#### ORIGINAL RESEARCH

## A Cross-Sectional Study on Potential Ovarian Volume and Related Factors in Women with Polycystic Ovary Syndrome from Infertile Couples

Nguyen Sa Viet Le 10<sup>1,2</sup> Minh Tam Le 10<sup>2,3</sup> Nguyen Dac Nguyen 10<sup>2,3</sup> Nhu Quynh Thi Tran 10<sup>3</sup> Quoc Huy Vu Nguyen 10<sup>2</sup> Thanh Ngoc Cao 10<sup>2,3</sup>

<sup>1</sup>Department of Assisted Reproduction, Hue Central Hospital, Hue, Vietnam; <sup>2</sup>Department of Obstetrics and Gynecology, Hue University of Medicine and Pharmacy, Hue University, Hue, Vietnam; <sup>3</sup>Center for Reproductive Endocrinology and Infertility, Hue University of Medicine and Pharmacy, Hue University, Hue, Vietnam **Purpose:** This study was designed to explore the value of ovarian volume (OV) measured by transvaginal ultrasound and its relationship with anthropometry and serum hormonal levels in a polycystic ovary syndrome (PCOS) population.

**Patients and Methods:** A total of 119 women with PCOS from infertile couples were recruited in this cross-sectional study. On days 2–4 of the menstrual cycle, transvaginal ultrasound examinations were performed, and hormonal profiles were measured. PCOS diagnosis was based on the Rotterdam 2003 criteria and classified into four phenotype groups. The PCOS group (study group) and the non-PCOS group (control group) were compared.

**Results:** The mean age of the participants was  $32.66\pm4.10$  years compared to  $33.99\pm4.78$  years in 273 cases (69.6%) without PCOS. The mean OV was statistically larger in the PCOS group than in the non-PCOS group (7.65 $\pm3.23$  mL vs  $6.08\pm3.67$  mL, p < 0.001) and positively correlated with serum anti-Mullerian (AMH) and luteinizing hormone (LH) levels (r=0.30; p < 0.001 and r=0.23; p < 0.001, respectively), and weakly and inversely correlated with age (-0.182, p < 0.001). The area under the receiver operating characteristic (ROC) curve of OV in the diagnosis of PCOS was 0.613 (0.557–0.670, 95% CI).

**Conclusion:** The enlarged OV is remarkable in women with PCOS and is related to AMH and LH concentrations. Although the diagnostic potential of PCOS is substantially low, OV alone may contribute to predicting the severity of PCOS and better performance for the diagnosis of PCOS phenotypes.

**Keywords:** ovarian volume, anti-Mullerian hormone, PCOS phenotypes, diagnose, infertility

#### Introduction

Polycystic ovary syndrome (PCOS) is the most common endocrine disorder. It can affect 6% of women in the reproductive age according to the diagnostic criteria of National Institutes of Health (NIH) and up to 8–13% of these women according to the Rotterdam criteria.<sup>1</sup> This heterogeneous disorder is characterized by hyperandrogenism, chronic anovulation, and the presence of polycystic ovaries on ultrasound.<sup>2</sup> PCOS diagnosis is done based on the presence of oligo-anovulation and androgen excess only according to the NIH criteria, after ruling out all other reasons for anovulatory infertility.<sup>3</sup> In 2003, the European Society for Human Reproduction and Embryology (ESHRE) and the American Society for Reproductive Medicine enlarged the consensus criteria by including polycystic ovarian morphology on ultrasoundas the third diagnostic criterion and further requires the presence of at least two of three criteria for

Correspondence: Minh Tam Le Email leminhtam@huemed-univ.edu.vn

793

© 2021 Le et al. This work is published and licensed by Dove Medical Press Limited. The full terms of this license are available at https://www.dovepress.com/terms.by you hereby accept the firms. Non-commercial uses of the work are permitted without any further permission from Dove Medical Press Limited, provided the work is properly attributed. For permission for commercial uses of the work are permitted without any further permission from Dove Medical Press Limited, provided the work is properly attributed. For permission for commercial uses of the work are permitted without any further permission from Dove Medical Press Limited, provided the work is properly attributed. For permission for commercial uses of this work, please see paragraphs 4.2 and 5 of our Terms (http://www.dovepress.com/terms.php). a diagnosis of PCOS.<sup>4</sup> The Rotterdam 2003 also expanded the NIH 1990 definition creating two new phenotypes. Classical phenotypes include phenotype A (full-blown syndrome PCOS); phenotype B (non-PCOS PCOS) while two additional phenotypes include phenotype C (ovulatory PCOS) and phenotype D (non-hyperandrogenic PCOS). More recently, the 2018 International evidence-based Guidelines on the diagnosis and management of PCOS endorsed the Rotterdam criteria for PCOS diagnosis.<sup>5</sup> Although an ultrasound is not strictly needed in diagnosing PCOS, it is still recommended to identify the complete phenotype and may be useful for other indications in PCOS.<sup>5</sup>

