# **ORIGINAL ARTICLE**

# Association of Postmenopausal Bleeding with Endometrial Carcinoma

SAFIA PERVEEN<sup>1</sup>, MUHAMMAD SULEMAN AZHAR<sup>2</sup>, FATIMA SHAH<sup>3</sup>, FIZZA MAHMOOD<sup>4</sup>, USAMA MAHMOOD<sup>5</sup>

<sup>1</sup>Associate Professors of Obstetrics & Gynaecology, Sahiwal Teaching Hospital and Medical College, Sahiwal

<sup>2</sup>4th Year MBBS Student, Lahore Medical & Dental College, Lahore

<sup>3</sup>Medical Officer, Ali Shirazi Poly Clinic Hospital, Sahiwal

<sup>4</sup>House Officer, Sahiwal Teaching Hospital and Medical College, Sahiwal

<sup>5</sup>Medical Officer, DHQ, Bahawalnagar

Correspondence to Dr. Safia Perveen, E-mail: drsaafia@hotmail.com Cell: 0300-6723575

#### **ABSTRACT**

Aim: To find the association of postmenopausal bleeding with endometrial carcinoma.

Study design: Cross-sectional analytical study

Place and duration of study: Department of Obstetrics & Gynaecology, Sahiwal Teaching Hospital and Medical College, Sahiwal from 1st January 2019 to 31 December 2021.

Methodology: One hundred women suffering from post-menopausal bleeding were enrolled. An endometrial biopsy was used for removing a piece of the tissue from the inner lining of the uterus and was conducted in suspicious women for confirming endometrial carcinoma by microscopic histopathological reporting. The frequency of endometrial carcinoma in post-menopausal

Results: The mean age was 56.2+0.5 years. The recurrent bleeding was observed in 7% cases and obesity was seen in 50 cases. Transvaginal ultrasonography was performed on women due to high suspicions. Out of the total 8 cases, 87.5% had a uterine thickness >5mm with no history of hormone replacement therapy. Incidence of endometrial carcinoma was 7% among postmenopausal women.

Conclusion: Postmenopausal bleeding was commonly observed in obese women and in those who were on hormonal replacement therapy. Incidence of endometrial carcinoma was 7% in postmenopausal bleeding females.

Keywords: Endometrial carcinoma; Women; Diagnosis; Developed countries

## INTRODUCTION

Endometrial carcinoma (EC) is the commonest cancer type among women of developed countries. Its prevalence is around five percent of all the cases of cancer with a morbidity rate as high as 2% among women, globally1. In countries of North America and Europe the incidence of EC is much higher than other developed countries. The attributing factors in EC formation can be aging, early menarche age, obesity, late menopause, post-menopausal estrogen and nulliparity<sup>2</sup>. Unfortunately the incidence of EC has ascended in the recent years instead of decreasing with an expectation of further increase in coming decade<sup>3-10</sup>

Majority of the EC are diagnosed at local staging and can be cured surgically. The survival rate is also around 5 years in more or less 95% of the cases. However, in cases of later staging such as stage IV the five-year survival rate is around 16-45% 11-13 The strategies which can result into early EC detection has yet not been efficiently studied with none of a screening test available for screening population for EC.14 Initiating early detection for EC which target all the women having high risk can help increasing the survival rate in such women.

Post-menopausal bleeding is a most prominent symptom of with a finding of 2/3rd women of all those voiding gynaecological setting to be suffering from it. The protocol for diagnosis of EC combines with transvaginal ultrasonography of such women and hysteroscopy, biopsy and curettage. Postmenopausal bleeding is also linked with non-cancerous condition as endometrial-polyps or un-scheduled bleeding in hormone replacement therapy cases.<sup>15</sup> Still the risk of endometrial cancer remains around 3% in women of western countries having postmenopausal bleeding.16

Present study was designed to find the association of postmenopausal bleeding with endometrial carcinoma. Results of this study will reveal the suspected women that should get evaluated on an early basis for prompt treatment and to increase their life expectancy. This study will also highlight the importance of timely diagnosis of this deadly disease.

Received on 03-01-2022 Accepted on 17-03-2022

#### MATERIAL AND METHODS

After permission from Institutional Ethical Review Board, this cross-sectional analytical research design women suffering from post-menopausal bleeding were enrolled. The study was conducted at Department of Obstetrics & Gynaecology, Sahiwal Teaching Hospital and Medical College, Sahiwal from 1st January 2019 to 31 December 2021.A total of 100 women were between the age of 50-65 years and women who were menopausal and were suffering from bleeding were examined in detail for their endometrial status. Suspected women were undergoing transvaginal ultrasonography. Those women who were taking hormone replacement therapy and their endometrium were >8mm with irregularities in thickness were considered suspicious for endometrial carcinoma, while those women who were not taking hormone replacement therapy and their endometrium were >5mm with irregularities in thickness were considered suspicious for endometrial carcinoma. An endometrial biopsy was used for removing a piece of the tissue from the inner lining of the uterus and was conducted in suspicious women for confirming endometrial carcinoma by microscopic histopathological reporting. The age, post-menopausal age, bleeding frequency, BMI, USG details and histopathological reports were recorded. The frequency endometrial carcinoma in post-menopausal women was analyzed. The SPSS version 25.0 was used for the analysis in terms of frequencies and percentages through chi square tools, mean and standard deviation and also t test for quantitative variables. P value less than 0.05 was considered as significant.

