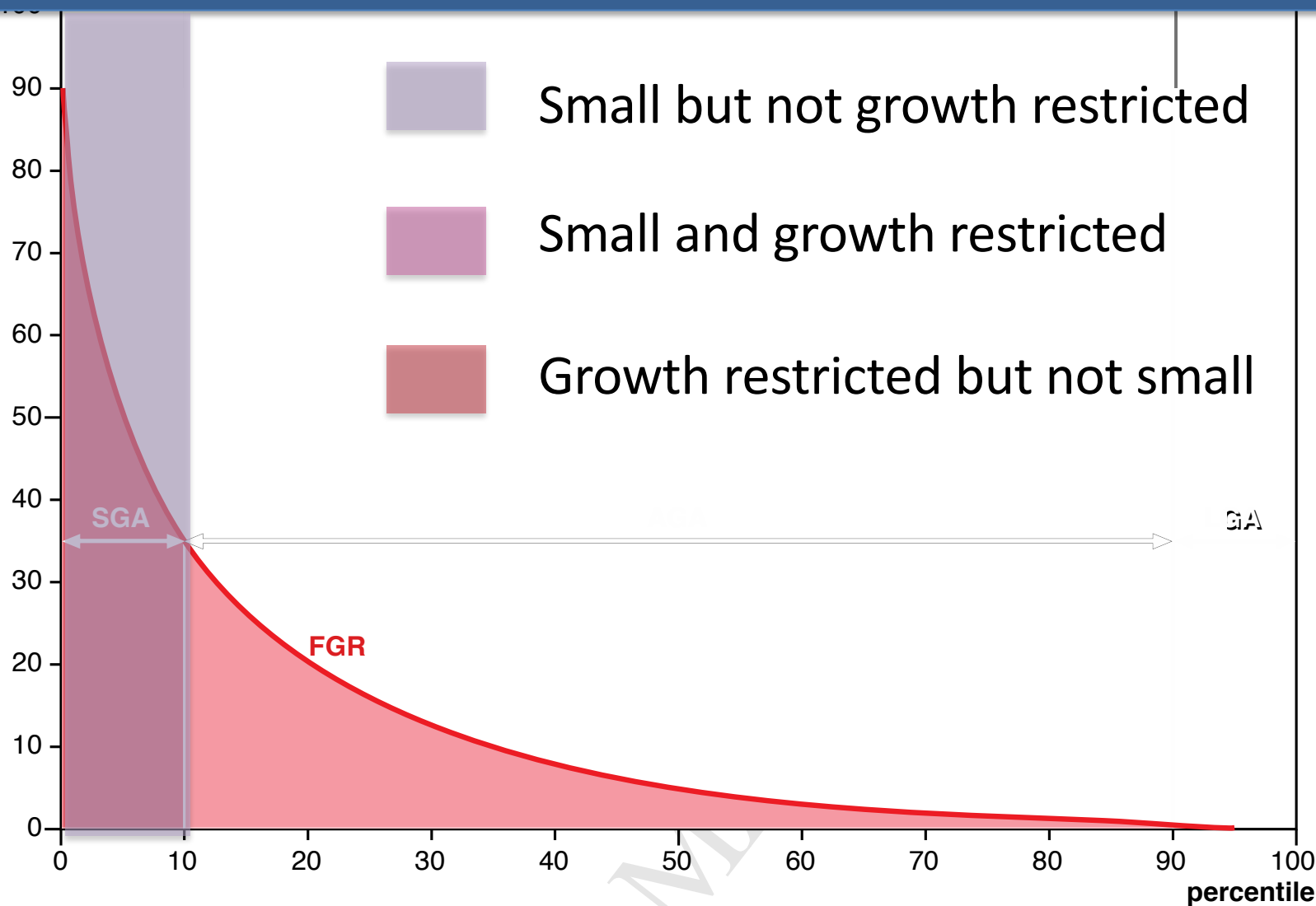


# Choosing a growth standard

Clare Whitehead

# Optimising detection of FGR not SGA



***Which chart you choose will determine what % of babies in your population are SGA and FGR*** Ganzevoort, AJOG 2018

# Which growth chart: birthweight vs fetal weight

Babies in the NICU are not the same.....

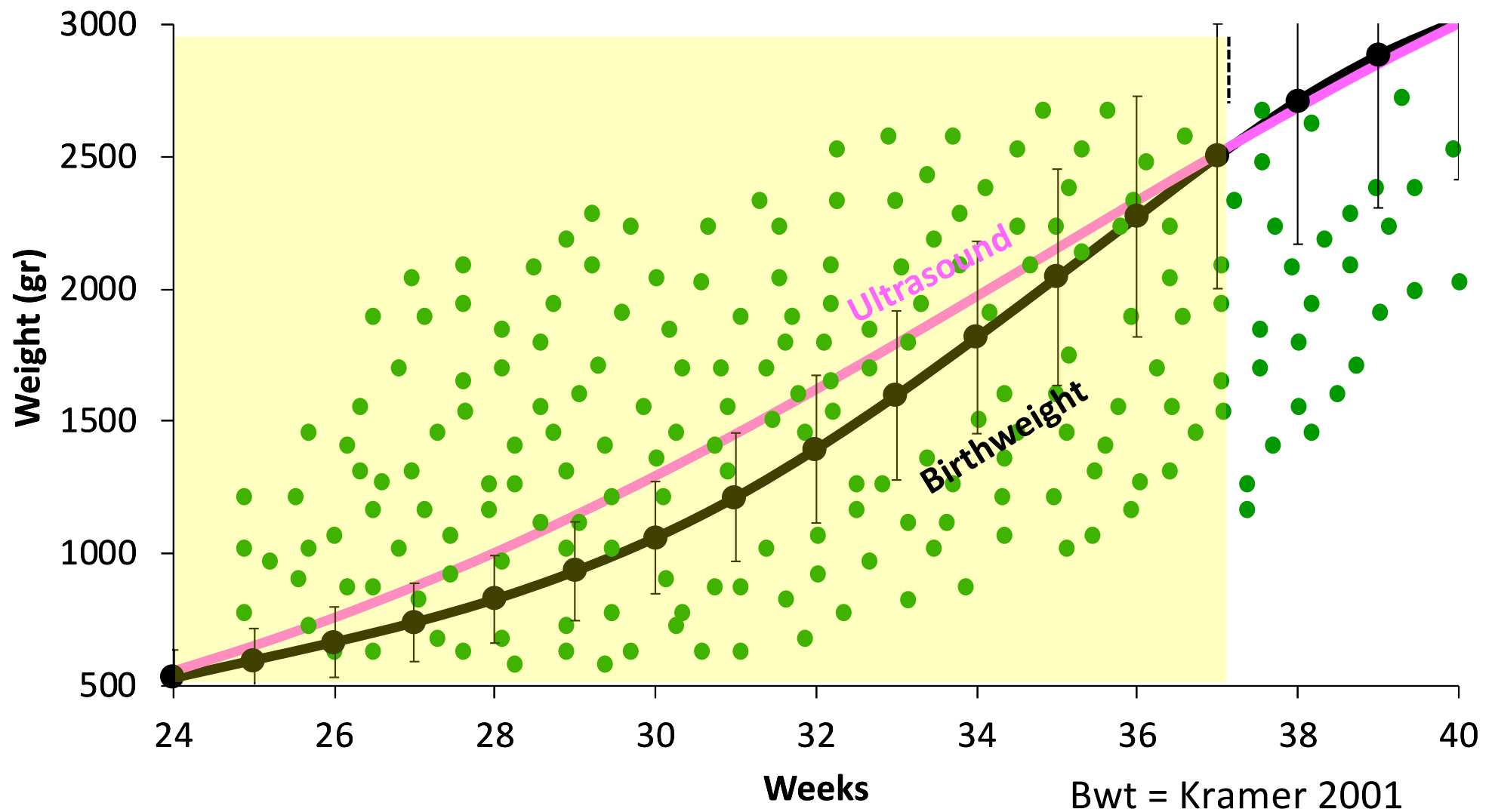


As babies that remain in-utero until term!



