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The relationship between gestational weight gain, maternal upper-body subcutaneous fat changes and infant birth size: a pilot observational study amongst women with obesity

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1 **The relationship between gestational weight gain, maternal upper-body subcutaneous fat**
2 **changes and infant birth size: a pilot observational study amongst women with obesity.**

3 Abstract

4 Background: It is widely acknowledged that maternal obesity and excessive gestational weight gain
5 (GWG) are associated with increased risk of fetal macrosomia and recent studies have suggested a role
6 for the timing and composition of GWG.

7 Aims: To examine the effect of the rate of change in GWG and maternal upper-body subcutaneous fat on
8 neonatal anthropometric outcomes in a pilot observational study amongst women with obesity.

9 Study design: Expectant women with a body mass index (BMI) > 30 kg/m² at first antenatal appointment
10 were recruited at 12 weeks gestation. Maternal height, weight and skinfold thickness (SFT)
11 measurements were collected at baseline and repeated at 28 and 36 weeks gestation. Following delivery,
12 World Health Organisation (WHO)-UK infant birthweight z-scores were calculated, and infant
13 anthropometric measurements were obtained.

14 Results: The sum of upper body SFT measurements increased in mid-pregnancy (0.08 ± 0.71 mm/week)
15 and decreased in late pregnancy (-0.04 ± 1.17 mm/week). After adjustment for maternal age, BMI and
16 parity, mid- but not late- pregnancy GWG was positively associated with infant birthweight z-score
17 ($p < 0.05$), while mid- but not late-pregnancy changes in the sum of SFT were inversely associated with
18 infant birthweight z-score ($p < 0.01$).

19 Conclusions: The present study suggests that mid- rather than late-pregnancy changes in weight and
20 upper-body subcutaneous fat are associated with infant birthweight. Further research is required in
21 larger, more diverse populations to explore whether pregnancy interventions aiming to improve maternal
22 and offspring health can be personalised beyond BMI and GWG.

23 **Key words: Pregnancy, maternal obesity, body composition, skinfold thickness, birthweight.**

24

25 Abbreviations: BMI - body mass index; GWG – gestational weight gain; UK - United Kingdom; UME –
26 upper arm muscle area estimate; UFE – upper arm fat area estimate; FM – fat mass; FFM – fat free mass.

27 Introduction

28 Obesity has become a worldwide epidemic, and in the United Kingdom (UK), it is estimated that half
29 of women of childbearing age are living with overweight or obesity [1] with the prevalence of
30 maternal obesity increasing, as defined by trimester one body mass index (BMI) [2]. It is well
31 documented that maternal obesity has significant health implications for both mother and baby,
32 increasing the risk of complications during pregnancy and labour [3]. Offspring of mothers with
33 obesity are also more likely to be born large for gestational age or macrosomic [3–5] which
34 predisposes infants to adiposity and obesity during infancy and childhood [6,7]. Excessive gestational
35 weight gain (GWG) carries similar risks to maternal obesity for both maternal and neonatal
36 outcomes [8,9], and postpartum weight retention increases the risk that women will enter their next
37 pregnancy with obesity [10].

38 Due to the lack of evidence-based guidelines, the National Institute for Health and Care
39 Excellence does not currently make recommendations for GWG amongst the UK population [11]. In
40 the United States (US), the Institute of Medicine (IOM) has published recommendations for GWG,
41 which were updated in 2009 to include BMI-specific guidelines [12]. Although the IOM
42 recommendations were designed for the US population, the recommendations are largely based on
43 evidence derived from the US and Europe, thus, the IOM recommendations have been adopted in
44 many other countries worldwide and are widely reported in the literature [13,14]. As well as total
45 GWG, the IOM recommend “normal” weekly rates of GWG for the second and third trimesters.

