Role of Hysteroscopic EvaluationinAbnormal Uterine Bleeding and its histopathological co-relation.

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Abstract: The present study was conducted to assess the feasibility and accuracy of hysteroscopy in evaluation of intrauterine abnormalities in women with abnormal uterine bleeding and to correlate it with histopathological evaluation.50 women in the reproductive, perimenopausal and postmenopausal age group, up to the age of 65 years presenting with AUB were included as per the inclusion and exclusion criteria. All patients underwent a hysteroscopic examination followed by a biopsy and histopathological evaluation. Hysteroscopic findings were noted and compared with histopathological findings.sensitivity, specificity, PPV, NPV and accuracy of hysteroscopy was calculated.

Results: Maximum age incidence was between 41-50 years (46%).Of the 100 patients, maximum number of women (66%) had symptoms for less than 6 months, 13 patients (26%) had symptoms for 6 months to 1 year and 4 patients (8%) had symptoms for more than 1 year.Out of total 50 patients, 42% presented with Menorrhagia. The second commonest presentation was metrorrhagia (32%). There were 7 cases (14%) with Polymenorrhea and 14 (28%) patients with post-menopausal bleeding and 6% had intermenstrual spotting.Of the 50 patients, 46 cases (924%) were Multiparous, and 4 cases (8%) were Nulliparous. Abnormal findings were seen in 35 patients (70%), while in the remaining 15 patients (29%), no abnormality was detected (negative hysteroscopic view). The most common abnormality was endometrial Hyperplasia (20%cases), followed by Endometrial Polyps (18%) and submucosal fibroids (16%). There were also 7 cases (14%) of endometrial atrophy and 1 case of endometrial carcinoma. Of the 50 cases, 23 (46%) were normal on histopathology. The most common abnormality observed was polyp (20% cases), followed by submucosal fibroids (16%), endometrial Hyperplasia (14%) and endometrial carcinoma (4%). The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose polyp was 90%, 100%, 100% and 97.6% respectively. All the case of fibroids were correctly diagnosed by hysteroscopy, thus sensitivity, specificity, PPV and NPV of Hysteroscopy were all 100%.

The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose endometrial hyperplasia was 85.7%, 90.7%, 60% and 97.5% respectively. The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose endometrial carcinoma was 50%, 100%, 100 and 98% respectively.

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I. Introduction

Abnormal uterine bleeding (AUB) is a term used to describe any change in the duration of a menstrual cycle, the amount or duration of blood loss. Up to 33% of women referred to gynecological outpatient clinics present with AUB and the proportion rises in perimenopausal and postmenopausal women (1, 2). Although AUB may be due to various causes, FIGO has classified the aetiologies into nine main categories, namely (PALM-COEIN): polyp, adenomyosis, leiomyoma, malignancy and hyperplasia, coagulopathy, ovulatorydysfunction, endometrial, iatrogenic, and not yet classified. The components of PALM group are discrete(structural)entities that can be measured visually with imaging techniques and/ or histopathology, whereas the COEIN group is related to entities that are not defined by imaging or histopathology (nonstructural) (3). AUBaffects a woman adversely with respect to her social life, quality of living and thereby her efficiency. Diagnosing the aetiology of AUB can be difficult even with a thorough history and clinical examination. Various diagnostic techniqueslike dialatation and curettage (D&C), , transvaginal ultasonography and saline infusion sonography and hysteroscopy have evolved over a period of time to determine the etiology of abnormal uterine bleeding. The judicious use of hysteroscopy to manage this medical entity adds a new dimension in handling this often-perplexing problem. This technique has replaced dilatation and curettage (D&C), which is a blind technique with a high diagnostic failure rate (4,5) Hysteroscopy as an outpatient procedure is an important diagnostic tool in AUB.Although, the major role of out-patient hysteroscopy in the management of AUB is diagnostic, there is scope for simple operative procedures such as polypectomy and targeted endometrial biopsy. Since , Hysteroscopy and its directed biopsy is more accurate than dilatation and curettage, it is considered an accurate 'gold standard' in uterine pathology(6,7)The present study was thus taken upto assess

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the feasibility and accuracy of hysteroscopy in evaluation of intrauterine abnormalities in women with abnormal uterine bleeding and to correlate it with histopathological evaluation.

