

Ovarian function and ovarian blood supply following premenopausal abdominal hysterectomy

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Abstract

Introduction: The issue of conserving the ovaries at hysterectomy in premenopausal women with benign gynecologic disease has been the subject of considerable controversy. Some clinicians prefer prophylactic oophorectomy in premenopausal women during hysterectomy to prevent future development of malignant changes in conserved ovaries. Other clinicians prefer to conserve apparently normal ovaries, because bilateral oophorectomy in premenopausal women results in an abrupt imbalance, sudden onset of menopausal symptoms, decreased libido, increased cardiovascular risk and osteoporosis.

Material and methods: Two hundred and twenty multipara women (who had completed their families), with benign uterine pathology were included in this prospective study for abdominal hysterectomy with bilateral ovarian preservation. Pre-operative vaginal ultrasound, Doppler studies, diagnostic hysteroscopy and endometrial biopsy were done followed by laboratory studies including Anti-mullerian hormone (AMH), follicle stimulating hormone (FSH) and estradiol for all studied women. Doppler studies, AMH, FSH and estradiol were repeated 6 and 12 months post-operative for assessment of the ovarian function and ovarian blood supply after hysterectomy.

Results: Pre-operative AMH, FSH and estradiol of the studied women were statistically insignificant compared to AMH, FSH and estradiol 6 and 12 months post-operative. Twelve months post-operative right and left ovarian volumes (6.92 ± 0.18 and 6.85 ± 0.19 cm³, respectively) were significantly larger than pre-operative right and left ovarian volumes (6.19 ± 0.22 and 5.86 ± 0.23 cm³, respectively), and, 12 months post-operative right and left ovarian pulsatility indices (2.92 ± 0.15 and 2.96 ± 0.16 cm/s, respectively) were significantly lower than pre-operative right and left ovarian pulsatility indices (3.45 ± 0.19 and 3.36 ± 0.2 cm/s, respectively). Eight (3.6%) cases of the studied women developed an ovarian cyst 6 months after hysterectomy, 3 were spontaneously resolved and the remaining 5 (2.27%) cases underwent exploratory laparotomy.

Conclusions: There is no evidence of ovarian dysfunction affecting conserved ovaries one year after hysterectomy in premenopausal women as evident by AMH, FSH and estradiol. Furthermore, an increased ovarian volume and reduced ovarian pulsatility indices indicate a possible increase in ovarian blood supply, and preserved non-compromised ovarian function.

Key words: ovarian, function, blood supply, premenopausal, abdominal hysterectomy.

Introduction

The issue of conserving the ovaries at hysterectomy in premenopausal women with benign gynecologic disease has been the subject of considerable controversy [1-5]. Some clinicians prefer prophylactic oophorectomy in premenopausal women during hysterectomy to prevent future development of malignant changes in conserved ovaries [6-16].

Other clinicians prefer to conserve apparently normal ovaries, because bilateral oophorectomy in premenopausal women results in an abrupt imbalance, sudden onset of menopausal symptoms, decreased libido, increased cardiovascular risk and osteoporosis [17-23].

There is conflicting evidence regarding the effect of premenopausal hysterectomy on the conserved ovarian function. Some research indicates an increase in incidence and severity of menopausal symptoms following removal of the uterus despite the ovaries remaining in place, which may be due to reduction in ovarian blood flow and follicular atresia [6]. Animal studies (rat models) concluded that hysterectomy with ovarian conser-

vation results in an abrupt imbalance, sudden onset of menopausal symptoms, decreased libido, increased cardiovascular risk and osteoporosis [17-23].

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vation could preserve a normal hormonal milieu but might accelerate the onset of early menopause [24, 25]. This prospective study was designed to detect the effect of premenopausal abdominal hysterectomy on the ovarian function and ovarian blood supply.

