

1 **Association between mode of delivery and body mass index at 4-5**
2 **years in White British and Pakistani children: the Born in Bradford**
3 **birth cohort**

4 **Eleanor Ralphs^{1*}, Lucy Pembrey¹, Jane West², Gillian Santorelli²**

5 ¹London School of Hygiene and Tropical Medicine, Keppel Street, London, UK

6 ²Bradford Institute for Health Research, Bradford Teaching Hospitals Foundation
7 Trust, Bradford, UK

8 *Corresponding author: Eleanor Ralphs (email: eleanorr@sky.com)

9 **Abstract**

10 **Background**

11 Globally, it is becoming more common for pregnant women to deliver by caesarean section
12 (CS). In 2020, 31% of births in England were CS, surpassing the recommended prevalence
13 of CS. Concerns have been raised regarding potential unknown consequences of this mode
14 of delivery.

15 Childhood adiposity is also an increasing concern. Previous research provides inconsistent
16 conclusions on the association between CS and childhood adiposity. More studies are
17 needed to investigate the consequences of CS in different populations and ethnicities.
18 Therefore, this study investigates the association between mode of delivery and BMI, in
19 children of 4-5 years and if this differs between White British (WB) and Pakistani ethnicities,
20 in Bradford UK.

21 **Methods**

22 Data were obtained from the Born in Bradford (BiB) cohort, which recruited pregnant women
23 at the Bradford Royal Infirmary, between 2007-2010. For these analyses, a sub-sample
24 (n=6410) of the BiB cohort (n=13858) was used.

25 Linear regression models determined the association between mode of delivery (vaginal or
26 CS) and BMI z-scores at 4-5 years. Children were categorised as underweight/healthy
27 weight, overweight and obese, and logistic regression models determined the odds of
28 adiposity. Effect modification by ethnicity was also explored.

29 **Results**

30 Multivariable analysis found no evidence for a difference in BMI z-score between children of
31 CS and vaginal delivery (0.005 kg/m², 95% CI= -0.062–0.072, p=0.88). Neither was there
32 evidence of CS affecting the odds of being overweight (OR=1.05, 95% CI=0.86–1.28,

33 p=0.65), or obese (OR=0.98, 95% CI=0.74–1.29, p=0.87). There was no evidence that
34 ethnicity was an effect modifier of these associations (p=0.97).

35 **Conclusion**

36 Having CS, compared to a vaginal delivery, was not associated with greater adiposity in
37 children of 4-5 years in this population. Concerns over CS increasing adiposity in children
38 are not supported by the findings reported here using the BiB study population, of both WB
39 and Pakistani families.

40 **Background**

41 Delivery by caesarean section (CS) is increasing globally. Using data from 150 countries
42 from 1990 to 2014, longitudinal analysis suggests that CS represent 18.6% of all births (1).
43 In England, CS rates rose from 23% to 31%, between 2004 and 2020 (2).

44 Research suggests CS rates are increasing due to protective effects against fetal death (3)
45 and to avoid adverse impacts of macrosomia in obese and diabetic mothers (4). Also CS is
46 sometimes perceived as more convenient, less painful and more profitable for private
47 hospitals (4). Contributions to such CS rates additionally arise from the cohort of women
48 who have had one previous CS (5).

49 However, the rise in CS has aroused alarm due to the lack of knowledge on the short- and
50 long-term risks. The World Health Organisation recommends CS should ideally only be
51 undertaken when medically necessary and that CS rates higher than 10% are not
52 associated with reductions in maternal and newborn mortality rates (6). Some evidence
53 suggests those who have undergone CS have just over twice the odds of severe maternal
54 morbidity, compared to those experiencing vaginal deliveries (3). CS has also been
55 associated with other complications, such as a higher risk of immune and metabolic
56 disorders in children (7), and offspring overweight and obesity (8–11) . The latter
57 complication will be investigated in this report.

58 Overweight and obesity in England was prevalent in 22.6% of children aged 4-5 years, in
59 2018-2019; more specifically 23.1% in White British (WB) children and 19.9% in Pakistani
60 children (12).

61 In the first six months of life, the colonisation and diversity of gut microbiota is associated
62 with the mode of delivery (13). Those born vaginally have a higher abundance of
63 Bifidobacteria and Bacteroides than those born by CS (13). These bacteria genera have a
64 protective effect against being overweight as they are well equipped to obtain nutrients from
65 breast (or formula) milk oligosaccharides (14). Additionally, there is evidence to suggest the

66 gut microbiota of a child born by CS is more abundant in *Staphylococcus aureus*, which has
67 been associated with the development of obesity (14,15). It is important to note that
68 guidelines endorse the use of prophylactic antibiotics for women undergoing CS, to prevent
69 wound infection (16).

70 However, in children over the age of six months, there is very weak evidence of an
71 association between mode of delivery and gut microbiota (13,17), suggesting that the
72 protective effect of vaginal deliveries against adiposity attenuates through early childhood.
73 This conflicts with some evidence of an association between mode of delivery and BMI
74 found in adult life (18), highlighting additional mechanisms might explain this association.

75 Previous research presents mixed results. One systematic review concluded children
76 delivered by CS had higher odds of being overweight or obese at 0-8 years (pooled odds
77 ratio from 10 studies=1.32, 95% CI=1.15–1.51) (8). Another review also determined CS
78 children to be at higher risk of being obese at 2-18 years (pooled risk ratio from 19
79 studies=1.34, 95% CI=1.18–1.51) (9).

80 Nine other studies provide evidence to suggest delivering by CS increases the risk of child
81 adiposity (19–27). However, three studies have found no evidence of differences in child
82 BMI between CS and vaginal deliveries (28–30).

