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Study of Endometrial Aspiration Cytology and Its Correlation with HPE in Cases of Dub

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Abstract

Background: Endometrial aspiration cytology has long been undermined, but has now emerged as a powerful yet cost effective tool for diagnosing DUB in resource limited settings.

Purpose: The current study compares the accuracy of EAC with HPE as gold standard in diagnosis of DUB in women. It also determines the various morphologic patterns of endometrium in DUB.

Methods: 70 women diagnosed as DUB were subjected to endometrial aspiration with 4mm Karman's cannula and MR syringe prior to D&C and HPE sampling. Data was compiled & analysed with SPSS 18 package. Patients were analysed based on socioeconomic, clinical parameters; cytological diagnosis was compared against HPE diagnosis based on sensitivity, specificity, diagnostic accuracy.

Result: EAC showed a sampling adequacy of 90%. The commonest lesion detected was proliferative endometrium followed by secretory endometrium and hyperplasia without atypia. Sensitivity for detecting proliferative endometrium was 100%, specificity 89.2% and accuracy of 89.2% against gold standard HPE. Sensitivity for detecting secretory endometrium was 100%, specificity 92.8% and accuracy was 88.5%. Sensitivity and specificity of detecting hyperplasia by aspiration cytology was 64% and 100% respectively. Accuracy was 90%. Sensitivity of aspiration cytology in detecting adenocarcinoma was 50% and specificity was 100% in this study with Accuracy of detection being 97% in this study.

Conclusion: *Endometrial aspiration is an effective, useful and a minimally invasive procedure comparing with gold standard HPE.*

Background

Defined as abnormal bleeding from the uterus in the absence of any organic pathology of the genital tract, DUB is one of the most frequently encountered conditions in the gynecological practice and accounts for approximately 10% of all new patients. Endometrial interpretation is valuable not only to find the etiology of DUB but also to rule out any organic cause for abnormal uterine bleeding. Exfoliative cytology has a unique place in the study of female genital tract lesions. Endometrial aspiration study has attempted to overcome the intrinsic weakness of vaginal and cervical smears in diagnosis of endometrial pathology. This study attempts to bust the myth of utter inadequacy of EAC as a first line, cost effective, painless, minimally invasive diagnostic tool for diagnosing DUB compared to

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gold standard HPE studies, which albeit being marginally more accurate is laborious and costly.

Aims of the Present Study

The current study aims to evaluate the efficacy of endometrial aspiration cytology vs D&C in the diagnosis of dysfunctional uterine bleeding by correlating with histopathology. Additionally, we try to determine the various morphologic patterns of endometrium in DUB.

Materials and Method

70 cases of clinically diagnosed DUB at Dept of OBGYN at JNIMS, a tertiary care centre, with their informed consent, were recruited in our cross sectional study from September 2017 to March 2019. Endometrial aspiration with 4mm diameter Karman's cannula and MR syringe prior to D&C/Hysterectomy for HPE study was done. Various variables like age, address, religion, symptoms, parity, haemoglobin, USG finding, cytology finding, HPE finding were entered in Microsoft excel 2019. Data were checked for consistency and accurateness. Data was analysed using SPSS 18 and were tabulated in mean and percentages. Sensitivity, specificity and accuracy was calculated for each method.

Results and Discussion

Our study shows majority of the patients were in the age group 40-50 years (50%) followed by 30-40 years (42.9%) and >50 years in 7.1% of cases (mean age of 42.5 \pm 6.3 years) with anaemia in 70% of cases. In the study by Patil P et al¹, 80% of women belonged to age group of 40 - 49 years which is in concordance with this study. It was comparable to the mean age (44 years) of 51 patients selected for the study by Liza et al².

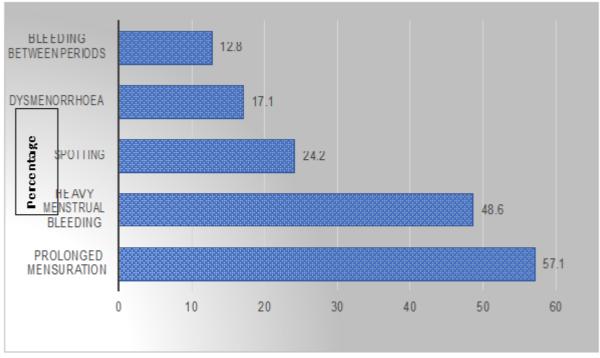


Figure 1: Distribution of the respondents by symptoms

Prolonged menstruation was the commonest symptom in 57% of cases; followed by heavy menstrual bleeding (48.6%), spotting (24.2%), dysmenorrhoea (17.1%) and intermenstrual bleeding (12.8%). Same was observed in the study by Devi LS et al³.

