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BMJ Open The relation between birthweight, childhood body mass index, and overweight and obesity in late adolescence: a longitudinal cohort study from Norway, The Tromsø Study, Fit Futures

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ABSTRACT

Objectives Childhood overweight/obesity is associated with later overweight/obesity. However, the association between birth weight and later overweight/obesity has not been established. The aim of this study was to investigate the relation between both birth weight and childhood body mass index (BMI), and adolescent overweight/obesity in a Norwegian population.

Methods The Tromsø Study - Fit Futures is a population-based cohort study conducted in 2010-2011 and 2012-2013 in Tromsø, Norway. A representative sample of 961 adolescents participated. Longitudinal anthropometric data were obtained from the Medical Birth Registry of Norway, childhood health records at 2-4 and 5-7 years of age, and repeated measurements at 15–18 and 18–20 years of age. Outcome was delined as normal weight (adult BMI < 25 kg/m²) or overweight/ obese (adult BMI ≥2 5 kg/m²) at 15–20 years of age according to international age- and sex-speciac cut-off values for children. Associations were investigated using generalised estimating equations.

Results In adjusted analyses, a 1-SD (586 g) higher birth weight was associated with a higher OR for overweight/obesity at 15-20 years of age (OR 1.25, 95%) Cl 1.06 to 1.48). Childhood BMI was also associated with overweight/obesity at 15-20 years of age: a 1-SD (1.35 kg/m²) increase in BMI at age 2-4 years rendered an OR of 1.66 (95% Cl 1.40 to 1.96); a 1-SD (1.83 kg/ m²) increase in BMI at age 5–7 years rendered an OR of 3.23 (95% Cl 2.56 to 4.07). When compared with normal-weight children, those with severe overweight/ obesity in childhood (adult BMI ≥27 kg/m²) showed stronger associations with overweight/obesity at 15-20 years of age: OR3.01 (95%Cl 1.47 to 6.18) and OR 11.51 (95%Cl 6.63 to 19.99) at ages 2-4 and 5-7, respectively.

Conclusion Associations between birth weight and overweight/obesity at 15-20 years of age were modest, whereas the in uence of BMI at 2-4 and 5-7 years on overweight/obesity at 15-20 years was moderate to strong.

Strengths and limitations of this study

- ► The main strengths of this study are its populationbased design and access to longitudinal data from birth (1992-1994 cohort) to 18-20 years of age.
- ► The high attendance rate and representative sample from a Norwegian adolescent population minimise selection bias.
- Data from the national Medical Birth Registry of Norway and objectively measured height and weight were used to calculate body mass index at all ages, reducing the risk of information bias.
- Missing data from childhood measurements were handled with a multiple imputation technique.
- Alimitation of this study is the lack of information on potential confounding factors such as parental and lifestyle factors at birth and childhood.

INTRODUCTION

The increasing prevalence of overweight/ obesity among children worldwide is a major health concern due to several related immediate and long-term health problems.¹⁻⁷ A moderate degree of tracking (maintenance of certain risk factors over time) has been reported from childhood to adolescence and into adulthood, indicating that individuals experience only small changes in weight class positions throughout their life.^{6 8 9} However, the question remains whether birth weight is a predictor of later overweight/ obesity. In a review by Brisbois et al¹⁰, birth weight did not emerge as an early marker for adult overweight/obesity, but a recent review reported consistent associations between high birth weight and overweight later in childhood. 11 Data from Norway also revealed strong associations between birth

weight and overweight/ obesity at 7–8 years of age. 12 13 From a health perspective, it is of interest to investigate whether high birth weight also tracks into adolescence/early adulthood.

Lately, the focus of this investigation has shifted to early-life factors, especially genetic factors that influence obesity and later health problems. ^{14–16} Information on the degree of tracking of high birth weight and body mass index (BMI) during early childhood is important for health authorities and healthcare workers who are planning preventive interventions to halt the overweight epidemic. Effective treatment of childhood obesity has proven to be very challenging ¹⁷; therefore preventive efforts are of major importance. However, the appropriate age at which to initiate such efforts is still a matter of discussion. ¹¹⁴

To answer these questions, we aimed to explore the associations between both birth weight and childhood BMI and overweight/obesity in adolescence in a population-based cohort of Norwegian adolescents born in 1992–1994, a period with high mean birth weight in Norway.¹⁸

METHODS

Fit Futures is an expansion of The Tromsø Study, a population-based study from Northern Norway with repeated health surveys among adults. All first-year students in all upper-secondary schools in the municipalities of Tromsø and Balsfjord in 2010–2011 (n=1117) were invited to Fit Futures 1 (FF1). Of these invited students 1038 (92.9%) attended. Detailed information on FF1 and its youth cohort has already been presented. 19 20 A follow-up study, Fit Futures 2 (FF2), was conducted 2 years later during 2012–2013, and re-invited all participants from FF1. The present study consists of the 961 participants (492 boys and 469 girls) in FF1 who were aged <18 years at FF1 (born 1992–1994). Among these, 659 had anthropometric measurements available at 18–20 years of age in FF2.

Longitudinal anthropometric data were obtained through linkage to the Medical Birth Registry of Norway (MBRN) and childhood health records. Each student's unique personal identification number was used to link to the MBRN, from which information on birth weight and length, gestational age, and other variables related to birth were obtained. Height, weight, age, and date of measurements at two time points were collected from childhood health records. In Norway, regular health controls by public health nurses, including measurement of height and weight, are offered for all children in accordance with national preventive health programme guidelines. Most of our participants had their height and weight measured at 2 and 6 years of age. If data were missing for those exact ages or there were supplementary measurements, the measurement closest to the 2- and 6-year birthday was recorded. The exact age of the participants at the time measurements were taken varied slightly;

therefore the age groups are reported as 2–4 (mean age 2.6 years) and 5–7 years of age (mean age 6.0 years).

A total of 411 included participants (43%) had one or more variables missing from the MBRN and/or child-hood health records (5% missing birth weight, 29% and 23% missing height/ weight at 2–4 and 5–7 years of age, respectively). Reasons for missing the childhood measurements were change of residency and measurements outside the age limits. Three hundred and two (31%) participants had missing data at FF2. A flow chart shows the study population and exclusions/ missing information (figure 1).

The Regional Committee for Medical and Health Research Ethics, North Norway (REK nord) approved FF1, FF2 and the present study (Reference number: 2014/1397/REK nord). All students and parents/guardians of students <16 years of age gave written informed consent.

