

WEBAPPENDIX: WHO AMANHI Gestational Age Study

Machine Learning Models to Assess Newborn Gestational Age in Low-Middle Income Countries: Findings from a multi-country, prospective cohort study

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Web Table 1: Detailed description of the Study Sites and Procedures

| | Bangladesh | Ghana | Pakistan (Karachi) | Tanzania (Pemba) | Zambia |
|---|--|---|---|---|--|
| Site Location/ Description | Rural areas of Sylhet district | Rural areas in central Ghana | 1 peri-urban community of Bin Qasim town and 1 peri-urban community of Korangi town, Karachi | Pemba Island in the Zanzibar archipelago | 6 rural districts in Southern Province |
| Pregnancy surveillance | 2-monthly by trained community health workers (CHWs) | Monthly fieldworkers (FW) | 3-monthly by FWs | 6-weekly by CHWs | Facility ANC enrollment |
| Population area¹ | 88,000 | 147,000 | 97000 | 72,000 | 25,000 |
| Ultrasonography | Fujifilm Sonosite M-Turbo® ultrasound system (Bothel, WA, US). Scans performed by trained MD sonographers. | Portable SonoSite™ S180 machines (SonoSite™, Inc, Bothel, WA, US). Scan performed by hospital midwives. | Mindray 7 Ultrasound System. Scan done by trained sonologist. | Sonoscape s6. Scans performed by trained sonographer in district hospitals. | Philips Sono Diagnost 260; Scans performed by trained sonographers. |
| % Facility Delivery¹ | 43.7 | 79.9 | 64.3 | 99 (?) | 97.1 |
| % with newborn visit completed <72 hours | 71.1% | 99.2% | 80.0% | 92.7% | 82.6% |
| Infant Weighing Scale | TANITA BD-585 Pediatric Scale Digital weighing scale. Precision 10 gm. | Salter suspension scale (non-digital) with sling. Precision 100 gm. | LAICA Electronic/digital infant scale. Precision 5 gm. | Seca 374 (capacity 44lb Digital infant weighing scale Precision 10 gm | GPC Gps092 Medical hanging scale (Delhi, India). Precision 100 grams. |
| Health worker type performing newborn assessment | Locally recruited women (non-clinical) with at least 10 grade education received 6 weeks training on maternal and newborn health; additional 7 days training on newborn assessment | Locally recruited non-clinical field supervisors with secondary school education. Trained for 7 days on newborn assessment. | Locally recruited women with at least 12 grade education received 6 month training on maternal and newborn care; additional 7 days training on newborn assessment | Trained health worker (health assistant) with 2 years formal health training. Initial 7 days training for newborn assessment. | Locally recruited women (non-clinical) with 12 years of schooling; Trained for 3 days on newborn assessment. |
| % SGA¹ | 41.8 | 33.4 | 35.5 | 9.6 | 17.8 |
| % LBW¹ | 25.6 | 11.8 | 22.5 | 4.6 | 6.5 |
| NMR¹ | 37.8 (35.5-40.2) | 29.1 (26.9-31.2) | 50.1 (46.9-53.4) | 16.0 (14.3-17.8) | 14.5 (13.0-16.0) |

¹Population-based rates, timing, and causes of maternal deaths, stillbirths, and neonatal deaths in south Asia and sub-Saharan Africa: a multi-country prospective cohort study. Lancet Glob Health 2018; 6(12): e1297-e308.

Web Table 2. Diagnostic accuracy of Ballard, LMP, and AMANHI models for identification of <37 week and <34 week infants

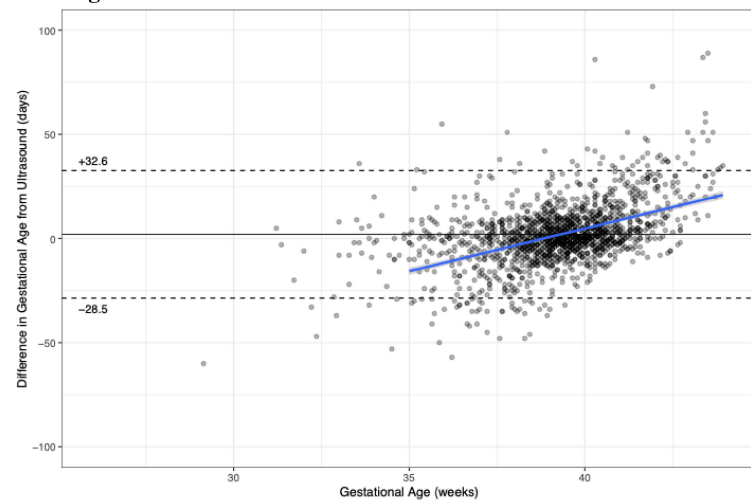
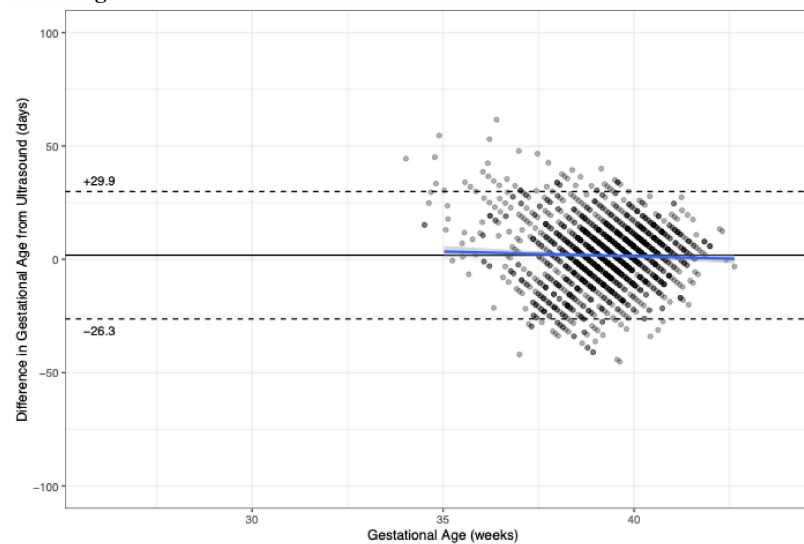
Gold standard dated by early pregnancy ultrasound, additional clinical thresholds maximizing sensitivity for identification of preterm births.

