



Use of magnetic resonance imaging in the diagnosis of fetal vertebral abnormalities in utero: a single-center retrospective cohort study

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Background: Magnetic resonance imaging (MRI) has been used increasingly as an adjunct examination to ultrasound (US) for the evaluation of fetal anomalies. The purpose of this study was to determine whether the accuracy and confidence of diagnosing fetal vertebral anomalies are improved with MRI. We also assessed whether fetal MRI provides additional information when diagnosing fetal vertebral anomalies.

Methods: We performed a single-center, retrospective study of 127 pregnant women with fetuses suspected of having vertebral anomalies on US examination; women also underwent fetal MRI scanning. Comparisons of diagnostic accuracy and confidence were made between MRI and US for the identification of fetal vertebral anomalies. We also assessed any additional information provided by MRI. McNemar's paired binomial test, chi-square test, or Fisher's exact test were used to compare the diagnostic ability between MRI and US. In all cases, postnatal or postmortem imaging findings were used as reference standards.

Results: A total of 127 participants were recruited between December 2015 and January 2021. Fetal vertebral anomalies were detected in 63.8% (81/127) cases and found to be negative in 36.2% (46/127) of cases at follow up. The diagnostic accuracy of vertebral anomalies was 46.9% (38/81) for US and 84.0% (68/81) for MRI [difference, 37.1%; 95% confidence interval (CI): 27% to 48%; $P < 0.001$]. Both MRI and US were concordant and correct in 36.2% (46/127) of fetuses; MRI provided additional information for 16.5% (21/127) of fetuses, and corrected US diagnoses of 36.2% (46/127) of fetuses; both MRI and US were not consistent with postnatal findings in 10.2% (13/127) of fetuses, and the remaining fetus (0.8%, 1/127) was diagnosed correctly using US but failed to be diagnosed by MRI. Diagnoses were reported with high confidence using MRI in 95.3% (121/127) of cases and 73.2% (93/127) using US.

Conclusions: Fetal vertebral MRI improves the accuracy and confidence of diagnosing fetal vertebral anomalies. This finding indicates that fetal MRI supplements the information provided by US and that MRI may be a good complement in selected fetuses, when US can either not achieve a definite diagnosis or there is doubt regarding its reliability. Thus, MRI may be used to inform prenatal counseling and management decisions.

Keywords: Fetal imaging; fetal vertebra; ultrasound; fetal vertebral anomalies

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Introduction

Fetal vertebral anomalies result from abnormal embryonic development during gestational weeks 4–8 (1) and often lead to asymmetric spinal growth, which is classified as congenital scoliosis or kyphosis. The prognosis of fetal vertebral anomalies is related to the type, site, the number of the affected vertebra, and the associated anomalies. Accurate antenatal diagnosis of vertebral anomalies is essential for planning of postnatal follow up to optimize outcomes (2).

Ultrasound (US) is the primary screening method for fetal skeletal evaluation (3). However, certain fetal and maternal factors such as oligohydramnios, advanced gestational age, unfavorable fetal position, maternal abdominal wall scarring, or maternal obesity (4) can undermine the quality of US images. Previous studies have demonstrated a low detection rate in antenatal diagnosis of skeletal anomalies (5–8), especially where there is isolated vertebral involvement without a spinal curvature deformity.

Over the past several years, magnetic resonance imaging (MRI) has been used increasingly as an adjunct examination to US for the evaluation of fetal anomalies (9,10) and has proven useful in imaging fetal spinal canals and cord pathologies (11) with ultrafast spin-echo T2-weighted imaging sequences (12–17). However, literature regarding the use of MRI for the diagnosis of the musculoskeletal system (18–21) and, in particular, bony spinal structures, is scarce and mainly focuses on postmortem imaging examinations without vertebral developmental abnormalities (22–26) or case reports (27,28). Recently, with the improvement and modification of fetal imaging of the susceptibility-weighted imaging (SWI) sequence, the application of fetal MRI for bony spinal structures has attracted mounting attention (29,30). Moreover, at our center (Shandong Provincial Hospital Affiliated to Shandong First Medical University), along with conventional sequences, SWI has been routinely applied to all fetuses suspected of vertebral anomalies on US. In this article, we aimed to compare the accuracy and confidence of diagnosing fetal vertebral anomalies between MRI and US and determine whether MRI can be used successfully as a complementary imaging method in combination with US to optimize prenatal counseling and postnatal management.

We present the following article in accordance with

the Standards for Reporting Diagnostic accuracy studies (STARD) reporting checklist (available at <https://qims.amegroups.com/article/view/10.21037/qims-21-1070/rc>).

Methods

Study population

This retrospective study was conducted in accordance with the Declaration of Helsinki (as revised in 2013) and approved by the Institutional Review Board of Shandong Provincial Hospital Affiliated to Shandong First Medical University. Patients were recruited consecutively, and all participants provided written informed consent. The inclusion criteria was fetuses suspected to have vertebral anomalies on US. The exclusion criteria included women with contraindications to MRI; fetuses without definite follow-up results and incomplete images or poor image quality. A total of 269 second or third trimester pregnant women carrying a fetus with suspected vertebral anomalies (screened by US) were recruited to the study between December 2015 and January 2021. All participants had inconclusive or uncertain findings on US. Of these 269 cases, 128 had no confirmed follow-up results, and 14 cases were excluded, leaving a sample population of 127 fetuses with suspected vertebral anomalies. The participants were then divided into 2 groups: a <28-week gestational age (GA) group (n=68) and a ≥28-week GA group (n=59). The reasons for using a GA of 28 weeks as the dividing point were that problems raised by parents and clinicians at mid-pregnancy needed to be addressed, and image quality improves with GA (30). *Figure 1* shows the selection characteristics of the patients.

Imaging acquisition

The US examinations were performed at the Department of Ultrasound, Shandong Provincial Hospital Affiliated to Shandong First Medical University using a Voluson E10 (GE Healthcare, Waukesha, WI, USA), Philips EPIC 5 or 7 (Philips Healthcare, Best, The Netherlands), or a UGEO WS80A (Samsung, Seoul, Korea) ultrasound unit outfitted with a 3.5–5.0 MHz frequency probe using standard US imaging techniques (31), and static two-dimensional (2D)-US or three-dimensional (3D)-US were used depending on

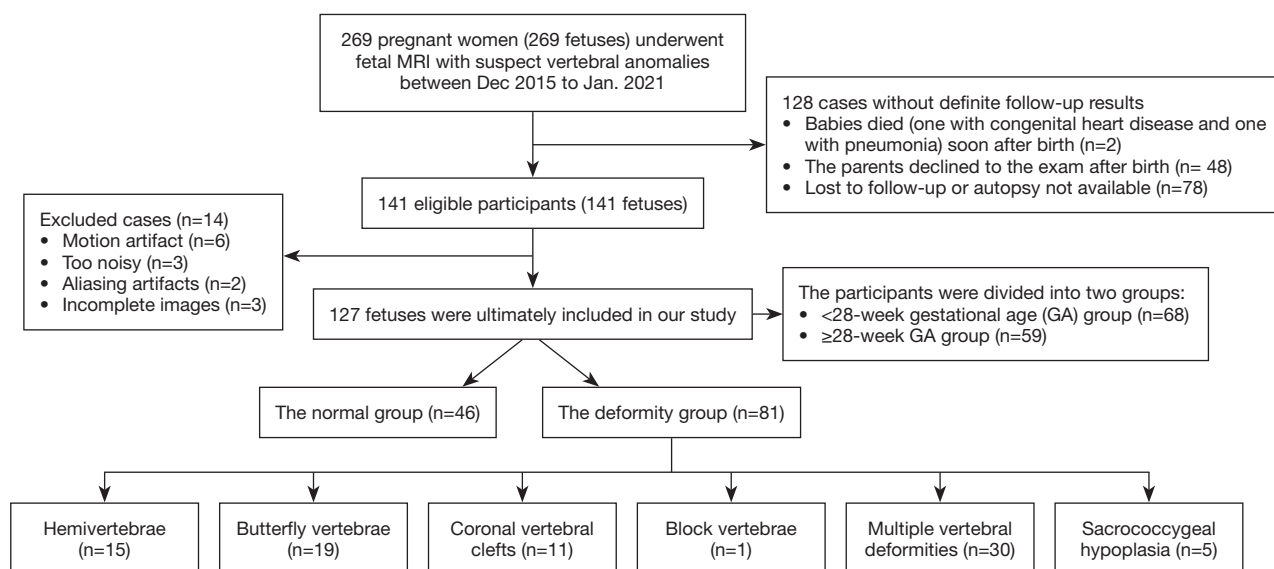


Figure 1 Flowchart of patients with selection data and fetal MRI included in the analysis. MRI, magnetic resonance imaging.

clinical demand. The 3 planes of imaging commonly used to assess the fetal spine from the cervical region through the coccyx included the coronal, parasagittal, and transverse planes. In the transverse plane, it was stipulated that all 3 ossification centers should be visualized, and the centers of the neural arches should be parallel or converging (31). The parallel configuration is particularly noticeable when the fetus is in a decubitus position with respect to the transducer (32). In the longitudinal plane, the spine has a ‘railroad track’ appearance, with gradual widening towards the fetal head and gradual tapering in the sacrum (33). Fetal MRI examinations were performed within 3 days of the anomalous US findings. All MRI examinations were performed on a 1.5-T MAGNETOM Amira (Siemens, Shenzhen Magnetic Resonance, Ltd., Shenzhen, China) with an 18-channel spine coil and a 13-channel body coil positioned over the lower pelvic area. All cases were imaged in the supine or left-lateral position. The MRI protocol for fetal spine imaging consisted of 2D half-fourier acquisition single-shot turbo spin echo (HASTE), true fast imaging with steady-state (TrueFISP), SWI in 3 orthogonal planes, and a T1-weighted ultrafast sequence in at least one plane (usually sagittal). The SWI was performed immediately after the HASTE or TrueFISP sequences by simply copying the slice position to reduce the chance of fetal motion.

