

# Implementing the INTERGROWTH-21<sup>st</sup> fetal growth standards in France: a ‘flash study’ of the Collège Français d’Echographie Foetale (CFEF)

J. J. STIRNEMANN\*†, N. FRIES‡, R. BESSIS‡, M. FONTANGES‡, R. MANGIONE‡  
and L. J. SALOMON\*†‡

\*Assistance Publique-Hôpitaux de Paris, Hôpital Necker-Enfants Malades, Paris, France; †EA FETUS, 7328, Université Paris-Descartes, Paris, France; ‡Collège Français d’Echographie Foetale, CFEF, France

**KEYWORDS:** biometry; congenital abnormalities; cross-sectional studies; epidemiology; fetal development; growth charts; pregnancy; ultrasound

## ABSTRACT

**Objectives** To assess potential differences in fetal size between the French population and the international population from the INTERGROWTH-21<sup>st</sup> (IG-21<sup>st</sup>) Project and to measure the impact of switching to the IG-21<sup>st</sup> reference standards for fetal size.

**Methods** This was a nationwide cross-sectional study of fetal ultrasound biometry. Low-risk singleton pregnancies were recruited prospectively within the network of the national French College of Fetal Ultrasound, CFEF, over a 6-week period. Further selection was performed based on the criteria of the IG-21<sup>st</sup> Project in order to obtain a comparable population. Head circumference (HC) was used as the main fat-free skeletal measure of growth for comparison of French fetal size with that of the IG-21<sup>st</sup> population. The impact of switching to the IG-21<sup>st</sup> fetal growth standards was quantified by comparing Z-scores calculated using the IG-21<sup>st</sup> standards with those calculated using locally derived reference ranges for HC, abdominal circumference (AC) and femur length (FL).

**Results** Following selection, 4858 cases were analyzed. The distribution of HC demonstrated clear similarity between our French population and the IG-21<sup>st</sup> population: our observed centile curves closely matched those of IG-21<sup>st</sup> and the Z-scores were close to 0 across gestational age. The IG-21<sup>st</sup> standards performed as well as did locally derived charts in terms of screening for small-for-gestational age by AC, while they identified significantly fewer small FL values than were expected and than did the locally derived charts.

**Conclusions** Under strict selection criteria, fetal size in France is similar to that of the international population

used in the IG-21<sup>st</sup> Project. The discrepancies in FL are unlikely to impact on prenatal management. Therefore, switching from locally derived reference ranges to the IG-21<sup>st</sup> standards appears to be a safe option. Copyright © 2016 ISUOG. Published by John Wiley & Sons Ltd.

## INTRODUCTION

Screening for intrauterine growth restriction (IUGR), mainly by ultrasound measurements, is one of the most important components of antenatal care<sup>1</sup>. Unfortunately, the antenatal detection rate of IUGR confirmed at birth is low, even in high-risk subpopulations<sup>2,3</sup>.

The low detection rate can be explained in part by both the choice of growth chart, from amongst the more than 80 available, on which to base judgments about the fetal measurements<sup>4</sup> and the lack of international consensus regarding the diagnostic criteria for IUGR. This is in contrast with the pediatric evaluation of growth in infants and children, as the World Health Organization (WHO) has produced a single set of international growth standards for use worldwide<sup>5,6</sup>.

Conceptually similar to the WHO Child Growth Standards, the International Fetal Growth Standards were published in 2014 as part of the INTERGROWTH-21<sup>st</sup> Project<sup>7</sup>. These standards were developed from multi-ethnic populations worldwide, whose health, nutrition and care needs were largely met. As opposed to previous locally produced references, international standards have the potential to improve the detection of growth disturbances and, as a result, perinatal outcomes, by standardizing the diagnostic approach to IUGR and macrosomia.

Correspondence to: Prof. L. J. Salomon, Hôpital Necker-Enfants Malades, 149, rue de Sèvres, 75743 Paris Cedex 15, France (e-mail: laurentsalomon@gmail.com)

Accepted: 5 August 2016

The present cross-sectional study was undertaken to: (1) evaluate whether fetal biometry measurements from pregnant French women, using inclusion criteria similar to those of the INTERGROWTH-21<sup>st</sup> Project, produce patterns comparable to the INTERGROWTH-21<sup>st</sup> standards and (2) quantify the implications of using the INTERGROWTH-21<sup>st</sup> standards in the French population.