Live birth charts will systematically underdiagnose FGR prior to term

# Which growth chart: birthweight vs fetal weight





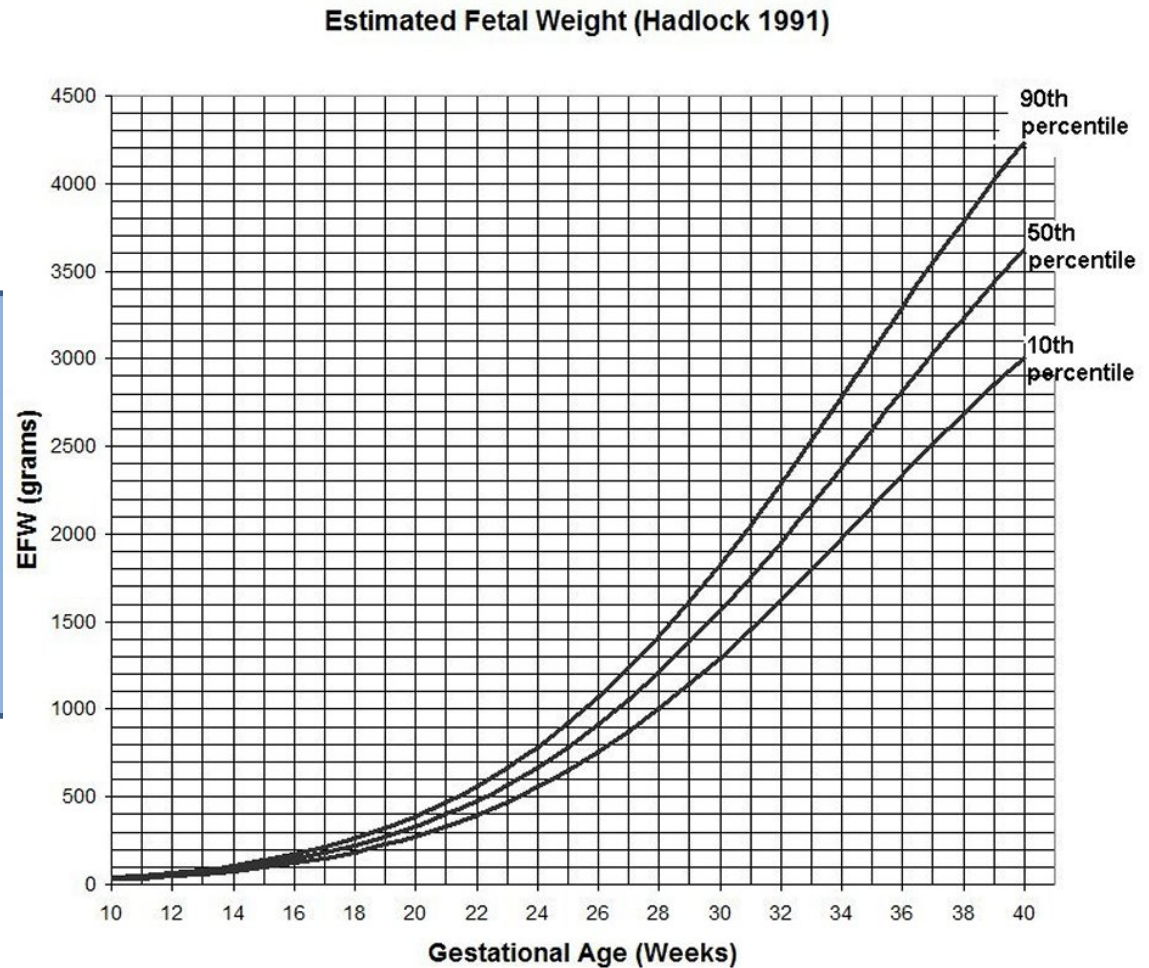
# Which growth chart: *Standard vs Reference charts*

Growth *standards* charts describe how a baby *should* grow.....based on data from *only healthy pregnancies*

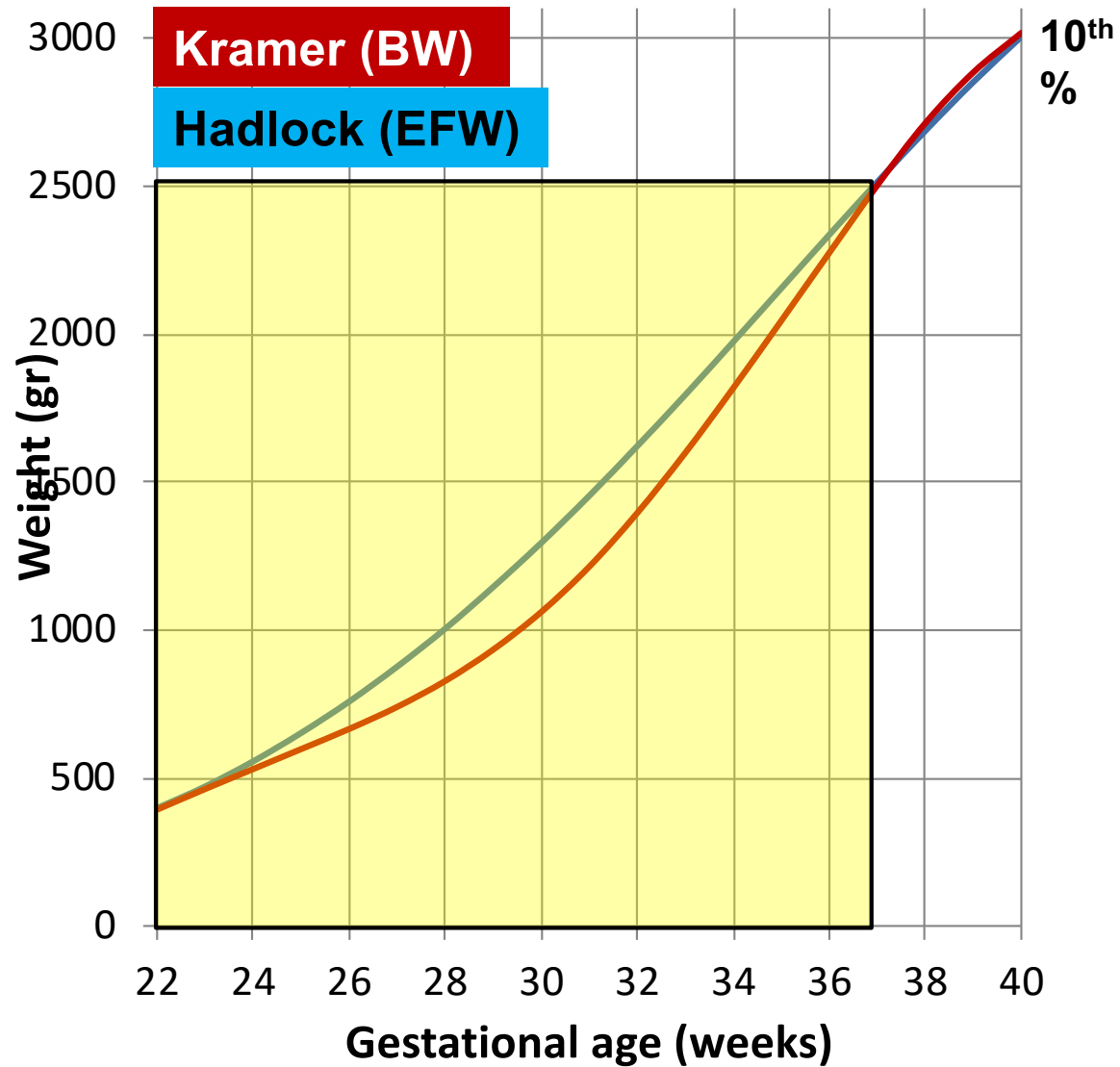
Whereas growth *reference* charts describe how *all babies in a population* grow including those that subsequently develop complications

# Hadlock Charts

- **USA 1991**
- 392 women
- All caucasian
- Single center in Texas
- Only 1 USS per fetus



# Growth charts



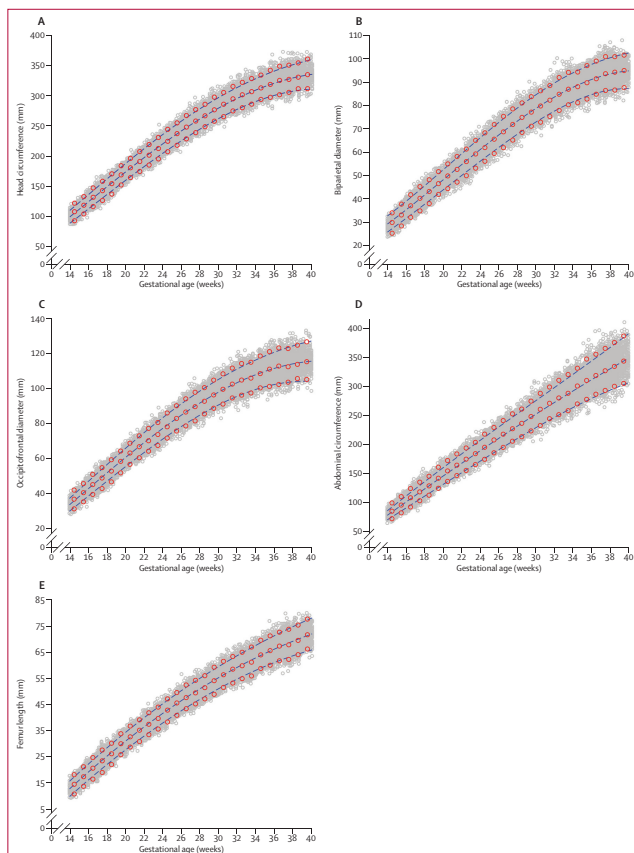
# Intergrowth 21



## International standards for fetal growth based on serial ultrasound measurements: the Fetal Growth Longitudinal Study of the INTERGROWTH-21<sup>st</sup> Project

Lancet 2014

*Aris T Papageorgiou, Eric O Ohuma, Douglas G Altman, Tullia Todros, Leila Cheikh Ismail, Ann Lambert, Yasmin A Jaffer, Enrico Bertino,*



**Figure 2: Fitted 3rd, 50th, and 97th smoothed centile curves of fetal measurements**  
Fitted 3rd (bottom dashed line), 50th (middle dashed line), and 97th (top dashed line) smoothed centile curves for fetal head circumference (A), fetal biparietal diameter (B), fetal occipitofrontal diameter (C), fetal abdominal circumference (D), and fetal femur length (E) measured by ultrasound according to gestational age. Open red circles show empirical values for each week of gestation and open grey circles show actual observations.