Material and methods

Two hundred and twenty (220) women with benign uterine pathology were included in this study for abdominal hysterectomy with bilateral ovarian preservation which was conducted in Ahmadi, Kuwait Oil Company (KOC) and Sabah Maternity Hospitals, Kuwait, from January 2012 to January 2014. Women included in this study were 38-45 years old, multipara (had completed their families), with benign uterine pathology. Women planning future pregnancy or with other gynecological problems indicating alternative surgery such as endometriosis or cervical intraepithelial neoplasia (CIN) or ovarian masses were excluded from this study and women who had one or both ovaries removed in a previous surgery or malignant pelvic pathology or follicle stimulating hormone (FSH) of more than 20 IU/ml (on at least two sessions) were also excluded from this study. Women were included in this study after informed consent, proper counseling and approval of the study protocol by the institute's ethics committee. After complete history, physical examination, preoperative vaginal ultrasound, Doppler studies, diagnostic hysteroscopy and endometrial biopsy were done for all studied women, followed by pre-operative laboratory studies including Anti-mullerian hormone (AMH), FSH and estradiol. Doppler studies, AMH, FSH and estradiol were repeated 6 and 12 months post-operative for assessment of the ovarian function and blood supply after abdominal hysterectomy. Blood samples were collected from women included in this study pre-operative, 6 and 12 months after hysterectomy in plain tubes, centrifuged and serum samples were stored at -20°C . AMH, FSH and estradiol levels in serum were measured by electrochemiluminescence immunoassay (ECLIA) using Elecsys Kits (Roche Diagnostics, Mannheim, Germany). Ovarian volume was measured in the pelvic infundibulum by transvaginal ultrasound and Doppler ultrasonography was used to assess the ovarian flow velocity waveform by measuring the Pulsatility Index (PI; cm/s) pre-operative, and 6 and 12 months after hysterectomy. All examinations were conducted during the follicular phase of the menstrual cycle by the same investigator, in the afternoon. In hysterectomized women, the follicular phase was determined by the absence of both a dominant follicle and corpus luteum [18]. Doppler ultrasonography of the ovarian arteries was done at the follicular phase of the menstrual cycle because flow is reduced and similar in both ovaries during this period [18].

Transvaginal ultrasound and Doppler studies were done using Philips HD9 (Philips International; Amsterdam; Netherlands) with a two-dimensional endo-vaginal convex probe 4-9 MHz by a sonographer who was blinded to the patients' criteria.

Baseline measures included transverse (T), antero-posterior (AP), and longitudinal (L) diameters of both ovaries and ovarian volume (V), was estimated using the formula $V (\text{cm}^3) = T (\text{cm}) \times AP (\text{cm}) \times L (\text{cm}) \times 0.52$. Left and right ovarian artery flow in the pelvic infundibulum was visualized with the color Doppler technique and the typical velocity spectrum of this vessel was determined. Blood flow impedance was expressed as the PI (cm/s). The PI values were calculated electronically according to the formula $PI = \text{peak systolic velocity} - \text{minimum diastolic velocity} / \text{mean flow velocity}$.

Sample size justification

The required sample size was calculated using G*Power software version 3.17 for sample size calculation (*Heinrich Heine Universität, Düsseldorf, Germany), setting α -error probability at 0.05, power ($1 - \beta$ error probability) at 0.95 % and effective sample size (w) at 0.3. The effective size (w) was calculated as follows: $w = \sqrt{\chi^2 / N}$, where χ^2 is the chi-square test and N is the total sample size. The number of participants needed to produce a statistically acceptable figure was 220 women.

Statistical analysis

Data were collected, tabulated, then statistically analyzed using the Statistical Package for Social Sciences (SPSS) computer software version 18. Numerical variables were presented as mean and standard deviation (\pm SD), while categorical variables were presented as a number and percentage. Student t -test was used for comparison between groups as regards quantitative variables. A difference with a p value < 0.05 was considered statistically significant.

Results

Mean age of premenopausal women included in this study for hysterectomy was 42.3 ± 8.7 years, parity was 4.9 ± 1.6 and body mass index (BMI) was 32.1 ± 2.07 kg/m². Hysterectomy was indicated for studied women due to fibroid uterus 131 (59.6%) cases (causing pelvic-abdominal mass 51 [23.2%] cases, menorrhagia 47 [21.4%] cases, pelvic pain or pressure 33 [15%] cases), heavy menstrual bleeding (HMB) with failed medical and hormonal treatment 52 (23.6%) cases and polymenorrhea 37 (16.8%) cases. The pre-operative histology of endometrium samples showed secretory endometrium in 87 (39.5%) cases, proliferative endometrium in

Tab. I. Indications of hysterectomy for the studied population and preoperative histology of endometrial samples

Variables	Number (%)
Indication of hysterectomy	
Fibroid uterus	131 (59.6%)
Pelvic-abdominal mass	51 (23.2%)
Menorrhagia	47 (21.4%)
Pelvic pain or pressure symptoms	33 (15%)
Heavy menstrual bleeding (HMB)	52 (23.6%)
Polymenorrhea	37 (16.8%)
Pre-operative histology of endometrium samples	
Secretary endometrium	87 (39.5%)
Proliferative endometrium	69 (31.4%)
Simple hyperplasia	36 (16.4%)
Complex hyperplasia without atypia	28 (12.7%)

69 (31.4%) cases, simple endometrial hyperplasia in 36 (16.4%) cases and complex hyperplasia without atypia in 28 (12.7%) cases (Table I).

