

# Uterine Artery Doppler Velocimetry of Uterine Leiomyomas in Nigerian Women

## Dopplervelocimetria da artéria uterina de leiomiomas uterinos em mulheres nigerianas

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Abstract **Objective** To describe the blood flow velocities and impedance indices changes in the uterine arteries of leiomyomatous uteri using Doppler sonography. Methods This was a prospective, case-control study conducted on 140 premenopausal women with sonographic diagnosis of uterine leiomyoma and 140 premenopausal controls without leiomyomas. Pelvic sonography was performed to diagnose and characterize the leiomyomas. The hemodynamics of the ascending branches of both main uterine arteries was assessed by Doppler interrogation. Statistical analysis was performed mainly using non-parametric tests. **Results** The median uterine volume of the subjects was 556 cm<sup>3</sup>, while that of the controls was 90.5 cm<sup>3</sup> (p < 0.001). The mean peak systolic velocity (PSV), end-diastolic velocity (EDV), time-averaged maximum velocity (TAMX), time-averaged mean velocity (Tmean), acceleration time (AT), acceleration index (AI), diastolic/systolic ratio (DSR), diastolic average ratio (DAR), and inverse pulsatility index (PI) were significantly higher in the subjects (94.2 cm/s, 29.7 cm/s, 49.1 cm/s, 25.5 cm/s, 118 ms, 0.8, 0.3, 0.6, and 0.8 respectively) compared with the controls (54.2 cm/s, 7.7 cm/s, 20.0 cm/s, 10.0 cm/s, 92.0 ms, 0.6, 0.1, 0.4, and 0.4 respectively); p < 0.001 for all values. **Keywords** Conversely, the mean PI, resistivity index (RI), systolic/diastolic ratio (SDR) and leiomyoma impedance index (ImI) of the subjects (1.52, 0.70, 3.81, and 3.81 respectively) were ► uterine artery significantly lower than those of the controls (2.38, 0.86, 7.23, and 7.24 respectively); embolization p < 0.001 for all values. Doppler **Conclusion** There is a significantly increased perfusion of leiomyomatous uteri that is ultrasonography most likely due to uterine enlargement. Resumo **Objetivo** Descrever as velocidades do fluxo sanguíneo e as alterações dos índices de impedância nas artérias uterinas leiomiomatoso utilizando a ultrassonografia Doppler. Métodos Estudo prospectivo, caso-controle, realizado em 140 mulheres pré-menopáusicas com diagnóstico ultrassonográfico de leiomioma uterino e em 140 controles na pré-menopausa sem leiomiomas. A ultrassonografia pélvica foi realizada para

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diagnosticar e caracterizar os leiomiomas. A hemodinâmica dos ramos ascendentes de ambas as artérias uterinas principais foi avaliada por meio de interrogatório Doppler. A análise estatística foi feita principalmente por meio de testes não paramétricos.

**Resultados** A média do volume uterino dos das pacientes foi de 556 cm<sup>3</sup>, enquanto a dos controles foi de 90,5 cm<sup>3</sup> (p < 0,001). A média de velocidade de pico sistólico (VPS), a velocidade diastólica final (VDF), a velocidade máxima do tempo médio (VMTM), a velocidade média do tempo médio (VMdTM), o tempo de aceleração (TA), o índice de aceleração (IA), a relação diástole/sístole (RDS), a proporção diastólica média (PDM) e o índice de pulsatilidade (IP) inversa foram significativamente maiores em pacientes (94,2 cm/s, 29,7 cm/s, 49,1 cm/s, 25,5 cm/s, 118 ms, 0,8, 0,3, 0,6 e 0,8, respectivamente) do que nos controles (54,2 cm/s, 7,7 cm/s, 20,0 cm/s, 10,0 cm/s, 92,0 ms, 0,6, 0,1, 0,4 e 0,4, respectivamente); p < 0,001 para todos os valores. Por outro lado, o IP médio, o índice de resistividade (IR), a relação sístole/diástole (RSD) e o índice de impedância (II) nas pacientes (1,52, 0,70, 3,81 e 3,81, respectivamente) foram significativamente mais baixos do que os dos controles (2,38, 0,86, 7,23 e 7,24, respectivamente); p < 0,001 para todos os valores.

#### Palavras-chave ► leiomioma

- embolização da artéria uterina
- ultrassonografia com Doppler

**Conclusão** Existe um aumento significativo da perfusão dos úteros leiomiomatosos, que provavelmente se deve ao alargamento uterino.

