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# Comparison of the resistive indices obtained in the uterine artery and the ophthalmic artery in preeclamptic and normotensive patients in Doppler US

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## **Keywords**

preeclampsia, resistive index, uterine artery, ophthalmic artery, Doppler

#### Abstract

Introduction: Preeclampsia is a medical condition characterized by hypertension and proteinuria during pregnancy, with the symptoms generally manifesting in the 3<sup>rd</sup> trimester. Hypertension brings hemodynamic changes; it is therefore expected that arterial blood flow velocity waveforms will be different in the uterine and ophthalmic arteries in preeclampsia. Vascular changes do occur in preeclampsia, which in turn induces hemodynamic changes. **Aim:** To compare mean values of the resistive index of the ophthalmic and uterine arteries in patients with preeclampsia and normotensive individuals in Doppler US. Methods: In this cross-sectional observational study, ultrasound of the uterine and ophthalmic arteries was performed in 60 pregnant women in the 2nd and 3rd trimester of pregnancy to compare the resistive index of these arteries in preeclamptic and normotensive individuals. All the patients, i.e. 30 preeclamptic and 30 normotensive pregnant women, took part in this study voluntarily. The study was approved by the institutional review board (IRB) of the University of Lahore. **Results:** The mean resistive index of the uterine artery was  $0.50 \pm 0.08$  in normotensive participants and  $0.64 \pm 0.09$  in preeclamptic women, with the p-value < 0.001. The mean resistive index of the ophthalmic artery was  $0.70 \pm 0.05$  in normotensive participants and  $0.63 \pm 0.04$  in preeclamptic women, with the p-value < 0.001. Conclusions: There was a significant negative correlation between the resistive index of the uterine and ophthalmic arteries among the patients with preeclampsia and a significant positive correlation among normotensive individuals. Preeclampsia could be easily diagnosed with Doppler ultrasound based on hemodynamic changes in response to vascular changes in the ophthalmic and uterine arteries.

## Introduction

Preeclampsia is one of the leading causes of maternal morbidity throughout the world(1). It is a pregnancy disorder characterized by high blood pressure and often a significant amount of protein in urine(2). The incidence of preeclampsia is from 5–10%<sup>(3,4)</sup>, and its prevalence is 2–8%<sup>(5)</sup>. The possible causes of preeclampsia include: an exaggerated inflammatory response by endothelial cells, which intrinsically increases the local blood supply to the affected area, (6) and the activation of monocytes<sup>(7,8)</sup> and granulocytes as the maternal inflammatory response<sup>(9)</sup>. An increase in maternal diastolic pressure during pregnancy can also lead to preeclampsia<sup>(10)</sup>. The possible interventions in preeclampsia include therapies with melatonin<sup>(11)</sup> to decrease blood pressure (BP), corticosteroids (in severe preeclampsia), anticonvulsants (to prevent seizers in severe cases) as well as bed rest, hospitalization and delivery. The recent and most popular theory suggests that severe hypertension increases the limits of cerebral autoregulation and leads to vasodilatation with breakthrough brain edema. Endothelial damage is recognized as a major feature in the pathophysiologic mechanism of preeclampsia and as a possible risk factor for posterior reversible encephalopathy syndrome. Studies suggest that posterior reversible encephalopathy syndrome associated with substantial endothelial damage may develop without a relevant increase in blood pressure(12).

This means that identification of cerebral overflow in patients with preeclampsia with the help of ophthalmic artery Doppler ultrasound may be a marker of the risk of cerebral hemorrhage and may be able to determine the severity in patients with preeclampsia<sup>(13)</sup>. On the other hand, the uteroplacental circulation is crucial for a normal pregnancy outcome. Elevated resistive index (RI), pulsatility index (PI) or systolic to diastolic (S/D) ratios and the presence of a diastolic notch are considered as abnormal uterine artery flow velocity disorders. Impaired trophoblastic invasion of the maternal spiral arteries is associated with increased risk for maternal complications of pregnancy, such as pregnancy-induced hypertension, preeclampsia, placental abruption, poor fetal outcome, intrauterine growth restriction (IUGR) and small for gestational age (SGA) infant<sup>(14)</sup>. Numerous tests, including cold pressor test, have been found to identify mothers at risk of preeclampsia<sup>(15)</sup>.