## METHODS

Flash studies<sup>8,9</sup> are pragmatic, short and very focused studies, conducted without modifying routine clinical practice and at no extra cost. They have both a scientific and an educational purpose and are conducted in France across the countrywide network of sonographers who are members of the French College of Fetal Ultrasound (College Français d'Échographie Foetale (CFEF)).

We invited sonographers first to take an online training course ([www.cfef.org](http://www.cfef.org)) that reviewed the aims of the study, the inclusion criteria, the methodology for taking the measurements and the biometric quality control criteria<sup>10,11</sup>. Only sonographers who completed the course and passed the final test were eligible to participate in the study.

Pregnant women contributed with a single measure and were included prospectively and consecutively over a fixed study period of 6 weeks. Those included had a singleton pregnancy without congenital malformations and were scanned between 8 September 2014 and 18 October 2014, based on a routine indication: in France, three scans are planned for all pregnancies, at 11–14 weeks, 21–24 weeks and 30–34 weeks, as part of standard antenatal care. Pregnancy dating was based on crown–rump length measurement in the first trimester, as recommended by the French College of Obstetrics and Gynaecology, CNGOF<sup>12,13</sup>.

The following information was collected: parity, gravidity, maternal age, body mass index (BMI), socioprofessional category, presence of comorbidities, smoking, alcohol consumption, gestational age at scan and fetal head circumference (HC), abdominal circumference (AC) and femur length (FL). Fetal measurements were taken as described elsewhere<sup>10,11</sup>. Briefly, HC was measured in an axial plane of the fetal head at the level at which the continuous midline echo is broken by the septum pellucidum in the anterior third, using the ellipse tool with calipers placed on the outer edges of the skull. AC was measured in an axial plane of the fetal abdomen, just above the level of the cord insertion, using the ellipse tool. FL was measured on a plane showing the entire femoral diaphysis, with both ends clearly visible and at an angle < 45° to the horizontal. From 30 weeks' gestation onwards, particular care was taken not to include the epiphysis.

These measurements, collected prospectively, constituted our primary database. Within this dataset, a subsample of women was selected who met, as closely as possible, the strict inclusion criteria of the Fetal Growth Longitudinal Study (FGLS) of the INTERGROWTH-21<sup>st</sup> Project<sup>7</sup>.

Briefly, the FGLS participants were selected first at the level of a geographic area and then at the individual level within each study site<sup>7</sup>. The principal geographic-level criteria were met because all deliveries in France occur in health institutions, all of which are located at an altitude of 1600 m or lower. For the individual-level criteria, similar to the FGLS, we included women who were at low risk of adverse maternal outcome (e.g. absence of smoking, diabetes, hypertension, renal disease, maternal symptoms of pre-eclampsia or other medical conditions associated with fetal growth disturbances) and perinatal outcome (patients not referred for suspicion of fetal anomaly, history of fetal anomaly or IUGR). In the final analysis we included only women aged  $\geq 18$  years and  $\leq 35$  years, who had BMI  $\geq 18.5$  kg/m<sup>2</sup> and  $< 30$  kg/m<sup>2</sup>, who had no clinically relevant obstetric or gynecological history and who initiated antenatal care before 14 weeks' gestation. We also excluded *a-priori* measurements below or above  $-5$  and  $+5$  SD, respectively.

