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1	Associations of body mass index, physical activity and sedentary time with blood pressure in
2	primary school children from south-west England: a prospective study
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26 Abstract

Elevated blood pressure in children is a significant risk factor for the development of 27 cardiovascular disease in adulthood. We examined how children's body mass index (BMI), 28 physical activity and sedentary time at ages 9 and 11 are associated with blood pressure at age 29 30 11. Data were from 1283 children from Bristol, UK, who participated in the study aged 11 31 years, 797 of whom also participated in the study aged 9 years. Child height, weight and blood pressure were measured, and children wore accelerometers for five days, from which moderate-32 33 to-vigorous-intensity physical activity and sedentary minutes per day were derived. Multiple imputation of missing data and adjusted linear and logistic regression models were used to 34 35 examine associations. Child BMI at 11 years was cross-sectionally associated with higher systolic and diastolic blood pressure (mean difference [95% confidence interval]: 0.91 [0.32 to 36 37 1.50] mm Hg and 1.08 [0.54 to 1.62] mm Hg, respectively, per standard deviation (SD) of 38 BMI). BMI at age 9 was also positively associated with diastolic blood pressure at age 11 (1.16 mmHg per two years [0.49 to 1.84], per SD of BMI). For girls, sedentary time at age 9 years 39 40 was associated with increased odds of having high systolic blood pressure at age 11 (odds ratio: 1.08 [1.01 to 1.16], per 10 minutes per day). There was no evidence of associations between 41 sedentary time and blood pressure among boys. Similarly, there was little evidence that 42 43 physical activity was associated with blood pressure in either cross-sectional or prospective analyses. Effective strategies are needed to prevent excess bodyweight among children in order 44 to reduce cardiovascular disease risk. 45

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51 Introduction

52 Elevated blood pressure in children is a significant risk factor for the development of cardiovascular disease in adulthood. Increasing numbers of children and young people are 53 being diagnosed with hypertension [1-3], with global prevalence rates of at least 3% among 54 55 asymptomatic children and adolescents [2]. Higher childhood blood pressure tracks through to 56 adulthood [4] and is positively associated with the development of cardiovascular disease later in life and risk of premature mortality [5-10]. Globally, children are also more likely to be 57 58 categorised as overweight or obese than ever before [11, 12]. In 2017-2018, approximately one third of eleven-year-olds in England were categorised as overweight, while one fifth were 59 60 categorised as obese [11]. Children whose overweight and obesity persist into adulthood are at an increased risk of hypertension, type 2 diabetes, dyslipidemia, and carotid-artery 61 62 atherosclerosis compared to children who are normal weight [13]. Available evidence has 63 generally demonstrated strong positive associations between body mass index (BMI) and blood 64 pressure in children and adolescents [14-24]. However, the majority of studies were cross-65 sectional [15-19] or examined prospective associations with blood pressure measured at only a single time-point [21-24]. Prospective studies with blood pressure measures at baseline and 66 67 follow-up, therefore enabling adjustment for baseline blood pressure, would be valuable to gain 68 a better understanding of whether higher BMI is a risk factor for higher blood pressure in children. 69

Trend data from the UK suggests that BMI explained ~15% of the increases in systolic blood pressure between 1980-2008, with associations weakening over time [14]. Thus, there is value in examining other factors that may be associated with the rising levels of blood pressure in children in recent years. Physical inactivity is positively associated with higher blood pressure and cardiovascular disease in adulthood [25, 26], and regular physical activity has the potential to lower blood pressure and reduce BMI among adults [26-28]. 76 There is a lack of information about the association between physical activity and blood 77 pressure in children. Several studies have reported inverse associations between physical activity and systolic and/or diastolic blood pressure, with most showing this association is 78 independent of BMI [29-40]. It is proposed that physical activity produces more favourable 79 80 vascular health profiles among children [38, 41]. In a review of the literature published in 2007 [20], it was suggested that engaging in 40 minutes of moderate-to-vigorous-intensity physical 81 82 activity (MVPA) on 3-5 days per week was required to improve vascular function and reduce blood pressure in obese children. However, other studies have reported no association between 83 84 physical activity and blood pressure in children [24, 42-45]. The majority of studies to date 85 were cross-sectional and used child or parental report of physical activity [31, 33, 36, 39, 41, 43, 45], which may be subject to misclassification, particularly in relation to physical activity 86 87 intensity and duration. We previously published cross-sectional analyses using baseline data 88 from the present study, finding no association between physical activity and sedentary time with blood pressure in 9-year-old children [24]. Therefore, there is a need for more prospective 89 90 studies with measures of objectively-assessed physical activity and sedentary time to further 91 understand key determinants of variation in child blood pressure.

In this study, we investigated the cross-sectional and prospective associations of BMI
and accelerometer-assessed physical activity and sedentary time measured at age 9 and 11 with
blood pressure at age 11.