II. Material and methods

Study Design

Hospital Based Screening Test Study

Study Duration

July 2015 - June 2017

Study Area

Department of Obstetrics and Gynecology, MGM Hospital and medical college, Navi Mumbai.

Sampling Technique & Sample Size:

Consecutive type of non-probability sampling was used for selection of study subjects. A total of 50 consecutive cases of AUB were taken up for the study after informed consent.

Inclusion criteria

- 1. Age >/= 18 years
- 2. Patients not on any extrinsic hormonal therapy
- 3. Normal PAP smear findings
- 4. No obvious extra-uterine pathology on USG

Exclusion criteria

- 1. Abnormal PAP smear findings
- 2. Acute Pelvic Infection (positive swab test)
- 3. Pregnancy
- 4. Anaemia
- 5. Coagulation disorders

Study Methodology

All 50 patients in reproductive, perimenopausal and postmenopausal age groupupto the age of 65 years presenting with AUB were included after informed consent. Demographic details and clinical history was taken for all patients. General, systemic and local pelvic examination to record the size and position of the uterus, its mobility and the presence of any cervical or adnexal masses was also carried out. After the necessary preoperative investigations all patients were admitted and standard preoperative preparations were carried out.

Hysteroscopy followed by guided biopsywas performed for all patients under total intravenous anesthesia (TIVA). Patient was put in lithotomy position and the findings of examination under anesthesia (EUA) were recorded. We used a rigid continuous flow panoramic hysteroscope, 25 cm in length, 2.9 mm in diameter with an outer sheath of 4 mm and a 30-degree fibro optic lens (Karl Storz, Germany). A fibro optic cable was connected to the light source and to the hysteroscope. Hysteroscopy was performed with normal saline as distention media.

Hysteroscope was inserted inside the uterus gradually after negotiating the external os and cervical canal. once the cavity was entered, a panoramic view of the uterine cavity was taken to exclude uterine malformations or a deformed cavity. Examination was then done systematically, first the fundus, anterior, posterior and lateral walls of the uterus ending by visualization of the uterotubal junctions.

In case of any intrauterine pathology detected, the shape, size and site of it was noted. The thickness, color, vasculature and consistency of the mucous membrane covering the uterine cavity was observed and recorded.

DiagnosticCriteria for

Endometrial hyperplasia: (8)

- i) Increased endometrial thickness
- ii) Non-homogenous endometrial regeneration
- iii) Increased vascularisation
- iv) Polypoid formation
- v) Cystic dilatation
- vi) Necrotic areas

Endometrial carcinoma(8)

- i) Irregular, polylobular, delicate excrescences which are partly necrotic or bleeding,
- ii) Vascularization is also irregular and anarchic.

Biopsy or gross specimen of pathological lesion if any was obtained and sent for histopathological evaluation.

Statistical Analysis

All the collected data was entered in Microsoft Excel Sheet 2007. The data was then transferred and analyzed using SPSS ver. 21. Qualitative data was represented in the form of frequency and percentage while quantitative data was represented using Mean +/- S.D. Appropriate statistical evaluation was carried out as per the type and distribution of data. A p-value of < 0.05 was taken as level of significance

III. Results

Table 1. Distribution of subjects based on Age group

| Age Group (years) | N | % |
|-------------------|----|--------|
| 20-30 | 3 | 6.0% |
| 31-40 | 11 | 22.0% |
| 41-50 | 23 | 46.0% |
| > 50 | 13 | 26.0% |
| Total | 50 | 100.0% |

In the present study, maximum age incidence was between 41-50 years (46%). The youngest patient in this study was 22 years old and the oldest was 68 years old.

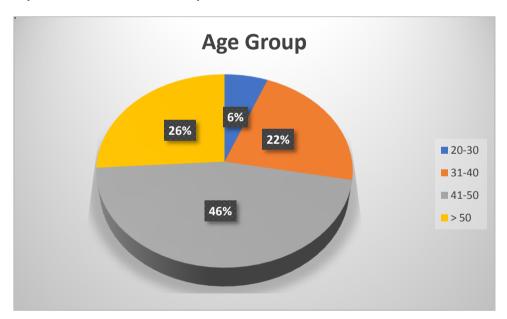


Table 2. Distribution of subjects based on duration of symptoms

| Duration of Symptoms | N | % |
|-----------------------------|----|--------|
| < 6 months | 33 | 66.0% |
| 6 -12 months | 13 | 26.0% |
| > 12 months | 4 | 8.0% |
| Total | 50 | 100.0% |

Of the 100 patients, maximum number of women (66%) had symptoms for less than 6 months, 13 patients (26%) had symptoms for 6 months to 1 year and 4 patients (8%) had symptoms for more than 1 year.