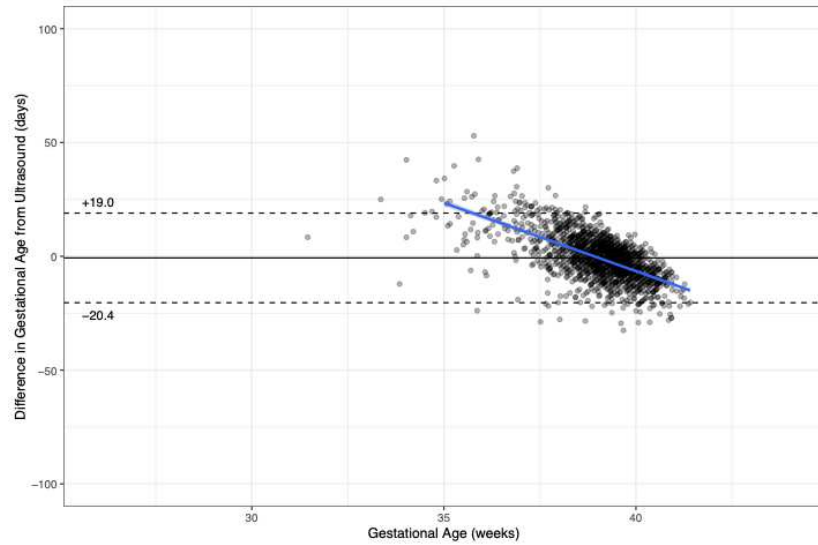
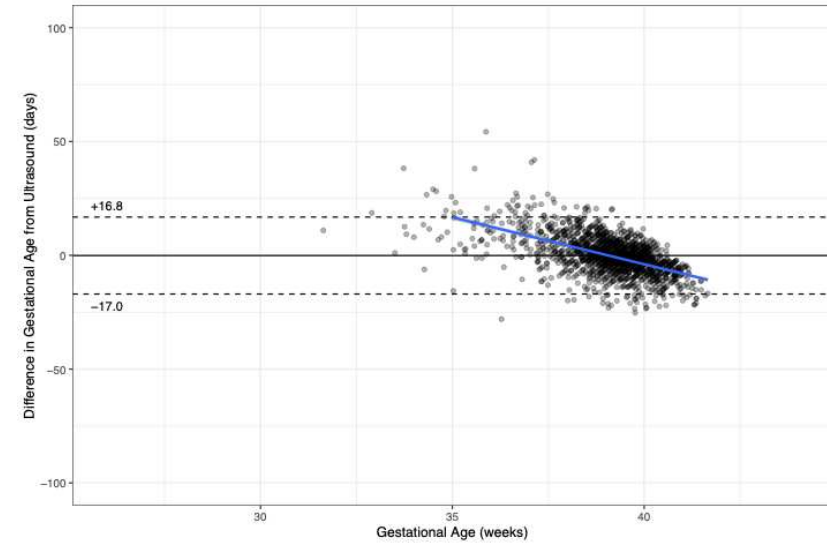
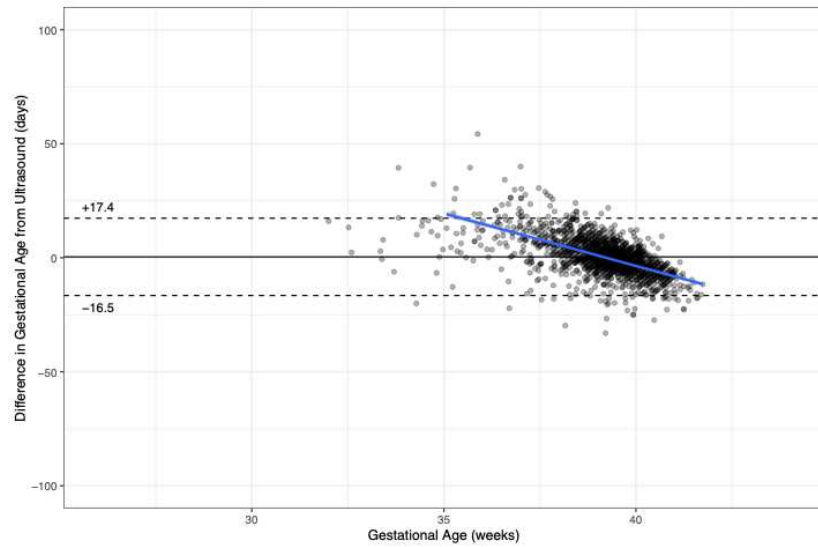
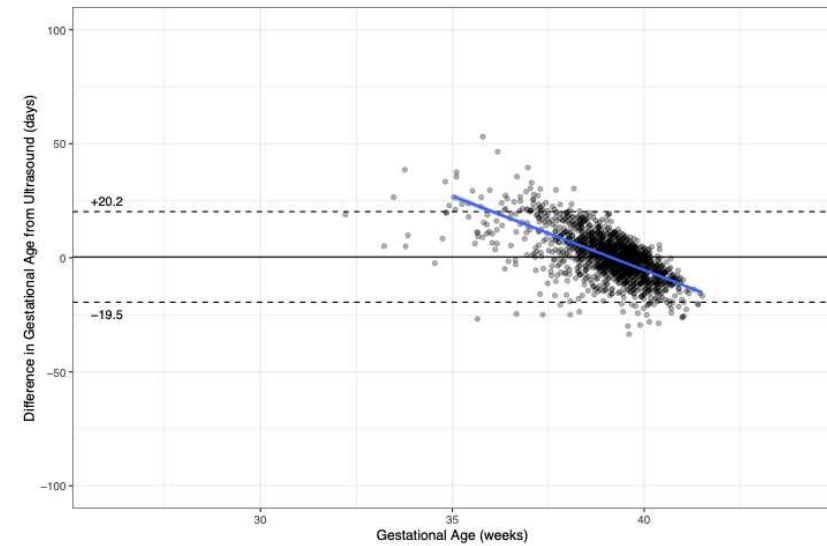
| Model | AUC | Threshold selection | Se | Sp | PPV | NPV | LRP | LRN |
|----------------------------------|------|---------------------|------|------|------|------|-------|------|
| Classify <37 weeks | | | | | | | | |
| LMP | 0.81 | Youden Index | 0.76 | 0.76 | 0.20 | 0.96 | 2.38 | 0.51 |
| LMP | 0.81 | 95% sensitivity | 0.95 | 0.23 | 0.09 | 0.98 | 1.23 | 0.22 |
| LMP | 0.81 | 90% sensitivity | 0.90 | 0.36 | 0.11 | 0.98 | 1.66 | 0.22 |
| LMP | 0.81 | 85% sensitivity | 0.85 | 0.61 | 0.14 | 0.98 | 2.17 | 0.25 |
| Ballard exam | 0.74 | Youden Index | 0.63 | 0.71 | 0.15 | 0.96 | 2.20 | 0.52 |
| Ballard | 0.74 | 95% sensitivity | 0.95 | 0.25 | 0.09 | 0.98 | 1.26 | 0.20 |
| Ballard | 0.74 | 90% sensitivity | 0.90 | 0.37 | 0.10 | 0.98 | 1.42 | 0.27 |
| Ballard | 0.74 | 85% sensitivity | 0.85 | 0.45 | 0.11 | 0.97 | 1.55 | 0.33 |
| Model A (10-characteristics) | 0.88 | Youden Index | 0.78 | 0.82 | 0.25 | 0.98 | 4.35 | 0.27 |
| Model A | 0.88 | 95% sensitivity | 0.95 | 0.45 | 0.12 | 0.99 | 1.74 | 0.11 |
| Model A | 0.88 | 90% sensitivity | 0.90 | 0.64 | 0.16 | 0.99 | 2.48 | 0.16 |
| Model A | 0.88 | 85% sensitivity | 0.85 | 0.73 | 0.20 | 0.98 | 3.15 | 0.21 |
| Model B (10-characteristics+LMP) | 0.91 | Youden Index | 0.82 | 0.85 | 0.30 | 0.98 | 5.40 | 0.21 |
| Model B | 0.91 | 95% sensitivity | 0.95 | 0.52 | 0.13 | 0.99 | 1.96 | 0.10 |
| Model B | 0.91 | 90% sensitivity | 0.90 | 0.71 | 0.19 | 0.99 | 3.07 | 0.14 |
| Model B | 0.91 | 85% sensitivity | 0.85 | 0.81 | 0.26 | 0.99 | 4.49 | 0.19 |
| Model C (BW+LMP) | 0.88 | Youden Index | 0.76 | 0.84 | 0.27 | 0.98 | 4.83 | 0.29 |
| Model C | 0.88 | 95% sensitivity | 0.95 | 0.42 | 0.11 | 0.99 | 1.65 | 0.12 |
| Model C | 0.88 | 90% sensitivity | 0.90 | 0.60 | 0.15 | 0.99 | 2.27 | 0.17 |
| Model C | 0.88 | 85% sensitivity | 0.85 | 0.72 | 0.19 | 0.98 | 3.03 | 0.21 |
| Model D (BW+HC) | 0.84 | Youden Index | 0.74 | 0.79 | 0.22 | 0.97 | 3.53 | 0.33 |
| Model D | 0.84 | 95% sensitivity | 0.95 | 0.38 | 0.11 | 0.99 | 1.54 | 0.13 |
| Model D | 0.84 | 90% sensitivity | 0.90 | 0.54 | 0.13 | 0.99 | 1.97 | 0.18 |
| Model D | 0.84 | 85% sensitivity | 0.85 | 0.65 | 0.16 | 0.98 | 2.40 | 0.23 |
| Classify <34 weeks | | | | | | | | |
| LMP | 0.94 | Youden Index | 0.88 | 0.84 | 0.05 | 1.00 | 5.58 | 0.14 |
| LMP | 0.94 | 95% sensitivity | 0.95 | 0.74 | 0.03 | 1.00 | 3.65 | 0.07 |
| LMP | 0.94 | 90% sensitivity | 0.90 | 0.82 | 0.04 | 1.00 | 5.13 | 0.12 |
| LMP | 0.94 | 85% sensitivity | 0.85 | 0.87 | 0.06 | 1.00 | 6.57 | 0.17 |
| Ballard | 0.89 | Youden Index | 0.81 | 0.80 | 0.03 | 1.00 | 4.04 | 0.24 |
| Ballard | 0.89 | 95% sensitivity | 0.95 | 0.53 | 0.02 | 1.00 | 2.02 | 0.09 |
| Ballard | 0.89 | 90% sensitivity | 0.90 | 0.67 | 0.02 | 1.00 | 2.70 | 0.15 |
| Ballard | 0.89 | 85% sensitivity | 0.85 | 0.75 | 0.03 | 1.00 | 3.39 | 0.20 |
| Model A (10-characteristics) | 0.94 | Youden Index | 0.84 | 0.93 | 0.10 | 1.00 | 12.28 | 0.17 |
| Model A | 0.94 | 95% sensitivity | 0.95 | 0.62 | 0.02 | 1.00 | 2.47 | 0.08 |
| Model A | 0.94 | 90% sensitivity | 0.90 | 0.83 | 0.05 | 1.00 | 5.39 | 0.12 |
| Model A | 0.94 | 85% sensitivity | 0.85 | 0.92 | 0.09 | 1.00 | 10.93 | 0.16 |
| Model B (10-characteristics+LMP) | 0.96 | Youden Index | 0.87 | 0.95 | 0.12 | 1.00 | 15.91 | 0.14 |
| Model B | 0.96 | 95% sensitivity | 0.95 | 0.72 | 0.03 | 1.00 | 3.38 | 0.07 |
| Model B | 0.96 | 90% sensitivity | 0.90 | 0.90 | 0.08 | 1.00 | 9.11 | 0.11 |
| Model B | 0.96 | 85% sensitivity | 0.85 | 0.96 | 0.16 | 1.00 | 22.01 | 0.16 |
| Model C (BW+LMP) | 0.96 | Youden Index | 0.88 | 0.93 | 0.10 | 1.00 | 12.88 | 0.13 |
| Model C | 0.96 | 95% sensitivity | 0.95 | 0.78 | 0.04 | 1.00 | 4.25 | 0.06 |
| Model C | 0.96 | 90% sensitivity | 0.90 | 0.91 | 0.08 | 1.00 | 9.56 | 0.11 |
| Model C | 0.96 | 85% sensitivity | 0.85 | 0.95 | 0.14 | 1.00 | 18.71 | 0.16 |
| Model D (BW+HC) | 0.93 | Youden Index | 0.82 | 0.92 | 0.08 | 1.00 | 9.72 | 0.19 |
| Model D | 0.93 | 95% sensitivity | 0.95 | 0.56 | 0.02 | 1.00 | 2.18 | 0.09 |