Pre-operative AMH (1.75 ± 4.61 ng/ml) of the studied women was statistically insignificant compared to AMH 6 and 12 months post-operative (1.78 ± 2.45 and 1.81 ± 2.19 ng/ml, respectively) and pre-operative FSH (7.98 ± 5.7 IU/ml) was statistically insignificant compared to FSH 6 and 12 months post-operative (8.26 ± 5.4 and 8.55 ± 6.2 IU/ml, respectively), also, pre-operative estradiol (129 ± 57.3 pg/ml) was statistically insignificant compared to estradiol 6 and 12 months post-operative (134.5 ± 66.2 and 139.3 ± 77.1 pg/ml, respectively) (Table II).

Tab. II. Preoperative and postoperative Anti-mullerian hormone (AMH), follicle stimulating hormone (FSH), estradiol, ovarian volume, ovarian Pulsatility Index (PI) of the studied population

Variables	Preoperative	6 months postoperative	12 months postoperative	P value (95% CI) test used
AMH (ng/ml), mean ± SD	1.75 ± 4.61	1.78 ± 2.45	1.81 ± 2.19	P1 = 0** (CI: -0.71; -0.03; 0.65), <i>t</i> test P2 = 0** (CI: -0.73; -0.06; 0.61), <i>t</i> test
FSH (IU/ml), mean ± SD	7.98 ± 5.7	8.26 ± 5.4	8.55 ± 6.2	P1 = 0.21** (CI: -1.31; -0.28; 0.75), <i>t</i> test P2 = 0.89** (CI: -1.68; -0.57; 0.54), <i>t</i> test
Estradiol (pg/ml), mean ± SD	129.0 ± 57.3	134.5 ± 66.2	139.3 ± 77.1	P1 = 0.98** (CI: -17.0; -5.5; 0.06), <i>t</i> test P2 = 0.99** (CI: -22.9; -10.3; 2.39), <i>t</i> test
Right ovarian volume (cm ³), mean ± SD	6.19 ± 0.22	6.75 ± 0.25	6.92 ± 0.18	P1 = 0.97** (-0.60; -0.56; -0.51), <i>t</i> test P2 = 0.001* (-0.82; -0.79; -0.75), <i>t</i> test
Left ovarian volume (cm ³), mean ± SD	5.86 ± 0.23	6.57 ± 1.12	6.85 ± 0.19	P1 = 1** (-0.86; -0.71; -0.55), <i>t</i> test P2 = 0.002* (-1.02; -0.99; -0.95), <i>t</i> test
Right ovary Pulsatility Index (cm/s), mean ± SD	3.45 ± 0.19	3.12 ± 0.21	2.92 ± 0.15	P1 = 0.9** (0.29; 0.33; 0.36), <i>t</i> test P2 = 0.002* (0.48; 0.53; 0.56), <i>t</i> test
Left ovary Pulsatility Index (cm/s), mean ± SD	3.36 ± 0.2	3.07 ± 0.21	2.96 ± 0.16	P1 = 0.9** (0.25; 0.2; 0.32), <i>t</i> test P2 = 0.0005* (0.36; 0.4; 0.43), <i>t</i> test

**Non-significant, *Significant

P1 – *p* for preoperative values compared to 6 months postoperative values, P2 – *p* for preoperative values compared to 12 months postoperative values, *t* test – Student *t*-test, cm/s – cm/second

Pre-operative right and left ovarian volumes (6.19 ± 0.22 and 5.86 ± 0.23 cm³, respectively) were statistically insignificant compared to 6 months post-operative right and left ovarian volumes (6.75 ± 0.25 and 6.57 ± 1.12 cm³, respectively), and were significantly smaller compared to 12 months post-operative right and left ovarian volumes (6.92 ± 0.18 and 6.85 ± 0.19 cm³, respectively) (Table II).

