

# SHORT FEMUR LENGTH FOR EFW WORKING GROUP- NOV 2019



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# Background

- FL is only routine long bone measurement
  - ▣ *Many scanning guidelines recommend the systematic documentation of the presence and symmetry of all extremities*
  
- Considerations short FL in mid-second trimester
  - ▣ Normal variant greater majority ( ~ two-thirds)
  - ▣ Significant FPR up to 13% noted on re-measurement
  - ▣ Fetal growth restriction
  - ▣ Aneuploidy, Skeletal Dysplasia
  - ▣ Other

## Hypothesis : Why Short FL in mid trimester?

- Etiologies are likely multifactorial
- Some Hypothesis include:
  - ▣ Early adaptive response to chronic hypoxia and placental dysfunction
  - ▣ Disorder of vascular origin such as pregnancy induced hypertension or pre-eclampsia
  - ▣ Alteration in secretion of growth factors
  - ▣ FGF (fibroblast growth factor) receptor may be altered

# Define short FL

- Short FL
  - Definition is below the 5<sup>th</sup> percentile which roughly corresponds to below the -2SD
  - Prospective Danish population study, the 5<sup>th</sup> percentile corresponded to the mean minus 1.645 SD, led authors to suggest that cutoff value closer to -2 SD instead of the 5<sup>th</sup> percentile should be considered to minimize overdiagnosis
    - Isolated short FL at the 2<sup>nd</sup> mid-trimester anatomic scan associated with a higher risk of chromosomal anomalies, in particular trisomy 21, and a higher risk for delivery of a small for gestational age infant and early preterm delivery
- Markedly shortened FL
  - In Kurtz et al a markedly short FL ( $\geq 5$  mm below the -2 SD line [equivalent to  $> -4.3$  SD]) was associated with a high likelihood of a skeletal dysplasia, whereas a mildly shortened FL (within 4 mm of the -2 SD line [between -2 and -4 SD below the mean]) in combination with normal interval growth was unlikely to be associated with skeletal dysplasia

Association of isolated short femur in the mid-trimester fetus with perinatal outcome. Weisz B, David AL, Chitty L, Peebles D, Pandya P, Patel P, Rodeck CH *Ultrasound Obstet Gynecol.* 2008;31(5):512. Usefulness of a short femur in the utero detection of skeletal dysplasias. Kurtz et al. *Radiology* 1990;177(1) 197.

# Short FL – Non-Isolated

- Literature is limited to retrospective studies
- Isolated short FL is associated with FGR
  - 40% Papageorgiou, 2008
    - Details: <5<sup>th</sup> percentile, Severe IUGR defined as AC < 5<sup>th</sup> percentile with abnormal UA Doppler requiring delivery before 37wk
    - High risk severe IUGR requiring PTB accompanied by PET in 1/3
    - Short FL and association of aneuploidy depend on prevalence, thus this population which had all NT found no isolated cases with T21
    - Non-isolated short FL were 1/3 of this group
  - 39% Todros 2004
    - Details: < 10<sup>th</sup>ile, retrospective 86 cases with overall 32.5% normal, 46.5% structural abnormality, 21% SGA
  - 43% Vermeer 2013
    - Details: < 5<sup>th</sup> percentile,, 112 cases with 78% isolated short FL
    - Non-isolated 6 aneuploidy, 12 MFA, 1 genetic
    - Isolated 43% IUGR with LR 1.2

# Isolated Short Femur

D'ambrosio, Valentina, et al. "Midtrimester isolated short femur and perinatal outcomes: A systematic review and meta-analysis." *Acta obstetrica et gynecologica Scandinavica* 98.1 (2019): 11-17.

- Singleton pregnancies 18-28 weeks with **isolated short FL** <5<sup>th</sup> percentile
- 6 retrospective studies, total 3078 cases, control of 222,303 (normal FL)
  - **14.2% IUGR or SGA prevalence vs 5.2% controls, odds ratio ~ 4x**
    - 438/3078
  - Higher incidence low BW 22.10% vs control group: 8.57%, odds ratio 3.24
  - Higher incidence of low APGAR, fetal demise, NICU admission, neonatal death
- **Conclude:**
  - Significant association short FL, IUGR, SGA & adverse perinatal outcome
  - Conservative counsel as 61% normal outcome
    - Consider closer monitor maternal BP, increase fetal surveillance
    - If placental dysfunction Doppler may aid distinguish “inherent short FL” vs at risk group
    - Need larger prospective trials
      - Limitations: sample size study group 1.4%; although same definition short FL used different biometric charts
      - Could not exclude entirely other reasons for referral

# Next Steps:

- Review maternal/parental history, serum tests & markers FGR
- Re-measure to confirm short FL ( 13% FPR)
- Measure all long bones
- Define pattern over time ( 3-4 weeks)
  - Normal interim growth albeit along line below normal percentiles but along same growth curve may be constitutional
  - If FL over this interval falls further from the mean, consider severe FGR or skeletal dysplasia
  - Mildly shortened femur is between -2D and -4SD below mean for GA
  - Markedly shortened femur is  $> 4$  SD below mean for GA higher association with skeletal dysplasia

