Cerebro-placental ratio as a prognostic factor of fetal outcome in women with hypertensive disorders complicating pregnancy during third trimester

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ABSTRACT

Background: CPR is emerging as an important predictor of adverse pregnancy outcome and helps in management of high risk pregnancy. Therefore we undertook the study to find the correlation of CPR with perinatal outcomes in women with hypertensive disorder complicating during third trimester.

Methods: 128 patients with hypertensive disorder of pregnancy, ≥32 weeks of singleton gestation, were randomly selected during their hospital visit. They were subjected to USG Doppler study to calculate MCA/UA pulsatility index-CPR. The CPR<1 was considered abnormal and >1 as normal. These results were compared with the perinatal outcome and adverse fetal outcome.

Results: The present study revealed that the incidence of adverse outcomes like Apgar score <7 (36.5%), still birth (15.9%), NICU admission (69.8%) and LBW i.e. <2500 gm (68.3%) were significantly higher in abnormal CPR than normal CPR.

Conclusions: Abnormal CPR is valuable in predicting the outcome of hypertensive disorders in pregnancy. CPR is an easy procedure which can be included in the routine antenatal sonographic evaluation to predict poor perinatal outcome and to detect or recognize those fetuses at risk.

Keywords: Cerebro placental ratio, Hypertensive disorders of pregnancy, Middle cerebral artery, Umbilical artery

INTRODUCTION

Hypertensive disorders represent the most common maternal and perinatal complication of pregnancy. It has been reported that the incidence of this disorder is between 5 and 10 percent. Hypertensive disorders associated with pregnancy along with haemorrhage and infection form a deadly triad that contributes greatly to maternal as well as perinatal morbidity and mortality.¹²

Preeclampsia is associated with increased vascular resistance and decreased uteroplacental perfusion resulting in increased incidence of fetal hypoxia. In such condition, fetal modifications occur in retort to non-performance of the maternal-placental nutrients supply so as to meet the foetal demands.³⁴

In recent decades, routine antenatal ultrasonography is commonly used in assessment of anthropometric measurement of umbilical cord and placenta and also to detect the intrauterine growth retardation (IUGR) and congenital anomalies. Yet women with high risk pregnancies like hypertensive disorder of pregnancy and diabetes mellitus were showing the increase incidence of maternal and fetal morbidity. Various biochemical tests used in screening of high risk population for preeclampsia have lower positive predictive values, high cost and less patient compliance.⁵
Doppler sonography especially cerebro-placental ratio (CPR) is a rapid non-invasive test that provides valuable information about hemodynamic status of the fetus and is an efficient diagnostic tool to evaluate blood flow alterations in placenta-umbilical and feto-cerebral circulation. The CPR is calculated by dividing the Doppler pulsatility index from the middle cerebral artery (MCA) into the umbilical artery (UA). CPR is emerging as an important predictor of adverse pregnancy outcome and helps in management of high risk pregnancy. Therefore we undertook the study to find the correlation of CPR with perinatal outcomes in women with hypertensive disorder complicating during third trimester.

**METHODS**

This cross-sectional study was conducted among 128 pregnant women who underwent a routine antenatal sonogram at the Department of Radiodiagnosis, JSS medical college, Mysuru, Karnataka, India, during the study period November 2015 to July 2017. The study was approved by the Institutional Ethical Committee and written informed consent was obtained from the participants.

**Inclusion criteria**

Our inclusion criteria were singleton pregnant women with hypertensive disorder of pregnancy and gestational age at the time of study ≥28 weeks.

**Exclusion criteria**

We excluded pregnancies completed with co-morbid conditions like gestational diabetes, IUGR, oligohydramnios, polyhydramnios and intrauterine death (IUD).

About 128 antenatal women were selected after consideration of inclusion and exclusion criteria, their preliminary data were recorded and were subjected to ultrasound scan for fetal anthropometric parameters, fetal and placental position, fetal age and heart rate, umbilical artery pulsatility index (UI), middle cerebral artery pulsatility index (MI). CPR will be calculated from Middle cerebral artery pulsatility index and Umbilical artery pulsatility index.

A CPR was defined as normal and abnormal when its ratio was >1.0 and <1.0 respectively. Based on the above statement, cases were divided into normal group (CPR>1.0), and abnormal group (CPR<1.0). Follow up with respect to perinatal outcome and to detect the fetuses at risk.

The mean and standard deviation of MCA PI and UA PI were calculated for different gestational age groups. The data obtained were statistically analysed using SPSS software version 20.0. Association between cerebro-placental ratio and other categorical variables were analysed using chi-square test. Pearson Correlation coefficient was used to show correlation between cerebro-placental ratio and other quantitative variables like birth weight.