Based on ultrasonographic evidence for diagnosis of PCOS, the Rotterdam consensus criteria in 2004 defined polycystic ovarian morphology (PCOM) as follicle number per ovary (FNPO) with a threshold of  $\geq 12$  follicles measuring 2-9 mm in diameter (mean of both ovaries) and/or increased ovarian volume (OV) of  $\geq 10$  mL.<sup>4</sup> The 2018 ESHRE PCOS guideline group suggested a threshold of >20 FNPO with or without an OV  $\ge$  10 mL in either ovary using transvaginal ultrasound transducers with a frequency bandwidth of 8 MHz to diagnose PCOM.<sup>5</sup> It is worth noting that there are no changes to an OV of ≥10 mL the diagnosis of PCOM even imaging techniques have considerably advanced.6 However, FNPO is still recommended over OV in PCOS diagnosis over OV because of its higher predictive performance and lesser variability.<sup>6</sup> However, the levels of intra-observer reliability and inter-reliability were higher when assessing OV than FNPO and follicle number per section.<sup>7</sup> Therefore, OV is still a potential diagnostic tool for detecting PCOS. Several studies have investigated the relationship between OV and PCOS,8 obesity,9 insulin resistance,<sup>10</sup> androgen activity,<sup>11,12</sup> metabolic syndrome,<sup>13</sup> and AMH levels<sup>12,14,15</sup> in PCOS patients. Furthermore, to date, while an ovarian volume of >10mL is a part of the Rotterdam criteria for PCOS, several research groups have suggested various cut-off points for OV due to variation in population characteristics. The aim of this study was to examine the value of OV measured by transvaginal ultrasound and its relationship to anthropometry and serum hormonal levels in the PCOS population.

### **Patients and Methods** Study Population

This cross-sectional study was carried out at the Hue Center for Reproductive Endocrinology and Infertility, Vietnam, from January 2019 to December 2020. Infertile women who were diagnosed with PCOS followed the Rotterdam criteria 2003<sup>4</sup> were recruited for the study. Exclusion criteria were the presence of ovarian diseases (ovarian cyst/tumor or endometrioma), history of adnexal surgery, ovarian failure, history of hormonal contraception use, or any hormonal treatment within three months prior to enrollment. Ethical approval for this study was obtained from the Ethics Committee of the Hue University of Medicine and Pharmacy. The study was conducted in accordance with the Declaration of Helsinki. Written informed consent was obtained from all individual participants included in the study.

The sample size (n) required for this study was calculated based on the equation  $n=Z_{\alpha/2}^2 \times P \times (1-P)/\Delta^2$ , where  $\alpha$ =0.05,  $\Delta$ =0.05, and  $Z_{\alpha/2}$ =1.96. The estimated prevalence of PCOS in infertile women (*P*) was 40.9%.<sup>16</sup> The total sample size required in this study was 372.

## Clinical and Hormonal Tests and Ultrasonography Assessment

All participants were asked about their menstrual cycle, medical, gynecologic, and obstetric history. Oligomenorrhea was defined as having fewer than 8 menstrual cycles per year, the absence of 3-6 consecutive menstrual cycles per year or the length of menstrual cycle greater than 35 days. Clinical examinations, including measurement of body weight, height, waist and hip circumference, and evaluation for signs of hyperandrogenism were performed. Body mass index (BMI) was calculated by dividing body weight in kilograms by the square of patient height in meters. Waist and hip circumferences measurement were made on exposed skin around the abdomen at the levels of navel and pubic symphysis, respectively, in the standing position. Hirsutism was visually graded using a modification of the Ferriman and Gallwey scoring system (mFG). We defined clinical hirsutism as mFG  $\geq$ 5 using the mFG cut-off criterion for the Asian PCOS population.<sup>17</sup> All questions were done through a demographic questionnaire.