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96 Methods

Data used in the present study were from B-PROACT1V, a longitudinal study
examining physical activity and sedentary behaviours of children and parents as children
progress through primary school. The study has been described in detail elsewhere [46-49].
Briefly, in 2012-2013, data were collected from 1299 Year 1 children (median age: 6 years)

101 who were recruited from 57 schools in Bristol, UK, and the surrounding area. In 2015-2016, 102 data were collected from 1223 Year 4 children (median age: 9 years) from 47 of the original schools. In 2017-2018, 50 of the original schools were re-recruited and data were collected 103 104 from 1296 Year 6 children (median age: 11 years). In total, 2132 children participated, of whom 105 958 participated at one time-point, 662 at two time-points, and 512 at three time-points. The current study used data from the 1283 children who provided at least two blood pressure 106 107 measurements at age 11, since blood pressure was only measured at age 9 and 11, and the age 108 9 blood pressure data have been presented previously [24]. This included 797 children who 109 also participated in the study at age 9 years. Ethical approval for the study was granted from 110 the School for Policy Studies Research Ethics Committee at the University of Bristol, and written parent consent was provided for both child and parent participation. 111

Blood pressure

113 Blood pressure was measured in the child's school by trained fieldworkers using an 114 Omron 907 Professional Blood Pressure Monitor [50] with a small or medium cuff (OMRON 115 Corporation, Kyoto, Japan). After three minutes of rest, measurements were taken, using the 116 appropriate cuff size, three times, one-minute apart, in the left arm while the child was seated. 117 Of the 1296 children who participated at age 11, we have included 1283 (99%) children who provided at least two blood pressure measurements at age 11, in the current study. The mean 118 119 of all available blood pressure measurements was used in the analyses. In our main analyses 120 we examined associations with difference in mean blood pressure as a continuous variable. We also examined associations with high systolic and diastolic blood pressure, defined as the 121 systolic or diastolic blood pressure that was $\geq 95^{\text{th}}$ percentile using US children's age, sex and 122 height standardized blood pressure references [51]. US reference charts were used because of 123 the lack of such population references for UK children. 124

125 Body mass index

126 Children's weight and height were measured at the children's schools by trained 127 fieldworkers. Weight was measured using a SECA 899 digital scale to the nearest 0.1kg, and 128 height was measured using a SECA Leicester stadiometer to the nearest 0.1cm (HAB 129 International, Northampton). BMI was derived as weight (kg)/height (m²) and converted to an 130 age- and sex-specific standard deviation score based on UK 1990 growth centiles [52].

131 Accelerometer-assessed physical activity

MVPA and sedentary time of children were measured using ActiGraph wGT3X-BT 132 133 (Pensacola, FL, USA) accelerometers. Children were asked to wear an accelerometer on their 134 waist during all hours they were awake for five consecutive days, including weekend days. 135 Accelerometer data were processed using Kinesoft (v3.3.75; Kinesoft, Saskatchewan, Canada). 136 At least three valid days of data were required for inclusion in analysis, and a valid day was defined as \geq 500 minutes of data, after excluding intervals of \geq 60 minutes of zero counts (for 137 the latter allowing up to two minutes of interruptions during the 60 minutes). The Evenson [53] 138 population-specific cut points for children were used to derive the average number of MVPA 139 140 and sedentary minutes per day.

141 **Observed confounders**

Of the data available in this study we considered child's sex, age and height, household 142 socioeconomic position and parental history of high blood pressure as key confounders given 143 their known influences on BMI, physical activity, sedentary behaviour and blood pressure. 144 145 Dietary factors are likely to influence blood pressure, but we did not have information on this. Parental behaviours, such as smoking and alcohol consumption, may be mimicked by their 146 147 children and hence they (or these behaviours in the child) may also confound the associations we are exploring. We considered that at age 9-11 years very few children would be smoking 148 or drinking alcohol to an extent that would influence their blood pressure, and so did not 149 150 consider these further. At least one of the children's parents were recruited to the study. Parents 151 completed a questionnaire requesting information about their child's sex and date of birth. 152 Where the child's date of birth was missing (8.4% of children), the median age was assigned (11.0 years at Year 6). While replacing missing data with average sample values can introduce 153 154 bias, as children were from a single school year we felt this was appropriate. Parents were also asked if either of the child's biological parents had ever been informed that they had high blood 155 pressure. As indicators of socioeconomic position, parents were asked to report the highest 156 157 level of education in the home, with the following response options: 'up to GCSEs/ GCEs/ O Levels or equivalent' (qualifications usually obtained in several subjects at age 16, the 158 minimum legal education leaving age in the UK up to 2015), 'A Levels/ NVQs/ GNVQs' 159 (qualifications usually obtained at age 18), 'Degree/ Diploma/ HNC/ HND or equivalent' and 160 'Postgraduate degree (MSc, PhD)'. This was combined across time-points to produce a single 161 162 indicator of highest household education.