### ***Ultrasound based growth standard: “Optimal fetal size”***

4321 low risk women from 8 countries included in final chart (Brazil, UK, Italy, Oman, USA, China, India & Kenya)

Linked with WHO infant and childhood growth charts up to age 2 yrs

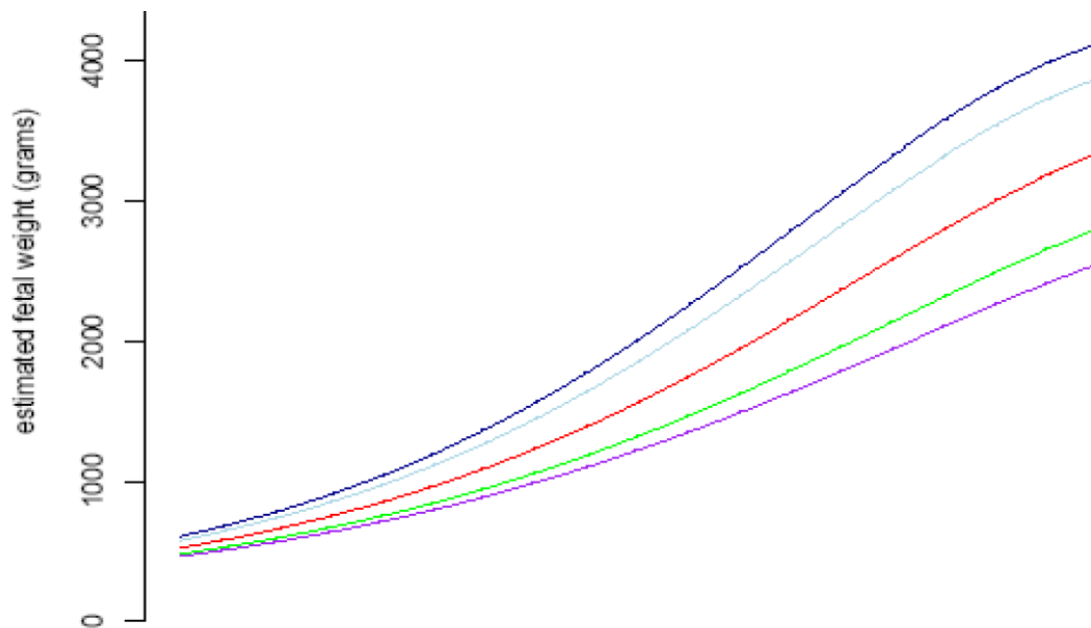
Dating scan < 14 weeks then scanned every 5 weeks to 42 weeks

# Intergrowth 21

## International Estimated Fetal Weight Standards of the INTERGROWTH-21<sup>st</sup> Project

J. Stirnemann<sup>a</sup>, J. Villar<sup>b\*</sup>, L.J.Salomon<sup>a</sup>, E. Ohuma<sup>b,c</sup>, P. Ruyan<sup>d</sup>, D.G. Altman<sup>c</sup>, F. Nosten<sup>e</sup>, R. Craik<sup>b</sup>, S. Munim<sup>f</sup>, L. Cheikh Ismail<sup>b</sup>, F.C. Barros<sup>g,h</sup>, A. Lambert<sup>b</sup>, S. Norris<sup>i</sup>, M. Carvalho<sup>j</sup>, Y.A. Jaffer<sup>k</sup>, J.A.Noble<sup>l</sup>, E.Bertino<sup>m</sup>, M.G Gravett<sup>n</sup>, M Purwar<sup>o</sup>, C. Victora<sup>h</sup>, R. Uauy<sup>p,q\*</sup>, Z. Bhutta<sup>r\*</sup>, S. Kennedy<sup>b\*</sup>, A.T. Papageorgiou<sup>b\*</sup>, for the International Fetal and Newborn Growth Consortium for the 21<sup>st</sup> Century (INTERGROWTH-21<sup>st</sup>).

UOG 2017



$$\log(\text{EFW}) = 5.1 - 54.1 \times (\text{AC}/100)^3 - 95.8 \times (\text{AC}/100)^3 \times \log(\text{AC}/100) + 3.1 \times (\text{HC}/100)$$

**ONLY HC and AC included !**

# Intergrowth 21

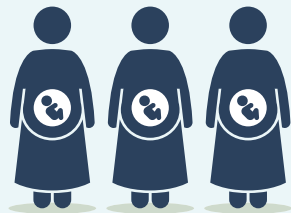
## INTERGROWTH



### LOCATION

#### 8 Countries

Brazil, China, India, Italy, Kenya, Oman, U.K. and U.S.

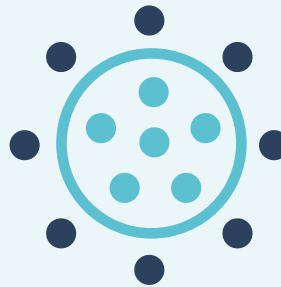


### RACE & ETHNIC

One overall growth chart

No statistical testing for differences among countries

- One size fits all
- Skeletal growth not affected
- AC, EFW, Bwt – differ due to environment
- Not fetal sex specific – inc in calculator



### INCLUSION/ EXCLUSION

Exclusion of pregnancy complications and fetal factors such as congenital anomalies and stillbirth

Supernormal

2 yrs – normal ND



### ANALYTIC APPROACHES

Data transformation: none

Model assumptions: linear mixed models with location and scale assumptions, assuming a normal distribution of the fetal growth trajectories

Smoothing technique over gestational age: second-degree fractional polynomials

- All measurements blinded, standardised equipment, trained sonographers

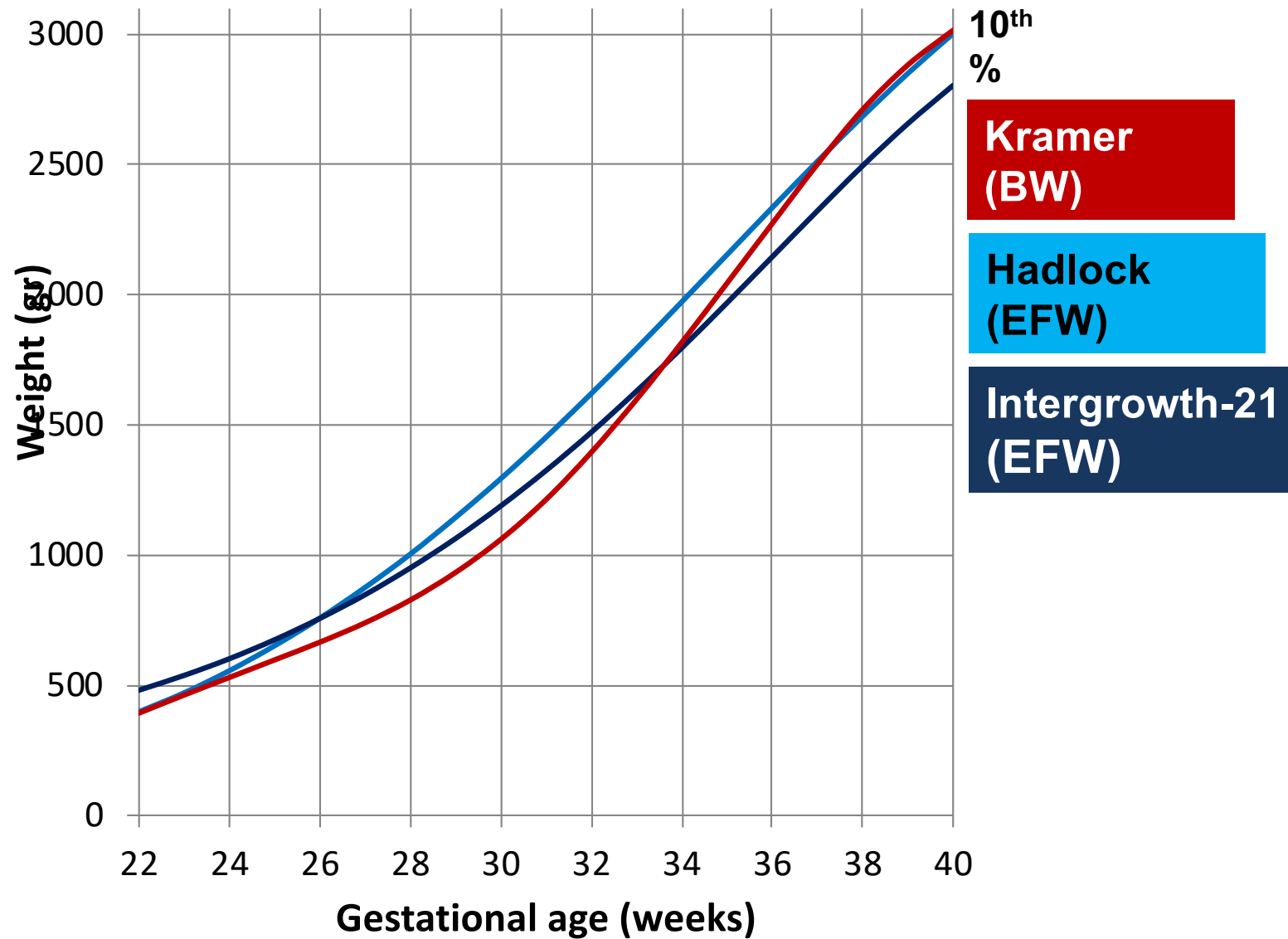


### ESTIMATED FETAL WEIGHT

Created a new formula<sup>12</sup> based on only HC and AC, making the comparison of EFW less meaningful