The following table elucidates the commonest endometrial patterns in some other contemporary studies.

Studies	Commonest Endometrial patterns Percent			
Rao et al^4	Proliferative endometrium 4			
Tripathy et al^5	Secretory endometrium	45%		
Hemaalatha et al ⁶	Secretory endometrium	54%		
Morse et al^7	Proliferative endometrium	50%		
Perween et al ⁸	Proliferative endometrium	46.15%		
Present study	Proliferative endometrium	34%		

Table: Table showing commonest endometrial findings in some studies:



Figure 2: Distribution of the respondents by BMI

80% of cases were of normal BMI, followed by overweight (BMI: 25-29.9) in 14.2% of cases and obese in 5.8% (BMI>30) as shown above. Pre aspiration Speculum examination & USG screen were done in all patients. In USG, bulky uterus was found in 55.7% of cases followed by thickened endometrium in 27.1% of cases, and 17.2% had no aberrant find. On cervical speculum examination half of them were healthy, erosion in 28.6%, cervicitis in 28.6% and enlarged cervix in 7.1%.

Table 1: Distribution of the respondents by cytology and histopathologic finding:

Finding	Cytology	HPE	Consistent	Inconsistent
	n(%)	n(%)		
Proliferative endometrium	29(41.4)	24(34.2)	24	5
Secretory endometrium	15(21.4)	14(20.0)	14	1
Mixed (irregular)	10(14.2)	12(17.1)	10	2
Endometrial hyperplasia without atypia	9(12.8)	14(20.0)	9	5
Adenocarcinoma	2(2.1)	4(4.2)	2	2
Inadequate	5(7.3)	2 (2.1)	2	3
Total	70 (100.0)	70 (100.0)	-	-

In cytology, 29 were diagnosed as proliferative vs 24 in HPE study, 15 were secretary vs 14 on HPE, 10 were mixed as compared to 12 by HPE, 9 were hyperplasia vs 14 by HPE and adenocarcinoma in 6 cases vs 4 cases by HPE. 7 cases (8%) had inadequate sample.

Below we break down each endometrial picture, and compare accuracy of EAC with gold standard HPE study.

Cytology	HPE			Fisher exact test
	(gold standard)			
	Proliferative	Non proliferative	Total	
	n(%)	n(%)	N(%)	
Proliferative	24 (100.0)	5 (10.8)	29(41.4)	Value-56.691 df-1
Non proliferative	0 (0.0)	41(89.2)	41(58.6)	p-0.000
Total	24 (100.0)	46(100.0)	70(100.0)	

Table 2: Distribution of the respondents by cytology and histopathologic finding

In cytology, 29 were proliferative but the gold standard HPE diagnosed 24, so 5 cases (3.4%) were false positive. Calculated sensitivity for detecting proliferative endometrium was 100%, specificity was 89.2% and accuracy was 89.2% against gold standard HPE. In the study by Kaur et at⁹ similar finding was noted as HPE diagnosed 20 out of 23 cases detected by

cytology with a sensitivity of 100%, specificity of 96% and accuracy of 96.84%. Similar finding was noted in the study by Patel P et al^{58} and Handa U et al^{10} . Thirty-four of 38 cases diagnosed as proliferative cytologically matched with their corresponding histopathology in the study by Baxi SN et al^{11} .

Table 3: Distribution of the respondent	ts by cytology and histopathologic finding
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Cytology	HPE			Fisher exact test
	(gold standard)			
	Secretory	Non secretory	Total	
	n(%)	n(%)	N(%)	
Secretory	14 (100.0)	1 (1.8)	15(21.4)	Value-75.2
Non secretory	0 (0.0)	55(98.2)	41(78.6)	df-1
Total	14 (100.0)	56(100.0)	70(100.0)	p-0.000

In case of secretory endometrium, 15 were diagnosed by cytology but HPE diagnosed only 14, with a false positivity of 6.6%. Sensitivity for detecting secretory endometrium was 100%, specificity was 92.8% and accuracy was 88.5%. In

the study by Kaur et at⁹, it was noted that cytology could diagnosed 16 out of 17 cases detected by HPE with a sensitivity of 94.4%, specificity of 100% and accuracy of 98.9%.