Doppler ultrasound and color Doppler imaging (CDI) are non-invasive, fast and easy to perform in the evaluation of the uterine and placental blood flow<sup>(16)</sup>. Moreover, they produce highly reliable measures in the ophthalmic artery<sup>(17)</sup>. Besides, there is no evidence that diagnostic ultrasound has produced any harm to humans. Ultrasound is then neither harmful nor expensive, and produces accurate results<sup>(18)</sup>. Vascular changes do occur in preeclampsia and induce hemodynamic changes which can be easily evaluated with Doppler ultrasound. As arterial diseases are almost generalized, it has been postulated that if the uterine artery is affected by

preeclampsia, the ophthalmic artery might be affected as well<sup>(19)</sup>.

This study was performed to compare the resistive index of the uterine artery versus the ophthalmic artery in patients with preeclampsia in Doppler ultrasound. Doppler ultrasound used for early diagnosis of preeclampsia could contribute to reduced morbidity by enabling proper patient management.

### Material and methods

This cross-sectional, observational study was conducted at the Gilani Ultrasound Clinic in Lahore, Pakistan, to compare the resistive index of the uterine artery with that of the ophthalmic artery using ultrasound in pregnant women in the 2<sup>nd</sup> and 3<sup>rd</sup> trimester. The Board of Studies and the Institutional Review Board (IRB) of the University of Lahore approved the study protocol. The sample of 60 participants was calculated with a sample power formula. For the purpose of comparison, 30 normotensive and 30 preeclamptic pregnant women were recruited. Participation was voluntary and written consent was obtained from the patients or their guardians. Patients with chronic hypertension, non-cooperative individuals, patients whose uterine arteries were not visualized due to abdominal gases or large fetus, and patients with any eye disease or impaired vision were excluded. The ultrasound machine used in the study (Toshiba Xerio) was equipped with a linear probe of 7-14 MHz for ophthalmic artery examination and with a curvilinear probe of 3–6 MHz for uterine artery assessment. The American Institute of Ultrasound in Medicine (AIUM) guidelines for obstetrics were observed during examinations<sup>(20)</sup>. The gestational age was calculated in weeks by ultrasound and from the last menstrual cycle. History regarding proteinuria, hypertension, family history and previous history of preeclampsia were taken from the patients. The uterine artery was localized in the supine position from either left or right to avoid fetal parts near the cervix. Spectral waveforms were taken in the longitudinal view, and the measurements were recorded for known preeclamptic and normotensive individuals. The ophthalmic arteries were localized, and spectral waveforms were taken with the help of a linear transducer. RI of the ophthalmic and uterine arteries was calculated, and the acquired data was evaluated with the help of the Statistical Package for the Social Sciences version 24 (SPSS 24, IBM, Armonk, NY, United States of America). Mean and standard deviation values were calculated for age as well as resistive index of the uterine and ophthalmic arteries. The data was normally distributed. Correlations of RI of the uterine and ophthalmic arteries in normal and preeclamptic patients were assessed using the Pearson's correlation.

#### Results

The mean age of the normotensive participants was  $25.40 \pm 3.8$  years with the range from 19 to 35 years.

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**Tab. 1.** Mean and standard deviation values for age as well as resistive index of the uterine and ophthalmic arteries among 30 normotensive and 30 preeclamptic participants

Variables	Study groups	Mean	Std. deviation
Age (years)	Preeclamptic	25.4	3.79
	Normotensive	25.8	3.33
Resistive index of uterine artery	Preeclamptic	0.65	0.09
	Normotensive	0.50	0.08
Resistive index of ophthalmic artery	Preeclamptic	0.63	0.05
	Normotensive	0.71	0.06

**Tab. 2.** Pearson's correlation of the resistive index of the ophthalmic artery and the uterine artery among 30 normotensive and 30 preeclamptic pregnant women

Correlations						
Study groups			RI	RI		
			of uterine	of ophthalmic		
			artery	artery		
Preeclamptic	RI of uterine artery	Pearson Correlation	1	-0.450*		
		Significant (2-tailed)		0.013		
Normotensive	RI of uterine artery	Pearson Correlation	1	0.524**		
		Significant (2-tailed)		0.003		

<sup>\*</sup> Correlation is significant at the 0.05 level (2-tailed).