On day 2–4 of natural cycle or on day 2–4 of progesterone withdrawal in case of oligo or amenorrhea condition, transvaginal ultrasound examination was performed by the same experienced physician by ultrasonography (ALOKA ProSound SSD-3500, Hitachi, Japan) using a vaginal probe of 7 MHz to evaluate each participant's antral follicle number and OV. Ovaries were scanned from the inner to the outer margin ovaries in both the transverse and sagittal planes. Three dimensions of each ovary were measured, and the total number of antral follicles that were 2–9 mm in diameter were counted. The OV was estimated for each ovary using the  $\pi/6 \times (D1 \times D2 \times D3)$  formula. D presented the longest diameter of each ovary dimension (long, anterior-posterior, and transverse sections).

Blood samples were collected for basic hormonal testing on the same day as the ultrasound examinations were performed. Basic hormonal profiles including follicle-stimulating hormone (FSH), LH, estradiol, and prolactin were measured by an immunoradiometric assay (IRMA). AMH levels were determined by an Elecsys Roche System using electrochemiluminescence (ECLIA) technology. All tests were performed at the laboratory center of Hue University Hospital.

The PCOS diagnosis followed the Rotterdam criteria 2003 with at least two of the following three features (study group): (i) oligo-and/or anovulation, (ii) clinical and/or biochemical signs of hyperandrogenism, and/or (iii) polycystic ovaries by transvaginal ultrasound scan (more than 12 follicles in the 2–9 mm range in each ovary and/or OV > 10 mL).<sup>4</sup> The remaining participants were infertile women without PCOS (control group). PCOS patients were classified into four phenotype groups according to the NIH consensus panel 2012.<sup>18</sup> Phenotype A included patients with hyperandrogenism (HA), ovulatory dysfunction (OD), and polycystic ovaries (PCO) (HA +OD+PCO); Phenotype B included those with HA + OD; phenotype C included those with HA + PCO; and phenotype D included those with OD + PCO.

#### Data Analysis

The data are presented as proportions and mean  $\pm$  standard deviation (SD). Categorical data were assessed for normal distribution using the Shapiro-Wilk test. The Student's t-test or Mann Whitney U-test was used to compare the differences between the PCOS and control groups, and four phenotype groups of PCOS. Dichotomous variables were compared using two-tailed chi-square or Fisher exact tests, where appropriate. Spearman correlation was used for an analysis of the association between AMH levels and OV. Statistical significance was set at p < 0.05. The Youden index was used to identify the best threshold values for AMH levels and OV. ROC curves were constructed to assess the diagnostic ability of AMH and OV. Sensitivity against 1- specificity was plotted at each threshold level, and the area under the curve (AUC) was calculated. AUC represents the probability of correctly identifying controls and patients with PCOS. A value of 0.5 indicated that the result was not better than random. All statistical data were analyzed using SPSS Statistics Version 24.0 software (SPSS Inc., Chicago, IL, USA).

#### Results

The general characteristics, anthropometry, hormonal tests, and ultrasound scans of all recruited populations, presented in the PCOS and non-PCOS groups, are reported in Table 1. Mean age of the 119 participants was 32.66±4.10 years compared to  $33.99\pm4.78$  years in 273 cases without PCOS. Women with PCOS more commonly had primary infertility, higher LH and LH:FSH ratios, and lower FSH values. Serum AMH levels were 2-fold higher in the PCOS group than that in the control group (7.35±5.17 vs 3.48±3.17 ng/mL). There were no statistically significant differences in the waist-hip ratio, mean plasma estradiol, and mean prolactin concentrations in either group. The mean OV was calculated as 7.48±3.11 mL in the PCOS group and 6.58±3.69 mL in the non-PCOS group (p < 0.001). Significant differences were observed on both sides of the ovaries.

Table 2 shows the distribution of anthropometry, clinical and hormonal profiles, and ultrasound findings among the different phenotypes. The prevalence of classical phenotype A (HA+ OD+ PCO), phenotype B (HA + OD), phenotype C (HA + PCO), and phenotype D (OD+PCO) were 5.04%, 1.68%, 13.45%, and 79.83% in women with PCOS, respectively. The four subgroups showed similar clinical, physical, and hormonal characteristics. However, mFG was higher in subgroup C than in the other subgroups (p < 0.01). Serum AMH concentration and mean OV seemed to be lower in phenotype B than in other phenotypes, but the differences did not reach statistical significance.

