged sitting with light-intensity walking improves postprandial**
18 **glycemia, but breaking up sitting with standing does not**

19

20 **Abstract**

21 **Objectives:** To explore the effects of breaking up prolonged sitting time with standing or
22 light-intensity walking on a range of cardiometabolic risk markers.

23 **Design:** A randomized three-period, three-treatment acute crossover trial.

24 **Method:** Ten non-obese adults took part in three trials: 1) uninterrupted sitting; 2) seated
25 with 2-min bouts of standing every 20 min; and 3) seated with 2-min bouts of light-intensity
26 walking every 20 min. Two standardised test drinks (total 80.3 carbohydrate, 50 g fat) were
27 provided after an initial 1-h period of uninterrupted sitting. Plasma glucose and blood
28 pressure were assessed hourly to calculate total area under the curve (AUC). Total
29 cholesterol, HDL, and triglycerides were assessed at baseline and 5-h. ANOVAs were used
30 to explore between-trial differences.

31 **Results:** Glucose AUC was lower in the activity-break condition compared to the
32 uninterrupted sitting and standing-break conditions: mean AUC 18.5 (95% CI 17, 20), 22.0
33 (20.5, 23.5), and 22.2 (20.7, 23.7) mmol.L/5-h, respectively, $p<0.001$; no difference between
34 uninterrupted sitting and standing-break conditions ($p>0.05$). Systolic and diastolic blood
35 pressure AUC did not differ significantly between conditions, nor did responses in lipid
36 parameters ($p>0.05$).

37 **Conclusions:** This study suggests that interrupting sitting time with frequent brief bouts of
38 light-intensity activity, but not standing, imparts beneficial postprandial responses that may
39 enhance cardiometabolic health. These findings may have importance in the design of
40 effective interventions to reduce cardiometabolic disease risk.

41 **Keywords:** exercise; blood glucose; sedentary lifestyle; postprandial period

42 **Introduction**

43 A modern day sedentary lifestyle (prolonged sitting) may be a significant contributor to
44 hypokinetic disease risk¹. Individuals with high levels of sedentary time may have a 112%,
45 147%, 90%, and 49% increased relative risk of diabetes, cardiovascular events,
46 cardiovascular mortality, and all-cause mortality, respectively². To reduce disease risk,
47 interventions typically focus on engagement in moderate-to-vigorous physical activity
48 (MVPA)³. However, sedentary behaviour in itself is a risk factor for comorbidities and
49 mortality regardless of physical activity level^{2,4}.

50 Government guidelines recommend engagement in ≥ 150 min/wk of MVPA
51 accumulated in bouts of ≥ 10 min⁵. However, improvements in postprandial glycemia occur
52 following light-intensity activity similar to that observed following moderate- and vigorous-
53 intensity activity⁶. Observational data show that frequent interruptions to sitting time
54 (transition from sedentary to an active state for ≥ 1 min) are beneficially associated with
55 metabolic risk⁴. The mean duration of these breaks was approximately 4 min, which was
56 characterised by light-intensity physical activity⁴. Importantly, these relationships persisted
57 after accounting for MVPA, suggesting that frequent short breaks in sitting time may impart
58 unique benefit to health. Indeed, experimental data show interrupting prolonged sitting with
59 short bouts of walking improves postprandial glucose and insulin levels^{7,8}.

60 Interrupting sitting with standing could also impart health benefits⁹, although
61 experimental studies in humans is lacking. A combination of 4-h standing and 2-h walking
62 per day for four days improves fasting lipid levels and insulin sensitivity compared to
63 vigorous-intensity exercise for 1-h per day¹⁰. However, the independent effects of standing
64 were not explored and the potential for interrupting sitting with standing should be
65 investigated.

66 This study therefore investigates the acute effects of interrupting sitting with standing
67 or light-intensity walking on cardiometabolic risk markers in healthy adults.

68

69 **Methods**

70 The study was approved by the Sport Science and Physical Activity departmental ethics
71 review board at the University of Bedfordshire and conformed to the Declaration of Helsinki.
72 Written, informed consent was obtained from participants before any testing procedures
73 following a verbal and written explanation of the nature and risks involved in the
74 experimental procedures.