46 Recent studies conducted amongst women have tended to observe stronger positive
47 associations between GWG in the second trimester [15,16] or early GWG (before the end of the
48 second trimester) and infant birth size outcomes [17–19]. Studies examining the relationship
49 between maternal body composition assessed via bioelectrical impedance analysis and infant
50 birthweight have tended to observe a positive association between maternal fat-free mass (FFM),
51 but not maternal fat mass (FM) [19–21]. With the exception of one study [17], these studies were
52 conducted amongst women across all BMI ranges and tend to report estimates of FM or FFM at

53 single time-points rather than changes during pregnancy, which makes it difficult to establish
54 trimester-specific recommendations, particularly for women with obesity. A 2020 National Academy
55 of Medicine discussion paper focusing on GWG amongst women with obesity highlights that many
56 studies report total GWG over pregnancy, rather than patterns of GWG and correlations between
57 GWG and fetal growth [22]. A recent study conducted amongst 72 women with obesity observed
58 that individual differences in total GWG were predominantly explained by changes in FM, as
59 assessed by air displacement plethysmography, with gains in FM significantly lower amongst women
60 with Class III obesity, than amongst those with Class I and II [23]. In addition, when examined by
61 trimester, FM was found to increase in the second trimester, and decrease in the third, whilst GWG
62 in the third trimester was attributed to FFM accumulation and fetal growth. However, this study did
63 not examine the relationship between these patterns of GWG, FM and FFM accrual and infant birth
64 size and there appears to be a lack of observational studies conducted amongst women with obesity
65 that examine these associations. The time periods examined also vary considerably between studies,
66 with some looking at early versus late pregnancy, and others looking at trimesters, which makes
67 comparison difficult. Studies examining rates of GWG at frequent assessments are therefore useful
68 in order to increase our understanding of the importance of GWG during different stages of
69 pregnancy, facilitate comparison of GWG amongst pregnancies or varying durations, and to enable
70 the development of transferable recommendations.

71 The aim of the present study was therefore to identify whether there is a relationship
72 between trimester-specific rates of GWG or upper-body skinfold thickness (SFT) measurements and
73 infant birthweight and anthropometrics at birth. SFT measurements were chosen to assess
74 subcutaneous fat stores, which traditionally accumulate up to the end of the second trimester and
75 are subsequently mobilised in the third trimester to support maternal metabolism and rapid fetal
76 growth. In addition, callipers are a simple, quick, portable and cost-effective tool that could be used
77 in addition to weighing scales by health professionals caring for pregnant women, enabling
78 personalised care beyond BMI [24]. SFT measurements could also be used as an additional outcome

79 measure to assess the success of future pregnancy interventions aiming to reduce GWG and infant
80 macrosomia if a relationship is observed between changes in maternal upper body subcutaneous fat
81 and infant outcomes. As women with obesity are at increased risk of delivering a LGA or macrosomic
82 baby, but at lower risk of multiple pregnancy complications than women with Class III obesity [3],
83 the study focuses on women with Class I and Class II obesity (pre-pregnancy BMI ≥ 30 kg/m² and <40
84 kg/m²). To our knowledge, this is the first study to report the association between GWG, changes to
85 maternal upper-body SFT and infant birth size amongst women with obesity in the UK. It is
86 therefore difficult to form a hypothesis in terms of how changes in maternal upper-body SFT will
87 affect infant birth size, however, based on previous work examining changes in maternal weight, FM
88 and FFM in women of all weights, we hypothesise that mid-pregnancy GWG will be more strongly
89 associated with infant birthweight than late-pregnancy GWG, and that we will observe reductions in
90 upper body subcutaneous fat in late pregnancy.

91 Methodology

92 [Recruitment of women](#)

93 Women aged between 18 and 40 years of age, with a BMI ≥ 30 and <40 kg/m² at booking and
94 pregnant with a singleton pregnancy were eligible to take part in the study. Women meeting
95 inclusion criteria were identified from their antenatal booking notes and approached by the
96 researcher at their 12 week dating scan. Ethical approval was obtained from the NHS Health
97 Research Authority National Research Ethics Service and local Research and Development approval
98 was obtained from University Hospitals Plymouth NHS Trust.

