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Article

Fetal Splenic Artery Pulsatility Index May Predict the Need for Neonatal Intensive Care in Gestational Diabetes Class A1 Cases

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Abstract: The fetal splenic artery pulsatility index (PI) is a parameter that reflects fetal well-being and has been used as a predictor of adverse pregnancy outcomes. The aim of this study was to investigate the predictive value of splenic artery PI in GDM class A1 cases for NICU admission. In this prospective case controlled study, only sixty single pregnancy cases diagnosed with gestational diabetes mellitus class A1 were evaluated. Fetal splenic artery Doppler parameters Peak systolic velocity (PSV), Pulsatility index (PI), Resistivity index (RI) and End diastolic velocity (EDV) were measured in all cases. The rate of requirement of need of neonatal intensive care unit was noted. In cases with need fetal intensive care, fetal splenic PI index was found to be statistically significantly lower than in healthy cases without it (0.94 ± 0.29 vs. 1.70 ± 0.53 , respectively, $p < 0.001$, Student-t test.). When the fetal splenic PI cutoff value was selected as 1.105 cm^3 , the sensitivity was calculated as 97.9% and the specificity as 58.3% for predicting the need for fetal intensive care (AUC 0.968, $p < 0.001$, 95% CI 0.929–0.998). The use of low fetal splenic artery PI parameter is a significant and good indicator for predicting the need for fetal intensive care according to the binary logistic regression analysis result ($p = 0.006$). This study suggests that evaluation of fetal splenic artery Doppler in mothers with GDM may be used to predict neonates requiring NICU. Therefore, we recommend obstetricians to use this simple, rapid, and valuable evaluation of fetal splenic artery Doppler, and to alert the neonatologist that NICU may be required.

Keywords: gestational diabetes; neonatal intensive care unit; prenatal sonography; splenic artery doppler

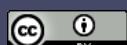
1. Introduction

Gestational diabetes (GDM), characterized by high blood sugar levels during pregnancy that affects both maternal and newborn outcomes, is a common complication of pregnancy [1]. GDM is often classified into two categories, Class A1 and Class A2, with Class A1 being the milder form that can typically be managed with diet and lifestyle modifications. Despite being a milder form, GDM Class A1 cases can still present challenges in predicting and managing neonatal outcomes, particularly in identifying neonates who may require neonatal intensive care unit (NICU) admission [2].

Fetal Doppler ultrasound, a non-invasive imaging technique, is a valuable tool in assessing fetal well-being and identifying potential complications [3]. The fetal splenic artery PI, Doppler-derived parameter, has been used to evaluate fetal hemodynamics and perfusion [4,5]. Some studies have demonstrated a correlation between abnormal fetal splenic artery PI and adverse neonatal outcomes in various pregnancy complications, such as fetal growth restriction [6–10].

In this prospective study, it was investigated whether fetal splenic artery Doppler measured in cases followed for gestational diabetes class A1 predicts the need for neonatal intensive care. By identifying neonates at risk of requiring NICU care, healthcare professionals can better prepare and allocate resources, ensuring optimal care for both mothers and their babies.

2. Materials and Methods



In this study, 75 single pregnancy cases diagnosed with gestational diabetes mellitus class A1 who applied to our high-risk pregnancy outpatient clinic were evaluated. in one year period. Ethics committee approval was obtained.

The study group were aged 18-40 years, who presented at the polyclinic for routine antenatal follow-up at third trimester of pregnancy (>28 weeks) and had the diagnosis of gestational diabetes class A1. Only the pregnant women at 30-34 weeks of gestation (including the limit weeks) were included in the study. In the first examination of the pregnant cases, it was determined that they did not contain any high-risk pregnancy factors other than gestational diabetes, and liver, kidney, thyroid and hematological test results were within normal limits. Gestational diabetes mellitus (GDM) diagnosis was made with 75 g oral glucose tolerance test (OGTT). The 75 gram OGTT was used for diagnosing diabetes in pregnant individuals. The two-hour 75 gram OGTT was diagnostic of GDM when one glucose value was elevated. According to this test, fasting blood glucose was measured following an 8-hour fasting period, and then a solution containing 75 grams of glucose was drunk. Blood glucose was measured 1 hour and two hours after drinking the solution. GDM was diagnosed when fasting blood glucose was 92 or 1st hour blood glucose was 180 or second hour blood glucose was 153 mg/dL and above. Only the cases that were euglycemic with diet were accepted as GDM Class A1.