The correlations between clinical characteristics and hormonal profiles with mean OV are shown in Table 3. AMH and OV were positively correlated (right OV: r=0.28; left OV: r=0.26; mean OV: r=0.30; p < 0.001). Similarly, LH and OV had a weaker positive relationship (right OV: r=0.19; left OV: r=0.22; mean OV: r=0.23; p < 0.001). In addition, OV had a weak negative relationship with age and FSH levels.

The diagnostic potency of the OV compared to the AMH assay was calculated using the ROC procedure. The AUC of the OV to predict PCOS was 0.613 (0.557–0.670, 95% CI) (Figure 1A). At the optimal cut-off (4.57mL), the sensitivity and specificity for OV were 92.74% and 33.3%, respectively. When the threshold value was set at 6.03 mL, the sensitivity and specificity for OV were 61.3% and 50.5%, respectively (Table 4). The ROC analysis of the AMH test was more informative for the diagnosis of PCOS, with an AUC of 0.787 (0.741–0.834; 95% CI) (Figure 1B). The ideal cut-off value for AMH level was determined to be 4.195 ng/mL. The

Characteristics		PCOS (n=119)	Non – PCOS (n=273)	р
Age (yrs)		32.66 ± 4.10	33.99 ± 4.78	0.042
Menarche (yrs)		13.37 ± 0.92	13.74 ± 1.39	0.002
Duration of infertility (yrs	5)	4.27 ± 2.46	4.09 ± 2.80	0.010
Type of infertility	Primary	90 (75.6)	177 (64.8)	0.045
	Secondary	29 (24.4)	96 (35.2)	
BMI (kg/m <sup>2</sup> )		21.31 ± 2.80	20.82 ± 2.56	<0.001
mFG score		0.55 ± 1.09	0.14 ± 0.46	<0.001
WHR	WHR		0.83 ± 0.07	0.762
FSH (mlU/mL)		6.07 ± 1.46	7.41 ± 3.49	<0.001
LH (mIU/mL)	LH (mIU/mL)		6.23 ± 3.43	<0.001
LH/FSH		1.57 ± 0.85	0.95 ± 0.63	<0.001
E2 (pg/mL)	E2 (pg/mL)		40.22 ± 40.76	0.137
Prolactin (µIU/mL)		410.45 ± 243.46	432.26 ± 318.07	0.060
AMH (ng/mL)		7.35 ± 5.17	3.48 ± 3.17	<0.001
Ovarian volume (cm <sup>3</sup> )	Right ovarian	8.18±3.73	6.69 ± 4.17	<0.001
	Left ovarian	6.78±3.40	6.46 ± 4.11	<0.001
	Mean volume	7.48±3.11	6.58 ± 3.69	<0.001

Table I General Characteristics, Anthr	opometry, Hormonal Profiles a	nd Ovarian Ultrasound Findings
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Notes: Data presented in Mean ± standard deviation or number (percentage).

Abbreviations: PCOS, polycystic ovary syndrome; yrs, year; BMI, body mass index; mFG, modified Ferriman Gallwey; WHR, weight hip ratio; FSH, follicle stimulating hormone; LH, luteinizing hormone; E2, estradiol; AMH, anti-Mullerian hormone.

sensitivity and specificity at this cut-off point were 73.1% and 70.7%, respectively.

## Discussion

This study was designed to explore the value of OV measured by transvaginal ultrasound in the diagnosis of PCOS and its relation to anthropometry and hormonal profiles, especially AMH. This study revealed that in Vietnamese women from infertile couples with PCOS, the right OV was 8.25±4.03 mL and the left OV was 7.06±3.37 mL. The mean volume of the two ovaries was 7.65±3.23 mL; which was smaller in comparison with results from several previous studies, reported by Carmina et al (9.6±3.2 mL),<sup>19</sup> Ndoua et al  $(12.02\pm3.36 \text{ mL})$ .<sup>14</sup> and Nylander et al (9.40 $\pm 3.62$  mL).<sup>20</sup> However, our findings were consistent with data of Korean PCOS patients, who had a right OV of 7.9  $\pm 3.6$  mL and left OV of 6.7 $\pm 3.1$  mL.<sup>12</sup> In the Asian population, lower OV in women with PCOS was mentioned previously<sup>21</sup> and was supposed to differ among women in various ethnic subgroups of women.<sup>22,23</sup> Furthermore, ovarian volume varies throughout the reproductive life of a woman, that is, it reaches its maximal size during adolescence, slowly declines during adulthood, and rapidly shrinks during menopause.<sup>24,25</sup> Therefore, the difference in age could explain the difference in mean OV between studies.

