ORIGINAL ARTICLE

What accounts for ethnic differences in newborn skinfold thickness comparing South Asians and White Caucasians? Findings from the START and FAMILY Birth Cohorts

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OBJECTIVE: South Asians are a high-risk group for type 2 diabetes and coronary heart disease. We sought to determine ethnic differences in newborn adiposity comparing South Asians (SA) to White Caucasians (Whites).
METHODS: Seven hundred ninety pregnant women (401 SA, 389 Whites) and their full-term offspring from two birth cohorts in Canada were analyzed. Pregnant women completed a health assessment including a 75-g oral glucose tolerance test to assess for dysglycemia. Birthweight, length, waist and hip circumference, and triceps and subscapular skinfold thickness (a surrogate measure of body adiposity) were measured in all newborns. Multivariate regression was used to identify maternal factors associated with newborn skinfold measurements.
RESULTS: South Asian women were younger (30.1 vs 31.8 years, $P < 0.001$), their prepregnancy body mass index was lower (23.7 vs 26.2, $P < 0.0001$) and gestational diabetes was substantially higher (21% vs 13%, $P = 0.005$) compared with Whites. Among full-term newborns, South Asians had lower birthweight (3283 vs 3517 g, $P = 0.0001$), had greater skinfold thickness (11.7 vs 10.6 mm; $P = 0.0001$) and higher waist circumference (31.1 vs 29.9 cm, $P = 0.0001$) compared with Whites. Risk factors for newborn skinfold thickness included South Asian ethnicity (standardized estimate (s.e.): 0.24; $P < 0.0001$), maternal glucose (s.e.: 0.079; $P = 0.04$) and maternal body fat (s.e.: 0.14; $P = 0.0002$).
CONCLUSIONS: South Asian newborns are lower birthweight and have greater skinfold thickness, compared with White newborns, and this is influenced by maternal body fat and glucose. Interventions aimed at reducing body fat prior to pregnancy and gestational diabetes during pregnancy in South Asians may favorably alter newborn body composition and require evaluation.

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INTRODUCTION

People who originate from the Indian subcontinent, known as South Asians, suffer a high prevalence of type 2 diabetes and cardiovascular disease.[1,2] Although both sociocultural and genetic factors have been implicated as potential explanations, more recently attention has turned to early-life influences as determinants of these adult diseases.[3]

Previous studies that compared South Asians living in India with those in the United Kingdom reported that South Asian newborns had lower birthweight and had relatively more adipose tissue compared with White Caucasian (White) newborns of the same gestational age.[3,4] This has been termed the 'thin-fat' phenotype and may represent an early feature of increased cardiometabolic risk, as there is a growing body of evidence associating lower birthweight with abdominal obesity, hypertension, future type 2 diabetes and cardiovascular disease.[5,6] Interestingly, this phenotype is present among South Asian immigrants to the United Kingdom and persists across multiple generations.[7]

Health system databases in Canada indicate greater maternal gestational diabetes (GDM) in South Asian women,[8,9] and that South Asian newborns have lower birthweight compared with White newborns.[10] These two observations are at odds with each other because elevated maternal glucose is generally associated with higher birthweight or macrosomia.[11]

To further understand these observations, among women with full-term pregnancies recruited in two contemporary birth cohorts in Ontario, Canada, we sought to investigate: (1) the ethnic differences in newborn skinfold thickness; and (2) the antenatal maternal factors which influence these measurements.

PATIENTS AND METHODS

Eligibility and recruitment

Participants from the FAMILY and START prospective birth cohorts were potentially eligible for this analysis. Both studies were Research Ethics Board approved and all participants provided written informed consent.

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Eligible participants for this analysis were required to have a full-term pregnancy that is, ⩾ 37 weeks, have delivered a single offspring, and have the maternal baseline assessment conducted between 22 and 30 weeks gestational age. White participants were selected from the FAMILY cohort and were recruited from the greater Hamilton area in Ontario, Canada between 8 October 2002 and 8 July 2009. South Asian participants were from the START cohort and recruited from the Peel region, Ontario, Canada between 11 July 2011 and 20 September 2013. Detailed recruitment and baseline measures are reported in detail for both cohorts in prior publications.[12,13]