Data/measurements

Trained study nurses performed anthropometric measurements in FF1 and FF2, following standardised procedures. Participants wore light clothing and no footwear. Height and weight were measured to the nearest 0.1 cm and 0.1 kg, respectively, on an automatic electronic scale/ stadiometer (Jenix DS 102 stadiometer, Dong Sahn Jenix, Seoul, Korea) at FF1 (15–17 years of age; mean age 16.6 years) and FF2 (18–20 years of age; mean age 18.6 years).

Height and weight were used to calculate BMI (weight/ height²; kg/m²) at 2-4, 5-7, 15-17 and 18-20 years of age and participants were classified into weight classes: underweight (adult BMI <18.5 kg/m²), normal weight (adult BMI ≥18.5 to <25 kg/m²), overweight (adult BMI \geq 25 to $\langle 30 \text{ kg/m}^2 \rangle$, and obese (adult BMI $\geq 30 \text{ kg/m}^2 \rangle$). BMI reference values for every half-year and the International Obesity Taskforce (IOTF) age- and sex-specific cut-off values were used for children 2–18 years of age.²¹ The WHO index for adults was used at age >18 years.²² Due to the small proportion of obesity at 2-4 and 5-7 years of age participants were also classified into the following weight classes: normal weight, light overweight (adult BMI ≥ 25 to $< 27 \text{kg/m}^2$), and severe overweight/ obesity (adult BMI ≥27 kg/ m²). The childhood BMI variables were used as predictors in the analyses.

When used as an outcome, weight classes at 15–17 and 18–20 years of age were dichotomised as normal weight (adult BMI $<25 \,\mathrm{kg/m^2}$) or overweight/obesity (adult BMI $\ge25 \,\mathrm{kg/m^2}$).

Birth weight was divided into low (<2500 g), normal (≥2500 to <4500 g) and high birth weight (≥4500 g) according to the WHO definition. The ponderal index (PI)—birth weight (kg) divided by the cube of birth length (m) (kg/m³)—was divided into tertiles. Age- and sex-specific weight and BMI SD scores (SDS) were calculated using LMS coefficients corresponding to the Norwegian growth reference. Growth status at birth was categorised as small for gestational age (<10th

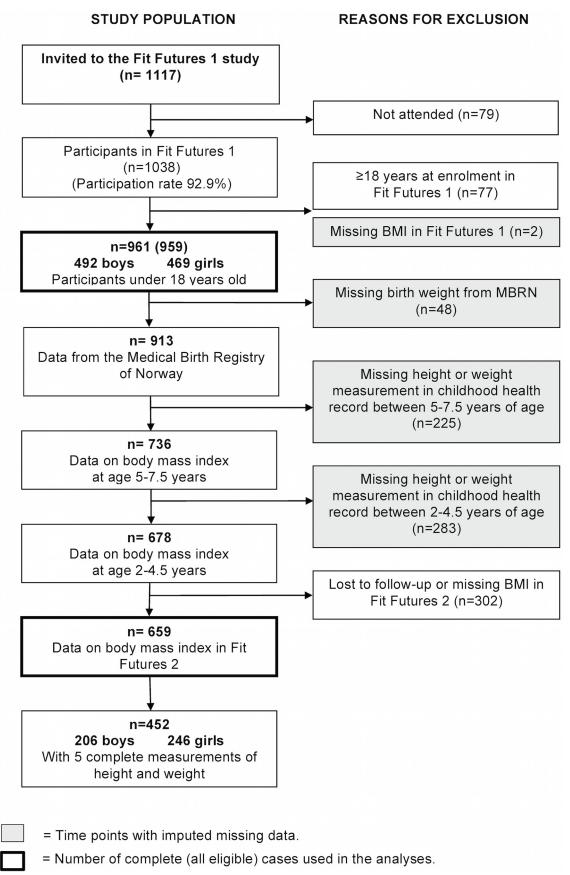


Figure 1 Flow chart of the study population, The Tromsø Study, Fit Futures 2010–2011 and 2012–2013. BMI, body mass index; MBRN, Medical Birth Registry of Norway.

percentile), appropriate for gestational age, and large for gestational age (>90th percentile) based on birth weight and gestational age and according to a national reference standard of births in 1987–1998.²⁴

Girls were categorised into three stages of pubertal maturation: early (<12.5 years), intermediate (12.5–13.9 years), and late (≥14.0 years), based on age at menarche specified in self-administered questionnaires. Pubertal maturation in boys was classified as barely started, underway, and completed based on a validated self-reported measure.²⁵

Statistics

Characteristics of the study population are presented as means and SD or numbers and percentages, by sex and weight class at 15–17 years of age. Differences between groups were assessed by t-test for continuous variables and by the χ^2 test for categorical variables. Associations between BMI at different ages were assessed with Spearman's rank correlations.

Tracking of birth weight or childhood BMI and weight class into adolescence was estimated as odds ratios (ORs) for being overweight/obese in adolescence (15-17 and 18–20 years of age, as a combined endpoint). Generalised estimating equations (GEE) were used with a logit link function and an unstructured correlation matrix in crude and multivariable models. Predictors were birth weight (per 1-SD increase), birth weight SDS, PI in tertiles, BMI (per 1-SD increase), BMI SDS and weight class at 2-4 and 5-7 years of age. Covariates included in the multivariable models were gestational age/ age, maternal age, and sex. All models were run in the total cohort and by sex. Covariates tested but not included in the final models were: multiple births, caesarean section, maternal disease or diabetes, time between measurements, and participation in FF2. Potential interaction between sex and birth weight or childhood BMI was assessed by including crossproduct terms in the models. Pubertal maturation was tested as a potential mediator. However, as it did not affect the coefficients and interaction terms were not significant, pubertal maturation was not included in the final models. Linearity was assessed by visual inspections of scatterplots, with birth weight and BMI as continuous variables (see online supplementary figures 1-3). No sign of a non-linear relationship was observed, but we analysed the predictors both as continuous and categorical variables.

Due to the relatively large number of missing explanatory variables, we performed multiple imputations on data that were missing at birth, 2–4 years and 5–7 years of age using chained equations generating 20 duplicate datasets. (see online supplementary document). The estimates from the 20 imputed datasets were combined into an overall estimate with corresponding SE using Rubin's rule. (26 27)

Multiple imputations and statistical analyses were carried out using Stata/MP 14.1 for Mac (Stata Corp, College Station, TX, USA). The level of statistical significance was set to two-sided p values <0.05.