Cases with any systemic disease, use of drugs other than vitamins and iron, multiple pregnancy, smoking, biochemical/hormonal/hematological test disorders, and cases diagnosed with congenital fetal anomaly were not included in the study. Cases with abnormal uterine, umbilical, middle cerebral artery Doppler findings were not included in the study. 15 cases not fulfilled the inclusion criteria were excluded from the study. 11 cases who did not come for pregnancy follow-ups and 4 cases who developed any pregnancy complications were excluded from the study.

In the study, all participating women provided informed consent. Key information collected included age, gravidity, parity, and body mass index (BMI), as well as sonographic measurements of fetal biometry, such as biparietal diameter (BPD), abdominal circumference (AC), femur length (FL), and estimated fetal weight (EFW). Gestational age was verified using first trimester ultrasound data.

During the Doppler assessment, the patient was positioned on the examination table in a supine posture. It was measured when the fetus was inactive. A standard AC axial section view was obtained. Stomac pocket was visualized. Color Doppler was placed on the upper part of the stomach pocket and the splenic artery was visualized at the entrance to the spleen. Later, it was observed that the trace progressed towards the truncus celiacus and abdominal aorta. Color Doppler was placed at the entrance to the spleen with a 0-degree insomination angle, without the splenic artery entering the spleen. Fetal splenic artery Doppler parameters Peak systolic velocity (PSV), Pulsatility index (PI), Resistivity index (RI) and End diastolic velocity (EDV) were measured in all cases (Figures 1 and 2).

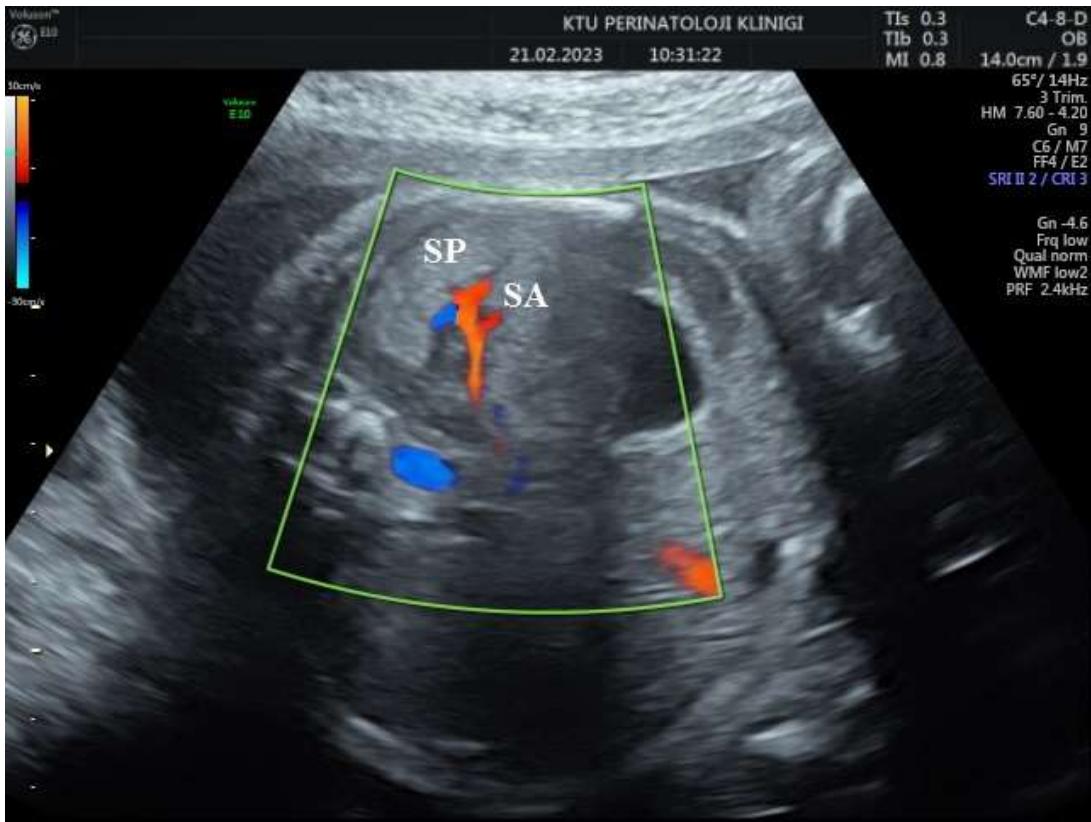


Figure 1. Two-dimensional ultrasonographic imaging of the 30-week-old fetus shows fetal spleen (SP), splenic artery (SA).