The mean age of the preeclamptic participants was  $25.9 \pm 3.3$  years with the range from 20 to 33 years. The mean resistive index of the uterine artery was 0.50  $\pm$  0.08 in normotensive participants, and 0.64  $\pm$  0.09 in preeclamptic patients, with the p-value < 0.001. The mean resistive index of the ophthalmic artery was 0.70  $\pm$  0.05 among normotensive participants and 0.63  $\pm$ 0.04 in preeclamptic patients, with the p-value < 0.001; details are summarized in Tab. 1 and Tab. 2. A significant negative correlation of 0.450 with the *p*-value of 0.013 was found between the resistive index of the uterine and ophthalmic arteries in preeclamptic participants, and a positive correlation of 0.524 with the *p*-value of 0.003 was found between the resistive index of the uterine and ophthalmic arteries among normotensive patients, as shown in Tab. 2 and Fig. 1.

# Discussion

Ultrasound techniques are used to predict early signs of preeclampsia by providing blood flow readings in Doppler imaging. The resistive index of the uterine artery is increased in hypertensive patients at risk of preeclampsia. Conversely, the resistive index of the ophthalmic artery is decreased in preeclamptic patients. According to a study by C A de Oliveira *et al.*, conducted in 2013 to determine changes in ophthalmic artery Doppler indices in hypertensive disorders during pregnancy,  $^{(21)}$  it was estimated that the resistive index was  $0.75 \pm 0.05$  among normotensive pregnant women

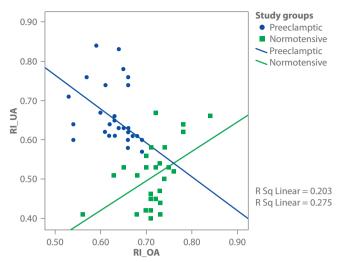
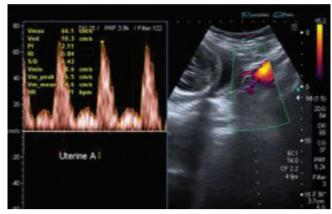


Fig. 1. Scatter plot represents the comparison of the resistive index (RI) of the ophthalmic artery (OA) and the uterine artery (UA) in preeclamptic and normotensive pregnant women

and  $0.63 \pm 0.09$  in preeclamptic women. The study was conducted among seventy-three patients to confirm the status of the resistive index of the ophthalmic artery as a predictor of preeclampsia in the second trimester of pregnancy. All the patients at risk of preeclampsia were included. They were evaluated with Doppler imaging of the ophthalmic artery at 24 to 28 weeks of pregnancy and monitored until the end of pregnancy to verify the occurrence of preeclampsia. ROC curves were created to determine the predictive characteristics of the ophthalmic artery resistive index. Of the observed patients, 14 developed preeclampsia while 59 remained normotensives until delivery. The patients who developed preeclampsia had lower ophthalmic artery resistive indices  $(0.682 \pm 0.028)$  than those who remained normotensive (0.700  $\pm$  0.029; the p-value was within the range of significance: p-value = 0.044). Low resistance in the ophthalmic artery was considered a risk factor for the development of preeclampsia with the area under the curve of 0.694 and the confidence interval from 0.54 to 0.8. It was concluded that low resistance of the ophthalmic artery was related to preeclampsia<sup>(22)</sup>. In the present study, the mean resistive index of the ophthalmic artery measured in normotensive pregnant women and in preeclamptic women was very similar to that reported in the aforementioned study, as shown in Tab. 1 and Fig. 1, Fig. 2 and Fig. 3.

