



## Comparison of Efficacy in Diagnosis of Breast Lesions by FNAC, Frozen Section and Histopathological Study in Tertiary Care Hospital

Authors

**Dr Deshpande S.A.<sup>1</sup>, Dr Makode A.J.<sup>2</sup>**

Dr Shankarrao Chavan Medical College and Hospital, Vishnupuri, Nanded, India

### Abstract

**Background:** In women, malignancy of breast imposes significant reduction in life span. So, FNAC is a widely accepted cytological technique in the early diagnosis of palpable breast lesions. For making further appropriate therapeutic decision intra-operative frozen section is used. Ultimately, histopathological study remains the confirmatory test in final diagnosis of breast lesions. There have been many studies of comparison of efficacy of FNAC, Frozen section and histo-pathological study which has been shown to be high in many centres.

**Aim:** To compare cytological and histopathological diagnosis of breast lesions and Frozen section and histopathological diagnosis of breast lesions and to establish accuracy of each of them as an diagnostic aid.

**Material and Methods:** FNAC of 51 cases of palpable breast lesions were evaluated retrospectively between January 2018 to June 2019. Only those cases which underwent frozen section and subsequent histopathological follow-up for confirmative diagnosis were included. FNAC, Frozen section diagnosis and Histopathological diagnoses then correlated to assess the efficacy in diagnosis of palpable breast lesions. Males and non-palpable breast lesions were excluded.

**Results:** The present study showed sensitivity, specificity, accuracy, positive predictive value, and negative predictive values of FNAC in relation to Histopathological diagnosis of breast lesions as 94.44%, 84.85 %, 88.24%, 77.27%, and 96.55% respectively. Also the present study showed sensitivity of 88.89%, specificity of 87.89%, positive predictive value of 80.00%, negative predictive value of 93.55% and accuracy of 88.24% on Frozen section in relation to Histopathological study.

**Conclusion:** FNAC is thus a rapid, cost-effective highly sensitive and highly specific first minimal invasive method in diagnosing breast lumps. But Frozen is also sensitive and specific and has also a definite role in settings like difficult cytology, evaluation of lumpectomy margins and intra operative nodal status. But the histopathological study is still considered as a gold standard in the final diagnosis.

**Keywords:** FNAC; Frozen section; Histopathological study; palpable breast lesions; Malignancy of Breast.

### Introduction

Breast cancer is the second most prevalent cancer among Indian women, the first being cervical cancer. The prognosis of breast cancer is primarily dependent on the extent of disease and also early

diagnosis.<sup>(1)</sup> FNAC though not entirely specific, is an important source of information in patients with breast lumps. In particular; it confidently allows exclusion of breast cancer and other more common disease and is useful in planning a

surgical approach to the lesion<sup>(2)</sup>. It is the most common diagnostic modality for palpable breast lumps<sup>(3)</sup>. FNAC is simple, safe and quick procedure with high sensitivity and specificity<sup>(4,5)</sup>. The accuracy rate of FNAC in diagnosing breast cancer is up to 96%<sup>(6,7)</sup>.

The frozen section technique for intra-operative pathologic diagnosis has been used for more than 100 years. The introduction of the cryostat in 1960 established intraoperative frozen section examination as a highly reliable procedure for the rapid histological evaluation of tissue specimen during surgery<sup>(8)</sup>.

Though Fine Needle Aspiration Cytology (FNAC) has diminished the demand for frozen section evaluation of breast lumps, yet in certain situations, particularly when FNAC fails, the need for frozen section persists.

But Frozen Section it should not be used to replace paraffin embedded tissue technique. Comparatively, Frozen Section is still inferior to the later due to its various limitations. The sampling of specimen is limited and there are technical difficulties of getting good quality sections and staining of tissue and it will all influence the interpretation of the section by the pathologist.