99 Following recruitment, verbal and written informed consent were obtained from women,
100 and the first study visit occurred between 12 and 14 weeks gestation. Further visits occurred at the
101 end of the second trimester at approximately week 28 of gestation (visit 2), and at the end of the
102 third trimester at approximately week 36 of gestation (visit 3). A single researcher performed all
103 measurements at all study visits in order to reduce inter-observer error.

104 Maternal anthropometric measurements

105 In order to examine GWG throughout pregnancy, maternal weight was measured at each visit using
 106 the same digital scales for each woman throughout the duration of the study. GWG was recorded as
 107 a simple difference between weight at each study visit to give a crude value for GWG in each
 108 trimester and a ‘total’ GWG for the study duration.

109 Weekly rates of GWG were calculated for each woman, based on the difference in weight between
 110 study visits, divided by the number of weeks (and days) between visits. Reporting GWG in this way
 111 accounts for gestation, and facilitates comparison against the IOM guidelines (Table 1). Rates of
 112 GWG were calculated for the second trimester (between visits 1 and 2, defined as mid-pregnancy
 113 GWG), the third trimester (between visits 2 and 3, defined as late pregnancy GWG) and over the
 114 study duration (between visits 1 and 3, defined as total pregnancy). Women were further classified
 115 as achieving ‘insufficient’, ‘adequate’ or ‘excessive’ GWG according to their rate of GWG between
 116 each of these time points according to IOM guidelines (Table 1) [12].

117 *Table 1 IOM recommendations for total and rate of weight gain during pregnancy, by pre-pregnancy BMI*

	Total Weight gain	Rate of weight gain, 2nd and 3rd trimester
Pre-pregnancy BMI	Range in kg	Mean (range) in kg / week
Underweight (<18.5 kg/m²)	12.5 – 18.0	0.51 (0.44-0.58)
Normal weight (18.5 – 24.9 kg/m²)	11.5 – 16.0	0.42 (0.35–0.50)
Overweight (25.0 – 29.9 kg/m²)	7.0 – 11.5	0.28 (0.23-0.33)
Obese (≥30 kg/m²)	5.0 – 9.0	0.22 (0.17-0.27)

118
 119 Maternal upper-body SFT of the biceps, triceps and subscapular were assessed at each
 120 anthropometric visit according to the methods described by Kannieappan et al. [25] and the
 121 International Society for the Advancement of Kinanthropometry (ISAK) [26] using Harpenden
 122 callipers (British Indicators, Sussex, England) by a single researcher in order to minimise inter-

123 observer error. A full set of all three SFT measurements was completed in order to reduce the
124 effects of skinfold compressibility prior to repeating a second measurement at each site. If the
125 difference was greater than 7.5%, a third measurement was taken according to ISAK
126 recommendations. SFT for each site was reported in mm as the mean of two measurements, or the
127 median of three measurements [26]. In addition, SFT measurements were reported as the sum of all
128 three measurements rather than entered into an equation as recommended by numerous authors
129 to assess changes in body composition over time and to reduce systematic error associated with
130 equations, which are particularly difficult to validate in a pregnant population [27,28]. Rate of
131 change to the sum of these three SFT measurements over the study duration, as well as in mid-
132 pregnancy and late-pregnancy were calculated (in mm/week) to adjust for length of gestation for
133 each woman.

134 [Infant outcomes](#)

135 Information about the infants was collected from hospital notes, or through measurements made by
136 the researcher. This included gestational age at delivery, method of delivery, the incidence of any
137 complications, infant gender, birthweight and head circumference. Birthweight and head
138 circumference were used to calculate the z-scores from UK-WHO reference values for term infants
139 [29,30] using the LMS method [31] (LMS Growth Programme v2.77, Medical Research Council, UK)
140 which adjusted for infant gender. Crown-heel length was measured by the researcher using a mobile
141 measuring mat (Seca 210, Hamburg, Germany). This measurement was taken as soon after delivery
142 as possible and recorded to the nearest 5 mm. Length z-scores were also calculated from UK-WHO
143 reference values for term infants [29,30] using LMS software and were adjusted for infant gender
144 and age at assessment.

