

Study of endometrial histopathology in patients with abnormal uterine bleeding in a tertiary care Centre.

¹Dr. Nilesh Tatkare, Associate Professor, Dept of Pathology, K. J. Somaiya Medical College, Sion, Mumbai.

²Dr. Meherrituja Palve, Assistant Professor, Dept of Pathology, K. J. Somaiya Medical College, Sion, Mumbai.

³Dr. Smita Sawant, Professor, Dept of Pathology, K. J. Somaiya Medical College, Sion, Mumbai.

⁴Dr. Kalpana Hajirnis, Professor & HOD, Dept of Pathology, K. J. Somaiya Medical College, Sion, Mumbai.

Corresponding Author: Dr. Kalpana Hajirnis, Professor & HOD, Dept of Pathology, K. J. Somaiya Medical College, Sion, Mumbai.

How to citation this article: Dr. Nilesh Tatkare, Dr. Meherrituja Palve, Dr. Smita Sawant, Dr. Kalpana Hajirnis, “Study of endometrial histopathology in patients with abnormal uterine bleeding in a tertiary care Centre”, IJMACR- March - 2023, Volume – 6, Issue - 2, P. No. 430 – 437.

Open Access Article: © 2023, Dr. Kalpana Hajirnis, et al. This is an open access journal and article distributed under the terms of the creative commons attribution license (<http://creativecommons.org/licenses/by/4.0>). Which allows others to remix, tweak, and build upon the work non-commercially, as long as appropriate credit is given and the new creations are licensed under the identical terms.

Type of Publication: Original Research Article

Conflicts of Interest: Nil

Abstract

Introduction: Abnormal uterine bleeding (AUB) is one of the commonest presenting complaints in females of varying age groups. AUB is defined as change in frequency of menstruation, duration of flow or amount of blood loss. It can be caused by various inflammatory, benign or malignant lesions. Microscopic examination of the endometrium aids in the diagnosis of AUB. We did this study to analyse the various histopathological patterns of endometrium in AUB and to observe the incidence of various pathologies in women of different age groups.

Materials and methods: This was a retrospective study done in Department of Pathology on samples received by procedures like D & C, endometrial biopsy of patients complaining of AUB over a period of 3 years from 2019-2021. This histomorphology of the endo-

metrium was studied. The analysis was done in the form of percentages and proportions and represented as tables where necessary.

Results: We studied a total of 271 samples. Maximum samples were received from perimenopausal females (45.86%, n=122). Iatrogenic endometrium was seen in 27% cases (n=72). Proliferative endometrium was seen in 18.42 % (n=49) cases. Secretory endometrium was seen in 22.55 % (n=60) cases. Basal endometrium was seen in 1.12 % cases (n=3). Hyperplasia without atypia was seen in 13.15 % (n=35) cases. Hyperplasia with atypia was seen in 0.75 % (n=2) cases. Endometrial carcinoma was seen in 3.38% (n=9) cases. 6.76 % samples (n=18) were inadequate to opine. 1.87 % samples (n=5) showed products of conception. Endometrial polyp was seen in 5.63% (n=15) cases. 1.12% cases (n=3) showed menstrual endometrium. In the

reproductive age group (<40 years), normal cyclical endometrium was most commonly seen on histopathology. Perimenopausal females (41-50 years) had normal cyclical endometrium followed by iatrogenic endometrium and hyperplasia. Malignancies were most commonly seen in the postmenopausal (>51 years) age group.

Conclusion: Histopathological examination of endometrial samples has diagnostic importance in evaluation of AUB. There is significant correlation between age and endometrial pathologies. Endometrial study hence plays critical role in evaluation and further management of AUB

Keywords: AUB, Endometrial histopathology, Pre and postmenopausal, iatrogenic change, hyperplasia, carcinoma.

Introduction

Abnormal uterine bleeding (AUB) is one of the most common gynecological problems faced by many females of varying age groups. The prevalence of AUB in women between menarche and menopause is around 9-14%. The reported prevalence of AUB in India is around 17.9%. AUB is defined as the change in frequency of menstruation, duration of flow or amount of blood loss. The mean duration of menstruation is 4.7 days and average blood loss per cycle is 35ml.

