

ANALYSIS OF WOMEN WITH ENDOMETRIAL HYPERPLASIA MEASURED BY U/S IN THE CONTEXT OF SOCIO-DEMOGRAPHIC AND OTHER RISK FACTORS IN LAST FIVE YEARS IN AL-HILLA CITY IN IRAQ

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ABSTRACT

Objective:-Assessment of presence of risk factors of this disease, year's through highly associated prevalence of the endometrial uterine hyperplasia from 2014-2018 & age-related prevalence of endometrial hyperplasia, in perimenopausal patients with abnormal vaginal bleeding.

Keywords: risk factors, endometrial hyperplasia, vagina bleeding, perimenopausal

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INTRODUCTION

Its pathophysiology is related with the exciting and uninterrupted motivation of the endometrial layers of the uterus ⁽¹⁾ through increase in estrogen levels united with insufficient levels of progesterone ^(2, 3). The risk factors for endometrial hyperplasia comprise obesity, chronic anovulation, polycystic ovarian syndrome, etc ^(4, 5) tamoxifen only treatment, and estrogen-only hormone treatment ⁽⁵⁾.

Endometrial uterine hyperplasia has been categorized into:-

- 1- Simple ENH: - comprised of minor endometrial glandular accumulating & with low risk for development to CA.
- 2- Simple AENH:- comprised of endometrial layer with simple glandular accumulating & cytological atypia & maximum risk for development to CA ⁽⁶⁾.
- 3- Complex EH:- comprised of more endometrial glandular accumulating & intermediary risk of development to endometrial carcinoma.

4- Complex AENH:-consist of endometrial layer with complex glandular accumulating & cytological atypia and the maximum danger of development to carcinoma⁽⁶⁾.

Although the statement that endometrial uterine carcinoma was a highest public gynecologic malignancy in creation, absolutely after breast carcinoma, with an incidence of 23.2/100,000 ladies⁽⁷⁾. Endometrial uterine hyperplasia not only prompts to endometrial carcinoma but its offering with clinical symptoms such as abnormal uterine bleeding, polymenorrhea & menorrhagia, often lead to urgency and outpatient assessments^(6,11,12). Endometrial uterine hyperplasia (EH) is a pathological disorder described by hyperplastic deviations in endometrial glandular and stromal configurations lining of the uterine cavity⁽⁸⁾. Unopposed estrogen lead to motivation of the endometrium causes proliferative glandular epithelial deviations, including glandular transformation, causing inconsistently designed, sporadically scattered glands. The most common risk factors for a progress to (endometrial carcinoma) include Diabetes, obesity, unbalanced estrogen treatment, PCOs, tamoxifen therapy and grand multipara & nulliparity...etc⁽⁴⁾. Female patient, her family and the healthiness staff tolerate an expense & load of diagnostic assessments then medical as well surgical treatment (including taking endometrial biopsy, hystoscopic examination, possible prolonged progestogen treatment, D&C, & hysterectomy)^(6, 8,9). Here were no predictable screening methods presented for recognition of endometrial carcinoma & her precursor scratches earlier. Cancers frequently are distinguished after history taking from the patients about abnormal uterine bleeding then examination & investigation^(10,11). Estimations a prevalence for endometrial uterine hyperplasia need take into attention the diagnoses of it was complete only between women who had endometrial specimen. Owing to the disturbing nature of endometrial specimen, rarely we do endometrial cultures on patients had no complain^(12, 13). Results from this study suggest that half of the ladies with abnormal uterine bleeding, had simple and complex one. Explanation of that was the several forms of endometrial hyperplasia necessity take into justification, that documented diagnostic challenges for endometrial tissues and the disagreement about the pathologic cataloging of endometrial hyperplasia and well-differentiated endometrial cancer^(16,21). Incorrectness in the histopathologic analyses of endometrial hyperplasia had been well demarcated, as well as concerns for associated endometrial carcinoma⁽¹⁴⁾. The WHO94 plan is a most frequently used by pathologists, but transitioning to the endometrial intraepithelial neoplasia (EIN) terms could be more advantage to clinical organization^(1, 12, 16).