Our data were compared to the INTERGROWTH-21<sup>st</sup> standards<sup>7</sup> using a method similar to that described in the original studies that generated the WHO Child Growth and INTERGROWTH-21<sup>st</sup> standards<sup>7,14</sup>. To assess the similarity of size between French fetuses and those used to create the INTERGROWTH-21<sup>st</sup> standards, we decided to use HC, the main fat-free skeletal measure of linear growth, for comparison<sup>14–16</sup>. First, the charts obtained from fitting an 'INTERGROWTH-like' fractional polynomial regression model (i.e. the same functional form defined by the set of fractional polynomial powers) to our newly collected data were compared visually to those from the INTERGROWTH-21<sup>st</sup> standards<sup>7</sup>. Similarities in the French and INTERGROWTH-21<sup>st</sup> populations with respect to the fetal HC were identified based on standardized differences, at various gestational-age windows. Standardized differences were defined as the difference between the mean of our French sample and the mean of the pooled INTERGROWTH-21<sup>st</sup> data, expressed as units of the pooled SD observed in the corresponding INTERGROWTH-21<sup>st</sup> study. Standardized differences were therefore similar to a Z-score. A standardized difference value < 0.5 was prespecified as reflecting adequately that the French population could have been included in the pooled dataset used to construct the international standards<sup>7,14</sup>.

We also evaluated the impact of switching from the locally derived reference charts that are presently recommended for use in France<sup>17,18</sup> to the INTERGROWTH-21<sup>st</sup> standards<sup>7</sup> by calculating the mean and SD of computed Z-scores based on these two different references. A mean Z-score < 0.5 was again considered to demonstrate good concordance between observed and reference-based predicted values. A SD of Z-scores close to 1 (i.e. between 0.8 and 1.2) was considered to reflect good concordance between observed and reference-based predicted dispersion of values. Finally, we calculated the proportions of fetuses below the 3<sup>rd</sup> and 10<sup>th</sup> centiles and above the 90<sup>th</sup> and 97<sup>th</sup> centiles, using both the INTERGROWTH-21<sup>st</sup> and the

locally derived charts. Statistical analyses were performed using Stata 9.2 for Windows (StataCorp LP, College Station, TX, USA), Statistica (StatSoft, Inc., 2001) and R (Foundation for Statistical Computing, Vienna, Austria; www.R-project.org).

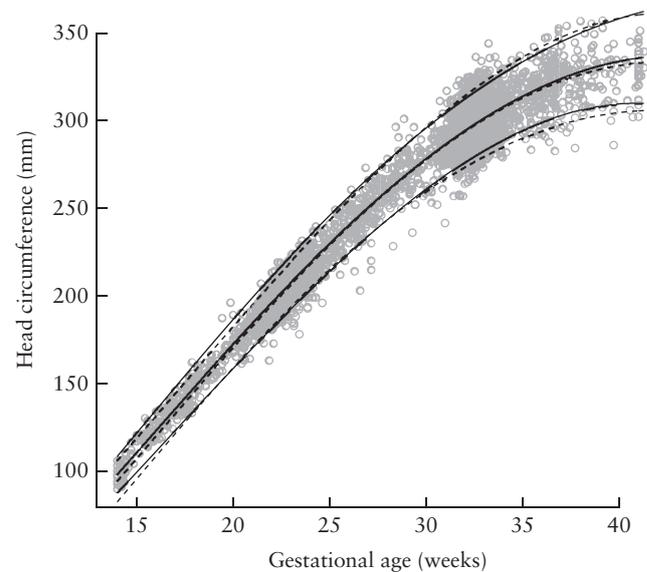
## RESULTS

There were 160 sonographers who agreed to participate, of whom 120 fulfilled the inclusion criteria of the study. During the study period they performed a total of 8784 scans, of which 1689 were excluded for at least one of the following reasons: maternal condition ( $n = 1090$ ); maternal symptoms ( $n = 311$ ); history of previous small baby, suspected fetal anomaly or IUGR ( $n = 599$ ). A further 1248 (14.2%) were excluded because maternal age was  $> 35$  years ( $n = 1220$ ) or  $< 18$  years ( $n = 28$ ), and 961 (10.9%) were excluded because BMI was  $< 18.5 \text{ kg/m}^2$  ( $n = 193$ ) or  $> 30 \text{ kg/m}^2$  ( $n = 768$ ). Lastly, 28 examinations were excluded because measurement values were below or above  $-5$  or  $+5$  SD, respectively.