#### Introduction

Leiomyomas/fibroids are benign uterine tumors **that are more common in black people**. Gray-scale ultrasonography is a reliable method of diagnosing these tumors, and it can be supplemented with Doppler mode to provide additional hemodynamic information.

Though the pulsatility index (PI) is currently the preferred Doppler index in Obstetrics and Gynecology because it is sensitive to alterations in waveform shape and it analyzes the entire cardiac cycle,<sup>1</sup> several other useful Doppler indices have been documented in the literature. Previous studies reported increased peak systolic velocity (PSV) and time-averaged maximum velocity (TAMX), as well as diminished pulsatility index (PI), resistivity index (RI), and systolic-diastolic ratio (SDR) in myomatous uteri compared with controls.<sup>2–6</sup>

The purposes of this study are: to compare the changes within the main uterine artery of myomatous uteri and normal uteri using several Doppler indices (in addition to the traditional ones); to explore the relationship between these changes in uterine volume and the clinical symptoms; and to compare the main uterine artery Doppler changes in women with recurrent leiomyoma (after a previous myomectomy) and those with primary tumors (without previous surgical or medical intervention).

## Methods

This prospective, non-randomized, case-control study was performed over a one-year period in the Department of Radiology of our institution as approved by the Ethics in Research Committee of the hospital. The study participants were consecutively recruited, and consisted of 140 premenopausal women with sonographic features of uterine leiomyomas, as previously described by other authors,<sup>4,7,8</sup> and 140 premenopausal controls with normal leiomyoma-free uteri **and normal endometrial stripe thickness**. Recurrent leiomyoma was defined as the presence of a dominant leiomyoma larger than 4 cm in size following previous myomectomy.<sup>9</sup> The exclusion criteria were: pregnancy, post-menopausal status (cessation of menses for more than 12 months), recent childbirth (less than 1 year prior to presentation), **adenomyosis/ endometriosis**, and coexisting pelvic pathologies, like ovarian tumors and pelvic inflammatory disease. Informed consent was obtained from all participants.

The participants' demographic and gynecologic histories were obtained, including age, parity, and history of previous myomectomy. Their body mass index was also determined.

All the participants underwent sonography via the transabdominal route using MINDRAY real-time ultrasound machine model DC-7 (Shenzhen Mindray Bio-medical Electronics, Nanshan, Shenzhen, China) with a convex transducer (with a frequency of 3.8-5.0 MHz) and Doppler functionality.

Transabdominal sonography was performed by applying the transducer to the abdominopelvic region to scan the uterus in at least two planes (transverse and longitudinal planes). The following sonographic parameters were recorded: the uterine length, the anteroposterior diameter, and the transverse diameter, which were used to estimate the uterine volume using the ellipsoid formula. The length of the uterus was measured in the longitudinal plane from the fundus to the external cervical os; its anteroposterior diameter was measured perpendicular to the plane of the length, while the transverse diameter was measured from cornu to cornu on a transverse image. The volume of the largest leiomyoma nodule was similarly determined using the prolate ellipse formula. The uterine volume and dominant leiomyoma volume were divided into large and small if they were larger or smaller than 200 cm<sup>3</sup> respectively.<sup>5</sup> The other sonographic parameters recorded included: leiomyoma type (intramural, subserous, submucous, panmural, pedunculated, or in more than one site) and presence of degenerative changes (cystic, calcific, or mixed).