163 Statistical analysis

164 Means and proportions were used to examine the characteristics of the cross-sectional and prospective samples. Linear regression models were used to examine the cross-sectional 165 166 associations between the child's BMI z-score, MVPA and sedentary time and systolic and 167 diastolic blood pressure at age 11. Linear regression models were also used to examine the prospective associations between BMI, MVPA and sedentary time at age 9 and systolic and 168 diastolic blood pressure at age 11. Model 1 was unadjusted and in the prospective models, we 169 170 adjusted for baseline (age 9 years) levels of blood pressure (systolic or diastolic respectively). 171 In Model 2, we additionally adjusted for highest household education, child age and height, 172 child sex (for models with all children), parent reported high blood pressure, BMI z-score (for activity measures), and accelerometer wear time (for activity measures). Due to a previous 173 study that used B-PROACT1V data finding a negative association between MVPA and BMI 174 175 as children age [50], the models with activity measures were adjusted for BMI z-score to

176 examine the independent effect of physical activity on blood pressure. Logistic regression 177 models were used to explore cross-sectional and prospective associations with odds of having high systolic and diastolic blood pressure. We undertook all analyses with girls and boys 178 179 combined and explored differences between them by running analyses separately and exploring 180 evidence for statistical interaction between sex and each exposure. To account for children being recruited via schools, robust standard errors, which took account of the school-level 181 182 clustering, were used for all models. All analyses were performed using Stata version 15.0 (StataCorp, 2015). 183

184 **Dealing with missing data**

Among the 1283 children who were eligible for inclusion in the cross-sectional and the 185 797 eligible for the prospective analyses, there were small amounts of missing data for risk 186 187 factors, and/or confounders (Table 1). This varied from 0 (e.g., for child blood pressure at age 11, child age and sex) to 17.2% (for parental high blood pressure reported at age 11) in the 188 189 cross-sectional analyses and 14.8% (for parental high blood pressure reported at age 9) in the 190 prospective analyses. To enable us to include information from all study participants in our 191 analysis, and thus potentially increase statistical power and minimise selection bias, we used 192 multiple imputation of missing data using chained equations. Imputation was completed separately for cross-sectional and prospective analyses. Thus, for the cross-sectional analyses 193 194 we imputed data for the 1283 children who participated at age 11 and provided at least two 195 blood pressure measurements. For prospective analyses, which examined associations between 196 BMI, physical activity or sedentary time at age 9 with blood pressure at age 11, we imputed to 197 the 797 children who took part at both time points.

All child accelerometer measures, measurements of blood pressure, and characteristics that were potential predictors of missingness (child age, sex, BMI, highest household education, parental high blood pressure, and the child's school) at either year, were included

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in the multiple imputation models. Children's classification of high systolic and diastolic blood
pressure were imputed passively. Twenty imputed datasets were created using 20 cycles of
regression switching and combined regression coefficients across imputed datasets using
Rubin's rules [54].

Regression analyses were repeated restricting to children who had complete data on all exposures, outcomes and covariables in cross-sectional and prospective analyses (N= 1025 and 655 respectively). The results were very similar between the main analyses with multiple imputed datasets and the complete case analyses for both cross-sectional and prospective analyses and, therefore, the results of the complete case analyses are not presented but are available from the authors on request.

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212 **Results**

The characteristics of children who provided at least two measures of blood pressure at age 11, and the subset who additionally took part at age 9, in the observed and imputed datasets are shown in Table 1. The subset that took part in both years were comparable to the whole age 11 sample, and in both years the distributions of all characteristics were very similar in imputed and observed data. The mean systolic blood pressure for 11-year-old children was 104.65 mm Hg (7.8% had high systolic blood pressure) and mean diastolic blood pressure was 68.70 mm Hg (10.7% had high diastolic blood pressure).

Table 1. Characteristics of the children who participated in the study at age 11 years, and those who participated at both age 9 and age 11 years in the observed and multiple imputation data