The aim of this study was to analyze the efficacy of Fine Needle Aspiration Cytology (FNAC), frozen section and histopathological study of breast lumps. The results have been compared with published literature and the utility of these three diagnostic modalities for breast lumps has been reassessed in the present scenario.

### Materials and Methods

This research included total 51 female patients of age group 11-80 years clinically presenting with palpable breast lesions referred to the Department of Pathology, for FNAC, frozen section and histopathological evaluation and a prospective study was conducted from January 2018 and June 2019.

FNAC procedure was performed according to the standard protocol<sup>(9)</sup>. Only diagnosed cases were

included in the study. Cases did not undergo surgery were excluded from the study.

The interpretation of the slides was done by the cytopathologist and treatment is decided by the surgeon from the cytological diagnosis.

All the breast specimens received in normal saline (0.9%) solution immediately from the operation theatre without fixation in formalin were subjected for frozen section. Following this they were fixed in 10% formalin for 24 hrs and processed for H&E staining for histopathological examination.

### Results

In our study, all the cases in present study were female patients and no case of male patient was found.

In the present study, the age group ranged from 11-80 years. The most common age group presenting with breast lesions was 31-40 years.

The left side showed a higher frequency of breast lumps with 26 cases (50.98%) while the right side was affected in 24 (47.05%) cases and 1(1.96%) case of Bilateral side. In the present study, 48 (94.11%) patients presented with palpable breast lump followed by 2(3.92%) cases of palpable axillary lymph nodes and 1 (1.96%) case of nipple discharge with 35(68.62%) cases presented within 6 months of duration of complaints followed by 19-24 months of duration of complaints.

Following FNAC, out of 51cases, 29(56.86%) cases presented with malignant breast lesions and 22(43.13%) cases with benign breast lesions. In the present study, among benign lasions, 18(35.29%) cases are of fibroadenoma followed by 3(5.88%) cases of phylloides tumor and 1(1.96%) case of duct ectasia.29 (56.86%) cases are of malignant type with Infiltrating duct carcinoma as the most common lesion.

On frozen section 31(60.78%) cases are of malignant breast lesion and 20(39.21%) cases are of benign type. In the present study, 16 (31.37%) cases are of fibroadenoma followed by 3(5.88%) cases of phylloides tumor and 1(1.96%) case of duct ectasia on frozen section study and

31(60.78%) cases are of malignant breast lesion. Frozen section study of all the resected surgical margins of all malignant breast tumors are free of tumor tissue.

On Histopathology, 18(35.29%) cases are of benign breast lesions and 33(64.70%) cases are of malignant breast lesions in the present study.

Histopathological study of all resected margins of carcinoma breast are free of tumor tissue. In the present study, 14(27.45%) cases are of fibroadenoma followed by 3(5.88%) cases of phylloides tumor and 1(1.96%) case of duct ectasia on paraffin section study and 33(64.70%) cases are malignant.

**Table 1:** Showing Cytohistological correlation of breast lesion diagnosis (n=51)

Sr No	Cytological Diagnosis	Number of cases	Histopathological Diagnosis			
			Benign			Malignant
<b>1</b>	<b>Benign</b>		Fibro Adenoma	Duct Ectasia	Phylloides tumor	Infiltrating duct carcinoma
A	Fibroadenoma	18	14			
B	Duct ectasia	01		01		
C	Phylloides tumor	03			03	
<b>2</b>	<b>Malignant</b>					
A	Infiltrating duct carcinoma	29				33
	<b>Total</b>	<b>51</b>	<b>18</b>			<b>33</b>

On cytohistopathology correlation, 4 cases which were benign on FNAC found to be malignant on Histopathology study.