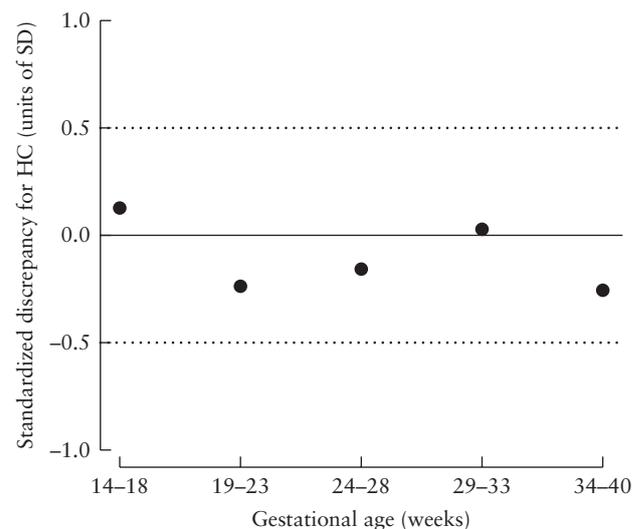
From the original 8784 scans, our analysis thus included data from 4858 (55.3%) independent ultrasound examinations in low-risk women and fetuses across gestation, i.e. an 'INTERGROWTH-21<sup>st</sup> FGLS-like' population. Figure 1 presents the INTERGROWTH-21<sup>st</sup> standard for HC, the main, fat-free, skeletal measure of linear growth used in the INTERGROWTH-21<sup>st</sup> Project for comparisons across populations<sup>7</sup>, superimposed on the individual HC values for our French subpopulation. The centiles estimated from our French subpopulation using the same statistical methodology as that of the FGLS (i.e. same set of fractional polynomial powers) are also superimposed. Visual inspection of the two sets of centiles demonstrates clear similarity between the two populations, suggesting that fetal HC size in the French population is comparable to that observed in the FGLS. This was further confirmed by analyzing the standardized differences of HC, which remained close to 0 and within  $\pm 0.5$  when computed at different gestational-age windows across pregnancy (Figure 2).

To assess the impact of switching from local charts to the INTERGROWTH-21<sup>st</sup> charts, we also estimated the individual Z-scores for AC, FL and HC in our sample of the French population based on each of the two sets of charts<sup>7,17</sup>. The mean Z-scores remained between  $-0.5$  and  $+0.5$  for all measurements at all gestational-age windows for both the French locally-derived charts and the INTERGROWTH-21<sup>st</sup> charts (Table 1) in all except two of the 15 comparisons for each of the two charts.

In any distribution of Z-scores the expected SD is 1; we therefore compared the actual SDs obtained using the INTERGROWTH-21<sup>st</sup> charts and local charts. The INTERGROWTH-21<sup>st</sup> charts gave a SD of 1.01 for HC, 0.99 for AC and 1.00 for FL, and the local charts gave SDs of 0.75, 0.75 and 0.78, respectively; thus, the INTERGROWTH-21<sup>st</sup> charts were closer to the expected population distribution than were the local charts. This was further confirmed by the fact that two SD values



**Figure 1** Observed values of head circumference among 4858 fetuses at low risk of adverse maternal and perinatal outcome from current overall prospective French national flash study, with 3<sup>rd</sup>, 50<sup>th</sup> and 97<sup>th</sup> centiles estimated using an 'INTERGROWTH-21<sup>st</sup>-like' model (dashed curves) compared with INTERGROWTH-21<sup>st</sup> international growth charts (solid curves)<sup>7</sup>.



**Figure 2** Standardized differences in head circumference (HC) between those obtained in current French national flash study and INTERGROWTH-21<sup>st</sup> charts<sup>7</sup> across gestational age.

out of 15 comparisons were beyond the prespecified cut-off for the INTERGROWTH-21<sup>st</sup> charts *vs* 12 values using the locally derived charts (Table 1). This resulted in different proportions of fetuses below the 3<sup>rd</sup> and 10<sup>th</sup> centiles and above the 90<sup>th</sup> and 97<sup>th</sup> centiles, using each chart (Table 2). Specifically, the INTERGROWTH-21<sup>st</sup> standard identified a proportion of HCs below the 3<sup>rd</sup> and 10<sup>th</sup> centiles closer to the expected value than did the locally derived charts. Despite the differences in the observed mean and SD of the distribution of Z-scores according to each chart, the proportions of ACs below the 3<sup>rd</sup> and 10<sup>th</sup> centiles according to each chart were

