Reference Values of Fetal Peak Systolic Velocity in the Middle Cerebral Artery at 19–40 Weeks of Gestation in Nepalese Population

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Aims: The purpose of this study was to construct new reference range for fetal middle cerebral artery peak systolic velocity (MCA-PSV) in uncomplicated pregnancy at 19-40 weeks of gestation.

Methods: This was a prospective cross-sectional study involving 400 singleton pregnancies between 19 and 40 weeks of gestation without any known risk factors of adverse pregnancy outcome who were referred for routine obstetric examination. The protocol included the doppler examination of fetal middle cerebral artery (MCA) within 2 mm after its origin from the internal carotid artery and data were used to construct the normograms and percentile fitted curves of each doppler parameter for different gestational age.

Results: Among 400 singleton uncomplicated pregnancies between 19 and 40 weeks of gestation maximum number of pregnancies (10%) was at 19 weeks of gestation and minimum (2.5%) was at 31 weeks. The fetal peak systolic blood flow in the MCA showed significant correlation with period of gestation. Mean MCA-PSV was 22.35 ± 3.05 at 19 weeks of gestation which increased to 67.73 ± 9.92 at 40 weeks. The MCA-PSV showed continuous increment with increasing gestational age

Conclusions: Continuous increment in the peak systolic volume with advancing gestational age was obtained which was consistent with the previous studies done by various authors. The percentile fitted values and normograms will be valuable for the serial measurement of the peak systolic volume of the middle cerebral artery for complicated pregnancies.

Keywords: fetal middle cerebral artery, fetal peak systolic velocity, pregnancy.

INTRODUCTION

Introduction of doppler velocimetry to obstetrics offered a non-invasive method of assessing fetoplacental circulation.1 Advance of ultrasonogaphy have revolutionized prenatal diagnosis of fetal anemia and intraterine-growth-restriction (IUGR).2 Use of fetal middle-cerebral-artery peak-systolic-velocity (MCA-PSV) for diagnosis of fetal anemia has reduced the number of invasive tests in the assessment of red cell alloimmunised pregnancies by >70%.3 Measurement of fetal MCA-PSV has a predictable relationship with fetal hemoglobin.4-5 The decreasing red-cell-mass leads to decreased blood viscosity and increased cardiac output resulting in increased fetal arterial-PSV. MCA-PSV is mainly used for the prediction of fetal anemia6-7 however; a preliminary study8 has reported increased MCA-PSV in IUGR fetuses.

Detection of truly high-risk pregnancy needs a strong tool with suitable reference ranges, the accuracy of which is important. Several fetal MCA-PSV reference ranges are currently in use3 however none are available for Nepalese population. This study was conducted to establish the reference values for fetal MCA-PSV in uncomplicated pregnancies between 19-40 weeks gestation in Nepalese population.

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METHODS

This was a prospective cross-sectional study done in the Department of Radiology and Imaging, Tribhuvan University Teaching Hospital, Nepal from September 2009 to August 2010. Four hundred singleton pregnancies between 19-40 weeks of gestation without any known risk factors of adverse pregnancy outcome who were referred for routine obstetric ultrasonography were studied. Those with undetermined period of gestation, maternal smoking, multiple pregnancy, diagnosed fetal abnormality in current pregnancy, previous history of preeclampsia, intrauterine growth retardation, abruptio placenta or preterm delivery, history of any pre-existing medical condition (such as hypertension, diabetes mellitus, renal disease), risk of developing fetal anemia including Rhesus negative women and those not willing to give consent for the study were excluded. Gestational age was determined from ultrasound examination before 20 weeks of gestation or from last menstrual period. While calculating the gestational age, those at three days and below were taken in lower gestational week and those after four days and above were taken in higher gestational week.

The doppler examination of fetal middle cerebral artery was done within 2 mm after its origin from the internal carotid artery with the angle of insonation kept at <20 degrees in all cases. The data obtained were compiled and analyzed using standard statistical analysis. SPSS 16.0 and Microsoft Excel were utilized for the data analysis and presentation. Pearson correlation was used to see the relation of doppler parameters with gestational age. Linear regression equations were calculated and percentile fitted values were obtained for the parameters at different gestational age. Normograms and percentile fitted curves were obtained.

RESULTS

During the period of this study 400 singleton uncomplicated pregnancies between 19-40 weeks of gestations were enrolled. The maternal age ranged from 18 to 36 years with the mean age being 24.94 ± 3.49 years. Majority of the patients (10%) were at 19 weeks of gestation and the least number of pregnancies (2.5%) were at 31 weeks.

Table 1. Mean and standard deviation of middle cerebral artery peak systolic velocity at different gestational age (n=400).

| Gestational Age (weeks) | Mean MCA-PSV | Standard Deviation |
|-------------------------|--------------|--------------------|
| 19                      | 22.2888      | 3.14365            |
| 20                      | 23.4823      | 3.62789            |
| 21                      | 23.6881      | 3.66769            |
| 22                      | 24.4428      | 3.74018            |
| 23                      | 25.0800      | 4.69530            |
| 24                      | 27.6214      | 2.98443            |
| 25                      | 28.4177      | 3.97432            |
| 26                      | 33.4629      | 4.06673            |
| 27                      | 36.1331      | 3.09927            |
| 28                      | 38.8000      | 6.07492            |
| 29                      | 42.8677      | 5.64531            |
| 30                      | 46.9617      | 8.39603            |
| 31                      | 48.2060      | 6.73701            |
| 32                      | 52.3294      | 10.56475           |
| 33                      | 53.3341      | 10.12149           |
| 34                      | 55.8285      | 11.25555           |
| 35                      | 59.1872      | 8.11738            |
| 36                      | 61.0894      | 9.75593            |
| 37                      | 62.2396      | 9.94632            |
| 38                      | 64.2150      | 9.10504            |
| 39                      | 64.5700      | 9.65823            |
| 40                      | 67.8408      | 9.67187            |