The imaging parameters for the HASTE, TrueFISP, and SWI sequences were as follows: (I) HASTE: repetition time/echo time (TR/TE)=1,300/93 ms; flip angle =180°; field of view

(FOV) =308.8 mm × 380 mm; matrix =198×256; bandwidth =698 Hz/pixel; slice thickness =4.0 mm; gap =0; number of slices =15; spatial resolution =1.5 mm × 1.5 mm × 4.0 mm; acquisition time =21 s; and free breathing. (II) TrueFISP: TR/TE=4.06/1.76 ms; flip angle =79°; FOV =310 mm × 380 mm; matrix =198×304; bandwidth =685 Hz/pixel; slice thickness =4.0 mm; gap =0; number of slices =20; spatial resolution =1.3 mm × 1.3 mm × 4.0 mm; acquisition time =12 s; and free breathing. (III) SWI: TR/TE =85/12.40 ms; flip angle =15°; FOV =244.8 mm × 300 mm; matrix =166×256; bandwidth =80 Hz/pixel; slice thickness =3.0 mm; gap =0; number of slices =8; reconstructed spatial resolution =0.6 mm × 0.6 mm × 3.0 mm; acquisition time =26 s; and two breath-holds, each over a period time of 13 s. The entire MRI examination, including setup, was completed within 30 min. No fetal or maternal sedation was used for any of the examinations, and specific absorption rate limits were at or below the recommended levels.

Outcome reference diagnoses

The results of postnatal or postmortem imaging [including X-ray, computerized tomography (CT), and MRI] were considered the diagnostic gold standard for US and MRI detection of vertebral anomalies. Whenever possible, MRI scans or X-rays were performed during follow up to avoid further radiation exposure, except when multiple vertebral malformations required 3D CT reconstruction for a

definitive diagnosis. The MRI examinations of the spine were performed on a 3.0-T system (MAGNETOM Skyra, Siemens Healthcare, Erlangen Germany) or 1.5-T system (MAGNETOM Amira, Siemens Healthcare, Germany). The imaging protocols included T2-weighted imaging (T2WI) in the axial, coronal, and sagittal planes and T1-weighted imaging (T1WI) in the sagittal plane. Spinal CT examinations in cases (supine position) suspected of spinal deformities were performed on a 128-channel, dual source, multidetector-row CT scanner (Siemens Definition Flash, Siemens Healthcare, Germany) as per standard of care. Normalized effective doses for all cases were determined from dose length product (DLP) using conversion factors for CT imaging of the trunk (children >1 to <3 years, 0.028; ≤1 year, 0.044) (34).

Image interpretation and diagnostic accuracy

For fetal vertebral anatomical orientation, the most caudal fetal rib corresponded to the T12 level (35), and the superior aspect of the iliac crest corresponded to the L5 level (35).

All fetal MRI images were assessed by 2 radiologists (** and **, with 5 and 10 years' experience in prenatal and pediatric radiology, respectively), who were blinded to the postnatal results. The US studies were interpreted by 2 sonographers (** and **, with more than 15 years' experience in the prenatal diagnosis of congenital anomalies, respectively). Postnatal or postmortem images were assessed by ** and ** (both with 5 years' experience in prenatal and pediatric radiology). Discrepancies were resolved through consensus. The radiologists were made aware of the diagnoses from US results before the fetal MRI study was completed. In cases with multiple anatomical diagnoses, all diagnoses had to be reported accurately in imaging interpretation to be classified as correct. The diagnostic accuracies for US and MRI studies were determined using the following equation:

$$Accuracy = \frac{(true\ positives + true\ negatives)}{Total\ number\ of\ cases} \quad [1]$$

where true positives and negatives were established at follow-up assessments.

Diagnostic confidence

Assessment of diagnostic confidence in this study was purely

descriptive. The level of confidence for each diagnosis of a vertebral anomaly was determined using a 5-point Likert scale (36): “very unsure” (10% certain), “unsure” (30% certain), “equivocal” (50% certain), “confident” (70% certain), and “highly confident” (90% certain). Assessments of the diagnostic confidence were performed by comparing the level of confidence of an US diagnosis and a fetal MRI diagnosis with the accuracy of diagnosis obtained from follow-up examination. Diagnostic confidence of the dominant diagnosis from the MRI and US images (that most likely to influence prognoses) derived from the Likert scales were converted to high confidence (70% and 90%) or low confidence (10%, 30%, and 50%) (37).

Statistical analysis

Statistics analyses were conducted using the software package SPSS 22.0 (IBM Corp., Armonk, NY, USA). Diagnostic accuracy was calculated for both GA groups (<28 and ≥28 weeks) and for the sample as a whole with a McNemar's paired binomial test. Diagnostic accuracies were compared between MRI and US for the subgroups using a chi-square test or Fisher's exact test. A P value (2-tailed) of less than 0.05 was considered to indicate a significant difference.

Results

Participant characteristics

Paired MRI and US data were available with follow-up results for 127 of the pregnant women in this study. The possible non-random sample of cases without follow-up results (n=128) favored either modality (MRI or US), and 14 cases were excluded because of incomplete MRI images or poor image quality, which contributes to bias in the case of US. Of the 127 pregnant women, 116 cases carried pregnancies to term (mean GA, 28±4 weeks), and 11 cases resulted in a termination of pregnancy (mean GA, 27±3 weeks). Characteristics of the study population are shown in *Table 1*.

Follow-up results and diagnostic performance

Postnatal and postmortem imaging results confirmed that 36.2% (46/127) of the fetuses had normal vertebra (*Figure 2A-2F*), and 63.8% (81/127) had vertebral anomalies at follow up. The 81 specific vertebral anomalies included

Table 1 Characteristics of the study population

| Characteristics | Follow-up results available (n=127) | Follow-up results unavailable (n=128) | Excluded cases (n=14) |
|--------------------------------------------|----------------------------------------|------------------------------------------|--------------------------|
| Maternal age at diagnosis (years) | 29.8 (4.3) | 27.6 (4.5) | 25.8 (3.2) |
| Gestational age at diagnosis | | | |
| Mean age (weeks) | 28.2 (3.8) | 30.9 (3.2) | 24.3 (2.8) |
| <28 weeks | 68 (53.5) | 61 (47.7) | 10 (71.4) |
| ≥28 weeks | 59 (46.5) | 67 (52.3) | 4 (28.6) |
| Fetal presentation | | | |
| Head | 101 (79.5) | 100 (78.1) | 13 (92.9) |
| Breech | 25 (19.7) | 28 (21.9) | 1 (7.1) |
| Transverse | 1 (0.8) | 0 | 0 |
| Pregnancy options | | | |
| Continued pregnancy | 116 (91.3) | 50 (39.1) | 9 (64.3) |
| Termination of pregnancy/lost to follow up | 11 (8.7) | 78 (60.9) | 5 (35.7) |
| Age at follow up | | | |
| Postnatal age at imaging (years) | 2.3 (1.4) | – | – |
| Postmortem age at imaging (weeks) | 26.5 (3.8) | – | – |
| US site | | | |
| Our center | 84 (66.1) | 90 (70.3) | 10 (71.4) |
| Other centers | 43 (33.9) | 38 (29.7) | 4 (28.6) |

Data are mean (SD) or n (%). US, ultrasound; our center refers to Shandong Provincial Hospital Affiliated to Shandong First Medical University.

