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Original Research Article

Evaluation of women with postcoital bleeding by clinical examination, papsmear, colposcopy and histopathology of cervix

Srilakshmi Yarlagadda¹*, Rangarao Diddi², Prasuna J. L. Narra¹

¹Department of Obstetrics and Gynecology, ²Department of Pathology, Dr. Pinnamaneni Siddhartha Institute of Medical Sciences and Research Foundation, Krishna, Andhra Pradesh, India

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***Correspondence:** Dr. Srilakshmi Yarlagadda, E-mail: ysiri531@gmail.com

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ABSTRACT

Background: Postcoital bleeding refers to spotting or bleeding per vagina that occurs after intercourse and is not related to menstruation. Although there are multiple benign etiologies to this complaint, the most serious cause of postcoital bleeding is cervical cancer (3-5.5%). Cervical cancer is the most common and preventable genital cancer of women. It has a long premalignant phase and with an ideal screening test with good sensitivity and specificity, we can diagnose and treat premalignant cervical lesions, preventing cervical cancer. The present study is aimed to evaluate the women with postcoital bleeding by clinical examination, pap smear, colposcopy and guided biopsy to detect premalignant cervical lesions and carcinoma cervix.

Methods: This was a prospective and retrospective study conducted from April, 2016 to March, 2018 for a period of 24 months in the Department of Obstetrics and Gynaecology on 100 women with postcoital bleeding at Dr. Pinnamaneni Siddhartha Institute of Medical Sciences and Research Foundation (Dr. PSIMS&RF). After clinical examination, these women were subjected to pap smear, colposcopy and guided biopsy. The findings were correlated with histopathology of cervix. The sensitivity, specificity, positive predictive value, negative predictive value and accuracy of both papsmear and colposcopy were calculated.

Results: Sensitivity of papsmear was 71.42%, specificity-86.20%, PPV-78.95%, NPV-80.64% and accuracy-80%. Sensitivity of colposcopy was 90.47%, specificity-89.65%, PPV-86.36%, NPV-92.85% and accuracy-90%.

Conclusions: These results establish colposcopy as an effective screening test for carcinoma cervix. Histopathology of suspected cervical lesion in colposcopy remains the gold standard for definitive diagnosis.

Keywords: Cervical cancer screening, Colposcopy, Postcoital bleeding, Papsmear, Premalignant lesions of cervix

INTRODUCTION

Post coital bleeding refers to spotting or bleeding per vagina that occurs after intercourse and is not related to menstruation.¹ The prevalence of postcoital bleeding ranges from 0.7-9% of menstruating women. Spontaneous resolution has been documented in 51% of premenopausal women who are naturally menstruating after 2 years with no further signs of recurrence.² About 30% of women with postcoital bleeding also experience abnormal uterine bleeding and 15% have dyspareunia.^{3,4}

There are multiple etiologies among which most common are benign cervicitis, ectropion or cervical polyps. CIN or carcinoma can also cause post coital bleeding.⁵The prevalence of cervical cancer in women with post coital bleeding is 3-5.5% and prevalence of CIN is 6.8-17.8%.⁶⁻ ¹⁰ Cervical cancer is the second most common cancer in the women throughout the world.

Cervical cancer kills approximately 2,30,000 women annually and the vast majority of deaths occur in developing countries. Carcinoma cervix is the 5th most common cancer related cause of death among women in the developing world and it is the leading cause of cancer death in women in these countries.⁵ Most women with post coital bleeding have benign diseases like cervicitis, endometritis, cervical polyps, ectropion, vulvar atrophy, benign vascular neoplasms like AVH, hemangiomas and lymphangiomas and sexual abuse.

Benign growths	Infection	Genital/ Vulvar lesions	Benign conditions	Malignancy	Trauma
Endometrial polyps	Cervicitis	HSV lesions	Vaginal atrophy	Cervical cancer	Sexual abuse
Cervical polyps	PID	Syphilis	Pelvic organ prolapse	Endometrial cancer	Foreign bodies
Cervical ectropion	Endometritis	Chancroid	Benign vascular neoplasms	Vaginal cancer	
	Vaginitis	LGV lesions	Endometriosis		
		Condyloma accuminata			

Table 1: Common causes of postcoital bleeding.