RESULTS

Overall mean birth weight was 3530 g and the proportion of participants with high birth weight (≥4500 g) was 4.5%. Characteristics of the study population at birth and at mean ages 2.6, 6.0, 16.6 and 18.6 years are presented by sex (table 1). The prevalence of overweight/obesity was 14.0% and 18.5% in girls, 8.8% and 10.9% in boys at 2–4 and 5–7 years of age, respectively. At 15–17 and 18–20 years of age, 20.6% and 20.9% of girls, and 23.4% and 28.0% of boys, were overweight/obese. The prevalence of obesity was <1.5% at 2–4 years and increased to 6.6% in girls and 8.1% in boys at 18–20 years of age (table 2).

Of the study population, 47.0% (452) had five complete measurements. The imputed dataset and the observed-cases dataset were similar with respect to main characteristics and prevalence rates of overweight/obesity. Comparisons between the datasets are presented in (see online supplementary tables 1-2). The distribution of missing childhood data between sexes and weight classes at 15–17 or 18–20 years of age is presented in (see online supplementary table 3).

We explored sex-specific differences in anthropometric measurements from birth throughout childhood between participants classified as normal weight and overweight/ obese at 15-17 years of age (table 3). Girls classified as overweight/ obese at 15-17 years of age had higher mean birth weight (158g, p<0.05) than their normal weight female counterparts. Differences in mean BMI and mean weight also increased with age both among girls and boys classified as overweight/obese at 15-17 compared with their normal weight counterparts. Mean differences in BMI were 0.84 and 2.28 kg/m² among girls and 0.48 and 1.61 kg/m² among boys at 2-4 and 5-7 years of age, respectively (p<0.05). Mean differences in weight were 1.0 and 3.8 kg among girls at 2-4 years and 5-7 years of age, respectively, and 0.3 and 2.7kg among boys at 2-4 years and 5-7 years of age, respectively (only significant for boys at age 5-7) (table 3).

PI at birth and BMI in childhood were positively (p<0.001) correlated with BMI in adolescence. Spearman's Rho between PI at birth and BMI at 15–17 and 18–20 years of age were 0.13 and 0.12, respectively. Corresponding values for BMI at age 2–4 were 0.32 and 0.26, and for BMI at age 5–7 they were 0.57 and 0.51. BMI at 15–17 years of age was highly correlated with BMI at 18–20 years of age (Spearman's Rho 0.85; p<0.001).

Tracking analysis

In multivariable GEE analyses, a 1-SD (586 g) increase in birth weight was associated with higher odds of overweight/ obesity at 15–20 years of age (OR 1.25, 95% CI 1.06 to 1.48). Infants in the highest tertile of birth weight had significantly higher odds of later overweight/ obesity compared with those in the lowest tertile, but not compared with the mid tertile (adjusted OR for highest vs lowest tertile of PI 1.56, 95% CI 1.08 to 2.24) (table 4). Repeating the analysis with growth categories as a predictor gave results similar to those obtained with

Table 1 Characteristics for girls and boys of the Fit Futures cohort at birth and four ages up to 18–20 years, The Tromsø Study, Fit Futures, observed cases

Characteristics	Girls			Boys			
	n	Mean/%	(SD)	n	Mean/%	(SD)	p Value [†]
Birth				<u> </u>			
Birth weight (g)	443	3455.0	(576.7)	470	3601.0	(590.0)	<0.001
Birth length (cm)	419	49.4	(2.3)	454	50.2	(2.3)	<0.001
Gestational age (weeks)	393	39.7	(1.8)	430	39.6	(2.1)	0.176
Preterm birth (before GA week 37)	15	3.8%		23	5.4%		0.292
Ponderal index (kg/m³)	419	28.73	(2.81)	454	28.45	(2.75)	0.141
Body mass index (kg/m²)	419	14.19	(1.49)	454	14.28	(1.52)	0.338
Birth weight SDS [‡]	443	-0.58	(1.74)	470	-0.24	(1.32)	0.001
Body mass index SDS [‡]	419	0.06	(1.05)	454	-0.02	(1.04)	0.265
Birth weight group:	443			470			0.015
Low birth weight (<2500 g)	20	4.5%		17	3.6%		
Normal birth weight (2500–4500 g)	412	93.0%		423	90.0%		
High birth weight (≥4500 g)	11	2.5%		30	6.4%		
Size for gestational age:	443			470			0.554
Small for gestational age	49	11.1%		46	9.8%		
Appropriate for gestational age	356	80.4%		375	79.8%		
Large for gestational age	38	8.6%		49	10.4%		
Maternal age at birth	443	28.4	(5.3)	441	28.1	(5.3)	0.995
2–4 years of age							
Age (years)	328	2.6	(0.4)	350	2.6	(0.4)	0.952
Height (cm)	328	91.2	(4.6)	350	92.7	(4.7)	<0.001
Weight (kg)	328	13.5	(1.7)	350	14.1	(1.8)	<0.001
Body mass index (kg/m²)	328	16.17	(1.39)	350	16.37	(1.30)	0.053
Weight SDS [‡]	328	-0.12	(1.11)	350	0.16	(1.16)	0.001
Body mass index SDS [‡]	328	-0.04	(1.17)	350	-0.07	(1.07)	0.217
5-7 years of age							
Age (years)	354	6.0	(0.4)	384	6.1	(0.4)	0.253
Height (cm)	354	116.7	(5.3)	384	118.3	(5.3)	<0.001
Weight (kg)	352	21.8	(3.8)	384	22.2	(3.5)	0.143
Body mass index (kg/m²)	352	15.96	(2.04)	384	15.82	(1.74)	0.304
Weight SDS [‡]	352	0.0004	(1.10)	384	0.11	(1.02)	0.162
Body mass index SDS [‡]	352	-0.06	(1.10)	384	-0.03	(0.98)	0.616
15–17 years of age							
Age (years)	469	16.6	(0.4)	492	16.6	(0.4)	0.325
Height (cm)	467	164.9	(6.5)	492	176.9	(6.7)	<0.001
Weight (kg)	467	60.9	(11.5)	492	70.2	(14.4)	< 0.001
Body mass index (kg/m²)	467	22.39	(3.96)	492	22.38	(4.17)	0.952
Pubertal maturation, §girls:	464			_	_	_	
Early (<12.5 years)	147	31.7%					
Intermediate (12.5–13.9 years)	212	45.7%					
Late (≥14.0 years)	105	22.6%					
Pubertal maturation, §boys:	_	_	_	387			
Barely started (PDS 2.0-2.9)				69	17.8%		
							Continued

Continued

Table 1 Continued							
Characteristics	Girls			Boys			
	n	Mean/%	(SD)	n	Mean/%	(SD)	p Value [†]
Underway (PDS 3.0-3.9)				285	73.6%		
Completed (PDS 4.0)				33	8.5%		
18–20 years of age							
Age (years)	363	18.6	(0.4)	296	18.7	(0.3)	0.105
Height (cm)	363	165.7	(6.5)	296	179.1	(6.5)	< 0.001
Weight (kg)	363	63.2	(12.0)	296	75.2	(14.6)	<0.001
Body mass index (kg/m²)	363	23.02	(4.22)	296	23.42	(4.18)	0.228

^{*}In observed data n is varying from 659 to 961.