83 A further search found one study to address the effects of ethnicity on the association of
84 mode of delivery and child adiposity. The study found differing race-specific effects of CS
85 with body size at 2 years between African American and non-African American mothers (26).
86 In children of African American mothers, CS was associated with a significantly higher odds
87 of obesity, whereas no association was found in children of non-African American mothers
88 (26). It was suggested that ethnic differences in the developing gut microbiome or epigenetic
89 structure, could be the cause of the effect modification (26).

90 In this paper, studies were cited if they reported effect estimates for the association between
91 the mode of delivery (CS compared with vaginal delivery) and overweight or obesity in
92 children. The age range defining childhood was 2–18 years.

93 **Rationale for this study**

94 There is limited published research on the direct association of mode of delivery and child
95 BMI, at the age of children starting school (4-5 years). This association has not been
96 investigated in UK South Asian mothers compared to WB mothers.

97 The aim of this study was to determine if there is any association between mode of delivery
98 (CS and vaginal delivery) and BMI at 4-5 years of age, in the Born in Bradford (BiB) cohort,
99 and if this differs between ethnicities (WB and Pakistani).

100

101 **Methods**

102 **Study design**

103 Born in Bradford (BiB) is a longitudinal multi-ethnic birth cohort study. BiB aims to investigate
104 parent and child wellbeing by examining physiological, environmental and genetic factors in
105 the City of Bradford (31). Bradford is situated in the north of England; it is ethnically diverse
106 and has high levels of socio-economic deprivation. BiB recruitment occurred from
107 September 2007 to December 2010. Women who attended the Bradford Royal Infirmary at
108 26-28 weeks gestation for a routine glucose-tolerance test, which is offered to all women
109 booked to give birth in Bradford, were invited to join the study and 87% of those approached
110 agreed to participate. It is at this point that women were weighed, and their height measured.
111 Weight at first antenatal clinic assessment (median 12 weeks' gestation) was abstracted
112 from the antenatal records and this weight, together with height measured at recruitment,
113 was used to calculate the woman's early pregnancy BMI (kg/m^2).

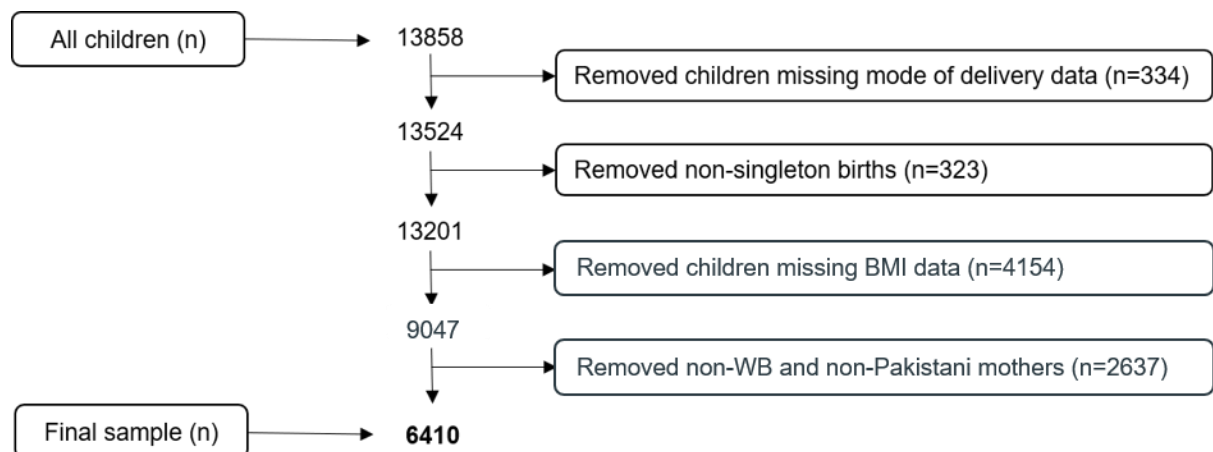
114 The BiB population is broadly representative of the maternal population in Bradford (31).
115 12453 pregnant women gave consent to be involved with the study. One woman could

116 contribute more than a single pregnancy, resulting in a total of 13776 pregnancies being
117 reported, which gave rise to 13858 children. A baseline questionnaire was conducted at
118 recruitment. The mother self-reported most variables, including ethnicity, socioeconomic
119 indicators, alcohol-related and smoking habits. In this analysis, the mother's self-reported
120 ethnicity was used to define their child's ethnicity. School nurse teams took anthropometric
121 measurements of children in reception class (aged 4-5) as part of the National Child
122 Measurement Programme (NCMP). Where anthropometric measurements were missing
123 from NCMP data, it was possible to obtain some of these measurements from Primary Care
124 and Child Health Records at this age. Of the non-missing anthropometric data, 82.2%
125 originated from NCMP, 10.2% from Primary Care, 7.6% from Child Health Records. A
126 subgroup of mothers was followed up for data on breastfeeding, at 6, 12, 18, 24 and 36
127 months. This subgroup was part of BiB1000, a nested cohort of the BiB prospective birth
128 cohort (32).

129 **Study population**

130 From the total 13858 children enrolled in BiB, 7448 were excluded due to not meeting the
131 inclusion criteria of having data on mode of delivery, BMI at 4-5 years, singleton birth and
132 being WB or Pakistani. As approximately a third of the children enrolled in BiB are missing
133 outcome data (BMI at 4-5 years), a characteristics table comparing the final study population
134 (n=6410) and those with missing BMI at 4-5 years (n=4154) is presented in the
135 supplementary material. There is no suggestion of selection bias at this stage.