Pre-operative right and left ovarian pulsatility indices (3.45 ± 0.19 and 3.36 ± 0.2 cm/s, respectively) were statistically insignificant compared to 6 months post-operative right and left ovarian pulsatility indices (3.12 ± 0.21 and 3.07 ± 0.21 cm/s, respectively), and were significantly higher compared to 12 months post-operative right and left ovary pulsatility indices (2.92 ± 0.15 and 2.96 ± 0.16 cm/s, respectively) (Table II).

Eight (3.6%) cases of the studied women developed an ovarian cyst 6 months after hysterectomy, 3 were spontaneously resolved and the remaining 5 (2.27%) cases underwent exploratory laparotomy which revealed 3 cases of serous cystadenoma and 2 cases of a paraovarian cyst.

Discussion

In this study, there is no evidence of ovarian dysfunction affecting conserved ovaries one year after abdominal hysterectomy in premenopausal women as evident by AMH, FSH and estradiol.

There is conflicting evidence regarding the effect of premenopausal hysterectomy on the ovarian function. Some research indicates an increase in incidence and severity of menopausal symptoms and ovarian failure following removal of the uterus, despite the ovaries

remaining in place [26], whereas others report no decrease in ovarian function [26].

The advance of menopause age after hysterectomy is related to an increased rate of follicular atresia (surgical removal of uterus will increase follicular atresia in conserved ovaries) [27]. The presence of uterus would inhibit follicle depletion or atresia and its surgical removal at reproductive age would accelerate follicular loss, atresia and subsequent accelerated menopause [27].

Other hypothesis, the increased prevalence of ovarian failure after hysterectomy, is due to stretch and thrombosis of ovarian blood vessels with a subsequent reduction in ovarian blood supply [17, 27, 28].

Deng *et al.* concluded that hysterectomy with the conservation of bilateral/unilateral ovaries may have some influence on the ovarian function [17], also, Ahn *et al.* concluded that total abdominal hysterectomy accelerates ovarian dysfunction and women treated with total abdominal hysterectomy are at risk of early menopause [6].

On the contrary, Ylikorkala and Viinikka studied pituitary-ovarian function in 2 women with congenital absence of the uterus and vagina (Mayer-Rokitansky-Kuster-Hauser syndrome) and concluded that presence or absence of the uterus does not affect the ovarian function [29].

In this study, pre-operative AMH, FSH and estradiol were statistically insignificant compared to AMH, FSH and estradiol 6 and 12 months after abdominal hysterectomy, also, Chalmers *et al.*, concluded that there is no evidence of compromise of the ovarian function, as reflected in FSH levels, within 2 years of hysterectomy [19].

Findley *et al.* concluded that laparoscopic hysterectomy \pm salpingectomy with ovarian preservation does not appear to have any short-term deleterious effects on ovarian reserve, as measured by the AMH level [30].

Morelli *et al.* compared women treated with total laparoscopic hysterectomy (TLH) plus bilateral salpingectomy, with women treated by TLH without adnexectomy and they found no significant difference between two groups regarding AMH, FSH, antral follicle count (AFC), mean ovarian diameters and peak systolic velocity [31].

Recently, Venturella *et al.* has concluded that OvAge is one of the first reliable attempts to create a new method able to identify ovarian reserve [32, 33].

Although, Ishii *et al.* found that fifteen of 33 patients became climacteric after premenopausal radical hysterectomy for stage IB and II cervical cancer with ovarian preservation, they also found a significant correlation between ovarian dysfunction after radical hysterectomy and age [34]. Petri Nahás *et al.* found that ovarian volumes were greater 6 and 12 months after total abdominal hysterectomy compared to controls and they found reduced PI of ovarian vessels of hysterectomized women compared to controls [18].

They concluded that the reduced PI of ovarian vessels of hysterectomized women indicates decreased resistance with a subsequent increased ovarian blood flow in hysterectomized women compared to controls, also, in this study, 12 months post-operative right and left ovarian volumes (6.92 ± 0.18 and 6.85 ± 0.19 cm³, respectively) were significantly larger than pre-operative right and left ovarian volumes (6.19 ± 0.22 and 5.86 ± 0.23 cm³, respectively), and, 12 months post-operative right and left ovary pulsatility indices (2.92 ± 0.15 and 2.96 ± 0.16 cm/s, respectively) were significantly lower than pre-operative right and left ovarian pulsatility indices (3.45 ± 0.19 and 3.36 ± 0.2 cm/s, respectively) [18].