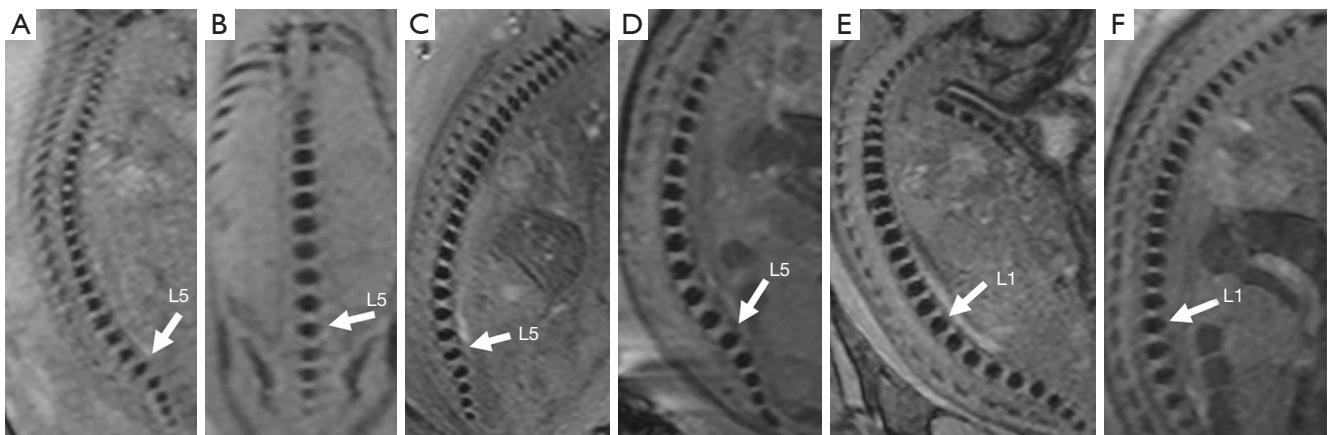


Figure 2 Fetuses with normal vertebral structures. (A) Case 36, 24 weeks; (B) Case 32, 26 weeks; (C) Case 26, 29 weeks +5 d; (D) Case 97, 31 weeks +1 d; (E) Case 43, 34 weeks +6 d; (F) Case 27, 36 weeks. SWI images showed excellent depiction (anatomical location indicated by the arrows) between the bone and surrounding soft tissues, and image quality appears to be improved with increasing gestational ages. SWI, susceptibility-weighted imaging.

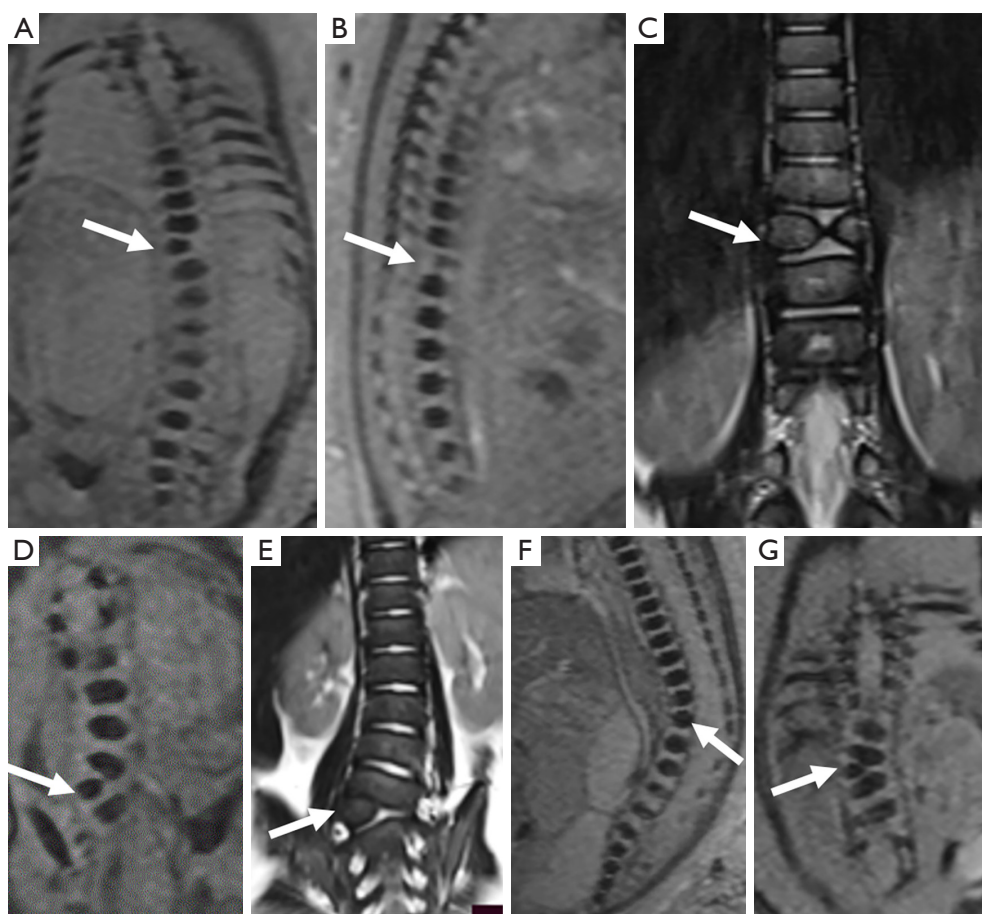


Figure 3 Fetuses with vertebral anomalies. (A-C) Case 56, 31 weeks, T11 was wedge-shaped in coronal and sagittal SWI images (A, B, arrows), diagnosed hemivertebra on prenatal MRI and proved to be butterfly vertebra (C, arrow) at 3 years on postnatal coronal T2WI; (D-E) Case 101, 33 weeks, L5 hemivertebra on prenatal MRI (D, arrow), consistent with postnatal coronal T2WI (E, arrow) at 7 months; (F-G) Case 96, 39 weeks, fetus suspected sacrococcygeal vertebra irregularity in morphology by US showed L2 hemivertebra (G, arrow) with kyphosis (F, arrow) in SWI images. SWI, susceptibility-weighted imaging; MRI, magnetic resonance imaging; T2WI, T2-weighted imaging.

46 isolated vertebral anomalies: butterfly vertebra (n=19) (Figure 3A-3C), hemivertebra (n=15) (Figure 3D-3G), coronal clefts vertebra (n=11) (Figure 4A-4C), block vertebra (n=1) (Figure 4D-4F); 30 multiple vertebral anomalies (Figure 5A-5E); and 5 sacrococcygeal hypoplasias (Figure 6A-6E). Among these, 13 cases had coexisting anomalies: filum terminale (n=1), syringomyelia (n=1), imperforate anus (n=2), meningocele (n=1), congenital undescended scapula syndrome (n=1), caudal degeneration syndrome (n=5), segmental spinal dysgenesis (n=1), and scoliosis (n=1).

The diagnostic accuracy of US and MRI was 46.9% (38/81) and 84.0% (68/81), respectively, in the deformity group, 19.6% (9/46) and 97.8% (45/46), respectively, in the negative group, and 37.0% (47/127) and 89.0%

(113/127), respectively, in the overall population. Using MRI, diagnostic accuracy was significantly improved by 33.3% in the <28-week group and by 41.6% in the ≥ 28 -week group ($P < 0.001$) in the deformity group. The same was true for the negative group and the overall sample (both $P < 0.001$). Moreover, the diagnostic accuracy of US in the <28-week GA group was higher than in the ≥ 28 -week GA group in all groups, but the difference was not statistically significant ($P = 0.397, 0.892, 0.710, 0.312, 0.296, \text{ and } 0.775$, respectively) (Table 2). In the subgroup-specific analysis of the anomalies detected using US compared with MRI, the accuracy of the MRI diagnoses was significantly higher than those of the US diagnoses in isolated vertebral anomalies ($P < 0.001$), hemivertebra

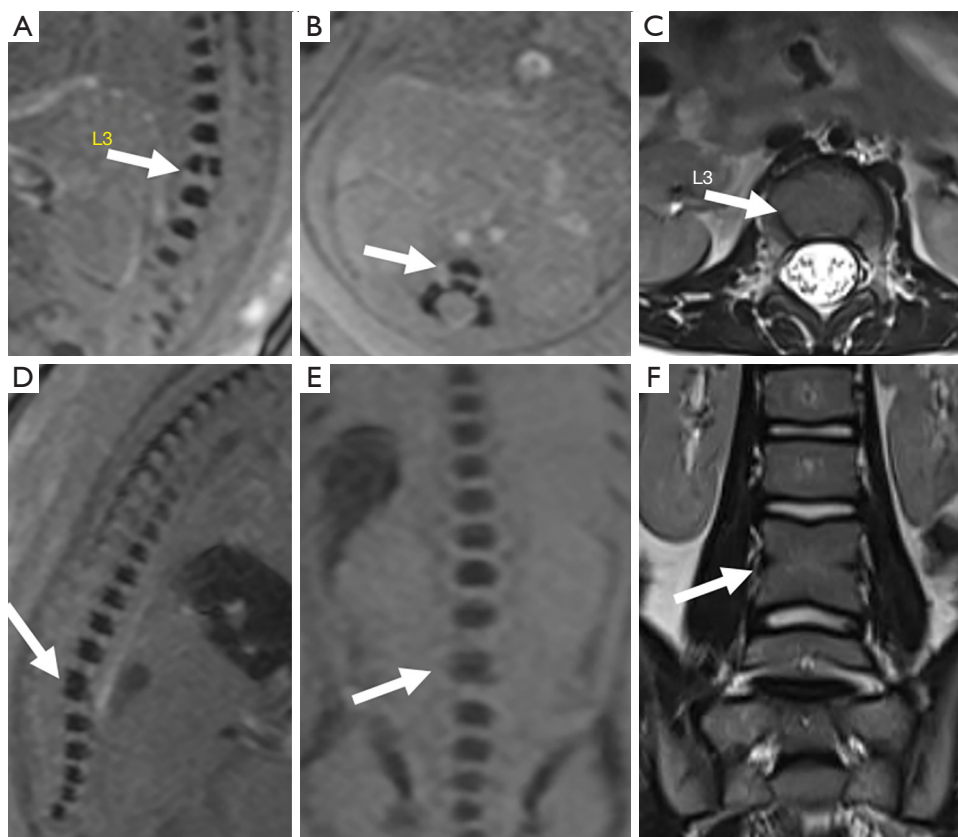


Figure 4 Fetuses with vertebral anomalies. (A-C) Case 72, L3 coronal vertebral cleft at 25 weeks +6 d, demonstrated as a hyperintensity cleft band in sagittal (A, arrow) and axial (B, arrow) SWI images. Postnatal axial T2WI (C, arrow) showed cleft disappear at 7 months; (D-E) Case 90, L3-4 block vertebra at 25 weeks +3 d in sagittal (D, arrow) and coronal (E, arrow) SWI images, corresponding to postnatal coronal T2WI finding (F, arrow) at 10 months. SWI, susceptibility-weighted imaging; T2WI, T2-weighted imaging.