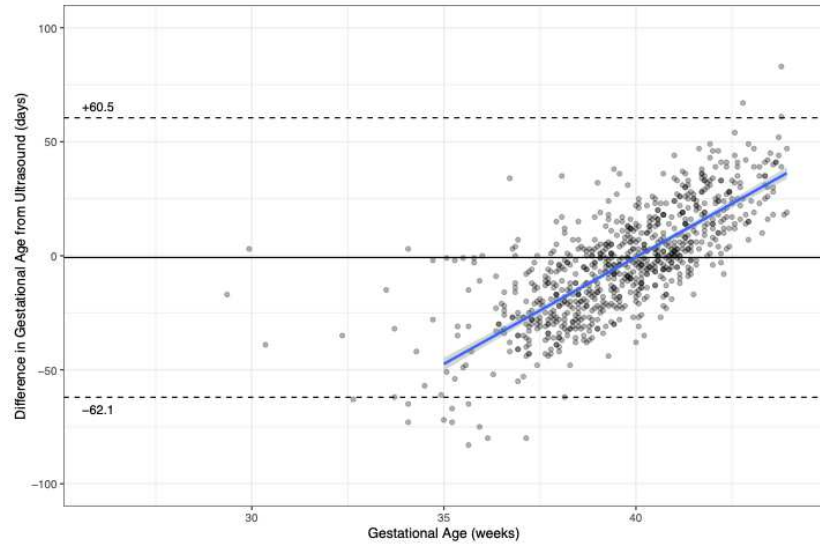
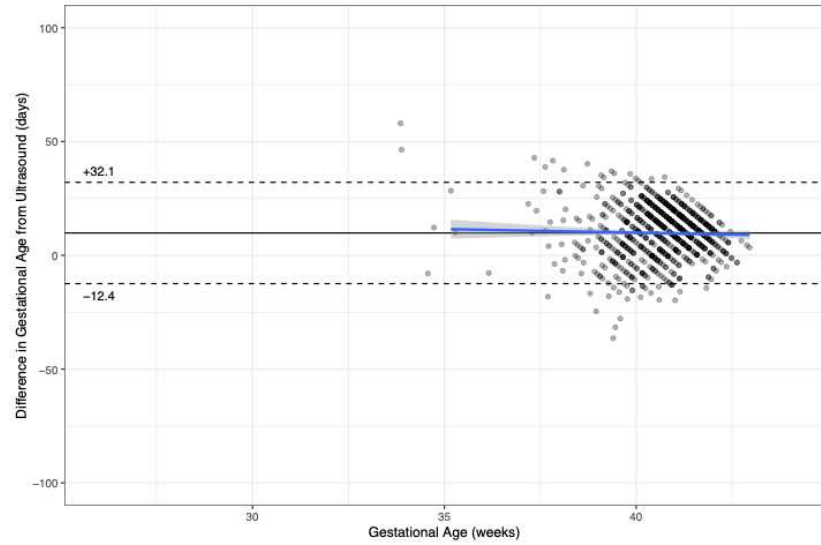
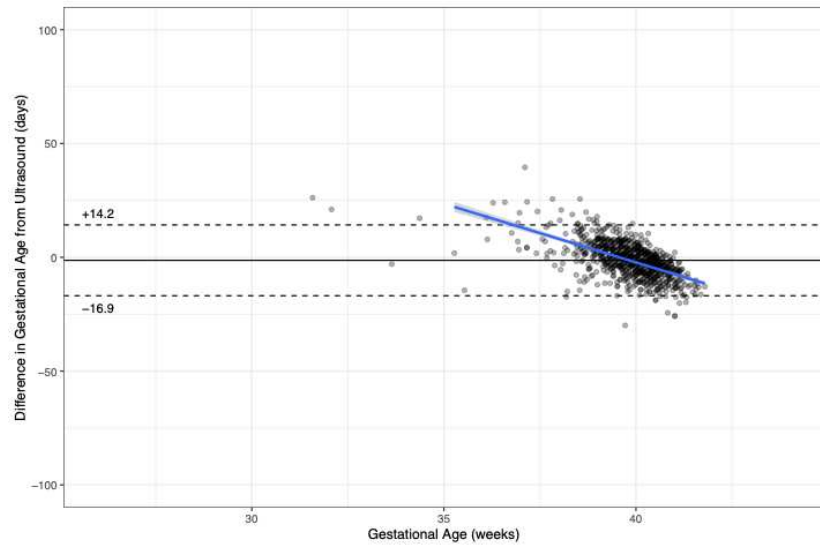
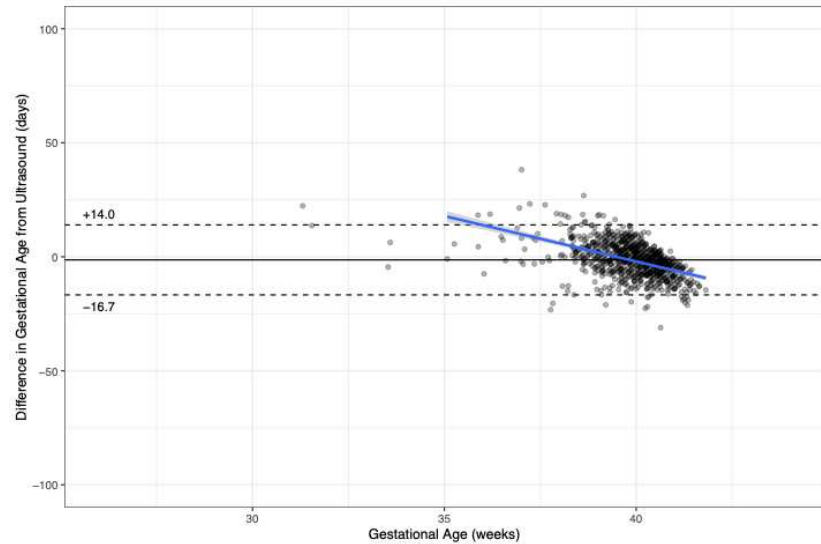
| | | | | | | | | |
|---------|------|-----------------|------|------|------|------|------|------|
| Model D | 0.93 | 90% sensitivity | 0.90 | 0.78 | 0.04 | 1.00 | 4.10 | 0.13 |
| Model D | 0.93 | 85% sensitivity | 0.85 | 0.88 | 0.06 | 1.00 | 7.19 | 0.17 |

Web Table 3. Stratified analysis by SGA vs AGA Status

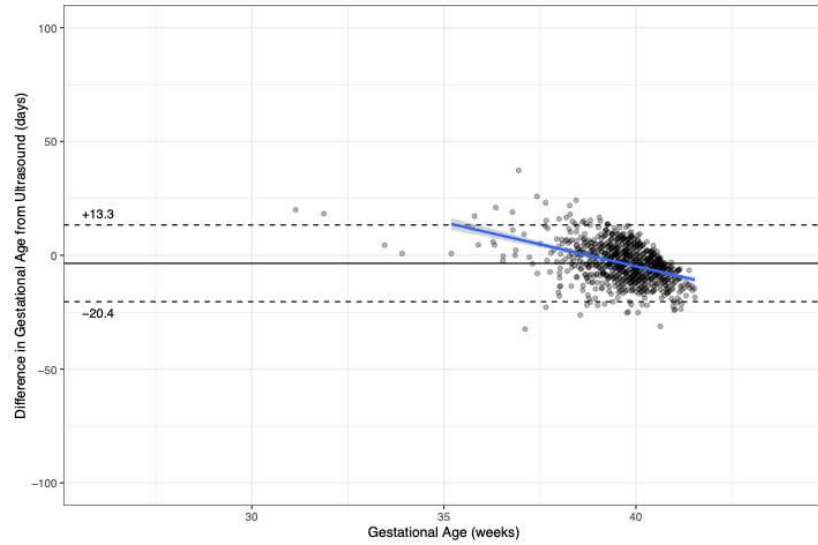
| Model | SGA | | | AGA | | |
|---|----------------------|----------------------------------|--------|----------------------|----------------------------------|--------|
| | Mean Difference | Bland Altman Limits of Agreement | BA LOA | Mean Difference | Bland Altman Limits of Agreement | BA LOA |
| LMP | -0.64 (-1.52, -0.25) | (-40.6, 39.3) | 40.0 | -1.42 (-2.54, -1.98) | (-41.2, 37.3) | 39.2 |
| Ballard | 4.85 (4.31, 5.39) | (-19.7, 29.4) | 24.6 | 10.53 (10.19, 10.86) | (-13.0, 34.1) | 23.5 |
| Model A (10 characteristics) | -5.05 (-5.38, -4.71) | (-20.1, 10.0) | 15.0 | 1.66 (1.43, 1.88) | (-14.4, 17.7) | 16.0 |
| Model B (10 characteristics+LMP) | -4.03 (-4.35, -3.72) | (-18.2, 10.1) | 14.2 | 1.31 (1.10, 1.52) | (-13.3, 15.9) | 14.6 |
| Model C (BW+LMP) | -4.49 (-4.83, -4.15) | (-19.8, 10.9) | 15.4 | 1.50 (1.29, 1.72) | (-13.8, 16.8) | 15.3 |
| Model D (BW+HC) | -5.61 (-5.95, -5.27) | (-20.9, 9.7) | 15.3 | 1.93 (1.68, 2.17) | (-15.1, 19.0) | 17.0 |

