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

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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±4.10 years compared to 33.99±4.78 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±3.23 mL vs 6.08±3.67 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.1 This heterogeneous disorder is characterized by hyperandrogenism, chronic anovulation, and the presence of polycystic ovaries on ultrasound.2 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.3 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 ultrasound as the third diagnostic criterion and further requires the presence of at least two of three criteria for...
a diagnosis of PCOS. 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. 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.

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 ≥12 follicles measuring 2–9 mm in diameter (mean of both ovaries) and/or increased ovarian volume (OV) of ≥10 mL. The 2018 ESHRE PCOS guideline group suggested a threshold of >20 FNPO with or without an OV ≥ 10 mL in either ovary using transvaginal ultrasound transducers with a frequency bandwidth of 8 MHz to diagnose PCOM. 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. However, FNPO is still recommended over OV in PCOS diagnosis over OV because of its higher predictive performance and lesser variability. However, the levels of intra-observer reliability and inter-reliability were higher when assessing OV than FNPO and follicle number per-section. Therefore, OV is still a potential diagnostic tool for detecting PCOS. Several studies have investigated the relationship between OV and PCOS, obesity, insulin resistance, androgen activity, metabolic syndrome, and AMH levels 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 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 \(n = \frac{Z_{\alpha/2}^2 \times P \times (1-P)}{\Delta^2}\), 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%. 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 ≥ 5 using the mFG cut-off criterion for the Asian PCOS population.

All questions were done through a demographic questionnaire.

On day 2–4 of natural cycle or on day 2–4 of progestrone 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 electrochemi-
 luminescence (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). The remaining participants
were infertile women without PCOS (control group).
PCOS patients were classified into four phenotype groups
according to the NIH consensus panel 2012.18 Phenotype
A included patients with hyperandrogenism (HA), ovula-
tory dysfunction (OD), and polycystic ovaries (PCO) (HA
+ OD + PCO); Phenotype B included those with HA + OD;
phenotype C included those with HA + PCO; and phy-
type D included those with OD + PCO.

Data Analysis

The data are presented as proportions and mean ± 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 con-
structed 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 \( 33.99±4.78 \) 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 statis-
tically 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, respec-
tively. 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 hor-
monal 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
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), 19 Ndoua et al (12.02±3.36 mL), 14 and Nylander et al (9.40±3.62 mL). 20 However, our findings were consistent with data of Korean PCOS patients, who had a right OV of 7.9±3.6 mL and left OV of 6.7±3.1 mL. 21 In the Asian population, lower OV in women with PCOS was mentioned previously 21 and was supposed to differ among women in various ethnic subgroups of women. 22,23 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. 24,25 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. 12,21,26,27 The ovaries of PCOS patients generally showed an increased number of small antral follicles and different degrees of thecal cell hyperplasia and hyper trophy, stromal hyperplasia and hypertrophy, and cortical thickening, resulting in an increase in the OV. 28,29 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; 30 6.7; 27 7.0; 31,32 and 7.5 mL 26) 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. 11,24,25,33 Our data revealed that the diagnostic

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>PCOS (n=119)</th>
<th>Non – PCOS (n=273)</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (yrs)</td>
<td>32.66 ± 4.10</td>
<td>33.99 ± 4.78</td>
<td>0.042</td>
</tr>
<tr>
<td>Menarche (yrs)</td>
<td>13.37 ± 0.92</td>
<td>13.74 ± 1.39</td>
<td>0.002</td>
</tr>
<tr>
<td>Duration of infertility (yrs)</td>
<td>4.27 ± 2.46</td>
<td>4.09 ± 2.80</td>
<td>0.010</td>
</tr>
<tr>
<td>Type of infertility</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Primary</td>
<td>90 (75.6)</td>
<td>177 (64.8)</td>
<td>0.045</td>
</tr>
<tr>
<td>Secondary</td>
<td>29 (24.4)</td>
<td>96 (35.2)</td>
<td></td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>21.31 ± 2.80</td>
<td>20.82 ± 2.56</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>mFG score</td>
<td>0.55 ± 1.09</td>
<td>0.14 ± 0.46</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>WHR</td>
<td>0.82 ± 0.06</td>
<td>0.83 ± 0.07</td>
<td>0.762</td>
</tr>
<tr>
<td>FSH (mIU/mL)</td>
<td>6.07 ± 1.46</td>
<td>7.41 ± 3.49</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>LH (mIU/mL)</td>
<td>9.37 ± 5.31</td>
<td>6.23 ± 3.43</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>LH/FSH</td>
<td>1.57 ± 0.85</td>
<td>0.95 ± 0.63</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>E2 (pg/mL)</td>
<td>43.75 ± 26.55</td>
<td>40.22 ± 40.76</td>
<td>0.137</td>
</tr>
<tr>
<td>Prolactin (µIU/mL)</td>
<td>410.45 ± 243.46</td>
<td>432.26 ± 318.07</td>
<td>0.060</td>
</tr>
<tr>
<td>AMH (ng/mL)</td>
<td>7.35 ± 5.17</td>
<td>3.48 ± 3.17</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Ovarian volume (cm³)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Right ovarian</td>
<td>8.18±3.73</td>
<td>6.69 ± 4.17</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Left ovarian</td>
<td>6.78±3.40</td>
<td>6.46 ± 4.11</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td>Mean volume</td>
<td>7.48±3.11</td>
<td>6.58 ± 3.69</td>
<td>&lt;0.001</td>
</tr>
</tbody>
</table>