**Table 2:** Showing statistical analysis of FNAC diagnosis of breast lesions in relation to paraffin section diagnosis

FNAC		Histopathological Diagnosis		Total
		Benign	Malignant	
	<b>Benign</b>	17 (a)	05 (b)	22
	<b>Malignant</b>	01 (c)	28 (d)	29
<b>Total</b>		<b>18</b>	<b>33</b>	<b>51</b>

a :- True Positive ; b :- False Positive ; c :- False Negative ; d :- True Negative

On statistical analysis of cytohistopathology correlation, 5 cases are False Positive on FNAC and 1 case found to be False Negative on Histopathology study.

**Table 3:** Showing sensitivity, specificity, positive predictive value, negative predictive value of FNAC in relation to histopathological study of breast lesions

Sr No.	Parameter	VALUE (%)
1	Sensitivity	<b>94.44</b>
2	Specificity	<b>84.85</b>
3	Positive predictive value	<b>77.27</b>
4	Negative predictive value	<b>96.55</b>
5	Accuracy	<b>88.24</b>

**Table 4:** Showing statistical analysis of Frozen Section diagnosis in relation to paraffin section diagnosis of breast lesions

Frozen section	Histopathological Diagnosis		Total
	Benign	Malignant	
Benign	16 (a)	04 (b)	20
Malignant	02 (c)	29 (d)	31
Total	<b>18</b>	<b>33</b>	<b>51</b>

a :- True Positive ; b :- False Positive ; c :- False Negative ; d :- True Negative

On statistical analysis of Frozen Section and histopathology correlation, 4 cases are False Positive on Frozen section and 2 cases are found to be False Negative on Histopathology study

**Table 5:** Showing sensitivity, specificity, positive predictive value, negative predictive value of Frozen Section in relation to histopathological study of breast lesions

Sr No	Parameter	VALUES %
1	Sensitivity	<b>88.89</b>
2	Specificity	<b>87.88</b>
3	Positive predictive value	<b>80.00</b>
4	Negative predictive value	<b>93.55</b>
5	Accuracy	<b>88.24</b>

**Table 6:** Showing total number of cases on FNAC, Frozen Section and Histopathological diagnosis of breast lesions (n=51)

Sr No	Diagnosis	Benign	Malignant	Total
1	<b>FNAC</b>	22	29	51
2	<b>Frozen Section</b>	20	31	51
3	<b>HPE</b>	18	33	51

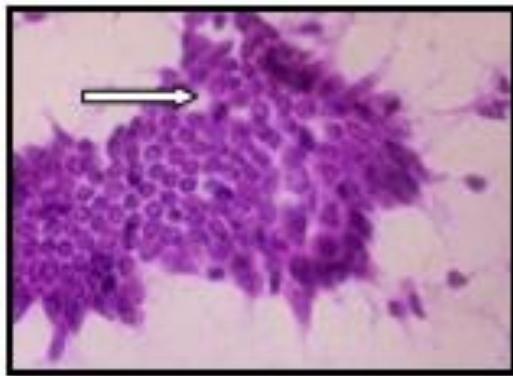


Fig No.1:- Photomicrograph of FNAC slide, Fibroadenoma of Breast (H & E ; 40x) tightly cohesive cluster of monolayered ductal epithelial cells(white arrow) along with Bipolar cell in the [ackground.

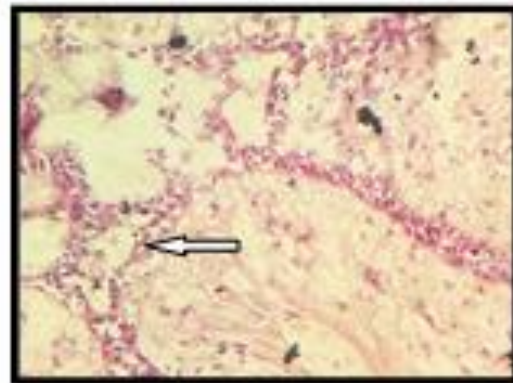


Fig. 2.; Photomicrograph of Frozen section slide, Fibroadenoma ( H & E ; 40 X)show tumor comprised of slit like or dilated ducts (white arrow) having two cell types lining i.e. inner cuboidal epithelial & outer myoepithelial. Stroma is abundant.