Maternal measurements

During the second trimester, participants who did not have pre-existing diabetes completed a 75-g oral glucose tolerance test (OGTT). Individuals with pre-existing diabetes did not have the OGTT, but provided a fasting blood sample. GDM was defined using International Association of Diabetes and Pregnancy Study Groups criteria[14] as fasting glucose ⩾ 5.1 mmol l^{-1} or 1 h post OGTT glucose ⩾ 10.0 mmol l^{-1} or 2 h post OGTT glucose of ⩾ 8.5 mmol l^{-1} and area under the curve (AUC) glucose was used as a continuous measure of glucose response to a 75-g OGTT.[15] All participants completed detailed health questionnaires including self-reported personal medical and family history, social and cultural questions, and previously validated ethnic-specific food frequency questionnaires.[16] Physical activity in pregnancy was defined as hours of active sport per week during pregnancy as reported by the mother at her initial visit.

Social disadvantage was determined using a previously validated index which includes employment, marital status and income.[17] Physical measurements using a protocol standardized between both cohorts included resting blood pressure using an oscillometric device, body weight, height, hip circumference and skinfold thickness. Maternal body fat in pregnancy was estimated by adding the skinfold thickness from triceps and subscapular sites.[18] Participants also self-reported their prepregnancy weight, and gestational weight gain was calculated by subtracting the prepregnancy weight from the end pregnancy weight prior to delivery.

Delivery information and newborn measurements

Type, duration and outcomes of labor were obtained from hospital charts by trained research assistants using a standardized protocol. Measurements of newborns were completed by trained research assistants using a standardized protocol and 90% were taken within 24 up to 96 h after delivery. Specifically, newborn length was collected using the O'LEARLY length board. Birthweight was obtained from the hospital chart in 98% of newborns, or if unavailable, the newborn was weighed using the hospital newborn scale at the time of the birth visit. Head circumference was measured using a nonstretchable measuring tape. Waist and hip circumferences were measured using an OHAUS nonstretchable tape with an attached spring balance and the time of last feed prior to the measurements was not recorded. Skinfold thickness were taken in triplicate and recorded to the nearest mm using the HOLTAIN calipers in START (0.2 mm) and the LANGE calipers in FAMILY, (0.5 mm). The intraclass correlation (reliability) of the skinfold measurements was calculated and was 0.98 for subscapular skin folds and 0.96 for triceps skinfolds overall.[19] Gestational weight gain was reported as a percent weight gain relative to the participant's prepregnancy weight. Ponderal index—a measure of neonatal leanness was calculated as birthweight per length[3].[3]

Statistical considerations

We compared ~ 400 subjects per group to have 90% power to detect a difference of 10% in skinfold thickness between the ethnic groups. Statistical analyses were performed using SAS, version 9.2 (SAS Institute, Cary, NC, USA). Means, s.d. and proportions are presented for descriptive variables. Between group comparisons were made using ANOVA or χ^2-tests. Comparisons between South Asian and White newborns were adjusted for newborn sex and gestational age. A multiple regression model

Table 1. Maternal baseline characteristics			
	South Asian	*White Caucasian*	*P-value*
N per group	401	389	
Maternal age (years)	30.1 (4.0)	31.8 (5.0)	< 0.001
Years in Canada	8 (7)	30 (7)	< 0.0001
Gestational age at study enrolment (years)	26.7 (1.3)	26.9 (1.7)	0.03
Primiparous (%)	38.7	43.7	0.15
Smoked in pregnancy (%)	0	13.3	< 0.0001
Vegetarian (%)	40.5	0.8	< 0.0001
Hours of active sport per week	1.7 (2.6)	2.2 (3.0)	0.01
Family history of diabetes (%)	38.7	19.0	< 0.0001
Personal history of prepregnancy DM (%)	0.5	3.1	0.006
Social disadvantage index			
High (%)	18.9	5.6	< 0.0001
Moderate (%)	36.4	14.7	
Low (%)	44.7	79.7	
Currently employed (%)	52.8	83.2	< 0.0001
Annual household income ⩾ 50 K (%)	40.8	80.2	< 0.0001
High school education (%)	99.2	97.9	0.12
Prepregnancy weight, kg	62.2 (11.3)	71.1 (17.7)	< 0.0001
Height, cm	162.1 (6.5)	164.9 (6.2)	< 0.0001
Prepregnancy BMI, kg m^{-2}	23.7 (4.3)	26.2 (6.3)	< 0.0001
Gestational weight gain, kg	13.8 (6.3)	15.0 (5.3)	0.01
GWG / prepregnancy weight	24%	22%	0.17
Tricep skinfold thickness, mm	28.2 (6.8)	31.1 (12.1)	< 0.001
Subscapular skinfold thickness, mm	23.1 (6.9)	23.6 (10.2)	0.46
Tricep+subscapular skinfold thickness thickness, mm	51.2 (12.5)	54.3 (20.9)	0.01
Gestational DM–IADPSG and recorded in chart (%)	20.8	13.3	0.005
Received insulin during pregnancy (%) (Use of insulin at baseline visit/current use of insulin)	0.8	2.3	0.07
Area under curve glucose, mmol × h	13.3 (2.8)	12.3 (2.3)	< 0.0001