The Doppler interrogation was performed as previously described by other authors.<sup>10,11</sup> The transducer was placed longitudinally in the midline above the symphysis pubis. A longitudinal section of the uterus and the cervical canal was obtained to identify the internal cervical os. The transducer was then moved from side to side to the lateral border of the uterus until the para-cervical vascular plexus was seen. The color Doppler function was activated, and the uterine artery was identified at the level of the junction between the uterine body and the cervix, as it starts to make its ascent to the uterine body. Measurements were taken at this point, before the uterine arteries branch into the arcuate arteries. The pulsed Doppler gate (sample volume = 1 mm) was placed in the center of the vessel, and a Doppler insonation angle of less than 60 degrees was used to obtain a measurement line parallel to the arterial wall. The Doppler signal was updated until there was a clear and consistent waveform. Doppler velocimetry was repeated three times each in both the right and left uterine arteries, with an interval of at least 2 minutes between the measurements. The average value of the two arteries was used for analysis. The measurements were obtained after the visualization of at least three to five consecutive arterial waveforms of similar amplitude. Blood velocity waveforms with the lowest RI or highest velocities were recorded.<sup>12–14</sup> A spectral analysis was performed electronically (auto-trace) from a smooth curve fitted to the average waveform over three to five cardiac cycles to obtain the following parameters: peak systolic velocity (PSV), enddiastolic velocity (EDV), systolic/diastolic ratio (SDR or S/D ratio = PSV/EDV),<sup>6</sup> resistivity index (RI= [PSV – EDV]/PSV),<sup>15</sup> pulsatility index (PI = [PSV - EDV]  $\div$  [{PSV + EDV}]/2}),<sup>16</sup> time-averaged maximum velocity (TAMX), time-averaged mean velocity (Tmean), and acceleration time (AT). The diastolic/systolic ratio or end-diastolic ratio (DSR or D/S ratio = EDV/PSV),<sup>17</sup> impedance index (ImI =  $[S \times D]/D^2$ ),<sup>18</sup> inverse PI [= 1/PI],<sup>19</sup> diastolic average ratio (DAR = EDV/ TAMX),<sup>20</sup> and the acceleration index  $(AI = PSV/AT)^{21}$  were also calculated from the electronically generated indices.

The color and pulsed Doppler parameters, including the high pass filter, the sample volume and the velocity scale were optimized for the detection of slow flow.<sup>21,22</sup> Low filtration (color wall filter of 50–100 Hz) was used to detect the diastolic flow in the arteries.<sup>23</sup> The color box was kept small, covering only the area of interest, and the Doppler gain was set just below noise level. All of the participants were scanned by the first author.

The study data was analyzed with the Statistical Package for the Social Sciences (SPSS), version 20, for Windows (IBM Corp., Armonk, NY, US). The Mann-Whitney U test was used to compare the means because the uterine volumes and the accompanying Doppler indices were skewed and not normally distributed. Due to similar reasons, the Spearman correlation analysis was used to determine the correlation between the variables. The chi-squared test was used to compare two categorical variables, while the Kruskal-Wallis test was used when more than two variables were compared. The level of statistical significance was set at  $p \leq 0.05$ .

#### Results

A total of 280 premenopausal women were recruited consecutively for this prospective, case -control study comprising 140 subjects with uterine leiomyoma and 140 leiomyoma-free controls. **The subjects' general characteristics are displayed in ~ Table 1.** 

Nineteen subjects (13.6%) were asymptomatic, while 121 (86.4%) had various leiomyoma-related symptoms. Menorrhagia with abdominal swelling and/or pain was the most common presenting symptom, which was observed in 24 (17.1%) subjects. However, menorrhagia alone, and in various combinations with other symptoms, like infertility and dysmenorrhea, was a feature in 70 (50%) subjects. Fourteen subjects (10%) had recurrent leiomyoma after previous myomectomy, with a median time interval of 5 years (range = 2–15 years) after surgery. Eight (57.1%) of these presented  $\leq$  5 years post-myomectomy, while 6 (42.9%) of them were > 5 years post-myomectomy.

The subjects had a median uterine volume of 556 cm<sup>3</sup> (range = 37-9,384 cm<sup>3</sup>) which was significantly larger (p < 0.001) than that of the controls, who had a median uterine volume of 90.5 cm<sup>3</sup> (range = 21-304 cm<sup>3</sup>). When a

Variable	Patients	Controls	Statistic	df	р
	n = 140	n = 140			
Age in years					
Mean $\pm$ SD*	37.9 ± 7.4	30.5 ± 8.3	7.9	278	< 0.001
n (%)**					
< 20	0 (0.0)	13 (9.3)	46.0	4	< 0.001
20–29	18 (12.9)	51 (36.4)			
30-39	68 (48.6)	56 (40.0)			
40-49	45 (32.1)	18 (12.9)			
$\geq$ 50	9 (6.4)	2 (1.4)			
Parity, <i>n (%)</i>					
Nulliparous	79 (56.4)	70 (50.0)	2.3	3	0.5
Primiparous	21 (15.0)	19 (13.6)			
Multiparous (2–4)	32 (22.9)	43 (30.7)			
Grand multip. (> 4)	8 (5.7)	8 (5.7)			
Weight (Kg)	66.8 ± 15.2	63.7 ± 16.0	1.7	278	0.1
Height (m)	1.61 ± 0.06	1.6 ± 0.1	-0.1	278	0.9
BMI (kg/m²)	25.7 ± 5.3	24.6 ± 6.1	1.7	278	1.0