75 Ten healthy (free of any known metabolic or cardiovascular disease) participants with
76 no contraindications to physical exercise took part (7 men, 3 women; mean age, 24.0 ± 3.0 y;
77 mean BMI, 26.5 ± 4.3 kg/m²) in this randomised repeated measures cross-over design study.
78 Participants attended a familiarisation session where they became accustomed to the light-
79 intensity walking speed and familiarised with use of the Borg Rate of Perceived Exertion
80 (RPE)¹¹. During the familiarisation, RPE was recorded to ensure walking speed was
81 equivalent to light-intensity activity for each participant (RPE of 6-9). Participants then made
82 three separate visits to the laboratory to complete the 5-h trial conditions in a randomized
83 order: 1) uninterrupted sitting; 2) sitting interrupted by standing breaks; and 3) sitting
84 interrupted by light-intensity walking breaks. Because an acute bout of physical activity may
85 enhance insulin sensitivity for up to 72 h¹², a minimum wash-out period of 6 days between
86 each condition was used to eliminate potential carryover effects of the activity conditions.

87 Participants were instructed to refrain from any exercise, alcohol, or caffeine for 24 h
88 prior to each of the trial conditions. They attended the laboratory at 0900 h after an overnight
89 fast and sat for 1-h to achieve a steady state before a resting blood sample and blood
90 pressure measures were taken. Two standardised test drinks with a total of 80.3 g
91 carbohydrate were then consumed: 1) 75 g carbohydrate (100% dextrose monohydrate
92 powder; Thornton & Ross Ltd, UK) in 200 mL of water; energy, 273 kcal, and 2) 100 mL
93 drink consisting of 50 g fat (Calogen; Nutricia, UK); nutritional components were energy, 467
94 kcal; fat, 50.0 g; saturated fat, 5.3 g; carbohydrate, 4.3 g; sugars, 4.0 g; protein, nil; fibre, nil;
95 and sodium, 7.0 mg. The fat and protein content were included to 1) better simulate a mixed
96 meal and 2) help slow the ingested glucose production to spread the plasma glucose
97 responses over more of the 5-h treatment period¹³. Following consumption, the 5-h testing

98 period commenced. Participants were guided through each trial and supervised at all times
99 by a member of the research team to ensure full compliance with the protocols. Hourly blood
100 samples were collected and hourly blood pressure readings taken prior to the standing or
101 activity bouts during those respective conditions.

102 The trial conditions were as follows:

- 103 1. *Uninterrupted sitting*: participants remained seated throughout the experimental
104 period and were instructed to minimise excessive movement, only rising from the
105 chair to void.
- 106 2. *Sitting + standing breaks*: participants rose from the seated position every 20 min
107 throughout the experimental period (three breaks per hour) and stood as still as
108 possible for 2 min. They then returned to the seated position. This procedure was
109 undertaken on 14 occasions, providing a total of 28 min standing.
- 110 3. *Sitting + light-intensity activity*: participants rose from the seated position every 20
111 min and completed 2-min bouts of light-intensity walking on a motorised treadmill
112 (Woodway PPS55 Med-i, GmbH, Germany) with a level surface at 3.2 km/h,
113 providing a total of 28 min activity. They then returned to the seated position.

114

115 Participants watched television or DVDs; read books, magazines, or newspapers; or worked
116 on a laptop computer throughout the three conditions. Activity intensity during the sitting +
117 activity breaks was monitored at the completion of each activity bout using the Borg RPE
118 scale. Mean \pm SD (range: min–max) RPE was 6.7 \pm 0.9 (6–9).

119 Stature was measured to the nearest 0.1 cm using a stadiometer (Horltaim Ltd,
120 Crymych, UK) and body weight to the nearest 0.1 kg using electronic weighing scales
121 (Tanita Corp., Tokyo, Japan). Blood pressure was measured in a seated position using an
122 automatic device (Omron M5-I automated oscillatory device; Omron Matsusaka Co. Ltd.,
123 Matsusaka, Japan). Blood samples were obtained using a finger prick method and analysed
124 immediately. Glucose was determined hourly using the YSI 2300 STAT plus glucose and
125 lactate analyser (YSI Inc., Yellow Springs, Ohio, USA). The YSI uses a steady state

126 measurement methodology, where membrane based glucose oxidase catalyses the
127 oxidation of glucose to gluconic acid and hydrogen peroxide. The difference between the
128 sample generated plateau current and the initial baseline current is proportional to the
129 glucose concentration. The YSI was calibrated at the start of every day and every 45 min
130 thereafter. Total cholesterol, HDL, and triglycerides were obtained at baseline and 5-h and
131 determined using the Reflotron® Plus system (Roche Diagnostics, F. Hoffmann-La Roche
132 Ltd, Burgess Hill, UK). Reflotron® plus is a compact reflectance photometer for fully
133 automatic evaluation of Reflotron® tests. The instrument takes charge of all functions such
134 as heating, automatic calibration, test execution and evaluation and calculation of results.
135 The instrument has information on test principle and wavelength for each test and measuring
136 ranges. The YSI and Reflotron® systems were maintained according to manufacturers'
137 recommendations.