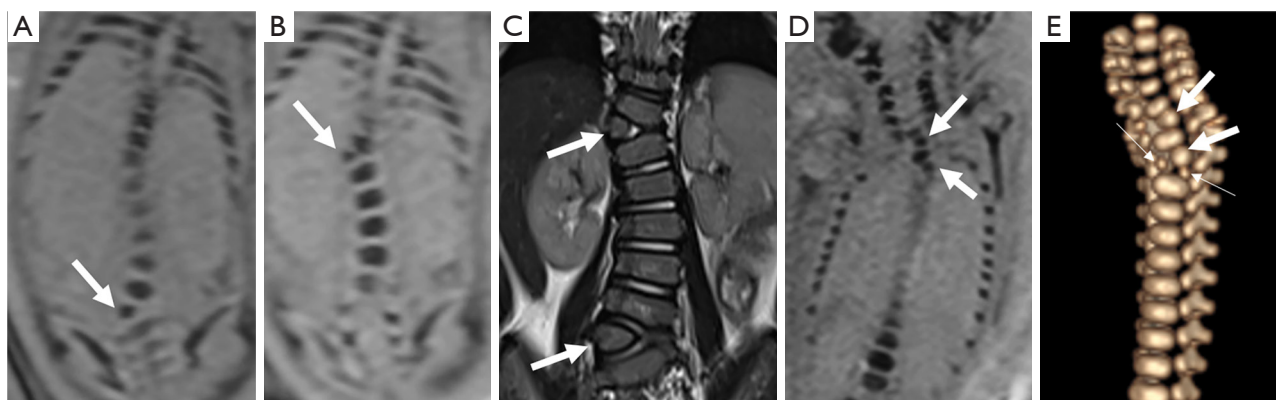


Figure 5 Multiple vertebral anomalies. (A-C) Case 106, L5, T11 hemivertebra at 27 weeks in prenatal coronal SWI images (A, B, arrows), consistent with postnatal coronal T2WI (C, arrows) at 20 months; (D-E) Case 118, C6, T1 hemivertebrae were diagnosed in prenatal coronal SWI at 25 weeks (D, E, thick arrows) and missed diagnosis of T2 butterfly vertebra, which displayed in postmortem CT (E, thin arrows). SWI, susceptibility-weighted imaging; T2WI, T2-weighted imaging; CT, computed tomography.

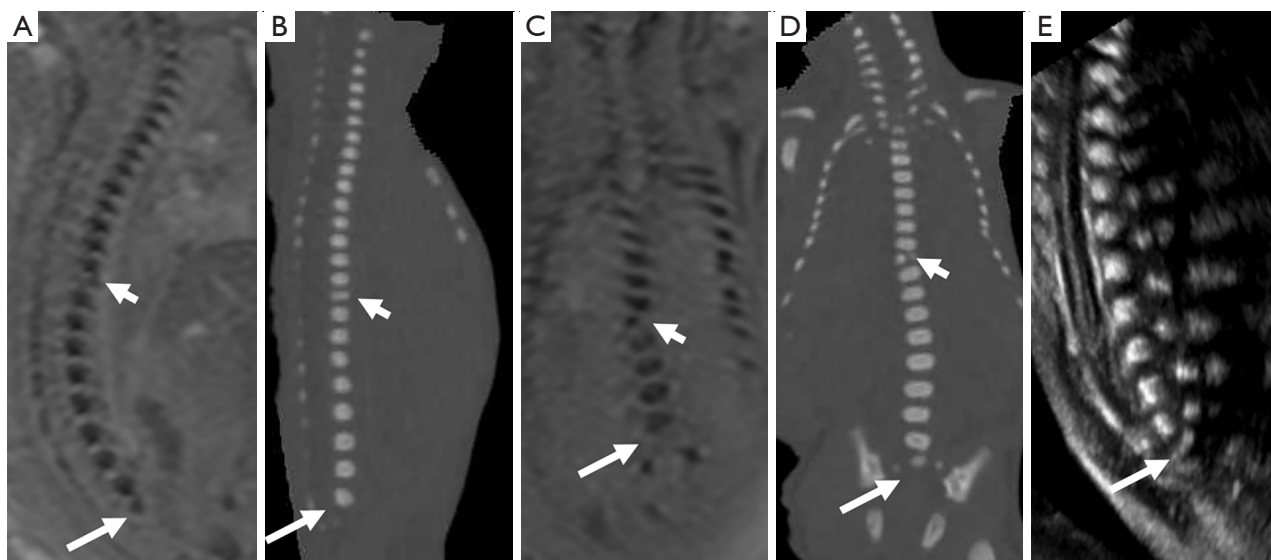


Figure 6 Sacrococcygeal hypoplasia. (A-E) Case 124, (A, C) the spine ends at S1, and the sacrum was not visible below S1 (long arrows) in sagittal and coronal SWI images at 26 w and complicated with T9 hemivertebrae (short arrows). (B, D) These findings were consistent with the postmortem CT (arrows). (E) US shows dysplasia of the sacrococcygeal vertebrae (arrow); however, the T9 hemivertebrae was missed in prenatal diagnosis. SWI, susceptibility-weighted imaging; CT, computed tomography; US, ultrasound.

Table 2 Diagnostic accuracy of ultrasound and fetal magnetic resonance imaging by gestational age in 127 cases

| Groups | Gestational age | US correct | MRI correct | Percentage difference (95% CI) | P value* (95% CI) |
|------------------------|----------------------------------|--------------------|--------------------|--------------------------------|---------------------|
| Deformity group (n=81) | Combined (n=81) | 38/81 (46.9%) | 68/81 (84.0%) | 37.1% (27%, 48%) | <0.001 (0.08, 0.35) |
| | <28 weeks (n=45) | 23/45 (51.1%) | 38/45 (84.4%) | 33.3% (20%, 47%) | <0.001 (0.07, 0.52) |
| | ≥28 weeks (n=36) | 15/36 (41.7%) | 30/36 (83.3%) | 41.6% (26%, 58%) | <0.001 (0.05, 0.43) |
| | Percentage difference (95% CI)** | 9.4% (-12%, 30%) | 1.1% (-15%, 18%) | NA | NA |
| | P value** (95% CI) | 0.397 (0.28, 1.65) | 0.892 (0.28, 3.03) | NA | NA |
| Negative group (n=46) | Combined (n=46) | 9/46 (19.6%) | 45/46 (97.8%) | 78.2% (62%, 87%) | <0.001 (0, 0.04) |
| | <28 weeks (n=23) | 5/23 (21.7%) | 22/23 (95.7%) | 74.0% (48%, 87%) | <0.001 (0, 0.12) |
| | ≥28 weeks (n=23) | 4/23 (17.4%) | 23/23 (100%) | 82.6% (58%, 93%) | <0.001 (NA) |
| | Percentage difference (95% CI)** | 4.3% (-19%, 27%) | 4.4% (-10%, 21%) | NA | NA |
| | P value** (95% CI) | 0.710 (0.18, 3.28) | 0.312 (NA) | NA | NA |
| Overall sample (n=127) | Combined (n=127) | 47/127 (37.0%) | 113 /127 (89.0%) | 52.0% (41%, 61%) | <0.001 (0.04, 0.14) |
| | <28 weeks (n=68) | 28/68 (41.2%) | 60/68 (88.2%) | 47.0% (32%, 59%) | <0.001 (0.04, 0.23) |
| | ≥28 weeks (n=59) | 19/59 (32.2%) | 53/59 (89.8%) | 57.6% (41%, 69%) | <0.001 (0.02, 0.15) |
| | Percentage difference (95% CI)** | 9.0% (-7.8%, 25%) | 1.6% (-10%, 13%) | NA | NA |
| | P value** (95% CI) | 0.296 (0.33, 1.41) | 0.775 (0.28, 2.61) | NA | NA |

*, McNemar's test between US and MRI correct diagnoses. **, Chi-square test or Fisher's exact test between <28 weeks and ≥28 weeks group. US, ultrasound; MRI, magnetic resonance imaging; 95% CI, 95% confidence interval; NA, not applicable.

