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1 **Physical Activity and Pediatric Obesity: a Quantile Regression Analysis**

2

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5 Accelerometry Database (ICAD) Collaborators.

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23

24 **Abstract**

25 **Purpose:** We aimed to determine if moderate-to-vigorous physical activity (MVPA) and
26 sedentary behavior (SB) were independently associated with body mass index (BMI) and waist
27 circumference (WC) in children and adolescents.

28 **Methods:** Data from the International Accelerometry Database (ICAD) were used to address our
29 objectives (N=11,115; 6-18y; 51% female). We calculated age and gender specific body mass
30 index (BMI) and waist circumference (WC) Z-scores and used accelerometry to estimate MVPA
31 and total SB. Self-reported television viewing was used as a measure of leisure time SB.
32 Quantile regression was the used to analyze the data.

33 **Results:** MVPA and total SB were associated with lower and higher BMI and WC Z-scores,
34 respectively. These associations were strongest at the higher percentiles of the Z-score
35 distributions. After including MVPA and total SB in the same model the MVPA associations
36 remained, but the SB associations were no longer present. For example, each additional hour per
37 day of MVPA was not associated BMI Z-score at the 10th percentile ($b=-0.02$, $P=0.170$), but was
38 associated with lower BMI Z-score at the 50th ($b=-0.19$, $P<0.001$) and 90th percentiles ($b=-0.41$,
39 $P<0.001$). More television viewing was associated with higher BMI and WC and the associations
40 were strongest at the higher percentiles of the Z-score distributions, with adjustment for MVPA
41 and total SB.

42 **Conclusions:** Our observation of stronger associations at the higher percentiles indicate that
43 increasing MVPA and decreasing television viewing at the population-level could shift the upper
44 tails of the BMI and WC frequency distributions to lower values, thereby lowering the number of
45 children and adolescents classified as obese.

46 **Key Words:** adolescent, accelerometry, ICAD, BMI, television

47

48 **Introduction**

49 Over 34% of adults and 16% of children in the U.S. are obese (29). These levels are
50 approximately 3-fold higher compared to the levels observed in the 1960s (16), and similar
51 increases in obesity prevalence have been observed globally (28). Obesity increases the risk of
52 the leading chronic diseases (38), tracks from childhood into adulthood (37), and obesity related
53 metabolic co-morbidities are increasingly being diagnosed in childhood (30). It is therefore
54 particularly important to determine contributors to child and adolescent obesity to address any
55 related short and long term health problems.

56

57 While the data are not conclusive (15), there is evidence that secular declines in moderate-to-
58 vigorous physical activity (MVPA), and increases in sedentary behavior (SB), coincided with the
59 rise in childhood obesity levels (3). It has therefore been hypothesized that increasing MVPA
60 and decreasing SB in childhood could help to reduce the prevalence of childhood obesity.

61 However, the epidemiological evidence relies heavily on self-reported measures of MVPA and
62 SB (22, 31), with fewer studies using accelerometry estimated MVPA and SB to study childhood
63 obesity. Importantly, different accelerometer methodologies were used across these latter studies
64 making it challenging to draw comparisons and conclusions (10, 17, 33). Furthermore, there is
65 little-to-no evidence that accelerometry estimated total SB is independently associated with
66 pediatric obesity (19, 23, 36); yet, television viewing, the most common leisure time SB has been
67 robustly associated with pediatric obesity (5, 26, 35). Additional research is therefore needed to
68 understand the complexity of physical activity energy expenditure, SB and pediatric obesity.

69

70 In addition, statistical methods that extend beyond the analysis of the mean of obesity-related
71 phenotypes are needed to build upon past studies (14). The upper percentiles of the distribution
72 for such phenotypes are of interest in the context of childhood obesity (e.g., 95th body mass index
73 [BMI] percentile). Quantile regression is a statistical method that can model the median and any
74 other percentile of a continuous variable, enabling investigators to test how predictors (e.g.
75 MVPA and SB) affect the shape of obesity-related phenotype distributions (e.g. BMI and waist
76 circumference [WC]) (4). Importantly, this method does not categorize participants into obese
77 and non-obese groups (e.g. <95th percentile versus ≥95th percentile), which has the following
78 drawbacks (1): i) assumes that all participants within a category are homogeneous (e.g., 10th
79 percentile similar to 90th percentile), ii) assumes participants at category boundaries are very
80 different rather than very similar (e.g. 94th percentile more similar to 10th percentile than the 95th
81 percentile).