- Mean Bwt ~600g < in India
- In Canada high rates of LGA, low rates of SGA

# Growth charts



# WHO

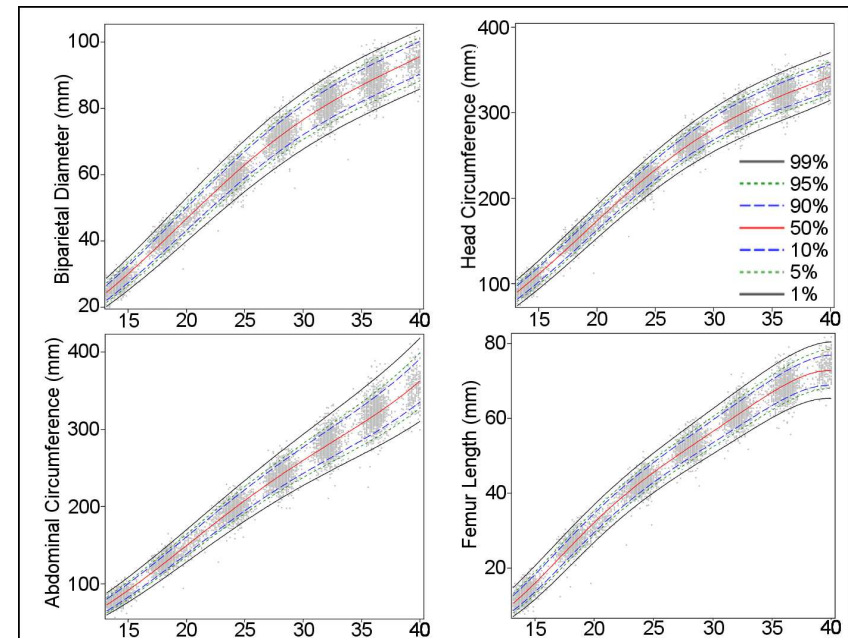
- ***Ultrasound based growth standard:***  
“Optimal fetal size”
- 1387 low risk women from 10 ultrasound centres included in final chart (Argentina, Brazil, DRC, Denmark, Egypt, France , Germany, India, Norway & Thailand)
- Dating scan < 14 weeks then scanned x 7
- EFW calculated using Hadlock eqn
- Presented pooled data – 1 chart

## RESEARCH ARTICLE

The World Health Organization Fetal Growth Charts: A Multinational Longitudinal Study of Ultrasound Biometric Measurements and Estimated Fetal Weight

Torvid Kiserud<sup>1,2\*</sup>, Gilda Piaggio<sup>3,4\*</sup>, Guillermo Carroli<sup>5</sup>, Mariana Widmer<sup>6\*</sup>, .

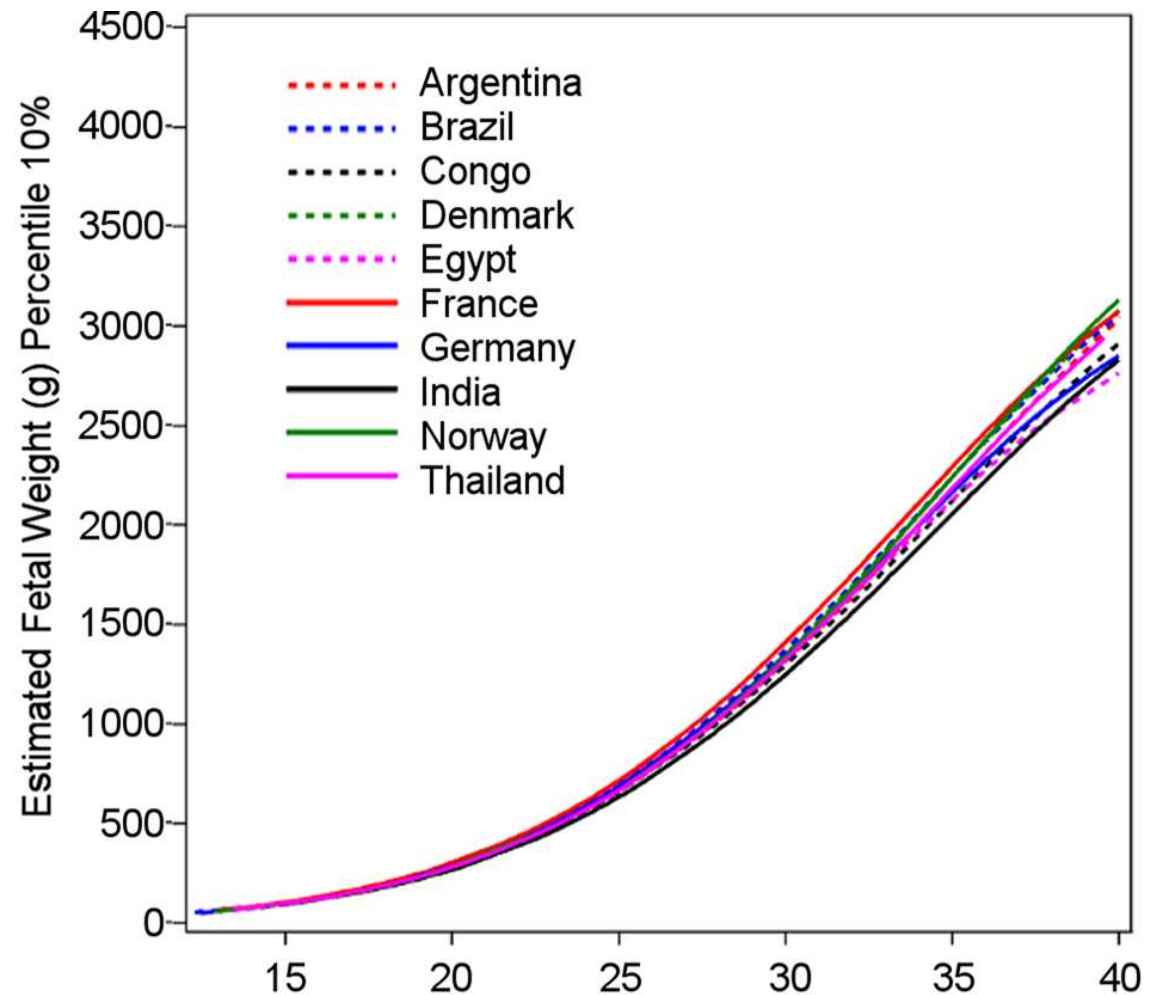
PLOS Med 2017





# WHO

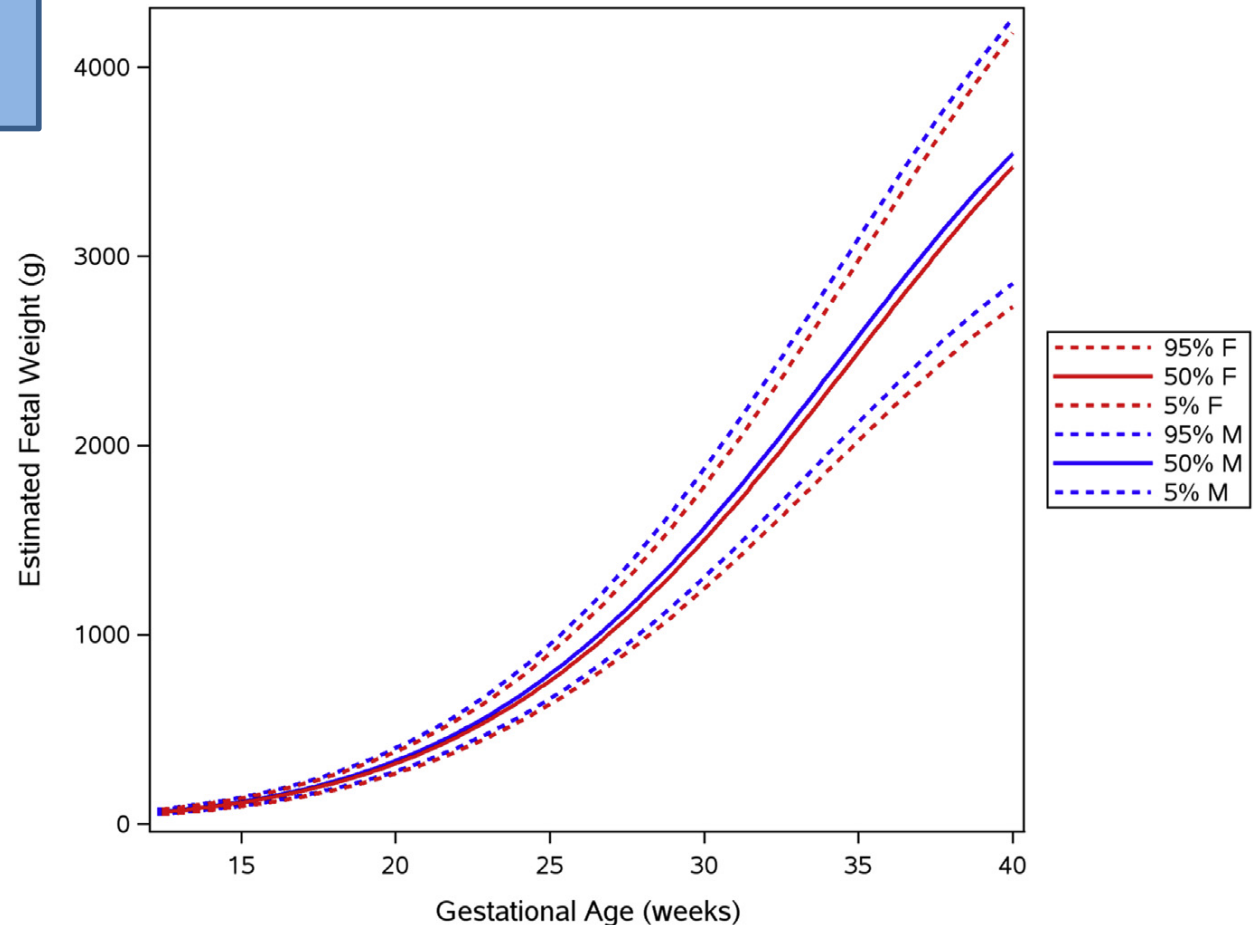
- **Country variation**
- Country not self reported race
- Sample size for each country ~150
- ~300 patients contribute to each geographical region (Asia, Africa)
- High rates of early term CS in S. America – limited term growth data
- Birthweight ~ 500g less in India compared to Norway



