

A Prospective Study of Doppler Velocimetry in Pregnancy-induced Hypertension in a Rural Population of a Developing Country

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ABSTRACT

Background: Pregnancy-induced hypertension (PIH) remains a great challenge to obstetricians. Doppler velocimetry can detect fetal compromise much before other antepartum tests. **Aim:** The aim of this study is to detect the changes of uterine artery, umbilical artery and middle cerebral artery in PIH by Doppler velocimetry. **Subjects and Methods:** This prospective study was conducted on hundred subjects with PIH. Doppler studies were carried, and parameters recorded in uterine, umbilical and middle cerebral artery (MCA) were Systolic/Diastolic ratio, Resistance Index, Cerebro-Placental Index (CPI). Fetal outcomes were monitored. Statistical analysis was performed using Epi Info™ software (Version 3.5.1, CDC, Atlanta). Test for significance was done with student's t-test and Chi-square where applicable. A P-value of <0.05 was considered as significant. **Results:** Among the 100 subjects, 76 (76%) of fetuses had abnormal and 24% normal umbilical artery Doppler velocimetry; 62% had abnormal and 38% normal MCA Doppler velocimetry; 64% fetuses had abnormal and 36% normal CPI. In 95% of subjects having abnormal umbilical Doppler studies, caesarean section had to be done for acute fetal distress. Incidence of caesarean section was 61% in abnormal MCA group and 63% in abnormal CPI group. Among 14 patients who had abnormal uterine artery Doppler, four developed pre-eclampsia, 2 IUGR. In patients with notches in uterine artery Doppler, 38% developed pre-eclampsia, 38% had IUGR, 13% babies were still born and 25% of newborns required NICU admission. In umbilical artery Doppler, when S/D ratio was abnormal, 60% developed pre-eclampsia, 40% had IUGR and 40% of newborns had to be admitted in NICU. **Conclusion:** Doppler study for fetal surveillance in pregnancy-induced hypertension is a very useful device and abnormal umbilical artery and uterine artery velocimetry seems to have worse pregnancy outcomes in the present study. Notch as a single parameter is the best indicator with highest sensitivity and positive predicative values. However, combination of parameters is the best indicator.

KEY WORDS: Doppler study, fetomaternal outcome, pregnancy-induced hypertension

INTRODUCTION

Pregnancy-induced hypertension (PIH) remains a great challenge to obstetricians. Impaired utero-placental blood flow in PIH may result in intrauterine growth restriction (IUGR), placental abruption and intrauterine fetal death (IUFD).^[1-3] Doppler velocimetry of uterine artery, umbilical artery and middle cerebral artery can detect fetal defects much before any other antepartum test.

The blood flow characteristics can be quantified by various Doppler indices like the systolic/diastolic ratio (S/D ratio = Peak systolic velocity/End diastolic velocity),

resistance index (RI = Peak systolic velocity- End diastolic velocity/Peak systolic velocity), Cerebro-placental index (CPI = Middle cerebral artery resistance/umbilical artery resistance) and pulsatility index (PI = Peak systolic velocity – End diastolic velocity/Mean velocity).^[2-5]

Forty percent of women with bilateral notches of uterine arteries and high pulsatility index have chances of developing pre-eclampsia, while in 45%, there is chance of developing IUGR.^[2-4] Color Doppler is an excellent tool for non-invasive hemodynamic monitoring of PIH patients. It helps to identify the fetuses at risk and predict perinatal morbidity and mortality.^[2-6]

The present study was conducted in a tertiary care hospital catering a rural population in a developing country to find

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the incidence of PIH and detect the changes by Doppler velocimetry of uterine artery, umbilical artery, and middle cerebral artery. Early detection of abnormalities by Doppler may improve perinatal outcome and decrease fetal and maternal morbidity and mortality. The majority of adverse perinatal outcomes in developing countries are placental-associated diseases. Doppler evaluation predicts most occurrences of early-onset preeclampsia and intrauterine growth restriction, and its use in these pregnancies improves perinatal outcomes.