Consider referral to center with expertise

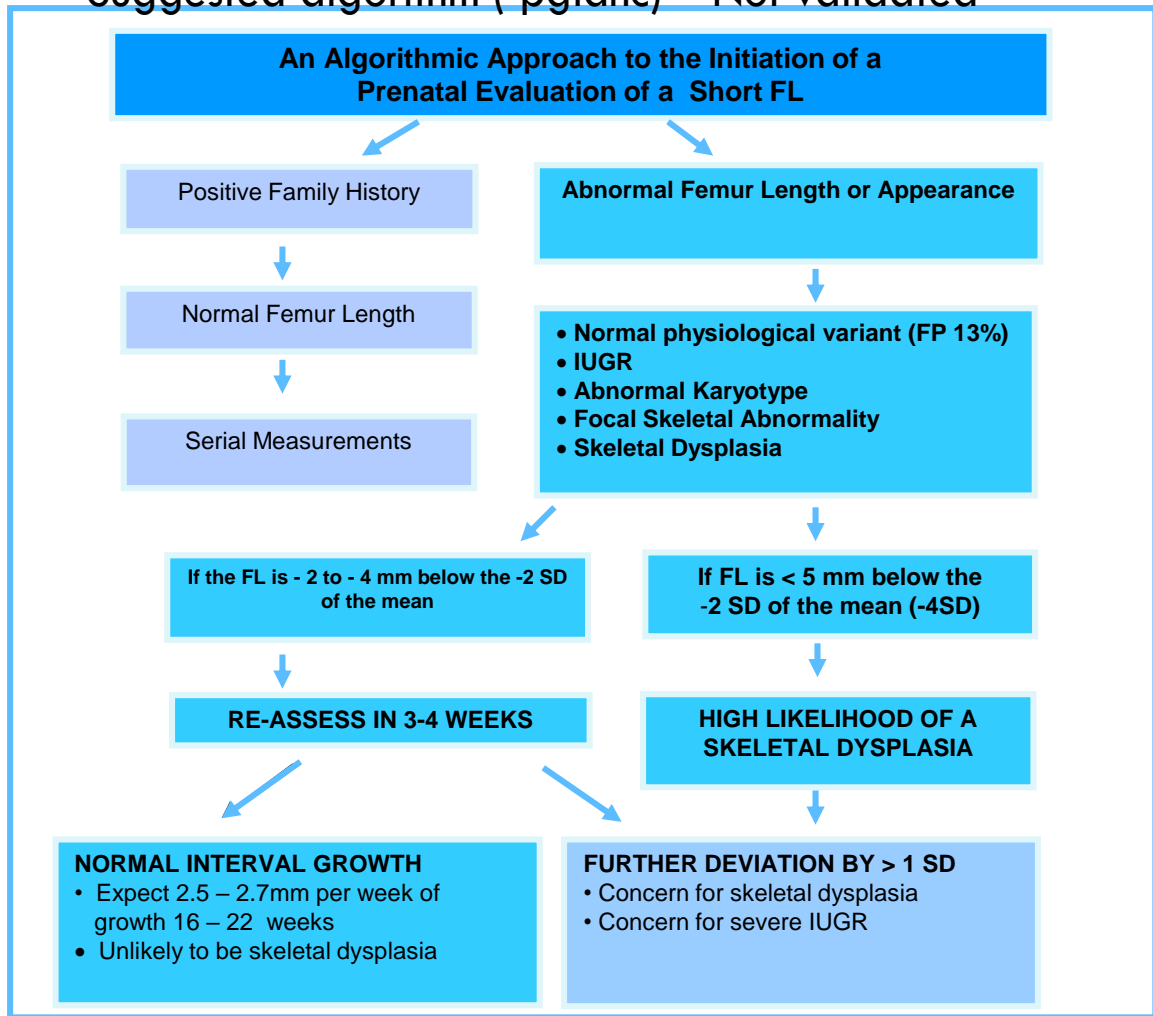
<https://www.uptodate.com/contents/approach-to-prenatal-diagnosis-of-the-lethal-skeletal-dysplasias?search=lethal%20skeletal%20dysplasia&source=>

## Conclusions: Isolated short FL at time mid-second trimester study

- Associated with an increased risk SGA, IUGR, adverse perinatal outcome in the range of 3-4x
- Consider increased surveillance for maternal hypertension and fetus well being
- Need larger prospective trials to refine our algorithms and management protocols



# Suggested algorithm ( pglanc) – Not validated



## General Reference

[https://www.uptodate.com/contents/approach-to-prenatal-diagnosis-of-the-lethal-skeletal-dysplasias?search=lethal%20musculoskeletal&source=search\\_result&selectedTitle=1~150&usage\\_type=default&display\\_rank=1#H478122598](https://www.uptodate.com/contents/approach-to-prenatal-diagnosis-of-the-lethal-skeletal-dysplasias?search=lethal%20musculoskeletal&source=search_result&selectedTitle=1~150&usage_type=default&display_rank=1#H478122598)

# THANK YOU.

Please address questions and/or referrals pertaining to potential fetal skeletal dysplasia to [phyllis.glanc@sunnybrook.ca](mailto:phyllis.glanc@sunnybrook.ca)