	Children who participated at age 11				Children who participated at ages 9 and 11			
		Observed data	Imputed (N=1283)	Observed data		Imputed (N=797)		
Characteristic	Ν	Mean (SD) or %	Mean (SD) or %	Ν	Mean (SD) or %	Mean (SD) or %		
Systolic blood pressure at age 11 (mm Hg)	1283	104.65 (10.91)	104.65 (10.91)	797	104.60 (10.96)	104.60 (10.96)		
Diastolic blood pressure at age 11 (mm Hg)	1283	68.70 (9.52)	68.70 (9.52)	797	68.72 (9.59)	68.72 (9.59)		
High systolic blood pressure at age 11	1283			797				
No		92.2%	92.2%		92.3%	92.3%		
Yes		7.8%	7.8%		7.7%	7.7%		
High diastolic blood pressure at age 11	1283			797				
No		89.3%	89.3%		89.1%	89.1%		
Yes		10.7%	10.7%		10.9%	10.9%		
Body mass index (z-score) at age 11	1279	0.35 (1.16)	0.35 (1.16)	795	0.32 (1.18)	0.32 (1.18)		
MVPA (mins/day) at age 11	1229	58.13 (22.52)	58.11 (22.56)	772	57.94 (22.38)	57.84 (22.47)		
Sedentary (mins/day) at age 11	1229	464.40 (68.66)	464.49 (68.86)	772	462.73 (66.07)	462.79 (66.28)		
Highest household education	1180			773				
Up to GCSE/O level		20.5%	20.8%		19.0%	19.1%		
A level/ NVQ		26.2%	26.3%		26.0%	26.0%		
Degree/ HND		36.6%	36.3%		37.8%	37.7%		
Higher degree (MSc/PhD)		16.7%	16.6%		17.2%	17.2%		
Parent/s had high blood pressure	1062			679				
No		83.9%	83.3%		84.5%	83.7%		
Yes		16.1%	16.7%		15.5%	16.3%		
Systolic blood pressure at age 9 (mm Hg)	-	-	-	786	106.22 (12.12)	106.05 (11.70)		
Diastolic blood pressure at age 9 (mm Hg)	-	-	-	786	70.64 (11.17)	70.50 (10.74)		
High systolic blood pressure at age 9				786				
No	-	-	-		86.1%	86.5%		
Yes	-	-	-		13.9%	13.5%		
High diastolic blood pressure at age 9				786				
No	-	-	-		80.5%	81.0%		
Yes	-	-	-		19.5%	19.0%		
Body mass index (z-score) at age 9	-	-	-	796	0.29 (1.06)	0.30 (1.06)		
MVPA (mins/day) at age 9	-	-	-	760	62.44 (22.52)	62.21 (22.57)		
Sedentary (mins/day) at age 9			-	760	431.89 (60.44)	433.22 (60.64)		

222 BMI and blood pressure

223 The cross-sectional associations of BMI with systolic and diastolic blood pressure at age 11 are shown in Table 2. A one standard deviation increase in BMI was associated with 224 225 increases of 0.91 and 1.08 mm Hg for systolic and diastolic blood pressure, respectively, in the confounder-adjusted models. The positive associations between BMI and systolic blood 226 227 pressure were stronger among boys than girls. There was also some evidence to suggest that BMI at age 11 was associated with increased odds of having high systolic blood pressure 228 229 among boys, but not in girls or in the overall sample (Table 3). A one standard deviation increase in BMI was associated with 27% increased odds of high diastolic blood pressure in 230 the overall sample. Evidence for associations between BMI and odds of having high diastolic 231 232 blood pressure were stronger among girls than boys.

Table 2. Cross-sectional associations of body mass index, physical activity and sedentary time with blood pressure at age 11 in the imputed data (N=1283)

Exposure	Systolic blood pressure (mm Hg) at age 11			Diastolic blood pressure (mm Hg) at age 11			
	Coef	ficient (95% Confidence Inte	erval)	Coef	ficient (95% Confidence Inte	erval)	
	All (N=1283)	Boys (N=611)	Girls (N=672)	All (N=1283)	Boys (N=611)	Girls (N=672)	
BMI z-score at age 11	(per						
SD of BMI)							
Model 1	1.30 (0.75 to 1.84)	1.53 (0.66 to 2.39)	1.09 (0.36 to 1.82)	1.15 (0.67 to 1.63)	1.11 (0.28 to 1.94)	1.20 (0.66 to 1.74)	
Model 2	0.91 (0.32 to 1.50)	1.21 (0.28 to 2.15)	0.62 (-0.19 to 1.42)	1.08 (0.54 to 1.62)	1.10 (0.21 to 2.00)	1.00 (0.35 to 1.64)	
P value for sex interaction	0.43			0.99			
MVPA at age 11							
(per 10 mins/day)							
Model 1	0.07 (-0.25 to 0.38)	0.12 (-0.28 to 0.52)	-0.06 (-0.48 to 0.36)	-0.15 (-0.45 to 0.15)	0.06 (-0.33 to 0.45)	-0.26 (-0.66 to 0.14)	
Model 2	0.15 (-0.17 to 0.47)	0.29 (-0.10 to 0.68)	-0.03 (-0.45 to 0.40)	0.05 (-0.24 to 0.33)	0.19 (-0.18 to 0.56)	-0.14 (-0.55 to 0.26)	
P value for sex	0.45			0.22			
interaction	0.45			0.25			
Sedentary time at age	e 11 (per						
10 mins/day)							
Model 1	-0.03 (-0.15 to 0.09)	-0.08 (-0.22 to 0.06)	0.03 (-0.13 to 0.19)	0.02 (-0.07 to 0.12)	-0.01 (-0.12 to 0.10)	0.04 (-0.10 to 0.18)	
Model 2	-0.01 (-0.18 to 0.15)	-0.08 (-0.29 to 0.13)	0.05 (-0.14 to 0.23)	0.08 (-0.05 to 0.21)	0.01 (-0.16 to 0.17)	0.15 (-0.03 to 0.33)	
P value for sex interaction	0.34			0.56			