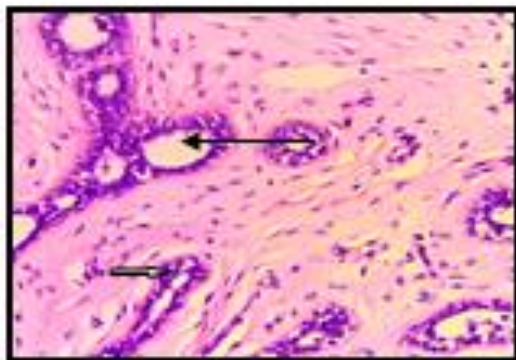


Fig No.3: Photomicrograph of Paraffin section slide, Fibroadenoma ( H& E ; 40 X)show tumor comprised of intracanalicular (white arrow) and pericanalicular pattern (black arrow) of ducts having two cell types lining i.e. inner cuboidal epithelial & outer myoepithelial. Stroma is abundant.

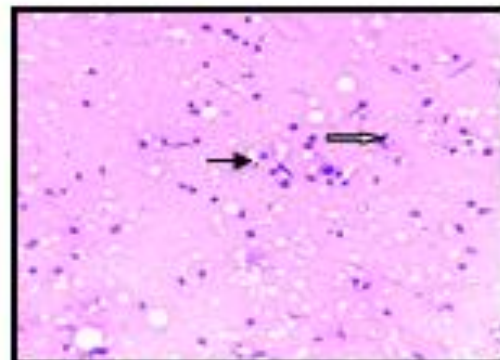


Fig No. 4...; Photomicrograph of FNAC slide, Duct ectasia. ( H& E ;40x) show areas of necrosis along with very few ductal epithelial cells(white arrow) and inflammatory infiltrate comprising of lymphocytes,cyst macrophages(black arrow).

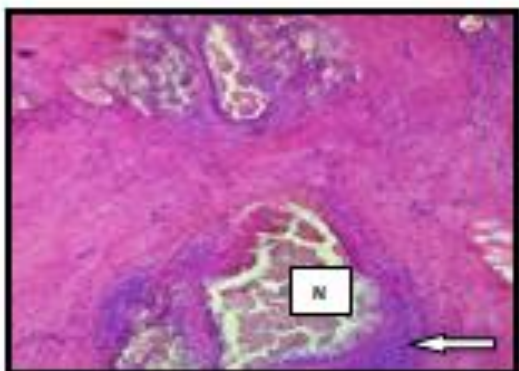


Fig No. 5 : Photomicrograph of Frozen Section slide, Duct ectasia (H &E ;10x) shows dilated ducts filled with necrosis(N)and inspissated secretions surrounded by inflammatory cells(white arrow) and ductal epithelial cells.

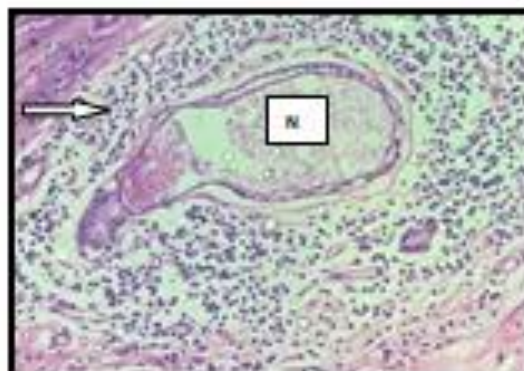


Fig No. 6 : Photomicrograph of Paraffin Section slide, Duct ectasia(H & E; 40x) shows dilated duct filled with necrotic material(N)and surrounded by dense fibrocollagenous tissue and lymphocytic infiltrate(white arrow).