# WHO

## Sex specific

- Males 4.5% bigger



# WHO

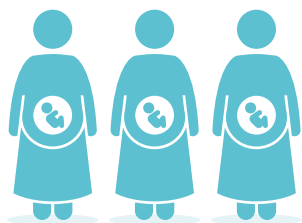
## WHO FETAL



### LOCATION

#### 10 Countries

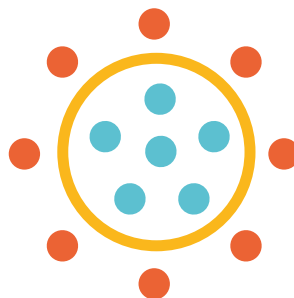
Argentina, Brazil, Democratic Republic of the Congo, Denmark, Egypt, France, Germany, India, Norway, and Thailand



### RACE & ETHNIC

One overall growth chart  
Fetal growth showed natural variation, differing significantly between countries which largely followed ethnic distribution

1 postnatal chart WHO MGRS

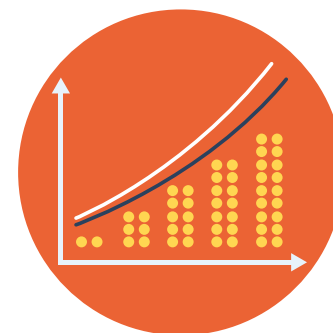


### INCLUSION/ EXCLUSION

Only optimal health inclusions

No complication excluded (no impact on percentiles)

Reflects clinical practice better



### ANALYTIC APPROACHES

Data transformation: log

Model assumptions:  
Quantile regression without distributional assumptions

Smoothing technique over gestational age: polynomial functions

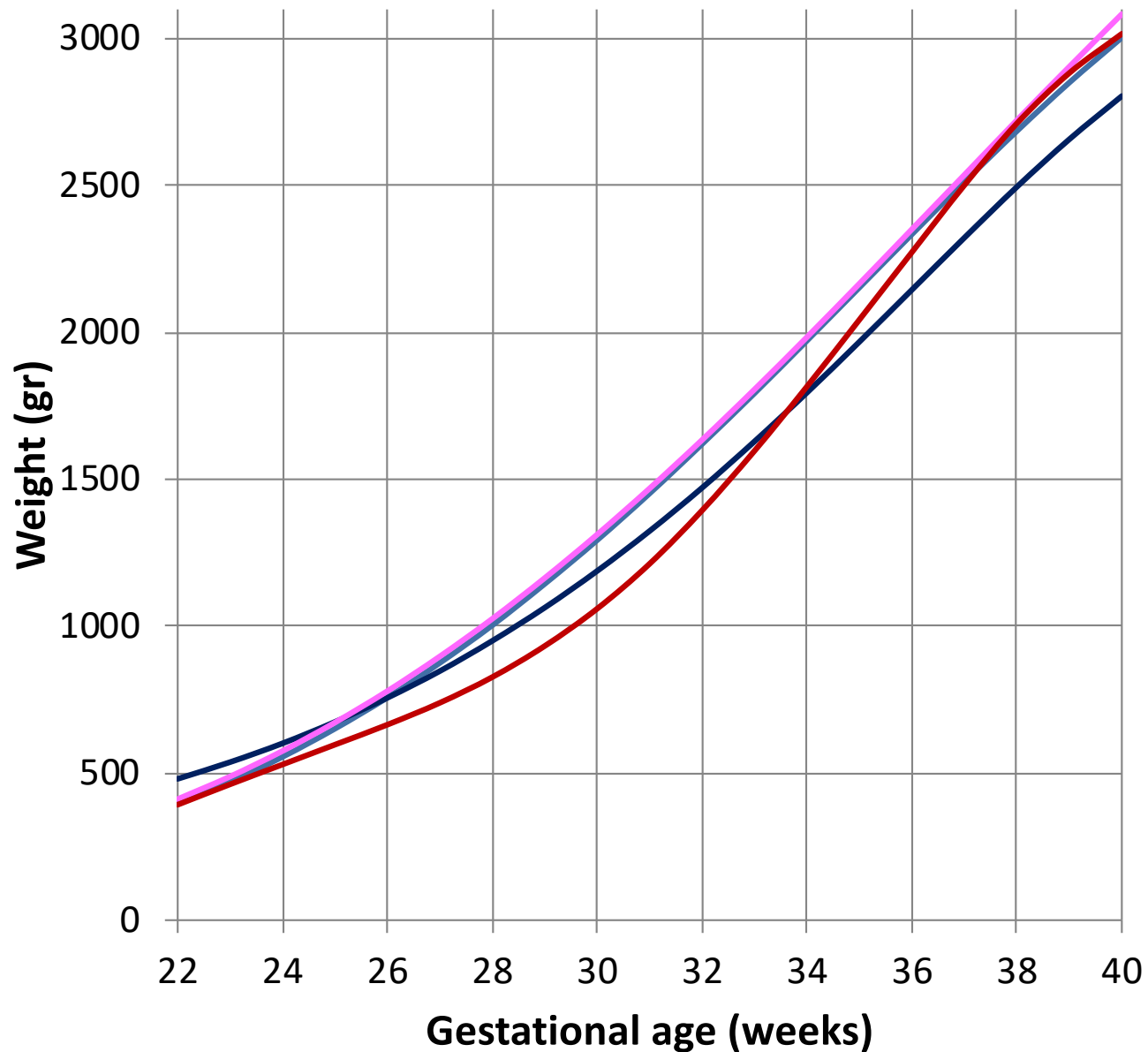
Measurements not blinded – clinically revealed