MATERIAL AND METHODS

The Incidence of endometrial hyperplasia in ladies presenting with abnormal uterine bleeding were rise in last few years, so we do observational study on inpatients & outpatients clinic in obstetrics and gynecological department in Babylon teaching hospital and some private clinics after taking their permission. Endometrial uterine hyperplasia (EH) is defined as exciting growing for the glands of endometrium & the type of EH determined on the histopathological description of the specimen changes then congregated according to the old and recent classification of the WHO, but nowadays the recent one is more commonly used than the other. One hundred fifty patients who presented with abnormal uterine bleeding were recorded and prevalence of endometrial hyperplasia assessed in them after taking an endometrial sampling either by dilatation and curettage or hystoscopic sampling according to patients wishes and availability of the methods. These ladies were above 30 years old in period between 2014 till 2018. The endometrial thickness were measured by U/S

then take permission from them to do diagnostic dilatation and curate some under GA and the other by hystroscope to take endometrial biopsy then send the sample for histopathological examination to detect the type of hyperplasia according to WHO criteria (previous & recent classification) and do analysis for the results whether simple, complex, atypia or presence of incite malignancy.

RESULTS

Table 1 shows the mean age of women with endometrial hyperplasia was 48.51 ± 8.05 ranging from (30-66). More than half of women with endometrial hyperplasia live in urban area (57.3%), while women who live in rural area 42.7%. Regarding occupation, 58.0% of women were employed in comparison with housewives which represented 42.0% of women.

Table 1: Distribution of socio-demographic factors of women with endometrial hyperplasia

| Variables | Mean±SD | Range |
|------------|------------|---------|
| Age(years) | 48.51±8.05 | (30-66) |
| Residence | | |
| Urban | 86 | 57.3% |
| Rural | 64 | 42.7% |
| Total | 150 | 100% |
| Occupation | | |
| Employed | 87 | 58% |
| Housewife | 63 | 42% |
| Total | 150 | 100% |

Table 2:- shows endometrial thickness for women with abnormal uterine bleeding that was measured by ultrasound was 15.5 ± 10.1 . Three quarter of them was multiparous (78.0%)

Table 2: Distribution of variables of women with abnormal uterine bleeding

| Variables | Mean±SD | Range |
|----------------------------|-----------|--------|
| Endometrial thickness (mm) | 15.5±10.1 | (2-37) |

| | | |
|-------------|-----|------|
| Gravida | | |
| Multiparous | 117 | 78% |
| Nulliparous | 33 | 22% |
| Total | 150 | 100% |

Table 3:- shows old age as a risk factor for AUB 34.7% followed by PCO, grand multiparty and bleeding tendency in a percentage of 19.3%, 16.7% and 15.3% respectively.

Table 3: Distribution of risk factors for abnormal uterine bleeding

| Variables | Number | Percentage |
|-------------------|--------|------------|
| Old age | 52 | 34.7% |
| Polycystic ovary | 29 | 19.3% |
| Grandmultiparty | 25 | 16.7% |
| Bleeding tendency | 23 | 15.3% |
| Diabetes Mellitus | 15 | 10.0% |
| Tamoxifinuse | 6 | 4.0% |
| Total | 150 | 100.0% |

Notes:- grand multiparous means herparity 5 and above

Figure 1 shows that the most common cause of abnormal uterine bleeding was endometrial hyperplasia which represented 50.0% followed by endometrial aplasia and bleeding tendency in a percentage of 34.7% and 15.3% respectively.