[1] The FIGO Working Group on Menstrual Disorders has classified the various causes for AUB into structural/organic lesions and non-structural entities. [1] In the absence of medical causes, endometrial sampling and subsequent histopathological examination are the gold standard for the diagnosis of causes of AUB. This study was carried out to analyse the various histopathological patterns of endometrium in AUB and to observe the

incidence of various pathologies in women of different age groups.

Materials and methods

This was a retrospective study conducted in the Department of Pathology in a tertiary care Centre. The endometrial samples received for histopathology from Department of Obstetrics and Gynecology over a period of 3 years were analysed. The endometrial samples were subjected to routine tissue processing after fixation in 10% buffered neutral formalin and the H&E-stained slides prepared were evaluated under light microscope. The endometrial samples were grouped in various age groups like reproductive (18-40 years), perimenopausal (41-50 years) and postmenopausal (>50 years) and correlated with histomorphological pattern. Inclusion criteria: All samples received by procedures like D & C, endometrial biopsy of patients complaining of AUB over a period of 3 years from 2019-2021. Exclusion criteria: Hysterectomy samples. Histopathological findings were divided into two broad categories, AUB due to non-organic and organic causes. Non-organic includes proliferative, secretory, disordered proliferative and atrophic. Organic causes include endometrial carcinoma, tubercular endometritis, endometrial polyp and endometrial hyperplasia. Histopathological findings were correlated with the clinical presentations of the patient.

Statistical analysis Statistical package for social sciences (SPSS - Version 20) was used to carry out the statistical analysis of data. The analysis was done in the form of percentages and proportions and represented as tables where necessary.

Results

We analysed a total of 271 samples from three years 2019 to 2021. We found the age wise distribution of the

samples. Of these, 43.17% (n = 117) were in the reproductive age group (< 40 years), 47.2% (n=128) were in the perimenopausal age group (41-50 years) and 9.6% (n=26) were in the postmenopausal age group (>50 years). Maximum samples were received from females belonging to the perimenopausal age group (45.86%, n=122).

Histomorphological patterns of the samples were studied. Iatrogenic endometrium was seen in 27% cases (n=72). Proliferative endometrium was seen in 18.42% (n=49) cases. Secretory endometrium was seen in 22.55% (n=60) cases. Basal endometrium was seen in 1.12% cases (n=3). Hyperplasia without atypia was seen in 13.15% (n=35) cases. Hyperplasia with atypia was seen in 0.75% (n=2) cases. Endometrial carcinoma was seen in 3.38% (n=9) cases. 6.76% samples (n=18) were inadequate to opine. 1.87% samples (n=5) showed products of conception. Endometrial polyp was seen in 5.63% (n=15) cases. 1.12% cases (n=3) showed menstrual endometrium.