### ESTIMATED FETAL WEIGHT

Calculated EFW from HC, AC and FL using the Hadlock 1985 formula<sup>26</sup>

# Growth charts



10<sup>th</sup>%

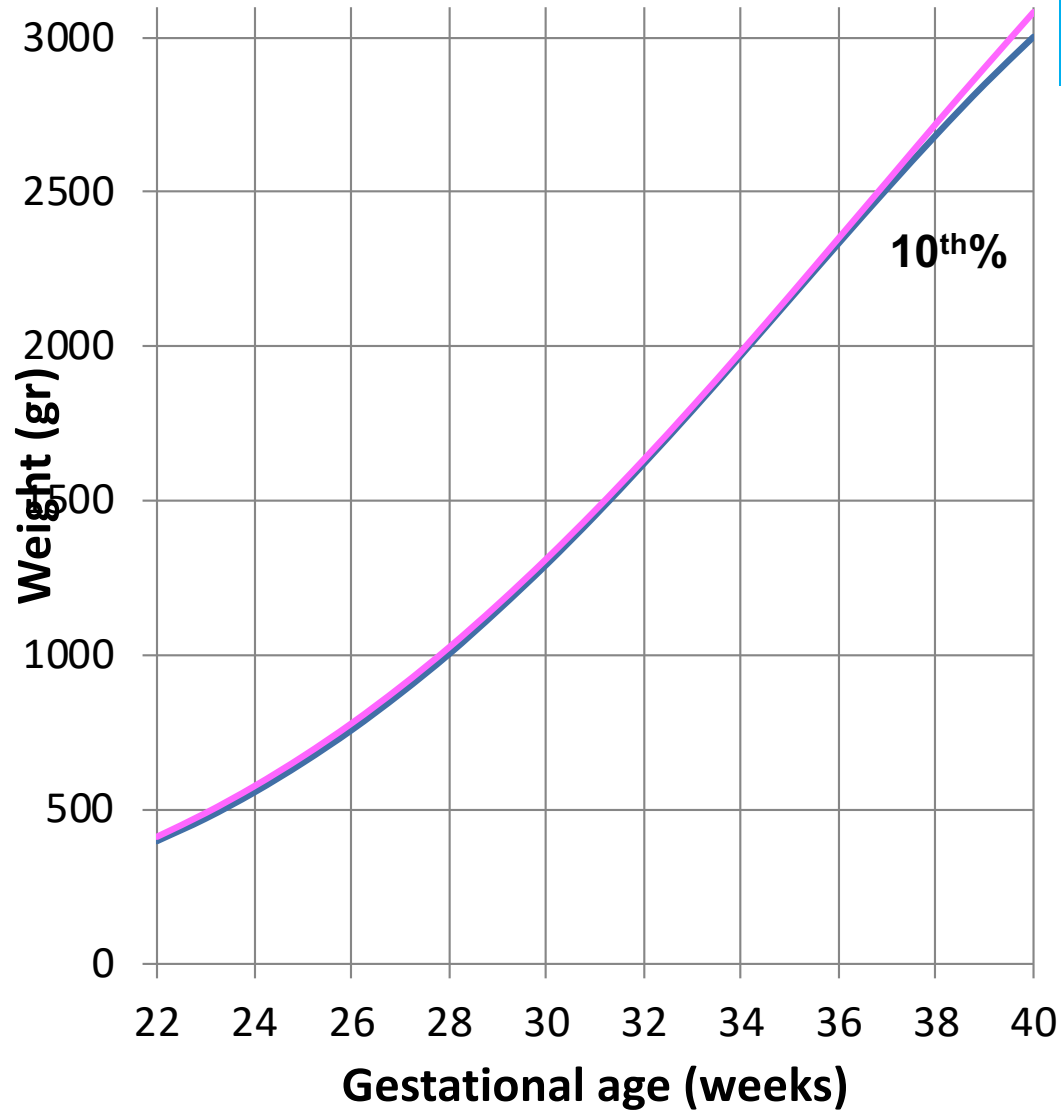
**Kramer (BW)**

**Hadlock (EFW)**

**Intergrowth-21 (EFW)**

**WHO (EFW)**

# Growth charts



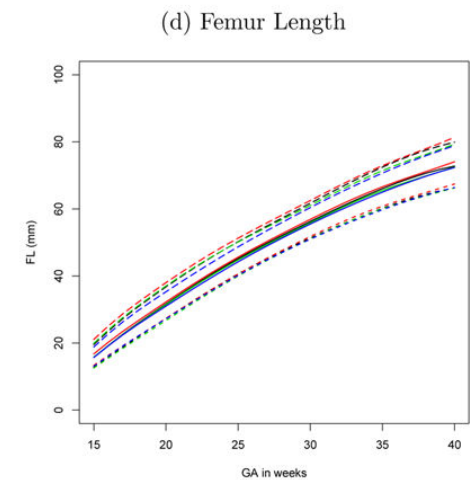
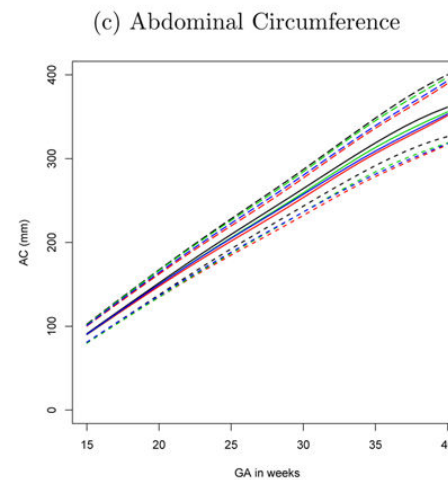
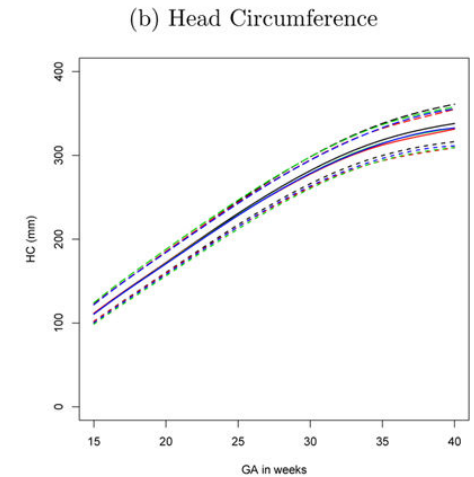
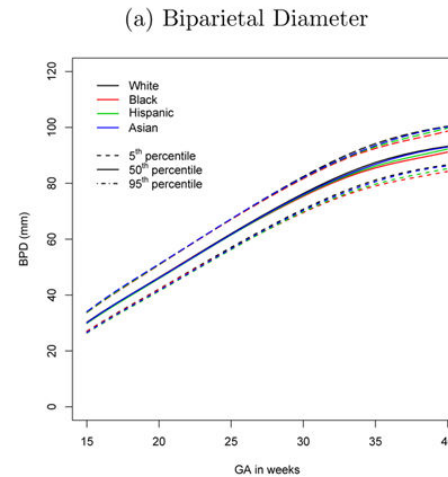
**Hadlock (EFW)**

**WHO (EFW)**

**10<sup>th</sup>%**

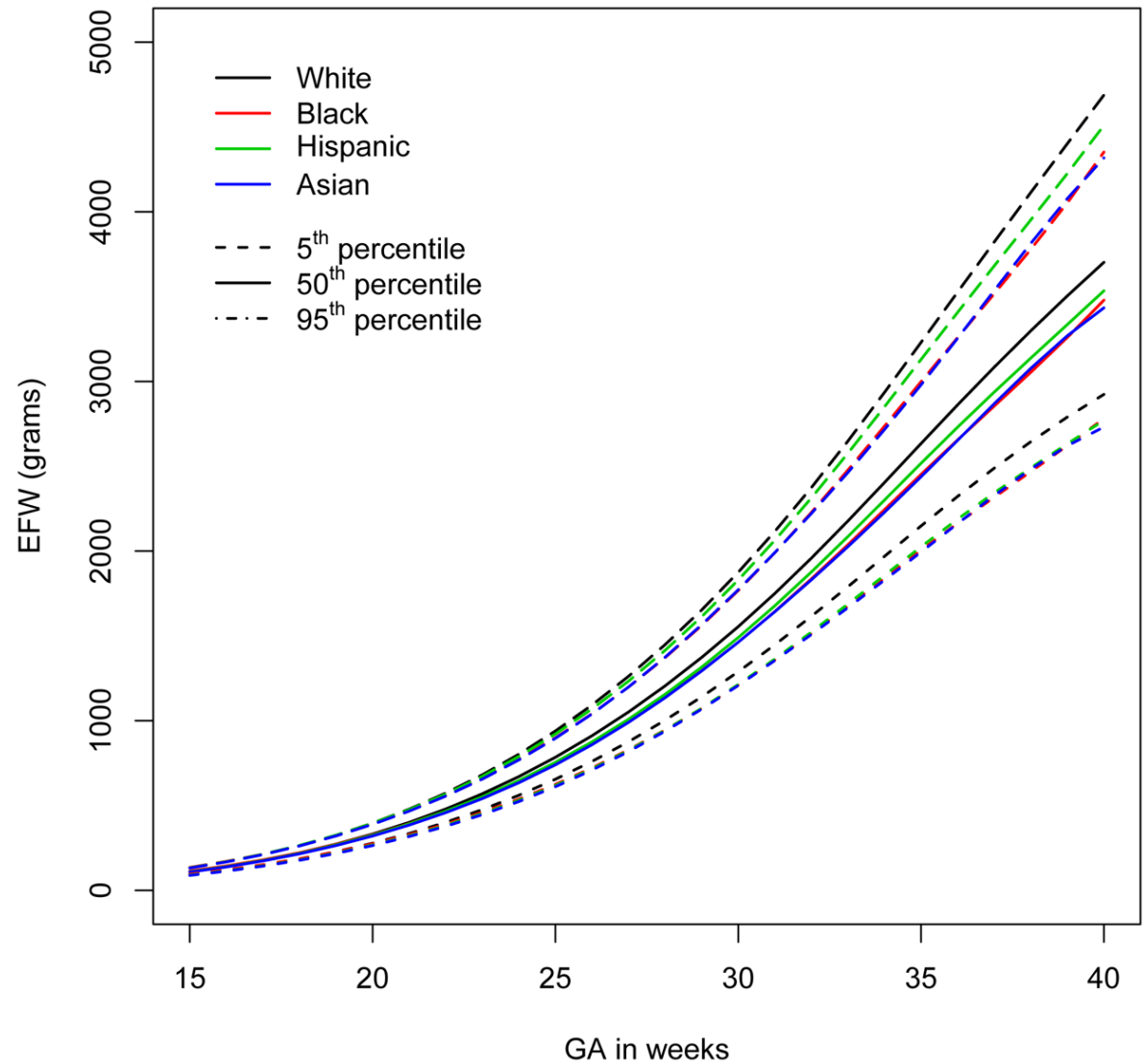
# NICHD

- ***Ultrasound based growth standard: “Optimal fetal growth”***
- Hypothesis: are racial/ethnic growth standards better
- 12 USA sites, 1733 women
- Dating scan 10-14 weeks then scanned x 5
- EFW calculated using Hadlock eqn