138 Sample size calculations were based on Dunstan et al⁷ who reported a 24%
139 reduction in 5-h positive incremental AUC (iAUC) when interrupting sitting with 2 min of light-
140 intensity walking every 20 min compared with uninterrupted sitting. 9 individuals were
141 required to achieve 90% power to detect the minimum effect size between the three
142 interventions, given a two-sided significance level = 5%.

143 Analyses were completed using SPSS version 19.0 (SPSS Inc., Chicago, IL.). Data
144 are presented as mean (95% CI). One-way ANOVA assessed between-trial condition
145 differences in pre-trial weight and cardiometabolic risk variables. Total area under the curve
146 (AUC) for each 5-h trial was calculated for glucose, systolic blood pressure, and diastolic
147 blood pressure using the trapezoidal method and between-trial condition differences
148 assessed using One-way ANOVA. Repeated measures ANOVA assessed differences
149 across conditions for pre- and post-trial lipid parameters. Estimates of effect size for
150 condition, partial eta squared (η^2), were calculated for each dependent variable. Statistical
151 significance was accepted as $p < 0.05$. Graphical representations of results are presented as
152 mean (SEM) to avoid distortion of the graphs.

153

154 **Results**

155 Biochemical and anthropometric data at baseline for each trial are shown in Table 1. There
156 were no significant differences for baseline values between trials.

157 Figure 1 shows glucose response over time during each of the trial conditions. A
158 significant effect of condition with a large effect size was observed ($F=8.59$, $p=0.001$,
159 $\eta^2=0.39$) for glucose AUC. As shown in Figure 2, after sitting + activity breaks (mean AUC,
160 18.5; 95% CI 17.0, 20.0 mmol.L/5-h) the glucose response to the test drink was 15.9% and
161 16.7% lower ($p<0.001$) compared to uninterrupted sitting (22.0; 20.5, 23.5 mmol.L/5-h) and
162 sitting + standing breaks (22.2; 20.7, 23.7 mmol.L/5-h), respectively.

163 There was no significant effect of condition and small effect size ($F=0.45$, $p=0.65$,
164 $\eta^2=0.03$) for systolic blood pressure AUC: mean AUC, 601.5 (95% CI 565.5, 637.5), 585.3
165 (549.3, 621.3), and 608.1 (572.1, 644.1) mmHg/5-h for uninterrupted sitting, sitting +
166 standing breaks, and sitting + activity breaks, respectively. There was no significant
167 condition effect but a medium effect size ($F=1.10$, $p=0.35$, $\eta^2=0.08$) for diastolic blood
168 pressure AUC: mean AUC, 356.7 (336.0, 377.4), 347.8 (327.1, 368.5), 335.6 (314.9, 356.3)
169 mmHg/5-h for uninterrupted sitting, sitting + standing breaks, and sitting + activity breaks,
170 respectively. Blood pressure responses over time are available as supplementary material
171 (Appendices 1 and 2).

172 There was no significant main effect for condition on changes in total cholesterol
173 ($F=0.01$, $\eta^2=0.00$), HDL ($F=0.09$, $\eta^2=0.01$), or triglycerides ($F=1.45$, $\eta^2=0.10$) from baseline to
174 5-h (Table 2), although a medium-large effect size was observed for triglycerides. There was
175 no significant interaction effect for total cholesterol ($F=1.79$, $p=0.19$, $\eta^2=0.12$; medium-large
176 effect size), HDL ($F=0.41$, $p=0.67$, $\eta^2=0.03$; small effect size), or triglycerides ($F=2.74$,
177 $p=0.08$, $\eta^2=0.17$; large effect size).

178

179 **Discussion**

180 The main finding of this study was that interrupting sitting time with short bouts of light-
181 intensity activity, but not standing, acutely lowers postprandial glycemia in healthy adults.

182 This supports recent observational and experimental data demonstrating the deleterious
183 health consequences of prolonged sitting and the potential benefits of frequent short bouts
184 of activity^{4,7,8}.