Model 1 is adjusted for clustering at the school level; Model 2 is additionally adjusted for child age and height at the age 11 data collection,

child sex (for models with all children), highest household education, parental high blood pressure, BMI z-score (for activity variables) and

accelerometer wear time (for activity variables).

Table 3. Cross-sectional associations of body mass index, physical activity and sedentary time with odds of high systolic and diastolic

blood pressure at age 11 in the imputed data (N=1283)

Exposure	High s	systolic blood pressure at	age 11	High diastolic blood pressure at age 11			
	Odds	Ratio (95% Confidence In	terval)	Odds Ratio (95% Confidence Interval)			
	All (N=1283)	Boys (N=611)	Girls (N=672)	All (N=1283)	Boys (N=611)	Girls (N=672)	
BMI z-score at age	11 (per SD						
of BMI)							
Model 1	1.13 (0.98 to 1.30)	1.20 (0.96 to 1.51)	1.08 (0.89 to 1.30)	1.25 (1.06 to 1.47)	1.21 (0.92 to 1.60)	1.28 (1.06 to 1.54)	
Model 2	1.13 (0.97 to 1.32)	1.32 (1.06 to 1.66)	1.00 (0.80 to 1.24)	1.27 (1.05 to 1.53)	1.28 (0.94 to 1.73)	1.24 (1.01 to 1.53)	
P value for sex	0.28			0.84			
interaction	0.38			0.84			
MVPA at age 11	(per 10						
mins/day)							
Model 1	0.94 (0.85 to 1.04)	0.99 (0.88 to 1.12)	0.88 (0.76 to 1.03)	0.96 (0.89 to 1.03)	1.01 (0.90 to 1.12)	0.96 (0.84 to 1.08)	
Model 2	0.96 (0.86 to 1.06)	1.02 (0.90 to 1.17)	0.90 (0.77 to 1.05)	1.01 (0.94 to 1.10)	1.05 (0.93 to 1.17)	0.98 (0.87 to 1.12)	
P value for sex	0.26			0.58			
interaction	0.28			0.38			
Sedentary time at a	age 11 (per						
10 mins/day)							
Model 1	1.01 (0.97 to 1.04)	1.00 (0.96 to 1.05)	1.01 (0.96 to 1.05)	0.99 (0.97 to 1.02)	0.99 (0.96 to 1.02)	0.99 (0.96 to 1.03)	
Model 2	1.03 (0.98 to 1.08)	1.01 (0.95 to 1.08)	1.04 (0.98 to 1.10)	1.01 (0.97 to 1.04)	1.00 (0.95 to 1.05)	1.01 (0.96 to 1.06)	
P value for sex	0.08			0.07			
interaction	0.98			0.97			

240 Model 1 is adjusted for clustering at the school level; Model 2 is additionally adjusted for child age and height at the age 11 data collection, child

sex (for models with all children), highest household education, parental high blood pressure, BMI z-score (for activity variables) and

242 accelerometer wear time (for activity variables).

243 In the prospective models, BMI at age 9 was positively associated with diastolic blood pressure at age 11 in the overall sample and for girls, but not boys (Table 4). A one standard 244 245 deviation increase in BMI at age 9 was associated with a 1.36 mm Hg increase in diastolic 246 blood pressure at age 11 for girls. There was some evidence for a positive association between 247 BMI at age 9 and systolic blood pressure at age 11 in the unadjusted model, but this association 248 was not evident in the adjusted model. In the prospective models for odds of having high blood pressure, BMI at age 9 was associated with increased odds of having high diastolic blood 249 250 pressure at age 11 in the overall sample and for girls (30% and 32% respectively, Table 5). 251 There was no evidence for associations between BMI at age 9 and odds of having high systolic 252 blood pressure at age 11, or with high diastolic blood pressure at age 11 among boys.