SUBJECTS AND METHODS

This prospective study was conducted in a time span of one year on hundred subjects in a teaching hospital in West Bengal, catering mainly a rural population of a developing country. Clearance of the Institutional ethical committee and consent of the patients were taken. The study was conducted in the department of Gynecology and Obstetrics in collaboration with the Department of Radiology and Physiology. Inclusion criteria: All antenatal patients with blood pressure $\geq 140/90$ after 20 weeks of gestation, with proteinuria 300 mg/24 h or edema or both. Exclusion criteria: Women with multiple pregnancies, chronic hypertension, cephalopelvic disproportion being the cause of previous caesarean section done, having fetus with congenital abnormality, history of antepartum hemorrhage, smoking, hepatic, renal, cardiac disease were excluded. During antenatal checkup after careful history taking, physical examination, blood tests (total count, differential count, erythrocyte sedimentation rate, hemoglobin percentage, bleeding time, clotting time, platelet count, random blood sugar, urea, creatinine, uric acid, VDRL tests, Hepatitis B surface antigen, HIV), Doppler studies were carried out. During Doppler studies parameters recorded were S/D ratio, RI in uterine, umbilical and middle cerebral artery (MCA), Pulsatility Index (PI = difference between the peak systolic and end-diastolic shift divided by the average shift (A) over the cardiac cycle. The flow velocity waveforms were considered abnormal if there existed an early diastolic notch in uterine artery (in either right or left uterine arteries) and S/D, RI exceeded 95th percentile of the range of reference; in umbilical artery if S/D, RI exceeded 95th percentile and if end-diastolic flow velocity was absent or reversed. S/D ratio of 3 was considered abnormal after 30th weeks of pregnancy. CPI ≤ 1 was considered abnormal.

Outcome measures recorded were: IUGR, IUFD, LBW (low birth weight), Gestational age at birth, Apgar score and Neonatal intensive care unit (NICU) admissions required at birth. In cases with meconium staining of liquor, caesarean section was done.

Statistical analysis was performed using Epi Info™ software (Version 3.5.1, CDC, Atlanta). Test for significance was done with student's *t*-test and Chi-square where applicable. *P* value $<0.05^*$ was considered as significant.

RESULTS

Among hundred PIH cases, 76% (76/100) of fetuses had abnormal and 24% (24/100) normal umbilical artery Doppler velocimetry (24 had UA S/D ratio <3 and in 76 UA S/D ratio was ≥ 3). Significant differences were seen in meconium staining of liquor, birth weight, NICU admission, Apgar score and number of caesarean section done among the two groups.

Sixty two percent (62/100) of the patients had abnormal and 38% (38/100) normal MCA Doppler velocimetry. Incidence of caesarean section was significantly increased. Among the 32 fetuses who had MCA S/D ratio ≥ 3 , 29 had poor fetal and neonatal outcome and out of 19 fetuses who had MCA S/D ratio <3 , seven had poor outcome.

Sixty four percent (64/100) of fetuses had abnormal and 36% (36/100) normal CPI. Table 1 shows that fetuses with CPI ≤ 1 had significantly lower mean gestational age at birth. Meconium staining of liquor during labor for which caesarean section was done was significantly higher ($P < 0.001$) among these fetuses. Caesarean section for acute fetal distress was ninety five percent in abnormal umbilical Doppler group. Incidence of caesarean section was sixty one percent in abnormal MCA group and sixty three percent among abnormal CPI group.

The incidence of low Apgar score (<7) at 5 minutes and NICU admission rate was significantly higher ($P < 0.01$) among fetuses who had CPI ≤ 1 . This group of fetuses also had significantly longer ($P < 0.01$) NICU stay. Out of 32 fetuses who had CPI values of 1 or less, 31 had poor fetal and neonatal outcome; while among 18 fetuses who had CPI > 1 , in 5 outcome was poor. Among 14 patients who had abnormal uterine artery Doppler, four developed pre-eclampsia, two IUGR. Abnormal CPI value had a sensitivity of 86.1%, specificity of 92.8%, positive predictive

Table 1: Neonatal outcome according to cerebroplacental index

Neonatal events	Cerebroplacental index		P value
	≤ 1 (n=32)	>1 (n=31)	
Gestational age at delivery (weeks)	34.7 (3.1)	36.7 (2.9)	0.33
Birth weight <2500 g (n=40)	31 (96.9%)	9 (50%)	$<0.001^*$
Meconium staining of liquor (n=33)	25 (78%)	8 (44%)	0.04*
Apgar score <7 at 5 min. (n=36)	29 (90.6%)	7 (38.9%)	$<0.001^*$
Caesarean section (n=24)	20 (62.5%)	4 (22.2%)	$<0.01^*$
NICU admission (n=24)	22 (68.8%)	2 (11.1%)	$<0.001^*$
NICU admission >7 Days (n=20)	18 (56.3%)	2 (11.1%)	$<0.01^*$

NICU – Neonatal intensive care unit; P value $<0.05^*$ significant

value of 96.9%, and negative predictive value of 72.2% and diagnostic accuracy of 88%.