In previous studies, a larger mean OV in women with PCOS was reported compared to that in the non-PCOS population.<sup>12,21,26,27</sup> The ovaries of PCOS patients generally showed an increased number of small antral follicles and different degrees of thecal cell hyperplasia and hypertrophy, stromal hyperplasia and hypertrophy, and cortical thickening, resulting in an increase in the OV.<sup>28,29</sup> Several studies conducted based on the Rotterdam consensus suggested a lower cut-off for OV ranging from 6.4 to 7.5 mL (6.4;<sup>30</sup> 6.7;<sup>27</sup> 7.0;<sup>31,32</sup> and 7.5 mL<sup>26</sup>) to increase the sensitivity of the polycystic ovary definition. These differences could be explained by the variation in population characteristics, particularly the ethnic factors, obesity, serum insulin level, and the methods used to determine the OV.<sup>11,24,25,33</sup> Our data revealed that the diagnostic

**Table 2** Comparison of General Characteristics, Anthropometry, Hormonal Profiles and Ovarian Ultrasound Findings Between PCOS

 Phenotypes

Characteristics	Phenotype A (HA+OD+PCO)	Phenotype B (HA+OD)	Phenotype C (HA+PCO)	Phenotype D (OD+PCO)	р
Prevalence	6 (5.04)	2 (1.68)	16 (13.45)	95 (79.83)	
Age (yrs)	30.50 ± 2.43	37.00 ± 12.73	32.31 ± 4.14	32.77 ± 3.95	0.433
Menarche (yrs)	13.33 ± 0.52	13.50 ± 0.71	13.75 ± 1.39	13.31 ± 0.84	0.042
Duration of infertility (yrs)	4.00 ± 3.03	4.00 ± 2.83	2.94 ± 2.29	4.52 ± 2.41	0.017
BMI	21.48 ± 3.11	20.12 ± 0.83	20.55 ± 1.74	21.45 ± 2.95	0.694
mFG score	0.33 ± 0.52	0	2.06 ± 1.81	0.32 ± 0.70	<0.001
WHR	0.81 ± 0.05	0.77 ± 0.07	0.80 ± 0.06	0.82 ± 0.06	0.540
FSH (mIU/mL)	6.10 ± 1.05	6.44 ± 2.21	6.71 ± 2.05	5.95 ± 1.35	0.551
LH (mIU/mL)	11.75 ± 7.72	7.57 ± 4.62	10.75 ± 5.85	9.03 ± 5.07	0.539
E2 (pg/mL)	57.63 ± 41.37	67.14 ± 45.63	58.39 ± 30.29	39.92 ± 23.46	0.017
Prolactin (µIU/mL)	455.77 ± 245.09	419.25 ± 250.67	413.84 ± 178.74	406.83 ± 255.53	0.888
AMH (ng/mL)	7.35 ± 6.28	2.73 ± 0.89	7.94 ± 7.33	7.35 ± 4.73	0.264
Ovarian volume (cm <sup>3</sup> )	7.48 ± 2.35	3.87 ± 1.38	7.15 ± 4.74	7.61 ± 2.82	0.059

**Notes**: Data presented in Mean ± standard deviation or number (percentage).

Abbreviations: PCOS, polycystic ovary syndrome; yrs, year; BMI, body mass index; mFG, modified Ferriman Gallwey; WHR, weight hip ratio; FSH, follicle stimulating hormone; LH, luteinizing hormone; E2, estradiol; AMH, anti-Mullerian hormone.