 Table 1
 Subjects' characteristics

Abbreviations: BMI, body mass index; df, degree of freedom. Notes: \*Independent samples *t*-test used to compare the means. \*\* Chi-squared test applied for the proportions. cut-off value of 200 cm<sup>3</sup> was used to divide the participants' uteri into large and small, 126 (90%) and 14 (10%) subjects had large and small uteri respectively, compared with 5 (3.6%) and 135 (96.4%) controls with large and small uteri respectively. This difference was statistically significant (p < 0.001).

Degenerative changes were observed in 69 (49.3%) subjects: 35 (25%) had cystic degeneration, 22 (15.7%) had calcific degeneration, and 12 (8.6%) had both cystic and calcific degenerations.

All the evaluated Doppler indices of the main uterine artery showed statistically significant differences between the subjects and controls (**- Table 2**). The main uterine arteries of the subjects had significantly higher PSV, EDV, TAMX, Tmean, AT, AI, DSR, DAR and inverse PI, but significantly lower PI, RI, SDR and ImI than the main uterine arteries of the controls.

All the main uterine artery Doppler indices, except the AI, showed statistically significant differences between the subjects with uterine volume  $\leq 200 \text{ cm}^3$  and those with uterine volume  $200 \text{ cm}^3$ . The latter group had higher flow velocities and lower resistance to flow than the former (**-Table 3**).

Symptomatic subjects (all symptoms inclusive) had significantly higher main uterine artery PSV, EDV, TAMX, and Al than the asymptomatic subjects. Furthermore, there were statistically significant differences in all the main uterine artery Doppler indices, except the AT and DAR, between the subjects with menorrhagia and those without. Those with menorrhagia had higher flow velocities and lower resistance to flow (**\sim Table 4**).

**Table 2** Comparison of main uterine artery Doppler indices of the subjects and controls

Doppler indices	Subjects	Controls	p <sup>†</sup>
	n = 140	n = 140	
PSV (cm/s)	94.2 ± 39.3	54.2 ± 16.4	< 0.001
EDV (cm/s)	29.7 ± 19.1	$7.7\pm2.6$	< 0.001
TAMX (cm/s)	$49.1\pm26.4$	$20.0\pm7.2$	< 0.001
Tmean (cm/s)	$25.5\pm16.4$	$10.0\pm4.2$	< 0.001
PI	$1.5\pm0.7$	$\textbf{2.4}\pm\textbf{0.4}$	< 0.001
RI	0.7 ± 0.1	$0.9\pm0.03$	< 0.001
SDR	3.8 ± 1.6	7.2 ± 1.5	< 0.001
AT (ms)	$117.9\pm28.0$	$92.0\pm22.8$	< 0.001
AI	0.8 ± 0.3	$0.6\pm0.2$	< 0.001
DSR	$0.3\pm0.1$	$0.1\pm0.03$	< 0.001
Iml	3.8 ± 1.6	7.2 ± 1.5	< 0.001
DAR	0.6 ± 0.2	$0.4\pm0.1$	< 0.001
1/PI	0.8 ± 0.3	0.4 ± 0.1	< 0.001

Abbreviations: 1/PI, inverse pulsatility index; AI, acceleration index; AT, acceleration time; DAR, diastolic average ratio; DSR, diastolic/systolic ratio; EDV, end-diastolic velocity; ImI, impedance index; PI, pulsatility index; PSV, peak systolic velocity; RI, resistivity index; SDR, systolic/ diastolic ratio; TAMX, time-averaged maximum velocity; Tmean, time-averaged mean velocity.

Note: † Mann-Whitney U test applied.