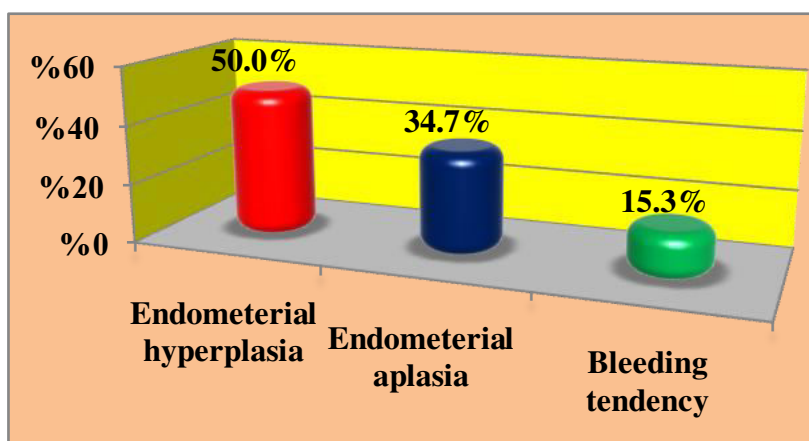


Figure 1: Distribution of the most common causes of uterine bleeding

Figure 2 shows that the percentage of ladies who had endometrial hyperplasia according to year that diagnosis in it, was increase in 2016, 2017 and 2018 in a percentage of 8.0%, 28.0% and 27.3% respectively.

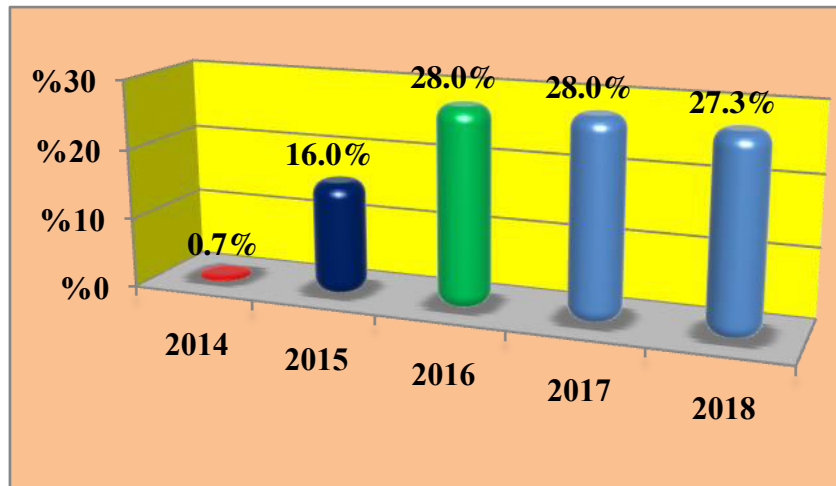


Figure 2: Distribution of women with endometrial hyperplasia according to year of disease diagnosis

Table 4 shows the higher percentage of AUB was hyperplasia (in all types) which represented 50.0% & the most common subtype of HP is Complex typical hyperplasia which represented 16.7% and the most common cause of AUB endometrial aplasia

Table 4: Distribution of causes of abnormal uterine bleeding

| Variables | Number | Percentage |
|------------------------------|------------|-------------|
| Simple typical hyperplasia | 10 | 6.6% |
| Simple atypical hyperplasia | 18 | 12.0% |
| Complex typical hyperplasia | 25 | 16.7% |
| Complex atypical hyperplasia | 22 | 14.7% |
| Aplasia | 48 | 32.0% |
| Others | 27 | 18% |
| Total | 150 | 100% |

Table 5 shows F test was conducted to show if there is a mean differences in age of women with AUB. According to causes (endometrial hyperplasia, endometrial aplasia and bleeding tendency), types of endometrial hyperplasia (simple typical endometrial hyperplasia, simple atypical endometrial hyperplasia, complex typical endometrial hyperplasia, complex atypical endometrial hyperplasia, aplasia and others) and

residence (urban and rural). There are significant mean differences in all circumstances, p value =0.008, <0.001 and 0.047 respectively.