**Table 1** Z-scores for study population of 4858 low-risk singleton French pregnancies estimated based on French fetal size charts (local)<sup>17</sup> and on INTERGROWTH-21<sup>st</sup> international growth charts (IG-21<sup>st</sup>)<sup>7</sup>

Gestational age	n	Mean of Z-scores						SD of Z-scores					
		Mean local			Mean IG-21 <sup>st</sup>			SD local			SD IG-21 <sup>st</sup>		
		HC	AC	FL	HC	AC	FL	HC	AC	FL	HC	AC	FL
14 to 18 weeks	196	-0.42	-0.67*	-0.66*	0.13	0.46	0.08	0.74*	1.05	0.78*	0.85	1.27*	0.96
19 to 23 weeks	2123	0.08	0.08	-0.27	-0.24	0.49	0.32	0.62*	0.65*	0.70*	0.82	0.90	0.92
24 to 28 weeks	333	0.19	-0.01	-0.26	-0.16	0.33	0.47	0.85	0.77*	0.86	1.16	1.07	1.13
29 to 33 weeks	1826	0.33	0.20	-0.36	0.03	0.43	0.59*	0.79*	0.73*	0.77*	1.14	0.98	0.98
34 to 40 + 6 weeks	380	0.38	0.13	-0.43	-0.25	-0.12	0.53*	0.86	0.93	1.07	1.19	1.13	1.25*
All	4858	0.18	0.09	-0.33	-0.11	0.41	0.44	0.75*	0.75*	0.78*	1.01	0.99	1.00

\*Mean values below -0.5 or above +0.5 and SDs below 0.8 or above 1.2 are indicated. AC, abdominal circumference; FL, femur length; HC, head circumference.

**Table 2** Percentages of 4858 low-risk French singleton pregnancies with fetus under 3<sup>rd</sup> or 10<sup>th</sup> centile or above 90<sup>th</sup> or 97<sup>th</sup> centile using international INTERGROWTH-21<sup>st</sup> prescriptive charts (IG-21<sup>st</sup>)<sup>7</sup> and using locally derived French charts (local)<sup>17</sup>

	< 3 <sup>rd</sup> centile (%)		< 10 <sup>th</sup> centile (%)		> 90 <sup>th</sup> centile (%)		> 97 <sup>th</sup> centile (%)	
	IG-21 <sup>st</sup>	Local	IG-21 <sup>st</sup>	Local	IG-21 <sup>st</sup>	Local	IG-21 <sup>st</sup>	Local
HC	3.71	0.70	11.20	2.28	7.95	6.81	2.66	1.50
AC	1.17	0.89	4.06	3.36	18.46	5.25	6.81	1.11
FL	1.03	2.08	3.13	9.18	19.06	2.37	7.43	0.56

AC, abdominal circumference; FL, femur length; HC, head circumference.

comparable. However, the proportions of ACs above the 90<sup>th</sup> and 97<sup>th</sup> centiles showed significant differences, both far from the expected value. Lastly, the proportion of small FLs was closer to the expected value using the locally derived chart than using the INTERGROWTH-21<sup>st</sup> standard, whereas the proportion of large FLs was overestimated using the INTERGROWTH-21<sup>st</sup> standard and underestimated using the locally derived chart.

## DISCUSSION

This study shows that, in a French population selected using similar criteria to those of the FGLS at the individual level, HC measures closely matched the INTERGROWTH-21<sup>st</sup> standard<sup>7</sup>. Indeed, the patterns of measurements obtained across gestation were very similar (Figure 1). This suggests that French fetuses from low-risk pregnancies do not differ from those included in the INTERGROWTH-21<sup>st</sup> Project.