Table 4. Prospective associations of body mass index, physical activity and sedentary time at age 9 with blood pressure at age 11 in the

²⁵⁴ imputed data (N=797)

Exposure	Systolic	blood pressure (mm Hg) a	t age 11	Diastoli	c blood pressure (mm Hg) a	t age 11	
	Coef	ficient (95% Confidence Inte	erval)	Coefficient (95% Confidence Interval)			
	All (N=797)	Boys (N=355)	Girls (N=442)	All (N=797)	Boys (N=355)	Girls (N=442)	
BMI z-score at age 9	(per						
SD of BMI)							
Model 1	0.98 (0.16 to 1.80)	1.05 (-0.19 to 2.29)	0.90 (-0.03 to 1.84)	1.18 (0.62 to 1.74)	0.83 (-0.12 to 1.78)	1.44 (0.77 to 2.10)	
Model 2	0.71 (-0.13 to 1.56)	0.79 (-0.55 to 2.12)	0.57 (-0.42 to 1.55)	1.16 (0.49 to 1.84)	0.85 (-0.24 to 1.93)	1.36 (0.59 to 2.13)	
P value for sex	0.70			0.33			
interaction							
MVPA at age 9 (per 10						
mins/day)							
Model 1	0.24 (-0.11 to 0.59)	0.30 (-0.22 to 0.82)	0.24 (-0.25 to 0.74)	-0.02 (-0.33 to 0.29)	-0.004 (-0.46 to 0.45)	0.20 (-0.37 to 0.76)	
Model 2	0.22 (-0.15 to 0.59)	0.32 (-0.19 to 0.84)	0.13 (-0.35 to 0.61)	0.03 (-0.28 to 0.35)	0.008 (-0.44 to 0.46)	0.09 (-0.41 to 0.59)	
P value for sex	0.73			0.85			
interaction							
Sedentary time at age	e 9 (per 10						
mins/day)							
Model 1	0.03 (-0.09 to 0.15)	-0.06 (-0.24 to 0.12)	0.12 (-0.05 to 0.29)	0.12 (0.002 to 0.24)	0.09 (-0.08 to 0.26)	0.13 (-0.03 to 0.29)	
Model 2	-0.02 (-0.21 to 0.17)	-0.11 (-0.38 to 0.15)	0.08 (-0.16 to 0.33)	0.06 (-0.14 to 0.25)	0.04 (-0.23 to 0.31)	0.09 (-0.14 to 0.32)	
P value for sex	0.23			0.54			
interaction							
5 Model 1 is adju	sted for systolic and dias	tolic blood pressure (res	pectively) at age 9 and	for clustering at the sch	ool level; Model 2 is add	litionally	
6 adjusted for chi	ild age and height at the a	ge 9 data collection chi	ld sex (for models with	all children) highest ho	usehold education nare	ntal high	
7 blood program	$\mathbf{PMI}_{\mathbf{Z}}$ googe at ago 0 (for	ge > dutu concetion, en	aggeleromater waar tin	α an emiliately, ingliest in	variables)		
o biobu pressure,	Bivii 2-score at age 9 (10)	activity variables) and	acceleronneter wear this	le al age 9 (101 activity)	variables).		
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263 Table 5. Prospective associations of body mass index, physical activity and sedentary time at age 9 with odds of high systolic and diastolic blood pressure at age 11 in the imputed data (N=797) 264

Exposure	High s	systolic blood pressure at	age 11	High diastolic blood pressure at age 11 Odds Ratio (95% CI)			
		Odds Ratio (95% CI)					
	All (N=797)	Boys (N=355)	Girls (N=442)	All (N=797)	Boys (N=355)	Girls (N=442)	
BMI z-score at age 9							
(per SD of BMI)							
Model 1	1.12 (0.89 to 1.40)	1.06 (0.81 to 1.39)	1.15 (0.83 to 1.58)	1.30 (1.10 to 1.53)	1.15 (0.84 to 1.58)	1.38 (1.13 to 1.67)	
Model 2	1.11 (0.86 to 1.43)	1.12 (0.83 to 1.53)	1.07 (0.76 to 1.51)	1.30 (1.06 to 1.59)	1.20 (0.82 to 1.76)	1.32 (1.04 to 1.68)	
P value for sex	0.70			0.53			
interaction	0.79			0.33			
MVPA at age 9							
(per 10 mins/day)							
Model 1	1.04 (0.92 to 1.17)	1.15 (0.92 to 1.43)	0.97 (0.81 to 1.15)	1.04 (0.92 to 1.16)	0.98 (0.80 to 1.20)	1.17 (1.00 to 1.38)	
Model 2	1.04 (0.91 to 1.19)	1.17 (0.95 to 1.44)	0.92 (0.78 to 1.08)	1.06 (0.95 to 1.19)	0.99 (0.83 to 1.19)	1.13 (0.97 to 1.31)	
P value for sex	0.10			0.27			
interaction	0.19			0.27			
Sedentary time at ag	ge 9 (per						
10 mins/day)							
Model 1	1.03 (0.99 to 1.07)	0.99 (0.93 to 1.05)	1.07 (1.02 to 1.13)	1.03 (1.00 to 1.06)	1.01 (0.96 to 1.06)	1.04 (1.00 to 1.07)	
Model 2	1.02 (0.96 to 1.09)	0.96 (0.86 to 1.06)	1.08 (1.01 to 1.16)	1.00 (0.94 to 1.06)	1.01 (0.92 to 1.10)	1.00 (0.93 to 1.07)	
P value for sex interaction	0.03			0.32			