($P < 0.001$), coronal clefts vertebra ($P = 0.011$), and multiple vertebral anomalies ($P = 0.002$). The detailed diagnostic accuracy data for the subgroup of fetuses with vertebral anomalies between different modalities within specific GA groups are shown in *Table 3*.

In addition, for the cases with US performed in our center (Shandong Provincial Hospital Affiliated to Shandong First Medical University) ($n = 84$) (*Table 1*), the diagnostic accuracy of MRI was significantly higher than that of US [90.5% (76/84) *vs.* 40.5% (34/84), respectively, $P < 0.001$]; and if the excluded cases ($n = 14$) were included assuming that US could have made the correct diagnosis, the diagnostic accuracy of MRI was significantly higher than that of US [80.1% (113/141) *vs.* 42.6% (60/141), respectively, $P < 0.001$].

In addition, our results showed that MRI and US were concordant and correct in 36.2% (46/127) of cases; MRI yielded findings additional to US in 16.5% (21/127) of cases [butterfly vertebra ($n = 3$), hemivertebra ($n = 7$), and multiple vertebral anomalies ($n = 11$)]; MRI was correct when US failed in 36.2% (46/127) of cases; US was correct and MRI failed in 0.8% (1/127) of cases; both MRI and US were not consistent with postnatal findings in 10.2% (13/127) of cases.

Agreements between US and MRI in the deformity group and the negative group were 45.7% and 19.6%, respectively. A detailed description of agreement and disagreement rates between prenatal US and MRI in different groups is provided in *Table 4*.

After MRI, 116 participants carried pregnancies to term, and 11 participants had a termination of pregnancy. It is worth mentioning that MRI corrected the US diagnosis in 36.2% (46/127) of these cases, including 78.3% (36/46) of the cases where the MRI did not find an abnormality and was justified as correct based on later follow up. In the cases of pregnancy termination, 6 had coexisting anomalies, and 5 had multiple vertebral anomalies.

Of the 10.2% (13/127) cases with incorrect diagnosis using MRI and US, there were 3 cases of butterfly vertebra misdiagnosed as hemivertebra anomalies, 9 cases of multiple vertebral anomalies that were missed or misdiagnosed on MRI, and 1 case (case 46) of a normal vertebra misdiagnosed as a hemivertebra anomaly (*Table 5*). A full summary of all fetal US and MRI imaging findings and follow-up diagnoses is shown in *Table S1*.

Diagnostic confidence

Figure 7 presents the proportions of correct and incorrect diagnoses made with high and low diagnostic confidence.

High-confidence diagnoses were made in 73.2% (93/127) of cases using US compared with 95.3% (121/127) of cases using MRI, an absolute difference of 22%. High-confidence US diagnoses were subsequently found to be incorrect in 44.9% (57/127) of patients compared with 9.4% (12/127) using MRI. The MRI yielded fewer low-confidence diagnoses than US (4.7% *vs.* 26.8%), of which 3.1% (4/127) were found to be correct on MRI, while 8.7% (11/127) were correct on US compared with follow-up results.

Discussion

In this study, we assessed the diagnostic accuracy and confidence of fetal MRI compared to US in 127 fetuses with a range of fetal vertebral anomalies. Several previous studies had used both MRI and US to image the fetal spine (5,38-40), but the sample sizes were small, and the study populations did not focus on fetal spine vertebral malformations. To the best of our knowledge, this is the largest study to date exploring the use of MRI and specifically targeting fetuses with vertebral anomalies. Based on postnatal and postmortem imaging findings, adding fetal MRI to the diagnostic pathway increased diagnostic accuracy to 84.4% for fetuses < 28 weeks GA ($P < 0.001$) and to 83.3% for fetuses ≥ 28 weeks GA ($P < 0.001$). Moreover, MRI provided additional information in 16.5% of the cases. These results indicated that MRI significantly increased the diagnostic accuracy of fetal vertebral pathologies compared to US alone.

Although previous studies assessed fetal skeletal diagnoses and spinal lesions using US or MRI, the diagnostic accuracy ranged from 40.9% to 67.9% (5,6) (7,39,41), and the included populations focused on long bone or spinal canal disease. In our study, we included 127 participants with inconclusive or uncertain fetal vertebral finding on US, where MRI achieved a better diagnostic performance. The higher diagnostic accuracy of MRI (84.0%) compared with US (46.9%) can be explained by the SWI sequence yielding a high contrast between bone and soft tissues but a low contrast between soft tissues (30). Moreover, the higher absolute difference in diagnostic accuracy between MRI and US for the older GA group (41.6%) compared with the younger GA group (33.3%) can be attributed to advancing GA, fetal vertebral body increases in volume, which are increasingly distinguishable from the surrounding soft tissues in SWI images, and amniotic fluid reductions as a result of GA (42) interfering with fetal movement. This increase in vertebral volume and decrease in amniotic fluid

Table 3 Diagnostic accuracy in descriptive categorization of findings between ultrasound and fetal magnetic resonance imaging and different gestational ages

| Postnatal/ postmortem final diagnosis | US | | <28 weeks | | ≥28 weeks | | P* | | Percentage difference* | | P* | | Percentage difference** | | P*** | |
|---------------------------------------------|---------------|---------------|---------------|---------------|------------------|---------------|--------------------|---------------------------------|------------------------|----------------------------------|--------------------|-----------------------------------|-------------------------|-------------------|--------------------|--|
| | correct | MRI correct | US correct | MRI correct | US correct | MRI correct | P* (95% CI) | Percentage difference* (95% CI) | P* (95% CI) | Percentage difference** (95% CI) | P* (95% CI) | Percentage difference*** (95% CI) | P*** (95% CI) | | | |
| Isolated vertebral anomalies (n=46) | 25/46 (54.3%) | 42/46 (91.3%) | 17/28 (60.7%) | 26/28 (92.9%) | 8/18 (44.4%) | 16/18 (88.9%) | 0.004 (0.02, 0.60) | 32.1% (10%, 51%) | 0.004 (0.02, 0.60) | 44.4% (14%, 66%) | 0.005 (0.02, 0.57) | 16.3% (-12%, 42%) | 0.280 (0.16, 1.72) | 4.0% (-13%, 26%) | 0.641 (0.08, 4.81) | |
| Butterfly vertebrae (n=19) | 12/19 (63.2%) | 15/19 (78.9%) | 10/12 (83.3%) | 10/12 (83.3%) | NA | 2/7 (28.6%) | 0.283 (0.11, 1.94) | NA | NA | 42.9% (-7%, 72%) | 0.109 (0.02, 1.63) | 54.8% (9%, 78%) | 0.017 (0.01, 0.75) | 11.9% (-23%, 49%) | 0.539 (0.05, 4.67) | |
| Hemivertebrae (n=15) | 7/15 (46.7%) | 15/15 (100%) | 5/9 (55.6%) | 9/9 (100%) | 44.4% (5%, 73%) | 2/6 (33.3%) | <0.001 (NA) | 53.3% (22%, 75%) | 0.023 (NA) | 66.7% (13%, 90%) | 0.014 (NA) | 22.2% (-24%, 57%) | 0.398 (0.05, 3.42) | NA | NA | |
| Coronal Clefts vertebra (n=11) | 6/11 (54.5%) | 11/11 (100%) | 2/6 (33.3%) | 6/6 (100%) | 66.7% (13%, 90%) | 4/5 (80.0%) | 0.011 (NA) | 45.5% (10%, 72%) | 0.014 (NA) | 20.0% (-26%, 62%) | 0.292 (NA) | 46.7% (-9%, 75%) | 0.122 (0.01, 2.0) | NA | NA | |
| Block Vertebral (n=1) | 0/1 (0%) | 1/1 (100%) | 0/1 (0%) | 1/1 (100%) | NA | 0/0 (0%) | 0.157 (NA) | NA | 0.157 (NA) | NA | NA | NA | NA | NA | NA | |
| Multiple vertebral anomalies (n=30) | 9/30 (30.0%) | 21/30 (70.0%) | 3/14 (21.4%) | 9/14 (64.3%) | 42.9% (6%, 67%) | 6/16 (37.5%) | 0.002 (0.06, 0.55) | 40.0% (15%, 59%) | 0.022 (0.03, 0.81) | 37.5% (3%, 62%) | 0.033 (0.04, 0.91) | 16.1% (-16%, 44%) | 0.338 (0.09, 2.32) | 10.7% (-21%, 40%) | 0.523 (0.12, 2.89) | |
| Sacroccygeal hypoplasia (n=5) | 4/5 (80.0%) | 5/5 (100%) | 3/3 (100%) | 3/3 (100%) | NA | 1/2 (50.0%) | 0.292 (NA) | 20.0% (-26%, 62%) | NA | 50.0% (-27%, 91%) | 0.248 (NA) | 50.0% (-19%, 91%) | 0.171 (NA) | NA | NA | |

*, comparisons between US and MRI; **, comparisons between <28 and ≥28 weeks in US; ***, comparisons between <28 and ≥28 weeks in MRI. US, ultrasound; MRI, magnetic resonance imaging; 95% CI, 95% confidence interval; NA, not applicable.