136 Mothers self-reporting an ethnicity other than WB or Pakistani were excluded as they were a
137 very heterogeneous ethnic group (429 = Indian, 288 = White Other, 253 = Bangladeshi, 226
138 = Black, 108 = Mixed-White and Black, 61 = Mixed-White and South Asian, 309 = other).
139 This left a sample size of 6410 children (Figure 1). The original BiB cohort had a similar
140 distribution of child BMI and mode of delivery to the study population.



141
142 **Figure 1. A flowchart describing the selection of the final study population.**

143 **Abbreviation: WB, White British.**

144 **Exposure (mode of delivery)**

145 Exposed participants were children who were delivered by elective or emergency CS.

146 Unexposed participants were children who were delivered vaginally; including normal,
147 forceps and ventouse extraction deliveries. Mode of delivery was recorded by a midwife or
148 paediatrician within the first six hours of life. Paper forms with handwritten notes were
149 entered into the routine eClipse electronic maternity record as neonatal data.

150 **Outcome (child BMI)**

151 BMI values of children aged 4-5 years, recorded as part of the NCMP by school nurse teams
152 (33) or obtained from Primary Care or Child Health Records, were transformed to a
153 standardised measure (z-scores). The z-scores were calculated using the LMS method. This
154 is prepared via an Excel spreadsheet, which can be obtained online for free (34). The LMS
155 growth application includes access to a 1990 UK (UK90) reference population. Using this
156 reference, each individual is assigned a z-score which adjusts for age, sex and the BMI
157 distribution for skewness (35). The UK90 reference group is recommended for population
158 monitoring and clinical assessment in children aged four years and over (36). It serves as an
159 anchor for comparison; it is used by the NCMP and has been used for other BiB studies
160 (37). Children with BMI z-scores above the 85th percentile were classified as overweight, and
161 those above the 95th percentile as obese (38).

162 **Sample size calculations**

163 Sample size calculations were conducted in OpenEpi (39). This study had a power of 99%,
164 determined from a post-hoc power calculation using the parameters from this study (vaginal
165 to CS ratio of 3.71 and prevalence of childhood overweight or obesity in vaginal births at
166 14.8%), and the odds ratio from a previous study (odds ratio of 2.10 (95% CI 1.36-3.23) of
167 obesity in children aged 7 years, by CS compared to vaginal birth) (10).

168 **Statistical analysis**

169 All analyses were conducted in Stata/IC 15.1. Figures were produced using RStudio version
170 1.3.1056. Variables that had good evidence (chi-squared tests, ANOVA, and judging
171 correlation to have approximately $p < 0.05$) to suggest they had an association with both
172 mode of delivery and z-scores, as well as not being on the causal pathway, met the criteria
173 to be potential confounders. The following variables were considered for assessment of
174 being potential confounders: maternal age; maternal BMI; maternal education; maternal job
175 status; maternal house tenure; maternal benefits received; maternal drinking of alcohol
176 during pregnancy or three months before; maternal smoking during pregnancy; parity;
177 maternal gestational diabetes; child gender; child birthweight; gestational period.

178 **Multivariable analysis**

179 The forward selection approach was used to create regression models. Potential
180 confounders were added individually according to their effect size. The covariate was
181 retained in the model if there was an appreciable (10%) difference in effect size of mode of
182 delivery on z-score.

183 The final multivariable linear regression model assessed the association between mode of
184 delivery and BMI z-score. Preliminary analysis confirmed the assumptions of the regression
185 were met; z-scores were normally distributed and lacked collinearity. BMI z-scores were also
186 categorised and logistic regression models performed to obtain odd ratios for being
187 overweight and obese in children delivered by CS. All the study population ($n=6410$)

188 contributed to the unadjusted regression modeling. Complete case analysis was used for the
189 adjusted regression modeling (n=6115).

190 **Effect modification**

191 Potential effect modification was judged by stratifying the final model by ethnicity to observe
192 the separate association of mode of delivery on z-score in WB and Pakistani children. A
193 formal test for effect modification was also conducted; a likelihood ratio test compared the
194 final model with a model which also included an interaction term between mode of delivery
195 and ethnicity.

196 **Missing data**

197 The number and proportion of patients missing data on descriptive variables was described.
198 Complete case analysis was used for the multivariate analysis. No imputation was
199 performed.

200 Approximately 85% of the study population had missing data on breastfeeding (n= 5439
201 missing). Due to the large proportion of missingness, breastfeeding was not assessed in this
202 study. Maternal parity (n= 273 missing) and maternal BMI (n= 275 missing) were missing for
203 about 4% of the study population.

204

205 **Results**

206 **Descriptive results**

207 Table 1 and 2 summarise the baseline characteristics of the study population stratified by
208 mode of delivery and ethnicity, also visualised in Figure 2. In this study, 21.3% (n=1361) of
209 babies were delivered by CS. There were more Pakistani mothers (54.6%, n=3502) than
210 WB mothers (45.4%, n=2908). Amongst Pakistani mothers, 19.8% had CS, whereas the CS
211 prevalence among WB mothers was higher at 23.0%. Most children were underweight/
212 healthy weight (84.7%), and fewer were overweight (10.0%) or obese (5.2%) (Table 2). The

213 mean BMI z-score was slightly higher in CS deliveries (0.32) than vaginal deliveries (0.22) at
214 age 4-5 years. Furthermore, the mean BMI z-score was higher in WB children (0.43) than
215 Pakistani children (0.08).