Model 1 is adjusted for systolic and diastolic blood pressure (respectively) at age 9 and for clustering at the school level; Model 2 is additionally 265

adjusted for child age and height at the age 9 data collection, child sex (for models with all children), highest household education, parental high 266 blood pressure, BMI z-score at age 9 (for activity variables) and accelerometer wear time at age 9 (for activity variables). 267

268 **Physical activity and blood pressure**

269 In the cross-sectional models, there was no strong evidence for associations between MVPA or sedentary time with systolic or diastolic blood pressure (Table 2), or odds of having 270 271 high systolic or diastolic blood pressure (Table 3), in any of the models. In the prospective 272 analyses, there was weak evidence for a positive association between sedentary time at age 9 and diastolic blood pressure at age 11, but this association was only evident in the unadjusted 273 model (Table 4). For girls, MVPA at age 9 was associated with increased odds of having high 274 275 diastolic blood pressure at age 11 in the unadjusted model only (Table 5). Similarly, there was 276 evidence for a positive association between sedentary time at age 9 and odds of having high 277 systolic blood pressure at age 11 among girls in both models. There was also weak evidence 278 (unadjusted model only) that sedentary time at age 9 was associated with increased odds of having high diastolic blood pressure at age 11 in the overall sample and among girls, but not 279 280 boys.

281

282 **Discussion**

In this study, we found small, but consistent, cross-sectional and prospective 283 associations of higher BMI with higher mean diastolic blood pressure and the likelihood of 284 having high diastolic blood pressure. Prospectively, a one standard deviation increase in BMI 285 286 at age 9 was associated with an increase of 1.16 mm Hg for diastolic blood pressure at age 11, 287 as well as 30% increased odds of high diastolic blood pressure at that age. These associations were stronger in girls compared to boys. For systolic blood pressure, the association with BMI 288 289 was only evident in the cross-sectional models at age 11. There was only very weak evidence 290 that children's physical activity or sedentary time at age 9 were prospectively associated with 291 blood pressure at age 11 years. These findings suggest that while greater BMI during middle childhood may influence the future risk of higher diastolic blood pressure, interventions aimed
at increasing physical activity and reducing sedentary time are unlikely to impact the
development of cardiovascular disease risk during childhood. Whereas, interventions to reduce
BMI, or prevent high BMI, have the potential to impact children's blood pressure. The findings
also suggest that childhood BMI might be a more important risk factor for higher diastolic than
systolic blood pressure.

298 This study adds prospective evidence, with blood pressure measured at two time-points, 299 and therefore the ability to adjust for baseline blood pressure, to the existing cross-sectional 300 and prospective studies suggesting that there is a positive association between BMI and blood 301 pressure among children [15-24]. A cross-sectional study with 3923 children aged 6-11 years 302 from southern Italy found BMI and waist circumference z-scores were positively associated 303 with blood pressure [16]. Another cross-sectional study of 1432 twelve-year-olds found a high BMI and large waist circumference (above the 90th percentile) were associated with higher 304 305 systolic and diastolic blood pressure levels and adverse blood cholesterol levels [22]. These 306 studies highlight the potential usefulness of both BMI and waist circumference measures for 307 identifying those at risk of future adverse cardiovascular risk profiles. A large prospective study 308 of 5235 children also from Bristol, UK (Avon Longitudinal Study of Parents and Children (ALSPAC)), found a one standard deviation (SD) increase in BMI at age 9-12 years was 309 310 associated with an increased risk of high systolic blood pressure (≥130mm Hg) at age 15-16 in 311 girls (odds ratio (OR): 1.23, 95% confidence interval (CI): 1.10 to 1.38) and boys (OR: 1.24, 312 95% CI: 1.13 to 1.37) [21]. However, they found no evidence for associations between BMI 313 and high diastolic blood pressure [21], which is contrary to our findings at somewhat younger 314 ages of an association with diastolic but not systolic blood pressure. This difference may be due to chance or to differences in age or birth cohort; children in the current cohort were born 315 316 in 2006-2007, while the ALSPAC participants were born in the 1990s, when population levels 317 of childhood obesity were lower [55]. The prospective associations between BMI with high 318 diastolic blood pressure in the present study were greater in magnitude than the previously published study of cross-sectional associations at age 9 in the same cohort (B-PROACT1V), 319 320 using the equivalent definition of high blood pressure and the same confounders [24]. This 321 suggests that excessive weight during childhood may be progressively associated with risk of 322 elevated diastolic blood pressure as children age, and this may be particularly the case for girls. 323 With replication and additional causal evidence, for example from methods such as within 324 sibship analyses or Mendelian randomization, this would suggest that effective obesity 325 prevention interventions, especially those that target girls, are needed from an early age. 326 However, it is also possible that the somewhat weaker association in the earlier cross sectional 327 analyses are chance findings.