Web Figure 2a-f. Bangladesh Bland Altman Curves**2a. Bangladesh LMP vs Ultrasound****2b. Bangladesh Ballard vs Ultrasound**

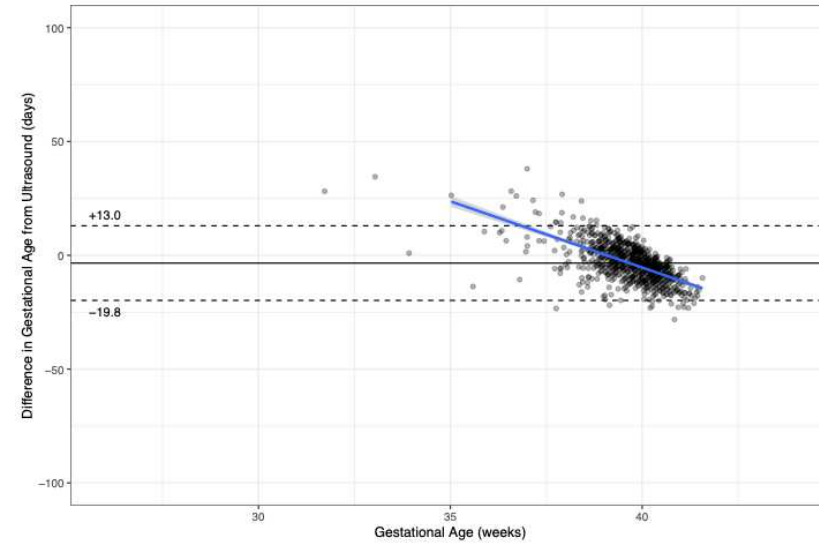
2c. Bangladesh Model A vs Ultrasound**2d. Bangladesh Model B vs Ultrasound****2e. Bangladesh Model C vs Ultrasound****2f. Bangladesh Model D vs Ultrasound**

Web Figure 3a-f. Ghana Bland Altman Curves**3a. Ghana LMP vs Ultrasound****3b. Ghana Ballard vs Ultrasound****3c. Ghana Model A vs Ultrasound****3d. Ghana Model B vs Ultrasound**

3e. Ghana Model C vs Ultrasound

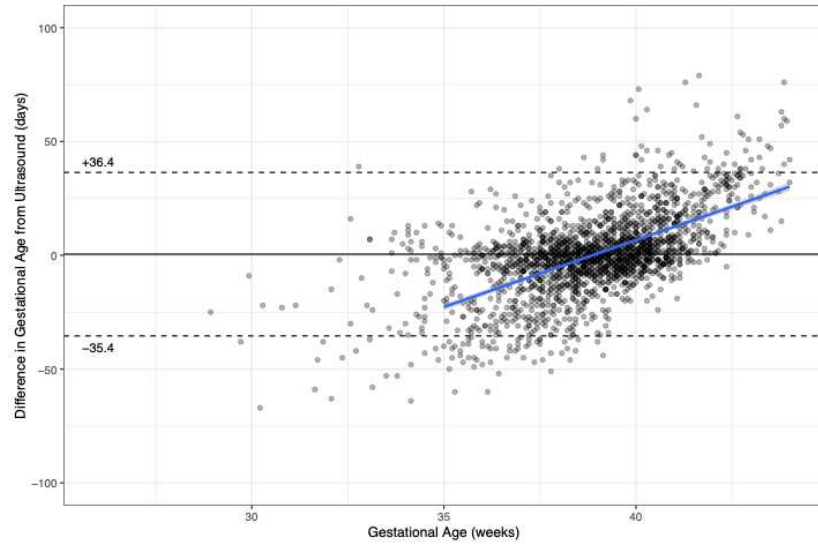


3f. Ghana Model D vs Ultrasound

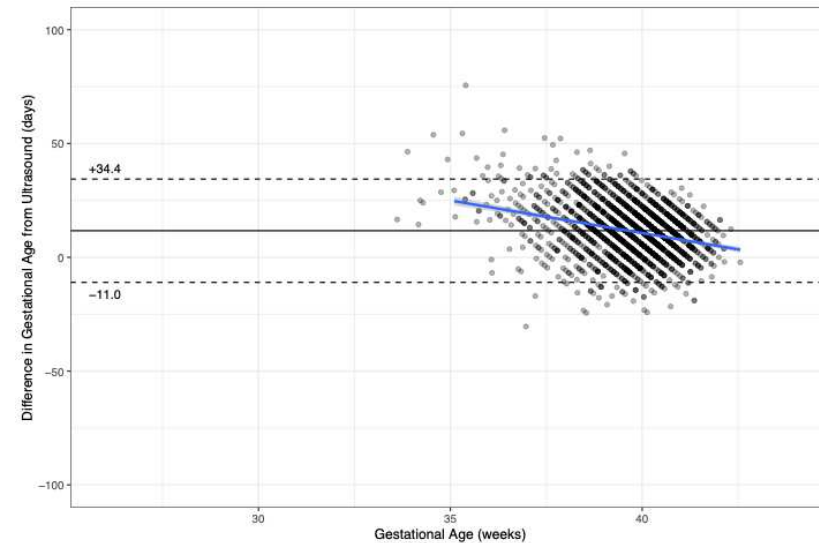


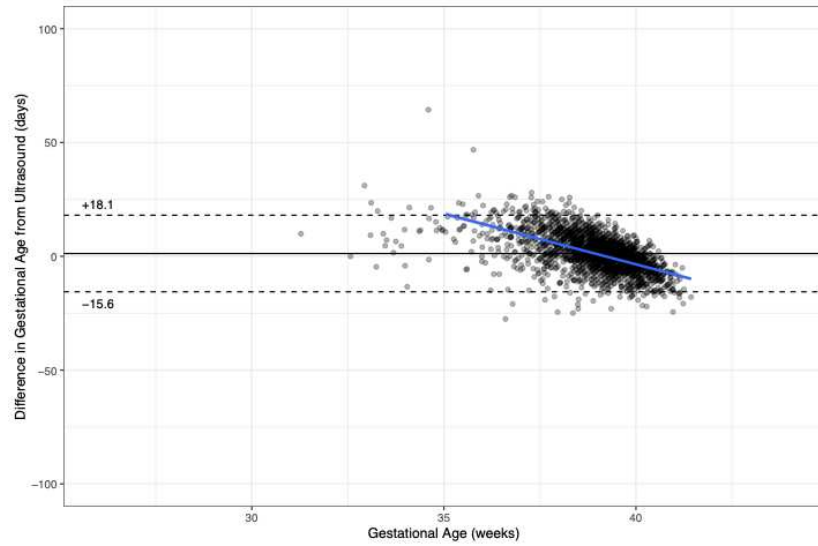
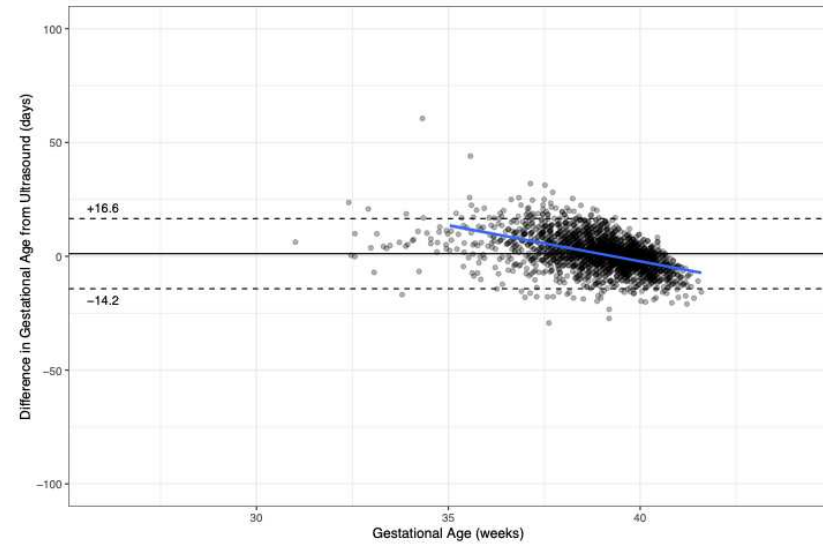
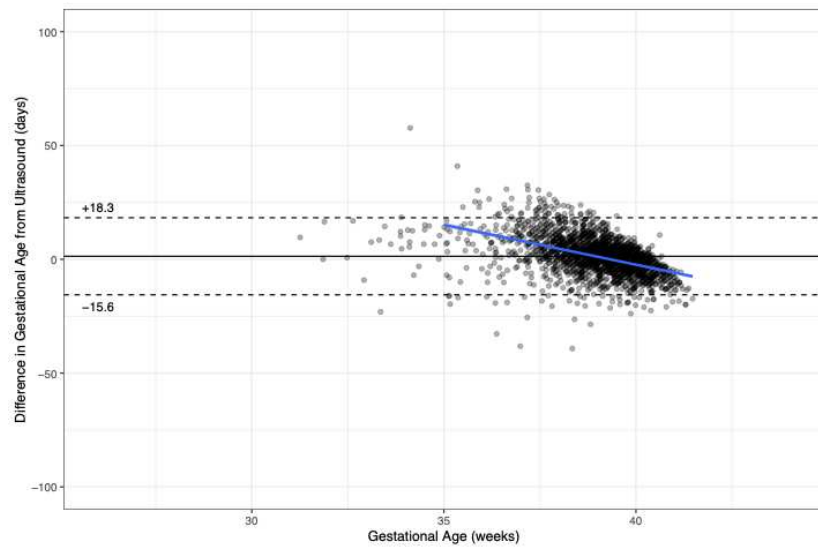
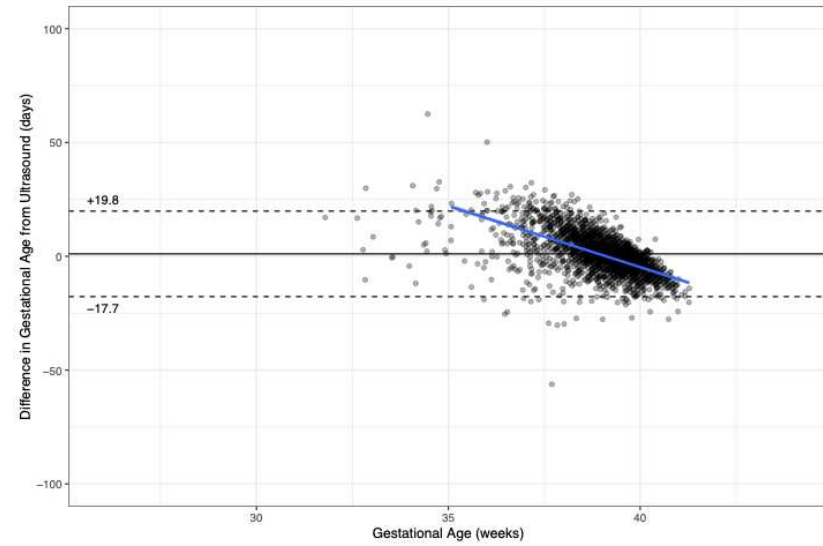
Web Figure 4a-f. Pakistan (Karachi) Bland Altman Curves