**RESULTS**

Total one hundred and twenty eight pregnant women were included in this study. The mean±SD of maternal age was 24.74±4.08 years, ranging from 18-37 years. The mean±SD of gestational age during delivery was 36.78±2.46 weeks, ranging from 32-41 weeks. The mean birth weight was 2.44 kg (SD=0.74), ranging from 0.60 to 3.60 kg. Table 1 shows the maternal and perinatal characteristics of study population.

**Table 1: Maternal and perinatal characteristics of study population (n=128).**

| Characteristics                          | n (%) |
|------------------------------------------|-------|
| Parity                                   |       |
| Primipara                                | 82 (64.1%) |
| Multipara                                | 46 (35.9%) |
| HTN disorder in pregnancy                |       |
| Eclampsia                                | 26 (20.3%) |
| Pre-eclampsia                            | 91 (71.1%) |
| Gestational HTN                          | 11 (08.6%) |
| Delivery                                 |       |
| Normal vaginal delivery                   | 26 (20.3%) |
| LSCS                                     | 102 (79.7%) |
| Fetal outcome                            |       |
| Live birth                               | 118 (92.2%) |
| Still birth                              | 10 (07.8%) |
| Apgar score at 5 minutes                 |       |
| ≤7                                       | 28 (21.8%) |
| >7                                       | 100 (78.2%) |
| Admission to NICU                        |       |
| Yes                                      | 79 (61.7%) |
| No                                       | 49 (38.3%) |

| Gestational age | No. of cases | MCA PI | UA PI |
|-----------------|--------------|--------|-------|
| 28              | 03           | 0.55   | 0.78  | 0.90  | 1.27  | 0.08  | 0.53  |
| 29              | 02           | 1.20   | 1.06  |       |       |       |       |
| 30              | 06           | 0.86   | 0.67  | 0.99  | 0.84  |       |       |
| 31              | 04           | 1.00   | 1.15  | 1.05  | 0.85  |       |       |
| 32              | 09           | 1.23   | 1.36  | 1.54  | 0.72  |       |       |
| 33              | 08           | 0.91   | 1.28  | 0.88  |       |       |       |
| 34              | 13           | 1.23   | 1.66  | 0.34  |       |       |       |
| 35              | 08           | 1.27   | 1.37  | 0.34  |       |       |       |
| 36              | 24           | 1.31   | 1.34  | 0.35  |       |       |       |
| 37              | 15           | 1.28   | 1.54  | 0.37  |       |       |       |
| 38              | 18           | 1.29   | 1.38  | 0.48  |       |       |       |
| 39              | 10           | 1.44   | 1.26  | 0.22  |       |       |       |
| 40              | 08           | 1.44   | 1.09  | 0.08  |       |       |       |
| Total           | 128          | 1.23   | 1.35  | 0.53  |       |       |       |
Table 2 shows descriptive statistics of MCA pulsatility index, UA pulsatility index based on gestational age (by USG). The middle cerebral artery pulsatility index (MCA PI) had a mean value of 1.22, with the scores ranging in between 0 to 1.96; The Umbilical artery (UA) pulsatility index had mean value of 1.34 (range 0-2.6).

Table 3: Perinatal outcome characteristics based on cerebro-placental ratio.

| Variables                              | Abnormal CPR (n=63) | Normal CPR (n=65) | P value |
|----------------------------------------|---------------------|-------------------|---------|
| Mode of delivery [n (%)]               |                     |                   |         |
| NVD                                    | 14 (22.2%)          | 12 (18.5%)        | 0.597   |
| LSCS                                   | 49 (77.8%)          | 53 (81.5%)        | (NS)    |
| Apgar score [n (%)]                    |                     |                   |         |
| <7                                     | 23 (36.5%)          | 05 (7.7%)         | 0.000   |
| >7                                     | 40 (63.5%)          | 60 (92.3%)        | (SIG)   |
| Fetal outcome [n (%)]                  |                     |                   |         |
| Alive                                  | 53 (84.1%)          | 65 (100.0%)       | 0.001   |
| Still birth                            | 10 (15.9%)          | 0 (0%)            | (SIG)   |
| NICU admission [n (%)]                 |                     |                   |         |
| Yes                                    | 44 (69.8%)          | 35 (53.8%)        | 0.063   |
| No                                     | 19 (30.2%)          | 30 (46.2%)        | (NS)    |
| Birth weight [n (%)]                   |                     |                   |         |
| <2500 gm                               | 43 (68.3%)          | 19 (58.7%)        | 0.000   |
| >2500 gm                               | 20 (31.7%)          | 46 (70.8%)        | (SIG)   |
| Gestational age at the time of delivery [n (%)] |             |                   |         |
| ≤37 weeks                              | 38 (60.3%)          | 30 (46.2%)        | 0.108   |
| >37 weeks                              | 25 (39.7%)          | 35 (53.8%)        | (NS)    |