According to previous studies, Doppler indices for the uterine artery decreased with advancing gestational age. Ademola Joseph Adekanmi *et al.* compared Doppler indices of the uterine and umbilical arteries among healthy Nigerian women with singleton pregnancies in the second and third trimesters. The authors found that normal values of the resistive index were  $0.53 \pm 0.18$  in the second trimester and  $0.48 \pm 0.20$  in the third trimester<sup>(23)</sup>. In another study by H. Valensise, the resistive index of the uterine artery above 0.58 was considered abnormal<sup>(24)</sup>. In the present study, the mean resistive index of the uterine artery in normotensive

<sup>\*\*</sup> Correlation is significant at the 0.01 level (2-tailed).



**Fig. 2.** Ultrasound image of a preeclamptic female at 33 weeks of pregnancy representing high RI (0.84) of the uterine artery

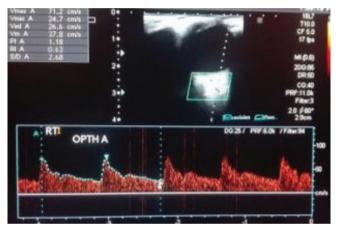


Fig. 3. Ultrasound image of a preeclamptic female at 33 weeks of pregnancy representing low RI (0.63) of the ophthalmic artery

pregnant women was 0.50 and in preeclamptic women: 0.64, which is in line with the results of the above-mentioned study by H. Valensise. The results of this study prove that the resistive index of the ophthalmic artery

is significantly correlated with the resistive index of the uterine artery in both normotensive and preeclamptic patients. A negative linear correlation was found, i.e. the resistive index increases in the uterine artery and decreases in the ophthalmic artery in preeclampsia. Blood flow in both uterine artery and ophthalmic artery was almost equally affected by preeclampsia, except for a few cases where any of the two remained normal. Both vessels can be used to predict the severe problem like preeclampsia to reduce maternal mortality and morbidity. In pregnant women, especially in the second and third trimesters, long examinations during which the sonographer manipulates the transducer over the abdomen to locate the uterine artery, may be uncomfortable. The ophthalmic artery can be easily located with the transducer, and the readings are obtained in a short time. That is why, in this research, we decided to compare the resistive index of the ophthalmic artery in confirmed preeclamptic and normotensive individuals.

#### Conclusions

There was a significant negative correlation between the resistive index of the uterine and ophthalmic arteries in the patients with preeclampsia, and a significant positive correlation was found for normotensive individuals. Preeclampsia could be easily diagnosed by Doppler ultrasound due to hemodynamic changes that occur in response to vascular changes in the ophthalmic and uterine arteries.

#### Conflict of interest

Authors do not report any financial or personal connections with other persons or organizations, which might negatively affect the contents of this publication and/or claim authorship rights to this publication.

#### References

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- 1. Say L, Chou D, Gemmill A, Tunçalp Ö, Moller A-B, Daniels J *et al.*: Global causes of maternal death: a WHO systematic analysis. Lancet Glob Health 2014; 2: e323–e333.
- Hobson SR, Lim R, Mockler JC, Gurusinghe S, Wallace EM: Role of activin a in the pathogenesis of endothelial cell dysfunction in preeclampsia. Methods Mol Biol 2018; 1710: 39–52.
- Rigó J Jr, Nagy B, Fintor L, Tanyi J, Beke A et al.: Maternal and neonatal outcome of preeclamptic pregnancies: the potential roles of factor V Leiden mutation and 5, 10 methylenetetrahydrofolate reductase. Hypertens Pregnancy 2000; 19: 163–172.
- Duckitt K, Harrington D: Risk factors for pre-eclampsia at antenatal booking: systematic review of controlled studies. BMJ 2005; 330: 565.
- Jeyabalan A: Epidemiology of preeclampsia: impact of obesity. Nutr Rev 2013; 71 Suppl 1: S18–S25.
- Redman CW, Sacks GP, Sargent IL: Preeclampsia: an excessive maternal inflammatory response to pregnancy. Am J Obstet Gynecol 1999; 180: 499–506.
- Medeiros LT, Peraçoli JC, Bannwart-Castro CF, Romão M, Weel IC, Golim MA et al.: Monocytes from pregnant women with pre-eclamp-