Five to eight percent of hysterectomized women require subsequent surgeries for benign ovarian diseases [5]. The post-hysterectomy ovarian cysts appear within the first post-operative year and spontaneously resolved in more than 50% of cases, and most of these cysts are functional cysts [35]. Zalel *et al.* found ovarian cysts in 50.7% of hysterectomized women (37/73) and Pete *et al.* found ovarian cysts in 9.2% of hysterectomized women (6/65) [36, 37].

Four women were lost during follow up (excluded from the study) and short duration of post-operative follow up (one year) were the two limitations faced during this study.

Conclusions

There is no evidence of ovarian dysfunction affecting conserved ovaries one year after hysterectomy in premenopausal women as evident by AMH, FSH and estradiol. Furthermore, an increased ovarian volume and reduced ovarian PI indicates a possible increase in ovarian blood supply, and preserved, non-compromised ovarian function.

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Disclosure

Authors declare no conflict of interest.

References

1. Yavuzcan A, Yıldız G, Çağlar M, et al. Which one is safer – performing a laparoscopic hysterectomy with a tissue fusion device involving diagnostic cystoscopy or traditional abdominal hysterectomy with ureteral dissection? *Videosurgery Miniinv* 2013; 8: 280-288.
2. Grabowski A, Korlacki W, Pasierbek M. Laparoscopy in elective and emergency management of ovarian pathology in children and adolescents. *Videosurgery Miniinv* 2014; 9: 164-169.
3. Wilczyński M, Cieślak J, Malinowski A. Supracervical hysterectomy – the vaginal route. *Videosurgery Miniinv* 2014; 9: 207-212.