Table 4 Rate of agreement and disagreement between prenatal ultrasound and magnetic resonance imaging

| MRI vs. US | Deformity group (n=81) | Negative group (n=46) | Overall sample (n=127) |
|-------------------------------------------------------------|------------------------|-----------------------|------------------------|
| MRI and US were concordant and correct | 37 (45.7%) | 9 (19.6%) | 46 (36.2%) |
| MRI showed additional findings to US | 21 (25.9%) | 0 | 21 (16.5%) |
| MRI correct, US failed | 10 (12.3%) | 36 (78.3%) | 46 (36.2%) |
| US correct, MRI failed | 1 (1.2%) | 0 | 1 (0.8%) |
| Both MRI and US were not consistent with postnatal findings | 12 (14.8%) | 1 (2.2%) | 13 (10.2%) |

Data are n (%). US, ultrasound; MRI, magnetic resonance imaging.

Table 5 Cases misdiagnosed and missed diagnoses using magnetic resonance imaging

| Case number | Age (years) | Gestational age (weeks + days) | US findings | MRI findings | Outcomes/follow-up |
|-------------|-------------|--------------------------------|----------------------------------------------------------------------|-----------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------|
| 46 | 37 | 25 | T11 hemivertebra | T11 hemivertebra | (-) |
| 51 | 23 | 27 | T10 hemivertebra | T10 hemivertebra | T10 butterfly vertebra |
| 54 | 38 | 29 | T5 hemivertebra | T5 hemivertebra | T5 butterfly vertebra |
| 56 | 25 | 31 | T11 hemivertebra | T11 hemivertebra | T11 butterfly vertebra |
| 66 | 23 | 32 | T1 irregularity in morphology | T1 hemivertebra | T1 hemivertebra; T3 butterfly vertebra |
| 67 | 31 | 27 | T3, T5, T7, T10 arranged irregularly | T7 butterfly vertebra T10 hemivertebra | T2, T3, T5, T6, T7, T10 butterfly vertebra |
| 110 | 36 | 24 | T4, T9 hemivertebra T8 butterfly vertebra L1-2 block vertebrae | T9 butterfly vertebra T4 hemivertebrae L1-2 block vertebrae | T8, T9 butterfly vertebra T4, L2 hemivertebrae |
| 113 | 29 | 21+6 | Multiple vertebral deformities; hemivertebra? | T5, T11 hemivertebra T10 butterfly vertebra T8-9, L3-4 block vertebra | T3, T5, T7, T11 hemivertebra T10 butterfly vertebra T8-9, L3-4 block vertebra |
| 114 | 26 | 29 | T6, T7 hemivertebrae? | T6, T7, T9 hemivertebra and/or butterfly vertebra | T5, T7, T8 butterfly vertebra T6 hemivertebra |
| 115 | 30 | 24 | T11, T12 arranged irregularly | T11-12 block vertebra | T3 hemivertebra T4, T8, T9 butterfly vertebra T11-12 block vertebra |
| 116 | 26 | 31 | Cervical vertebral arranged irregularly | Cervical vertebral body and appendix Arranged irregularly | The congenital undescended scapula syndrome (sprengel deformity) Cervical closed spina bifida |
| 118 | 28 | 25 | Multiple vertebral deformities with scoliosis in upper thoracic | C6, T1 hemivertebra | C6, T1 hemivertebra T2 butterfly vertebra |
| 120 | 23 | 29 | Vertebrae arranged irregularly in lower cervical and upper thoracic | T6-8 block vertebra, and/or hemivertebra | T3 butterfly vertebra T5-6, T11-12, L4-5 block vertebra |

US, ultrasound; MRI, magnetic resonance imaging.

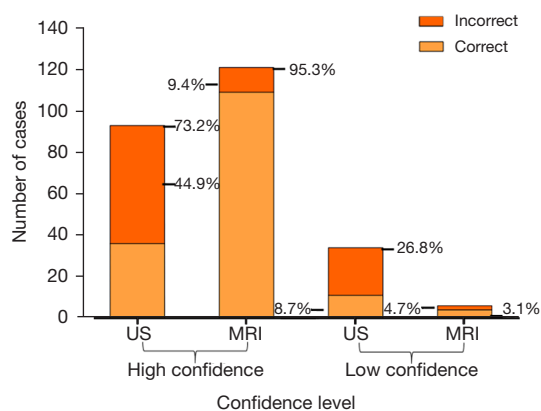


Figure 7 High (rated as 70% or 90%) and low (rated as 10%, 30%, or 50%) confidence diagnoses made using US and MRI in 127 patients in comparison with the follow-up results. US, ultrasound; MRI, magnetic resonance imaging.

contribute to better MRI definitions on image quality but weaken the accuracy of US.

In fetal US examinations, operational bias caused by ongoing ossification of the fetal skull, increasing physical size of the woman, decreasing amniotic fluid, and descent of the fetal head into the maternal pelvis with GA may decrease the diagnostic accuracy of US, especially in the ≥ 28 -week group in certain subgroups. In addition, acoustic shadowing in US from the overlying iliac wing, the scapula, or bones of the fetal arm make it difficult for US views, especially standard coronal views, to demonstrate or there are artifacts, and as such, the fetal sacrum, cervical, and upper thoracic spine may be difficult to view on coronal view (32). However, MRI images can be obtained in any orientation with large FOV and offset the shortcomings of US. The advent of 3D-US has made evaluation of the fetal spine more comprehensive. It allows for evaluation of the complete anatomy of the spine, which is not always possible in 2D imaging due to the curvature of the spine (43). However, given this was a retrospective analysis, 2D-US or 3D-US were alternately used depending on clinical requirements.

Use of MRI provided additional information relative to US in 16.5% of cases, mainly in cases of butterfly vertebra and hemivertebra, which in US are better imaged using coronal views rather than routine sagittal views, as discussed before. Another important finding of our work was that in 36.2% of cases, the US diagnosis was revised by MRI such that management and parental counseling changed

completely. Most of these revisions occurred in the negative group (36/46). Moreover, the negative group showed a wide disagreement diagnosis between MRI and US. It is important to understand that this may in part be explained by referral bias, because the participants included in our study were suspected of fetal vertebral anomalies using US instead of negative US.

Of the 81 cases with vertebral anomalies, 16.0% were incorrectly diagnosed by MRI, primarily in cases of butterfly vertebra and multiple vertebral anomalies. Butterfly vertebra results from 2 lateral centers of chondrification failing to fuse at the midline (44). When the bilateral ossification center is asymmetric, the condition can be misdiagnosed (case 56) or result in a missed diagnosis (case 118) by MRI. Moreover, in our study there were 4 cases of partially segmented hemivertebra (cases 91 to 94) diagnosed incorrectly with US but revised with MRI, which was of great importance to prenatal counseling and postnatal management. Hemivertebra could be classified into unsegmented hemivertebra, partially segmented hemivertebra, and fully segmented hemivertebra, on the basis of presence or absence of fusion to the vertebral bodies above and/or below (45). Partially segmented hemivertebra are less likely to cause curvature of the spine with good prognosis during postnatal growth (45) compared with fully segmented hemivertebra (cases 95 to 107). In addition, there were 11 cases of isolated coronal clefts and 11 cases of multiple coronal clefts in our study (cases 68 to 89). The clefts predominantly occurred in the lumbar region (16/22, 73%) and all of these clefts disappeared after birth, consistent with a previous study (46). Hence, coronal clefts in fetuses should not be interpreted as vertebra malformation, but as a physiological variation of vertebral body development. In addition, some diseases with low incidence that involve fetal vertebra, such as osteochondrodysplasias (47) and hypophosphatasia (48), were not encountered in our study.

Several studies have demonstrated the importance of diagnostic confidence in assessing an imaging technology (37,49). Ng and Palmer (50) explained the relevance of diagnostic accuracy and diagnostic confidence. In our study, the proportion of high-confidence diagnoses on MRI increased by 22% compared with US. An incorrect diagnosis made with high confidence can result in an inappropriate change in management, including termination of pregnancy. The MRI resulted in fewer high-confidence, incorrect diagnoses compared with US (9.4% *vs.* 44.9%, respectively) and resulted in fewer low-confidence diagnoses

(4.7% vs. 26.8%, respectively).

This study had several limitations. Firstly, the SWI sequence in our study was sensitive to motion artifacts, requiring a breath-hold scan. Often several scans were needed to cover the required volume, particularly for younger GA fetuses; secondly, the retrospective methodology may have led to an underestimation of the diagnostic power of US; thirdly, fetal spinal canal and cord pathologies were not included in our study; and lastly, there was a lack of reported sensitivity or specificity of MRI and US in patients, because cases were derived from abnormal US findings. Future studies should be multicenter, prospective, and randomized controlled design, including spinal canal cases.

Conclusions

Fetal vertebral MRI improves the accuracy and confidence of diagnosing fetal vertebral anomalies. Such improvements are likely to result in changes to counseling and clinical management of spinal anomalies.