- sia are polarized to a M1 phenotype. Am J Reprod Immunol 2014; 72: 5-13.
- Haram K, Mortensen JH, Mastrolia SA, Erez O: Disseminated intravascular coagulation in the HELLP syndrome: how much do we really know? J Matern Fetal Neonatal Med 2017; 30: 779–788.
- Barden AE, Beilin LJ, Ritchie J, Walters BN, Michael C: Does a predisposition to the metabolic syndrome sensitize women to develop preeclampsia? J Hypertens 1999; 17: 1307–1315.
- 10. Sibai BM: Diagnosis and management of gestational hypertension and preeclampsia. Obstet Gynecol 2003; 102: 181–192.
- Hobson SR, Gurusinghe S, Lim R, Alers NO, Miller SL, Kingdom JC et al.: Melatonin improves endothelial function in vitro and prolongs pregnancy in women with early-onset preeclampsia. J Pineal Res 2018; 65: e12508.
- Mazoor I, Bacha R, Gilalni SA: The significance of doppler ultrasound in the causes of varicose veins. J Vasc Ultrasound 2017; 41: 159–164.
- Bartynski W: Posterior reversible encephalopathy syndrome, part 2: controversies surrounding pathophysiology of vasogenic edema. AJN Am J Neuroradiol 2008; 29: 1043–1049.

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- Kurjak A, Chervenak FA: Donald School Textbook of Ultrasound in Obstetrics & Gynaecology: JP Medical Ltd; 2017.
- 15. Jani PS, Patel UM, Gandhi MR, Thakor NC, Kakani C: Placental laterality and uterine artery resistance as predictor of preeclampsia: a prospective study at GMERS Medical College, Dharpur-Patan, North Gujarat, India. Int J Res Med Sci 2017; 3: 1484–1487.
- Angelova M, Todorov I, Kovachev E: Early prognostic factors for the progress of preeclampsia – our experience in the period 2010–2011. Open Access Maced J Med Sci 2016; 4: 420–422.
- 17. Harris A, Williamson TH, Martin B, Shoemaker JA, Sergott RC, Spaeth GL *et al.*: Test/Retest reproducibility of color Doppler imaging assessment of blood flow velocity in orbital vessels. J Glaucoma 1995; 4: 281–286.
- ter Haar G: The new British Medical Ultrasound Society Guidelines for the safe use of diagnostic ultrasound equipment. Ultrasound 2010; 18: 50–51.
- Manzoor I, Bacha R, Gilani SA: Diagnostic accuracy of sonoelastography in different diseases. J Ultrason 2018; 18: 29–36.

- Zaman S, Gilani SA, Bacha R, Manzoor I, Hasan ZU: Correlation between portal vein diameter and craniocaudal length of the spleen. J Ultrason 2019; 19: 276–281.
- de Oliveira CA, de Sá RAM, Velarde LGC, daSilva FC, doVale FA, Netto HC: Changes in ophthalmic artery Doppler indices in hypertensive disorders during pregnancy. J Ultrasound Med 2013; 32: 609–619.
- Aquino LOd, Leite HV, Cabral AC, Brandão AH: Doppler flowmetry of ophthalmic arteries for prediction of pre-eclampsia. Rev Assoc Med Bras 2014; 60: 538–541.
- Adekanmi AJ, Roberts A, Adeyinka AO, Umeh EO, Anor F, Odo JC et al.: Normal second and third trimester uterine and umbilical doppler indices among healthy singleton gestation Nigerian women. West Afr J Radiol 2017: 24: 1.
- 24. Valensise H, Bezzeccheri V, Rizzo G, Tranquilli AL, Garzetti G, Romanini C: Doppler velocimetry of the uterine artery as a screening test for gestational hypertension. Ultrasound Obstet Gynecol 1993; 3: 18–22.

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