Table 5: Difference mean for women's age rendering to variables

| Variable | groups of study | N | Mean of age | F test | P-value |
|-----------|--|---------|-------------|--------------|-------------------|
| Age(year) | Causes of abnormal uterine bleeding | | | | |
| | Endometrial hyperplasia | 75 | 46.53±7.31 | 4.968 | 0.008* |
| | Endometrial aplasia | 48 | 58.87±7.91 | | |
| | Bleeding tendency | 23 | 49.65±9.28 | | |
| | Total | 150 | | | |
| | Types of abnormal uterine bleeding | | | | |
| | Simple typical hyperplasia | 10 | 38.2±5.43 | 7.418 | <0.001* |
| | Simple atypical hyperplasia | 18 | 47.56±5.53 | | |
| | Complex typical hyperplasia | 25 | 44.6±6.31 | | |
| | Complex atypical hyperplasia | 22 | 51.68±6.25 | | |
| | Aplasia | 48 | 58.44±8.11 | | |
| | Others | 27 | 50.59±8.81 | | |
| | Total | 150 | | | |
| | Residence | | | | |
| | Urban | 86 | 49.64±7.36 | 4.023 | 0.047* |
| Rural | 64 | 47±8.71 | | | |
| Total | 150 | | | | |

*P value ≤ 0.05 is significant

Table (6) demonstrates F test was conducted to show if there is a mean difference in endometrial thickness measured by ultrasound of ladies with uterine endometrial hyperplasia according to types for uterine endometrial hyperplasia (simple typical uterine hyperplasia, simple atypical uterine hyperplasia, complex typical uterine hyperplasia, complex atypical uterine hyperplasia, aplasia and others). There is significant mean difference p value <0.001.

Table 6: Mean difference of endometrial thickness measured by ultrasound according to variables

| Variable | Study groups | N | Mean±SD | F test | P-value |
|----------------------------|---|----|------------|----------------|-------------------|
| endometrial thickness (mm) | Types of endometrial hyperplasia | | | | |
| | Simple typical uterine hyperplasia | 10 | 21.6±2.75 | 115.491 | <0.001* |
| | Simple atypical uterine hyperplasia | 18 | 22.61±3.56 | | |
| | Complex typical uterine hyperplasia | 25 | 26.64±5.69 | | |
| | Complex atypical uterine hyperplasia | 22 | 24.64±5.5 | | |

| | | | | | |
|--|--|--|--|--|--|
| | | | | | |
|--|--|--|--|--|--|

*P value ≤ 0.05 is significant.

Table 7 displays Fisher exact test is conducted to show if there is an association between risk factors of endometrial hyperplasia (tamoxifen use, grand multiparty, PCO, DM, old age and bleeding tendency) and types of endometrial hyperplasia (simple typical uterine hyperplasia, simple atypical uterine hyperplasia, complex typical uterine hyperplasia, complex atypical uterine hyperplasia, aplasia and others).

Table 7: Association between risk factors and causes of abnormal uterine bleeding

| Risk factors | causes of abnormal uterine bleeding | | | | | | P value |
|-------------------|-------------------------------------|-------------------------------------|-------------------------------------|--------------------------------------|------------|------------|-------------------|
| | Simple typical uterine hyperplasia | Simple atypical uterine hyperplasia | Complex typical uterine hyperplasia | Complex atypical uterine hyperplasia | Aplasia | Others | |
| Bleeding tendency | 0(0.0%) | 0(0.0%) | 0(0.0%) | 0(0.0%) | 1(2.1%) | 22(81.5%) | <0.001* |
| DM | 1(10.0%) | 5(27.8%) | 1(4.0%) | 0(0.0%) | 0(0.0%) | 0(0.0%) | |
| Grand multiparty | 8(80.0%) | 6(33.3%) | 3(12.0%) | 8(36.4%) | 0(0.0%) | 0(0.0%) | |
| Old age | 0(0.0%) | 0(0.0%) | 0(0.0%) | 8(36.4%) | 47(97.9%) | 0(0.0%) | |
| PCO | 1(10.0%) | 5(27.8%) | 20(80.0%) | 0(0.0%) | 0(0.0%) | 0(0.0%) | |
| Tamoxifen use | 0(0.0%) | 2(11.1%) | 1(4.0%) | 3(13.6%) | 0(0.0%) | 0(0.0%) | |
| others | 0(0.0%) | 0(0.0%) | 0(0.0%) | 3(13.6%) | 0(0.0%) | 5(18.5%) | |
| Total | 10(100.0%) | 18(100.0%) | 25(100.0%) | 0(0.0%) | 48(100.0%) | 27(100.0%) | |
| | | | | 22(100.0%) |) |) | |
| | | | | | | | |