4. Ceccaroni M, Roviglione G, Pesci A, et al. Total laparoscopic hysterectomy of very enlarged uterus (3030 g): case report and review of the literature. *Videosurgery Miniinv* 2014; 9: 302-307.
5. Reich H. Issues surrounding surgical menopause. Indications and procedures. *J Reprod Med* 2001; 46 (3 Suppl): 297-306.
6. Ahn EH, Bai SW, Song CH, et al. Effect of premenopausal hysterectomy on ovarian function. *Yoseni Med J* 2002; 43: 53-58.
7. Rogala E, Nowicka A, Bednarek W, et al. Evaluation of the expression of the immunosuppressive enzyme – indoleamine 2,3-dioxygenase in ovarian cancer tissue. *Prz Menopauzalny* 2013; 17: 223-227.
8. Gottwald L, Danilewicz M, Fendler W, et al. The AgNORs count in predicting long-term survival in serous ovarian cancer. *Arch Med Sci* 2014; 10: 84-90.
9. Moulla A, Miliaras D, Sioga A, et al. The immunohistochemical expression of CD24 and CD171 adhesion molecules in borderline ovarian tumors. *Pol J Pathol* 2013; 64: 180-184.
10. Denel M, Marczak A. Panels of protein biomarkers and non-protein markers in the diagnosis of the ovarian cancer. *Prz Menopauzalny* 2013; 17: 404-408.
11. Tkaczuk-Włach J, Substyl M, Jakiel G. Biochemical markers for screening of ovarian cancer. *Prz Menopauzalny* 2013; 17: 442-445.
12. Smolarz B, Makowska M, Samulak D, et al. Association between polymorphisms of the DNA repair gene. RAD51 and ovarian cancer. *Pol J Pathol* 2013; 64: 290-295.
13. Lipińska N, Rubiś B. Telomerase as a target in diagnosis and treatment of cancer in postmenopausal women. *Prz Menopauzalny* 2013; 17: 478-483.
14. Marczak A, Bukowska B. New trends in the ovarian cancer treatment. *Prz Menopauzalny* 2013; 17: 489-492.
15. Dębska-Szmich S, Czernek U, Krakowska M, et al. Synchronous primary ovarian and endometrial cancers: a series of cases and a review of literature. *Prz Menopauzalny* 2014; 13: 64-69.
16. Marczak A, Denel M. Trabectedin as a single agent and in combination with pegylated liposomal doxorubicin – activity against ovarian cancer cells. *Contemp Oncol (Pozn)* 2014; 18: 149-152.
17. Deng CY, Tang DM, Yu Q, He FF. Effect of premenopausal hysterectomy on ovarian function. *Zhongguo Yi Xue Ke Xue Yuan Xue Bao* 2002; 24: 639-642 [Article in Chinese].
18. Petri Nahás EA, Pontes A, Nahas-Neto J, et al. Effect of total abdominal hysterectomy on ovarian blood supply in women of reproductive age. *J Ultrasound Med* 2005; 24: 169-174.
19. Chalmers C, Lindsay M, Usher D, et al. Hysterectomy and ovarian function: levels of follicle stimulating hormone and incidence of menopausal symptoms are not affected by hysterectomy in women under age 45 years. *Climacteric* 2002; 5: 366-373.
20. Oldenhave A, Jaszman LJ, Everaer WT, Haspelss A. Hysterectomized women with ovarian conservation report, more severe climacteric complaints than do normal climacteric women of the same age. *Am J Gynecol* 1993; 168 (3 Pt 1): 765-771.
21. Lemm MA, Skatba P. Biology of ovarian aging. *Prz Menopauzalny* 2013; 17: 231-234.
22. Bojar I, Witczak M, Stępnia A, et al. Cognitive functions measured with a battery of CNS VS tests and the subjective assessment of memory, concentration impairment and reduction in the quality of life in women after menopause. *Prz Menopauzalny* 2013; 17: 371-377.
23. Stępnia A, Kot K, Witczak M, et al. Impact of consumption of B-group vitamins on cognitive functions of women after menopause. *Prz Menopauzalny* 2013; 17: 464-471.
24. Ozdamar S, Ulger H, Sorkun HC, Mūderris I. Effects of hysterectomy on ovarian morphology and serum FSH level in rats. *Maturitas* 2005; 52: 60-64.
25. Tapisiz OL, Gungor T, Aytan H, et al. Does hysterectomy affect ovarian function? Histopathologic evaluation and serum FSH, inhibin A, and inhibin B levels in an experimental rat model. *Eur J Obstet Gynecol Reprod Biol* 2008; 140: 61-66.
26. Bhattacharya S, Mollison J, Pinion S, et al. A comparison of bladder and ovarian function two years following hysterectomy or endometrial ablation. *Br J Obstet Gynaecol* 1996; 103: 898-903.
27. Derksen JG, Brölmann HA, Wiegerinck MA, et al. The effect of hysterectomy and endometrial ablation on follicle stimulating hormone (FSH) levels up to 1 year after surgery. *Maturitas* 1998; 29: 133-138.
28. Korabel J, Krzysiek J. Assessment of the ovarian reserve in a group of perimenopausal women. *Prz Menopauzalny* 2013; 17: 333-338.
29. Ylikorkkala O, Viinikka L. Pituitary and ovarian function in women with congenitally absent uterus. *Obstet Gynecol* 1979; 53: 137-139.
30. Findley AD, Siedhoff MT, Hobbs KA, et al. Short-term effects of salpingectomy during laparoscopic hysterectomy on ovarian reserve: a pilot randomized controlled trial. *Fertil Steril* 2003; 100: 1704-1708.
31. Morelli M, Venturella R, Mocciano R, et al. Prophylactic salpingectomy in premenopausal low-risk women for ovarian cancer: *Primum non nocere*. *Gynecol Oncol* 2013; 129: 448-451.
32. Venturella R, Lico D, Sarica A, et al. OvAge: a new methodology to quantify ovarian reserve combining clinical, biochemical and 3D-ultrasonographic parameters. *J Ovarian Res* 2015; 8: 21.
33. Abdelazim IA, Belal MM, Makhlof HH. Antimüllerian hormone and antral follicle count as predictors of ovarian reserve and successful IVF. *Asian Pacific Journal of Reproduction (APJR)* 2012; 1: 89-92.
34. Ishii K, Aoki Y, Takakuwa K, Tanaka K. Ovarian function after radical hysterectomy with ovarian preservation for cervical cancer. *J Reprod Med* 2001; 46: 347-352.
35. Richlin SS, Rock JA. Ovarian remnant syndrome. *Gynaecol Endosc* 2001; 10: 111-117.
36. Zalel Y, Lurie S, Beyth Y, et al. Is it necessary to perform a prophylactic oophorectomy during hysterectomy? *Eur J Obstet Gynaecol Reprod Biol* 1997; 73: 67-70.
37. Pete I, Bösze P. The fate of the retained ovaries following radical hysterectomy. *Eur J Gynaecol Oncol* 1998; 19: 22-24.