Postcoital bleeding is the presenting complaint in 11% of women with cervical cancer.¹¹ The most important risk factor for carcinoma cervix include women who have been infected with a high-risk strain of Human Papilloma Virus (HPV), the virus believed to cause cervical cancer. Other risk factors include immunosuppression and smoking.

Table 2: Risk of cervical cancer in women with
postcoital bleeding.

Age (years)	Risk
20-24	1:44,000
25-34	1:5,600
35-44	1:2,800
45-54	1:2,400

The incidence of cervical cancer has been significantly decreased in the past decades due to enhanced screening for cervical cancer via cervical cytology either with or without testing for HPV and colposcopy to detect precancerous lesions of cervix.³ The long preclinical stage of cervical cancer during which precursor lesions can be treated conservatively and successfully make it an ideal target for screening.

The most common histopathological types of cervical cancer include Squamous cell carcinoma (69%)and Adenocarcinoma (25%).¹² Of the two types, Adenocarcinoma may be less likely to present with postcoital bleeding as lesions may be higher in cervical canal and protected from the trauma of intercourse.¹³ Women with postcoital bleeding who are found to have cervical cancer often are diagnosed with a higher stage of cancer than asymptomatic women.¹⁴ VAIN also causes postcoital bleeding and unusual vaginal discharge.¹⁵ Vaginal cancer is another gynaecological malignancy which causes postcoital bleeding. Primary vaginal

carcinoma is responsible for 3% malignant neoplasms of female genital tract. Primary vaginal carcinoma often be located on the posterior aspect of upper one-third of vagina which is in close proximity to cervix. It is believed that one of the most important risk factors for development of VAIN is from previous (or) concomitant cervical dysplasia. Post coital bleeding can be the presenting complaint in 90% of women with endometrial carcinoma. Primary malignant lymphoma and Non-Hodgkin's lymphoma of female genital tract can also be present with postcoital bleeding.

The objectives of this study are to screen women with postcoital bleeding for precancerous lesions and invasive carcinoma of cervix with papsmear, colposcopy and guided biopsy and to compare and correlate the findings of papsmear, colposcopy and histopathological examination of cervix.

METHODS

This prospective and retrospective study was done at Dr. PSIMS and RF, Chinoutapally, Andhra Pradesh, India, from April 2016-March 2018 for a period of two years on 100 women with postcoital bleeding who attended the Gynecology OPD. Patients details were noted regarding age, socioeconomic status, age at marriage, occupation of both patient and her husband, parity and usage of contraceptives. Detailed history was taken regarding postcoital menstruation, bleeding, dysmenorrhea, dyspareunia, intermenstrual bleeding, recurrent excessive white discharge per vagina associated with foul smell and itching, H/O DM. General and gynecological examination was done [examination of external genitalia, speculum and bimanual pelvic examination]

Papsmear denotes exfoliative cytology of cervix. It was taken from transformation zone. Endocervical smear was taken with cytobrush. If abnormal vaginal discharge was present, high vaginal swab for culture and sensitivity was sent. Papsmear was not taken during menses (or) vaginal douching, sexual intercourse, antibiotic taken in immediate past. The Bethesda classification system was used for cytological grading of pap smear. Because of low sensitivity [70-80%] and high false negative results with papsmear, it was complemented with colposcopy to reduce the false negative cases. Colposcopy is a simple, noninvasive optical method of visualisation of female lower genital tract under bright illumination and stereoscopic vision. It helps in locating the size and extent of suspicious areas over cervix and vagina by VIA, VILI and examination with green filter to study the vascular pattern. Abnormal colposcopic findings like acetowhite areas and unstained areas after lugol's iodine application were analyzed by Reid's Colposcopy Index.

Table 3: Reid's Colposcopy Index.