PI in tertiles. In adjusted analyses, being small for gestational age was associated with lower odds of overweight/obesity at 15–20 years of age compared with those appropriate for gestational age (OR 0.40, 95% CI 0.22 to 0.72). Infants large for gestational age did not have significantly higher odds of overweight/obesity compared with those appropriate for gestational age (adjusted OR for large vs. appropriate for gestational age 1.31, 95% CI 0.84 to 2.05) (data not shown).

Childhood BMI was associated with overweight/ obesity in adolescence (OR per 1-SD increase in BMI at age 2–4: 1.66, 95% CI 1.40 to 1.96; OR per 1-SD increase in BMI at age 5–7: 3.23, 95% CI 2.56 to 4.07). Children who were obese at 2–4 years of age had fivefold increased odds of becoming overweight/ obese in adolescence (OR 5.35, 95% CI 1.42 to 20.09) compared with their normal weight peers, while at 5–7 years of age the estimated OR was 15.59 (95% CI 6.60 to 36.85), compared with those

of normal weight. Cross-product terms between sex and birth weight/BMI were tested and no significant interaction was found (table 4).

Due to the small number of obese children, CIs were wide for these groups. Therefore additional analyses were performed with weight class in three groups as predictor. At 2–4 years of age, those with light overweight (adult BMI \geq 25 to <27 kg/ m² (n=52)) had an OR of 1.88 (95% CI 1.08 to 3.29) and those with severe overweight/obesity (adult BMI \geq 27 kg/ m² (n=25)) had an OR of 3.01 (95% CI 1.47 to 6.18) for overweight/obesity at 15–20 years of age, compared with those of normal weight. At 5–7 years of age the corresponding ORs were 4.96 (95% CI 2.82 to 8.73) for light overweight (n=48) and 11.51 (95% CI 6.63 to 19.99) for severe overweight/obesity (n=59) when compared with those of normal weight (data not shown).

Table 2	Weight classes* at four ages, The Tromsø Study, Fit Futures observed cases†								
	Age	2–4 ye	ars	5–7 yeaı	'S	15–17 y	ears	18–20 ye	ears
		n	%	n	%	n	%	n	%
Girls	Underweight	48	14.6	37	10.5	24	5.1	16	4.4
	Normal weight	234	71.3	250	71.0	347	74.3	271	74.7
	Overweight	41	12.5	49	13.9	70	15.0	52	14.3
	Obesity	5	1.5	16	4.6	26	5.6	24	6.6
	Total	328	100.0	352	100.0	467	100.0	363	100.0
Boys	Underweight	43	12.3	28	7.3	38	7.7	24	8.1
	Normal weight	276	78.9	314	81.8	339	68.9	189	63.9
	Overweight	27	7.7	27	7.0	79	16.1	59	19.9
	Obesity	4	1.1	15	3.9	36	7.3	24	8.1
	Total	350	100.0	384	100.0	492	100.0	296	100.0

^{*}Weight classes according to the International Obesity Taskforce (IOTF) age- and sex-specilic cut-off values for children 2–18 years of age and the WHO index for adults from age 18–20.^{21 22}

[†]p Value for sex difference was obtained by t-test or χ^2 test.

[‡]SDS, SD scores according to Norwegian reference data.²³

[§]Pubertal maturation is based on age of menarche in girls and according to Pubertal Development Scale (PDS) in boys; total score of four items of secondary sexual characteristics on a scale from 1 to 4 (sum of total score divided by 4). None had a score <2.0 in total score. ²⁵ GA, gestational age.

[†]In observed cases n is varying at different ages from 659 to 959.

Sex-special anthropometric characteristics from birth up to 15-17 years of age for normal weight and overweight/obese 15-17-year-olds in the Tromsø Study, Table 3

Fit Futures (n=961)								
	Girls				Boys			
	Normal weight	eight	Overweight/obese	ht/obese	Normal weight	eight	Overwei	Overweight/obese
	Mean	95 % CI	Mean	95% CI	Mean	95% CI	Mean	95% CI
Birth weight (g)	3418.0	(3358.1 to 3478.2)	3576.0	(3462.0 to 3689.5)	3597.0	(3535.5 to 3659.0)	3622.0	(3522.2 to 3721.4)
Birth weight SDS	-0.67	(-0.86 to -0.49)	-0.24	(-0.52 to -0.04)	-0.25	(-0.39 to -0.11)	-0.19	(-0.41 to 0.03)
Birth length (cm)	49.2	(48.9 to 49.4)	49.7	(49.2 to 50.1)	50.1	(49.9 to 50.4)	50.1	(49.7 to 50.6)
Ponderal index (kg/m³)	28.55	(28.24 to 28.85)	29.03	(28.44 to 29.62)	28.34	(28.04 to 28.64)	28.61	(28.14 to 29.08)
Weight 2-4 years (kg)	13.2	(13.0 to 13.4)	14.2	(13.8 to 14.6)	14.0	(13.8 to 14.2)	14.3	(13.9 to 14.7)
Weight SDS† 2-4 years	-0.23	(-0.36 to -0.09)	0.23	(-0.02 to 0.48)	60.0	(-0.04 to 0.22)	0.42	(0.14 to 0.69)
Height 2-4 years (cm)	8.06	(90.3 to 91.2)	91.8	(90.8 to 92.9)	92.6	(92.4 to 93.3)	92.3	(91.4 to 93.2)
BMI 2-4 years (kg/m²)	16.00	(15.84 to 16.16)	16.84	(16.53 to 17.15)	16.25	(16.10 to 16.40)	16.73	(16.43 to 17.04)
Weight 5-7 years (kg)	20.8	(20.5 to 21.1)	24.6	(23.6 to 25.5)	21.5	(21.2 to 21.8)	24.2	(23.4 to 25.1)
Weight SDS† 5-7 years	-0.27	(-0.37 to -0.17)	08.0	(0.57 to 1.02)	-0.09	(-0.19 to 0.001)	0.75	(0.54 to 0.97)
Height 5–7 years (cm)	116.2	(115.6 to 116.7)	117.5	(116.3 to 118.7)	117.9	(117.4 to 118.5)	118.9	(117.9 to 119.9)
BMI 5–7 years (kg/m^2)	15.41	(15.25 to 15.57)	17.69	(17.16 to 18.22)	15.44	(15.30 to 15.58)	17.05	(16.59 to 17.51)
Weight 15-17 years (kg)	56.8	(56.2 to 57.5)	76.8	(74.3 to 79.4)	64.4	(63.6 to 65.3)	88.9	(86.2 to 91.6)
Height 15-17 years (cm)	165.0	(164.4 to 165.7)	164.3	(162.9 to 165.6)	176.9	(176.2 to 177.5)	177.0	(175.8 to 178.3)
BMI 15–17 years (kg/m^2)	20.84	(20.65 to 21.03)	28.42	(27.59 to 29.25)	20.57	(20.36 to 20.78)	28.30	(27.59 to 29.01)