82

83 To address the research gaps related to physical activity energy expenditure, SB, and pediatric
84 obesity, and the focus on modeling the mean of obesity phenotypes, we analyzed standardized
85 data from the International Children's Accelerometry Database (ICAD), using quantile
86 regression. We specifically aimed to determine if time spent in MVPA and SB (total and leisure
87 time) were independently associated with BMI and WC Z-score frequency distributions in
88 children and adolescents.

89

90 **Methods**

91 *Study Design and Data Source*

92 The ICAD consortium pooled raw accelerometry data from international studies involving
93 children and adolescents, and used standardized methods to derive estimates of MVPA and SB
94 (33). Specifically, accelerometry data from 21 studies that used an ActiGraph accelerometer
95 (models: AM7164 and GT1M) were pooled, along with anthropometric and demographic data
96 when available. These studies were conducted between 1998 to 2009 in Europe, the U.S., Brazil
97 and Australia. For the purpose of our study, we included 6 to 18 year olds with complete
98 accelerometry, anthropometric and covariate data in the ICAD database (N=11,115). These
99 participants were originally enrolled in the following 8 studies: Avon Longitudinal Study of
100 Parents and Children (ALSPAC, Children's of the 90s Cohort, N=4,281) (8), Denmark European
101 Youth Heart Study (EYHS, N=1,108), Estonia EYHS (N=437), National Health and Nutrition
102 Examination Survey (NHANES) 2003-2004 (N=1,915), NHANES 2005-2006 (N=2,097),
103 Norway EYHS (N=286), Personal and Environmental Associations with Children's Health
104 (PEACH, N=640) and Portugal EYHS (N=351). All partners contributing data completed data-
105 sharing agreements and institutional review boards at each contributor's institution approved the
106 sharing of the data (33).

107

108 *Accelerometer Estimated MVPA and SB*

109 The majority of waking hours are spent in SB (20, 25). SB includes any waking behavior
110 characterized by low energy expenditure (1.0-1.5 metabolic equivalents [METs]), while in a
111 sitting or reclining posture (32). In contrast, a small proportion of waking hours are spent in
112 MVPA (39). In children, MVPA includes any physical activity that requires an energy
113 expenditure that is 4 times greater than the energy expended at rest (≥ 4 METs) (31).
114 Accelerometry can be used to provide estimates of free-living MVPA and SB. In our study,

115 standardized methods were used to derive estimates of MVPA and SB from the raw
116 accelerometer files, using commercially available software: KineSoft, version 3.3.20 (KineSoft,
117 Saskatchewan, Canada; www.kinesoft.org). All data files were integrated to a 1-minute epoch
118 and 60 minutes of consecutive zero counts were considered non-wear time (allowing for 2
119 minutes of non-zero interruptions). Participants providing at least 1 day of count data, for 10 to
120 20 hours, were included. Time spent in total SB was the average minutes per day $<100\text{cpm}$ and
121 time spent in MVPA was the average minutes per day $\geq 2,296$ (40). These SB and MVPA
122 cutpoints have been shown to have high sensitivity and specificity (40).

123

124 *Anthropometrics*

125 BMI correlates highly with total fat mass and WC correlates highly with visceral fat mass (2). In
126 our study, height and weight were measured by trained research staff, which allowed for the
127 calculation of BMI's (kg/m^2). WC (cm) was measured using metal anthropometric tapes, mid-
128 way between the lower rib margin and the iliac crest, at the end of a gentle expiration (in the
129 NHANES sample, the metal tape was placed just above the iliac crest at the midaxillary line).
130 Since BMI and WC increase with normal growth and physical development, age and sex specific
131 Z-scores were calculated, using the sample means and standard deviations from 8 ICAD studies,
132 and the Z-scores were included as outcomes in our study. We excluded those with BMI <12
133 kg/m^2 , $>50 \text{ kg}/\text{m}^2$ and BMI Z-scores >3.5 . Similarly, we excluded those with WC <40 cm or
134 >120 cm, and WC Z-scores >3.5 .