145 Infant anthropometric measurements were taken as close to birth as possible, in most
146 instances within 72 hours of delivery. Where this was not possible, for example, for infants who
147 spent longer than this on the neonatal intensive care, or transitional care units, measurements were

148 used after adjusting them for age. A model that calculated an infant upper arm fat area estimate
149 (UFE) and upper arm muscle area estimate (UME) was used. The equations are based on mid-upper
150 arm circumference and triceps skinfold, and has been previously validated against magnetic
151 resonance imaging in children [32]:

152 **$TUA = C^2 / (4\pi)$**

153 **$UFE = C \times (TS/2)$**

154 **$UME = TUA - UFE$**

155 TUA = total upper arm area; UFE = upper arm fat area estimate; UME = upper arm muscle area
156 estimate; C = mid upper arm circumference, TS = triceps skinfold (mm).

157 [Statistical analysis](#)

158 All data was entered into and analysed using SPSS (Statistics Package for the Social Sciences) for
159 Windows version 21 (IBM, Chicago USA). The level of significance was set to a probability $p < 0.05$ for
160 all statistical tests performed, and unless otherwise stated, data were presented as means \pm
161 standard deviation (SD).

162 Continuous outcome measures were inspected for normality and if this assumption was
163 met, parametric tests were performed. Pearson's correlation coefficients (r) were run to assess the
164 relationships between maternal changes in upper body subcutaneous fat and GWG. Multiple
165 regression was used to evaluate the extent to which maternal rates of GWG and SFT changes over
166 pregnancy influence infant birthweight z-scores, UFE and UME after adjustment for maternal age,
167 booking BMI and parity, and in the case of UFE and UME, also adjusted for infant sex and gestational
168 age. For all models there was independence of residuals, as assessed by a Durbin-Watson statistic of
169 approximately 2.0, homoscedasticity, as assessed by visual inspection of a plot of studentized
170 residuals versus unstandardized predicted values and there was no evidence of multicollinearity, as
171 assessed by tolerance values greater than 0.1. As the current study is the first, to the author's
172 knowledge, to examine the effect of the rate of change in upper body SFT measurements on

173 birthweight and infant body composition, there was no data on which to base an *a priori* power
 174 calculation. However, posthoc power calculations show that the large effect sizes observed for the
 175 multiple regression analyses reached 99% power, and thus, statistical analysis was adequately
 176 powered.

177 Results

178 A total of 75 women gave their consent to participate in the study, which was 31% of women
 179 approached. All women in the study identified their ethnicity as White Caucasian.

180 Data was collected for 75 women at visit one (12-14 weeks gestation), 65 women at visit two
 181 (28 weeks gestation), and 59 women at visit three (36 weeks gestation), with a total of 16 women
 182 lost to follow up between the first and last study visit. Womens' ages ranged from 19 years to 40
 183 years, with a mean age of 29.8 ± 4.8 years and mean BMI was 33.0 ± 1.9 kg/m². There were no
 184 significant differences in maternal descriptive characteristics nor birth outcomes obtained from
 185 notes between women completing the study and those who were lost to follow up, data not shown.