Parameters	Right Ovarian Volume		Left Ovarian Volume		Mean Volume	
	r	P value	r	P value	r	P value
Age (yrs)	-0.160	0.001	-0.162	0.001	-0.182	<0.001
BMI	0.123	0.015	0.048	0.343	0.098	0.054
mFG score	0.100	0.052	0.106	0.036	0.115	0.022
WHR	-0.019	0.705	-0.027	0.601	-0.026	0.612
FSH (mUI/mL)	-0.129	0.010	-0.087	0.084	-0.123	0.015
LH (mUI/mL)	0.193	<0.001	0.219	<0.001	0.233	<0.001
E2 (pg/mL)	-0.058	0.254	0.029	0.568	-0.017	0.731
Prolactin (µIU/mL)	-0.058	0.251	-0.019	0.713	-0.044	0.386
AMH (ng/mL)	0.281	<0.001	0.263	<0.001	0.307	<0.001

Abbreviations: PCOS, polycystic ovary syndrome; yrs, year; BMI, body mass index; mFG, modified Ferriman Gallwey; WHR, weight hip ratio; FSH, follicle stimulating hormone; LH, luteinizing hormone; E2, estradiol; AMH, anti-Mullerian hormone.

potential of OV was substantially low (AUC 0.613 [0.557–0.670]). At the cut–off point of 6.0 mL, a sensitivity of 61.3% and a specificity of 50.5% were obtained. The specificity of OV reached 57.5% when its cut-off value was 6.3 mL, but the sensitivity was lower (only 57.1%). These findings were different from several published studies that concluded that OV had satisfactory power for use in the diagnosis of PCOS.<sup>21,27,31,32</sup> The main reasons for this discrepancy are the differences in the study population and ethnicity. As mentioned above, our PCOS women had remarkably small ovaries and the difference range in OV between the PCOS and non-PCOS population was also lower (7.65±3.23 mL vs 6.08±3.67 mL, respectively).

Our study confirmed that AMH is a good diagnostic tool for PCOS. It is a much better predictor of PCOS than ovarian volume, with an AUC of 0.787 (0.741–0.834, 96% CI). AMH threshold at 4.2 ng/mL provided the best compromise, with a sensitivity of 73.1% and specificity of 70.7%. There is emerging evidence in the published literature to confirm the potential of AMH for the diagnosis of PCOS, with moderate to excellent sensitivity and specificity at cut-off points ranging from 3.94 ng/mL to 5 ng/mL.<sup>20,34–37</sup> A meta-analysis by Illiodromiti et al using ROC analysis figured out an AMH cutoff value of 4.7 ng/mL to obtain 82.8% sensitivity and 79.4% specificity.<sup>38</sup> It is important to highlight that the differences in ethnicity, age of study populations, criteria used to define

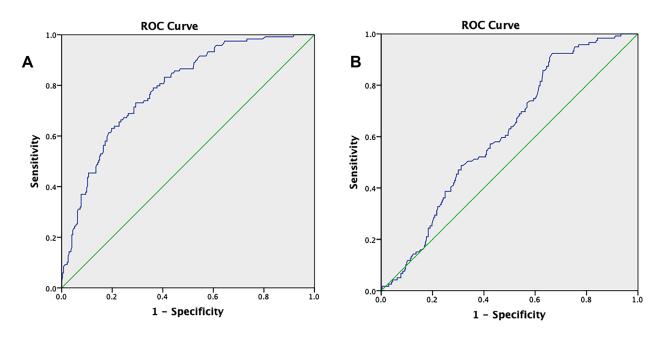


Figure I ROC curve of prediction PCOS by AMH (A) and by mean of ovarian volume (B). Diagonal segments are produced by ties.

PCOS, and type of AMH assays may influence on the results and proposed diagnostic cut-off.<sup>39–41</sup>

Published studies comparing the diagnostic ability of AMH measurement and ovarian imaging have reported conflicting results. Carmina et al carried out a retrospective matched controlled study and concluded that AMH did not appear to be as helpful as compared to FNPO and OV.19 They also assumed that FNPO was significantly sensitive in all phenotypes and was the single best criterion, supporting the essential role of ultrasound in PCOS diagnosis.<sup>19</sup> In contrast, Wongwanaruruk et al considered AMH to be a superior predictor of ultrasonographic imaging of ovarian morphology, as the latter is dependent on the quality of ultrasound scanners and sonographer experience.<sup>21</sup> However, the International evidence-based guideline for the assessment and management of PCOS 2018 still did not recommend the use of AMH as an alternative for the detection of PCOM or as a single diagnostic test.