**Table 3** Comparison of main uterine artery Doppler indices by uterine volume in the subjects

Doppler indices	Uterine volume		p <sup>†</sup>
	$\leq$ 200.0 cm <sup>3</sup>	> 200.0 cm <sup>3</sup>	
	n = 14	n = 126	
PSV (cm/s)	$65.1 \pm 14.6$	$97.4\pm39.9$	< 0.001
EDV (cm/s)	$12.8\pm5.7$	$31.6\pm19.1$	< 0.001
TAMX (cm/s)	$\textbf{23.8} \pm \textbf{7.9}$	$51.9 \pm 26.2$	< 0.001
Tmean (cm/s)	$12.7\pm4.7$	$26.9\pm6.7$	< 0.001
PI	$2.4\pm1.1$	$1.4\pm0.6$	< 0.001
RI	$0.8\pm0.1$	$0.7\pm0.1$	< 0.001
SDR	5.7 ± 1.7	$3.6 \pm 1.5$	< 0.001
AT (ms)	$98.5\pm30.0$	$120.1\pm27.1$	0.01
AI	$0.7\pm0.3$	$0.8\pm0.3$	0.20
DSR	$0.2\pm0.1$	$0.3\pm0.1$	< 0.001
Iml	5.7 ± 1.7	3.6 ± 1.5	< 0.001
DAR	$0.6\pm0.3$	0.6 ± 0.2	0.01
1/PI	0.5 ± 0.1	0.8 ± 0.3	< 0.001

Abbreviations: 1/PI, inverse pulsatility index; AI, acceleration index; AT, acceleration time; DAR, diastolic average ratio; DSR, diastolic/systolic ratio; EDV, end-diastolic velocity; ImI, impedance index; PI, pulsatility index; PSV, peak systolic velocity; RI, resistivity index; SDR, systolic/diastolic ratio; TAMX, time-averaged maximum velocity; Tmean, time-averaged mean velocity. † Mann-Whitney U test applied.

**Table 4** Effect of menorrhagia on main uterine artery Dopplerindices in the subjects

Doppler	Menorrhagia		p†
indices	Present (n = 70)	Absent (n = 70)	
PSV (cm/s)	$103.6\pm39.5$	$84.8\pm37.0$	0.001
EDV (cm/s)	34.1 ± 19.0	$25.3 \pm 18.3$	0.001
TAMX (cm/s)	$55.4\pm26.7$	$42.8\pm24.7$	0.001
Tmean (cm/s)	$28.8\pm14.5$	$22.2 \pm 17.6$	0.001
PI	$1.4\pm0.7$	1.6 ± 0.7	0.03
RI	0.7 ± 0.1	0.7 ± 0.1	0.02
SDR	3.5 ± 1.3	4.1 ± 1.8	0.03
AT (ms)	$116.2\pm28.4$	$119.6\pm27.7$	0.37
AI	0.9 ± 0.3	0.7 ± 0.3	0.001
DSR	0.3 ± 0.1	0.3 ± 0.1	0.03
Iml	3.5 ± 1.3	4.1 ± 1.8	0.03
DAR	0.6 ± 0.3	0.6 ± 0.2	0.14
1/PI	0.8 ± 0.3	$0.7\pm0.3$	0.03

Abbreviations: 1/PI, inverse pulsatility index; AI, acceleration index; AT, acceleration time; DAR, diastolic average ratio; DSR, diastolic/systolic ratio; EDV, end-diastolic velocity; ImI, impedance index; PI, pulsatility index; PSV, peak systolic velocity; RI, resistivity index; SDR, systolic/ diastolic ratio; TAMX, time-averaged maximum velocity; Tmean, time-averaged mean velocity.

Note: † Mann-Whitney U test applied.

Doppler indices	History of previous myomectomy		p <sup>†</sup>
	Present (n = 14)	Absent (n = 126)	
PSV (cm/s)	$122.1\pm62.7$	$91.1\pm34.8$	0.05
EDV (cm/s)	$42.0\pm32.4$	$\textbf{28.3} \pm \textbf{16.6}$	0.1
TAMX (cm/s)	$63.9 \pm 42.4$	$47.4\pm23.7$	0.2
Tmean	$36.7\pm30.6$	$24.3 \pm 13.7$	0.1
PI	$1.4\pm0.4$	$1.5\pm0.8$	0.9
RI	$0.7\pm0.1$	0.7 ± 0.1	0.5
SDR	$\textbf{3.4}\pm\textbf{0.9}$	3.9 ± 1.7	0.5
AT (mms)	$126.7\pm37.1$	$116.9\pm26.8$	0.5
AI	$0.9\pm0.3$	$0.8\pm0.3$	0.1
DSR	$0.3\pm0.1$	$0.3\pm0.1$	0.5
lml	$\textbf{3.4}\pm\textbf{0.9}$	3.9 ± 1.7	0.5
DAR	0.6 ± 0.1	0.6 ± 0.2	0.1
1/PI	$0.8\pm0.2$	$0.8\pm0.3$	0.9