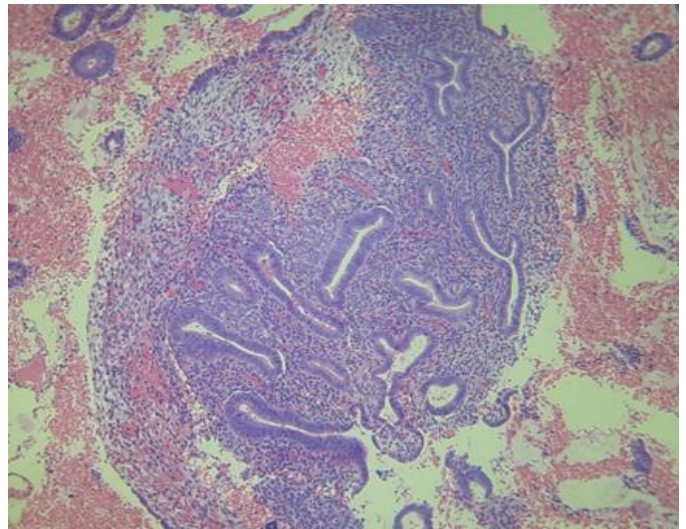


Fig 2. Proliferative Change

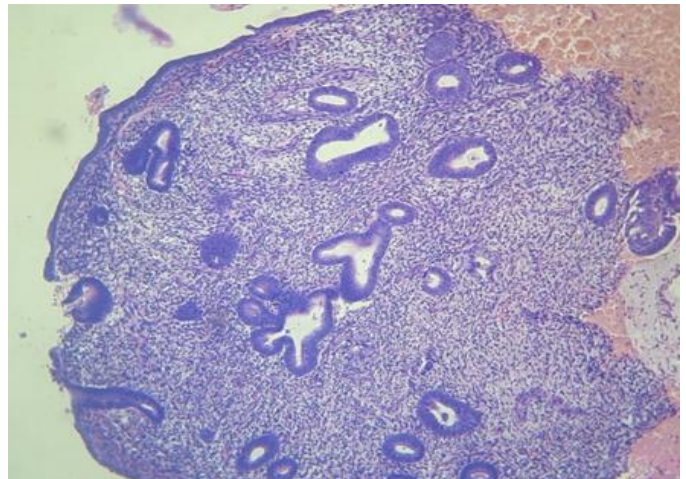


Fig 3. Iatrogenic changes

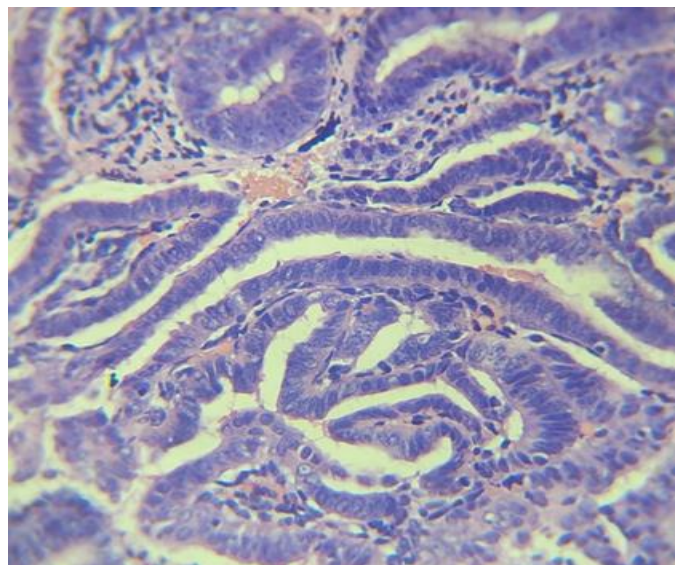


Fig 4. Hyperplasia without atypia

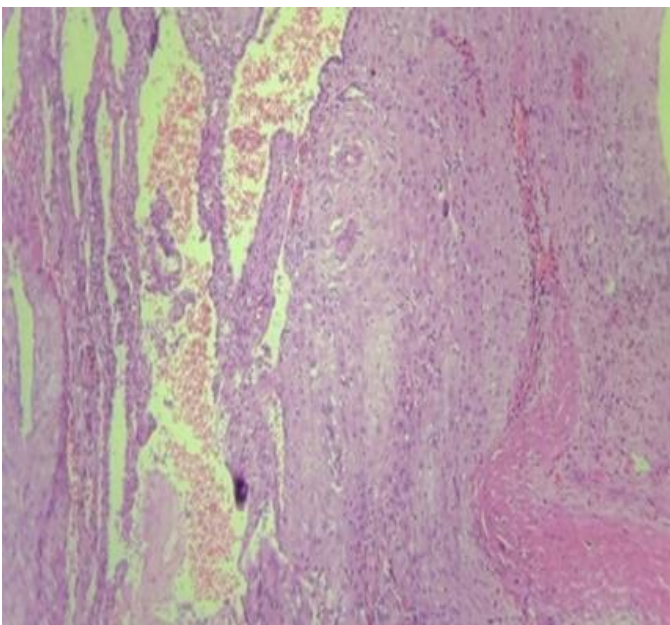


Fig 1. Decidual Change

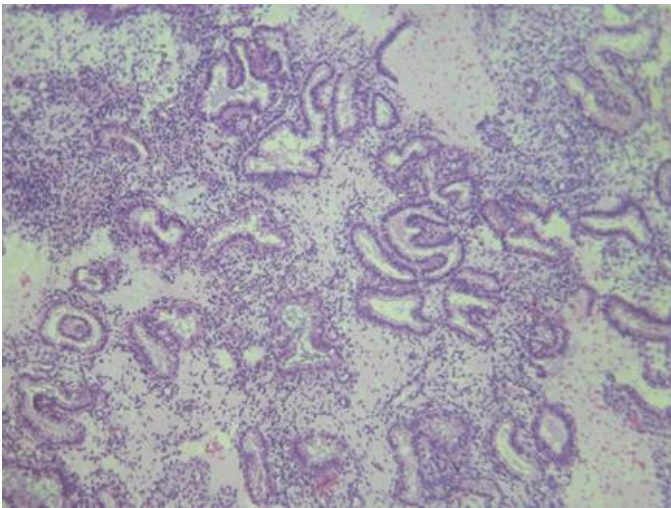


Fig 5. Secretory change

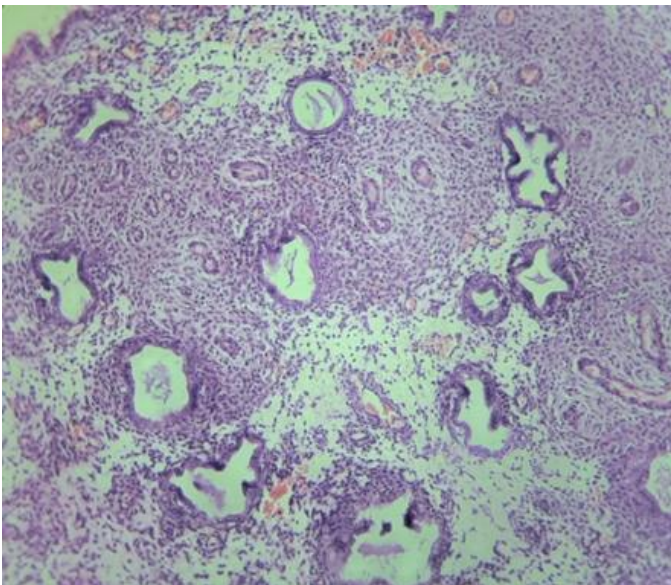


Fig 6. Endometrial polyp

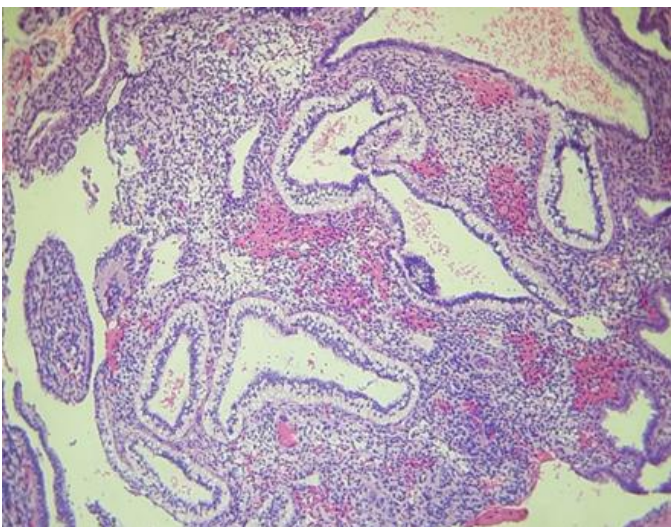


Fig 7. Secretory change

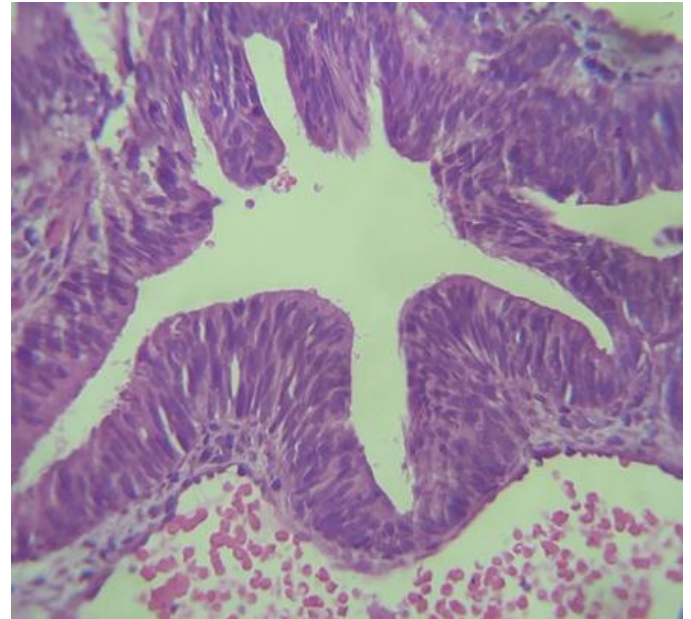


Fig 8. Hyperplasia with atypia

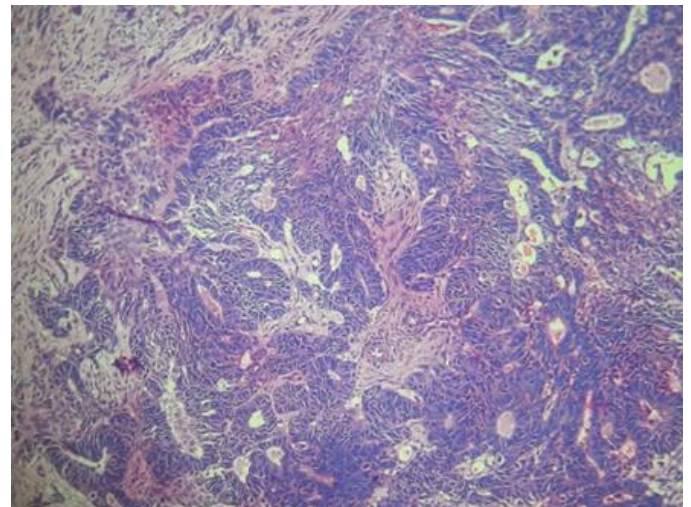


Fig 9. Endometrioid carcinoma

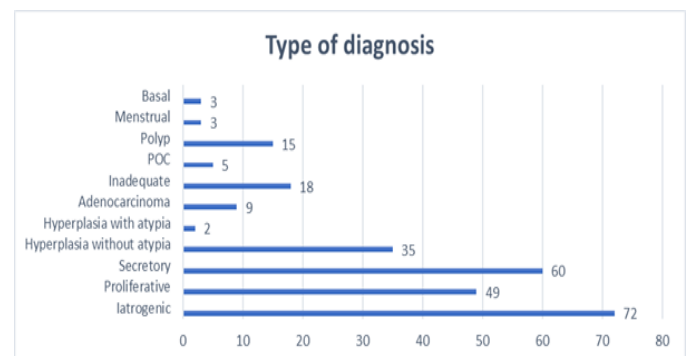


Fig 10. Analysis of the various endometrial patterns was done with respect to the various age groups.