Colposcopic signs	Zero point	One point	Two points
Colour	Low intensity acetowhitening Ning (not completely opaque; indistinct acetowhitening Ning, transparent or translucent acetowhitening Ning. Acetowhitening Ning beyond the margin of the transformation zone, pure snow-white colour with intense surface shine (rare)	Intermediate shade-gray/white colour and shiny surface (most lesions should be scored in this category)	Dull, opaque, oyster white; gray
Lesion margin and surface configuration	Microcondylomatous or micropapillary contoured flat lesions with indistinct margins. Feathered or finely scalloped margins. Angular, jagged lesions, 3 satellite lesions beyond the margin of transformation zone	Regular- shaped, symmetrical lesions with smooth, straight outlines	Rolled, peeling edges. Internal demarcation between areas of differing colposcopic appearance. A central area of high grade change and peripheral area of low grade change
Vessels	Fine /uniform calibre vessels closely and uniformly placed. Poorly formed patterns of fine punctuation and/mosaic. Vessels beyond the margin of transformation zone. Fine vessels within microcondylomatous or micropapillary lesions	Absent vessels	Well defined coarse punctuation or coarse mosaic
Iodine staining	Positive iodine uptake giving mahogany brown colour, negative uptake of insignificant lesion, i.e. yellow staining by lesion scoring 3 points or less on the first three criteria. Areas beyond the margins of transformation zone, conspicuous on colposcopy, evident as iodine negative areas (such areas are frequently due to parakeratosis).	Partial iodine uptake – variegated, speckled appearance.	Negative iodine uptake of significant lesion, i.e., yellow staining by a lesion already scoring four points or more on the first three criteria.

RCI-Reids Colposcopy Index, CIN- Cervical Intraepithelial Neoplasia. RCI (overall score): 0-2: Likely to be benign inflammatory, 3-5: Likely to be low grade lesion, 6-8: Likely to be high grade lesion, >8: Likely to be invasive carcinoma.

Inclusion criteria

• Sexually active women with postcoital bleeding.

Exclusion criteria

- Pregnant women.
- Women who underwent hysterectomy.
- Unmarried women.
- Diagnosed cases of carcinoma cervix.

Cervical biopsy was taken from abnormal areas on colposcopy. HPE of cervical biopsies was done by senior Pathologist at our institute.

Sensitivity, specificity, PPV, NPV and Accuracy of colposcopy in diagnosing lesions was calculated. Modified Reid's Index on colposcopy was validated with HPE of cervix. Hyteroscopy guided endometrial biopsy was done in women who also complain postmenopausal bleeding, menstrual abnormalities along with postcoital bleeding.

Statistical analysis

Sensitivity= TP/TP+FN, Specificity= TN/ TN +FP PPV=TP/TP+FP, NPV=TN/FN+TN Accuracy= TP+TN/Total

The results were analyzed statistically by chi-square test and by calculating the p-value.

RESULTS

Age

In present study, majority of women were between 31-40 years (57%). 23% were between 41-45 years and 15% were above 45 years.

Table 4: Age.

Age (years)	Number of cases
26-30	5
31-35	26
36-40	31
41-45	23
>45	15

Pap smear results

NILM was seen in maximum number of cases (62%). On pap smear, LSIL was observed in 22% and HSIL in 12% cases. Carcinoma was detected in 4% women on pap smear.

Correlation of age with pap smear

In present study, most of younger women had NILM & LSIL 93.54% (29/31). On papsmear, HSIL and carcinoma were seen in women above 36 yrs (20.25%).

Table 5: Pap smear results.

Pap smear	No. of cases	%
NILM	62	62
LSIL	22	22
HSIL	12	12
Carcinoma	4	4

NILM-Negative for Intraepithelial Lesion or Malignancy; LSIL-Low grade Squamous Intraepithelial Lesion; HSIL-High grade Squamous Intraepithelial Lesion

Table 6: Correlation of age with pap smear.

Papsmear	26-30	31-35	36-40	41-45	>45
NILM	4	19	20	12	7
LSIL	1	5	7	5	4
HSIL	0	2	3	4	3
Carcinoma	0	0	1	2	1

Distribution of colposcopic lesions

In present study, 56% had benign inflammatory lesions in colposcopy. 24% had low grade lesions and 14% had high grade lesions. 6% had invasive carcinoma in colposcopy.