328 In the current study, only one physical activity or sedentary time variable was 329 associated with blood pressure in the adjusted models. For girls, sedentary time at age 9 was 330 positively associated with the odds of having high systolic blood pressure at age 11 (OR: 1.08, 331 95% CI: 1.01 to 1.16), but the evidence of association was weak and there was no association 332 for boys, in the overall sample, or with blood pressure measured as a continuous variable. In contrast to our lack of association between physical activity and blood pressure, a study from 333 Birmingham, UK (cross-sectional analyses: N=512; two-year follow-up prospective analyses: 334 335 N=427), found total physical activity was inversely associated with diastolic blood pressure 336 cross-sectionally and prospectively; there was not strong evidence of association with systolic 337 blood pressure in either analysis [38]. Similarly, a cross-sectional study examining the cardiovascular health of 2049 9-to-10-year-old children from three UK cities, found total 338 339 physical activity level was inversely associated with lower diastolic blood pressure, but no evidence for an association with systolic blood pressure [30]. The contrast in findings between 340 341 these studies and the current study may be due to the differences between cohorts. The 342 Birmingham participants were younger at baseline (mean age 6.5 years, range 5.4-7.8 years) 343 and predominantly from a South Asian background [38], and the multi-city sample were also 344 more ethnically diverse [30] compared to the current study. In an earlier B-PROACT1V study 345 [24], there was no evidence that physical activity or sedentary time at 6 or 9 years were cross-346 sectionally or prospectively associated with systolic or diastolic blood pressure at 9 years. The present study adds to this evidence, demonstrating a lack of strong evidence of association 347 348 between physical activity and sedentary time with blood pressure at 11 years within the same cohort. These findings are in line with a Danish study that examined the cross-sectional 349 350 associations between objectively-assessed physical activity and metabolic syndrome among 351 589 8-10 year-old-children [44]. The study found no evidence of an association between physical activity and systolic or diastolic blood pressure [44]. The conflicting findings in the 352 353 literature suggest that other risk factors for higher blood pressure (e.g., BMI, genetic 354 influences) may be more important than physical activity in primary school aged children. They 355 also suggest that interventions to reduce BMI via physical activity and sedentary behaviour 356 may not directly reduce blood pressure in children.

357 In adults, a 2 mm Hg reduction in blood pressure is associated with a 6% reduction in coronary heart disease and a 15% reduction in stroke-related events [56]. In the present study, 358 359 the cross-sectional differences in systolic and diastolic blood pressure per standard deviation 360 of BMI were small (0.91 and 1.08 mm Hg, respectively), suggesting that relatively large 361 reductions in BMI are needed to reduce cardiovascular disease risk. The findings do indicate 362 that measuring BMI in primary-school children is a suitable and relatively low-cost measure 363 (compared to estimating body composition using dual energy x-ray absorptiometry) for 364 identifying those at risk of future adverse cardiovascular risk profiles. Prevention strategies are needed to shift the population distribution of childhood adiposity downwards. 365

366 Strengths and limitations

367 Strengths of this study include the measurement of blood pressure at two ages in 368 childhood, and the objective measurements of BMI and physical activity (via accelerometers), 369 allowing us to examine cross-sectional and prospective associations of these exposures at ages 370 9 and 11 years with blood pressure at age 11. Multiple imputation of missing data was used to 371 increase precision and reduce selection bias in our coefficient estimates [57]. Findings in the imputed data were consistent with the complete case analyses. Due to the observational nature 372 373 of the study, we were unable to exclude residual confounding, for example by dietary factors 374 such as salt, sugar or fat intake. Low birth weight is also associated with high blood pressure 375 later in life, but this information was not available in the present study [58]. The study sample 376 was relatively homogenous, primarily of White British origin from one area of the UK, which 377 limits the ability to extrapolate to other ethnic groups in more diverse areas of the UK.

378

379 Conclusions

The findings of the present study strengthen existing evidence suggesting that BMI may 380 be a risk factor in the development of high diastolic blood pressure during childhood. 381 Conversely, the amount of time that predominantly White British primary school aged children 382 383 spend being physically active or sedentary does not appear to be strongly associated with blood 384 pressure. These results suggest that interventions to prevent excessive bodyweight may be important in the prevention of cardiovascular disease risk during childhood. Current evidence 385 386 is limited on the effectiveness of physical activity interventions on BMI [59], and our results 387 suggest targeting this may not directly reduce blood pressure, therefore, future obesity prevention initiatives should target multiple components (e.g., physical activity, nutrition, and 388 emotional well-being), rather than focus on increasing physical activity in isolation. 389

390

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