

ORIGINAL RESEARCH

Cytopathological evaluation of breast lesions based on the international academy of cytology (IAC) yokohama system for reporting breast fine needle aspiration cytology (FNAC) in a regional cancer centre

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Received: 03 November, 2023

Accepted: 07 December, 2023

ABSTRACT

Introduction: FNAC, which is a minimally invasive procedure, has developed as the most accurate and cost-effective initial method for guiding the clinical management of the patients with breast lesions. The International Academy of Cytology (IAC), Yokohama system for breast fine needle aspiration cytology (FNAC) established a uniform, tired reporting system for breast FNAC specimens. Using this system, the cytopathologist can communicate breast FNAC interpretations to the referring clinician. Based on this objectives of our study is to standardize the reporting system of breast lesion by FNAC. To assign all reported breast lesions in one of the five diagnostic categories as defined by Yokohama system to maintain the uniformity. To correlate with histopathology whenever possible. To assess the risk of malignancy (ROM). **Materials and Methods:** Following institutional scientific and ethical committee approval, this study was undertaken in the Cytology division of Department of Pathology, Kidwai Memorial Institute of Oncology, over a period from 1-dec-2019 to 30-may-2021. This was a descriptive study. All smears from FNAC of breast lesions were interpreted according to IACY okohama system. **Results:** A total of 260 cases of breast lesions were taken during the study period. The demographics constituted 98.1% females and 1.90% males with an age range of 13-81 years and a mean age of 48.9. The most common lesions were malignant (52.3%), while benign constituted 28.8%. Histopathological analysis were available for 148/260 cases. **Conclusion:** FNAC is a reliable, well tolerated diagnostic modality which can be used in diagnosis of breast lesions. Following the IAC Yokohama system, FNA is a useful tool and requires specific training & ongoing experience. Therefore this newly proposed IACY okohama system for reporting breast cytopathology is a simple system that allows greater diagnostic clarity and, consequently, better communication between pathologist and treating clinician, also helps in predicting the ROM.

Keywords: IACY okohama, insufficient, benign, malignant, atypical, suspicious.

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INTRODUCTION

Breast cancer is the most common malignancy in woman worldwide with global incidence of about 2.1 million cases per year and in India, it is 1,62,468 per year¹. Breast lesions include a spectrum of lesions, from benign to malignant.

Fine needle aspiration is one of the primary investigation widely followed, as it is cost effective

and minimally invasive when compared to the core needle biopsy (CNB) which is gold standard². Fine needle aspirate is a simple, rapid, cost-effective, and minimally invasive test for both palpable and impalpable (under ultrasonographic guidance) breast lumps. However, now a days usage of core needle biopsy has gained popularity as it allows evaluation of histological grade as well as hormonal

status[estrogenreceptor(ER)/progesterone receptor (PR)/human epidermal growth factor receptor (Her2)]. Forthediagnosisofbreastcancer,FNAChasahighsensitivityof90%–95%andahigh positive predictive value ~100%³. It has a low false-negative rate related to low-grade ductal andlobularcarcinomaandaverylowfalse-positiveraterelatedtoFNAsoffibroadenomasand papillary lesions of the breast.⁴⁻⁶

Usingultrasonography(USG)guidanceandrapidon-siteevaluation(ROSE)optimizes the usage of FNAC. The cytological findings should be interpreted along with clinical and radiological findings in the “triple test.” With the triple test parameters, the sensitivity and specificity of FNAC is comparable to core needle biopsy.⁵

TheIACYokohamaBreastFNACReportingsystemhas been developed by a group of expert cytopathologists with assistance from surgeons, oncologists, and radiologists. It has been developed to have a standardized reporting system to improve the interpretation of breast cytology. It also aims at improving communication between the cytopathologist and clinician by linking reporting system with management options.⁷

ActuallytheInternationalAcademyofCytology(IAC) established a Yokohama system for breast fine needle aspiration cytology (FNAC) in 2016 intending to develop an internationally recognized and standardized reporting system that would define best practice guidelines for the use of FNAC in diagnosing breast lesions more consistently and accurately.

This guides the clinicians to proceed with management⁷. Our institute being a tertiary cancer center would be ideal to conduct such study. Based in this aim of our study is to standardize the reporting system of breast lesions by FNAC. Also to report all reported breast lesions in one of the five diagnostic categories as defined by Yokohama system to maintain the uniformity and to assess the risk of malignancy.