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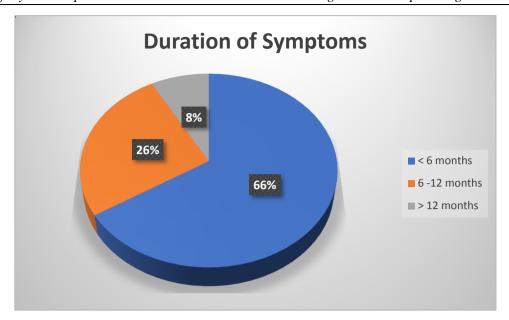


Table 3. Distribution of subjects based on presenting symptoms

| Symptoms | N | % |
|---------------------------|----|-------|
| Intermenstrual spotting | 3 | 6.0% |
| Menorrhagia | 21 | 42.0% |
| Metrorrhagia | 16 | 32.0% |
| Polymenorrhoea | 7 | 14.0% |
| Post- menopausal bleeding | 14 | 28.0% |

Out of total 50 patients, 42% presented with Menorrhagia. The second commonest presentation was metrorrhagia (32%). There were 7 cases (14%) with Polymenorrheaand 14 (28%) patients with post-menopausal bleeding and 6% had intermenstrual spotting.

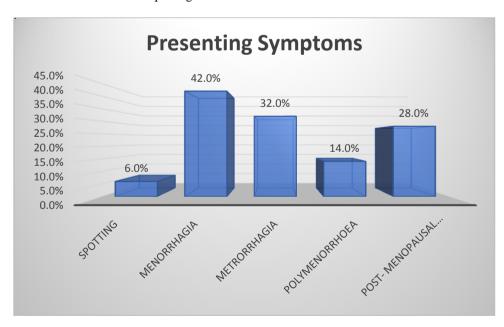
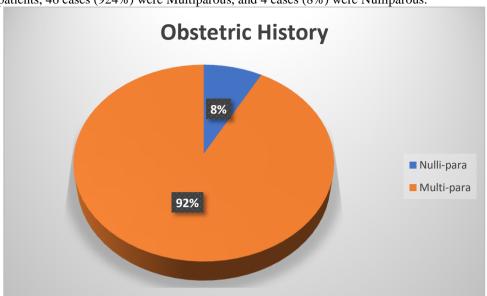


Table 4. Distribution of subjects based on Obstetric History

| Tuble 4. Distribution of Subjects bused on Obstetile History | | |
|--|----|--------|
| Obstetric History | N | % |
| Nulli-para | 4 | 8.0% |
| Multi-para | 46 | 92.0% |
| Total | 50 | 100.0% |



Of the 50 patients, 46 cases (924%) were Multiparous, and 4 cases (8%) were Nulliparous.

Table 5. Distribution of subjects based on Hysteroscopic findings

| Hysteroscopic Findings | N | % |
|------------------------|----|--------|
| Normal | 15 | 30.0% |
| Polyp | 9 | 18.0% |
| Submucosal Fibroid | 8 | 16.0% |
| Hyperplasia | 10 | 20.0% |
| Endometrial Carcinoma | 1 | 2.0% |
| Atrophy | 7 | 14.0% |
| Total | 50 | 100.0% |

Abnormal findings were seen in 35 patients (70%), while in the remaining 15 patients (29%), no abnormality was detected (negative hysteroscopic view). The most common abnormality was endometrial Hyperplasia (20% cases), followed by Endometrial Polyps (18%) and submucosal fibroids (16%). There were also 7 cases (14%) of endometrial atrophy and 1 case of endometrial carcinoma.