**Table 5** Effect of previous myomectomy on main uterine artery Doppler indices in the subjects

Abbreviations: 1/PI, inverse pulsatility index; AI, acceleration index; AT, acceleration time; DAR, diastolic average ratio; DSR, diastolic/systolic ratio; EDV, end-diastolic velocity; ImI, impedance index; PI, pulsatility index; PSV, peak systolic velocity; RI, resistivity index; SDR, systolic/ diastolic ratio; TAMX, time-averaged maximum velocity; Tmean, time-averaged mean velocity.

Note: † Mann-Whitney U test applied.

There were no statistically significant differences between all the main uterine artery Doppler indices of the subjects with previous myomectomy and of those without (**-Table 5**).

All the main uterine artery Doppler indices of the subjects showed a significant correlation with the subjects' uterine volume. The PSV, EDV, TAMX, and Tmean showed strong positive correlations with the uterine volume (correlation coefficient [r] 0.6), while the PI, RI, SDR, and the ImI showed moderate negative correlations with the uterine volume (r = -0.4 to -0.6). The main uterine artery PSV, EDV, TAMX, Tmean, PI, AT, DAR and inverse PI of the controls correlated weakly with the uterine volume (**-Table 6**).

Significantly higher main uterine artery flow velocity indices (except AI and DAR) and significantly lower resistance to flow were noted in subjects who had fibroids with degenerative changes compared with those without degenerative changes.

### Discussion

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Doppler sonography in Gynecology is an important adjunct to gray-scale sonography, with established roles in the assessment of infertility, uterine leiomyomas, pelvic arteriovenous malformations, hormone replacement therapy, etc.<sup>13,14</sup> In this study, the women with uterine leiomyomas showed a significant increase in uterine perfusion/blood flow compared with

**Table 6** Correlation\* between uterine volume and Doppler indices of the main uterine artery (UtA)

Doppler	Subjects (n = 140)		Controls (n = 140)	
indices	Correlation coefficient, r	P	Correlation coefficient, r	p
PSV (cm/s)	0.6	< 0.001	0.4	< 0.001
EDV (cm/s)	0.7	< 0.001	0.3	< 0.001
TAMX (cm/s)	0.7	< 0.001	0.4	< 0.001
Tmean (cm/s)	0.6	< 0.001	0.3	< 0.001
PI	-0.6	< 0.001	-0.3	< 0.001
RI	-0.5	< 0.001	-0.1	0.54
SDR	-0.5	< 0.001	-0.02	0.76
AT (ms)	0.3	< 0.001	0.4	< 0.001
AI	0.4	< 0.001	0.03	0.72
DSR	0.5	< 0.001	0.03	0.75
Iml	-0.5	< 0.001	-0.03	0.75
DAR	0.4	< 0.001	-0.2	0.01

Abbreviations: 1/PI, Inverse pulsatility index; AI, acceleration index; AT, acceleration time; DAR, diastolic average ratio; DSR, diastolic/systolic ratio; EDV, end-diastolic velocity; ImI, impedance index; PI, pulsatility index; PSV, peak systolic velocity; RI, resistivity index; SDR, systolic-diastolic ratio; TAMX, time-averaged maximum velocity; Tmean, time-averaged mean velocity. Note: \* Spearman correlation applied.

the controls, as evidenced by the elevation of uterine artery Doppler indices reflecting blood flow velocity (PSV, EDV, TAMX, Tmean, DAR, AT and AI) with increase in uterine volume, while the indices reflective of the degree of resistance to blood flow or vascular impedance (PI, RI, SDR, ImI, inverse PI) diminished with increase in uterine volume. This decrease in resistance to blood flow could be due to the uncoiling of the uterine artery with uterine enlargement in the leiomyomatous uterus.<sup>6</sup> Similar uterine artery uncoiling and diminution of resistance has been observed in enlarged gravid uteri.<sup>6</sup> The myometrial vasculature also enlarges<sup>24,25</sup> to supply the hypervascular networks present in uterine leiomyomas, which further decreases the resistance to blood flow.