186 Anthropometric measurements collected at each study visit are shown in Table 2.

187 *Table 2 Maternal anthropometric measurements*

	Visit 1 (n=75)	Visit 2 (n=65)	Visit 3 (n=59)
Weight (kg)	89.7 ± 8.7	95.1 ± 9.2	97.0 ± 9.9
Trimester-specific rate of GWG (kg/week)	-	0.3 ± 0.2	0.3 ± 0.4
Total pregnancy rate of GWG (kg/week)	-	-	0.3 ± 0.3
Proportion of women gaining in excess of IOM guidelines, n (%)	-	38 (59)	31 (54)
Arm circumference (cm)	35.2 ± 2.7	34.8 ± 2.8	34.3 ± 3.0
Triceps SFT (mm)	29.6 ± 5.2	30.4 ± 5.9	28.5 ± 5.7
Biceps SFT (mm)	18.9 {17.6-20.3} [†]	19.1 ± 6.0	18.0 ± 6.4
Subscapular SFT (mm)	34.2 ± 7.5	35.4 ± 9.0	34.5 ± 8.5
Sum of SFT (mm)	83.7 ± 13.6	85.0 ± 15.0	81.0 ± 17.1
Trimester-specific rate of change in SFT (mm/week)	-	0.1 ± 0.7	-0.1 ± 1.2
Total pregnancy rate of change in SFT (mm/week)	-	-	-0.1 ± 0.7

Data are mean ± SD unless otherwise stated.

[†]Mean calculated by back-transformation {CI}

Gestational weight gain (GWG), Institute of Medicine (IOM), Skinfold thickness (SFT).

188
 189 Table 2 shows that mean rate of GWG in mid-pregnancy, late pregnancy, and in total over
 190 pregnancy was 0.33 ± 0.23 , 0.29 ± 0.40 and 0.32 ± 0.26 kg/week, respectively. The proportion of
 191 women gaining in excess of IOM guidelines was 59 and 54% in mid- and late-pregnancy, respectively.
 192 In addition, Table 3 shows GWG class of obesity, and the proportion of women gaining weight in
 193 excess of the IOM guidelines. Mid-pregnancy GWG and rate of GWG were significantly higher
 194 amongst women in Class I (5.5 ± 0.5 kg and 0.4 ± 0.1 kg/week) than amongst women in Class II ($2.8 \pm$
 195 1.2 kg; $p = 0.026$ and 0.2 ± 0.1 kg/week; $p=0.024$), whereas GWG in late-pregnancy and over total
 196 pregnancy were not significantly different between women in the two classes of obesity ($P>0.05$).

197 *Table 3 Gestational weight gain by obesity class.*

	Class I Obesity (n=46)	Class II Obesity (n=11)
Mid-pregnancy GWG (kg)	5.5 ± 0.5	2.8 ± 1.2
Mid-pregnancy rate of GWG (kg/week)	0.4 ± 0.1	0.2 ± 0.1
Proportion of women gaining in excess of IOM guidelines in mid-pregnancy, n (%)	31.0 (67.4)	4.0 (36.4)
Late-pregnancy GWG (kg)	2.4 ± 0.5	2.1 ± 1.0
Late-pregnancy rate of GWG (kg/week)	0.3 ± 0.1	0.3 ± 0.1
Proportion of women gaining in excess of IOM guidelines in late-pregnancy, n (%)	26.0 (56.5)	5.0 (45.5)
Total GWG (kg)	8.1 ± 1.0	4.9 ± 1.9
Total pregnancy rate of GWG (kg/week)	0.3 ± 0.1	0.2 ± 0.1
Proportion of women gaining in excess of IOM guidelines over total pregnancy, n (%)	28.0 (60.1)	5.0 (45.5)

Data are mean \pm SD unless otherwise stated.

Gestational weight gain (GWG), Institute of Medicine (IOM), Skinfold thickness (SFT).

198
 199 The rate of GWG was significantly and positively associated with the rate of SFT changes in
 200 mid-pregnancy ($r=0.467$), late-pregnancy ($r=0.478$) and over total pregnancy ($r=0.609$; all $p<0.01$).
 201 Changes in SFT were highly variable over pregnancy for the study population with an overall trend

202 for a reduction in upper body subcutaneous fat between early and late pregnancy, although this did
 203 not reach statistical significance ($p=0.071$).

204 Information concerning the delivery of infants, was available for 74 infants, of which three
 205 were pre-term (<37 weeks gestation) and excluded from analysis. Infant anthropometric
 206 measurements acquired from hospital notes and from the researcher's home visit are shown in
 207 Table 4.