135

136 *Covariates*

137 As children age, MVPA declines, SB increases and the likelihood of being obese increases (25,
138 27); we therefore included age in years as a covariate. Also, males tend to accumulate more
139 MVPA, less SB and are less likely to obese compared to females (20, 27); we therefore included
140 sex as a covariate. Similarly, white children engage in more MVPA, less SB and are less likely to
141 be obese compared to their non-white counterparts (20); we therefore included race (white or
142 non-white) as a covariate. Children belonging to lower income households spend less time in
143 MVPA, more time in SB and are more likely to be obese compared to children belonging to
144 higher income households (20, 29); we therefore included household income quartile as a
145 covariate. Finally, we included study year as a covariate to control for any secular changes in the
146 outcomes and predictors from 1998 to 2009, and study as a covariate to control for any cultural
147 differences across the international studies.

148

149 *Subsample with screen-based SB*

150 Television viewing is the most common leisure-time SB (6), so we additionally tested for
151 association between this particular SB and BMI Z-score and WC Z-score. The participants were
152 categorized into the following groups based on their reported television viewing: <1 h/d, 1-2 h/d,
153 3-4 h/d or >4 h/d. Only a subsample of the participants included in our study had television
154 viewing data (N=6,712), from the following studies: Denmark European Youth Heart Study
155 (EYHS, N=1,091), Estonia EYHS (N=431), National Health and Nutrition Examination Survey
156 (NHANES) 2003-2004 (N=1,914), NHANES 2005-2006 (N=2,096), Norway EYHS (N=272),
157 Personal and Environmental Associations with Children's Health (PEACH, N=637) and Portugal
158 EYHS (N=271).

159

160 *Statistical Analysis*

161 To describe our sample, we presented means and standard deviations for the continuous variables
162 and frequencies and percentages for the categorical variables. We used the International Obesity
163 Task Force age- and gender-specific cutoffs to describe the proportion of normal, overweight and
164 obese participants (11). To address the aims of our study we used quantile regression, modeling
165 BMI Z-score and WC Z-score as the dependent variables. This statistical method models the
166 median of outcome variables (rather than the mean) and any other percentile across the
167 frequency distribution without the need to categorize participants. We first tested if MVPA (hour
168 per day, hr/d) was associated with BMI Z-score and WC Z-score, adjusting for the covariates
169 (model 1a). We then tested if total SB (hr/d) was associated with BMI Z score and WC Z-score,
170 adjusting for the covariates (model 1b). Thereafter, we included both MVPA and total SB in the
171 same model (model 2) to determine if the MVPA and total SB associations were independent of
172 one another. We tested for associations at the following BMI Z-score and WC Z-score
173 percentiles, to capture the lower, mid and upper parts of the frequency distributions: 5th, 10th,
174 15th, ..., 85th, 90th, and 95th percentiles. We repeated model 2 with the addition of television
175 viewing as the main predictor for the sub-sample. All statistical analyses were completed using
176 Stata 12.1 (StataCorp LP, College Station, TX) using the *sqreg* command with 100 bootstrap
177 samples to calculate standard errors. The *grqreg* command was used to graph key findings. We
178 also performed sensitivity analyses using UK and US population reference data to calculate BMI
179 Z-scores (12, 18); and stratified the analyses by sex, race and household income categories. We
180 also performed sensitivity analyses to re-examine the WC Z-score association with the NHANES
181 data removed, given the different WC measurement approached used in these studies.

182

183

184 **Results**

185 Our primary analytical sample included 11,115 participants aged 6 to 18 years; 51% were
186 female, 71% were white and 6% were obese (Table 1). The sample included approximately equal
187 proportions of children belonging to low, middle and high-income households. The average time
188 spent in MVPA was 55 minutes per day and the average time spent in total SB was 6.4 hours per
189 day. The percent of participants providing 3 or more days of accelerometry data was 91%. The
190 characteristics of the sub-sample (N=6,712) with television viewing data were similar to the full
191 sample; however, a large majority of those with missing television viewing data were white. The
192 participants most commonly reported watching television for 3-4 hours per day.

193

194 *MVPA and BMI/WC Z-scores*

195 More time spent in MVPA was associated with lower BMI and WC Z-scores, and these
196 associations were strongest at the upper percentiles of the Z-score distributions (Table 2). For
197 example, at the 10th percentiles each additional hour of MVPA was associated with a 0.01
198 ($P=0.248$) lower BMI Z-score and with a 0.04 ($P=0.018$) lower WC Z-score (Table 2, models
199 1a); whereas, at the 90th percentiles each additional hour of MVPA was associated with a 0.35
200 ($P<0.001$) lower BMI Z-score and with a 0.36 ($P<0.001$) lower WC Z-score (Table 2, models
201 1a). These associations remained after adjustment for total SB for BMI Z-score (10th percentile
202 $\beta=-0.02$, $P=0.167$; 90th percentile $\beta=-0.41$, $P<0.001$) and WC Z-score (10th percentile
203 $\beta=-0.02$, $P=0.329$; 90th percentile $\beta=-0.36$, $P<0.001$) (Table 2, models 2). The 90th
204 percentile beta coefficients were stronger than the 10th percentile coefficients for BMI Z-score
205 (beta difference: 0.39; 95% confidence interval (CI): 0.31-0.48; P -value: <0.001) and WC Z-