216

217 **Table 1. Baseline characteristics, of the study population, stratified by mode of**
 218 **delivery.**

		Vaginal (n= 5049)		Caesarean (n= 1361)		P value
		n	%	n	%	
Child BMI categories (n= 6410)	Underweight/Healthy weight	4303	85.22	1129	82.95	0.039
	Overweight	490	9.70	153	11.24	
	Obese	256	5.07	79	5.80	
Child BMI z-score (n= 6410)	Mean		0.22		0.32	0.003
	SD		1.11		1.15	
Ethnicity (n= 6410)	White British	2239	44.35	669	49.16	0.002
	Pakistani	2810	55.65	692	50.84	
Maternal age (years) (n= 6410)	Mean		27.12		28.77	<0.0001
	SD		5.53		5.73	
Maternal BMI at early pregnancy categories (n= 6135)	Underweight/Healthy weight	2502	51.83	517	39.53	<0.0001
	Overweight	1414	29.29	395	30.20	
	Obese	911	18.87	396	30.28	

Maternal BMI at early pregnancy (kg/m²) (n= 6135)	Mean		25.67		27.67	<0.0001
	SD		5.46		6.26	
Maternal education (n= 6395)	<5 GCSE equivalent	1230	24.41	272	20.04	<0.001
	5 GCSE equivalent	1718	34.10	437	32.20	
	A-level equivalent	695	13.80	181	13.34	
	Higher than A-level	1058	21.00	362	26.68	
	Foreign unknown/other	337	6.69	105	7.74	
Maternal job status (n= 6400)	Currently employed	2029	40.24	678	49.93	<0.001
	Previously employed	1517	30.09	368	27.10	
	Never employed	1496	29.67	312	22.97	
Maternal house tenure	Owns outright	807	16.02	207	15.25	0.055
	Mortgage	2518	49.97	718	52.91	

(n= 6396)	Private landlord	792	15.72	172	12.68	
	Social housing	522	10.36	145	10.69	
	Rent free/other	400	7.94	115	8.47	
Maternal benefits received (n= 6385)	Yes	2237	44.51	496	36.50	<0.001
	No	2789	55.49	863	63.50	
Maternal drinking of alcohol during pregnancy or 3 months before (n= 6393)	Yes	1570	31.17	495	36.42	0.001
	No	3465	68.79	863	63.50	
Maternal smoking during pregnancy (n= 6396)	Yes	856	16.99	210	15.45	0.176
	No	4181	83.01	1149	84.55	
	Never	211	27.47	59	29.06	
Parity (n= 6137)	Primiparous	1802	37.22	586	45.22	<0.001
	Multiparous	3039	62.78	710	54.78	
Maternal gestational	Yes	342	6.78	147	10.83	<0.001
	No	4705	93.22	1210	89.17	

diabetes						
(n= 6404)						
Child gender	Male	2548	50.47	718	52.76	0.134
	Female	2501	49.53	643	47.24	
(n= 6410)						
Child birthweight (g)	Mean		3244.10		3215.84	0.089
	SD		503.61		675.73	
(n= 6410)						
Gestational period (days)	Mean		277.67		273.90	<0.0001
	SD		11.09		15.34	
(n= 6410)						
Gestational period (weeks)	Preterm (<37 weeks)	208	4.12	117	8.60	<0.0001
	Term (≥37 weeks)	4841	95.88	1244	91.40	
(n= 6410)						

P values to provide the level of statistical evidence on the difference between mode of delivery; obtained from chi-squared tests or ANOVA, where appropriate. Abbreviations: n, sample size; BMI, body mass index; SD, standard deviation.

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Table 2. Baseline characteristics, of the study population, stratified by ethnicity.

		All		White British (n= 2908)		Pakistani (n= 3502)		P value
		n	%	n	%	n	%	
Child BMI categorised (n= 6410)	Underweight/	543	84.74	243	83.73	2997	85.5	0.041
	Healthy	2		5			8	
	weight							
	Overweight	643	10.03	335	11.52	308	8.79	
	Obese	335	5.23	138	4.75	197	5.63	
Child BMI z-score (n= 6410)	Mean		0.24		0.43		0.08	<0.0001
	SD		1.12		0.97		1.21	
Maternal age (years) (n= 6410)	Mean		27.47		27.03		27.84	<0.0001
	SD		5.61		6.09		5.15	
Maternal BMI at early pregnancy categorised (n= 6135)	Underweight/		49.2		45.7		52.0	<0.0001
	Healthy	3019	1	1269	8	1750	4	
	weight							
	Overweight	1809	29.4	810	29.2	999	29.7	
		9		2		1		
	Obese	1307	21.3	693	25.0	614	18.2	
			0		0		6	
Maternal BMI at early pregnancy (kg/m²) (n= 6135)	Mean		26.09		26.80		25.50	<0.0001
	SD		5.69		5.97		5.39	
Maternal education (n= 6395)	<5 GCSE	150	23.49	577	19.87	925	26.5	<0.001
	equivalent	2					0	
	5 GCSE	215	33.70	102	35.40	1127	32.2	

	equivalent	5		8			8	
	A-level	876	13.70	458	15.77	418	11.9	
	equivalent						7	
	Higher than	142	22.20	560	19.28	860	24.6	
	A-level	0					3	
	Foreign	442	6.91	281	9.68	161	4.61	
	unknown/other							
Maternal job status (n= 6400)	Currently employed	270	42.30	192	66.21	783	22.4	<0.001
		7		4			1	
	Previously employed	188	29.45	775	26.67	1110	31.7	
		5					7	
	Never employed	180	28.25	207	7.12	1601	45.8	
		8					2	
Maternal house tenure (n= 6396)	Owens outright	101	15.85	119	4.10	895	25.6	<0.001
		4					3	
	Mortgage	323	50.59	151	52.31	1717	49.1	
		6		9			7	
	Private landlord	964	15.07	654	22.52	310	8.88	
	Social housing	667	10.43	451	15.53	216	6.19	
	Rent free/other	515	8.05	161	5.54	354	10.1	
							4	
Maternal benefits received (n= 6385)	Yes	273	42.80	106	36.73	1670	47.8	<0.001
		3		3			4	
	No	365	57.20	183	63.27	1821	52.1	
		2		1			6	
Maternal drinking of alcohol during pregnancy	Yes	206	32.29	205	70.72	10	0.29	<0.001
		5		5				
	No	432	67.67	848	29.18	3480	99.7	
		8					1	