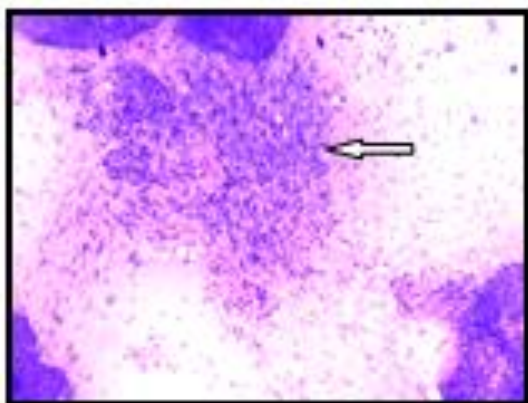


Fig. 7: Photomicrograph of FNAC slide, phylloides tumor (H &E ;10x)shows hypercellular smear showing stromal hypercellularity(white arrow) scattered ductal epithelial cells in the background.

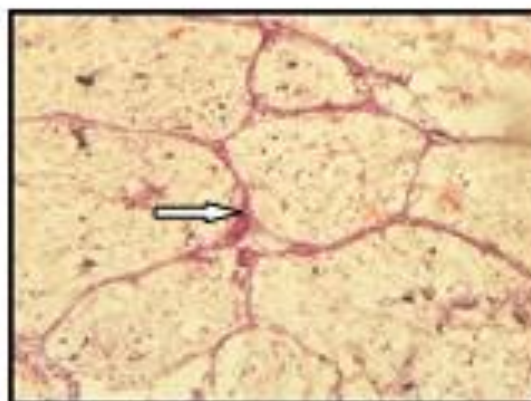


Fig. 8: Photomicrograph of Frozen section slide, phylloides tumor (H &E ;10x)show tumor comprised of exaggerated intracanalicular growth pattern with leaf-like projection(white arrow) into dilated ducts. Stroma is abundant fibrocollagenous.



Fig. 9: Photomicrograph of paraffin section slide, phylloides tumor (H & E ;10x) show tumor comprised of exaggerated intracanalicular growth pattern with leaf-like projection (white arrow) into dilated ducts. Stroma is abundant, fibrocollagenous.

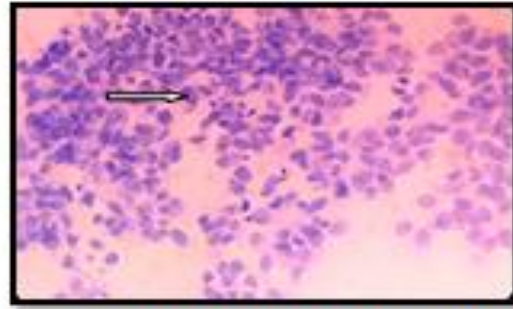


Fig. 10 : Photomicrograph of FNAC slide, IDC (H & E ; 40 X) shows large ductal epithelial cells (white arrow) showing hyperchromatic, pleomorphic nuclei and scanty cytoplasm on haemorrhagic background

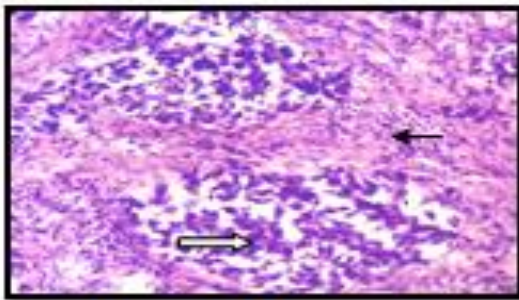


Fig. 11 : Photomicrograph of Frozen section slide, IDC (H & E ;40 X) shows tumour tissue (white arrow) comprised of groups, strands and gland-like structures formed by large, round to oval cells having large, hyperchromatic pleomorphic nuclei and scanty cytoplasm. Stroma is dense fibrocollagenous (black arrow).