Table 7: Distribution of colposcopic lesions.

Reid's Colposcopic score	No. of cases	%
0-2 (Benign inflammatory)	56	56
3-5 (low grade lesions)	24	24
6-8 (high grade lesions)	14	14
>8 (invasive carcinoma)	6	6

Correlation between pap smear and colposcopy

Majority of women who had NILM on pap smear had benign inflammatory lesions in colposcopy (87.09%) (54/62). 7 women had low grade lesions.

Table 8: Correlation between pap smear and colposcopy.

Pap smear	Benign inflammatory lesions in colposcopy (0-2)	Low grade lesions in colposcopy (3-5)	High grade lesions in colposcopy (6-8)	Invasive carcinoma lesions in colposcopy (>8)
NILM (62)	54	7	1	0
LSIL (22)	2	16	4	0
HSIL (12)	0	1	9	2
Carcinoma (4)	0	0	0	4

Chi- square value- 105.146, p value <0.00001 which is significant.

Only 1 out of 62 women who had NILM on pap smear had high grade lesion detected in colposcopy.

HSIL was observed on pap smear in 12 women.

Out of these, one woman had low grade lesion in colposcopy (8.33%), 9 women (75%) had high grade lesions and 2(16.6%) had lesions suggestive of carcinoma.

4 women whose pap smear showed carcinoma had invasive carcinoma lesions in colposcopy also.

Histopathological analysis

In present study, HPE of cervix showed chronic cervicitis with metaplasia in 58% cases. Mild dysplasia was observed in 17% cases and moderate-severe dysplasia was seen in 15% cases. 10% had invasive Squamous Cell Carcinoma in present study

Correlation between pap smear and histopathology of cervix

Out of 22 women who had LSIL on pap smear, 7 women had moderate-severe dysplasia of cervix in HPE (31.81%).

5 women whose pap smear showed HSIL had Squamous Cell Carcinoma in HPE (41.66%). Both papsmear and HPE of cervix turned out to be Squamous Cell Carcinoma in 4 cases.

Table 9: HPE analysis.

HPE Report	No. of cases	%
Chronic cervicitis with metaplasia	58	58
Mild dysplasia (LSIL)	17	17
Moderate-severe dysplasia (HSIL)	15	15
Squamous cell carcinoma	10	10

Table 10: Correlation between papsmear and histopathology of cervix.

Papsmear	Chronic cervicitis with metaplasia in HPE	Mild dysplasia in HPE	Moderate to severe dysplasia in HPE	Squamous cell carcinoma in HPE
NILM (62)	50	10	2	0
LSIL (22)	8	6	7	1
HSIL (12)	0	1	6	5
Carcinoma (4)	0	0	0	4

chi-square value- 69.157, p value <0.00001 which is significant.

Sensitivity and specificity of pap smear

Table 11: Sensitivity and specificity of pap smear.

Papsmear	Positive HPE	Negative HPE
Positive (38)	30 (TP)	8 (FP)
Negative (62)	12 (FN)	50 (TN)
TP True positive	FD False positive	EN False negative TN True

TP-True positive, FP-False positive, FN-False negative, TN-True negative

In present study, the sensitivity of papsmear for detecting low grade lesions and above came out to be 71.42%, specificity 86.20%, PPV 78.95%, and NPV 80.64% respectively. Accuracy of papsmear in present study was 80%.

Table 12: Sensitivity and specificity of pap smear.

Papsmear	Percentage
Sensitivity	71.42
Specificity	86.20
Positive predictive value	78.95
Negative predictive value	80.64
Accuracy	80

Table 13: Correlation between colposcopy and histopathology of cervix.

Colposcopy Grade	HPE-chronic cervicitis with metaplasia	HPE-Mild dysplasia (LSIL)	HPE-Moderate- severe dysplasia (HSIL)	HPE-carcinoma
Benign (56)	52	3	1	0
Low grade lesion (24)	6	14	3	1
High grade lesion (14)	0	0	11	3
Invasive ca (6)	0	0	0	6

Chi- square value: 112.83, p value < 0.00001 which is significant.