185 Prior to this study, the effect of interrupting sitting time with standing had not been
186 explored. Studies in animals and humans show impaired insulin action from reduced
187 standing and nonexercise ambulation⁹, while increases in standing time are associated with
188 reduced mortality rates¹⁴. It is postulated that energy demands to meet the requirements of
189 standing and the transition from sitting to standing may increase substrate utilisation that has
190 beneficial metabolic effects⁹. Improvements in insulin sensitivity have been reported
191 following four consecutive days of 4-h walking and 2-h standing daily compared to energy-
192 matched 1-h vigorous exercise daily or sitting for a full day¹⁰. Insulin action also increased
193 over a 24-h period when sitting time was replaced with standing and low- to moderate-
194 intensity walking¹⁵. However, these studies did not isolate the effects of standing from
195 walking and the role of standing alone cannot be inferred. Recent evidence in office workers
196 showed lower postprandial glucose AUC during an afternoon of working at a standing desk
197 compared to an afternoon of seated work and this was concomitant with a 174 kcal increase
198 in energy expenditure¹⁶. However, the current study revealed short frequent bouts of
199 standing had no effect on cardiometabolic risk markers over a 5-h period. These findings
200 may suggest that standing needs to be accrued in longer duration bouts or that a minimum
201 threshold increase in energy expenditure is required. As energy expenditure was not
202 measured the latter cannot be inferred from the current study.

203 In the current study, postprandial glucose AUC was lowered approximately 16%
204 when sitting was interrupted with 2 min of light-intensity walking every 20 min compared to
205 sitting + standing breaks or uninterrupted sitting. This is smaller than the 24-30% reduction
206 observed in overweight/obese adults when sitting was interrupted with light- or moderate-
207 intensity walking⁷. Dunstan et al⁷ also examined a 5-h postprandial period and interrupted
208 sitting with 2 min of activity every 20 min, thus disparities in magnitudes of change between
209 studies may be attributable to sample characteristic differences. Another study showed

210 interrupting sitting with 1 min 40 s bouts of moderate-intensity walking every 30 min reduced
211 glucose iAUC by 37% over a 9-h period compared to uninterrupted sitting⁸. These data
212 provide compelling support for interrupting sitting time with physical activity.

213 Postprandial hyperglycemia is a cardiovascular risk factor in people with Type 2
214 diabetes and even in nondiabetics^{17,18}. Reducing postprandial hyperglycemia improves
215 inflammation and endothelial function and reduces carotid intima-media thickness^{19,20} and
216 was also associated with a 49% relative risk reduction in the development of cardiovascular
217 events in individuals with impaired glucose tolerance²¹. The data from this current study and
218 others^{7,8} suggests that frequently interrupting sitting time with brief bouts of activity may be a
219 simple and effective approach to ameliorate such consequences of postprandial
220 hyperglycemia. Individuals who spend their working hours at a desk could reduce their
221 postprandial glycemia by rising from their chair and walking up and down corridors or
222 walking to other office areas to speak to colleagues instead of emailing or phoning.
223 Development of smartphone applications that frequently remind people to get up and move
224 around could also be a step forward in helping to interrupt prolonged sitting time.

225 The activity breaks in the current study accumulated to a total of 28 min of light-
226 intensity activity. In overweight adults, no differences in postprandial glucose AUC were
227 observed the morning after performing an acute 30 min bout of moderate-intensity aerobic or
228 resistance exercise, although insulin AUC was significantly lower²². Furthermore, a 45 min
229 moderate-intensity exercise session performed 30 min prior to a test meal resulted in lower
230 glucose (6%) and insulin (20%) AUC compared to no exercise and performing the same
231 exercise 17-h beforehand²³. It thus appears that interrupting sitting time with frequent short
232 bouts of activity has at least similar potential to continuous exercise bouts in lowering
233 postprandial glycaemia. Although the mechanisms for this response are not clear, it is
234 postulated that increases in carbohydrate oxidation may be important as this would augment
235 clearance of glucose from the blood⁸. The frequent nature of the activity bouts may have
236 also maintained increased permeability of muscle cells to glucose²⁴ and/or alter the
237 expression of genes that regulate translocation of the glucose transporter protein GLUT-4²⁵.