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MATERIALS AND METHODS

This is descriptive study undertaken at cytology division, Department of pathology at Kidwai Memorial Institute of Oncology (KMIO) for a period of 18 months from 2019 to 2021, to evaluate the cytomorphology of different breast lesions. Institutional approval has been obtained to conduct the study from the hospital ethics committee.

All patients clinically and radiologically diagnosed as breast lesions referred to the cytology division of KMIO for routine and image guided FNAC were included in the study. Clinical and radiological data relevant to the case were collected from the patient and case records. Sample size was 260.

All cases with breast lesion that are referred to Cytology division who were clinically and radiologically evaluated for breast lesions and are will

include informed/written consent were included in the study. Whereas cases not willing to give informed/written consent and review cases from elsewhere for which FNAC was done outside were excluded.

Clinical details like age, gender, radiological findings, and provisional clinical diagnosis were recorded from case files. The anatomical site and size of mass lesions documented by Ultrasonography (USG), mammography were considered. Informed consent before FNA was obtained from the parents/caretakers of patients.

Routine FNA was done in palpable lesions while USG guidance was used for small and difficult lesions. FNAs were performed by trained cytopathologists with the assistance of an interventional radiologist in the presence of a clinician for clinical monitoring. Two to four smears were made for each case depending on the material aspirated. Smears for Papanicolaou's (PAP) stain were immediately fixed. Air-dried smears were stained with May-Grunwald Giemsa (MGG) stain.

Breast FNAC's done was reclassified based on the newly proposed IAC Yokohama System, into five categories and was compared with histopathology, ROM was assessed whenever possible.

Socio demographic variables like the age and gender and the cytological analysis consisting of cytomorphology diagnosis, histopathology diagnosis were reconsidered as relevant variables for the analysis. Descriptive analysis was carried out by mean and standard deviation for quantitative variables, frequency and proportion for categorical variable and the final analysis was done with the use of Statistical Package for Social Sciences (SPSS) software version 23.0.

RESULTS

A total of 260 cases of breast FNA (including both routine and USG guided FNA) were performed during this period. The mean age of the study population was 48.9 years (age range: 13-81 years). The most common being 40 to 49 years age group with 83 patients followed by 50 to 59 years age group. 54 cases were above 60 years in age. Of 260 subjects, there were 255 females constituting to 98.1% and 5 males constituting to 1.90%.

The study group showed equal involvement of right and left breast lesions in 124 subjects (47.7%) each respectively. Bilateral breast involvement seen in 12 subjects (4.60%). Of the 260 subjects involved in this study, 121 subjects (46.5%) underwent routine FNA and 139 subjects went through USG guided FNA (53.5%)

All cases were interpreted cytologically and subjected into Yokohama category. The most common interpretation was category V in 136 cases constituting to 52.3% and second is category II in 75 cases constituting to 28.8%. 27 cases showed sparse cellularity or too poorly smeared or fixed to allow a cytomorphological diagnosis and were interpreted as category I constituting to 10.4%. There were

15(5.80%) cases interpreted as category IV and 7 (2.70%) cases interpreted as category III.

Table 1:-Distribution of Yokohama diagnostic category of study subjects.

S.No.	Yokohama Category	No. of subjects(%)
1	Category-I	27(10.4)
2	Category-II	75(28.8)
3	Category-III	07(2.70)
4	Category-IV	15(5.80)
5	Category-V	136(52.3)
Total		260(100.0)

Histopathological correlation was available for 148 cases of 260 subjects involved. Of the correlated 148 cases, 118 cases (80%) were category V, 26 cases (17.9%) were benign, 1 case (0.7%) of category I, III, IV respectively.

Table 2:-Distribution of HPE results of study subjects.

Yokohama category	HPE categories					Total no. of study subjects
	I	II	III	IV	V	
I	0	6	0	0	9	27
II	1	17	0	0	8	75
III	0	0	1	0	1	7
IV	0	3	0	1	5	15
V	0	0	0	0	95	136
Total	1	26	1	1	118	260

Respective ROM for each category was calculated ROM is higher for the category V with 69.8%.

Table 3:-Distribution of ROM of study subjects.

Yokohama category	No of cases(%)	Benign	Malignant	ROM(%)
I	27(10.4)	06	09	33.3
II	75(28.8)	17	08	11.1
III	07(2.70)	00	01	14.3
IV	15(5.80)	03	05	33.3
V	136(52.3)	00	95	69.8

In our study 118 of the 136 subjects interpreted as malignant under category V of Yokohama were falling in the age group of 40->60 years and 57 cases of the 75 subjects interpreted as benign, category II of Yokohama were falling in the age group of 20- 49 years. Malignant category, category V of Yokohama has 136 cases of which 134 cases (52.3%) were female and 2 cases (40.0%) were male.