206 score (beta difference: 0.34; 95% CI: 0.24-0.45; P -value: <0.001). Visual representation of the
207 MVPA association with WC Z-score is presented in Figure 1, and the linear regression
208 coefficients for MVPA and the mean WC Z-score (model 1a: $\beta=-0.27$, $P<0.001$ and model 2: $\beta=-$
209 0.26 , $P<0.001$) are plotted for comparison.

210

211 *Total SB and BMI/WC Z-scores*

212 More time spent in total SB was associated with higher BMI and WC Z-scores, and these
213 associations were strongest at the upper percentiles of the Z-score distributions (Table 2). For
214 example, at the 50th percentiles each additional hour of SB was associated with a 0.03 ($P<0.001$)
215 higher BMI Z-score and with a 0.04 ($P<0.001$) higher WC Z-score (Table 2, models 1b);
216 whereas, at the 90th percentiles each additional hour of SB was associated with a 0.05 ($P=$
217 $P=0.003$) higher BMI Z-score and with a 0.08 ($P<0.001$) higher WC Z-score (Table 2, models
218 1b). However, these associations did not hold after adjustment for MVPA (Table 2, models 2).
219 Visual representation of the total SB association with WC Z-score is presented in Figure 1, and
220 the linear regression coefficients for total SB and the mean WC Z-score (model 1b: $\beta=0.06$,
221 $P<0.001$; model 2: $\beta=0.01$, $P=0.504$) are plotted for comparison.

222

223 *Television Viewing and BMI/WC Z-scores*

224 In the subsample with complete television viewing data, more time spent watching television
225 was associated with higher BMI and WC Z-scores, and these associations were strongest at the
226 upper percentiles of the Z-score distributions, and remained after adjustment for total SB and for
227 MVPA (Figure 2 and Supplementary Table 1). For example, at the 10th percentiles, relative to <1
228 hour per day of television viewing, 3-4 hours per day was associated with a 0.08 ($P=0.005$)

229 higher BMI Z-score and with a 0.08 ($P=0.022$) higher WC Z-score; whereas, at the 90th
230 percentiles, relative to <1 hour per day of television viewing, 3-4 hours per day was associated
231 with a 0.34 ($P<0.001$) higher BMI Z-score and with a 0.35 ($P<0.001$) higher WC Z-score.

232

233 *Sensitivity Analyses*

234 We repeated our analysis of accelerometry estimated MVPA and total SB using BMI Z-scores
235 calculated using reference population data from the UK and the US. The MVPA and total SB
236 associations remained similar to our findings that used internal means and standard deviations to
237 calculate BMI Z-scores (Supplementary Table 2). We also repeated our analysis by sex, race and
238 household income categories. The MVPA and total SB associations with BMI and WC Z-scores
239 remained similar within each group (Supplementary Table 3). The WC Z-score associations
240 remained similar with the NHANES data removed (Supplementary Table 4).

241

242 **Discussion**

243 This is the largest study to date to use standardized accelerometry methods and quantile
244 regression to study the association between physical activity energy expenditure and childhood
245 obesity. We found that more time spent in MVPA was independently associated with lower BMI
246 and WC. In contrast, total SB was not independently associated with BMI or WC in our sample
247 of children and adolescents. However, time spent in a specific leisure time SB, television
248 viewing, was independently associated with higher BMI and WC. Interestingly, our quantile
249 regression analysis revealed that the associations between MVPA, television viewing and
250 BMI/WC Z-scores were non-linear; these exposures were most influential at the upper
251 percentiles of the BMI and WC frequency distributions. These non-linear associations indicate

252 that if more children were to meet the physical activity guidelines, and limit their television
253 viewing hours, then the prevalence of childhood obesity could be reduced by specifically shifting
254 the upper percentiles of the BMI and WC frequency distributions to lower values.