NICHD

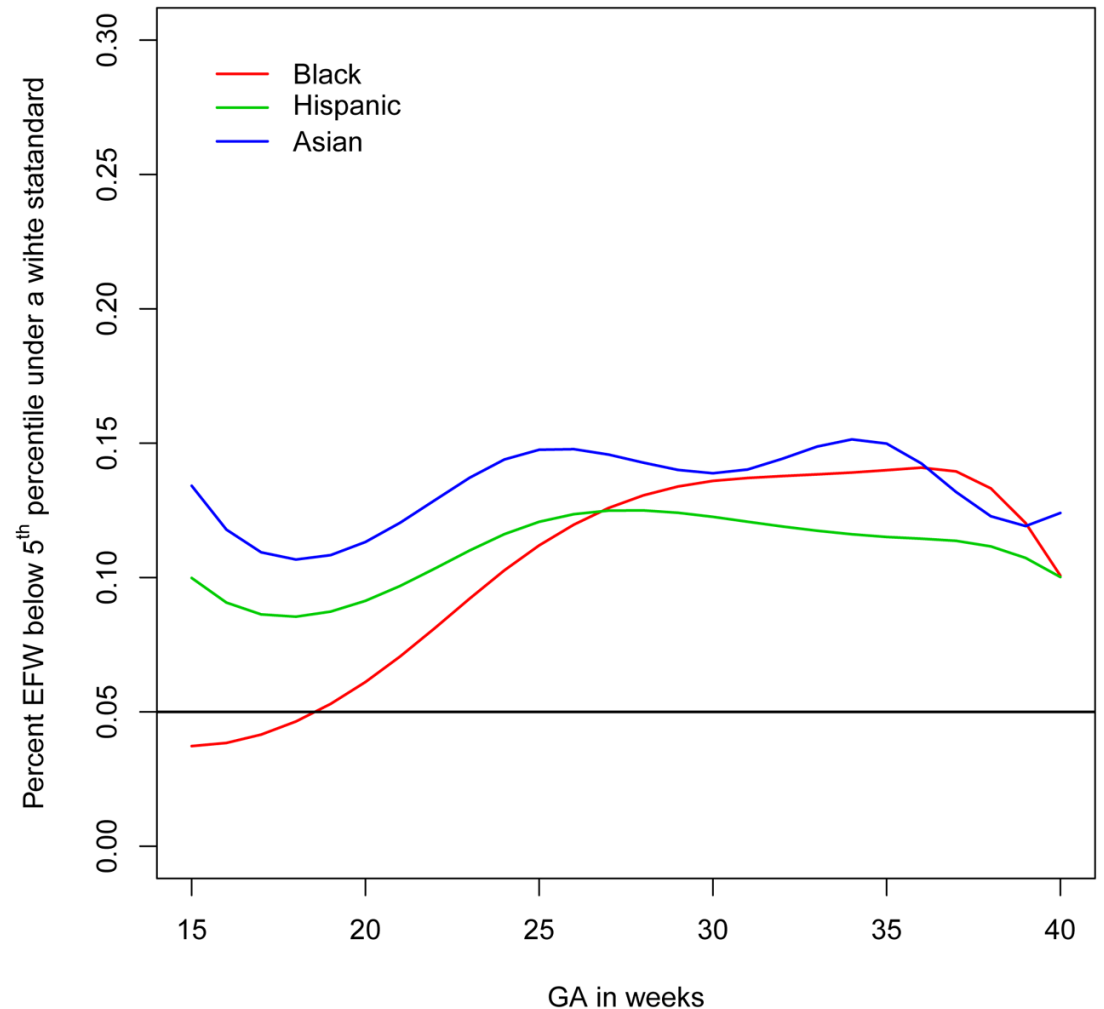
- *Race based differences in EFW*
- *Differences in maternal height, weight, leg length, body composition (fat) based on race*



# NICHD

***Risk of misclassification  
on FGR dependent on  
race – up to 15%***

But higher rate of  
adverse outcome  
(stillbirth, PTB) in Black  
and Hispanic groups





# NICHD

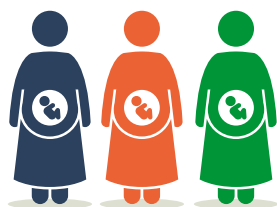
## NICHD



### LOCATION

#### 12 U.S. Sites

New York [2], New Jersey, Delaware, Rhode Island, Massachusetts, South Carolina, Alabama, Illinois, and California [3]



### RACE & ETHNIC

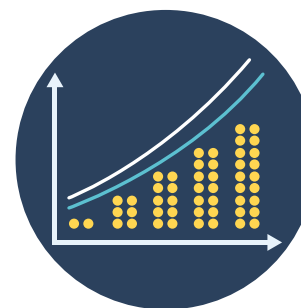
Highly statistically significant differences in fetal growth by race/ethnicity

Racial/ethnic-specific derived standards



### INCLUSION/ EXCLUSION

*A priori* exclusion of pregnancy complications, preterm delivery < 37 weeks' gestation, stillbirth and fetal factors including all structural anomalies and karyotype abnormalities



### ANALYTIC APPROACHES

Data transformation: log

Model assumptions: linear mixed models, assuming a normal distribution of the fetal growth trajectories (after log transformation)

Smoothing technique over gestational age: cubic splines

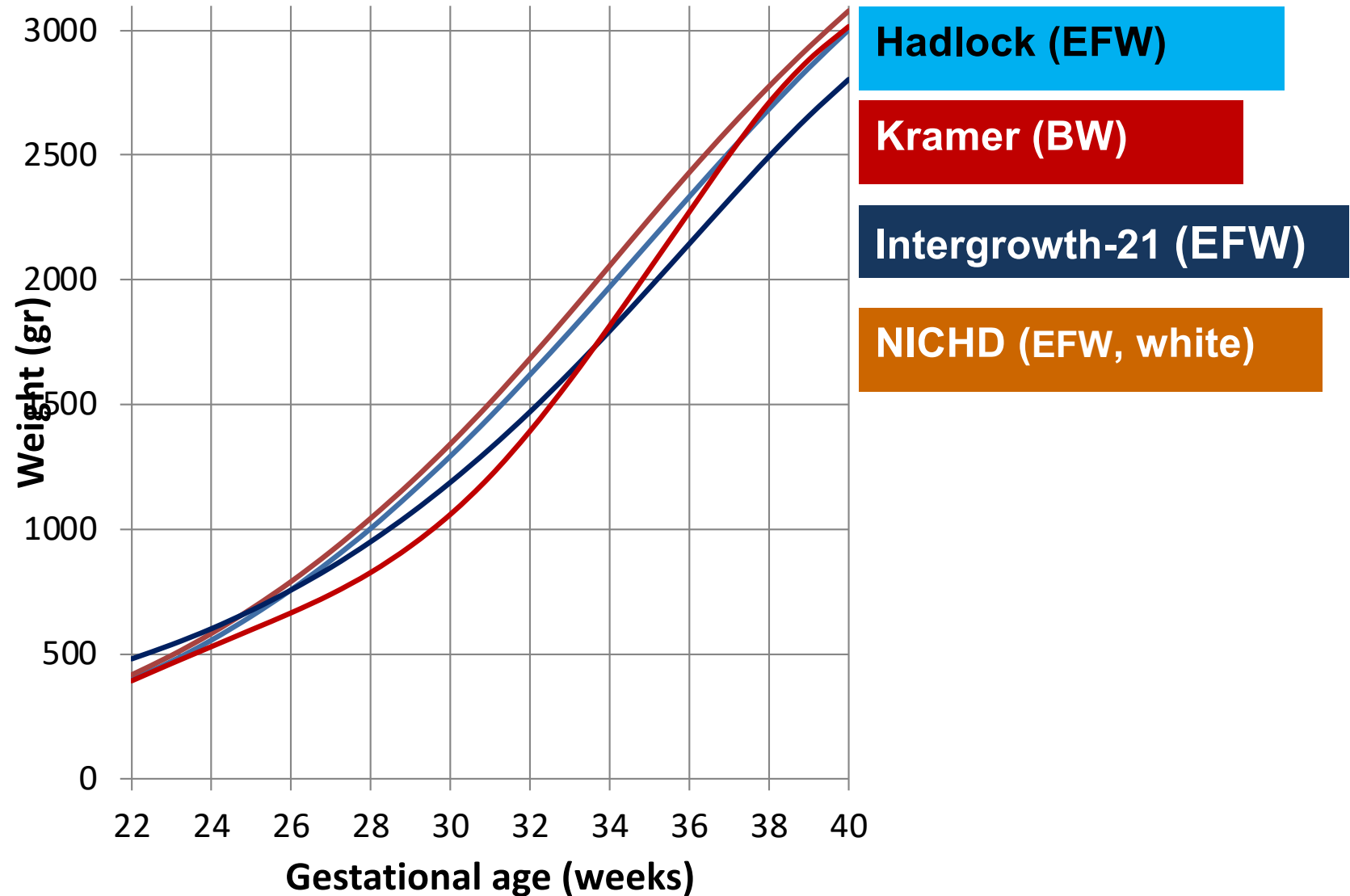


### ESTIMATED FETAL WEIGHT

Calculated EFW from HC, AC and FL using the Hadlock 1985 formula<sup>26</sup>

Different chart for:  
White  
Non Hispanic black  
Hispanic  
Asian/Pacific Islander

# Growth charts



# Customised GROW charts



## Gestation Network

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Gestational Age](#)

[Estimating  
Fetal Weight](#)

[Calculating  
Scan Errors](#)

[Reference  
Literature](#)

### Growth Charts

- **Country specific – not Canada**
- International charts available
- Fetal weight charts
- Term optimal weight and multiple regression analyses