4a. Karachi LMP vs Ultrasound



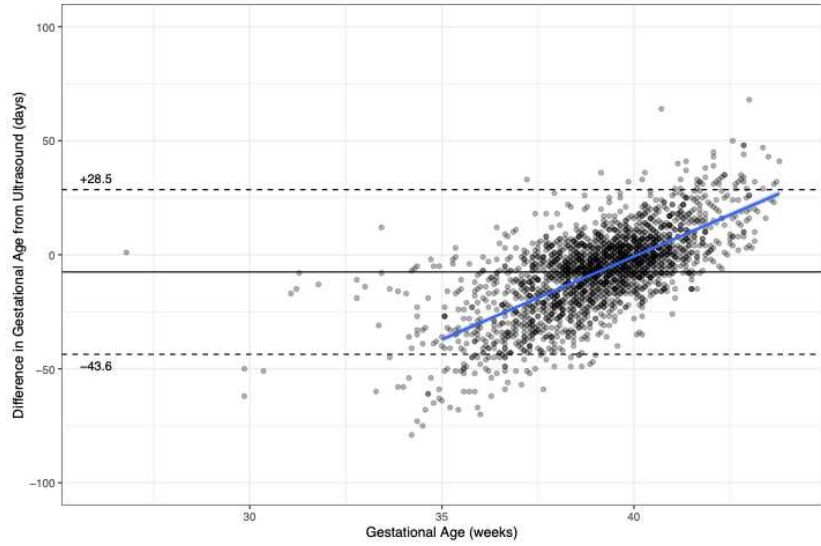
4b. Karachi Ballard vs Ultrasound



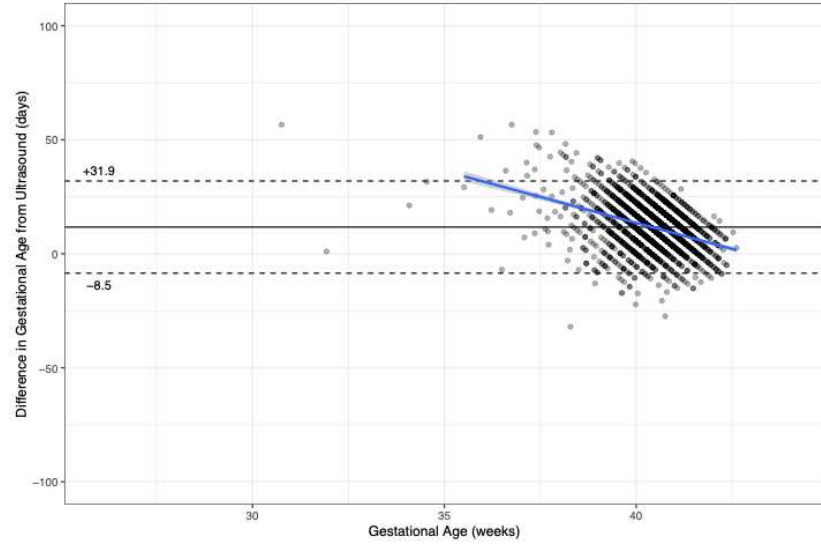
4c. Karachi Model A vs Ultrasound**4d. Karachi Model B vs Ultrasound****4e. Karachi Model C vs Ultrasound****4f. Karachi Model D vs Ultrasound**