Our results suggest that the routine implementation in France of such standards, based on a normative approach and international sampling from geographically and ethnically diverse populations, would be legitimate<sup>7</sup> as a first level of screening. Furthermore, considering the multiethnic characteristics of present-day populations in the country, it is the most practical tool to be used during routine antenatal care. This also confirms the international nature of skeletal fetal growth, when health, nutrition and care needs are met, as is the case for the general French population. Finally, the experience gained in the implementation, in more than 140 countries, of the postnatal WHO Child Growth Standards, whose design and methods were almost identical to those

of the INTERGROWTH-21<sup>st</sup> Project, also favors the widespread use of the new INTERGROWTH-21<sup>st</sup> standards<sup>7</sup>.

Our study shows that both the international standards and the locally derived charts appropriately describe mean fetal size, as mean Z-scores were close to zero in all cases. However, there were differences in the mean Z-score values between the two sets of charts and, consequently, discrepancies in the proportions of small and large measurements, which could be explained by several factors. A possible effect due to ethnic differences or differences in population background is still a matter of debate<sup>19,20</sup>, but is very unlikely since the INTERGROWTH-21<sup>st</sup> cohort includes countries such as Italy and the UK, with demographic characteristics very close to those of France. However, socioeconomic status has been shown to be a main determinant of growth<sup>21,22</sup> and this variable was not considered at an individual level in our population. The remaining differences could be attributed to ultrasound measurement methods. This is particularly relevant for the measurement of HC, which was previously obtained with calipers placed on the middle of the skull bone<sup>17</sup>, whereas recent recommendations<sup>12</sup> suggest caliper placement on the outer edge, as in the new INTERGROWTH-21<sup>st</sup> international standard<sup>7</sup>.

The greatest discrepancies were found for FL: compared with the locally derived reference, the INTERGROWTH-21<sup>st</sup> chart 'underestimated' the proportion of small femurs. In practice, this would have very little, if any, impact: it should not affect our ability to detect skeletal dysplasias, will not affect screening for IUGR, which relies on AC, HC or estimated fetal weight,

and has long been abandoned for screening for Down syndrome.

Another important factor that may explain the observed differences is the recent technological changes in ultrasound machines. This is particularly relevant for FL measurement<sup>23</sup>, because the thinning of the modern ultrasound beam leads to shorter measurements<sup>23</sup>. This may explain the lower mean FL values in the INTERGROWTH-21<sup>st</sup> standard<sup>7</sup> compared with older references constructed using data obtained with outdated equipment<sup>17</sup>.

Finally, all sonographers involved in the flash study were accustomed to working with the former French references (expected-value bias), and these same references were displayed on the screen of their ultrasound machine at the time at which measurements were being performed (in the INTERGROWTH-21<sup>st</sup> Project, sonographers were blinded to the measurements). It is therefore likely that this tended to bias measurements towards the values 'expected' based on the locally produced charts, possibly explaining the trend for the means of Z-scores to be closer to zero with the locally produced charts than with the new international standards.

The adequacy of the observed and expected SDs is critical and our results suggest better performance when using the new INTERGROWTH-21<sup>st</sup> standards. The locally produced reference ranges currently in use in France<sup>17</sup> are descriptive, i.e. they were derived from the general population, with no growth-related or outcome-related exclusion criteria, therefore capturing both normally and abnormally grown fetuses. The ability of these locally derived references to identify abnormal biometric measurements may therefore be hampered, reflected in the reported limited ability to pick up true IUGR cases in France<sup>18</sup>. Conversely, the new international prescriptive standards for fetal size<sup>7</sup>, which defines the expected fetal size under optimal conditions, have smaller dispersion (i.e. a SD closer to the theoretical value of 1) than do locally produced references and thus could improve the sensitivity of IUGR screening.

Limitations of this study, however, should be considered. The sonographers who volunteered and were eventually enrolled into this study may not fully represent the general population of sonographers. They performed biometric measurements and compared them to existing local references in a non-blinded fashion, which may have introduced a bias towards the expected values. Therefore, given the setting of our study, under real-life non-experimental conditions, we did not strictly reproduce the experimental setting of the INTERGROWTH-21<sup>st</sup> study. Using routine cross-sectional ultrasound data instead of longitudinal follow-up data may also have affected adversely our assessment of the INTERGROWTH-21<sup>st</sup> charts: (1) between the routine time-points, data were scarce; (2) since the study was not longitudinal, drop-outs were not recognized, which could have biased our population if such drop-outs were not random but related to growth disorders; (3) as this study aimed to measure

prenatal biometry adequacy, we did not collect perinatal outcomes.