*Analysed in the dataset with 20 imputations (multiple imputation), n=961 (469 girls and 492 boys). †SDS, SD scores according to Norwegian reference data.²³ BMI, body mass index. Numbers in italic type are signi

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0.020/0.01615 p Value¹⁴ Fit Futures 2010-11 and 2012-13 0.062 0.215 (1.08 to 1.77) (0.93 to 1.43) (0.95 to 1.33) (0.90 to 2.39) (0.69 to 1.87) (2.10 to 3.95) (1.05 to 1.67) (1.99 to 3.44) 95% CI Ref. Ī 1 Ī Ī Adjusted 1.12 1.39 1.32 2.62 1.13 1.47 2.88 OR 0. Ī Ī (1.12 to 1.80) (0.90 to 1.29) (0.92 to 1.22) (0.67 to 1.75) (0.89 to 2.32) (1.05 to 1.67) (2.02 to 3.70) (2.00 to 3.44) Odds ratios for overweight including obesity* in adolescence (15-20 years of age) overall and by sex. The Tromsø Study, 95% CI Ref. Ī ī Ī Ī Crude 1.42 1.08 1.06 1.33 2.63 1.08 1.44 2.73 OR 0.1 (0.99 to 1.46) (0.96 to 2.90) (1.60 to 2.58) (1.51 to 2.48) (2.82 to 5.53) (1.05 to 1.77) (0.92 to 2.95) (2.59 to 5.34) 95% CI Ref. Adjusted¶ 1 Ī Ī 1 2.03 1.20 3.95 1.37 1.65 1.67 1.93 3.72 OR 0. Ī Ī 1 1 (1.10 to 1.74) (1.03 to 1.44) (0.94 to 2.97) (1.55 to 2.47) (1.51 to 2.49) (2.80 to 5.57) (1.02 to 3.00) (2.57 to 5.37) 95% CI Ref. Ī 1 1 Ī Ī I Crude 1.39 1.75 1.96 1.94 3.95 1.22 3.72 1.67 В 0. Ī Ī 1 (6.60 to 36.85) (1.42 to 20.09) (3.37 to 8.48) (0.91 to 1.96) (1.08 to 2.24) (1.31 to 1.82) (2.53 to 3.81) (1.05 to 1.33) (1.40 to 1.96) (1.07 to 2.82) (2.56 to 4.07) (0.08 to 0.61) (1.06 to 1.48) (0.24 to 0.73) Overweight/obesity at 15-20 years of age 95% CI Adjusted‡§ Ref. Ref. Ref. 1.18 1.56 1.66 1.55 0.42 1.74 5.35 3.11 5.35 15.59 1.25 1.34 3.23 0.22 0.1 0. 0. OR (6.54 to 37.62) (1.44 to 20.98) (0.08 to 0.61) (3.34 to 8.25) (1.07 to 2.18) (1.09 to 2.84) (1.04 to 1.27) (0.90 to 1.89) (1.41 to 1.95) (0.24 to 0.73) (2.47 to 3.88) (2.53 to 3.82) (1.06 to 1.41) (1.32 to 1.83) 95% CI Ref. Ref. Ref. 1.15 0.42 Crude 9. 1.66 1.55 1.75 5.49 3.10 3.11 0.23 5.25 15.68 1.30 1.53 1.22 0.1 0. O_R ₹ Weight class at 2-4 years: Weight class at 5-7 years: Ponderal index in tertiles: Birth weight per SD Birth weight SDS 5-7 years of age 2-4 years of age Normal weight Normal weight Underweight **Jnderweight** BMI per SD BMI per SD Overweight Overweight **BMI SDS** BMI SDS Table 4 Medium Obese Birth High Γow

*Weight classes according to the International Obesity Taskforce (IOTF) age- and sex-speciac cut-off values for children 2–18 years of age and the WHO index for adults from age 18–20.21 22 Analysed in the dataset with 20 imputations (multiple imputation), n=961 (469 girls and 492 boys).

Variables at 2-4 and 5-7 years of age are adjusted for age, maternal age and sex, except age- and sex-speciac weight class and SDS, which is adjusted only for maternal age. Variables in sex special analysis are adjusted for gestational age/age and maternal age, except age- and sex-special SDS which is adjusted only for maternal age. :Variables at birth are adjusted for gestational age, maternal age and sex, except sex-special SDS, which is adjusted for gestational age and maternal age.

*p Value for sex difference (interaction term) in crude analysis.

to for trend in all cases: 0.020 in crude analysis, 0.016 in adjusted analysis.

BMI, body mass index (kg/m²); GEE, generalised estimating equations; Ref, reference group; SDS, SD scores according to Norwegian reference data.

DISCUSSION

In this population-based longitudinal study of Norwegian adolescents born in 1992-1994, we found a statistically significant, but modest, association between birth weight and overweight/obesity at 15-20 years of age. Tracking of BMI and overweight from childhood to adolescence was moderate. Children with more severe overweight/ obesity had considerably higher odds of later overweight/ obesity compared with those with light overweight or their normal weight peers. The prevalence of overweight/ obesity increased with age. More girls than boys were overweight/obese at 2-4 and 5-7 years of age, but at 15-20 years this shifted so that more boys than girls were overweight/obese. Tracking of overweight/obesity from birth and childhood into adolescence was generally stronger in girls than in boys; however, no significant interaction with sex was found.

Mean birth weight and the proportion with high birth weight (4.5%) were somewhat higher than national data from 1992 to 1994 birth records (3.7–4.2%). High mean birth weight and high prevalence rates of overweight/ obesity in children have been reported from the northernmost region of Norway. The prevalence of overweight/ obesity in childhood we observed was comparable with findings from other studies from Norway that included similar age groups 13 29; however, the prevalence of overweight/ obesity in adolescence was higher than figures from comparable Norwegian studies. In the prevalence of overweight/ obesity is generally lower in the Nordic countries compared with the USA and southern European countries, which was reflected in our study.