Table 4: Correlation between cerebro-placental ratio and various other variables in the Study (N=128).

| Parameter                              | Correlation coefficient | P value |
|----------------------------------------|-------------------------|---------|
| Gestational age at the time of delivery | 0.315                   | <0.0001 |
| Gestational age (USG)                  | 0.432                   | <0.0001 |
| Birth weight                           | 0.565                   | <0.0001 |
| Apgar score at 5 minutes               | 0.659                   | <0.0001 |

The cerebro-placental ratio (CPR) was calculated and showed a mean value of 0.92, ranging from 0 to 1.63; the value below 1.0 was considered as abnormal, and value above 1.0 was considered as normal. The adverse perinatal outcomes are LSCS, Apgar score <7 at 5 minutes, still birth, preterm delivery, NICU admission, and low birth weight (LBW <2500 gm) was observed in both the normal CPR and abnormal CPR. Table 3 shows the perinatal outcome characteristics based on the cerebro-placental ratio. Table 4 shows the correlation between CPR and other variables in the present study. The study population showed significant correlation between gestational age (both at the time of delivery and USG), birth weight, Apgar score at 5 min and cerebro-placental ratio values. All the values were statistically significant (p<0.01).
DISCUSSION

The primary goal of every obstetrician is to identify the pregnancies at risk and to prevent them from the perinatal morbidity and mortality. Every life in the earth has authority to born healthy both physically and mentally.

Doppler index of umbilical artery and middle cerebral artery especially pulsatility index was measured based on gestational age during third trimester. It is very difficult to use single cut off value for pulsatility index after 30 weeks, whereas CPR is very realizable to use as single cut off value after 30 weeks because cerebral-umbilical Doppler ratio does not vary significantly between 30th and 40th weeks as reported by Waldimiroff et al. This statement is in agreement with Arbeille et al. He suggested 1 as the cut off value and all values below 1 were considered abnormal. We classified our cases based on Arbeille et al.

In this present study it was shown that majority of the study population were primigravida. Among the pregnancy related hypertension, pre-eclampsia was the most common cause followed by eclampsia and gestational hypertension. The present study revealed that the incidence of adverse outcomes like Apgar score <7 (36.5%), still birth (15.9%), NICU admission (69.8%) and LBW i.e. <2500 gm (68.3%) were significantly higher in abnormal CPR than normal CPR.

Our observation is in concordance with Kant et al, who conducted the case control study to measure the CPR in normal and high risk pregnant women. Kant et al (2016) reported a high correlation between the abnormal CPR with caesarean section and NICU admission. In another study by Dagade et al clearly described an association of abnormal CPR with low birth weight, caesarean section, NICU admission and intrauterine death.

The middle cerebral artery pulsatility index (MCA PI) had a mean value of 1.22 (range 0 to 1.96). The Umbilical artery (UA) pulsatility index had mean value of 1.34 (range 0-2.6). The MCA PI is found to be significant independent predictive value of LBW. This observation was in accordance with observation of Lam et al.

A statistically significant positive correlation was observed between CPR and gestational age at the time of delivery ($r^2=0.315$, $P<0.0001$), gestational age by USG ($r^2=0.432$, $P<0.0001$), birth weight ($r^2=0.565$, $P<0.0001$) and Apgar score at 5 minutes ($r^2=0.659$ $P<0.0001$). In this study abnormal CPR had highest sensitivity (68.3%) for predicting low birth weight, 69.8% for NICU admission, and 77.8% for caesarean section and had highest specificity (100%) for predicting still birth, 70.8% for low birth weight. The sensitivity was comparable with that of Gramellini et al, Fong et al and Srilakshmi which accounts for 24%, 72.4% and 83.3% respectively.

Literature review reveals that the appropriate for gestational age (AGA) fetuses with a birth weight above the 10th percentile had the higher prevalence of adverse outcome.

Hence it could be justified that, primary care should be initiated for identifying fetal hypoxia instead of small for fetus where adverse outcome is due to consequence of placental impairment. Further screening should focus on the detection of pregnancies with low CPR rather than those with low estimated fetal weight.

The major limitation of our study was the low sample size. We suggest further studies with bigger sample sizes to validate the results of the present study.

CONCLUSION

The results of the present study revealed the relationship between cerebro-placental ratio and hypertensive disorder of pregnancies. CPR is an easy procedure which can be included in the routine antenatal sonographic evaluation to predict poor perinatal outcome and to detect or recognize those foetuses at risk irrespective of the findings of the individual values of MCA and UA. Such pregnancies should be closely followed up for the successful perinatal outcome.

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