In conclusion, our study demonstrates that a French pregnant subpopulation, selected using individual criteria closely matching those used in the INTERGROWTH-21<sup>st</sup> Project, produces fetal size patterns comparable to the INTERGROWTH-21<sup>st</sup> international standards in terms of skeletal ultrasound parameters. We suggest that these international standards can be used in France as part of routine ultrasound practice. However, it is of paramount importance to separate the choice of a standard based on data quality and reference population selection from any clinical screening procedures based on centile cut-offs. Therefore, the implementation of such standards deserves close monitoring: since they were derived from a highly selected population, they are likely to increase detection rates of growth abnormalities in the general population for a given centile cut-off. Therefore, biometry-based screening procedures should be adapted accordingly, based on the future appraisal of the implementation of these charts in the general population, in terms of detection rates and perinatal outcome.

## REFERENCES

1. Conde-Agudelo A, Papageorghiou AT, Kennedy SH, Villar J. Novel biomarkers for predicting intrauterine growth restriction: a systematic review and meta-analysis. *BJOG* 2013; **120**: 681–694.
2. Bricker L, Medley N, Pratt JJ. Routine ultrasound in late pregnancy (after 24 weeks' gestation). *Cochrane Database Syst Rev* 2015; **6**: CD001451.
3. Sylvan K, Ryding EL, Rydhstroem H. Routine ultrasound screening in the third trimester: a population-based study. *Acta Obstet Gynecol Scand* 2005; **84**: 1154–1158.
4. Ioannou C, Talbot K, Ohuma E, Sarris I, Villar J, Conde-Agudelo A, Papageorghiou A. Systematic review of methodology used in ultrasound studies aimed at creating charts of fetal size. *BJOG* 2012; **119**: 1425–1439.
5. de Onis M. Update on the implementation of the WHO child growth standards. *World Rev Nutr Diet*. 2013; **106**: 75–82. doi: 10.1159/000342550.
6. de Onis M. 4.1 The WHO Child Growth Standards. *World Rev Nutr Diet* 2015; **113**: 278–294.
7. Papageorghiou AT, Ohuma EO, Altman DG, Todros T, Cheikh Ismail L, Lambert A, Jaffer YA, Bertino E, Gravett MG, Purwar M, Noble JA, Pang R, Victora CG, Barros FC, Carvalho M, Salomon LJ, Bhutta ZA, Kennedy SH, Villar J, International Fetal and Newborn Growth Consortium for the 21st Century (INTERGROWTH-21st). International standards for fetal growth based on serial ultrasound measurements: the Fetal Growth Longitudinal Study of the INTERGROWTH-21st Project. *Lancet* 2014; **384**: 869–879.
8. Kuleva M, Fries N, Castaing O, Moeglin D, Salomon LJ. 'Flash study' on chorionicity determination from ultrasound images at 11–14 weeks' gestation in twin pregnancies. *Ultrasound Obstet Gynecol* 2013; **41**: 471–472.
9. Kuleva M, Castaing O, Fries N, Bernard J-P, Bussi eres L, Fontanges M, Moeglin D, Salomon LJ. A standardized approach for the assessment of the lower uterine segment at first trimester by transvaginal ultrasound: a flash study. *J Matern-Fetal Neonatal Med* 2016; **29**: 1376–1381.
10. Papageorghiou AT, Sarris I, Ioannou C, Todros T, Carvalho M, Pilu G, Salomon LJ, International Fetal and Newborn Growth Consortium for the 21st Century. Ultrasound methodology used to construct the fetal growth standards in the INTERGROWTH-21st Project. *BJOG* 2013; **120** (Suppl 2): 27–32.
11. Salomon LJ, Bernard JP, Duyme M, Doris B, Mas N, Ville Y. Feasibility and reproducibility of an image-scoring method for quality control of fetal biometry in the second trimester. *Ultrasound Obstet Gynecol* 2006; **27**: 34–40.
12. Salomon LJ. [How to date pregnancy?]. *J Gyn ecologie Obst etricque Biol Reprod* 2011; **40**: 726–733.
13. Papageorghiou AT, Kennedy SH, Salomon LJ, Ohuma EO, Cheikh Ismail L, Barros FC, Lambert A, Carvalho M, Jaffer YA, Bertino E, Gravett MG, Altman DG, Purwar M, Noble JA, Pang R, Victora CG, Bhutta ZA, Villar J, for the International Fetal and Newborn Growth Consortium for the 21st Century (INTERGROWTH-21st). International standards for early fetal size and pregnancy dating based on ultrasound measurement of crown–rump length in the first trimester of pregnancy. *Ultrasound Obstet Gynecol* 2014; **44**: 641–648.
14. Villar J, Papageorghiou AT, Pang R, Ohuma EO, Cheikh Ismail L, Barros FC, Lambert A, Carvalho M, Jaffer YA, Bertino E, Gravett MG, Altman DG, Purwar M, Frederick IO, Noble JA, Victora CG, Bhutta ZA, Kennedy SH, International Fetal and Newborn Growth Consortium for the 21st Century (INTERGROWTH-21st). The likeness of fetal growth and newborn size across non-isolated populations in the

