Recent changes in the pre- and postnatal growth trajectories of Offspring from Gestational Diabetic Mothers



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Introduction and objectives

Increased risk of short and longterm metabolic complications among Offspring of gestational diabetic mothers (OGDM)^{1,2}

Objectives To explore the growth trajectories of 'recent' and 'earlier' OGDM in Cambridge Baby Growth Study

(CBGS; UK population)

 To investigate whether 'recent' OGDM still exhibit a classical phenotype, with macrosomia and higher adiposity

Changing diagnostic criteria and subsequent more intensive management of gestational diabetes mellitus (GDM)³

> Some recent studies suggest that birth size of OGDM may be normalising^{4,5}



Methods Groups **GDM** was defined using •Visits: 0, 3, 12, 24 the same IADPSG criteria months on pregnancy OGTT³: Anthropometry •Controls (N=876) born in •>5.1 mmol/L at 0 minutes, outcomes: weight, 2001-2009, recruited antenatally length, and skinfolds •>10.0 mmol/L at 60 thicknesses (reflecting 'Earlier' OGDM (N=122) minutes, or born in 2001-2009, adiposity) •>8.5 mmol/L at 120 recruited antenatally minutes 'Recent' OGDM (N=98) Same study born in 2011-2015, recruited postnatally design **GDM Criteria** All from single maternity unit. GDM criteria were retrospectively applied.

Table 1 Maternal demographics

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	Controls	'Recent' OGDM	'Earlier' OGDM
	(Total N=876)	(Total N=122)	(Total N=98)
Age at birth (years)	33.4 <u>+</u> 4.2	33.6 <u>+</u> 5.1	33.4 <u>+</u> 4.4
Caucasian ethnicity	96%	76%*	98%
Index of multiple deprivation (IMD)	8.9 <u>+</u> 3.3	11.3 <u>+</u> 6.8**	9.1 <u>+</u> 3.6
Primiparous pregnancy	48%	52%	37%*
Height (cm)	166.1 <u>+</u> 7.2	162.7 <u>+</u> 6.8**	165.8 <u>+</u> 6.9
Pre-pregnancy BMI (kg/m²)	24.0 <u>+</u> 4.4	27.0 <u>+</u> 6.3**	26.6 <u>+</u> 5.6**
Fasting venous glucose (mmol/l)	4.2 <u>+</u> 0.3	4.8 <u>+</u> 0.8**	5.3 <u>+</u> 1.1**
60 minutes venous glucose (mmol/l)	6.5 <u>+</u> 1.4	10.6 <u>+</u> 1.5**	9.2 <u>+</u> 2.1**
Smoking during pregnancy	3.8%	9.4%*	8.2%*

Table 2. Birth characteristics

Controls	'Recent' OGDM	'Earlier' OGDM
(Total N=874)	(Total N=122)	(Total N=98)
40.0 <u>+</u> 1.3	38.9 <u>+</u> 0.9**	39.5 <u>+</u> 1.4**
28%	42%**	40%*
52%	54%	53%
3.523 <u>+</u> 0.481	3.303 <u>+</u> 0.472**	3.632 <u>+</u> 0.588
0.07 <u>+</u> 0.93	0.10 <u>+</u> 1.01	0.55 <u>+</u> 1.13**
51.5 <u>+</u> 2.4	50.0 <u>+</u> 2.0**	51.3 <u>+</u> 2.7
-0.05 <u>+</u> 0.93	-0.07 <u>+</u> 0.94	0.22 <u>+</u> 0.97*
25.9 <u>+</u> 3.2	26.3 <u>+</u> 2.7	26.7 <u>+</u> 3.2
24.6 <u>+</u> 6.0	20.0 <u>+</u> 3.6**	26.0 <u>+</u> 6.3
0.03 <u>+</u> 0.86	-0.41 <u>+</u> 0.61**	0.31 <u>+</u> 0.85*
15%	7%*	27%**
5%	4%	2%
	(Total N=874) 40.0 ± 1.3 28% 52% 3.523 ± 0.481 0.07 ± 0.93 51.5 ± 2.4 -0.05 ± 0.93 25.9 ± 3.2 24.6 ± 6.0 0.03 ± 0.86 15%	(Total N=874)(Total N=122) 40.0 ± 1.3 $38.9 \pm 0.9**$ $28%$ $42%***$ $52%$ $54%$ 3.523 ± 0.481 $3.303 \pm 0.472**$ 0.07 ± 0.93 0.10 ± 1.01 51.5 ± 2.4 $50.0 \pm 2.0**$ -0.05 ± 0.93 -0.07 ± 0.94 25.9 ± 3.2 26.3 ± 2.7 24.6 ± 6.0 $20.0 \pm 3.6**$ 0.03 ± 0.86 $-0.41 \pm 0.61**$ $15%$ $7%*$