# **RESULTS**

The mean age was 56.2±0.5 years. Majority of the women were between the ages of 56-60 years. The parity status of these women showed that 40% were nulliparous women suffering from postmenopausal bleeding (Table 1).

Within the postmenopausal women suffering from bleeding there were 24% those women who came for a re-visit due to postmenopausal bleeding continuity despite of initial treatment. The recurrent bleeding was observed in 7% cases. Fifteen were those who were obese on recurrent bleeding and 17 were on hormone replacement therapy. The obesity was seen in 50% women (Table 2)

The trans-vaginal ultrasonography was performed on 8 women due to high suspicion of endometrial carcinoma. These patients included recurrent bleeding as well as hormone replacement therapy cases. It was recorded that out of the total 8 cases 87.5% had a uterine thickness >5mm with no history of hormone replacement therapy in them. However, one woman who was on HRT had an endometrial thickness as > 8mm (Table 3).

Women having increased endometrial thickness were undergone endometrial biopsy. There were 3% patients who came for the first time with postmenopausal bleeding and were identified with endometrial carcinoma while 4% were those who had negative finding of EC on first visit but we're confirmed positive on their recurrent bleeding condition. The overall incidence of endometrial carcinoma 7% among postmenopausal women cases (Fig 1).

Table 1: Age and parity status of postmenopausal women (n=100)

Variable	No.	%
Age (years)		
50-55	9	9.0
56-60	70	70.0
>60	21	21.0
Parity		
Nulliparous	40	40.0
Multiparous	60	60.0

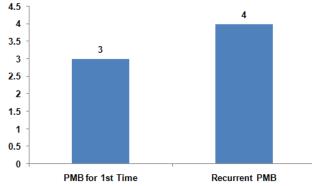
Table 2: Bleeding and obesity history of postmenopausal women (n=100)

PMB women	No.	%		
Re-visit required	24	24.0		
Recurrent bleeding	7	7.0		
Obesity				
Recurrent bleeding	15	15.0		
Hormone replacement therapy	17	17.0		

Table 3: Frequency of endometrial thickness (n=8)

Endometrial thickness	No.	%
>5mm in non-HRT	7	87.5
>8mm in HRT	1	12.5

Fig. 1: Incidence of endometrial carcinoma in postmenopausal women



## DISCUSSION

The recent ascend in endometrial carcinoma and its related mortality rate has raised a dire need of new strategy development for identifying and early detection of EC. Women who are at high risk of developing endometrial carcinoma need to be focused and properly addressed for initial detection to increase their survival rate and life expectancy. Nevertheless, this also assists in avoidance of unnecessary testing of females with low-risk of disease.<sup>17</sup>

The present study results lime lighted the fact that postmenopausal bleeding is strongly related with the endometrial carcinoma. Post-menopausal bleeding is noticed in 90% of endometrial carcinoma cases. However, in present study findings in all the postmenopausal bleeding women the incidence of endometrial carcinoma was only 7%. Current study is in

collaboration with previously reported data which elaborated the presence of endometrial carcinoma in 9% postmenopausal bleeding women<sup>18</sup>.

Effects of use of hormone replacement therapy, regional variance and obesity are significantly observed in a positive relation with endometrial carcinoma in postmenopausal women. Endometrial polyps can lead into bleeding which when suspected for endometrial carcinoma might show negative results on first visit. However, if the poly is left untreated it could result in cancerous form at later stages and thus represent endometrial carcinoma at second visit with recurrent bleeding. Lindenfeld and Langer<sup>19</sup> have described new guidelines for the workup of endometrial carcinoma in all high-risk patients especially in postmenopausal bleeding status.

Endometrial cancer screening is required bas mandatory practice by the available guidelines on EC. High risk population who is under threat of EC needs to be targeted for screening of EC. Among these women were PMB. The present study reassures this practice by identifying a significant percentage of PMB patients who suffer from endometrial carcinoma. Additional triage- testing is required for improving specificity related to EC identification among PMB women and also for avoiding unnecessary biopsies among women with low risk<sup>20</sup>.

There has been a decline in PMB risk post year 2000 however interpretations regarding results is highly significant for proper distinguishing between those women who are high risk population. There have been women with PMB and EC but there have also been those with benign findings and PMB. The number in both groups has escalated over time and might be strongly influenced by factors like hormone replacement therapy, obesity, changes in practices of clinical management as well as abnormal bleeding<sup>20</sup>.