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Footnote

Reporting Checklist: The authors have completed the STARD reporting checklist. Available at <https://qims.amegroups.com/article/view/10.21037/qims-21-1070/rc>

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. This retrospective study was conducted in accordance with the Declaration of Helsinki (as revised in 2013) and approved by the

Institutional Review Board of Shandong Provincial Hospital Affiliated to Shandong First Medical University. All patients gave written informed consent before participating in this study.

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Supplementary

Table S1 A full summary of all fetal diagnoses, imaging findings on US and MRI and postnatal diagnosis

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|--------------------------------------------------------------------------------|---------------|---------------------|
| 1 | 24y | 25w | the lumbosacral part slightly curved; hemivertebrae? | (-) | (-) |
| 2 | 29y | 26w | T1, T3 vertebra morphological irregularity | (-) | (-) |
| 3 | 29y | 21w | MRI examination recommended to exclude thoracic abnormalities | (-) | (-) |
| 4 | 29y | 24w | the thoracic vertebra small in size | (-) | (-) |
| 5 | 30y | 25w | MRI examination recommended to exclude middle and lower thoracic abnormalities | (-) | (-) |
| 6 | 34y | 25w+3d | T11, T12 and L1 vertebral bodies slightly smaller | (-) | (-) |
| 7 | 39y | 26w | MRI examination recommended to exclude thoracic abnormalities | (-) | (-) |
| 8 | 26y | 26w+1d | T1, T2 vertebrae arranged irregularly | (-) | (-) |
| 9 | 34y | 26w+3d | T2, T3 vertebrae slightly out of order | (-) | (-) |
| 10 | 32y | 27w | normal, MRI examination recommended | (-) | (-) |
| 11 | 29y | 29w | T2 slightly larger in size | (-) | (-) |
| 12 | 26y | 30w | normal, MRI examination recommended | (-) | (-) |
| 13 | 27y | 30w+5d | the upper thoracic vertebrae arranged irregularly | (-) | (-) |
| 14 | 33y | 36w | normal, MRI examination recommended | (-) | (-) |
| 15 | 24y | 38w | the thoracic block vertebrae | (-) | (-) |
| 16 | 21y | 38w+3d | the sacrococcygeal abnormalities | (-) | (-) |
| 17 | 31y | 28w+1d | the thoracic vertebrae arranged irregularly | (-) | (-) |
| 18 | 35y | 26w | T2, T3 vertebrae body small in size | (-) | (-) |
| 19 | 32y | 28w | T8, T10 irregularity in morphology | (-) | (-) |
| 20 | 29y | 22w+5d | normal, MRI examination recommended | (-) | (-) |
| 21 | 28y | 29w | T12 arranged irregularly | (-) | (-) |
| 22 | 30y | 24w+5d | T8, T9, T10, L3, L4, L5 displayed unsatisfactory | (-) | (-) |
| 23 | 27y | 35w | T8, T9 slightly smaller | (-) | (-) |
| 24 | 30y | 24w | echo of the L1/2 vertebral arch enhanced | (-) | (-) |
| 25 | 29y | 26w+6d | normal, MRI examination recommended | (-) | (-) |
| 26 | 31y | 29w+5d | L2, L3, L4 irregularity in morphology | (-) | (-) |
| 27 | 27y | 36w | normal, MRI examination recommended | (-) | (-) |
| 28 | 31y | 27w+5d | C7 transverse process too long | (-) | (-) |
| 29 | 27y | 31w | normal; MRI examination recommended | (-) | (-) |
| 30 | 38y | 35w | the sacrococcygeal abnormalities | (-) | (-) |
| 31 | 26y | 25w | T5, T8 vertebrae body small in size | (-) | (-) |

Table S1 (continued)

Table S1 (continued)

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|---------------------------------------------------------|------------------------|------------------------|
| 32 | 34y | 26w | parts of vertebrae arranged irregularly | (-) | (-) |
| 33 | 31y | 34w | T4 irregularity in morphology | (-) | (-) |
| 34 | 32y | 28w+2d | C7, T3, T4 coronal clefts vertebrae? | (-) | (-) |
| 35 | 26y | 24w | L5 coronal cleft vertebra | (-) | (-) |
| 36 | 26y | 24w | normal, MRI examination recommended | (-) | (-) |
| 37 | 33y | 29w+6d | L2 irregularity in morphology | (-) | (-) |
| 38 | 32y | 28w | T4, T5 arranged irregularly | (-) | (-) |
| 39 | 28y | 31w | T10 vertebrae body small in size | (-) | (-) |
| 40 | 31y | 30w | the sacrococcygeal vertebrae arranged irregularly | (-) | (-) |
| 41 | 33y | 27w+1d | the sacrococcygeal vertebrae arranged irregularly | (-) | (-) |
| 42 | 25y | 32w+2d | the lumbosacral vertebrae arranged irregularly | (-) | (-) |
| 43 | 37y | 34w+6d | the upper thoracic vertebrae irregularity in morphology | (-) | (-) |
| 44 | 32y | 27w | thoracic hemivertebra? | (-) | (-) |
| 45 | 23y | 34w | the sacrococcygeal vertebrae poorly showed | (-) | (-) |
| 46 | 37y | 25w | T11 hemivertebra? | T11 hemivertebra? | (-) |
| 47 | 39y | 27w | T12 butterfly vertebra | T12 hemivertebra? | T11 butterfly vertebra |
| 48 | 22y | 24w | T11 butterfly vertebra | T11 butterfly vertebra | T11 butterfly vertebra |
| 49 | 26y | 24w | T11 butterfly vertebra | T11 butterfly vertebra | T11 butterfly vertebra |
| 50 | 30y | 25w | L5 butterfly vertebra? | L5 butterfly vertebra | L5 butterfly vertebra |
| 51 | 23y | 27w | T10 hemivertebra | T10 hemivertebra | T10 butterfly vertebra |
| 52 | 31y | 27w | T8 butterfly vertebra | T8 butterfly vertebra | T8 butterfly vertebra |
| 53 | 32y | 23w | L4 butterfly vertebra | L4 butterfly vertebra | L4 butterfly vertebra |
| 54 | 38y | 29w | T5 hemivertebra | T5 hemivertebra | T5 butterfly vertebra |
| 55 | 37y | 25w+5d | T10 butterfly vertebra? | T10 butterfly vertebra | T10 butterfly vertebra |
| 56 | 25y | 31w | T11 hemivertebra | T11 hemivertebra | T11 butterfly vertebra |
| 57 | 34y | 29w+4d | C5/6, C6/7 intervertebral space narrowing | T4 butterfly vertebra | T4 butterfly vertebra |
| 58 | 22y | 35w+2d | sacrococcygeal vertebra arranged irregularly | T3 butterfly vertebra | T3 butterfly vertebra |
| 59 | 37y | 26w | T10 butterfly vertebra | T10 butterfly vertebra | T10 butterfly vertebra |
| 60 | 25y | 28w | T11 hemivertebra? | T11 butterfly vertebra | T11 butterfly vertebra |
| 61 | 34y | 33w | T10 butterfly vertebra? | T10 butterfly vertebra | T10 butterfly vertebra |
| 62 | 33y | 26w | T9 butterfly vertebra | T9 butterfly vertebra | T9 butterfly vertebra |
| 63 | 33y | 24w | L3 butterfly vertebra? | L3 butterfly vertebra | L3 butterfly vertebra |
| 64 | 33y | 36w+3d | T11 butterfly vertebra | T11 butterfly vertebra | T11 butterfly vertebra |

Table S1 (continued)

Table S1 (continued)

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|-----------------------------------------|--------------------------------------------|------------------------------------------------|
| 65 | 29y | 27w | T8 hemivertebra | T8 butterfly vertebra | T8 butterfly vertebra |
| 66 | 23y | 32w | T1 irregularity in morphology | T1 hemivertebra | T1 hemivertebra; T3 butterfly vertebra |
| 67 | 31y | 27w | T3, T5, T7, T10 arranged irregularly | T7 butterfly vertebra; T10 hemivertebra | T2, T3, T5, T6, T7, T10 butterfly vertebra |
| 68 | 34y | 25w | L4 butterfly vertebra? | L4 coronal vertebral cleft | disappear |
| 69 | 30y | 29w+4d | L3, L5 coronal vertebral clefts | L3, L5 coronal vertebral clefts | disappear, L2 level fatty filum terminale |
| 70 | 30y | 30w | L2, L3 coronal vertebral clefts | L2, L3 coronal vertebral clefts | disappear; T9-13 vertebral level syringomyelia |
| 71 | 32y | 33w | L2-L5 coronal vertebral clefts | L2-L5 coronal vertebral clefts | disappear |
| 72 | 32y | 25w+6d | L2 butterfly vertebra? | L2 coronal vertebral cleft | disappear |
| 73 | 22y | 26w | L3 butterfly vertebra? | L3 coronal vertebral cleft | disappear |
| 74 | 32y | 34w | L3 coronal vertebral cleft | L3 coronal vertebral cleft | disappear |
| 75 | 34y | 29w | T3-T6, L1-L3 irregularity in morphology | T4-6, L1-3 coronal vertebral clefts | disappear |
| 76 | 35y | 28w+4d | T12-L4 irregularity in morphology | T12-L1, L3-L5 coronal vertebral clefts | disappear |
| 77 | 26y | 28w | L2, L4 arranged irregularly | L2, L4, L5 coronal vertebral clefts | disappear |
| 78 | 32y | 28w | T4, T8, L1 coronal vertebral clefts | T4, T8, L1 coronal vertebral clefts | disappear |
| 79 | 31y | 27w+4d | L3 coronal vertebral cleft | L3 coronal vertebral cleft | disappear |
| 80 | 30y | 30w+2d | normal, MRI examination recommended | L3 coronal vertebral cleft | disappear |
| 81 | 26y | 28w | T10 irregularity in morphology | T10 coronal vertebral cleft | disappear |
| 82 | 29y | 26w | T1-T5, L2-L4 irregularity in morphology | L2 coronal vertebral cleft | disappear |
| 83 | 29y | 34w | T8 coronal vertebral cleft | T8 coronal vertebral cleft | disappear |
| 84 | 24y | 29w | T5-T7, L1-L5 irregularity in morphology | L2, L4 coronal vertebral clefts | disappear |
| 85 | 37y | 30w | L2-L5 irregularity in morphology | L2-5 coronal vertebral clefts | disappear |
| 86 | 36y | 26w | L3 irregularity in morphology | L3 coronal vertebral cleft | disappear |
| 87 | 22y | 27w+ 2d | L1 butterfly vertebra | L1, L3-5 coronal vertebral clefts | disappear |
| 88 | 30y | 34w | T8 vertebra small in size | T8 coronal vertebral cleft | disappear |
| 89 | 29y | 25w | T7, L2, L4 ossification center abnormal | L2, L3, L5 coronal vertebral cleft | disappear |