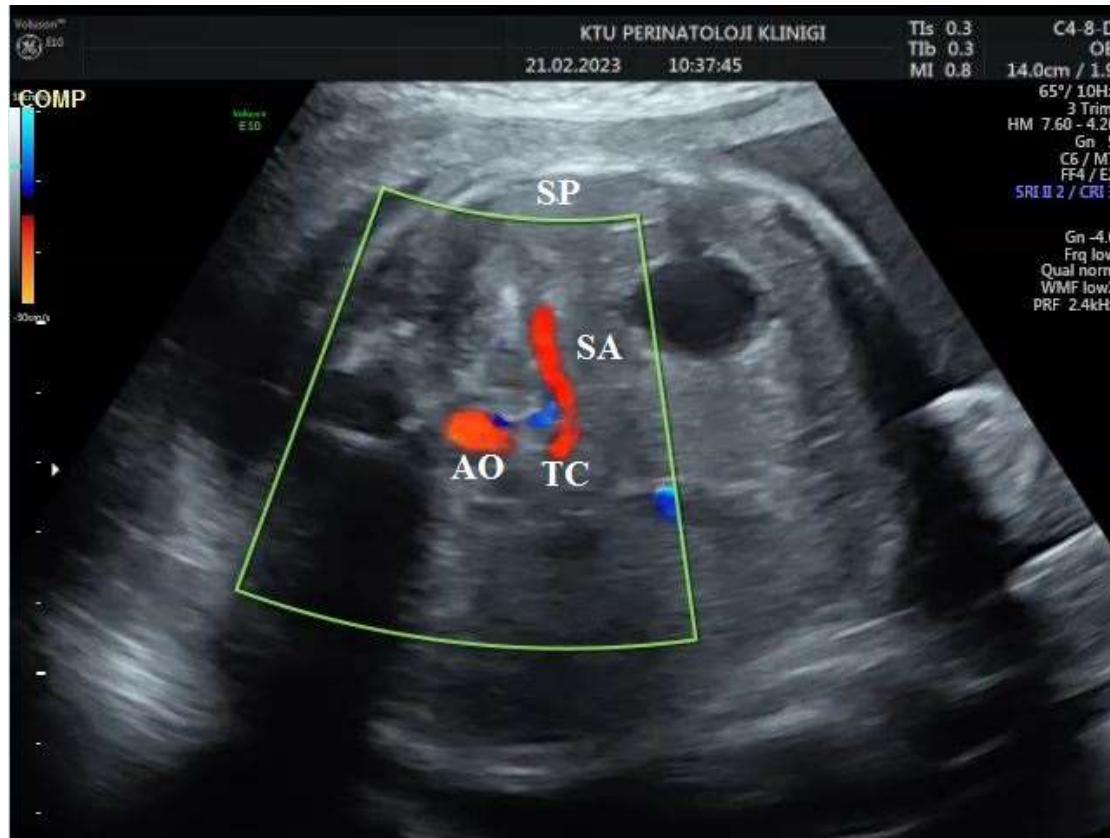


Figure 2. Two-dimensional ultrasonographic imaging in the axial plane of the 30-week-old fetus shows the origin of fetal splenic artery from truncus coeliacus (TC; truncus coeliacus, AO; Aorta, SA; Splenic artery, SP; Fetal Spleen).

All sonographic measurements were performed via the 2016 model VOLUSON GE E10 sonograph unit. All sonographic measurements were made at least twice by an experienced clinician. All cases were followed up until the end of pregnancy, and the birth data of the mother and the baby (fetal birthweight, fetal umbilical cord blood gas analysis) and the need for neonatal intensive care were recorded.

A total of 12 cases required neonatal intensive care due to respiratory distress, and their data were compared to those not needing intensive care.