Abbreviations: BMI, body mass index; DM, diabetes mellitus; GWG, gestational weight gain; IADPSG, International Association of Diabetes and Pregnancy Study Groups. Social Disadvantage Index, includes employment status, marital status and annual household income.[15]

was constructed to assess the independent predictors of newborn skinfold measurements adjusting for newborn age and sex; mothers taking insulin at baseline were excluded. Individual factors which were significant predictors of newborn skinfold measurements, adjusting for ethnicity, with a $P < 0.10$ was tested in a multivariable model. Backward stepwise regression was used to determine the final model with all variables significant at $P < 0.05$. Interactions between ethnicity and each significant main effect were tested.

RESULTS

Maternal comparisons

The baseline characteristics of mothers with full-term singleton pregnancies from the FAMILY ($n = 389$) and START ($n = 401$) cohorts are shown in Table 1. Briefly, compared with Whites, South Asian mothers were younger (30.1 vs 31.8 years, $P < 0.001$), and were mostly immigrants, having lived in Canada an average of 8 years. The majority of South Asian women were of Indian origin (77.3%) followed by Pakistani (16.3%), Bangladeshi (0.75%), Sri Lankan (2.7%) and other (2.5%). South Asian women were less likely to be smokers during pregnancy (0% vs 13%, $P < 0.0001$), more likely to be vegetarian (40.5% vs 0.8%, $P < 0.0001$), have a stronger family history of type 2 diabetes (38.7% vs 19.0%, $P < 0.0001$) and performed fewer hours of active sport per week
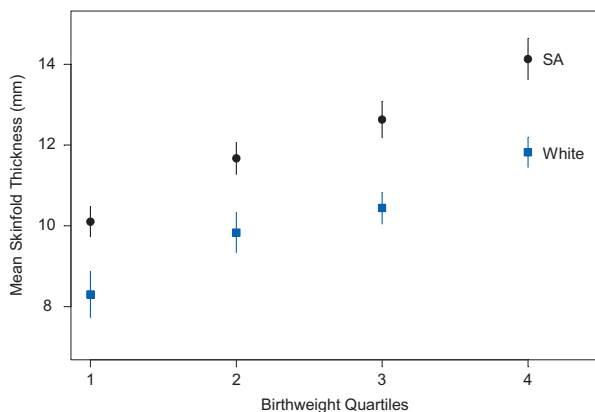

Figure 1. Mean skinfold thickness by birthweight quartiles comparing South Asian and White newborns adjusting for newborn age, sex and age at birth visit. Using continuous birthweight, South Asian newborns have higher skinfolds over all birthweights ($P < 0.0001$); the increasing trend in both ethnic groups is significant ($P < 0.0001$). Note that adjustment for GDM makes no appreciable difference in the results.

compared with White women (1.7 (2.6) vs 2.2 (3.0) h per week, $P = 0.01$). Despite having similar high school completion rates, South Asian women had greater social disadvantage (18.9% vs 5.6%, $P < 0.0001$) as reflected by their lower annual household income and employment.

Compared with White women, South Asians had lower weight (prepregnancy weight difference of 9 kg, $P < 0.0001$), were shorter (prepregnancy height difference: 3 cm, $P < 0.0001$), had a lower prepregnancy body mass index (23.7 vs 26.6 kg m^{-2}, $P < 0.0001$), and had lower triceps and subscapular skinfold thickness (51.2 vs 54.3 mm, $P = 0.01$). During pregnancy, the percent of weight gain relative to the prepregnancy weight comparing South Asian and White women was similar (24% vs 22%, $P = 0.17$; Table 1).