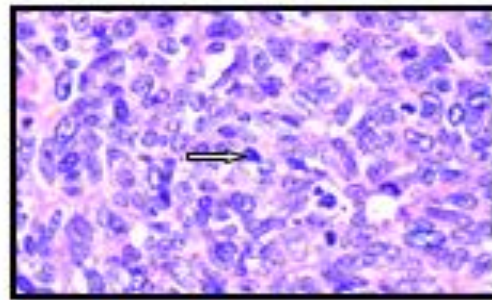


Fig. 12 : Photomicrograph of Paraffin section slide, IDC-High grade (H & E ;40 X) shows tumour tissue comprised of large, round to oval tumor cells having large, pleomorphic, hyperchromatic (white arrow) and at places vesicular nuclei with prominent nucleoli and scanty cytoplasm along with stroma between tumor cells.

## Discussion

Breast carcinoma is one of the commonest cancer among females in India preceded only by cervical cancer.<sup>(10,11)</sup> Early screening and diagnosis of breast lesions can aid in prevention as well as accurate management of the patients thus alleviating discomfort and anxiety in the process.<sup>(12,13)</sup> The present study confirms the clinical utility and efficacy of FNAC, Frozen Section and Histopathological study in the

evaluation of the patient with breast lesions and subclassify them as benign and malignant breast disease wherever possible.

In the present study, most of the patients were in the age group of 31-40 years followed by 51-60 years which was comparable with the study done by Shrestha et al<sup>(14)</sup>.

Most common side of involvement of breast lesion is left side, this finding correlates with the study done by Reddy<sup>(15)</sup> (1958) which showed

left side as the most common side of involvement in breast lesions. In our study we found one (1.96%) case of bilateral side breast involvement.

Most of the patients presented with palpable breast lumps (48 cases), 1 case of nipple discharge and 2 cases of palpable axillary lymph nodes.

In our study, most of the patients 35 cases (68.62%) presented with breast lesions within 0 to 6 months.

In the present study, on FNAC, the benign or non-malignant breast lesions were found in the 43.13% cases and malignant breast lesion in 56.86% cases, this result was similar to the study done by Sheikh et al.<sup>(16)</sup>

Rakhshindah Bajwa et al., found that among benign breast lesions (232 cases), fibroadenoma was the commonest benign breast lesion, 161 cases (69.39%)<sup>(17)</sup>.

In our study, FNAC showed 18 cases (35.29%) of fibroadenoma being the commonest benign lesion followed by 3 cases (5.88%) of phylloides tumor and 1 case (1.96%) of duct ectasia. Out of the 29

cases (56.86 %) of malignant breast lesions, all the cases are of infiltrating duct carcinoma as the most common malignant breast lesion.

On Frozen section study, 16 cases (31.37 %) of fibroadenoma being the commonest benign lesion followed by 3 cases (5.88%) of phylloides tumor and 1 case (1.96%) of duct ectasia. Out of the 31 cases (60.78 %) of malignant breast lesions, all the cases are of infiltrating duct carcinoma as the most common malignant breast lesion.

In the present study, surgically resected margins of all the malignant tumors are free of tumor tissue on frozen section and histopathological examination.

On HPE, 14 cases (27.45 %) of fibroadenoma being the commonest benign lesion followed by 3 cases (5.88%) of phylloides tumor and 1 case (1.96%) of duct ectasia. Out of the 33 cases (64.70%) of malignant breast lesions, all the cases are of infiltrating duct carcinoma as the most common malignant breast lesion.