Statistical Analysis

Data were analyzed using SPSS 13 software. The Levene test assessed variance homogeneity, while the Kolmogorov-Smirnov test evaluated normal distribution compatibility. Chi-square and Student's t-tests were employed for statistical analysis. To identify fetal splenic artery Doppler parameters for predicting neonatal intensive care requirements, ROC analysis was conducted. All data are presented as mean \pm standard deviation or percentage. The ability of fetal doppler parameters to predict neonatal intensive care needs was assessed using binary logistic regression analysis.

The correlation between the parameters was evaluated with the help of Pearson correlation test. The situation where the p value was below 0.05 was considered significant.

Ethical Approval

The study protocol was approved by current university ethic board (number 2020/37, date 28/01/2020).

3. Results

The data of 60 cases in total were evaluated. The demographic and sonographic factors of cases who need fetal intensive care and those not are shown in Table 1.

Table 1. The demographic and sonographic factors of cases who had the need for neonatal intensive care and who not.

Demographic/ Sonographic factors	Need for neonatal intensive care (n=12)	No need for neonatal intensive care (n=48)	p
Age (years)	32.17 \pm 4.91	31.38 \pm 5.01	0.627
Gravida (no.)	3.00 \pm 2.09	2.81 \pm 1.62	0.737
Parity (no.)	2.50 \pm 1.69	1.59 \pm 1.19	0.078
Body mass index (kg/m ²)	34.11 \pm 8.85	31.22 \pm 5.65	0.166
Gestational week at measurement	32.73 \pm 0.94	32.30 \pm 1.14	0.227
EFW at measurement (gr)	2176.75 \pm 627.65	2186.42 \pm 636.63	0.963
Gestational week at birth	38.25 \pm 1.14	38.02 \pm 2.12	0.715
Fetal Splenic artery PSV	34.58 \pm 9.04	37.02 \pm 14.78	0.587
Fetal Splenic artery S/D	4.16 \pm 0.49	4.23 \pm 1.68	0.796
Fetal Splenic artery PI	0.94 \pm 0.29	1.70 \pm 0.53	<0.001
Fetal Splenic artery RI	0.67 \pm 0.20	0.75 \pm 0.20	0.260
Fetal Splenic artery diameter	0.40 \pm 0.26	0.49 \pm 0.63	0.636
Caesarean section delivery (%)	41.7% (5/12)	41.7% (20/48)	0.624 ^a
Fetal gender (F/M)	5/7	23/25	0.467 ^a
Infant birthweight (gr.)	3397.08 \pm 516.25	3195.00 \pm 577.31	0.273
Fetal umbilical artery pH	7.28 \pm 0.40	7.36 \pm 0.04	<0.001
Fetal umbilical artery BE	-1.34 \pm 1.36	-1.18 \pm 1.81	0.782
APGAR score at 1 st minute	6.33 \pm 0.65	6.79 \pm 1.35	0.188
AGRAR score at 5 th minute	8.25 \pm 0.97	8.50 \pm 0.77	0.344

PSV; Peak systolic velocity, S/D; Systol/Diastole, PI; Pulsatility index, RI; Resistivity index, BE; Base excess. The Student t-test and aFisher's Exact Chi-square test were used in the statistical analyses.

Data are presented as mean \pm standard deviation values or as number of cases and percentage.

In cases with need fetal intensive care, fetal splenic PI index was found to be statistically significantly lower than in healthy cases without it (0.94 ± 0.29 vs. 1.70 ± 0.53 , respectively, $p < 0.001$, Student-t test,). When the fetal splenic PI cutoff value was selected as 1.105, the sensitivity was calculated as 97.9% and the specificity as 58.3% for predicting the need for fetal intensive care (AUC 0.968, $p < 0.001$, 95% CI 0.929-0.998, Figure 3). The use of low fetal splenic artery PI parameter is a significant and good indicator for predicting the need for fetal intensive care according to the binary logistic regression analysis result ($p = 0.006$).