Table 2. Percentile values (5th, 50th and 95th) of middle cerebral artery peak systolic velocity at different gestational age (n=400).

| Gestational Age | Percentile 5th | Percentile 50th | Percentile 95th |
|-----------------|----------------|-----------------|-----------------|
| 19              | 17.5380        | 21.8750         | 28.1860         |
| 20              | 18.0800        | 23.6150         | 29.3865         |
| 21              | 18.2000        | 24.1350         | 30.8600         |
| 22              | 19.1200        | 24.1900         | 33.1000         |
| 23              | 19.4100        | 25.4400         | 36.0100         |
| 24              | 23.4700        | 27.5300         | 36.6320         |
| 25              | 24.0400        | 28.6300         | 39.0800         |
| 26              | 27.5500        | 32.5700         | 40.3600         |
| 27              | 29.9400        | 36.4100         | 42.4700         |
| 28              | 31.4900        | 38.9700         | 50.8400         |
| 29              | 34.5200        | 42.8900         | 51.2200         |
| 30              | 35.9700        | 46.5150         | 56.4800         |
| 31              | 37.4300        | 48.7850         | 59.5800         |
| 32              | 38.7500        | 50.4250         | 67.6800         |
| 33              | 39.1500        | 51.6400         | 71.9300         |
| 34              | 40.4325        | 54.3950         | 76.7345         |
| 35              | 42.2330        | 58.1400         | 77.1370         |
| 36              | 42.9600        | 59.6650         | 80.0200         |
| 37              | 44.4550        | 63.5100         | 81.2525         |
| 38              | 45.7565        | 65.0450         | 84.2550         |
| 39              | 47.1160        | 66.7400         | 84.2690         |
| 40              | 47.6400        | 69.9600         | 84.4000         |
The fetal peak systolic blood flow in the middle cerebral artery showed significant correlation with period of gestation with a Pearson correlation (r) of 0.912 (p value: 0.0001). Mean MCA-PSV was 22.35 ± 3.05 at 19 weeks of gestation, which increased to 67.73 ± 9.92 at 40 weeks (Table 1). Table 2 shows the reference values of MCA-PSV with 5th, 50th and 95th percentile values. Linear regression was obtained for each Doppler parameter, which shows highly significant correlation (p value = 0.0001) between MCA-PSV and gestational age.

**DISCUSSION**

Cross-sectional observations were obtained in 400 uncomplicated singleton pregnancies between 19 and 40 weeks of gestation. Reference ranges for fetal MCA-PSV was constructed based on these observations. The MCA-PSV showed significant increase with increasing period of gestation with a significant positive correlation with period of gestation (r=0.912, p <0.0001). Although this confirms other studies\(^2\),\(^9\)-\(^13\) the reference values for MCA-PSV in this study varied from other studies, which are as follows.

Kurmanavicius J et al\(^12\) in a cross sectional study involving 331 normal pregnancies between 19-40 weeks also showed that MCA-PSV increases with increasing period of gestation and the 5th percentile of both the studies were similar however the 50th and 95th of our study were higher than the study conducted by Kurmanavicius J et al. Ebbing et al\(^10\) in a study with 161 singleton pregnancies between 19-40 weeks period of gestation also showed that the MCA-PSV increases with increasing gestational age but the reference values were lower than this study. Tongsong et al\(^11\) in a prospective cross-sectional descriptive study in which low-risk singleton pregnancies with accurate gestational age were recruited into the study showed a continuous increase in MCA-PSV over the period from 11-22 weeks. The 5th and 50th percentile of MCA-PSV in 19-22 weeks were lower than our study, however the 95th percentile were higher than our study.

Tan et al\(^2\) in a study conducted in 329 normal Asian pregnant women between 16-40 weeks of gestation comparing the reference values for MCA-PSV between Asian and non-Asian obstetric populations also showed that the MCA-PSV in Asian women increases with increasing period of gestation. The reference range in that study was lower than our study. This seems to suggest that there may be essential differences not only between Asian and non-Asian populations but also among various Asian countries, regions and ethnic groups. Thus this emphasizes the need to obtain the reference value for Nepalese population and also various ethnic groups among the Nepalese populations as using a non-local reference range could pick up more false-positives for anemia, especially in the later gestational ages. However, we note that these postulations have not been validated by a correlation of fetal and neonatal hemoglobin levels in our own population. It is thus hoped that this study will encourage more work to be done to ascertain and validate this local reference range for MCA-PSV, and for it to be utilized in the local setting in future for obstetric practice.
CONCLUSIONS

Reference ranges for peak systolic velocity of fetal middle cerebral artery were constructed for Nepalese population. Continuous increment in the peak systolic volume with advancing gestational age was obtained. The percentile fitted values and normograms were also obtained which will be valuable for the serial measurement of the peak systolic volume of the middle cerebral artery for complicated pregnancies.

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