South Asian women had significantly more gestational dysglycemia. Specifically using the IADPSG definition to diagnose GDM, South Asian mothers had almost a twofold higher rate of GDM than White mothers (20.8% vs 13.3%; $P = 0.005$) and higher AUC glucose (13.3 vs 12.3 mmol × h; $P < 0.0001$; Table 1). Fourteen women had pre-existing diabetes (2 SA vs 12 White) and 12 women were treated with insulin during pregnancy (SA: 3 vs White: 9).

Newborn comparisons

South Asian newborns had significantly lower birthweight (3283 vs 3517 g, $P < 0.0001$), and had significantly greater skinfold thickness (11.7 (0.1) vs 10.6 (0.1); $P < 0.0001$) compared with White newborns. Further, compared with White newborns, South Asian newborns had greater skinfold thickness across each quartile of birthweight (Figure 1). Interestingly South Asian newborns waist circumference, a measure of abdominal obesity in adults but not validated in newborns was significantly higher (31.1 vs 29.9 cm, $P = 0.0001$), although no difference in the waist to height ratio was observed between the groups (0.598 in SA vs 0.595 in Whites; $P = 0.40$; Table 2). Also, the ponderal index was substantially lower (23.3 vs 27.9, $P < 0.0001$) in South Asian than in White newborns. Furthermore, at any level of skinfold thickness, South Asian newborns had a lower ponderal index (that is, are thinner) than White newborns ($P < 0.0001$; Figure 2).

To determine the impact of increased maternal dysglycemia on newborn body composition, the AUC glucose was divided into tertiles, and the anthropometric characteristics of South Asian and White newborns were examined. Birthweight ($P < 0.0001$), skinfold thickness ($P = 0.006$) and ponderal index ($P < 0.0001$) increased progressively with increasing maternal glucose in both ethnic groups, but neonatal length ($P = 0.57$) did not (Figure 3).

Table 2. Newborn baseline characteristics[a]

	South Asian	White Caucasian	P-value
N per Group	400	389	
Male (%)	47.9	48.1	0.96
Gestational age at birth, weeks	39.3	39.6	0.002
Birthweight, g	3283 (22)	3517 (22)	< 0.0001
Newborn head circumference, cm	34.1 (0.1)	34.9 (0.1)	< 0.0001
Newborn length, cm	52.1 (0.1)	50.2 (0.1)	< 0.0001
Newborn Waist circumference, cm	31.1 (0.1)	29.9 (0.1)	< 0.0001
Waist to height ratio	0.598 (0.002)	0.595 (0.002)	0.40
Triceps skinfolds, mm	6.2 (0.1)	5.4 (0.1)	< 0.0001
Subscap skinfolds, mm	5.6 (0.1)	5.2 (0.1)	< 0.0001
Tricep+subscapular skinfolds, mm	11.7 (0.1)	10.6 (0.1)	< 0.0001
Ponderal index, kg m^{-3}	23.3 (0.2)	27.9 (0.2)	< 0.0001

Ponderal index defined as, Birthweight in kg per birthlength in m^3. [a]Means (s.e.) Adjusted for gestational age and sex.

Predictors of newborn skinfold thickness

Individual maternal factors, adjusting for ethnicity were first tested in regression models. Factors with a $P < 0.10$ (years in Canada, prepregnancy weight, tricep plus subscapular skinfolds thickness, AUC glucose, social disadvantage index and physical activity) as well as newborn male sex and gestational age were retained in a multiple regression analysis (Table 3). The final model showed that in addition to male sex and gestational age, South Asian ethnicity (standardized estimate (s.e.): 0.24; $P < 0.0001$), maternal skinfold thickness—a surrogate measure of maternal body fat (s.e.: 0.14; $P = 0.0002$)—and maternal AUC glucose (s.e.: 0.079; $P = 0.04$) were significantly associated with newborn skinfold thickness (Table 3).

DISCUSSION

South Asian newborns have lower birthweight and greater skinfold thickness—considered to represent 'fatness', when compared with White newborns. The skinfold thickness is independently influenced by maternal glucose and maternal body fat.