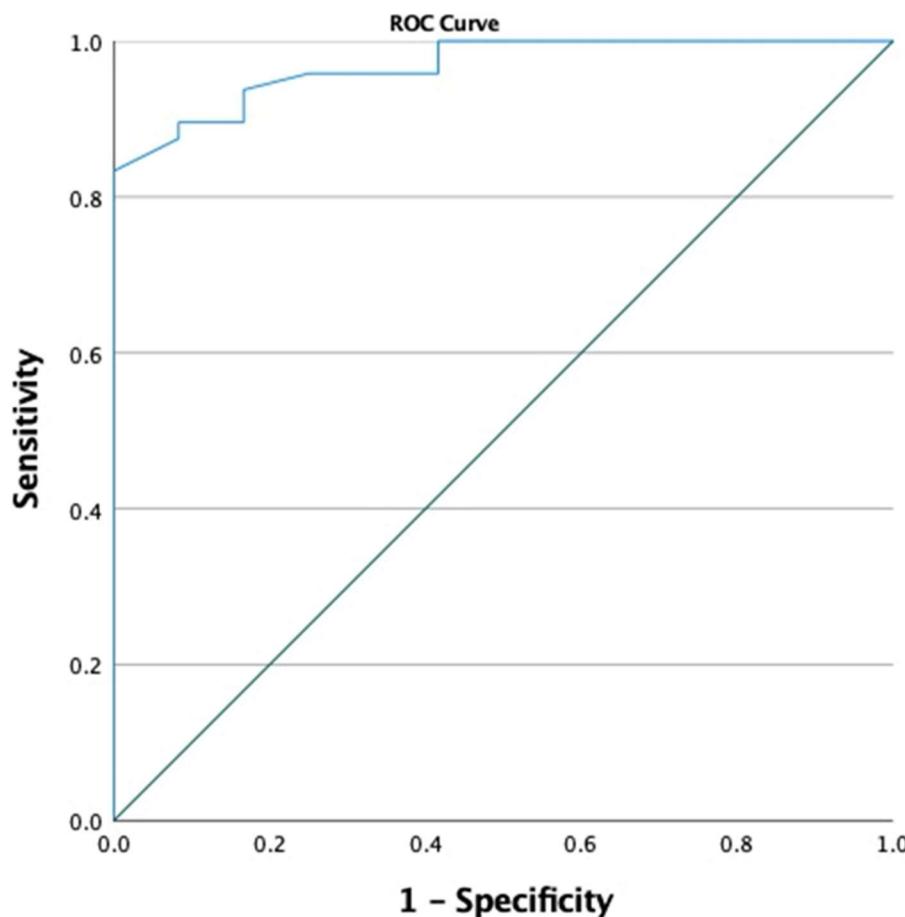


Figure 3. Receiver operating curve analysis data showing the relationship between calculated fetal splenic artery PI value and need for neonatal intensive care are presented.

A moderate positive correlation was found in terms of fetal umbilical cord pH and BE at delivery and prenatal fetal splenic doppler PI in all cases (Pearson correlation, $r=0.334$, $p=0.009$, Pearson Correlation test).

4. Discussion

This study evaluated the relationship between fetal splenic artery Doppler parameters and the need for NICU in pregnant women with GDM class A1. The results showed that a lower fetal splenic artery PI was significantly associated with an increased need for NICU, with a sensitivity of 97.9% and specificity of 58.3% for predicting the need for neonatal intensive care when the fetal splenic PI was taken as 1.105. The results also showed a moderate positive correlation between prenatal fetal splenic Doppler PI and fetal umbilical cord pH and BE at delivery.

The importance of fetal Doppler parameters in predicting adverse neonatal outcomes has been well-documented in the literature [7,11]. Previous studies have focused on umbilical and middle cerebral artery Doppler measurements in high-risk pregnancies, including those with GDM [12]. Limited research has been conducted on the relationship between fetal splenic artery Doppler parameters and neonatal outcomes, particularly in pregnancies complicated by GDM class A1. In the past decade, abnormal Doppler velocity patterns in various fetal vessels and their correlation with fetal outcomes have been extensively studied [13-15]. Recently, the pathophysiology of blood redistribution in relation to fetal growth deprivation has been evaluated [16,17]. Epidemiological evidence suggests that changes in blood distribution due to fetal growth deprivation may increase the risk of myocardial damage and abnormal liver function in newborns [18,19]. Decreased resistance in the splenic artery has been suggested as an indicator of fetal distress, and a decreased fetal liver circulation has recently been proposed as a predictor for the prognosis of fetal growth restriction [7,20].

The assessment of fetal well-being is crucial in prenatal care, particularly for high-risk pregnancies. Fetal splenic artery Doppler ultrasonography is a non-invasive technique that has been increasingly utilized for evaluating fetal well-being and predicting adverse perinatal outcomes. The technique measures blood flow velocities in various vessels, including the splenic artery, which supplies blood to the spleen, an organ essential for fetal immune system development. The Doppler indices, such as resistance index (RI), pulsatility index (PI), and systolic/diastolic ratio (S/D), can help assess blood flow and detect abnormalities in fetal circulation [21].