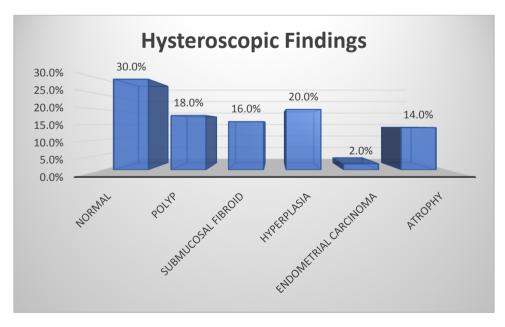


Table 6. Distribution of subjects based on Histopathological Diagnosis

| Histopathological Diagnosis | N | % | |
|-----------------------------|----|--------|--|
| Normal | 23 | 46.0% | |
| Polyp | 10 | 20.0% | |
| Submucosal Fibroid | 8 | 16.0% | |
| Hyperplasia | 7 | 14.0% | |
| Endometrial Carcinoma | 2 | 4.0% | |
| Total | 50 | 100.0% | |

Of the 50 cases, 23 (46%) were normal on histopathology. The most common abnormality observed was polyp (20% cases), followed by submucosal fibroids (16%), endometrial Hyperplasia (14%) and endometrial carcinoma (4%).

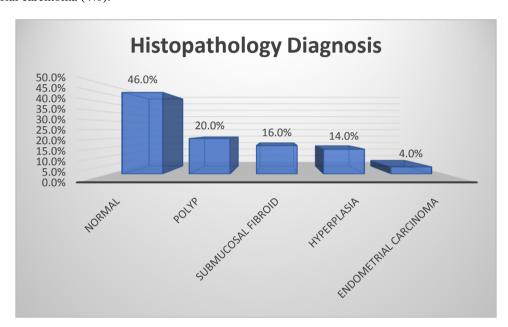


Table 7. Diagnostic efficacy of Hysteroscopy for polyps

| Hystonogoony Dolyn | Histopath - Polyp | | Total |
|----------------------|-------------------|----|-------|
| Hysteroscopy – Polyp | Yes | No | 10tai |
| Yes | 9 | 0 | 9 |
| No | 1 | 40 | 41 |
| Total | 10 | 40 | 50 |

| Parameters | % |
|---------------------------|--------|
| Sensitivity | 90.0% |
| Specificity | 100.0% |
| Positive Predictive Value | 100.0% |
| Negative Predictive Value | 97.6% |

The sensitivity, specificity, Positive PredictiveValueandNegative PredictiveValue of Hysteroscopy to diagnose polyp was 90%, 100%, 100% and 97.6% respectively.

Table 8. Diagnostic efficacy of Hysteroscopy for Submucous Fibroid

| Hysteroscopy - Submucos | Histopath - Submucos Fibroid | | Total |
|-------------------------|------------------------------|----|-------|
| Fibroid | Yes | No | |
| Yes | 8 | 0 | 8 |
| No | 0 | 42 | 42 |
| Total | 8 | 42 | 50 |

| Parameters | % |
|---------------------------|--------|
| Sensitivity | 100.0% |
| Specificity | 100.0% |
| Positive predictive Value | 100.0% |
| Negative Predictive Value | 100.0% |

All the case of fibroids were correctly diagnosed by hysteroscopy, thus sensitivity, specificity, Positive predictive Valueand Negative Predictive Valueof Hysteroscopy were all 100%.

Table 9. Diagnostic efficacy of Hysteroscopy for Endometrial Hyperplasia

| Hysteroscopy - Endometrial | Histopath - Endometrial Hyperplasia | | Total |
|----------------------------|-------------------------------------|----|-------|
| Hyperplasia | Yes | No | |
| Yes | 6 | 4 | 10 |
| No | 1 | 39 | 40 |
| Total | 7 | 43 | 50 |

| Parameters | % |
|---------------------------|-------|
| Sensitivity | 85.7% |
| Specificity | 90.7% |
| Positive Predictive Value | 60.0% |
| Negative Predictive Value | 97.5% |

The sensitivity, specificity, Positive Predictive Value and Negative Predictive Value of Hysteroscopy to diagnose endometrial hyperplasia was 85.7%, 90.7%, 60% and 97.5% respectively.