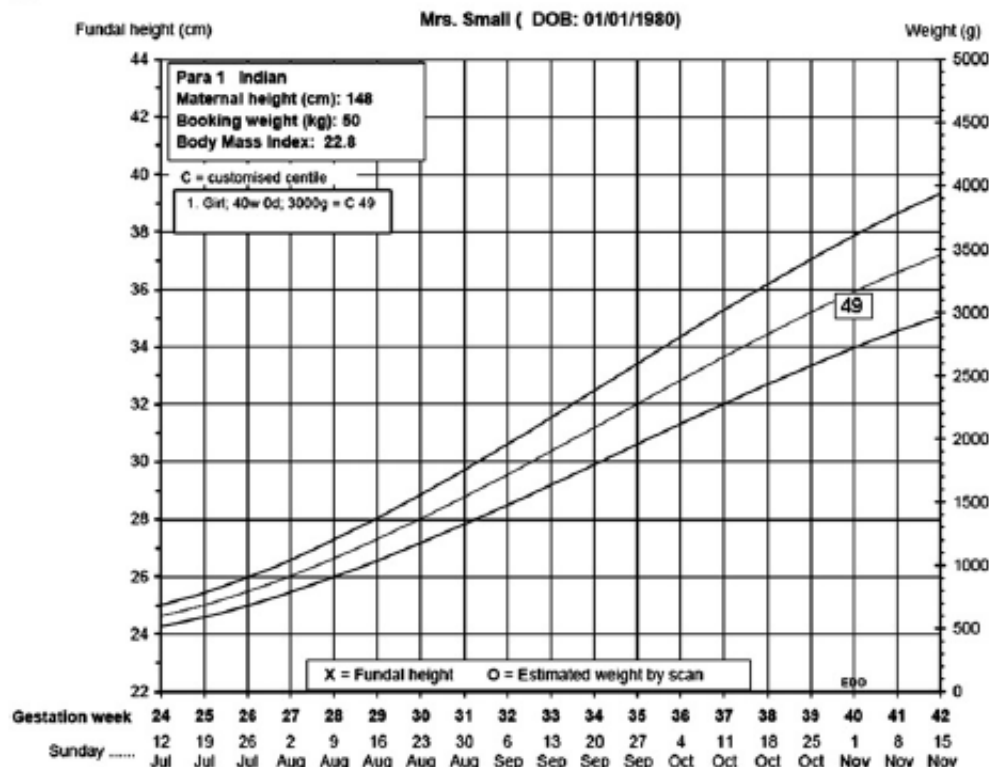
### Customise for:

- Maternal height
- Maternal weight
- Ethnicity
- Parity
- Fetal sex
- Country of residence

# Customised GROW charts

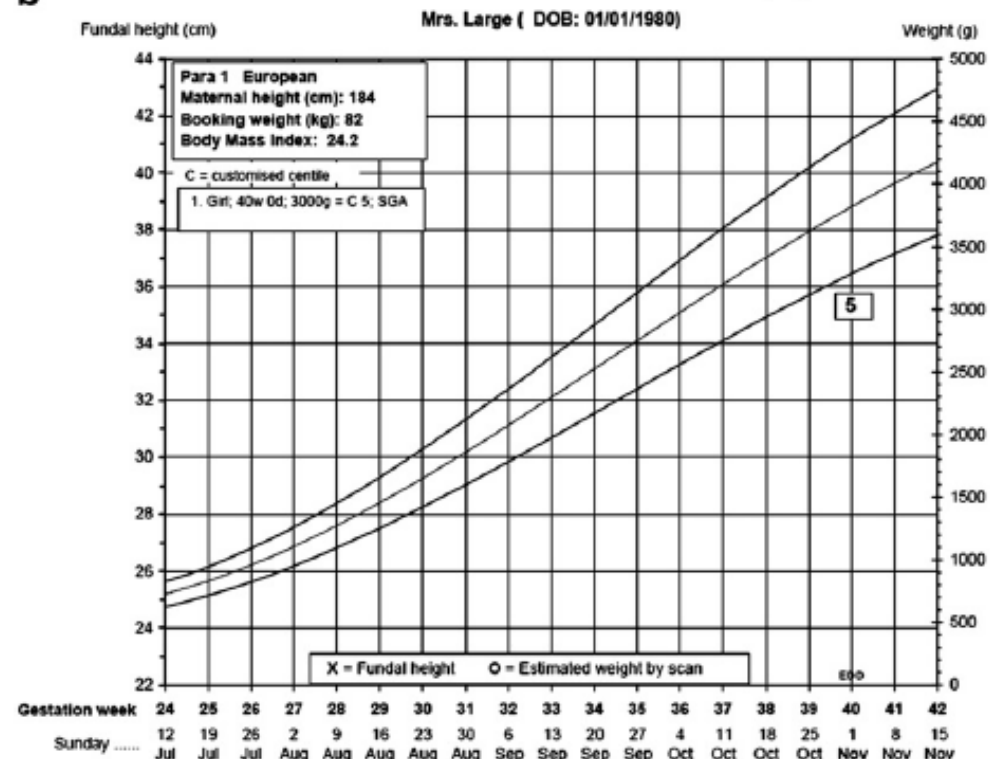
a

CUSTOMISED ANTENATAL GROWTH CHART v 7.5.1U (UK)



b

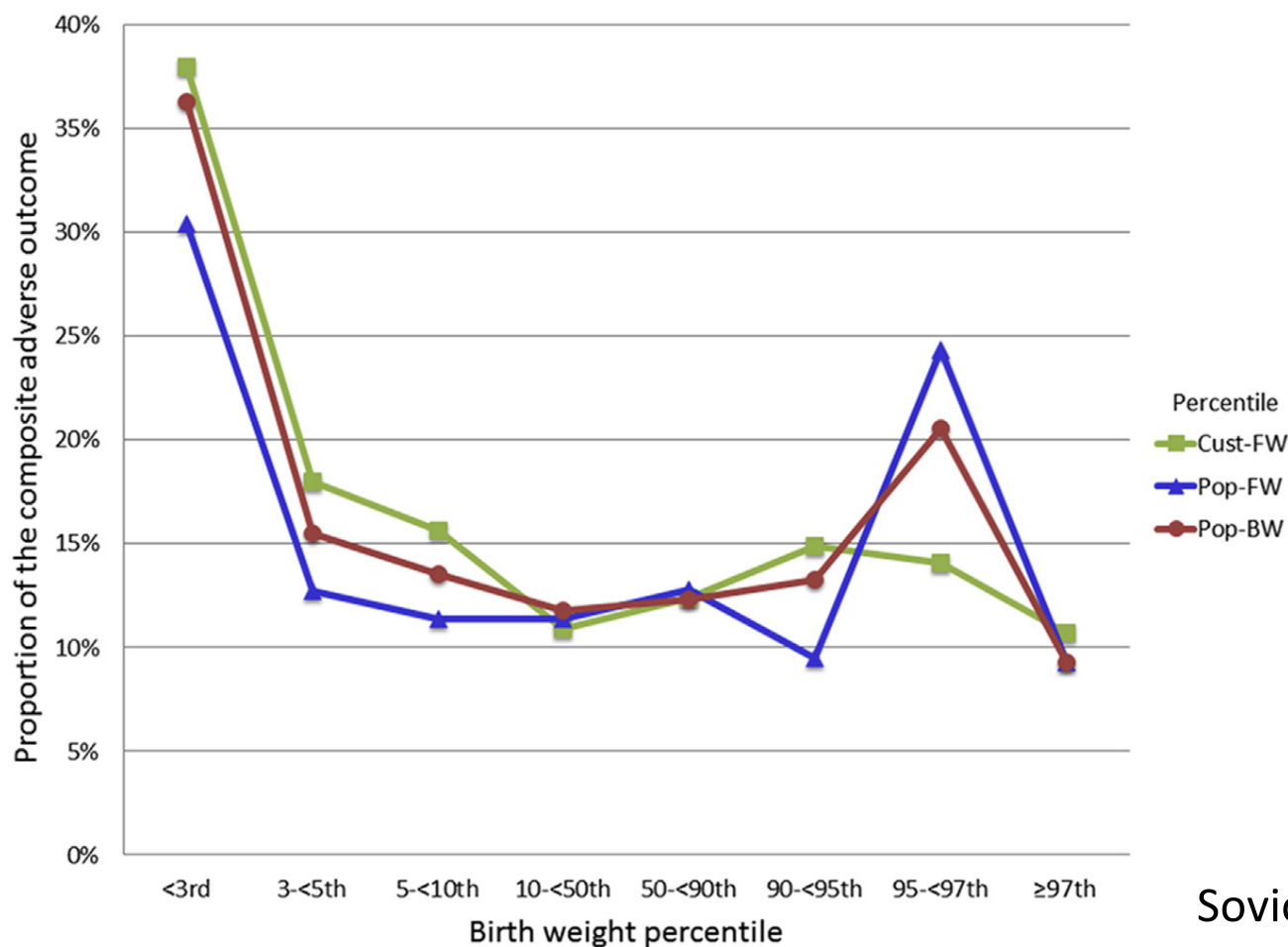
CUSTOMISED ANTENATAL GROWTH CHART v 7.5.1U (UK)



3 kg baby at 40 weeks

- **BMI**
- Better at identifying large mothers with FGR babies more at risk of FDIU
- Reduces interventions for low risk small mothers

# Customisation and adverse outcomes



Sovio, AJOG 2018

Proportion (%) of the composite adverse outcome in relation to birthweight percentile category.

Differences accounted for by use of fetal weight standard, prematurity and increased adverse outcomes in women with high BMI

# Self reporting ethnicity?



235 women – 16 categories  
50% used multiple descriptors  
13% couldn't identify their ethnicity  
Partners interpretation varied

Lockie, BJOG 2018