Web Figure 5a-f. Tanzania (Pemba) Bland Altman Curves

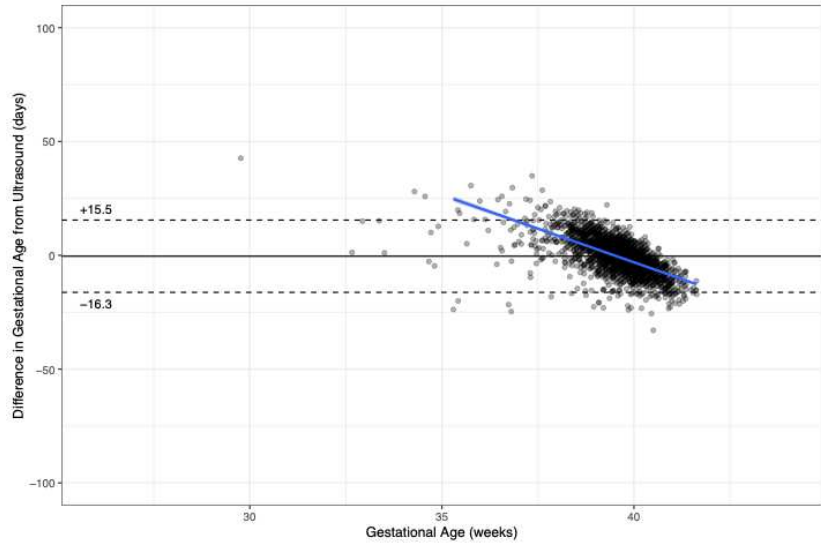
5a. Pemba LMP vs Ultrasound



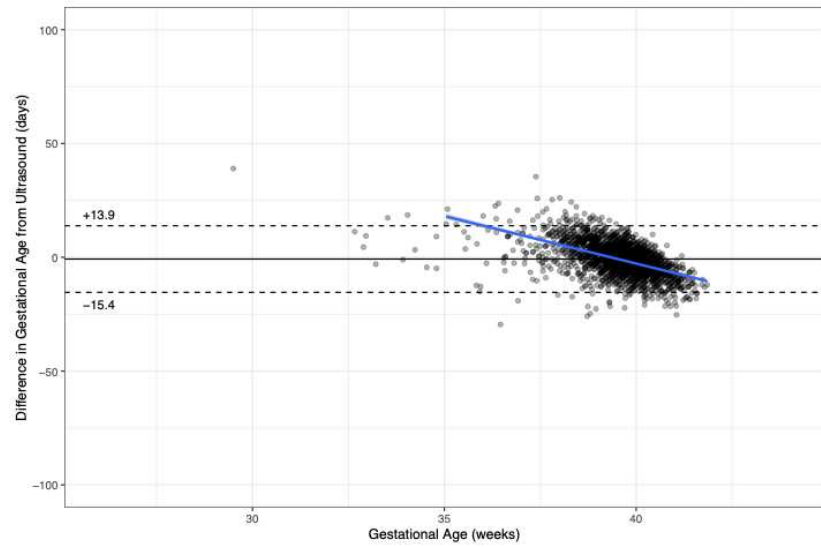
5b. Pemba Ballard vs Ultrasound



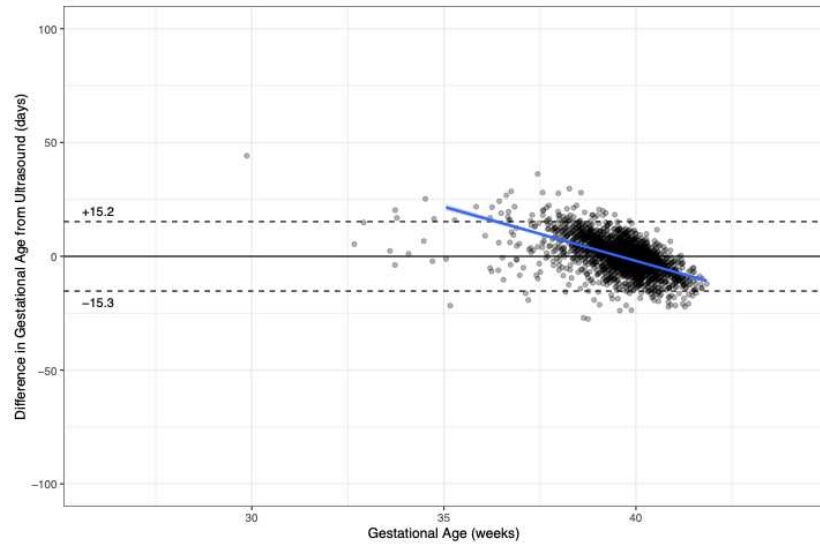
5c. Pemba Model A vs Ultrasound



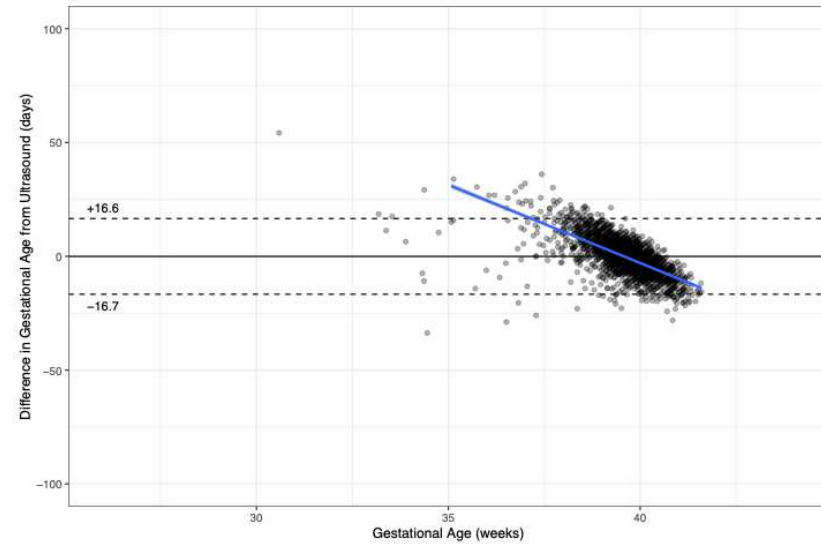
5d. Pemba Model B vs Ultrasound



5e. Pemba Model C vs Ultrasound

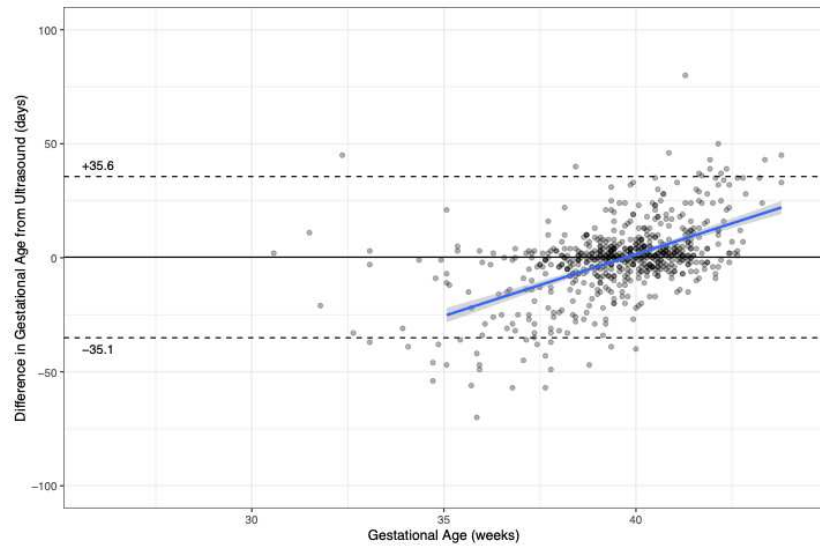


5f. Pemba Model D vs Ultrasound

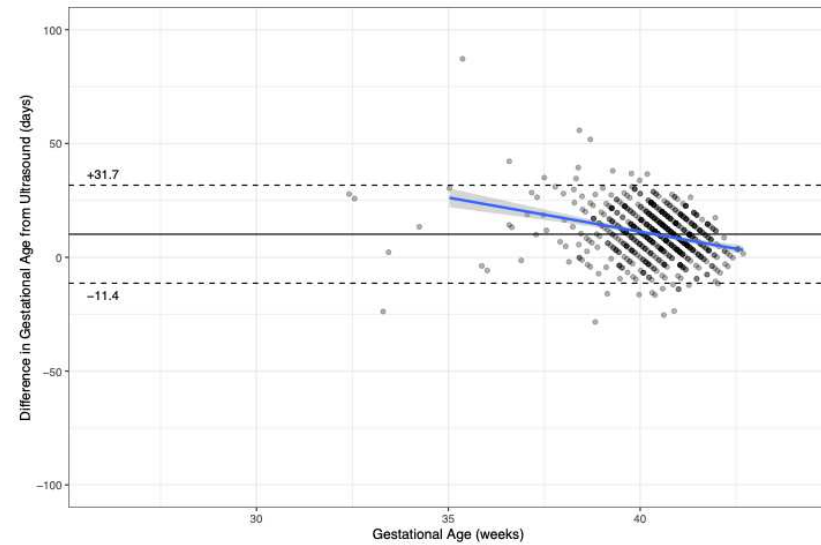


Web Figure 6a-f. Zambia Bland Altman Curves

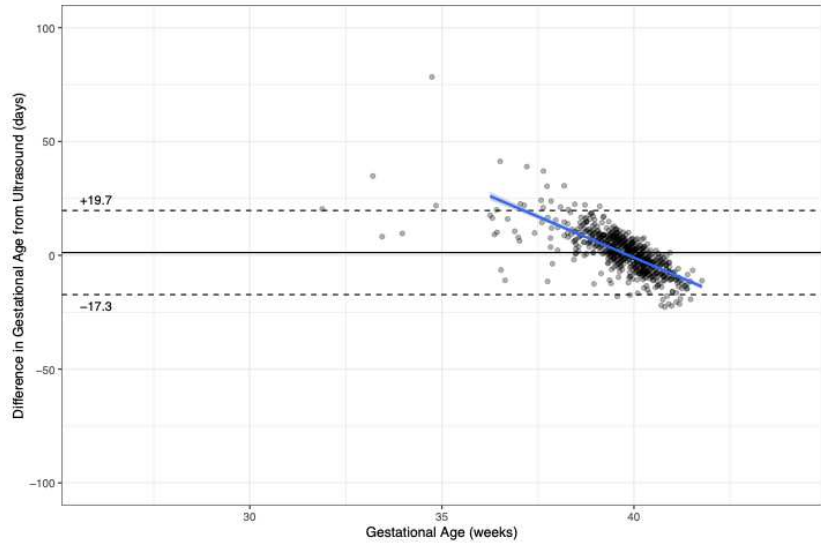
6a. Zambia LMP vs Ultrasound



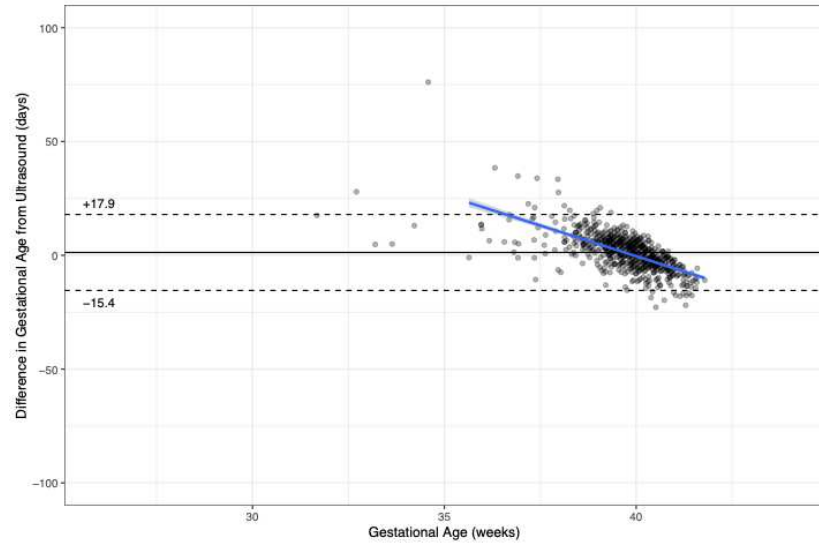