We identified intriguing differences in newborn body composition as South Asian offspring were lower birthweight, more

Figure 2. Newborn skinfold thickness by ponderal index (kg m^{-3}). White newborns have a higher ponderal index over all skinfolds ($P < 0.0001$); the increasing trend in both ethnic groups is significant ($P < 0.0001$). There is no significant interaction ($P = 0.19$).

adipose and had higher waist circumference compared to Whites. Lower birthweight and increased adipose tissue have been observed in some but not all prior comparisons of South Asian and White offspring.[3,4,20] We observed that across a range of birth weights, South Asians have significantly higher adipose tissue compared with Whites. South Asian ethnicity, mother's body fat and increased maternal glucose were positively associated with skinfold thickness—a surrogate measure of adiposity in newborns. However, collectively these factors accounted for only 10% of the variance of the newborn skinfold thickness, which suggests that additional factors including genetic and epigenetic factors may also be important determinants of this phenotype.[21,22]

Our observation that South Asian newborns have an increased waist circumference compared with White newborns may be important because increased waist circumference is associated with increased trunk fat mass in children aged 3 to 5 years,[23] insulin resistance in children ages 9 to 11 years of age,[24] cardiometabolic risk factors in children and adolescents[25] and is strongly correlated with visceral adiposity and cardiometabolic factors in adults.[1,22] Our observation is supported by a prior report using whole-body magnetic resonance imaging, which found that full-term South Asian newborns in India compared with White newborns in London, UK (within 2 weeks of birth) had increased abdominal adiposity (including more visceral, deep subcutaneous, and superficial subcutaneous fat) and less nonabdominal superficial subcutaneous fat.[26] However, the waist to height ratio was not different between South Asian and White newborns. Waist to height ratio has been proposed as a simple screen for visceral adiposity in adults, adolescents and children,[27,28] but to our knowledge has not been evaluated in full-term newborns. Therefore, although the waist circumference taken at birth may be a useful predictor of future cardiometabolic risk in the growing offspring, both waist circumference and waist to height ratio in newborns require validation in newborns and confirmation of their predictive abilities in longitudinal studies.

South Asian pregnant women were smaller stature yet had increased dysglycemia compared with White women. Thus, South Asian fetuses were exposed to greater hyperglycemia in utero compared with Whites. The expected physiologic response of the fetus to maternal dysglycemia is greater production of fetal insulin, and subsequently increased lean and adipose tissue
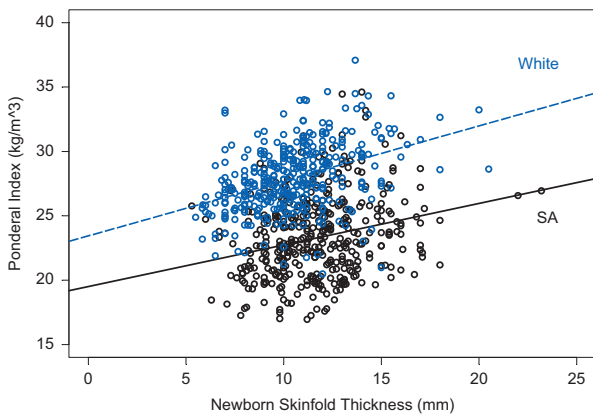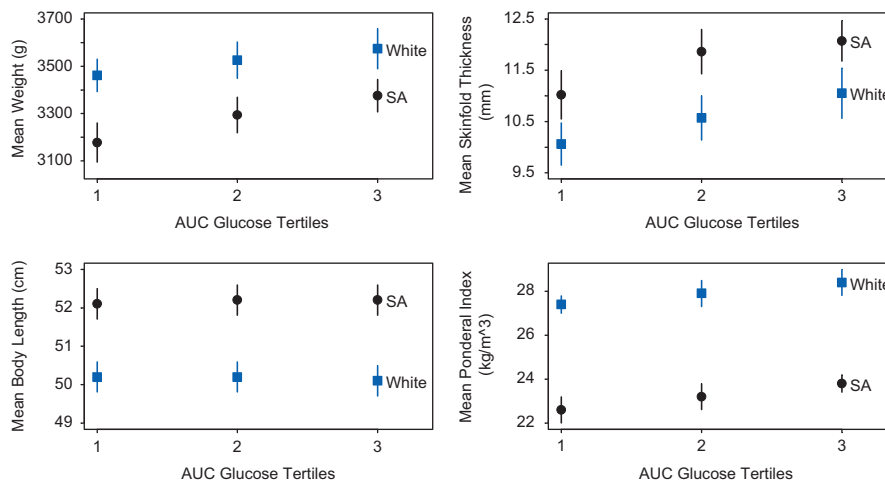

Figure 3. Newborn body composition by maternal AUC glucose tertiles (women receiving insulin during pregnancy excluded), adjusting for newborn sex and gestational age. Ethnic differences are significant for each newborn parameter ($P < 0.0001$ for each). Birthweight, skinfold thickness, and ponderal index increase with increasing AUC glucose (continuous measure) adjusting for newborn sex and gestational age ($P < 0.0001$, $P = 0.0006$, $P < 0.0001$ respectively); while there is no trend found with birthlength ($P = 0.57$).