Abuhamad et al. (1992) were the first to report the use of Doppler velocimetry in evaluating the main splenic artery of fetuses [6]. They noted a reduction in the splenic artery RI in a subset of fetuses classified as small for gestational age (SGA). The authors theorized that the associated chronic hypoxia led to the release of erythropoietin, which increased splenic flow to support increased erythropoiesis. This reduction in RI in some SGA fetuses was later confirmed by Capponi et al. in a subsequent study [22].

Several theories attempt to explain the reduced splenic artery pulsatility index (PI) observed in cases of fetal growth restriction. Recent research has shown that compensatory vasodilation in the splenic artery helps maintain low venous perfusion to the fetal liver [20,23]. The exact connection between a decreased splenic artery PI and fetal growth restriction remains unclear. However, one study discovered that a lower splenic artery PI correlated with higher perinatal mortality rates, lower Apgar scores, and metabolic acidosis. This suggests that severe fetal deprivation may lead to more significant hemodynamic changes in the spleen, potentially identifying fetuses at an increased risk of perinatal death [24].

In a study, it was determined that fetal splenic artery PI values were lower in fetuses small for gestational age compared to normal fetuses. This supports the argument that vascular resistance is lower in SGA fetuses [25].

In another study, fetal splenic artery Doppler measurement was performed in pregnancies with late-onset fetal growth restriction and similar results were reported. The researchers found that decreased splenic artery PI was significantly and positively correlated with a higher probability of experiencing adverse obstetric outcomes [3].

Neonatal outcomes of splenic artery Dopplers in gestational diabetic patients have not yet been studied. As a result, our study is the first of its kind in the literature. The current study demonstrates that fetal splenic artery PI could serve as a valuable indicator for predicting the need for neonatal intensive care in GDM class A1 pregnancies. A possible explanation for this finding could be that a lower PI reflects increased placental resistance and, consequently, impaired fetal blood flow, which could negatively impact fetal growth and wellbeing. This notion is supported by the observed moderate positive correlation between prenatal fetal splenic Doppler PI and fetal umbilical cord pH and BE at delivery, suggesting a relationship between splenic artery PI and fetal acid-base status.

The results of this study are consistent with previous findings that have demonstrated the utility of fetal Doppler measurements in predicting neonatal outcomes in high-risk pregnancies. However, the current study adds to the existing body of knowledge by specifically focusing on GDM class A1

pregnancies and examining the association between fetal splenic artery Doppler parameters and neonatal outcomes.

The most important limitation of our study is the small number of cases. Additionally, the study did not examine the long-term neonatal outcomes beyond the need for neonatal intensive care. Future research with larger sample sizes and follow-up of neonatal outcomes is warranted to further validate and expand upon these findings.

5. Conclusions

We believe that using fetal splenic artery Doppler measurements in mothers with gestational diabetes mellitus (GDM) can help predict which neonates may require NICU. We aimed to determine the best method for predicting NICU needs for infants born to mothers with GDM. As our study suggests, the evaluation of fetal splenic artery Doppler could be a brief and valuable answer to this question.

We kindly recommend this simple, rapid, and valuable evaluation of fetal splenic artery Doppler to obstetricians. By doing so, it may be a safe approach to alert the neonatologist that NICU might be required, allowing for better preparedness and resource allocation.

The results of this study suggest that fetal splenic artery PI could be a valuable indicator for predicting the need for neonatal intensive care in pregnant women with GDM class A1. This finding could have potential clinical implications, as it may help guide prenatal care and counseling for women with GDM, thereby potentially improving neonatal outcomes. Further research is needed to confirm and extend these findings in larger populations and to explore the long-term neonatal outcomes associated with fetal splenic artery Doppler parameters.

6. Patents

Authors contribution: All authors (MA, HA, ESGG, and SG) participated and contributed to data collection, literature review, and editing the manuscript.

Funding: Self-funding.

Institutional Review Board statement: The study protocol was approved by current university ethic board (number 2020/37, date 28/01/2020).

Informed Consent Statement: Informed consent was obtained from all subjects involved in the study.

Data Availability Statement: Any data set is available on request to the Corresponding author.

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Conflicts of Interest: The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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