Table 10. Diagnostic efficacy of Hysteroscopy for Endometrial carcinoma

| Hysteroscopy - Endometrial | Histopath - Endomet | Total | | |
|----------------------------|---------------------|-------|-------|--|
| Carcinoma | Yes | No | 10441 | |
| Yes | 1 | 0 | 1 | |
| No | 1 | 48 | 49 | |
| Total | 2 | 48 | 50 | |

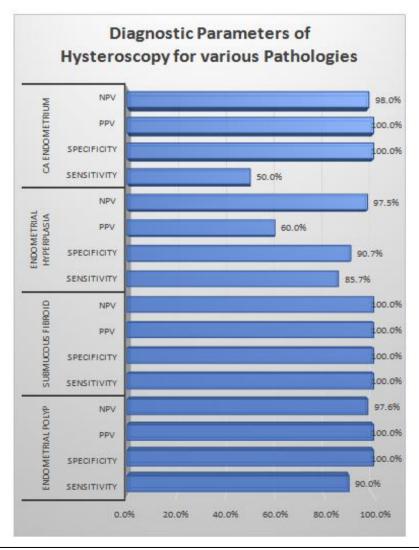
| Parameters | % | | |
|---------------------------|--------|--|--|
| Sensitivity | 50.0% | | |
| Specificity | 100.0% | | |
| Positive Predictive Value | 100.0% | | |
| Negative Predictive Value | 98.0% | | |

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The sensitivity, specificity, Positive Predictive Value and Negative Predictive Value of Hysteroscopy to diagnose endometrial carcinoma was 50%, 100%, 100 and 98% respectively.

Table 11. Diagnostic efficacy of Hysteroscopy for all pathologies

| Pathological lesions | Accuracy | Hysteroscopy |
|-------------------------|------------------------------|--------------|
| Endometrial Polyp | Sensitivity | 90.0% |
| | Specificity | 100.0% |
| | Positive Predictive Value | 100.0% |
| | Negative Predictive Value | 97.6% |
| Submucous Fibroid | Sensitivity | 100.0% |
| | Specificity | 100.0% |
| | Positive Predictive Value | 100.0% |
| | Negative Predictive Value | 100.0% |
| Endometrial Hyperplasia | Sensitivity | 85.7% |
| | Specificity | 90.7% |
| | Positive Predictive Value | 60.0% |
| | Negative Predictive Value | 97.5% |
| CA Endometrium | Sensitivity | 50.0% |
| | Specificity | 100.0% |
| | Positive Predictive Value | 100.0% |
| | Negative Predictive Value | 98.0% |



IV. Discussion

Hysteroscopy is an important method for the diagnosis of intrauterine pathology in several gynecologic complaints including abnormal uterine bleeding (AUB). Sensitivity and specificity of hysteroscopy is high in a large series of hysteroscopies performed in cases of Abnormal uterine bleeding (9-12). The various causes of AUB namely endometrial polyp, submucous myoma etc. are often missed on conventional diagnostic procedures like USG and dilatation and curettage (13). The present study was thus conducted to assess the feasibility and accuracy of hysteroscopy in evaluation of intrauterine abnormalities in women with abnormal uterine bleeding.

Age Distribution

In the present study, maximum age incidence was between 41-50 years (46%). The youngest patient in this study was 22 years old and the oldest was 68 years old.

Panda et al. (14) found that maximum age incidence was between 35-45 years in range between 25-70 years. In Gianninoto's (15) series, age range was 30-80 years and commonest incidence was between 30-45 years. Trotsenburg (16) reported maximum age incidence between 41-50 years. Similar observations were also made by other authors (9,10,11,12,17,18).

Clinical Presentation

In present study, out of total 50 patients, 42% presented with Menorrhagia. The second commonest presentation was metrorrhagia (32%). There were 7 cases (14%) with Polymenorrhea and 14 (28%) patients with post-menopausal bleeding and 6% had intermenstrual spotting.

Panda' et al. [14] series had 60% cases of menorrhagia followed by Polymenorrhagia and Metrorrhagia. Similar findings were also observed by Gita G et al.(17). and Sunitha et al(18).Goyal et al. in their study also observed menorrhagia as the commonest presenting symptom in the study population (58%) followed by metrorrhagia, menometrorrhagia and continuous bleeding >21 days(11).In a study, Chhikara A et al. assessed the accuracy of the hysteroscopic diagnosis with that of hystopathological reports obtained by endometrial sampling. The most common symptom was menorrhagia (40%) followed by metrorrhagia 38%, polymenorrhagia(12%) and postmenopausal bleeding(10%).(12)