Table S1 (continued)

Table S1 (continued)

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|----------------------------------------------------------------------|--------------------------------------------------------------------|----------------------------------------------------------------------------------------------------|
| 90 | 28y | 25w+3d | lumbar hemivertebra | L3/4 intervertebral space narrowing | L3-4 block vertebra |
| 91 | 44y | 24w | L4 irregularity in morphology | L4 hemivertebra with fusing to one side of L3 | L4 hemivertebra with fusing to one side of L3 |
| 92 | 34y | 27w | T12 irregularity in morphology | T10 hemivertebra with fusing to one side of T9 | T10 hemivertebra with fusing to one side of T9 |
| 93 | 30y | 24w | L2 hemivertebra? | L2 hemivertebra with fusing to one side of L3 | L2 hemivertebra with fusing to one side of L3 |
| 94 | 26y | 24w | L3 vertebra small in size | L2 hemivertebra with fusing to one side of L3 | L2 hemivertebra with fusing to one side of L3 |
| 95 | 27y | 26w | cervical vertebra hemivertebra | cervical hemivertebra | cervical hemivertebra |
| 96 | 34y | 39w | caudal vertebra irregularity in morphology | L2 hemivertebra | L2 hemivertebra |
| 97 | 27y | 31w+1d | thoracic block vertebra | T7 hemivertebra | T7 hemivertebra |
| 98 | 32y | 28w | scoliosis | T11 hemivertebra | T10 hemivertebra |
| 99 | 29y | 30w+3d | S1 hemivertebra? | S1 hemivertebra | S1 hemivertebra, imperforate anus |
| 100 | 29y | 27w | L1 hemivertebra | L1 hemivertebra | L1 hemivertebra |
| 101 | 31y | 33w | sacrococcygeal vertebra not seen | L5 hemivertebra | L5 hemivertebra |
| 102 | 35y | 34w | T12 hemivertebra | T12 hemivertebra | T12 hemivertebra |
| 103 | 30y | 26w | L2 hemivertebra | L2 hemivertebra | L2 hemivertebra |
| 104 | 31y | 27w | L3 hemivertebra | L3 hemivertebra | L3 hemivertebra |
| 105 | 26y | 26w | L4 hemivertebra | L4 hemivertebra | L4 hemivertebra |
| 106 | 20y | 27w | vertebra irregularity in morphology | T11, L5 hemivertebra | T11, L5 hemivertebra |
| 107 | 31y | 26w | L2 hemivertebra | T7, L2 hemivertebra | T7, L2 hemivertebra |
| 108 | 29y | 21w+6d | thoracic hemivertebra | multiple hemivertebra in thoracolumbar vertebra | T3, T5, T7, T11 hemivertebrae; T8 hemivertebra with fusing to one side of T9; L2, L3 large in size |
| 109 | 29y | 34w | T5 hemivertebra | T5 butterfly vertebra T9 hemivertebra | T5 butterfly vertebra T9 hemivertebrae |
| 110 | 36y | 24w | T4, T9 hemivertebra T8 butterfly vertebra L1-2 block vertebrae | T9 butterfly vertebra; T4 hemivertebra; L1-2 block vertebrae | T8, T9 butterfly vertebrae T4 hemivertebra L2-3 block vertebrae |
| 111 | 28y | 28w+5d | T12-L1 block vertebrae T9 butterfly vertebra | T12-L1 block vertebrae T9 butterfly vertebra | L1-L2 block vertebrae; T10 butterfly vertebra; imperforate anus |

Table S1 (continued)

Table S1 (continued)

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|---------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------|
| 112 | 31y | 24w | spina bifida with meningocele; multiple vertebral deformities | spina bifida with meningocele; tethered cord; multiple hemivertebra, butterfly vertebra in thoracolumbar vertebra; multiple intercostal space narrowing on one side | bifida with meningocele; T4 butterfly vertebra; T5-6, T11-12 block vertebra; T7-9, L3-5 hemivertebra; T7-8 rib fused |
| 113 | 29y | 21w+6d | multiple vertebral deformities; hemivertebra? | T5, T11 hemivertebra T10 butterfly vertebra T8-9, L3-4 block vertebra | T3, T5, T7, T11 hemivertebrae; T10 butterfly vertebra; T8-9, L3-4 block vertebrae; |
| 114 | 26y | 29w | T6, T7 hemivertebrae? | T6, T7, T9 hemivertebrae and/or butterfly vertebra | T5, T7, T8 butterfly vertebrae; T6 hemivertebra |
| 115 | 30y | 24w | T11, T12 arranged irregularly | T11-12 block vertebra | T3 hemivertebra; T4, T8, T9 butterfly vertebrae; T11-12 block vertebra |
| 116 | 26y | 31w | cervical vertebral arranged irregularly | cervical vertebral body and appendix arranged irregularly | the congenital undescended scapula syndrome (sprengel deformity); cervical closed spina bifida |
| 117 | 28y | 23w+2d | multiple vertebral deformities with scoliosis | T6, T9 butterfly vertebra; T8 hemivertebra | T6, T9 butterfly vertebra; T8 hemivertebra |
| 118 | 28y | 25w | multiple vertebral deformities with scoliosis in upper thoracic | C6, T1 hemivertebra | C6, T1 hemivertebra; T2 butterfly vertebra |
| 119 | 29y | 22w | multiple vertebral deformities with scoliosis | multiple vertebral deformities | C6, C7, T3, T4 hemivertebra T1-2 block vertebra; T5 butterfly vertebra |
| 120 | 23y | 29w | vertebrae arranged irregularly in lower cervical and upper thoracic | T6-8 block vertebrae and/or hemivertebrae | T3 butterfly vertebra; T5-6, T11-12, L4-5 block vertebrae |
| 121 | 27y | 31w | the sacrococcygeal vertebrae poorly showed | anterior sacral meningocele (ASM); sacrococcygeal vertebra dysplasia; | anterior sacral meningocele (ASM): sacrococcygeal vertebra dysplasia; Caudal degeneration syndrome |
| 122 | 32y | 29w | low position of the conus medullaris | sacral vertebra dysplasia tethered cord meningocele | sacral agenesis tethered cord meningocele; Caudal degeneration syndrome |
| 123 | 34y | 25w+6d | the sacrococcygeal vertebra poorly showed; anus imperforate | sacral vertebra below S2 not visible | sacral agenesis: sacral vertebra below S2 not visible; anus imperforate; Caudal degeneration syndrome |

Table S1 (continued)

Table S1 (*continued*)

| Case number | Age (years) | Gestational age (weeks + days) | US diagnosis before referral for MRI | MRI diagnosis | Outcomes /follow-up |
|-------------|-------------|--------------------------------|-----------------------------------------------------------------------|-------------------------------------------------------|--------------------------------------------------------------------------------------|
| 124 | 31y | 26w | the sacrococcygeal vertebrae showed poorly with thoracic hemivertebra | T9 hemivertebra sacral vertebrae below S1 not visible | T9 hemivertebra sacral vertebrae below S1 not visible; Caudal degeneration syndrome |
| 125 | 26y | 27w | the sacrococcygeal vertebra poorly showed | sacral vertebral below S1 not visible | sacral agenesis: sacral vertebrae below S1 not visible; Caudal degeneration syndrome |
| 126 | 21y | 30w | cervicothoracic vertebrae arranged irregularly | cervicothoracic vertebrae segmental spinal dysgenesis | cervicothoracic vertebrae segmental spinal dysgenesis |
| 127 | 27y | | scoliosis | scoliosis | scoliosis |

US, ultrasound; MRI, magnetic resonance imaging.