Birth weight as a risk factor for overweight/obesity at later ages has been extensively studied with varying results. 10 11 34 In the review by Brisbois et al 25 of 43 studies found an association between birth weight and adult BMI or overweight/obesity, but with no clear conclusion that birth weight was an early marker for adult overweight/ obesity.¹⁰ Six of seven studies in a review by Weng et al found significant and strong associations between high birth weight and childhood overweight.³⁴ In two recent studies from Norway, increased BMI SDS at birth was associated with overweight including obesity at 8 years of age (OR per 1-SDS increase in BMI at birth 1.8, 95% CI 1.6 to 2.0). 12 A moderate degree of tracking of body size from birth to 7 years of age was found in the Norwegian Mother and Child Cohort Study. 13 Our finding of a statistically significant association between birth weight and overweight/obesity in adolescence is in line with these studies. However, birth weight was only a modest predictor of later overweight/ obesity in our study, probably due to the longer timespan. Stronger tracking of overweight/ obesity over a shorter timespan has been reported. 12 13 34 In contrast to our findings, two newer population-based studies from Sweden, which were comparable to our study, did not find high birth weight to be associated with high BMI or obesity at 15 or 20 years of age. 35 36 Since

tracking coefficients are influenced by time, comparisons between studies are challenging.^{8 37}

The participants in our study were born in a period during which there was a higher mean birth weight in the Norwegian population. The percentage of infants with a birth weight over 4500 g increased in Norway from 1990 to 2000, before declining to the same level as in 1990. Is firth weight emerged as a strong predictor of later overweight/ obesity, one would expect to see a high degree of tracking in our study compared with studies of children with a lower mean birth weight, but the estimated tracking coefficients were only moderate. However, since the prevalence of overweight/ obesity in adolescence is relatively high in our study population, a birth cohort effect cannot be ruled out.

Among those classified as overweight/ obese at 15–17 years of age, mean weight was generally higher at birth and BMI was significantly higher at 2–4 years of age. The estimated tracking coefficients from 2–4 and 5–7 years of age were moderate among our overweight participants and stronger among those classified with severe overweight/ obesity. Our findings are in line with previous results from this cohort, as well as other studies and reviews. From this cohort, as well as other studies and reviews. BMI/ weight or BMI/ overweight at 2–4 years of age were not strong predictors of later overweight/ obesity; BMI/ weight class at 5–7 years of age were stronger predictors. Others have also observed a stronger degree of tracking from 5 to 6 years of age, especially among subjects classified as obese.

Weight status can fluctuate throughout childhood, ³² and we observed this in our study population. ³⁸ The proportion of those with a high birth weight (>4500 g) that were overweight/ obese at later ages varied between 17–34%; it was lowest at 5–7 years of age and highest in adolescence. The proportion of infants with a normal birth weight that were classified as overweight/ obese at later ages were 10–24%; this proportion was lowest at 2–4 years of age and highest in adolescence (data not shown).

The IOTF reference for BMI classification has low sensitivity but high specificity compared with other references.²¹³³ Hence, the estimated prevalence of overweight/ obesity based on the IOTF reference may be too low and could influence the predictive ability of the classification. On the other hand, a Swedish study comparing different references concluded that the IOTF reference would identify the fewest false positive cases.³³ We also calculated weight and BMI SDS based on the Norwegian reference population.²³ Mean weight SDS at 2-4 and 5-7 years of age was higher; however, mean BMI SDS was comparable to the reference population.²³ GEE analyses give a population-averaged estimate, which generally gives lower and valid estimates when the outcome is dichotomous.⁴⁰ We therefore consider the estimated tracking coefficients in our study to be representative of Norwegian children.

The strengths of this study are the population-based design of the Fit Future study and a high attendance rate, which minimise selection bias. In this region more than 90% of the population in the age group 16–19

years attend upper-secondary school (an entitlement in Norway). Birth variables from the MBRN and measured height and weight data at all ages assures good compliance and good quality data, which limits the risk of information bias. We have longitudinal data over a relatively long time-span from birth in the 1990s up to 18–20 years of age. This study brings updated data on tracking of overweight and obesity from a representative sample of Norwegian children and adolescents. The prevalence of overweight/ obesity at 15–20 years of age may not be a representative estimate for the entire Norwegian adolescent population or populations of different ethnicities. However, the associations found in the tracking analysis are considered to be comparable to other populations with similar characteristics.

The main limitation of this study was the missing data. The percentage of missing main variables from childhood did not differ between sexes or weight classes in adolescence (see online supplementary table 3). More boys than girls were missing at FF2 and we cannot rule out that they were not missing at random.²⁶ This leads to some uncertainty in the estimated prevalence of overweight/ obesity in childhood and at 18-20 years of age. However, analysis of the imputed dataset, the observed-cases dataset and only complete-cases dataset, all gave similar results. A recognised method was used to impute the missing childhood values.²⁷ Mean values were generally lower after imputation, compared with observed cases, but differences were small (see online supplementary table 1 and table 2). Childhood height and weight data are obtained from different health clinics and an overestimation of overweight/ obesity due to instrument error might occur. 41 However, potential misclassification bias is assumed to be non-differential. Factors known to affect overweight/obesity, such as maternal BMI, smoking habits, and socioeconomic status, were not registered in the MBRN for these children. 10 Nor did we have access to information on lifestyle factors from childhood, and the effect of these potential confounding factors cannot be excluded. BMI is a simple measure of obesity and does not separate between fat and muscle mass, although it is widely used in clinical practice and research. BMI in children varies with growth and results need to be interpreted with some caution. More sophisticated measures of body composition would give better measures of adiposity, which is more strongly linked to disease risk in adulthood. Unfortunately, such data were not available for the present study.

Several researchers have recommended early intervention and prevention efforts, some starting even before conception. The first 1000 days are suggested as a critical early-life period. Treatment of obesity among children/adolescents is challenging, which is why establishing healthy habits early in life is of the utmost importance. We aimed to shed light on the most appropriate age to start risk-based preventive efforts. Our findings do not indicate that most children at risk can be identified already at birth or at 2–4 years of age. However,

children with more severe overweight/ obesity and a high risk of later overweight/ obesity could be identified before 5-7 years of age. Development of overweight/ obesity is complex; it involves genetic and multiple environmental factors and is still not fully understood. 1 15 16 Since genetic factors can explain 40-75% of BMI variation in childhood and adolescence, 16 some degree of tracking is probably inevitable. Strong positive associations have been seen between parents and their adolescent offspring's BMI.⁴² Identifying children at risk of later overweight/obesity requires a family approach. The influential role of genetic factors varies with age, 16 and may be prevented by behavioural changes.¹⁵ Prevention of overweight/ obesity therefore requires efforts tailored at addressing parents and children at different ages. More research is needed into preventive and treatment programmes for families and children at different ages.