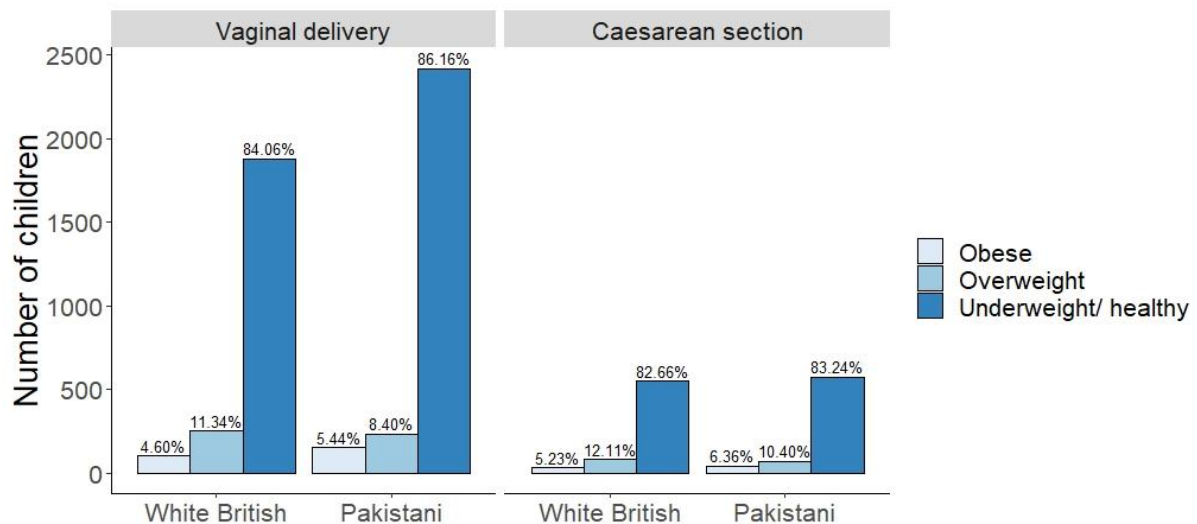
or 3 months								
before								
(n= 6393)								
Maternal smoking during pregnancy	Yes	106	16.67	961	33.08	105	3.01	<0.001
	No	6						
(n= 6396)		533	83.33	194	66.92	3386	96.9	
		0		4			9	
Parity	Primiparous	238	38.91	135	48.11	1037	31.1	<0.001
		8		1			5	
(n= 6137)		374	61.09	145	51.89	2292	68.8	
		9		7			5	
Maternal gestational diabetes	Yes	489	7.64	141	4.85	348	9.95	<0.001
	No	591	92.36	276	95.15	3151	90.0	
(n= 6404)		5		4			5	
Child gender	Male	326	50.95	148	51.13	1779	50.8	0.789
		6		7			0	
(n= 6410)		314	49.05	142	48.87	1723	49.2	
		4		1			0	
Child birthweight (g)	Mean	3238.10		3357.60		3138.86		<0.0001
	SD	544.78		550.53		519.56		
(n= 6410)								
Gestational period (days)	Mean	276.86		277.63		276.22		<0.0001
	SD	12.22		12.51		11.94		
(n= 6410)								
Gestational period	Preterm (<37 weeks)	325	5.07	157	5.40	168	4.80	
(n= 6410)		6085	94.9	2751	94.6	3334	95.2	0.300
			3		0		0	

P values to provide the level of statistical evidence on the difference between White

British and Pakistani ethnic groups; obtained from chi-squared tests or ANOVA, where appropriate. Abbreviations: n, sample size; BMI, body mass index; SD, standard deviation.

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224
 225 **Figure 2. A bar chart showing the number of children in the study population,**
 226 **stratified by mode of delivery, ethnicity and categorised BMI.**

227 **Mother's booking BMI**

228 Almost half the mothers in this study population were underweight or healthy weight
 229 (49.2%). Obesity was more prevalent in women who gave birth by CS (30.3%), compared to
 230 mothers giving birth vaginally (18.9%). WB mothers had a slightly higher BMI (+1.30 kg/m²)
 231 than Pakistani women. Leading to a higher prevalence of obesity in WB mothers (25.0%)
 232 than Pakistani women (18.2%).

233

234 **Family sociodemographic factors**

235 Mothers who had CS often had achieved a higher level of education than mothers with
 236 vaginal deliveries (26.7% of CS mothers, 21.0% of vaginal mothers). More Pakistani women
 237 achieved higher than A-level qualifications compared to WB women (24.6 vs 19.3%).

238 Most mothers were not currently employed (67.7%). Current employment was more
 239 common in women who had CS births, compared to those with vaginal births (49.9% and
 240 40.2%, respectively) and also more common in WB women (66.2%), compared to Pakistani
 241 women (22.4%). Additionally, a higher proportion of mothers having vaginal deliveries

242 (44.5%) received benefits than those having CS deliveries (36.5%). Receiving benefits was
243 more common in Pakistani mothers (47.8%) than WB mothers (36.7%).