238 There were no significant differences observed in pre- vs post-trial lipid parameters
239 across conditions. Unfortunately, hourly measures were not taken and analysis of AUC
240 responses thus not possible. A medium-large effect size was observed for triglycerides,
241 though, and a larger sample size may have provided sufficient power to detect statistically
242 significant difference between conditions. However, Peddie et al⁸ reported no differences in
243 postprandial triglyceride iAUC when interrupting sitting with frequent bouts of activity
244 compared to uninterrupted sitting over a 9-h period. Postprandial hypertriglyceridemia is
245 recognised to affect endothelial function, atherogenesis, and is associated with coronary
246 artery disease^{26,27}. A continuous bout of moderate-intensity walking performed 1-h²⁸ and 16
247 to 18-h²⁹ prior to meal consumption resulted in lower postprandial triglyceride AUC and this
248 type of intervention may be preferred. The current study did not test triglyceride AUC
249 responses and comparisons to the aforementioned studies are thus difficult.

250 This study found no significant differences between continuous sitting versus
251 interruptions with standing or walking for systolic or diastolic blood pressure AUC, although a
252 medium effect size was seen for diastolic. Lower systolic blood pressure has been observed
253 the day following completion of either a continuous 30 min bout or ten 3 min bouts (30 min
254 rest between each) of walking, thus suggesting how physical activity is accumulated may be
255 unimportant³⁰. More research is needed to further understand the effects of interrupting
256 sitting on blood pressure and, as outlined previously, should incorporate larger sample sizes
257 to ensure sufficient statistical power.

258 This study has some limitations. No measure of insulin was taken so it is difficult to
259 infer the effects of interrupting sitting time on insulin sensitivity. However, previous research
260 suggests improved postprandial insulin action in response to interrupting sitting with brief
261 bouts of light- or moderate-intensity walking^{7,8}. Participants were asked to refrain from any
262 exercise 24 h prior to trial days. However, an acute bout of exercise may enhance insulin
263 sensitivity for up to 72 h¹² and this should be considered in future studies. The small sample
264 size may have resulted in under-powered *p* values and a larger sample may have detected
265 significant effects in variables other than glucose. Energy expenditure was not measured

266 during the trials and this may have added important mechanistic insights regarding the
267 observed differences between conditions. This study compared the effects of 1 day
268 exposure of prolonged sitting to interrupted sitting and the consequences of long-term
269 exposure cannot be extrapolated. Furthermore, activity and standing breaks of fixed
270 frequency and duration were examined and it would be important to explore whether
271 variations in these variables impart differential effects on cardiometabolic risk markers.

272

273 **Conclusion**

274 Frequent brief interruptions to sitting time with light-intensity activity, but not standing,
275 imparts beneficial postprandial responses that may reduce risk of cardiometabolic disease.
276 These findings add support for a public health focus on reducing and breaking up sitting time
277 alongside recommendations currently in place for physical activity. Future research should
278 explore effective and sustainable interventions to break up sitting time to inform potentially
279 effective public health initiatives.

280

281 **Practical implications**

- 282 • Prolonged sitting has detrimental effects on postprandial glycemia
- 283 • Frequently interrupting sitting time with light-intensity activity, but not standing,
284 improves postprandial glycemia
- 285 • Health-benefiting physical activity can be performed in bouts of as little as 2 minutes
286 in duration

287

288 **Acknowledgements**

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Table 1: Participant characteristics before each trial

	Uninterrupted sitting	Sitting + standing breaks	Sitting + light-intensity activity breaks	<i>P</i> value
Weight (kg)	87.6 (71.2, 103.9)	87.8 (70.9, 104.6)	87.6 (71.3, 103.8)	1.00
Plasma glucose (mmol.L)	4.42 (4.09, 4.75)	4.32 (3.97, 4.67)	4.39 (4.04, 4.74)	0.89
Total cholesterol (mmol.L)	4.03 (3.34, 4.73)	3.95 (3.24, 4.65)	4.11 (3.32, 4.89)	0.94
HDL (mmol.L)	1.68 (1.20, 2.16)	1.52 (1.05, 1.99)	1.59 (1.22, 1.97)	0.85
Triglycerides (mmol.L)	0.83 (0.77, 0.90)	0.82 (0.76, 0.87)	0.87 (0.78, 0.96)	0.50
Systolic blood pressure (mmHg)	122.1 (113.9, 130.3)	117.1 (110.0, 124.2)	120.2 (109.5, 130.9)	0.66
Diastolic blood pressure (mmHg)	72.9 (64.9, 80.9)	73.9 (68.9, 78.9)	70.2 (62.1, 78.3)	0.70

Data presented as mean (95% CI). HDL, high-density lipoprotein cholesterol.

Figure legends

Figure 1: The effect of the three trial conditions on 5-h postprandial plasma glucose levels.