SUMMARY AND CONCLUSION

Birth weight did not have a strong influence on over-weight/ obesity at 15–20 years of age. Tracking of BMI from early childhood to late adolescence was moderate. Severe overweight/ obesity at 2–4 and 5–7 years of age was a strong predictor of later overweight/ obesity. However, the prevalence of overweight/ obesity increased with age. Prevention efforts should therefore address children of all ages.

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Contributors ASF, NE and AK contributed to the conception, study design and data acquisition of the Fit Futures study. ASF is the principal investigator of the Fit Futures study. E, NE and GS contributed to the conception, design, data collection, analyses and interpretation of data, as well as drafting the manuscript in the present study. Experiormed the statistical analyses. TW gave advice on statistical analysis and especially revised the manuscripts method and result sections. All authors have contributed substantially in interpretation of data and critically revising the manuscript. All authors have read and approved the Enal submitted manuscript.

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Patient consent Participants' consent: obtained.

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SUPPLEMENTARY DOCUMENT – DESCRIPTION OF THE MISSINGNESS PATTERN AND THE IMPUTATION MODEL USED FOR THE PAPER:

The relation between birthweight, childhood body mass index, and overweight and obesity in late adolescence: a longitudinal cohort study from Norway, The Tromsø Study, Fit Futures.

A total of 43% participants (411) were missing one or more of the height/length and weight variables from the birth registry (5% were missing birthweight) and/or childhood measurements (29% missing height/weight at 2-4 years of age, 23% missing height/weight at 5-7 years of age). In 31% (302) of participants data were missing from follow-up at Fit Futures 2, leaving 47.0% (452) of the core study population with five complete measurements.

Due to the relatively large number of missing explanatory variables, we performed multiple imputations to handle missing data. The missingness pattern was checked and found to be arbitrary, thus data was assumed to be missing at random (MAR). Multiple imputations were performed for nine variables from birth and childhood: birthweight, length, gestational age, exact age, and height and weight at 2-4 and 5-7 years of age. In addition, body mass index (BMI) for two girls at 15-17 years of age was imputed. Multiple imputations were performed by chained equations, generating twenty duplicate datasets. The missing values were replaced by imputed values based on the observed information from all five ages. The imputation model included all variables from the final generalised estimating equations analysis, and it included BMI at 15-17 and 18-20 years of age as outcome variables. Calculated variables. To account for a

significant interaction term between sex and BMI at 2-4 years of age, separate imputations were performed for boys and girls. To increase the predictive power of the imputation model, we included auxiliary variables: height, weight, waist circumference and exact age at Fit Futures 1, a dichotomous variable of missing or not at Fit Futures 2, and variables from the birth registry (maternal age at birth, caesarean section, multiple births, maternal disease or diabetes). Variables were imputed using linear regression for continuous variables and predictive mean matching (using 100 nearest neighbours) for variables that had a restricted range or were slightly skewed (the age variables, gestational age, weight, and BMI variables).

Participants with missing BMI at follow-up did not have a significantly higher BMI at Fit Futures 1 vs. those attending Fit Futures 2 (girls: 23.0 vs. 22.2 kg/m²; p= 0.06, boys: 22.7 vs. 22.2 kg/m²; p=0.20). Significantly more boys (39.8%) than girls (22.6%) were missing at Fit Futures 2. We chose not to use imputed missing BMI for those lost to follow-up in Fit Futures 2, since imputation of outcome variables are questionable and we cannot rule out the possibility of missingness that is not random (MNAR). Therefore all analyses performed on the imputed dataset include 961 subjects at 15-17 years of age and complete-case data for 659 subjects at 18-20 years of age.

Supplementary Table S.1. Comparison of the dataset with observed data and the dataset with imputed values (20 multiple imputations) for selected variables for girls and boys of the Fit Futures cohort at birth and four ages up to 18-20 years, The Tromsø Study, Fit Futures