Table 2 shows S/D ratio and notch had sensitivity of 60% and positive predictive value of 33.3% and 37.5%, respectively. Notch as a single parameter is the best indicator with highest sensitivity and positive predictive values. However, combination of parameters is the best indicator. Table 3 shows S/D ratio as the highest sensitivity and positive predictive value index in umbilical artery. Table 4 shows correlation of uterine artery Doppler with pregnancy outcomes. RI was abnormal in seven cases. Among them 29% patients developed pre-eclampsia, 14% developed PIH, 29% had IUGR, 29% had IUFD, 14% babies were still born and 25% of neonates had to be admitted in NICU. Nine patients had abnormal S/D ratio in uterine artery Doppler. Among them, 33% developed pre-eclampsia, 33% had IUGR, 22% had IUFD, 11% babies were still born and 22% of neonates had to be admitted in NICU. Eight patients had early diastolic notch in uterine artery Doppler. Among them 38% patients developed pre-eclampsia, 38% had IUGR, 22% had IUFD, 13% babies were still born and 25% of neonates had to be admitted in NICU. Table 5 shows correlation of umbilical artery Doppler with pregnancy outcomes. In umbilical artery Doppler, S/D ratio was abnormal in five cases. Sixty percent patients developed pre-eclampsia, 40% had IUGR, 20% had IUFD, 20% babies were still born and 40% neonates had to be admitted in NICU. RI was abnormal in eight cases. Twelve percent patients developed pre-eclampsia, 25% had IUGR, and 12% neonates had to be admitted in NICU. Diastolic flow was absent in one case. The patients who developed pre-eclampsia, had IUGR and IUFD.

DISCUSSION

PIH is a common cause of feto-maternal mortality affecting 10% of pregnant women and is associated with 22% perinatal deaths. Doppler studies in high-risk pregnancies are more beneficial in the management of perinatal and neonatal

outcomes.^[1-7]

In the present study, 76% of fetuses had abnormal and 24% normal umbilical artery Doppler velocimetry; 62% had abnormal and 38% normal MCA Doppler velocimetry; 64% fetuses had abnormal and 36% normal CPI. In 95% of subjects having abnormal umbilical Doppler studies, caesarean section had to be done for acute fetal distress. Incidence of caesarean section was 61% in abnormal MCA group and 63% in abnormal CPI group. Liberati *et al.*^[7] showed a mean uterine artery RI ≥ 0.66 (90th percentile) had better sensitivity than the placental and the non-placental uterine artery. The presence of a diastolic notch in the placental uterine artery increased sensitivity (31.7% for IUGR and 50.0% for PIH) and positive predictive value of the test.^[7] In the present study, S/D ratio and notch had sensitivity of 60% and positive predictive value of 33.3% and 37.5%, respectively, and S/D ratio had the highest sensitivity and positive predictive value index in umbilical artery. Sierszowski *et al.*^[8] observed the sensitivity of the notch in 20-24 week in the uterine artery velocimetry for the prediction of PIH and/or IUGR in the III trimester was 73%, 68% and the specificity 97.15%.

In studies of Bhatt *et al.*,^[9] 56% had abnormal S/D ratio in umbilical artery and/or uterine artery. The percentage of abnormal velocimetry seems to be higher in the present study. Sixty percent of these patients delivered IUGR babies. In patients with absent end diastolic velocity and reversed end diastolic velocity, perinatal mortality was 50%, and 50% had IUGR babies.^[9] Our study found decreased number of preterm births, obstetric interventions and improved outcome morbidity and mortality from the use of Doppler waveform analysis in early pregnancy. Evidence suggested that many pregnancy disorders originate at conception or in early gestation. A retrospective study was conducted by Frusca *et al.*^[10] on 344 hypertensive pregnant women. Abnormal uterine velocimetry was associated with a worse pregnancy outcome.^[10,11] Our study was a prospective study, while

Table 2: Value of uterine artery Doppler in predicting pre-eclampsia

Doppler test	True positive	False negative	False positive	True negative	Sensitivity	Specificity	Positive predictive value	Negative predictive value
Uterine artery								
S/D	3	2	6	89	60.0	93.7	33.3	97.3
RI	2	3	6	89	40.0	96.7	25.0	96.7
Notch	3	2	5	90	60.0	97.7	37.5	97.8
Combined	4	1	10	85	80.0	89.4	28.6	98.8

RI – Resistance index; S/D – Systolic/diastolic ratio

Table 3: Value of umbilical artery Doppler in predicting pre-eclampsia

Doppler test	True positive	False negative	False positive	True negative	Sensitivity	Specificity	Positive predictive value	Negative predictive value
SD ratio	2	3	3	92	40.0	96.8	40.0	96.8
RI	1	4	7	88	25.0	92.6	12.5	95.6
Combined	2	3	8	87	40.0	91.5	20.0	96.6