Correlation between colposcopy and histopathology of cervix

In present study, 56 women had benign inflammatory lesions in colposcopy. Out of them, 92.85% (52/56) had chronic cervicitis with metaplasia in HPE. 3 women had mild dysplasia (5.38%) and one woman (1.78%) had moderate-severe cervical dysplasia.14 women had high grade lesion in colposcopy. Out of them, 78.57% (11/14) had moderate-severe cervical dysplasia and 21.42% (3/14) had invasive Squamous Cell Carcinoma. All the women who had invasive Squamous Cell Carcinoma in histopathology also.

Sensitivity and specificity of colposcopy

In present study, the sensitivity of colposcopy for detecting low grade lesions and above came out to be 90.47%, specificity 89.65%, PPV 86.36%, and NPV 92.85% respectively. Accuracy of colposcopy in present study was 90%.

Table 14: Sensitivity and specificity of colposcopy.

Colposcopy	Histopathology	Histopathology
	Positive	Negative
Positive (44)	38 (TP)	6 (FP)
Negative (56)	4 (FN)	52(TB)

Table 15: Sensitivity and specificity of colposcopy.

Colposcopy	%
Sensitivity	90.47
Specificity	89.65
Positive Predictive Value	86.36
Negative Predictive Value	92.85
Accuracy	90

DISCUSSION

Though postcoital bleeding has multiple benign etiologies, it is the presenting complaint in 11% of women with cervical cancer. Cervical cancer has a long pre-invasive period and appropriate intervention in this period always leads to decrease in the incidence of this dreadful disease.

Proper evaluation of women with postcoital bleeding is necessary to rule out cervical cancer. Conventional papsmear has its own limitations like inadequancy of smear, non-inclusion of endocervical cells, air drying of slide and fixation problems. Colposcopy is an excellent means of evaluating abnormal cervical cytology. Ultimately, HPE of cervix is necessary to establish the definitive diagnosis. In present study, we carefully evaluated women with postcoital bleeding by clinical features, papsmear, colposcopy and guided biopsy. In present study, majority (57%) of women were between 31-40 years. 23% were between 41-45yrs and 15% were above 45 years. With increasing age, advanced lesions on pap test were observed.

In the study conducted by Shalini R et al the mean age of patients with invasive cancer was 41.3 years versus 32.9 years in patients with benign pathology, the difference being significant statistically.⁸ In the study by Himanshi G et al the mean age of patients with invasive cancer was 42.5 years versus 33.5 years in patients with benign pathology.¹⁶ On papsmear, NILM was seen in 62%, LSIL in 22%, HSIL in 12% and carcinoma in 4% women in present study. Himanshi G et al reported that 63% had inflammatory cytology, 15% had normal cytology, 6% had LSIL, 9% had HSIL and 7% had malignancy on papsmear in their study on women with postcoital bleeding.¹⁶ Manish Gupta et al reported in their study that 32% had inflammatory smear, 34.6% had LSIL and 18.7% had HSIL on papsmear in women with postcoital bleeding.17

In present study, 56% had benign inflammatory lesions in colposcopy. 24% had low grade lesions, 14% had high grade lesions and 6% had invasive carcinoma. With increasing age, advanced lesions in colposcopy were observed Manish Gupta et al reported that 34% had Grade-I lesions, 8% had Grade-II and 17% had Grade-III lesions in colposcopy in their study.¹⁷ Himanshi G et al reported 30% of women with normal colposcopy findings, 46% with Grade-I ,7% with Grade-II and 17% with Grade-III lesions in their study.¹⁶

In present study, we compared the results obtained with pap smear and colposcopy. NILM was seen in 62 women. 87.09% who had NILM on papsmear had benign inflammatory lesions in colposcopy, 11.29% women had low grade lesions and 1 out of 62 women (1.61%) who had NILM on pap smear had high grade lesion in colposcopy. LSIL was seen on papsmear in 22 women.