Characteristics			rls		ļ		ys	
		ed data ¹	Impute		Observe			d data ²
	Mean/n	(SD)/%	Mean/n	(SD)/%	Mean/n	(SD)/%	Mean/n	(SD)/%
Birth								
Birthweight (grams)	3454.8	(576.7)	3450.5	(574.8)	3601.0	(590.0)	3603.0	(586.7)
Birth length (cm)	49.4	(2.3)	49.3	(2.4)	50.2	(2.3)	50.1	(2.3)
Gestational age (weeks)	39.7	(1.8)	39.8	(1.8)	39.6	(2.1)	39.6	(2.1)
Ponderal index (kg/m³)	28.73	(2.81)	28.65	(2.88)	28.45	(2.75)	28.40	(2.78)
Body mass index (kg/m ²)	14.19	(1.49)	14.12	(1.55)	14.28	(1.52)	14.24	(1.54)
Birthweight SDS ³	-0.58	(1.74)	-0.58	(1.73)	-0.24	(1.32)	-0.24	(1.31)
Body mass index SDS ³	0.06	(1.05)	0.01	(1.12)	-0.02	(1.04)	-0.46	(1.06)
Birthweight group:								
Low birthweight (<2500 g)	20	4.5 %	22	5.0 %	17	3.6 %	17	4.0 %
Normal birthweight (2500-4500 g)	412	93 %	436	93.0 %	423	90.0 %	444	90.0 %
High birthweight (≥4500 g)	11	2.5 %	11.5	2.0 %	30	6.4 %	31	6.0 %
Small for gestational age	49	11.1%	64	13.5 %	46	9.8%	53	10.8 %
Appropriate for gestational age	356	80.4%	361	77.1 %	375	79.8%	385	78.3 %
Large for gestational age	38	8.6%	44	9.4 %	49	10.4%	54	10.9 %
Maternal age at birth	28.4	(5.3)	28.4	(5.3)	28.1	(5.3)	28.1	(5.3)
2-4 years of age								
Age (years)	2.6	(0.4)	2.5	(0.3)	2.6	(0.4)	2.6	(0.3)
Height (cm)	91.2	(4.6)	91.0	(4.4)	92.7	(4.7)	92.7	(4.4)
Weight (kg)	13.5	(1.7)	13.4	(1.6)	14.1	(1.8)	14.1	(1.7)
Body mass index (kg/m ²)	16.17	(1.39)	16.17	(1.38)	16.37	(1.30)	16.36	(1.31)
Weight SDS ¹	-0.12	(1.11)	-0.13	(1.12)	0.16	(1.16)	0.17	(1.15)
Body mass index SDS ³	0.04	(1.17)	0.03	(1.17)	-0.07	(1.07)	-0.08	(1.09)
5-7 years of age								
Age (years)	6.0	(0.4)	6.0	(0.4)	6.1	(0.4)	6.1	(0.4)
Height (cm)	116.7	(5.3)	116.4	(5.1)	118.3	(5.3)	118.2	(5.2)
Weight (kg)	21.8	(3.8)	21.6	(3.5)	22.2	(3.5)	22.1	(3.3)
Body mass index (kg/m ²)	15.96	(2.04)	15.88	(1.94)	15.82	(1.74)	15.81	(1.72)
Weight SDS ¹	0.0004	(1.10)	-0.05	(1.05)	0.11	(1.02)	0.11	(0.99)
Body mass index SDS ³	-0.06	(1.10)	-0,10	(1.09)	-0.03	(0.98)	-0.03	(1.00)
15-17 years of age								
Age (years)	16.6	(0.4)	-	-	16.6	(0.4)	-	
Height (cm)	164.9	(6.5)	164.9	(6.5)	176.9	(6.7)	-	
Weight (kg)	60.9	(11.5)	60.9	(11.4)	70.2	(14.4)	-	
Body mass index (kg/m²)	22.39	(3.96)	22,39	(3.95)	22.38	(4.17)	-	
18-20 years of age				No imput	ted values			
Age (years)	18.6	(0.4)	-	-	18.7	(0.3)	-	-
Height (cm)	165.7	(6.5)	-	-	179.1	(6.5)	-	-
Weight (kg)	63.2	(12.0)	-	-	75.2	(14.6)	-	-
Body mass index (kg/m ²)	23.02	(4.22)	-	-	23.42	(4.18)	-	-

⁻

¹ In observed data n is varying from 659-961 (328-469 girls and 296-492 boys)

 $^{^{2}}$ Dataset with 20 imputations n= 961(469 girls and 492 boys) in each imputations.

³ SDS= Standard deviation scores according to Norwegian reference data.[23]

SD = Standard deviation

^{- =} No imputed values, mean and SD are equal to the observed data.

Supplementary Table S.2. Weight classes 1 at four ages, The Tromsø Study, Fit Futures - Comparison of observed cases 2 and imputed datasets³

	2-4 years	ears	5-7 years	ears	15-17 years	years	18-20 years
	Observed cases	Imputed data	Observed cases	Imputed data	Observed cases	Imputed data	Observed cases 4
	n= 678	n= 961	n= 736	n= 961	n= 959	n= 961	n= 659
Underweight	14.6	14.8	10.5	10.7	5.1	5.1	4.4
Normal weight	71.3	72.5	71.0	72.4	74.3	74.4	74.7
Girls Overweight	12.5	11.3	13.9	13.1	15.0	14.9	14.3
	1.5	1.3	4.6	3.9	5.6	5.5	9.9
	100.0	100.0	100.0	100.0	100.0	100.0	100.0
Underweight	12.3	13.6	7.3	8.4	7.7	7.7	8.1
Normal weight	78.9	77.4	81.8	80.5	6.89	689	63.9
Boys Overweight	7.7	7.9	7.0	7.6	16.1	16.1	19.9
	1.1	1.0	3.9	3.5	7.3	7.3	8.1
	100.0	100.0	100.0	100.0	100.0	100.0	100.0

¹ Weight classes according to the International Obesity Taskforce (IOTF) age- and sex-specific cut-off values for children 2-18 years of age and the WHO index for adults from age 18-20.[21,22]
² In observed cases n is varying at different ages from 659-959.

³ N=961

⁴ No imputed values, mean and SD are equal to the observed data.

Supplementary table S.3. Distribution of missing data (%) total, by gender and weight classes in two categories at Fit Futures 1 and 2. The Tromsø Study, Fit Futures

missing weight obesity weight obesity obesity a.4 3.8 3.8 3.8 3.8 3.8 3.8 3.8 3.8 3.6 3.8 3.7 3.8 3.7 3.8 3.7 3.8 3.7 3.3 3.2 3.3 3.2 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3 3.3
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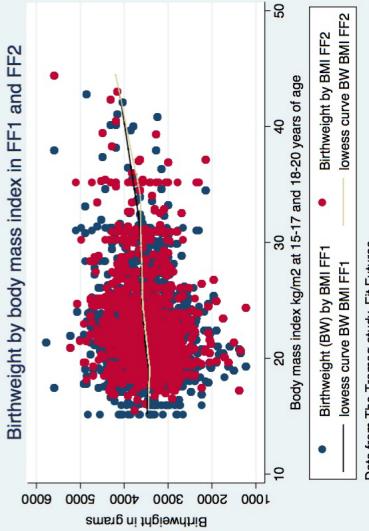
¹ % total missing among 961 cases.

² No significant difference between girls and boys in distribution of total missing for each of the five variables. Birthweight: girls., 5.5 % boys, 4.5 %; Length at birth: girls, 10.7% boys, 7.7%; Gestational age: girls, 16.2% boys 12.6%; BMI 2-4 years: girls, 30.1 boys, 28.9; BMI 5-7 years: girls 25.0% boys 22.0%.
³ Chi-square test, p=0.02 Significant differences between weightclasses in *Italic*.

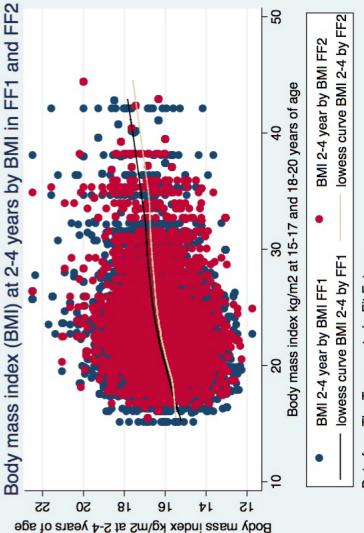
4 Chi-square test, p=0.16

⁵ Chi-square test, p=0.08

BMI, Body mass index kg/m²



Data from The Tromsø study, Fit Futures



Data from The Tromsø study, Fit Futures