It is important to note that postmenopausal bleeding has been found to be notably lower in cases with hormone replacement therapy as was also seen in the present study. In cases where the HRT consists of estrogen high doses the scenario becomes reversed as it leads to high carcinoma chances<sup>21</sup>.

## CONCLUSION

Postmenopausal bleeding is strongly related with the endometrial carcinoma. Postmenopausal bleeding was commonly observed in obese women and in those who were on hormonal replacement therapy. Incidence of EC was 7% in PMB females.

#### REFERENCES

- Ferlay J, Soerjomataram I, Dikshit R, et al. Cancer incidence and mortality worldwide: sources, methods and major patterns in GLOBOCAN 2012. Int J Cancer 2015; 136(5):E359-86.
- Setiawan VW, Yang HP, Pike MC, et al. Australian National Endometrial Cancer Study Group. Type I and II endometrial cancers: have they different risk factors? J Clin Oncol. 2013;31(20):2607-18.
- Jamison PM, Noone AM, Ries LA, Lee NC, Edwards BK. Trends in endometrial cancer incidence by race and histology with a correction for the prevalence of hysterectomy, SEER 1992 to 2008. Cancer Epidemiol Biomarkers Prev 2013;22(2):233-41.
- Torre LA, Islami F, Siegel RL, Ward EM, Jemal A. Global cancer in women: burden and trends. Cancer Epidemiol Biomarkers Prev 2017; 26(4):444-57.
- Warrko P, Sherman ME, Yang HP, Felix AS, Brinton LA, Trabert B. Recent changes in endometrial cancer trends among menopausal-age US women. Cancer Epidemiol. 2013; 37(4):374-7.
- Siegel RL, Miller KD, Jemal A. Cancer Statistics, 2017. CA Cancer J Clin 2017;67(1):7-30.
- Lortet-Tieulent J, Ferlay J, Bray F, Jemal A. International patterns and trends in endometrial cancer incidence, 1978-2013. J Natl Cancer Inst 2018;110(4):354-61.
- Lindemann K, Eskild A, Vatten LJ, Bray F. Endometrial cancer incidence trends in Norway during 1953-2007 and predictions for 2008-2027. Int J Cancer 2010;127(11):2661-8.
- Gaber C, Meza R, Ruterbusch JJ, Cote ML. Endometrial cancer trends by race and histology in the USA: projecting the number of new cases from 2015 to 2040. J Racial Ethn Health Disparities 2016.

- 10. Sheikh MA, Althouse AD, Freese KE, et al. USA endometrial cancer projections to 2030: should we be concerned? Future Oncol 2014;
- Weiderpass E, Antoine J, Bray FI, Oh JK, Arbyn M. Trends in corpus uteri cancer mortality in member states of the European Union. Eur J Cancer 2014;50(9):1675-84.
- Creasman WT, Odicino F, Maisonneuve P, et al. Carcinoma of the corpus uteri: FIGO 26th Annual Report on the Results of Treatment in Gynecological Cancer. Int J Gynaecol Obstet 2006; 95(suppl 1):S105-43.
- 13. UK CR Uterine cancer statistics.http://www.cancerresearchuk.org/health-professional/cancerstatistics/statistics-by-cancer-type/uterine-cancer. Updated July 3, 2017. Accessed January 18, 2018.
- Kitson SJ, Evans DG, Crosbie EJ. Identifying high-risk women for endometrial cancer prevention strategies: proposal of an endometrial cancer risk prediction model. Cancer Prev Res (Phila) 2017;10(1):1-13
- Van Hanegem N, Breijer MC, Khan KS, et al. Diagnostic evaluation of the endometrium in postmenopausal bleeding: an evidence-based approach. Maturitas 2011;68(2):155-64.

- 16. ACOG. The role of transvaginal ultrasonography in evaluating the endometrium of women with postmenopausal bleeding. Obstet Gynecol 2018; 131:e124-9.
- Clarke MA, Long BJ, Del Mar Morillo A, Arbyn M, Bakkum-Gamez JN, Wentzensen N. Association of Endometrial Cancer Risk With Postmenopausal Bleeding in Women: A Systematic Review and Metaanalysis. JAMA Intern Med 2018;178(9):1210-22.
- Trabert B, Wentzensen N, Yang HP, et al. . Is estrogen plus progestin menopausal hormone therapy safe with respect to endometrial cancer risk? Int J Cancer 2013;132(2):417-26.
- Lindenfeld EA, Langer RD. Bleeding patterns of the hormone replacement therapies in the postmenopausal estrogen and progestin interventions trial. Obstet Gynecol 2002;100(5, pt 1):853-63.
- Hickey M, Ameratunga D, Marino JL. Unscheduled bleeding in continuous combined hormone therapy users. Maturitas 2011; 70(4):400-3.
- van Hanegem N, Prins MM, Bongers MY, et al. The accuracy of endometrial sampling in women with postmenopausal bleeding: a systematic review and meta-analysis. Eur J Obstet Gynecol Reprod Biol 2016;197:147-55.