**Table 7** Showing comparison of statistical analysis of results of sensitivity, specificity & accuracy of FNAC diagnosis of breast disease between different studies (Literature Review)

ar	Author	Sensitivity (%)	Specificity (%)	Accuracy (%)
1	Rocha and Nadkarni <sup>(18)</sup> (1997)	93.8	98.21	---
2	Kamphausen BH et al <sup>(19)</sup> (2003)	90	100	---
3	Naggada <sup>(20)</sup> (2007)	95.7	98.7	97.7
4	Nguansangiam S et al <sup>(21)</sup> 2009)	92.5	90.2	91.2
5	Ahmed H.G.et al <sup>(22)</sup> (2009)	92.6	95.2	---
6	Pudasaini S et.al <sup>(23)</sup> (2011)	93.3	100	---
7	Prakash HM et al <sup>(24)</sup> (2011)	94.5	98	97
8	Sushma Yalavarthiet al <sup>(25)</sup> (2014)	100	88.5	---
9	Paramesh&Saha et al <sup>(26)</sup> (2015)	95.45	100	98.18
10	Daramola A Oet al <sup>(27)</sup> (2015)	95.4	88.9	---
11	Waghmare et al <sup>(28)</sup> (2016)	88.24	100	---
12	David E Ibikunle et.al <sup>(29)</sup> (2017)	99.4	100	---
13	Sanjay C Chauhan et.al <sup>(30)</sup> (2017)	98.24	98.93	---
14	Farida Begum et al <sup>(31)</sup>	--	--	98.1
15	Dr Vasundhara Gardas <sup>(32)</sup> (2018)	66.66	100	---
16	<b>Present study</b>	<b>94.44</b>	<b>84.85</b>	<b>88.24</b>

(---):- parameter not calculated in the study.

Our study shows 94.44% Sensitivity, 84.85% specificity and accuracy of 88.24% which is comparable with results obtained in other studies in the literature.



**Table 8** Comparison of sensitivity, specificity, positive predictive value, negative predictive value and accuracy of Frozen Section in diagnosis of breast disease between different studies (Literature Review)

Sr No	Author	Sensitivity (%)	Specificity (%)	PPV (%)	NPV (%)	Accuracy (%)
1	Hanzal E et al <sup>(33)</sup> (1992)	97.45	100	---	---	---
2	Hou MF et al <sup>(34)</sup> (1995)	99.1	100	---	---	---
3	Simonetta et al <sup>(8)</sup> (1995)	91.7	99.2	---	---	---
4	H Haeri et al <sup>(35)</sup> (2002)	95.4	92.4	---	---	100
5	PV Karve et al <sup>(36)</sup> (2005)	99.36	100	100	98.77	99.82
6	Nuzhat S et al <sup>(37)</sup> (2005)	99.3	93	96.6	99.3	99
7	Tarek MN, et al <sup>(38)</sup> (2008)	100	100	---	---	100
8	Andreas H. Brunner et al <sup>(39)</sup> (2009)	95	100	100	90	---
9	F.Caruso et al <sup>(40)</sup> (2011)	83	93	62	97	94
10	Shah Alam Sheikh et al <sup>(16)</sup> (2016)	97.37	100	100	---	98.57
11	<b>Present Study</b>	<b>88.89</b>	<b>87.88</b>	<b>80.00</b>	<b>93.55</b>	<b>88.24</b>

PPV: Positive Predictive Value; NPV:- Negative Predictive Value  
 (---):- parameter not calculated in the study.

In present study, sensitivity, specificity, positive predictive value, negative predictive value and accuracy of Frozen section is 88.89%, 87.88%, 80.00%, 93.55% and 88.24% respectively which is low as compared to those in other studies may be because of low sample size.

### Conclusion

To conclude, F.N.A.C. is a simple, rapid, cost effective, non-invasive procedure in the early diagnosis of breast lesions and not associated with any complications of the procedure. Despite increasing popularity and undisputed utility of FNAC, there are cases where frozen section still stands out as the method of choice for rapid diagnosis mainly for determining the resection margins of the lesion and the extent of metastasis in case of malignant lesion to ensure no residual

tumour mass thus helping in further treatment and follow-up of patients. Final histopathological study is required to accurately arrive at a definitive diagnosis along with IHC marker study which is considered as a gold standard for patient care.

Thus, a judicious selection of one or more of these modalities is essential in every patient presenting with a suspicious breast lump.

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