RI – Resistance index; S/D – Systolic/diastolic ratio

Table 4: Correlation of uterine artery Doppler with pregnancy outcome (%)

Uterine artery Doppler	Preeclampsia %	IUGR	IUFD %	Still birth	NICU admission %
S/D ratio (n=9)	33 (3/9)	33 (3/9)	22 (2/9)	11 (1/9)	22 (2/9)
RI (n=7)	29 (2/7)	29 (2/7)	29 (2/7)	14 (1/7)	14 (1/7)
Early diastolic notch (n=8)	38 (3/8)	38 (3/8)	38(3/8)	13 (1/8)	25 (2/8)

RI – Resistance index; S/D – Systolic/diastolic ratio; IUGR – Intrauterine growth restriction; IUFD – Intrauterine fetal death; NICU – Neonatal intensive care unit

Table 5: Correlation of umbilical artery Doppler with pregnancy outcome (%)

Umbilical artery Doppler	Preeclampsia %	IUGR %	IUFD %	Still birth %	NICU admission %
S/D ratio (n=5)	60 (3/5)	40 (2/5)	2 (1/5)	20% (1/5)	40 (2/5)
RI (n=8)	12 (1/8)	25 (2/8)	0	0	12 (1/8)
Absent diastolic flow (n=1)	100 (1/1)	100 (1/1)	100 (1/1)	0	0

RI – Resistance index; S/D – Systolic/diastolic ratio; IUGR – Intrauterine growth restriction; IUFD – Intrauterine fetal death; NICU – Neonatal intensive care unit

study by Frusca *et al.*^[10] was a retrospective one. Uterine artery sensitivity was 90% as compared to 40% sensitivity of combined parameters of umbilical artery in the present study, which is comparable to the above study.

Messawa *et al.*^[12] divided their subjects into group A (100) subjected to Doppler velocimetry and group B (100) without Doppler velocimetry. Preterm deliveries, preterm as well as full-term neonatal admissions were more frequent in group A than those in group B, i.e., (39% vs. 26%), (56% vs. 88%) (OR 0.2, 95% CI 0.04-0.7), and (30% vs. 57%) (OR 0.3, 95% CI 0.2-0.7), respectively. Similarly preterm and full-term neonatal deaths were rare in group A than those in group B, i.e., (9% vs. 78%) (OR 0.1, 95% CI 0.02-0.7) and (6% vs. 29%) (OR 0.2, 95% CI 0.03-1.8), respectively. Emergency caesarean section rate was rare in the subjects with normal Doppler than those with abnormal Doppler (48% vs. 100%) (OR 0.1, 95% CI 0.03-0.4) as well as in group B (48% vs. 82%) (OR 0.2, 95% CI 0.1-0.4). These outcomes are comparable to the present study.

Future studies can be conducted using Color Doppler imaging, the most appropriate and the commonly used method for the study of ocular circulation, especially during pregnancy. It enables the visualization and flow measurement of retro bulbar blood vessels. Hemodynamic studies have shown that general arteriolar vasoconstriction, which leads to hypoperfusion of target organs, including the eye area, remains the most significant pathological change in preeclampsia.^[13] The cardiovascular reactivity adaptation seen in normal pregnancy is absent in gestational hypertension.^[14] Thus it may be said that Doppler study for fetal surveillance in pregnancy-induced hypertension is a very useful device in reduction of perinatal morbidity and mortality by timely intervention.^[15-18]

Strengths of the study: The present study was conducted in a developing country with high incidence of maternal morbidity and mortality. Early detection of abnormal Doppler studies yielded better results in maternal of fetal outcomes as proper intervention in patient care was adopted.

Limitations of our study: The sample was drawn from one limited geographic area; the results cannot properly be generalized to the national population. Second, because of the cross-sectional design, this study had limited extrapolative value. A randomized comparison of routine versus highly selective Doppler waveform and biophysical profile usage may yield more conclusive results. We did not estimate PIGF. We are planning for a larger project in near future with the population at a large in different rural centers of our hospital in collaboration with the National Rural Health mission.

CONCLUSION

Doppler study for fetal surveillance in pregnancy-induced hypertension is a very useful device and abnormal umbilical artery and uterine artery velocimetry also seems to have worse pregnancy outcomes in the present study. Notch as a single parameter is the best indicator with highest sensitivity and positive predicative values. However, combination of parameters is the best indicator.

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