Hysteroscopy Findings

Abnormal findings were seen in 35 patients (70%), while in the remaining 15 patients (29%), no abnormality was detected (negative hysteroscopic view). The most common abnormality was endometrial Hyperplasia (20% cases), followed by Endometrial Polyps (18%) and submucosal fibroids (16%). There were also 7 cases (14%) of endometrial atrophy and 1 case of endometrial carcinoma. Overall percentage of abnormal findings observed on Hysteroscopy in abnormal uterine bleeding cases by various authors is given in table below.

| Author | Abnormal Hysteroscopy (%) | | |
|------------------------|---------------------------|--|--|
| Wamsteker et al. (19) | 58.5 | | |
| Gimpelso et al. (7) | 40 | | |
| Loffer et al. (20) | 51.44 | | |
| Sheth et al. (21) | 56 | | |
| Neumann et al. (22) | 44.8 | | |
| Panda et al. (14) | 53.4 | | |
| Gianninoto et al. (15) | 75 | | |
| de Wit AC et al. (23) | 75.8 | | |
| Present Series | 70 | | |

Etiology of Abnormal Uterine Bleeding

The most common abnormality observed on Histopathology was polyp (20% cases), followed by submucosal fibroids (16%), endometrial Hyperplasia (14%) and endometrial carcinoma (4%).

Panda (14) found endometrial hyperplasia in 28.3%, Wamsteker et al. (19) found endometrial polyp in 19%, endometrial hyperplasia in 12.2% and submucous myoma in 7.8%, Trotsenburg et al. (24) observed myomas and polyps in 14% and deLewit (23) reported myomas in 21% and polyps in 14.4%. Incidence of endometrial polyps has ranged from 9.1% in Hamous' (25) to 45.9% in Pasqualotto (26) series.

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Diagnostic Accuracy of Hysteroscopy

The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose polyp was 90%, 100%, 100% and 97.6% respectively. All the case of fibroids were correctly diagnosed by hysteroscopy, thus sensitivity, specificity, PPV and NPV of Hysteroscopy were all 100%. The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose endometrial hyperplasia was 85.7%, 90.7%, 60% and 97.5% respectively. The sensitivity, specificity, PPV and NPV of Hysteroscopy to diagnose endometrial carcinoma was 50%, 100%, 100% and 98% respectively.

The diagnostic accuracy of Hysteroscopy for endometrial polyp, submucous fibroid, endometrial hyperplasia and CA endometrium in cases of AUB is as follows:

| Study | Parameter | Polyp | Fibroid | Hyperplasia | Carcinoma |
|---------------------------|-------------|--------|---------|-------------|-----------|
| Mukhopadhay S et al. (27) | sensitivity | 71.4% | 100.0% | 50.0% | 50.0% |
| | specificity | 100.0% | 100.0% | 95.8% | 100.0% |
| Islam A et al(28) | sensitivity | 100.0% | 100.0% | 100.0% | 73.4% |
| | specificity | 100.0% | 100.0% | 100.0% | 100.0% |
| Chaudhari et al. (10) | sensitivity | 94.0% | 91.0% | 92.0% | 75.0% |
| | specificity | 96.0% | 95.0% | 92.0% | 98.0% |
| Chhikara et al. (12) | sensitivity | 80.0% | 100.0% | 66.6% | |
| | specificity | 97.5% | 100.0% | 90.2% | |
| Present Study | sensitivity | 90.0% | 100.0% | 85.7% | 50.0% |
| | specificity | 100.0% | 100.0% | 90.7% | 100.0% |

We thus observed that, Hysteroscopy showed a high sensitivity and specificity for most of the cases of AUB. We thus conclude that Hysteroscopy can increase the accuracy of clinical diagnosis and may even serve as an adjunct in the treatment of patients with specific intrauterine pathological conditions.

V. Conclusion

Hysteroscopy is a valuable tool for diagnosing endometrial inflammation, atrophic endometrium and focal lesions like endometrial polyps and submucosal fibroids. Although hysteroscopy is somewhat less reliable diagnostic modality for endometrial hyperplasia than other pathologies its diagnostic accuracy is still very high. We thus conclude that, sincehysteroscopy can increase the accuracy of clinical diagnosis it should be a part of diagnostic workup in cases of AUB . In addition hysteroscopy may even serve as an adjunct in the treatment of patients with specific intrauterine pathological conditions.

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