- INTERGROWTH-21st Project: the Fetal Growth Longitudinal Study and Newborn Cross-Sectional Study. *Lancet Diabetes Endocrinol* 2014; **2**: 781–792.
15. WHO Multicentre Growth Reference Study Group. Assessment of differences in linear growth among populations in the WHO Multicentre Growth Reference Study. *Acta Paediatr Suppl* 2006; **450**: 56–65.
  16. Villar J, Altman DG, Purwar M, Noble JA, Knight HE, Ruyan P, Cheikh Ismail L, Barros FC, Lambert A, Papageorgiou AT, Carvalho M, Jaffer YA, Bertino E, Gravett MG, Bhutta ZA, Kennedy SH, International Fetal and Newborn Growth Consortium for the 21st Century. The objectives, design and implementation of the INTERGROWTH-21st Project. *BJOG* 2013; **120** (Suppl 2): 9–26.
  17. Salomon LJ, Duyme M, Crequat J, Brodaty G, Talmant C, Fries N, Althuser M. French fetal biometry: reference equations and comparison with other charts. *Ultrasound Obstet Gynecol* 2006; **28**: 193–198.
  18. Vayssière C, Sentilhes L, Ego A, Bernard C, Cambourieu D, Flamant C, Gascoin G, Gaudineau A, Grangé G, Houfflin-Debarge V, Langer B, Malan V, Marcocelles P, Nizard J, Perrotin F, Salomon L, Senat M-V, Serry A, Tessier V, Truffert P, Tsatsaris V, Arnaud C, Carbone B. Fetal growth restriction and intra-uterine growth restriction: guidelines for clinical practice from the French College of Gynaecologists and Obstetricians. *Eur J Obstet Gynecol Reprod Biol* 2015; **193**: 10–18.
  19. Kato N. Ethnic differences in genetic predisposition to hypertension. *Hypertens Res* 2012; **35**: 574–581.
  20. Cooper RS, Kaufman JS, Ward R. Race and genomics. *N Engl J Med* 2003; **348**: 1166–1170.
  21. Bhandari N, Bahl R, Taneja S, de Onis M, Bhan MK. Growth performance of affluent Indian children is similar to that in developed countries. *Bull World Health Organ* 2002; **80**: 189–195.
  22. Stevens GA, Finucane MM, Paciorek CJ, Flaxman SR, White RA, Donner AJ, Ezzati M, Nutrition Impact Model Study Group (Child Growth). Trends in mild, moderate, and severe stunting and underweight, and progress towards MDG 1 in 141 developing countries: a systematic analysis of population representative data. *Lancet* 2012; **380**: 824–834.
  23. Økland I, Bjåstad TG, Johansen TF, Gjessing HK, Grøttum P, Eik-Nes SH. Narrowed beam width in newer ultrasound machines shortens measurements in the lateral direction: fetal measurement charts may be obsolete. *Ultrasound Obstet Gynecol* 2011; **38**: 82–87.