208 *Table 4 Infant outcomes (n=71)*

	n	Mean	SD
Gestation length (days) [†]	71	275.0	13.0
Infant gender male, n (%)	71	41 (58)	-
Vaginal delivery, n (%)	71	45 (63)	-
Birthweight, g	71	3497.0	461.0
Birthweight, z-score	71	0.1	0.9
Head circumference, cm [†]	55	35.0	2.0
Head circumference, z-score	55	0.4	1.3
Crown-heel length, cm	56	50.8	2.1
Crown-heel length, z-score	56	0.1	1.0
Infant arm circumference, cm [†]	56	10.0	1.5
Triceps skinfold thickness, mm	56	6.7	1.8
UME, cm ²	56	445.5	105.1
UFE, cm ²	56	334.4	108.2

†Median (IQR)

Standard deviation (SD), upper arm area muscle estimate (UME), upper arm area fat estimate (UFE).

209

210 A multiple regression was used to evaluate the extent to which maternal rates of GWG and
 211 changes in SFT in mid- and late-pregnancy influence infant birthweight z-scores, UME and UFE after
 212 adjustment for maternal age, booking BMI and parity. The models examining UFE and UME were
 213 additionally adjusted for infant sex and gestational age, which were already accounted for in the
 214 birthweight z-scores. The models statistically significantly predicted birthweight z-score ($p=0.016$)
 215 and UFE ($p = 0.017$), but not infant UME, which didn't quite reach statistical significance ($p =0.055$).
 216 As shown in Table 5, mid- but not late-pregnancy GWG was significantly and positively associated
 217 with infant birthweight z-score, while mid- but not late-pregnancy change in the sum of SFT was

218 significantly and inversely associated with infant birthweight z-score. Mid-pregnancy GWG was also
 219 positively associated with infant UFE, however, for late pregnancy this relationship was reversed.

220 *Table 5 Multiple regression coefficients.*

	Birthweight z-score (n=56)			UFE (n=53)		
	β	95% CI	p	β	95% CI	p
Mid-pregnancy GWG, kg/week	0.47	0.39 – 3.32	0.014	0.49	53.03 – 384.3	0.011
Mid-pregnancy change in the sum of SFT, mm/week	-0.50	-1.11 - -0.28	0.001	-0.28	-92.36– 3.39	0.068
Late-pregnancy GWG, kg/week	0.16	-0.39 – 1.12	0.329	-0.37	-187.68 - -13.06	0.025
Late-pregnancy change in the sum of SFT, mm/week	-0.24	-0.45 – 0.05	0.107	0.19	-10.87– 47.71	0.212

Multiple regression adjusted for maternal body mass index, parity, age, infant sex* and gestational age*

*UFE model only.

221

222 Discussion

223 This study is the first to our knowledge that has examined the impact of trimester-specific rates of
224 GWG and changes in upper-body skinfold measurements on infant anthropometric measurements
225 amongst women with obesity in the UK. Our findings suggest that mid- rather than late-pregnancy
226 changes in weight and upper-body subcutaneous fat are associated with infant birthweight and
227 upper body fat, suggesting that the second trimester may be an opportune window for pregnancy
228 interventions aiming to improve pregnancy and infant outcomes for women with obesity.

229 Mid-pregnancy GWG was positively associated with birthweight z-score, while no
230 association was observed for late-pregnancy GWG. These observations are consistent with others in
231 the literature. For example, Farah et al. [19] observed a positive correlation between GWG before
232 the third trimester and infant birthweight, with no association reported for GWG during the third
233 trimester in a cohort of 184 non-diabetic women in Ireland. Hivert et al [15] observed a positive
234 association between rate of GWG in all three trimesters and birthweight z-score amongst 979
235 mother-child pairs from the Project Viva cohort in the USA, with the largest effect size observed for
236 second trimester GWG, while Widen et al [16] observed that high rate of GWG, as defined by
237 tertiles, in the second trimester only was associated with higher infant birthweight and length
238 amongst 156 women in another USA cohort.