*P value ≤ 0.05 was significant. F: Fisher-exact test.

Data exploration

Facts admission and exploration is prepared by SPSS version 24 computer software (statistical package for social sciences), definite variables are existing for example incidences and percentages, constant variables are existing for example (mean ± standard deviation). Fisher exact test is as well used to display connotation between two definite variables. F-test is used to conclude the mean differences among groups. P value of ≤ 0.05 is reflected as significant statistically.

DISCUSSION

Our research's findings show that the EH percentage were 50% of women with abnormal uterine bleeding (simple typical EH percentage was 3.3% of ladies with hyperplasia of endometrium; simple atypical endometrial hyperplasia percentage was 6%. While complex typical was 7% and complex atypical was 7.35%, and this is higher than those resulted from other studies in the world. Prior studies that used information from 1985 to 2003, founded that typical EH: 0.121% lady-years; atypical EH: 0.017 woman-years) and carcinoma in situ 0.078% lady-years, correspondingly (Reed et al., 2009; Lacey et al., 2012)⁽¹⁶⁾. Causes of those variations were unidentified; Period when reports are achieved may disturb percentage. Prior researches are created on facts produced before 2002, at that time, old WHO classification was energetically used, however this report studied facts are produced next to new WHO classification⁽¹⁷⁾. Usage of old one had progressively weakened as a new classification was discovered, this might disturb the EH rate. Women age mean is 48.51 ± 8.05 years. EH rate subsequent to perimenopause, determined by Reed et al. is lower than reported in present study⁽¹⁸⁾. Different from unopposed estrogen, estrogen & progestogen mixture did not rise the EH rate⁽¹⁹⁾, signifying more researches were required. Additionally, age-linked endometrial hyperplasia or endometrial carcinoma incidences may differ between different places in the world. High BMI was a hazard cause for endometrial hyperplasia or endometrial carcinoma⁽²⁰⁾. Most common cause for abnormal uterine bleeding in perimenopausal period was endometrial hyperplasia; all causes significantly affected by age of the patients. Other causes of bleeding such as Bleeding tendency, DM, Grand multiparity, Old age, PCO and Tamoxifen use were significantly affect prevalence of EH&EC.

CONCLUSION

Incidence of EH proportions for all types according to WHO criteria considered in this report were slightly higher in last years. This increase till now not identified, may be increased due to infertility, fast food, large percent of women reach menopausal period more than previously due to developed care for elderly female, etc.

RECOMMENDATION

1- Additional reports at the same problem are mandatory in our city & then in our country to evaluate the real cause and rate of the type of endometrial hyperplasia, its sequels and the possible causes of its occurrence and increment of each one.

2- this disease was easily determined, treated and prevent its conversion to malignancy as can as possible. so preferably take an endometrial sampling from any menopausal women present with endometrial thickness > 5 mm.

ETHICAL CLEARANCE

The Research Ethical Committee at scientific research by ethical approval of both environmental and health and higher education and scientific research ministries in Iraq

CONFLICT OF INTEREST

The authors declare that they have no conflict of interest.

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