6b. Zambia Ballard vs Ultrasound



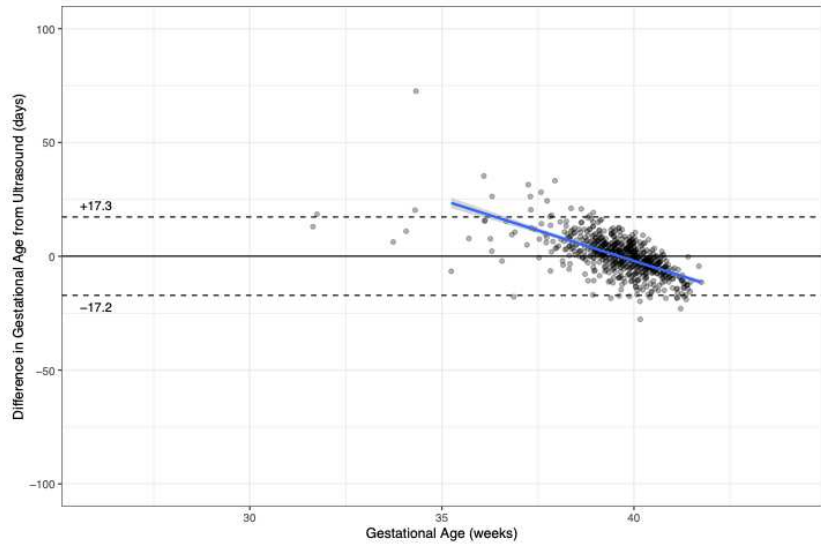
6c. Zambia Model A vs Ultrasound



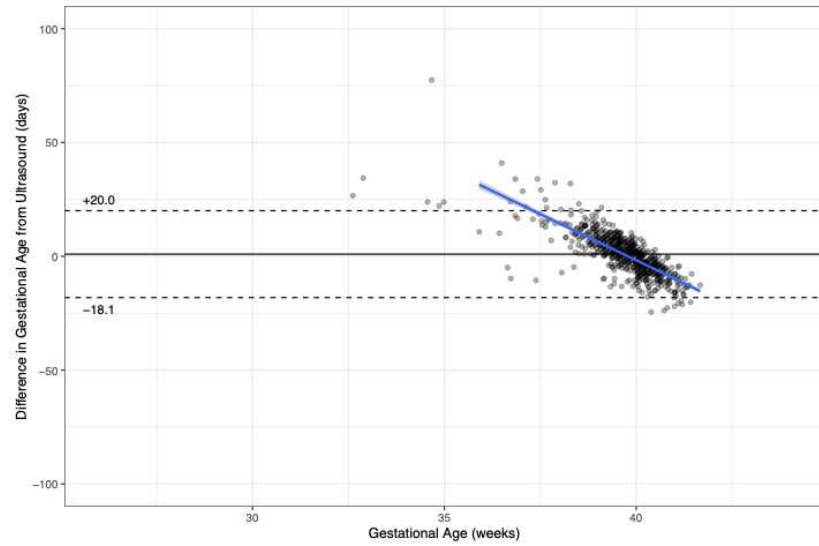
6d. Zambia Model B vs Ultrasound

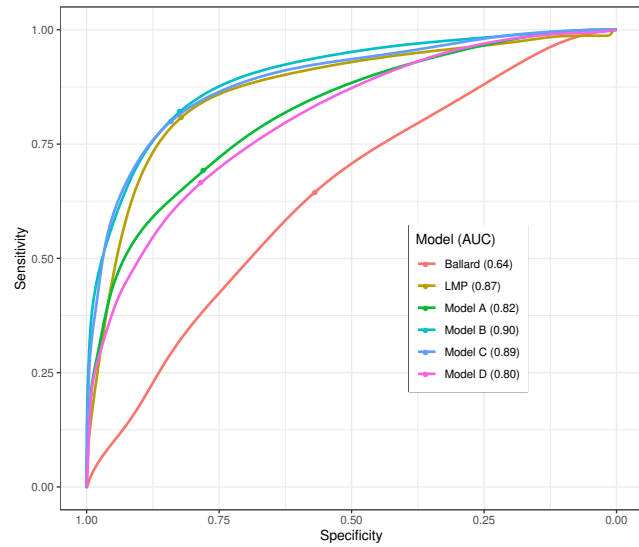
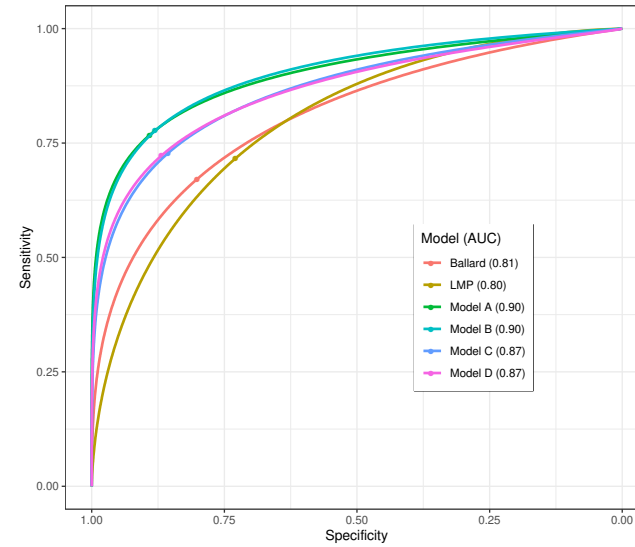
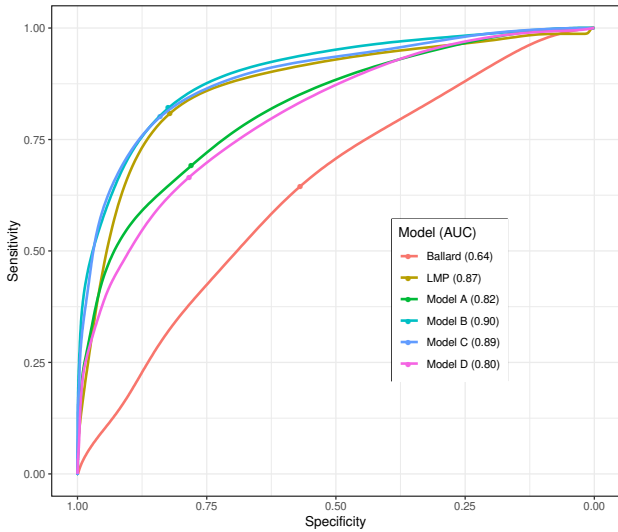
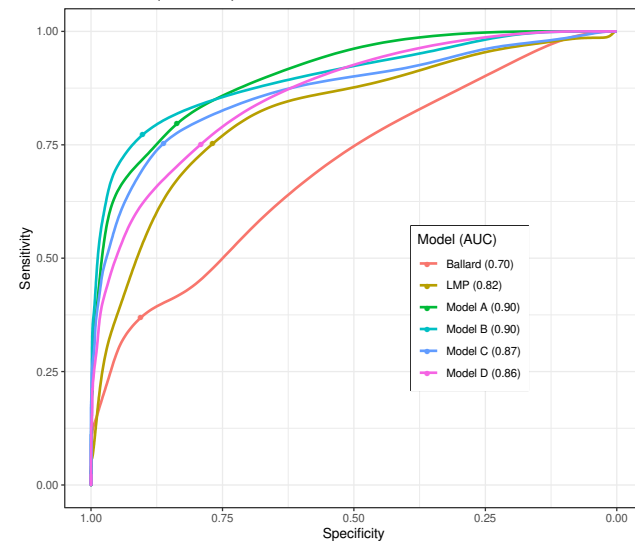


6e. Zambia Model C vs Ultrasound

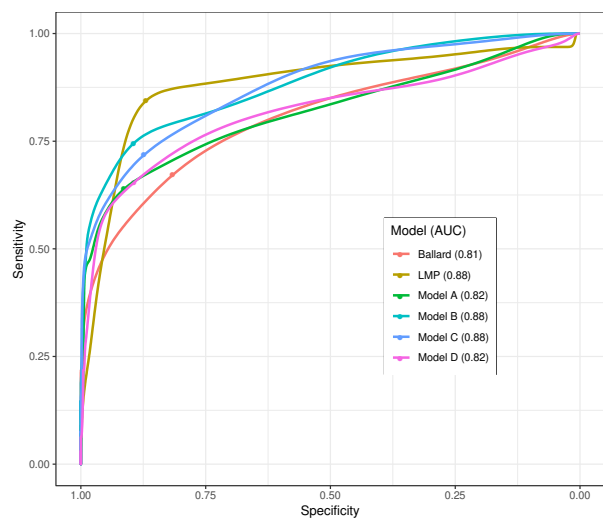


6f. Zambia Model D vs Ultrasound



Web Figure 7a-e. Receiver Operating Curves for the Identification of Preterm Births (<37 weeks)**WHO AMANHI Cohorts - Site Specific Data****7a. Bangladesh****7b. Ghana****7c. Pakistan (Karachi)****7d. Tanzania (Pemba)**