239 In terms of changes in maternal adiposity, there was large variation in the accumulation of
240 upper body fat amongst women in the cohort, and only mid-pregnancy changes in maternal upper-
241 body skinfold measurements were associated with infant birthweight, in an inverse direction. This is
242 in contrast to findings from Dodd et al [33] who estimated maternal FM from the same three sites
243 measured in the present study amongst 1582 overweight women in South Australia, and did not
244 observe any significant associations between total GWG, maternal percentage body fat, nor
245 individual SFT measurements and infant birthweight. Hediger et al [34] observed an inverse
246 relationship between change in triceps, but not subscapular SFT and infant birthweight, but this was

247 in late pregnancy, in contrast to the findings in the current study, where late pregnancy changes in
248 SFT did not contribute to the regression model. The study by Hediger et al. [34] was conducted in
249 608 women of all weights, and is more consistent with previous literature that suggests pregnant
250 women accrue subcutaneous fat in early and mid-pregnancy, and tend to experience a decrease in
251 SFT measurements in trimester three, when fat stores are mobilised to support rapid fetal growth
252 [35,36]. However, studies conducted amongst women with obesity suggest that subcutaneous fat
253 gains tend to be less than for women with a lower BMI [37,38]. A recent study conducted by Most et
254 al. [23] found individual differences in GWG amongst 54 women with obesity to be largely explained
255 by changes in maternal FM, with gains in FM significantly higher for women with Class I and II obesity
256 (who gained FM) compared with women with Class III obesity (who lost FM). Straughen and
257 colleagues [39] observed that subcutaneous fat declines measured by ultrasound in women with
258 overweight or obesity were more rapid from early through to late pregnancy than amongst women
259 with a healthy BMI. In a study conducted by Misra and Trudeau (2011), circulating leptin
260 concentration at the start of pregnancy was 1.8 times higher for women with obesity compared with
261 healthy weight women, but by the end of pregnancy it was just 1.2 times higher. These findings
262 suggest that metabolic adaptations to pregnancy amongst women with obesity are different to those
263 observed amongst healthy weight women and may explain our observation of an inverse
264 relationship between mid-pregnancy subcutaneous fat changes and infant birth size, despite
265 observing a positive relationship between mid-pregnancy GWG and infant birthweight. As already
266 discussed women tend to experience a decrease in subcutaneous fat in the final trimester, but based
267 on the observations from the present study, in combination with observations from the literature
268 examining women with obesity discussed above, it is possible that fat stores are perhaps mobilised
269 earlier, thus perhaps explaining, in part, the observations in the present study. In addition, the
270 positive relationship between mid-pregnancy GWG and birthweight may be driven by changes to
271 weight that exclude upper-body subcutaneous fat. For example increases in FFM, which would

272 incorporate TBW, the foetus, placenta and amniotic fluid, or changes to maternal FM at other
273 locations, such as visceral adipose tissue and breast tissue.

274 It is also important to acknowledge, that although it was a strength of the current study that
275 one researcher took all SFT measurements to reduce inter-observer error, this would not be the case
276 in clinical practice, if multiple health professionals were taking measurements, nor in larger research
277 studies, with multiple researchers. Further studies using methods of assessment that are able to
278 distinguish between the maternal and fetal unit are warranted, although achievement of this is likely
279 to require the use of four-compartment models, which is not generally feasible in larger cohort
280 studies. Widen and Gallagher [24] suggest further validation of portable methods such as BIA is
281 required with revised equations for use in pregnancy to account for changes in TBW and FFM
282 hydration during pregnancy, that can be effectively used in women from pre- to post-partum.

283 With regards to infant adiposity, higher mid-pregnancy GWG predicted infant UFE, while in
284 late pregnancy, an inverse relationship was observed between GWG and UFE. The model did not
285 significantly predict infant UME. These findings agree with others in the literature to an extent, with
286 one study reporting that women gaining 'excessive' weight in early pregnancy gave birth to babies
287 with significantly greater fat mass assessed via total body electrical conductivity, than those born to
288 women gaining 'excessive' GWG in late pregnancy [18]. In keeping with the present study, the
289 Norwegian STORK study used SFT measurements to assess infant subcutaneous fat and observed
290 that mid-pregnancy rate of GWG (15-28 weeks gestation) was the strongest independent predictor
291 of infant sum of SFT [40]. However, unlike the present study, no proxy for infant FFM was used, and
292 late-pregnancy GWG was not reported.