The present study demonstrated a positive correlation between OV and serum AMH and LH levels in women with PCOS. Some other Korean authors reported this relationship to be consistent with our results.<sup>12,42</sup> Ndoua et al demonstrated that AMH showed a good correlation with OV (r = 0.625, p = 0.0001).<sup>14</sup> In addition, we found a weak negative correlation between OV and age. This finding was also reported by Erdem et al (with r r-0.29, p < 0.05).<sup>43</sup> As OV in women aged 25–51 years reflects the number of primordial follicles,<sup>44</sup> it was not difficult to explain the negative correlation between OV and age, and the positive correlation between OV and AMH, as the latter was demonstrated to strongly correlate with the primordial follicle pool.<sup>45</sup>

As demonstrated in our data, there were no differences in age, BMI, WHR and hormonal profiles among four phenotypes of PCOS. Serum AMH level and mean ovarian volume were insignificantly lower in phenotype B than in other

Table 4 Diagnostic Value	s of Anti-Mullerian Hormone	and Ovarian Volume for	Polycystic Ovary	Syndrome
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Parameters	Threshold	Sensitivity	Specificity	AUC, p
Mean of ovarian volume	4.575	0.924	0.333	0.613 (0.557–0.670)
	6.030	0.613	0.505	P < 0.001
AMH (ng/mL)	4.195	0.731	0.707	0.787 (0.741–0.834) P < 0.001

Abbreviations: AUC, area under curve; AMH, anti-Mullerian hormone.

phenotypes. In fact, previous studies showed controversial results that phenotype A was the most severe form with more frequent prevalence of obesity, hyperandrogenism, insulin resistance, deranged lipid profile, metabolic syndrome and higher LH, LH-FSH ratio compared to others; whereas Phenotype D was associated with the least severe profile.<sup>46,47</sup> However, other studies suggested that no difference was observed in the clinical biochemical profiles of patients with different phenotypes.<sup>48,49</sup> Similar to our ultrasound findings, Clark et al, and Sachdeva et al have also described smaller ovarian volumes in phenotype B.<sup>46,50</sup> In addition to sample size, most researchers agreed that racial differences in different study populations, again, may explain these controversies. Indeed, our results seemed to be more consistent with those from studies on Chinese populations.<sup>48,49</sup>

The strength of our study was its relatively large sample size and its comparative cross-sectional design. Notably, although measurement of OV alone was insufficient to diagnose PCOS in the Vietnamese PCOS population, its correlation with AMH could be a contributor to PCOS severity. However, the present study only recruited women from infertile couples, which may have resulted in selection of a relatively young age group. Therefore, neither the women diagnosed with PCOS nor the controls can be considered representative of the general population. The impact of this select study population should be considered in the drawn conclusion. Additionally, this study was performed in a single center and due to the study population included women from infertile couples, the data did not represent the general population. Thus, our conclusions must be considered in light of these limitations. Further prospective studies with agematched controls are needed for more definitive conclusions.

### Conclusion

In summary, our data suggested that in women with PCOS from infertile couples, OV had a significant positive correlation with serum AMH and LH levels and a negative correlation with age. The diagnostic potential of OV was substantially low, and the measurement of OV by transvaginal ultrasound alone to diagnose PCOS should not be considered in Vietnamese infertile women. Alternatively, serum AMH levels seemed to have better performance in the diagnosis of PCOS in this population.

#### Abbreviations

OV, ovarian volume; PCOS, polycystic ovary syndrome; AMH, anti-Mullerian hormone; LH, luteinizing hormone; ROC curve, receiver operating characteristic curve; NIH, National Institutes of Health; PCOM, polycystic ovarian morphology; FNPO, follicle number per ovary; ESHRE, European Society for Human Reproduction and Embryology; BMI, body mass index; mFG, modification of the Ferriman and Gallwey scoring system; FSH, Follicle-stimulating hormone; IRMA, immunoradiometric assay; ECLIA, electrochemiluminescence; OD, ovulatory dysfunction, PCO, polycystic ovaries; SD, standard deviation; AUC, area under the curve.

### **Data Sharing Statement**

The dataset used and/or analyzed during the current study is available from the corresponding author upon reasonable request.

# Ethics Approval and Informed Consent

The study was approved by the Ethics Committee of the Hue University of Medicine and Pharmacy on November 9th, 2018 (approval number H2018/432). The present study was conducted in accordance with the Declaration of Helsinki. Informed consent form was obtained for all participants prior to participation in this study.

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### Disclosure

The authors alone are responsible for the content and writing of this article. The authors report no conflicts of interest for this work.

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Le et al

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