244 **Gestational factors**

245 The mean age of mothers who had a CS was 28.8 years old, which was 1.7 years older than
246 those who gave birth vaginally. Also, mothers having vaginal deliveries were more likely to
247 be multiparous (62.8%) compared to mothers having CS deliveries (54.8%). The difference
248 in mean gestation period between CS and vaginal deliveries was minimal (4 days
249 difference). A larger proportion of preterm births was experienced by mothers who had a CS
250 (8.6%), compared to mothers who gave birth vaginally (4.1%).

251 Only 10 out of 3490 Pakistani mothers (0.3%) drank alcohol during pregnancy or 3 months
252 before, whereas 70.7% of WB mothers reported alcohol consumption. Further to this,
253 mothers having CS were marginally more likely to have drunk alcohol during pregnancy or 3
254 months before (36.4% of CS mothers and 31.2% of vaginal mothers). More WB women
255 reportedly smoked during pregnancy than Pakistani women (33.1% of WB women, 3.0% of
256 Pakistani women).

257 Gestational diabetes was more prevalent in mothers having CS (10.8%) compared to those
258 having vaginal deliveries (6.8%). Additionally, prevalence was higher in Pakistani mothers
259 than WB mothers (10.0% and 4.9% respectively).

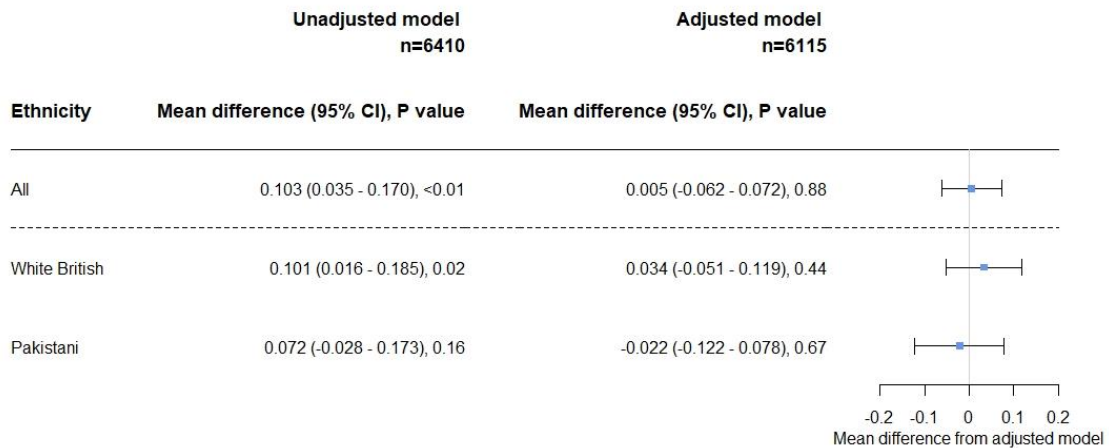
260 **Child factors**

261 Children born by CS had a mean birthweight 28.3 g lower than vaginal births. Irrespective of
262 mode of delivery, children with WB mothers had a higher mean birthweight than those with
263 Pakistani mothers (3357.6 g and 3138.9 g, respectively).

264 **Multivariable analysis**

265 The unadjusted linear regression calculated the predicted difference in z-score between
266 mode of delivery, the z-score being higher with CS (n=6410, difference= 0.103; 95% CI=

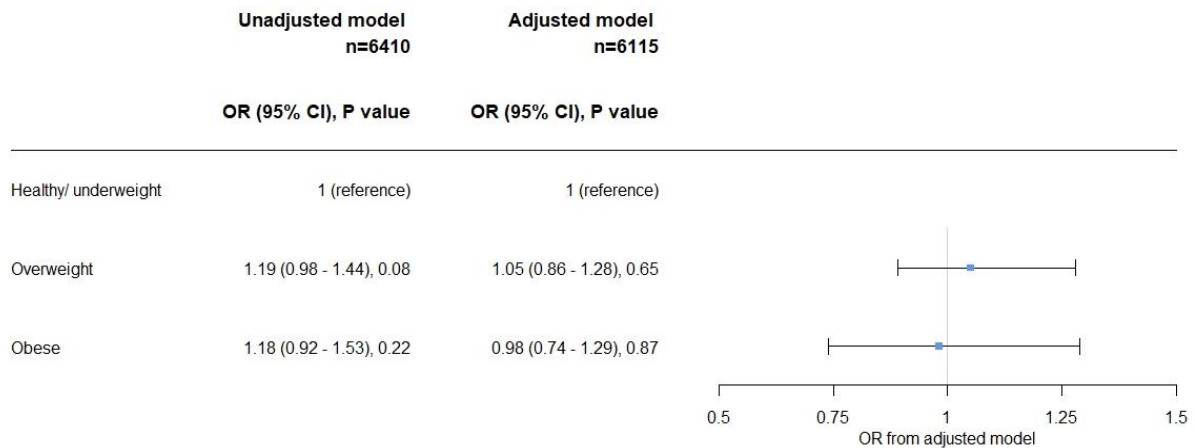
267 0.035–0.170) (Figure 3). The adjusted model calculated the predicted mean difference in z-
 268 score between mode of delivery, controlling for all factors which met the confounding criteria
 269 (ethnicity, maternal BMI (continuous), maternal job status and maternal drinking of alcohol
 270 during pregnancy or 3 months before), there was no difference in child BMI z-score (n=
 271 6115, difference= 0.005; 95% CI= -0.062–0.072) (Figure 3).