293 Although findings from the current study and previous work suggest that infant birth size
294 outcomes may be driven by changes in maternal weight and body composition, it is not clear
295 whether advising women to adhere to IOM recommendations will positively influence infant birth
296 size, particularly amongst women with obesity. This could explain why lifestyle interventions that
297 successfully reduce GWG do not tend to observe significant reductions in infant birthweight [41,42].

298 A recent observational study, conducted in Ireland, suggests that when infant birthweight is
299 subtracted from total GWG, the positive correlation between GWG and birthweight no longer exists
300 [43]. The authors argue that modifying GWG and maternal adiposity in women with obesity during
301 pregnancy is therefore unlikely to influence the growth of the baby, and that focus should move
302 from restricting GWG to encouraging a varied, balanced diet. However, it is important to note that
303 even if modifying GWG during pregnancy cannot alter infant birthweight, excess GWG is associated
304 with increased risk of other adverse outcomes [8,44] such as postpartum weight retention, which
305 increases the risk of women entering subsequent pregnancies with obesity, as well as their risk of
306 associated chronic diseases [10].

307 The current study is not without its limitations, and the primary limitation of the study is the
308 sample size of 75 women all of white ethnicity. This is due to the location of the study hospital,
309 where 95.4% of the population identify themselves as belonging to this group [45]. Future studies
310 investigating the relationship between maternal subcutaneous body fat changes and infant birth size
311 need to be conducted in other areas of the UK amongst women with obesity to determine whether
312 similar patterns are observed amongst more diverse populations. In addition, women were recruited
313 at the end of their first trimester, at their 12 week dating scan, and therefore, we were unable to
314 collect information concerning GWG and body composition changes from conception to week 12
315 gestation. Although some studies have indicated that minimal GWG occurs in the first trimester [12],
316 physiological changes such as growth of the uterus and breast tissue and plasma volume expansion
317 begin early in pregnancy. Studies examining maternal body composition changes early in pregnancy
318 are scarce, due to the difficulties recruiting women early in their pregnancy before their pregnancy
319 has been confirmed via dating scan, which occurs in the UK at 12 weeks gestation. Therefore,
320 although the majority of published studies appear to report stronger associations between mid-late
321 pregnancy GWG and FFM, observational studies examining changes in GWG and maternal body
322 composition in cohorts of women from pre-conception through to delivery are required in larger,

323 more ethnically diverse populations in order to confirm this, particularly amongst women with
324 obesity, where relatively little is known about very early changes in maternal body composition.

325 Despite methodological differences between the studies described above and limitations of
326 the present study, maternal changes in weight and body composition appear to play an important
327 role in the predication of infant birthweight and adiposity, especially in mid-pregnancy. To our
328 knowledge, this is the first study to have examined the relationship between the rate at which
329 maternal upper body subcutaneous fat changes over pregnancy in women with obesity and
330 highlights the need for a more personalised approach beyond BMI and total GWG to optimise
331 outcomes for mother and baby, particularly in the second trimester. However, the findings from this
332 pilot study amongst women with obesity, in combination with those in the literature amongst
333 women of all weights, do not consistently support the use of SFT measurements in addition to the
334 monitoring of GWG to assess risk of adverse birth size outcomes, but it is clear that more research is
335 warranted examining the relationship between GWG and changes in maternal body composition
336 amongst women with obesity. Future studies should examine the relationship between the
337 composition of GWG and infant body composition using body composition assessment methods that
338 can distinguish between the maternal and foetal unit, that are portable, and that can be used in a
339 clinical setting throughout pregnancy at frequent intervals.

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