Increase in blood flow velocity indices with increasing uterine volume was also noted in the controls, with weak statistical significance. This suggests that uterine enlargement rather than the mere presence of the leiomyomatous tumor is mainly responsible for the observed alteration in the uterine artery blood flow to the leiomyomatous uteri. Furthermore, women with a uterine volume  $> 200 \text{ cm}^3$  had significantly higher flow velocities and lower vascular ImIs than those with uterine volume  $\leq 200 \text{ cm}^3$ . This is similar to the results obtained by Farmakides et al<sup>6</sup> and Alatas et al.<sup>5</sup>

Leiomyomas with degenerative changes showed higher flow velocities and lower resistance to blood flow in the main uterine artery. This is in agreement with previous studies,<sup>2,26</sup> and could be due to the fact that degenerating leiomyomas are often large and rapidly growing, eventually outgrowing their blood supply. Such rapid growth probably requires greater blood supply to support the metabolic activities.

The study also found that the blood flow velocity indices of the main uterine artery were significantly higher in symptomatic subjects compared with asymptomatic subjects. When menorrhagia was considered as a specific symptom, those with menorrhagia had higher main uterine artery blood flow velocity indices and lower vascular resistance indices, which is suggestive of a "compensatory" increase in perfusion when menorrhagia occurs. These two observations suggest that the presence of menorrhagia leads to significant alterations in the main uterine artery Doppler indices. A similar pattern was documented by Hurskainen et al,<sup>27</sup> who reported a significant inverse correlation between the uterine artery PI and the amount of blood loss in menstruating women (those with lower uterine flow impedance bled more), and concluded that there likely is a significant relationship between uterine vascular tone and menorrhagia.<sup>27</sup>

Apart from being a black population study,<sup>22–30</sup> the markedly larger uterine volumes in this study could also be due to the fact that the majority of the patients in this environment present late, often after failure of alternative therapies, or due to the prohibitive cost of orthodox medical treatments. These markedly larger sizes may also explain the much lower Doppler indices in this study compared with those of some previous studies.<sup>2,4,5</sup> Moreover, the median uterine volume of the subjects in this study was 556 cm<sup>3</sup>, which is much higher than the reported mean uterine volumes of 128.5 cm<sup>3</sup>, 187.4 cm<sup>3</sup>, 276.2 cm<sup>3</sup>, 305 cm<sup>3</sup>, 312 cm<sup>3</sup>, and 381.16 cm<sup>3</sup> by Tsuda et al,<sup>31</sup> Alcazar et al,<sup>4</sup> Alatas et al,<sup>5</sup> Tranquart et al,<sup>32</sup> Danisman et al,<sup>14</sup> and Samani et al<sup>12</sup> respectively.

There were no significant differences between the main uterine artery Doppler indices of uteri with recurrent leiomyomas and those with primary tumors. This is most likely due to the fact that there was no statistically significant difference between the uterine volumes of the two groups; increase in uterine volume is the main contributor to the alterations in main uterine artery Doppler indices. Leiomyoma recurrence could be due to de novo growth of new tumors from previously normal myometrial cells post-myomectomy, or due to persistent undetected tumors (remnants) left unintentionally from an incomplete fibroid surgery.<sup>33</sup> Obed et al,<sup>33</sup> who defined recurrence as the presence of a leiomyoma of at least 2 cm in diameter upon ultrasound scan, reported an overall recurrence rate of 20.7% at 10 years post-myomectomy.

The limitations of our study include the fact that the sonographic diagnosis of leiomyoma was not confirmed with histology. However, several studies have shown the increased sensitivity and specificity of the sonographic diagnosis of leiomyomas with more modern ultrasound scanners. There was also difficulty in obtaining accurate measurements of very bulky, leiomyomatous uteri and huge pedunculated leiomyomas extending to the upper abdomen. Split-screen measurements were employed in such cases.

#### Conclusion

In conclusion, there is increased perfusion of the myomatous uterus, as evidenced by increased blood flow velocity indices and reduced vascular impedance indices in their main uterine artery compared with the controls. These changes are more likely due to increased uterine volume rather than the mere presence of myomas. No significant differences were detected between the main uterine artery Doppler indices of uteri with recurrent leiomyoma and those with primary tumors.

#### Contributions

Idowu BM, Ibitoye BO and Adetiloye VA contributed with the conception and design, data collection and analysis, interpretation of data, writing of the article, critical review of the intellectual content, and final approval of the version to be published.

Conflicts of Interest None to declare.

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