272 **Figure 3. Mean differences in BMI z-score depending on mode of delivery, at 4-5 years**
 273 **old, from White British and Pakistani ethnic groups.** The reference group had vaginal
 274 deliveries. Values obtained are β coefficients from unadjusted and adjusted linear regression
 275 models. P value acquired from t-tests. The adjusted model controls for: ethnicity, maternal
 276 BMI (continuous), maternal job status and drinking alcohol during or 3 months before
 277 pregnancy.
 278 pregnancy.

279 The proportion of variance in BMI z-scores explained by the mode of delivery was 7.49% for
 280 the adjusted model (adjusted R^2 value). The F ratio (71.76) shows how much variability the
 281 model can explain relative to how much it cannot explain. The standard error (0.034) did not
 282 differ between unadjusted and adjusted models, suggesting an absence of collinearity.

283 The adjusted logistic regression models with the outcome of overweight and obesity
 284 obtained odds ratios and confidence intervals of no strong statistical support for a difference
 285 in odds between mode of delivery (Figure 4).



286
287 **Figure 4. Results obtained from logistic regression models, for the association**
288 **between categorised BMI z-score in children delivered by caesarean section, at 4-5**
289 **years old, from White British and Pakistani ethnic groups.** Adjusted for: ethnicity,
290 maternal BMI (continuous), maternal job status and drinking alcohol during or 3 months
291 before pregnancy. Abbreviations: OR, odds ratio; CI, confidence intervals.

292 **Effect modification**

293 Stratified analysis suggests the adjusted association of mode of delivery on BMI z-score is
294 similar, irrespective of ethnicity (Figure 3). Additionally, there was no evidence of effect
295 modification from the likelihood ratio test ($p= 0.97$). When a test for effect modification was
296 performed on the categorised z-scores, similar results were obtained. There was weak
297 evidence of effect modification by ethnicity on the association between mode of delivery and
298 overweight ($p=0.14$), and no evidence for effect modification by ethnicity on the association
299 between mode of delivery and obesity ($p=0.79$).

300

301 **Discussion**

302 In this cohort study, it was found that undergoing a CS was not associated with an increased
303 risk of overweight and obesity in children, and there was no difference between ethnic
304 groups.

305 Mothers who had undergone a CS were generally of higher socio-economic status than
306 those who had vaginal deliveries; CS women were more educated, more likely to be
307 currently employed, more likely to have a stable housing situation and less likely to be
308 receiving benefits. Also, mothers who experienced CS had baseline characteristics to
309 suggest they had poorer health than mothers giving birth vaginally; CS mothers had a higher
310 mean BMI, were more likely to drink alcohol during pregnancy or 3 months before and more
311 prevalent gestational diabetes. The distribution of alcohol drinking varies vastly between
312 ethnicities, this is most likely due to religious beliefs (31). This explains the very low
313 prevalence of alcohol drinking and avoidance of smoking amongst Pakistani women.

314 The linear regression for the adjusted model offers no evidence for a difference in BMI z-
315 score between children born via CS and vaginal deliveries. The low adjusted R² value
316 suggests there are other variables which have an influence on the primary association. The
317 adjusted logistic regression models also suggest no evidence for children delivered by CS
318 having different odds of being overweight or obese, compared to children of vaginal
319 deliveries.

320 The stratified analysis and formal test for effect modification both indicate there is no
321 evidence that the association between mode of delivery and children's BMI z-scores varied
322 by ethnicity.

323 As discussed in the introduction, previous studies have varied interpretations. Two leading
324 systematic reviews suggest there is evidence that CS increases child BMI (8,9). However,
325 there were several studies which found no statistical association between mode of delivery
326 and child BMI. The findings from this paper are compatible with the latter studies mentioned.

327 Two out of three studies conducted in the UK concluded there was no 'statistical significant'
328 difference in risk of childhood overweight or obesity between modes of delivery, at 3 years
329 old (40) and 5 years old (28).

330 Furthermore, maternal BMI explained most of the observed association in this study and
331 was hence the main confounding factor. All previous studies cited here, looking at the
332 association between mode of delivery and BMI, also adjusted for this factor.

333 However, this study also differs with previous research. The other UK study found that CS
334 increased the odds of being overweight or obese, at 7 years old (19). This was a study
335 which used data from the Avon Longitudinal Study of Parents and Children (ALSPAC);
336 participants were recruited from the Avon area if they were born in 1991–1992.

337 Several confounders (child gender, gestational factors and child feeding patterns) were
338 adjusted for in the ALSPAC study but did not meet the confounding criteria (or the data were
339 unavailable) to be adjusted for in this study. There were also inconsistencies with other
340 factors adjusted for in this study compared to previous studies, such as not adjusting for
341 antibiotics during pregnancy (20). Different factors could have met the confounding criteria in
342 previous studies due to their population type, for example, by having a different BMI
343 distribution as the children were leaner.

344 Adjusting for ethnicity was not seen in previous research in the UK. Due to the large
345 proportion of Pakistani women in this BiB study, there was sufficient power to investigate
346 differences between WB and Pakistani ethnic groups, whereas this would not be possible in
347 studies like ALSPAC. As previous studies did not adjust for ethnicity, other variables could
348 have acted as confounders. Overall, the differences in study design, study population and
349 confounding adjustments could explain the inconsistent conclusions reached.

350 The large sample size used in this study allowed sufficient power to identify any meaningful
351 differences in association between BMI z-scores of two different modes of delivery.

352 Additionally, consistent statistical methodology with previous studies was used and there
353 was minimal recruitment bias due to the BiB study having a high recruitment rate of 87%.