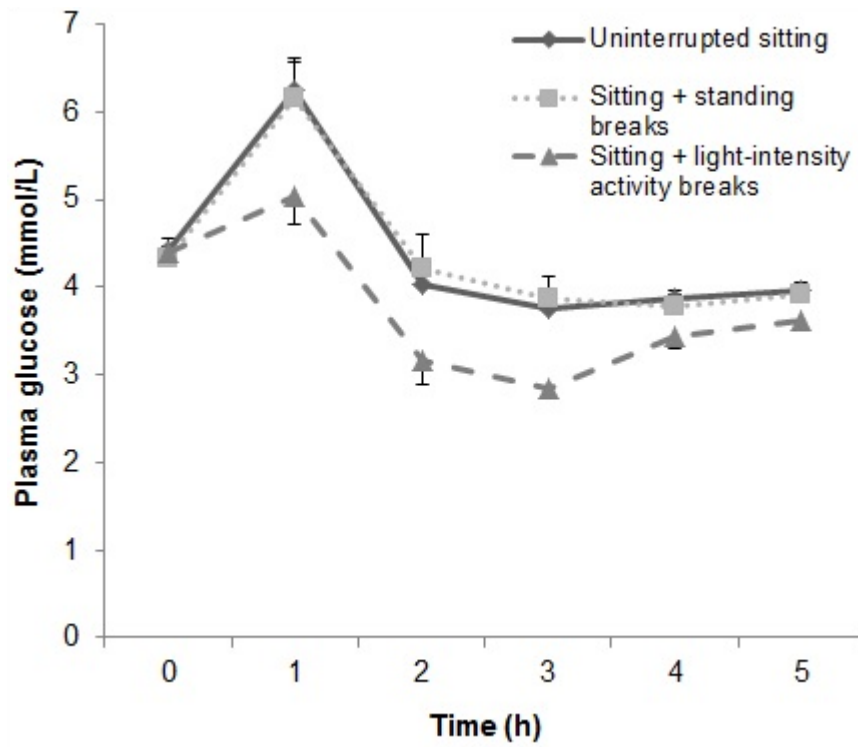
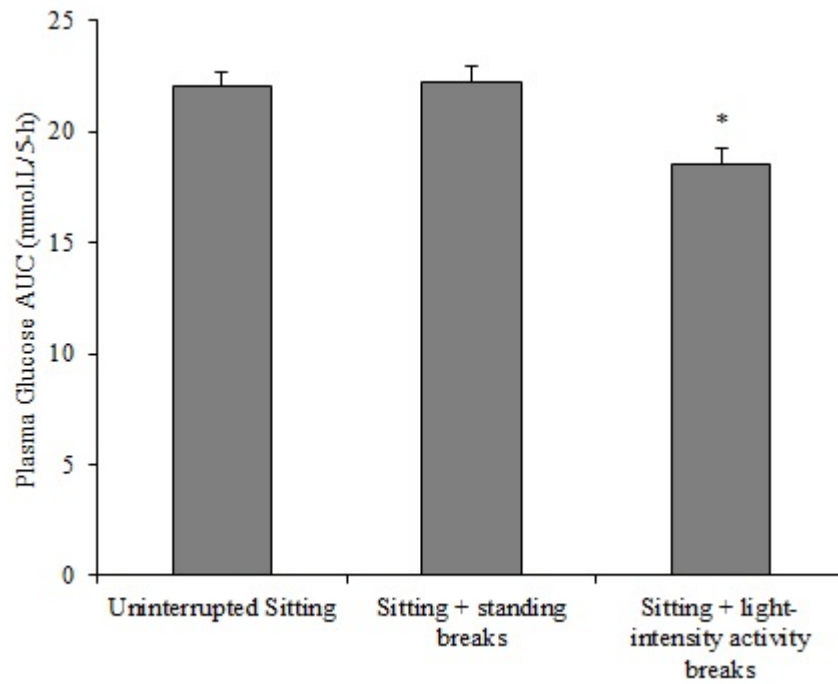


Figure 2: Postprandial plasma glucose area under curve (AUC) for the three trial conditions.

Data shown as mean and SE. *significantly different from sitting and sitting + standing breaks ($p < 0.001$).



Breaking up sitting time with physical activity

Appendix 1: The effect of the three trial conditions on 5-h systolic blood pressure levels.

Appendix 2: The effect of the three trial conditions on 5-h diastolic blood pressure levels