Out of them, 9.09% had benign inflammatory lesions in colposcopy,72.72% women had low grade lesions and 18.18% had high grade lesions in colposcopy. HSIL was observed on pap smear in 12 women. Out of them, 8.33% had low grade lesions, 75% had high grade lesions in colposcopy and 16.6% had carcinoma. 4 women whose pap smear showed carcinoma had invasive Squamous Cell Carcinoma in colposcopy also.

Cervical histopathology showed chronic cervicitis with metaplasia in 58% cases. Mild dysplasia was seen in 17% cases, moderate-severe dysplasia in 15% cases. 10% had invasive Squamous Cell Carcinoma in present study.

Gupta M et al reported that mild dysplasia of cervix was seen in 14.6% and moderate-severe dysplasia was seen in 10.7% of women with postcoital bleeding in their study.¹⁷ Himanshi G et al reported that 29% of women with postcoital bleeding had chronic cervicitis and 8% had invasive cervical carcinoma in HPE in their study.¹⁶

Shalini R et al reported that 5.5% of women with postcoital bleeding had invasive cancer cervix in their study whereas Rosenthal AN et al reported 4% cases with invasive cancer cervix.8,9 Allen et al reported that 1% of women with postcoital bleeding had carcinoma cervix in their study and Ashrafgangoei T et al reported 1.8% cases with invasive cancer cervix in women with postcoital bleeding in their study.^{19,21} Authors compared the results of papsmear with HPE. In present study, 62 women had NILM on papsmear. Out of them 80.64% had chronic cervicitis with metaplasia in HPE, 16.12% had mild dysplasia and 3.22% had moderate- severe dysplasia. 22 women had LSIL on papsmear. Out of them, 36.36% had chronic cervicitis with metaplasia, 27.27% had low grade lesions, 31.82% had moderate- severe dysplasia and 4.54% had carcinoma. 12 women had HSIL on papsmear. Out of them, 8.33% had mild dysplasia, 50% had moderate- severe dysplasia, 41.67% had carcinoma in HPE. All the women who had lesions suggestive of carcinoma on papsmear had invasive Squamous Cell Carcinoma in histopathology also.

Sensitivity of papsmear was 71.42%, specificity-86.20%, PPV-78.95%, NPV-80.64% and accuracy-80% in present study. Shalini R et al reported that papsmear had 56% sensitivity and 90% specificity in their study.⁸ Tehranian A et al reported that pap smear had 50% sensitivity and 86.5% specificity in their study on women with postcoital bleeding.³

We compared the results of colposcopy with HPE. In present study, 56 women had benign inflammatory lesions in colposcopy. Out of them 92.85% had chronic cervicitis with metaplasia, 5.36% had mild dysplasia and only 1 woman (1.78%) had moderate-severe dysplasia. 24 women had low grade lesions in colposcopy. Out of them, 25% had chronic cervicitis with metaplasia, 58.33% had mild dysplasia, 12.5% had moderate-severe dysplasia and 4.17% had carcinoma.

14 women had high grade lesions in colposcopy. Out of them 78.57% had moderate-severe cervical dysplasia and 21.42% had invasive squamous cell carcinoma. All the women who had lesions suggestive of carcinoma in colposcopy had invasive squamous cell carcinoma in histopathology also.

Sensitivity of colposcopy was 90.47%, specificity-89.65%, PPV-86.36%, NPV-92.85% and accuracy-90% in present study. Tehranian A et al reported that colposcopy had 79% sensitivity in their study on evaluation of women with postcoital bleeding.³

Present findings in this study show that colposcopy is more sensitive, specific and accurate than pap smear in detecting premalignant lesions of cervix and cervical carcinoma in women with postcoital bleeding. However, it should be confirmed with histopathology of cervical lesions for definitive diagnosis.

CONCLUSION

It is advisable to perform both papsmear and colposcopy in women with postcoital bleeding to detect premalignant cervical lesions and invasive carcinoma cervix. However, histopathology of suspected lesion in colposcopy remains the gold standard for definitive diagnosis in view of false positives and false negatives with papsmear and colposcopy.

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