Table 4: Distribution of the	respondents b	by cytology and	histopathologic	finding for malignancy

Cytology	HPE	HPE	
	(gold standard)		
	Adenocarcinoma	No cancer	
Adenocarcinoma	2 (50.0)	0 (0.0)	Value-35.8 df-1
No cancer	2 (50.0)	66 (100.0)	p-0.002
Total	4 (100.0)	66 (100.0)	

Unfortunately, Cytology could diagnose 2 out of 4 cases of adenocarcinoma (50%) and so 2 cases were missed. Sensitivity of aspiration cytology of detecting adenocarcinoma was 50% and specificity was 100% in this study. In the study by Byrne AJ^{12} for cytological diagnosis of endometrial cancer using endocyte endometrial sampler, they found that sensitivity was 90% which was a bit higher from this study and

specificity was 100% which is same with this study. Liza et al² study had a sensitivity of 81.63% and specificity of 83.3%. Cytologic sampling demonstrated a sensitivity of 78%, specificity of 96%, and positive predictive value of 78% and a negative predictive value of 96% for detection of endometrial abnormalities.¹³

Accuracy of detection adenocarcinoma by aspiration cytology was 97% in this study.

Similar finding was noted in the study by Robert R^{14} where accuracy of diagnosis by endometrial smear was 92.6%, by curettage was 98% and when both were used together accuracy was 100%. This was also noted in the study by Anderson et al¹⁵ where there was excellent

correlation of 96% between endometrial biopsies and curetting. Cytologic smears diagnostic accuracy of 93% was observed in the study by Sagar et al.¹⁶Accuracy of 100% was observed in Chakravarthy A¹⁷ in detecting adenocarcinoma.

Cytology	1	HPE	Fisher exact test
2 02	(gold	standard)	
	Hyperplasia	No hyperplasia	
	n(%)	n(%)	
Hyperplasia	9 (64.0)	0 (0.0)	Value-47.2 df-1
No hyperplasia	5 (36.0)	54 (100.0)	p-0.000
Total	14(100.0)	54 (100.0)	

Table 5: Distribution of the respondents by cytology and histopathologic finding for hyperplasia

Cytology could detect only 64% of hyperplasia and 36% were missed, also statistically significant. Regarding accuracy of hyperplasia it was 90% and this finding was similar to the finding by Chakravarthy A¹⁷ study which they got 87.5% of accuracy.

Average correlation between aspiration cytology and histopathology in this study was 93%. A brief comparative table ascribing correlation between EAC and HPE in some studies are delineated below:

Table 6: Comparative table of EAC vs HPEcorrelation in some studies:

Studies done	%correlation
Tripathy et al ⁵	97%
Hemalatha et al ⁶	94%
Rao et al ⁴	78%
Polson et al ¹⁸	78%
Present study	93%

Aspiration yield 7 cases (8%) with inadequate sample. This finding was almost similar to the finding by Sagar S et al^{16} where cytological smears were inadequate in 10.4% cases. Inadequacy of sample of 14% and 12.5% was observed in the study by Polson et al^{18} and Chakravarthy A^{17} respectively. So, sample adequacy was found in around 90%.

Conclusion

One of the major difficulties encountered in the

cytological study of the endometrium has been related to the inability to obtain a satisfactory and representative cellular sample consistently. Over different the years, authors opined that endometrial aspiration techniques was acceptable valuable method of assessing and the endometrium as an minimally invasive, almost painless and less time intensive OPD procedure. Our study corroborates this fact and proves beyond doubt that EAC faithfully mirrors the Endometrial findings in DUB as shown by gold standard HPE diagnosis.

Limitations of the Present Study

An added aspect which could have been studied is the diagnostic accuracy of EAC with different techniques of endometrial sampling; it could have provided a fruitful solution to ongoing efforts in bettering the EAC outcomes. Additionally, the statistical significance and applicability of any study can be magnified if a bigger pool or sample size, as is the case with our study too.

Conflict of Interest

The author(s) declare no conflict of interest, nor any affiliations or association of any entities or organization with financial motive and investment in this present study.

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