Table 1: Correlation of various endometrial patterns with age groups.

Age	Iatrogenic	Proliferative	Secretory	Hyperplasia Without atypia	Hyperplasia with atypia	Adenocarcinoma	Inadequate	POC	Polyp	Menstrual	Basal
11-20	0	0	0	0	0	0	0	1	0	0	0
21-30	5	7	6	2	0	0	0	3	0	0	0
31-40	23	22	25	14	0	1	5	1	3	0	0
41-50	43	20	27	16	1	2	9	0	4	3	3
51-60	1	1	2	2	1	2	3	0	6	0	0
61-70	0	0	0	1	0	3	0	0	2	0	0
71-80	0	0	0	0	0	0	1	0	0	0	0
81-90	0	0	0	0	0	1	0	0	0	0	0
Total	72	49	60	35	2	9	18	5	15	3	3

Discussion

AUB is defined as change in frequency of menstruation, duration of flow or amount of blood loss. Almost 25% of gynecological operations and 20 % of outpatient visits are due to AUB. [] Prevalence of AUB in women between menarche and menopause is around 9-14%. The reported prevalence of AUB in India is around 17.9%. [1] PALM-COEIN is a useful acronym provided by International Federation of Obstetrics and Gynecology to classify the etiologies of AUB. Causes of AUB are structural like polyps, adenomyosis, leiomyoma, malignancy and hyperplasia. Non-structural causes of AUB are coagulopathy, ovulatory dysfunction, endometrial disorders, iatrogenic and not otherwise classified. [5]

After ruling these out, D and C was done for diagnosis as well as a therapeutic procedure. The reported sensitivity of endometrial biopsy for detection of endometrial abnormalities has been as high as 96%. [1] We have studied the various histopathological patterns of endometrium in patients with AUB and correlated with the different age groups in which they presented.

The incidence of abnormal bleeding increases with increase in age.