Values are mean \pm SD, or %

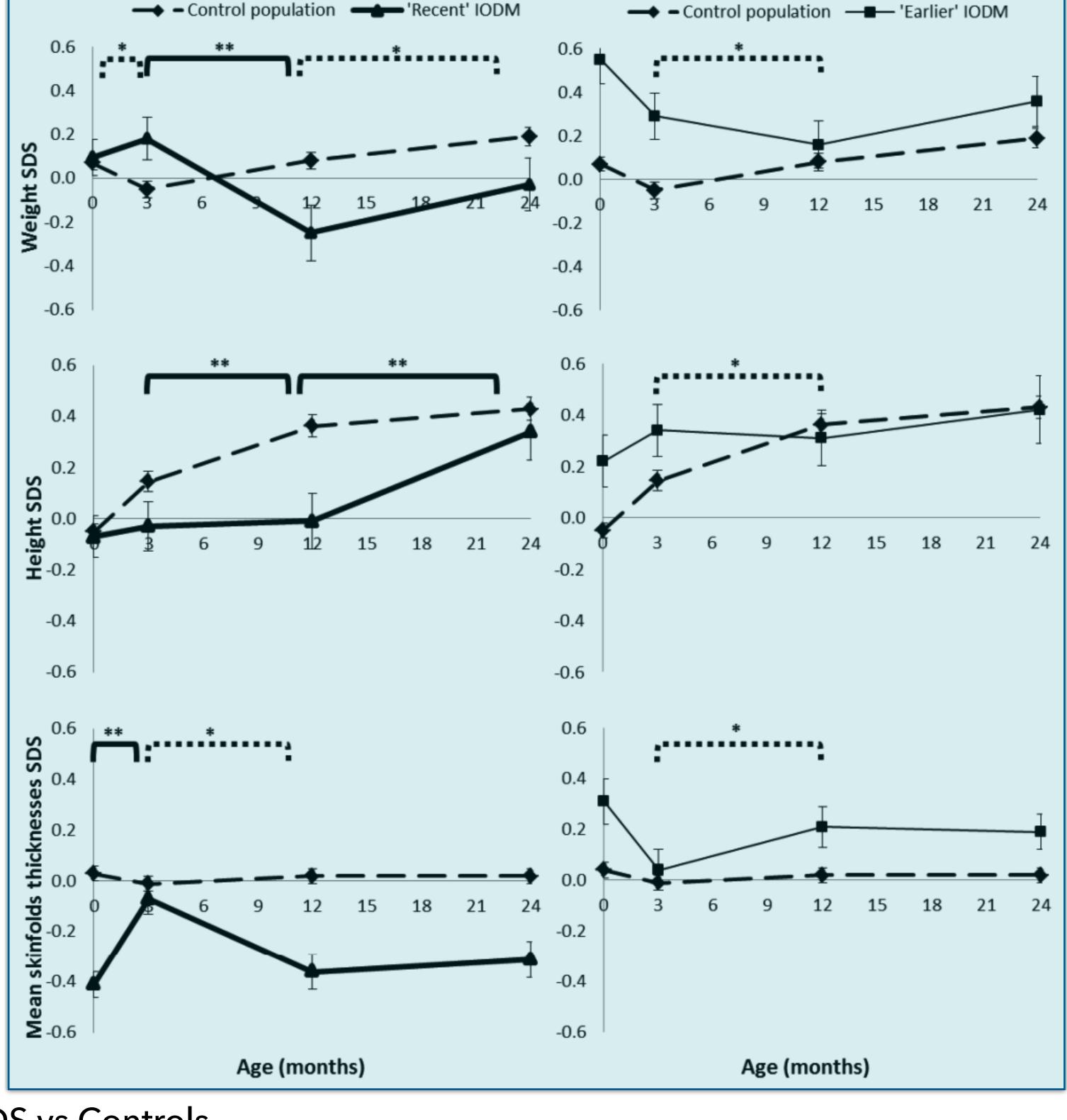
SDS, standard deviation score (for weight and length are calculated using the UK 1990 reference, for skinfolds using internal references)

- SGA, small for gestational age
- *p<0.05 vs. control group **p<0.005 vs. control group
- Weight-, length-, and mean skinfolds-SDS values are adjusted for gestational age, sex and postnatal age at measurement

Note: in control population, number of subjects for length and skinfolds measurements is 573

Results

Figure 1. Weight, length and skinfolds growth trajectories of 'recent' or 'earlier' OGDM compared to controls from birth to 2 years. Plotted values are mean \pm SEM, adjusted for sex, gestational age (birth and 3 months only), and age at measurement. Horizontal bars indicate statistically significant differences between OGDM and control groups for the displayed growth periods (* and dotted bar: p<0.05; ** and solid bar: p<0.001).



At birth:

- 'Earlier' OGDM had: higher weight-, length-, and mean skinfolds-SDS vs Controls
- 'Recent' OGDM had: similar birth weight- and length- SDS to controls, but lower mean skinfolds SDS

After birth:

- 'Earlier' OGDM had: lower gains in weight and length until 12 months
- 'Recent' OGDM had: higher weight- and skinfolds gains until 3 months, followed by lower gains in those parameters from 3-12 months, compared to controls. At 24 months, 'recent' OGDM remained less adipose than controls. At all time points 'recent' OGDM had lower growth measurements than 'earlier' OGDM.

Conclusions

'Recent' OGDM showed very different growth trajectories to the 'earlier' group: normalisation of weight and reduced adiposity at birth, followed by initial rapid weight gain but subsequent reduced growth postnatally. While avoidance of large size at birth may be advantageous, the longer-term health implications of these changing growth trajectories are uncertain.

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