Table 3. Predictors of newborn skinfold thickness

Factor	Single variable models adjusting for ethnicity		Full multivariable model[a]		Final multivariable model[b]	
	Standardized parameter estimate	P-value	Standardized parameter estimate	P-value	Standardized parameter estimate	P-value
Maternal characteristics						
South Asian	0.234	< 0.0001	0.326	< 0.0001	0.237	< 0.0001
Primiparous	− 0.034	0.34	NA			
Years in Canada	0.108	0.10	0.066	0.36		
Prepregnancy weight (kg)	0.161	< 0.0001	0.054	0.34		
Maternal height (m)	0.029	0.43	NA			
Tricep+subscapular skinfold (mother)	0.145	< 0.0001	0.117	0.03	0.141	0.0002
Gestational weight gain	0.021	0.57	NA			
AUC glucose (mmol×h)	0.119	0.001	0.054	0.19	0.079	0.04
Smoking during pregnancy	− 0.002	0.96	NA			
Vegetarian	− 0.045	0.27	NA			
Social Disadvantage Index (relative to low)						
Moderate	− 0.076	0.06	− 0.059	0.14		
High	− 0.046	0.24	− 0.025	0.54		
Physical activity	− 0.063	0.08	− 0.047	0.22		
Newborn Characteristics						
Male sex	− 0.097	0.006	− 0.090	0.02	−0.083	0.02
Gestational age at birth	0.059	0.10	0.084	0.03	0.082	0.02

[a]Factors with a univariate $P < 0.10$ were tested in a multiple regression model holding newborn age and sex in the model. [b]Backward stepwise regression methods were used to determine the final model with all variables significant at $P < 0.05$ holding newborn age and sex in the model ($R^2 = 0.10$).

growth, with the extreme hyperglycemic fetus becoming macrosomic.[11] Birthweight, adipose tissue and lean body mass increased among South Asian and White newborns with increasing maternal glycemia. Despite having a significantly higher burden of dysglycemia, South Asians were lower birthweight, yet had higher skinfold thickness in each AUC glucose tertile (Figure 3). Future investigations of fetal/newborn insulin and adipokines, such as leptin and adiponectin, genetic and epigenetic factors, may further elucidate the reasons for differences in skinfold measurements relative to lower birthweight among South Asians.[29,30]

Our study has a number of strengths. First, South Asians and White participants from the START and FAMILY birth cohorts both live in the same geographic regions of Southern Ontario, Canada. Second, standardized methods to assess hyperglycemia, maternal and infant anthropometry and health behaviors were used in both cohorts. These are advantages over previous transcontinental studies, which have indirectly compared South Asians with Whites. There are, however, some limitations of our interethnic comparisons. A potential for bias includes use of mother's self-reported prepregnancy weight, although this has been found to be reasonably reliable in previous birth cohorts.[31] Additional limitations include: the use of two different instruments to measure skinfold thickness in the two cohorts with no assessment of interobserver variability, timing of measurement of skinfold thickness ranged between 0 to 96 h after birth although adjustment for time and measurement did not change our results, and birth weights that were extracted from hospital records. Another potential limitation is that the time period during which the cohorts were recruited is not identical, as the Whites were recruited between 2002 and 2009 and the South Asians between 2011 and 2013. However, we do not believe that there has been any considerable secular change in lifestyle factors, that is, in availability of food or walking environment, or in health care delivery especially as Canada has a public health care system within which there is equal access to prenatal and hospital care for all landed immigrants and citizens.

CONCLUSION

South Asian newborns are lower birthweight, and have greater skinfold thickness compared with White newborns, and this is influenced by increased maternal body fat and glucose. Interventions aimed at reducing body fat prior to pregnancy and GDM during pregnancy in South Asians may favorably alter newborn body composition and require evaluation.

CONFLICT OF INTEREST

The authors declare no conflict of interest.

AUTHOR CONTRIBUTIONS

Researched data done by SA, MG, KM, KT, NA, DD, GW, CW, PS, KS; SA wrote the manuscript and researched the data; SA, MG, KS, AV, AK, PD Reviewed/edited the manuscript, and SA, MG, KT, KM, SM researched the data and contributed to discussion.

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