In our study the youngest patient was 20 years old and had retained products of conception. Eldest patient was 81 years old and had adenocarcinoma. In our study maximum patients were in the age group of 41-50 years, followed by 31-40 years. This was comparable with many other studies as by Doraiswamy et al, Bindroo et al, Sharma et al and Samal et al. [.] As women approach menopause, decline in the number of ovarian follicles and fluctuations in the estradiol levels cause the cycles to shorten and often become anovulatory leading to various patterns of abnormal bleeding.

The incidence of AUB was low between 51-60 years as compared to 41-50 years. This was similar to the study by Doraiswami et al, Samal et al. [6,9] This may be due to the fact that patients were evaluated and treated early in the course of the disease. In our study most patients showed normal cyclical endometrium on histology i.e proliferative, secretory, menstrual phase.

In our study, proliferative endometrium was seen in 18.42 % cases similar to the study by Dangal et al. [] Secretory endometrium was seen in 22.55 %cases. This

was similar to other studies by Kathleen et al and Iqbal et al. [,] Bleeding in the proliferative phase may be due to anovulatory cycles causing unopposed estrogen to act on the endometrium. Without sufficient progesterone to stabilize and differentiate the endometrium, the mucous membrane becomes fragile and sloughs irregularly. Endometrial study aids in the differentiation between ovulatory and anovulatory cycles.

The bleeding in secretory phase is due to ovulatory dysfunctional uterine bleeding. In ovulatory dysfunctional uterine bleeding, the main defect appears to be in the control of processes regulating the volume of menstrual blood loss, primarily decreased endometrial vasoconstriction and vascular hemostatic plug formation. [2]

Most patients which showed normal cyclical endometrium on histopathology were in the age group between 31- 50 years similar to study by Vani et al. [2] Iatrogenic endometrium was seen in 27% cases. Iatrogenic endometrium referred to endometrium showing effects of exogenous hormonal treatment. The histopathology shows small inactive glands set in a fibrotic stroma showing pseudo decidualization. It was higher than other studies. [8, 12,] The higher incidence may be due to the fact that most patients visit a primary healthcare before being referred to a tertiary Centre like ours where they are given hormonal treatment as first line of management for the bleeding. Most patients with this pattern were in the perimenopausal age group of 41-50 years. Hyperplasia without atypia was seen in 13.15 % cases and most patients belonged to the premenopausal age group of 41-50 years.

Atypia was seen only in 0.75 % cases. This was similar to the findings of Dangal et al, Veena et al and Baral et al. [10,] Endometrial hyperplasia is commonly seen in

perimenopausal age group due to failure of ovulation. Heavy bleeding is secondary to sustained level of estrogens. The epithelial overgrowth affects glands and stroma and there is abnormal vascularization leading to prolonged and excessive bleeding.[12] As is well known endometrial hyperplasia is a precursor of endometrial carcinoma with the overall risk of progression being 5-10%. [13] Hence, it is imperative to identify this lesion before its progression.

Endometrial carcinoma was seen in 3.38 % cases. In our study out of 9 cases, one case was of high grade serous papillary carcinoma and all others were endometrioid carcinoma. The incidence of endometrial carcinoma in our study was similar to the findings of Varun et al, Doraiswami et al and Iqbal et al. [3,6,12]

Even lower incidences were reported by Rajagopal et al, Vani et al, Kathleen et al and Prathippa et al. [1,2,6,13] When compared to the international studies, the low incidence of endometrial carcinoma in India can be attributed to the early child bearing age and multiparity of females which reduces the proliferative activity of endometrium.

Endometrial carcinoma can occur as a result of excess estrogenic stimulation and developing against a background of endometrial hyperplasia or de novo combined with insufficient progesterone levels.

Endometrioid endometrial carcinoma is the most common form of endometrial cancer. Similar to the findings of other studies, in our study too, maximum cases of endometrial carcinoma were seen in the post-Menopausal age group. [3,11,12,13] Benign endometrial polyp was seen in 5.63% cases. This was similar to the study by Kathleen et al. [11] Incidence of polyps was lesser in studies by Vani et al and Prathippa et al. [2,13] It was higher in the study by Varun et al and

Doraiswami et al. [3,6] Most cases of polyp have been observed in the age group of 51-60 years. Retained products of conception were seen in 1.87 % cases. In reproductive age group, complications of pregnancy should be ruled out in patients presenting with abnormal uterine bleeding. 6.76% samples were inadequate to opine in this study due to various reasons like inadequate sampling or the material showing only blood or mucous.