255

256 We have shown in previous U.S. based studies that lower accelerometry estimated MVPA and
257 higher television viewing are associated with greater increases in BMI over time during
258 childhood, and that these associations were strongest at the upper BMI percentiles (24, 26).

259 These findings are consistent with our present results and together support that increasing MVPA
260 and lowering television viewing could help to prevent childhood obesity, potentially by
261 increasing energy expenditure and correcting energy imbalance. However, there are contrasting
262 data suggesting that increases in weight status lead to lower MVPA and higher SB (21). We
263 cannot rule out reverse-causality in our cross-sectional study. We also cannot rule out that a
264 cyclical (bi-directional) relationship exists between physical activity energy expenditure and
265 weight status. Indeed, the stronger association between MVPA and television viewing at the
266 upper percentiles of the BMI/WC distributions could reflect such a cyclical relationship.

267

268 Alternatively, the stronger associations at the upper percentiles could be explained by gene-
269 environment interactions. It is well known that BMI and WC are heritable traits and a large
270 number of genetic loci are associated with higher BMI and WC (9). Specifically, individuals at
271 the upper percentiles of the BMI and WC distributions are more likely to be genetically
272 predisposed to obesity (26). If these individuals are exposed to lower MVPA or higher television
273 viewing, then greater increases in their BMI/WC could be due to their genetic susceptibility to

274 obesity, compared to those who are less likely to be genetically predisposed to higher BMI/WC
275 at the lower percentiles of the distribution.

276

277 It has been proposed that SB, independent of time spent in MVPA, is an obesity risk factor.
278 Studies that have estimated total SB using accelerometry have observed associations with
279 measures of childhood obesity (10, 17, 19, 23, 24, 36), but such associations have tended not to
280 hold after adjustment for MVPA (17, 19, 23, 36). We observed the same pattern in our study,
281 suggesting that total SB is not an independent risk factor for childhood obesity. In contrast, there
282 is convincing evidence that television viewing is associated with childhood obesity, independent
283 of MVPA (13, 26, 35). Television viewing is the most common leisure time SB, but it is also a
284 behavior associated with snacking and exposure to food advertisements (34). Therefore, the
285 television viewing associations may not be fully explained by the SB-energy expenditure
286 paradigm and could be partly explained by increases in energy intake. In addition, television
287 viewing is the most common activity before going to bed (7), and our television viewing
288 associations could be partly explained by reductions in total sleep time (26).

289

290 The strengths of our study include the large sample size and the standardized methods to
291 estimate MVPA and total SB using accelerometry. Given the large sample size, we were able to
292 perform sensitivity analyses and replicated our findings in key sub-groups. We used quantile
293 regression and the advantages of this analytical approach are increasingly being recognized for
294 investigating continuous variables in epidemiological studies (4). By using this method we
295 observed stronger associations between MVPA, total SB (not independent of MVPA) and
296 television viewing at the upper percentiles of the BMI and WC Z-score distributions; these

297 patterns of association across the percentiles would have been missed had we modeled the mean
298 Z-scores using linear regression. We used both BMI and WC as primary outcomes to make
299 inferences on overall fat mass and visceral fat mass, and the latter is of particular clinical
300 importance (2). Our study does have limitations. We adjusted for key confounders, but the ICAD
301 database does not have dietary or sleep data to include as covariates. Also, given the populations
302 of origin for this study a dichotomous race variable was used; it will be important to replicate our
303 findings in more diverse populations. The majority (91%) of participants provide 3 or more days
304 of valid accelerometry data. We included participants who provided 1 and 2 valid days
305 accelerometry data and the MVPA and total SB estimates may not be representative of habitual
306 patterns among these participants. We used a cross-sectional design and so were not able to
307 establish the temporality between our exposures and Z-score outcomes.

308

309 Preventing childhood obesity has the greatest potential to counter the short and long-term health
310 problems associated with obesity. We conclude that increasing MVPA and decreasing television
311 viewing in childhood could help to prevent obesity in early life. By using quantile regression we
312 showed that increasing the time children spend in MVPA and decreasing the time they spend
313 watching television could help to lower BMI's and WC's, especially for children in the
314 population with the higher BMIs and WCs for their age.