**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.

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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.

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. A meta-analysis by Illidromiti et al using ROC analysis figured out an AMH cut-off value of 4.7 ng/mL to obtain 82.8% sensitivity and 79.4% specificity. It is important to highlight that the differences in ethnicity, age of study populations, criteria used to define PCOS, and impaired fertility must be taken into consideration as possible contributing factors to these discordant results.

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

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

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.

Table 3: Correlation Between Anthropometry, Hormonal Profiles and Ovarian Volume

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Right Ovarian Volume</th>
<th>Left Ovarian Volume</th>
<th>Mean Volume</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>( r )</td>
<td>( P ) value</td>
<td>( r )</td>
</tr>
<tr>
<td>Age (yrs)</td>
<td>-0.160</td>
<td>0.001</td>
<td>-0.162</td>
</tr>
<tr>
<td>BMI</td>
<td>0.123</td>
<td>0.015</td>
<td>0.048</td>
</tr>
<tr>
<td>mFG score</td>
<td>0.100</td>
<td>0.052</td>
<td>0.106</td>
</tr>
<tr>
<td>WHR</td>
<td>-0.019</td>
<td>0.705</td>
<td>-0.027</td>
</tr>
<tr>
<td>FSH (mIU/mL)</td>
<td>-0.129</td>
<td>0.010</td>
<td>-0.087</td>
</tr>
<tr>
<td>LH (mIU/mL)</td>
<td>0.193</td>
<td>&lt;0.001</td>
<td>0.219</td>
</tr>
<tr>
<td>E2 (pg/mL)</td>
<td>-0.058</td>
<td>0.254</td>
<td>0.029</td>
</tr>
<tr>
<td>Prolactin (µIU/mL)</td>
<td>-0.058</td>
<td>0.251</td>
<td>-0.019</td>
</tr>
<tr>
<td>AMH (ng/mL)</td>
<td>0.281</td>
<td>&lt;0.001</td>
<td>0.263</td>
</tr>
</tbody>
</table>

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.
PCOS, and type of AMH assays may influence on the results and proposed diagnostic cut-off.\textsuperscript{39–41}

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.\textsuperscript{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.\textsuperscript{19} 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.\textsuperscript{21} 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.\textsuperscript{12,42} Ndoua et al demonstrated that AMH showed a good correlation with OV (\( r = 0.625, p = 0.0001 \)).\textsuperscript{14} In addition, we found a weak negative correlation between OV and age. This finding was also reported by Erdem et al (with \( r - r = -0.29, p < 0.05 \)).\textsuperscript{43} As OV in women aged 25–51 years reflects the number of primordial follicles,\textsuperscript{44} 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.\textsuperscript{45}

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

\begin{figure}
\centering
\includegraphics[width=\textwidth]{roc_curve.png}
\caption{ROC curve of prediction PCOS by AMH (A) and by mean of ovarian volume (B). Diagonal segments are produced by ties.}
\end{figure}

\begin{table}
\centering
\begin{tabular}{|c|c|c|c|c|}
\hline
Parameters & Threshold & Sensitivity & Specificity & AUC, \( p \) \\
\hline
Mean of ovarian volume & 4.575 & 0.924 & 0.333 & 0.613 (0.557–0.670) \\
& 6.030 & 0.613 & 0.505 & \\
AMH (ng/mL) & 4.195 & 0.731 & 0.707 & 0.787 (0.741–0.834) \\
\hline
\end{tabular}
\caption{Diagnostic Values of Anti-Mullerian Hormone and Ovarian Volume for Polycystic Ovary Syndrome}
\end{table}

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.46,47 However, other studies suggested that no difference was observed in the clinical biochemical profiles of patients with different phenotypes.48,49 Similar to our ultrasound findings, Clark et al, and Sachdeva et al have also described smaller ovarian volumes in phenotype B.46,50 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.48,49

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 age-matched 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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