Conclusion

Endo metrial sampling is a helpful diagnostic tool in the evaluation of AUB. Study of endometrium helps reveal many structural and functional causes of AUB which can be seen in the form of different his to patho logical patterns.

These endometrial pathologies have significant correlation with the age of the patient. Endo metrial hyperplasia is common in the peri Meno pausal age group and carcinomas are commonly seen in the Meno pausal age group. Therefore, endo metrial study must be done in the patients presenting with AUB to diagnose these conditions and provide timely intervention and reduce morbidity and mortality.

References

1. Rajagopal I, Thomas BM, Rao VNKR. Endometrial pathology in abnormal uterine bleeding. *Int J Res Med Sci* 2019; 7:3762-6.
2. Vani B. S, Vani R, Jijiya Bai P. Histopathological evaluation of endometrial biopsies and curettings in abnormal uterine bleeding. *Trop J Path Micro* 2019;5(4): 190-197.doi:10.17511/jopm. 2019.i04.02.
3. Varun N, Gupta N, Khan S. A retrospective study of endometrial his to patho logy in abnormal uterine bleeding patients. *Int J Reprod Contracept Obstet Gynecol* 2018; 7:4116-9

4. Sajitha K, Padma SK, Shetty KJ, Kishan Prasad HL, Permi HS, Hegde P. Study of histopathological patterns of endometrium in abnormal uterine bleeding. *CHRISMED J Health Res* 2014; 1:76-81
5. Munro MG, Critchley HO, Broder MS, Fraser IS. FIGO Working Group on Menstrual Disorders. FIGO classification system (PALM-COEIN) for causes of abnormal uterine bleeding in nongravid women of reproductive age. *Int J Gynaecol Obstet.* 2011 Apr 1; 113 (1):3-13.
6. Doraiswami S, Johnson T, Rao S, Rajkumar A, Vijaya Raghavan J, Panicker VK. Study of endometrial pathology in abnormal uterine bleeding. *J Obstet Gynaecol India.* 2011 Aug; 61 (4): 426-30. doi: 10.1007/s13224-011-0047-2. Epub 2011 Sep 22. PMID: 228 518 26; PMCID: PMC3295868.
7. Bindroo S, Garg M, Kaur T. Histopathological spectrum of endometrium in abnormal uterine bleeding. *Int J Reprod Contracept Obstet Gynecol* 2018; 7:3633-7
8. Sharma K, Rasania A. Clinicopathological spectrum of endometrial biopsies in a tertiary care center. *Int J Sci Res.* 2019; 8:4-7.
9. Samal R, Vaithy A, S, Habeebullah S. Clinico patho logical analysis of abnormal uterine bleeding in reproductive and post-menopausal women in a tertiary care Centre of south eastern part of India. *Indian J Obstet Gynecol Res* 2020;7(1):66-70
10. Dhangal G. A Study of Endometrium of Patients with Abnormal Uterine Bleeding at Chitwan Valley. *Katmandu Univ Med J* 2003;1(2):110-112.
11. Kathleen G, Patil MS, AS Anand, Histopathological correlation of endometrial samples in pre and post Meno pausal women with abnormal uterine bleeding, *J Diagn Pathol Oncol* 2019;4(1):32-38

12. Iqbal M. B, Kambale T, Khandelwal A, Koshy A, Banerjee B. Spectrum of Endometrial lesions in patients presenting with abnormal uterine bleeding. *Indian J Pathol Oncol.* 2018;5(4):587-591.
13. Prathipaa R, Divya J. Histopathological study of endometrial samples in abnormal uterine bleeding. *Indian J Pathol Oncol* 2020;7(4):567-570
14. Veena M, Chakraborty A, Tyagi S, Sharma R, Alam K. and Mohsin S. Endometrial changes in abnormal uterine bleeding. *J Obstet Gynaecol India* 1996; 33(4): 389-394.
15. Baral R and Pudasaini S. Histopathological pattern of endometrial samples in abnormal uterine bleeding. *J Pathol Nepal* 2011; 1:13-16.