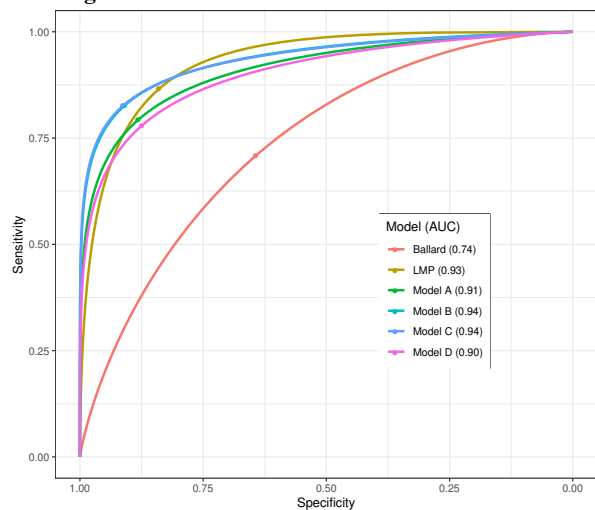
7e. Zambia



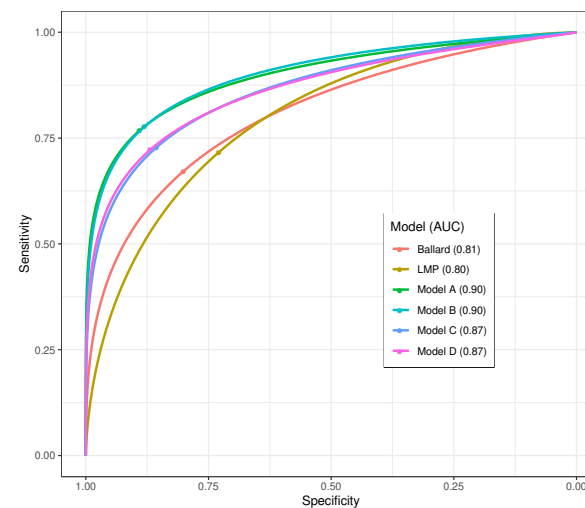
Web Figure 8.a-e Receiver Operating Curves for the Identification of Preterm Births (<34 weeks)

WHO AMANHI Cohorts - Site Specific Data

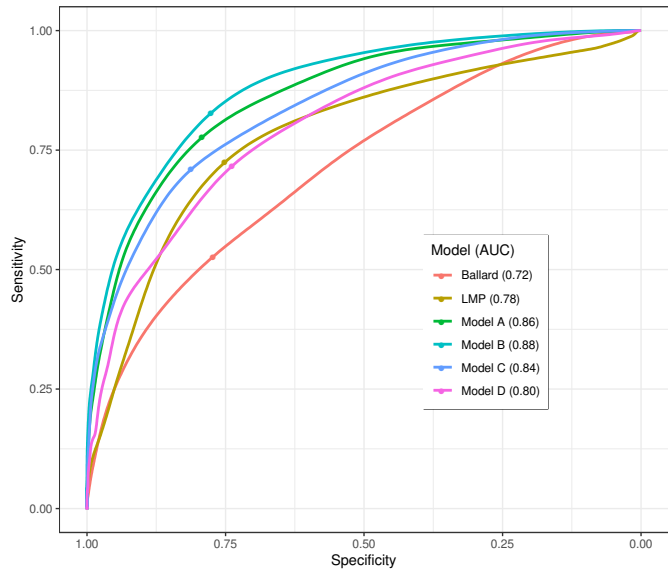
8a. Bangladesh



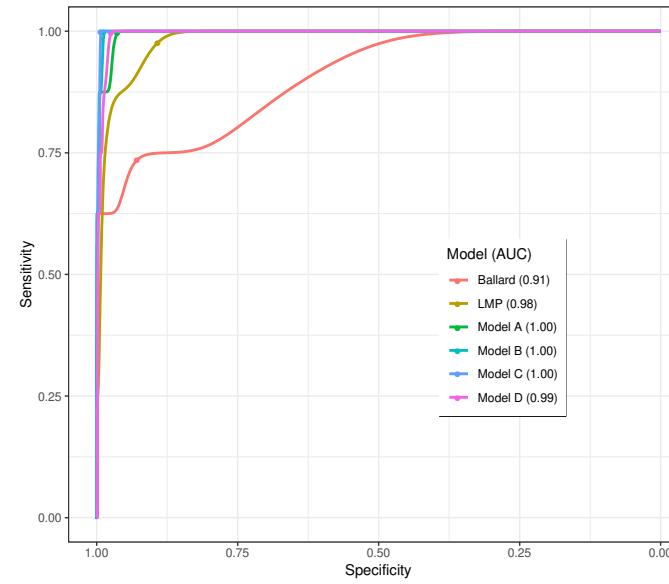
8b. Ghana



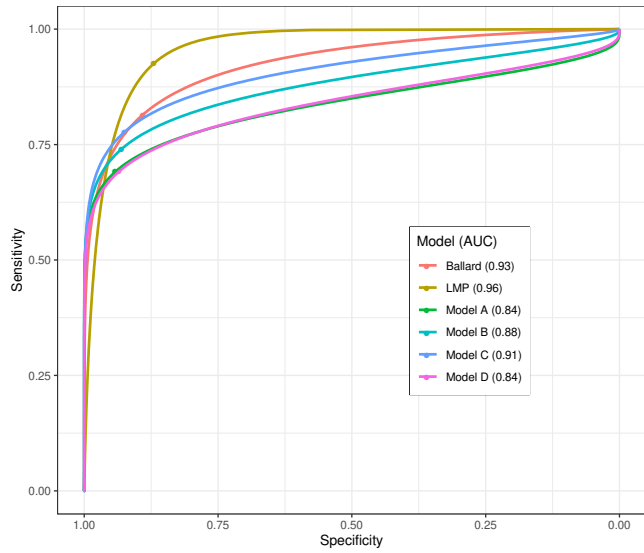
8c. Pakistan (Karachi)



8d. Tanzania (Pemba)

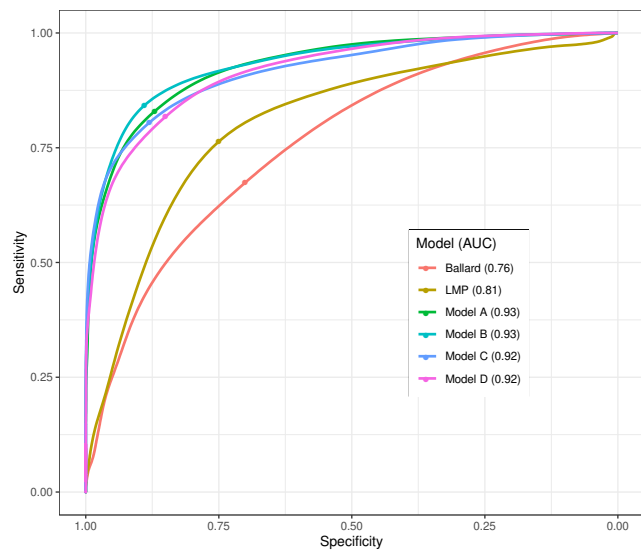


8e. Zambia

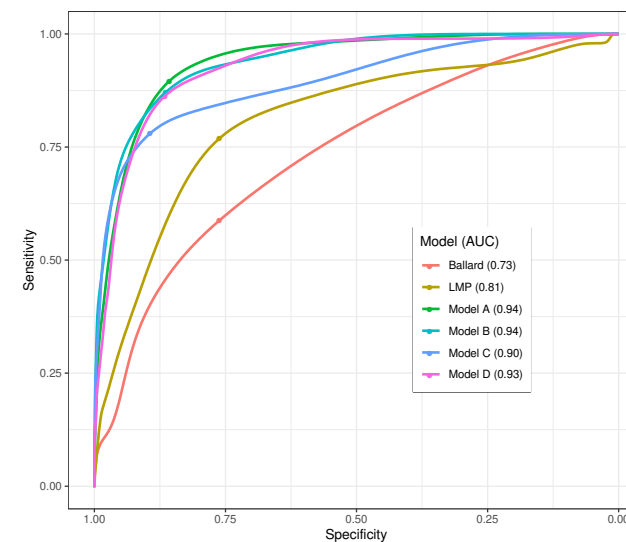


Web Figure 9a-d. ROC Curves by AGA and SGA status for classification of infants <37 weeks and <34 weeks

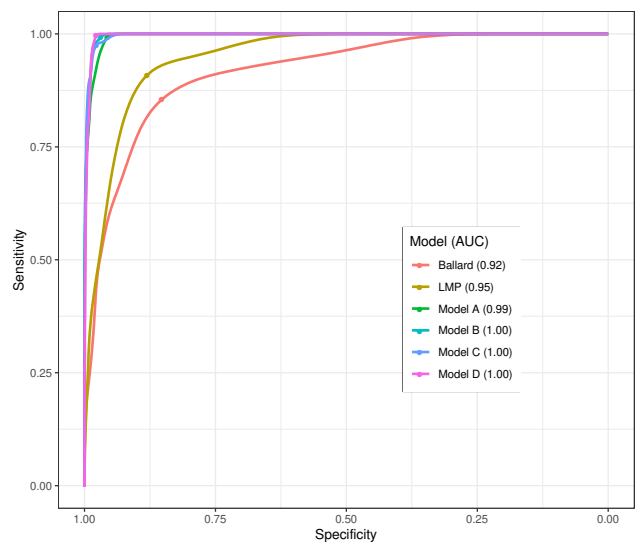
9a. Preterm (<37 weeks) classification among AGA infants



9b. Preterm (<37 weeks) classification among SGA infants



9c. Early preterm (<34 wk) classification among AGA infants



9d. Early preterm (<34 wk) classification among SGA infants