354 As CS and vaginal deliveries are very different procedures, in theory, there was no
355 opportunity for this to be incorrectly recorded. Hence no information bias, in the form of non-

356 differential misclassification, should have occurred. Furthermore, observer bias would not
357 arise when recording the child's BMI, as nurses taking anthropometric measurements at
358 ages 4-5 were blinded to information regarding the child's mode of delivery.

359 There is evidence to suggest BMI measurements systematically underestimate childhood
360 adiposity (41). This has also been specifically investigated in South Asians with evidence to
361 suggest that BMI additionally appears to underestimate adiposity in this ethnic group.
362 Despite South Asians being generally smaller and lighter, they seem to have greater relative
363 fatness compared to white European populations (42).

364 Most of the data on covariates were collected in the baseline questionnaire, completed by
365 the mother. As data were self-reported, information bias in the form of differential
366 misclassification could have occurred which would tend the results to overestimate or
367 underestimate the true association. An example would be smoking as this is a likely factor to
368 be underreported. Underreporting could underestimate the association between mode of
369 delivery and smoking, which would have led to it not being adjusted for in the final analysis.

370 There may be residual confounding which is obscuring the true effect of mode of delivery on
371 child BMI. The low adjusted R^2 value implies other factors could have an influence on the
372 association, therefore suggesting factors which were not included in the analysis explained
373 some of the association. These could be factors such as amount of exercise feeding pattern
374 of the child or breastfeeding. It would have been desirable to have considered breastfeeding
375 as a potential confounder but there was insufficient data to assess this.

376 The study was limited by approximately 85% of the study population having missing data on
377 breastfeeding due to the data being collected in a subgroup of women who participated in
378 the BiB1000, as described in the study design section. Parity and maternal BMI was also
379 missing for 4% of the final study population. There was no evidence of any statistically
380 significant difference in the distribution of mode of delivery or child BMI z-score, when
381 comparing: those with data on parity vs those missing parity; and those with data on

382 maternal BMI vs those missing maternal BMI. Therefore, there was no evidence of selection
383 bias based on the distribution of missing data in parity and maternal BMI.

384 Additionally, these results will be generalisable to other populations with similar demography
385 to Bradford. The CS rate in Bradford is not markedly different to the national rate, and
386 obesity at 4-5 years is very slightly above the national average (10.8% in Bradford, 9.9% in
387 England, in 2019-2020) (12). The results obtained from the Pakistani population may not be
388 generalisable to other South Asian groups.

389 **Conclusion**

390 Overall CS was not associated with an increased risk of overweight or obesity in children
391 aged 4-5 years in Bradford. Neither was there a difference in association seen between
392 White British or Pakistani children. To our knowledge, this is the first study to assess this
393 association, between these ethnicities, at this age.

394 As CS deliveries are becoming more common globally and health concerns have been
395 raised, the results from this study, combined with similar studies, should be informative to
396 prospective parents and healthcare advisors.

397 Data collection within the BiB cohort should be continued to provide more reliable estimates
398 of adiposity and to allow investigation at older ages. This will enable examination of whether
399 any association exists at subsequent ages between mode of delivery and later life adiposity.

400 As there is some uncertainty around how well BMI represents child adiposity, the use of
401 body fat centile curves should be explored instead. To do this, data on fat mass analysed
402 using a DXA scanner would be needed. A DXA scanner is an extremely accurate method for
403 analysing body composition, and could be used as a gold standard in the population. The
404 sensitivity and specificity of BMI against the DXA scanner in the population can be obtained
405 and incorporated in the interpretation.

406 Further research could also investigate if the method of fetus extraction acts as an effect
407 modifier on the association between mode of delivery and BMI. Vaginal deliveries can occur
408 by: natural vaginal birth, forceps assistance or by ventouse techniques. CS can occur by:
409 emergency, elective or semi-elective delivery. These studies would need to have a large
410 sample size to power the subgroup analyses.

411 Additionally, if the mechanism for any potential association is related to the developing gut
412 microbiota, then more studies could focus on the differences between gut microbiota
413 stratified by mode of delivery and by ethnicity. This is likely to involve a genetic approach
414 when looking at the differences between the WB and Pakistani population (17).

415 **Abbreviations**

416 ALSPAC, Avon Longitudinal Study of Parents and Children; BiB, Born in Bradford; BMI,
417 body mass index; CI, confidence intervals; CS, caesarean section; DXA, Dual-energy X-ray
418 absorptiometry; NCMP, National Child Measurement Programme; OR, odds ratio; WB,
419 White British.

420

421 **Declarations**

422 **Ethics approval and consent to participate**

423 All methods were carried out in accordance with relevant guidelines and
424 regulations. Informed consent was obtained from all participants or, if subjects are under 18,
425 from a parent or legal guardian. BiB ethical approval was granted by Bradford Research
426 Ethics Committee (Ref 07/H1302/112). Participating women consented at recruitment to
427 access to their routine primary and secondary health care records. Ethical approval for the
428 current study was received from the Ethics Committee at the London School of Hygiene and
429 Tropical Medicine.

430 **Consent for publication**

431 Not applicable.

432 **Availability of data and materials**

433 Data from the BiB study is available to researchers following approval from the Executive
434 Committee (<https://borninbradford.nhs.uk/research/how-to-access-data/>)

435 **Competing interests**

436 The authors declare that they have no competing interests.

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448 **Authors' contributions**

449 ER was responsible for analysing the data, interpreting the results and writing the first draft.
450 LP had input into design of the study, analysis plan, and interpreting the results. JW and GS
451 conceived the original research idea, were responsible for facilitating data extraction from
452 the Born in Bradford cohort. All authors contributed to further iterations of the draft. All
453 authors approved the final version.

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