315

316

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318

319

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380

381 **Conflicts of Interest**

382 The results of the present study do not constitute endorsement by ACSM

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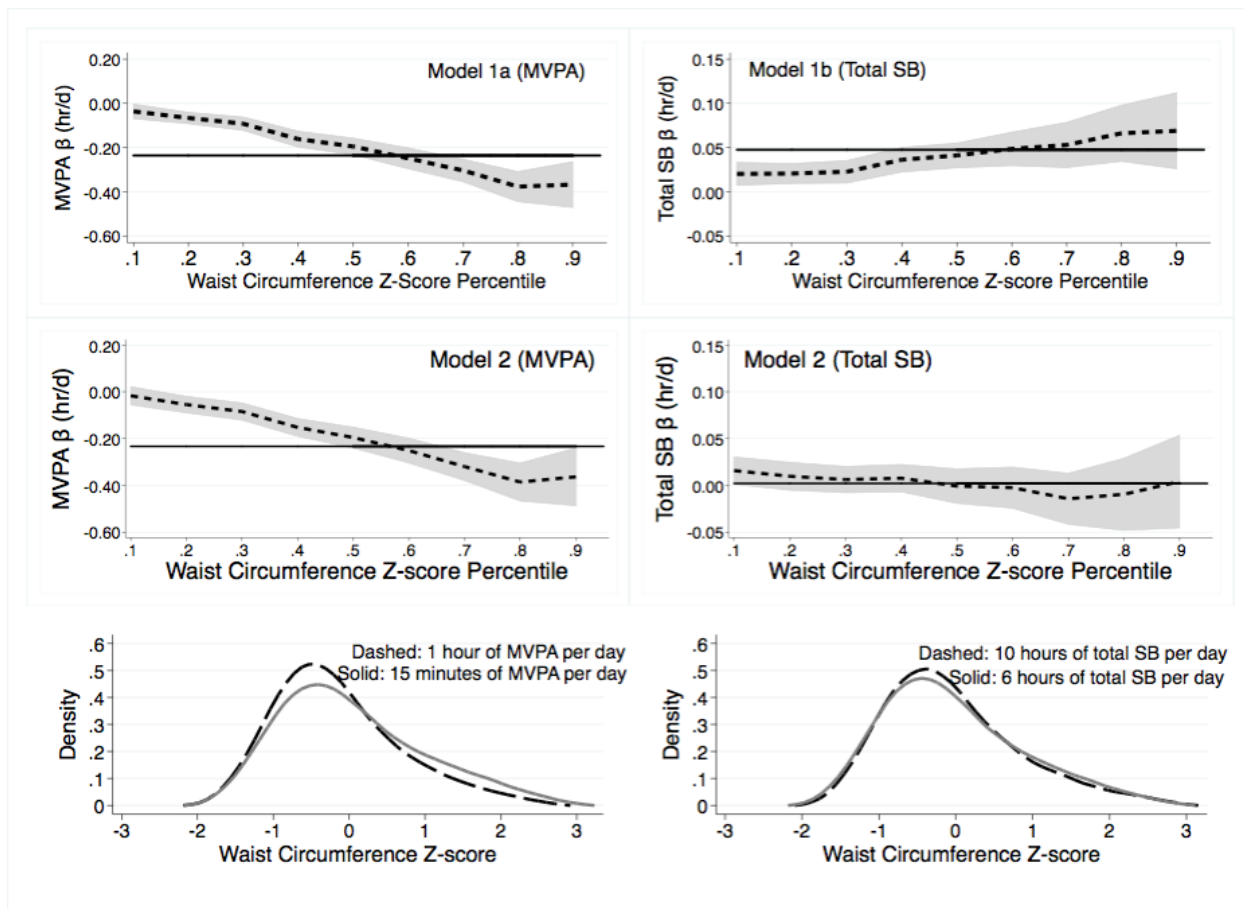
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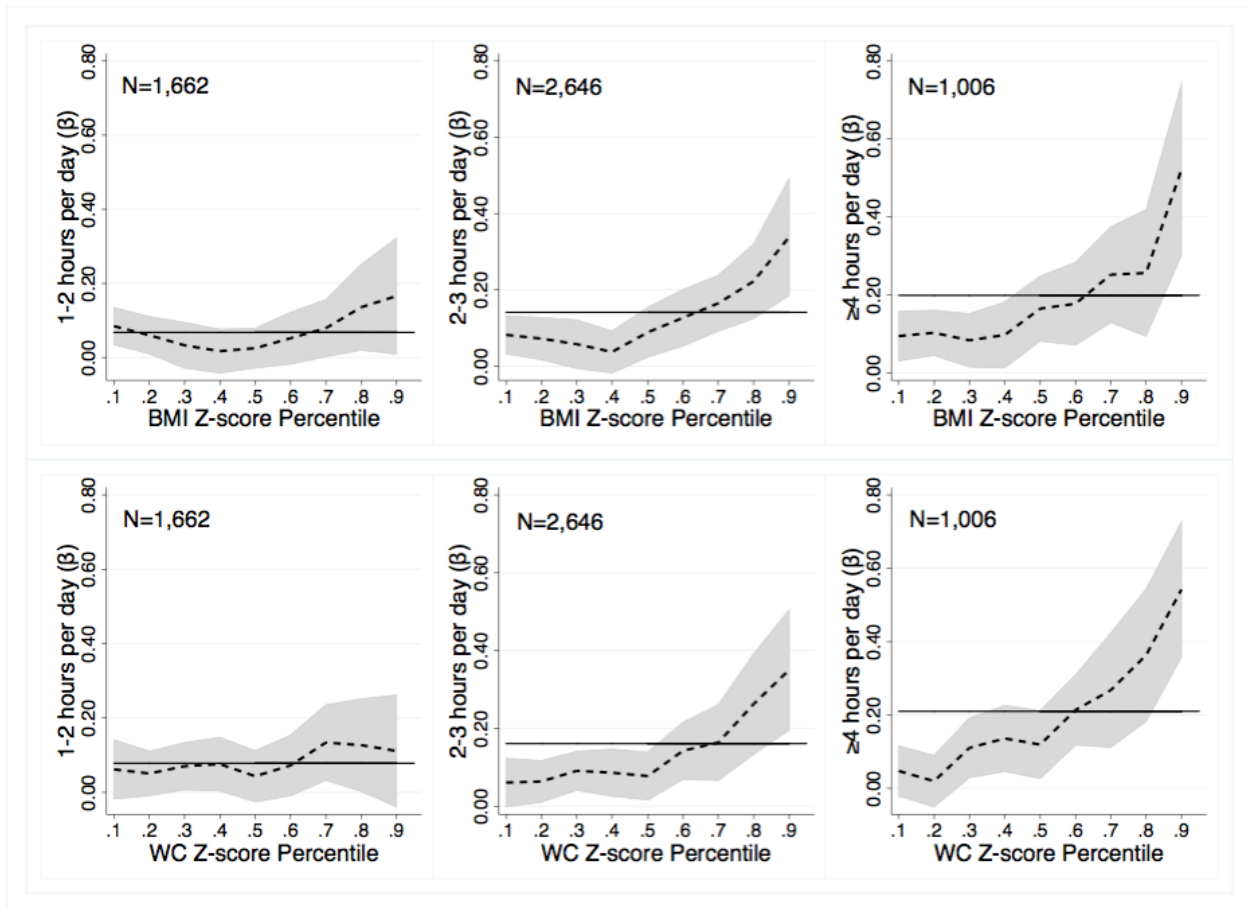
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503 **Figure 1.** Illustration of the quantile regression associations between moderate-to-vigorous
504 physical activity (MVPA, left column), total sedentary behavior (SB, right column) and waist
505 circumference (WC) Z-scores. The quantile regression findings from model 1a and model 1b are
506 presented in the top row; whereas the quantile regression findings from model 2 are present in
507 the second row. MVPA is the primary predictor in model 1a and total SB is the primary
508 predictor in model 1b; both models are adjusted for age, race, household income and
509 accelerometer wear time. In model 2, MVPA and total SB are included as predictors in the same
510 model along with age, race, household income and accelerometer wear time. For comparison,
511 the horizontal black lines represent the linear regression coefficients for the change in mean WC
512 Z-score per additional hour per day of MVPA or total SB. The distribution plots in the bottom
513 row are derived from model 2.



514

515 **Figure 2.** Television viewing associations with body mass index (BMI) Z-scores and waist
 516 circumference (WC) Z-scores using quantile regression. The <1 hour per day television viewing
 517 category is the referent group. The quantile regression models are adjusted for age, race,
 518 household income, moderate-to-vigorous physical activity, total sedentary behavior and
 519 accelerometer wear time. The horizontal black lines represent the linear regression coefficients
 520 for the change in mean BMI Z-scores and WC Z-scores for each television viewing category.

521

522 **Supplemental Digital Content**

523 Supplementary Table 1

524 Supplementary Table 2

525 Supplementary Table 3