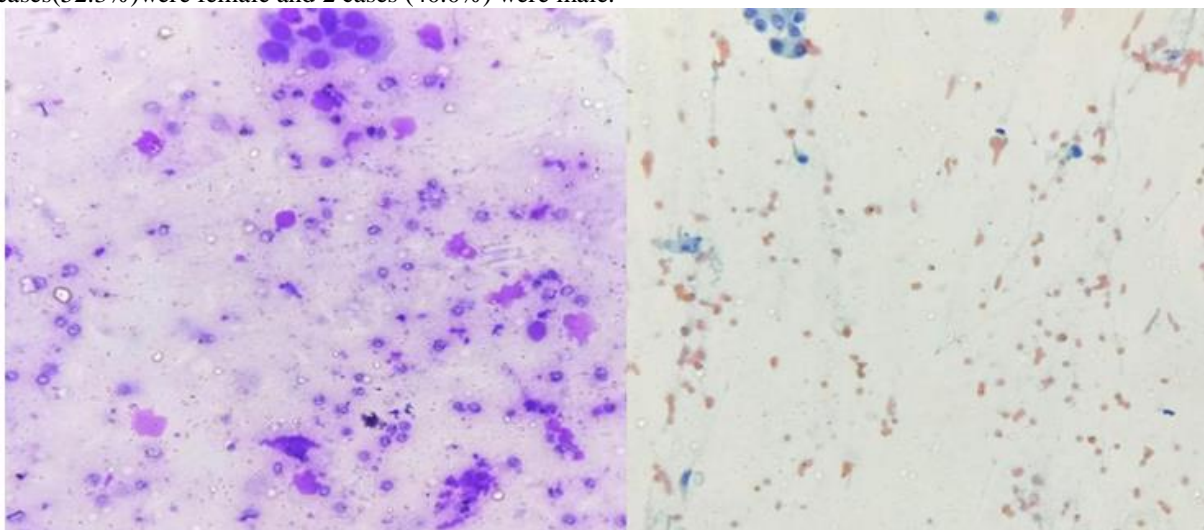


Figure 1: MCG and PAP stains smear of Yokohama category 1-Insufficient/inadequate

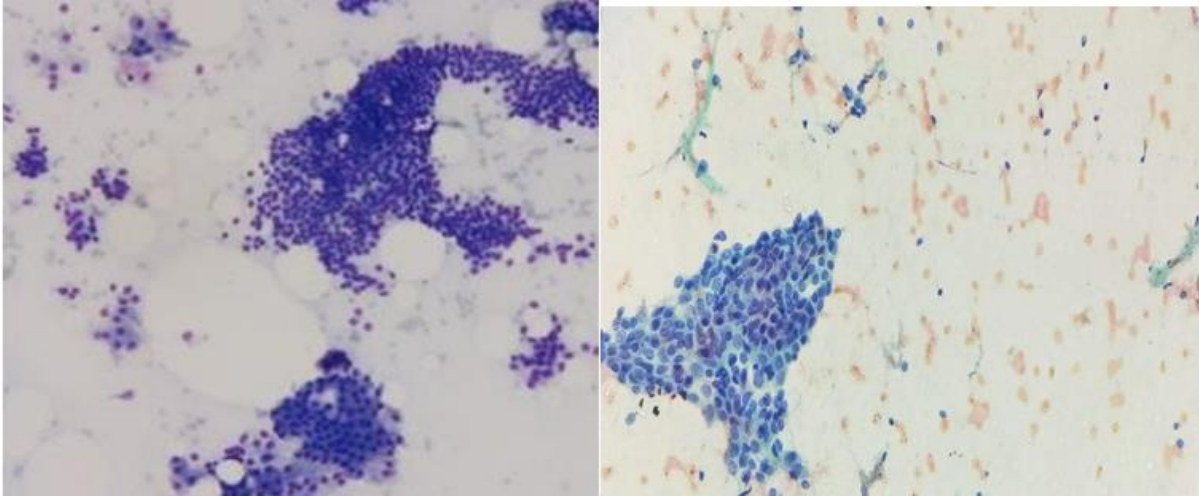


Figure2: MCG and PAP stains smear of Yokohama category 2- Fibroadenoma

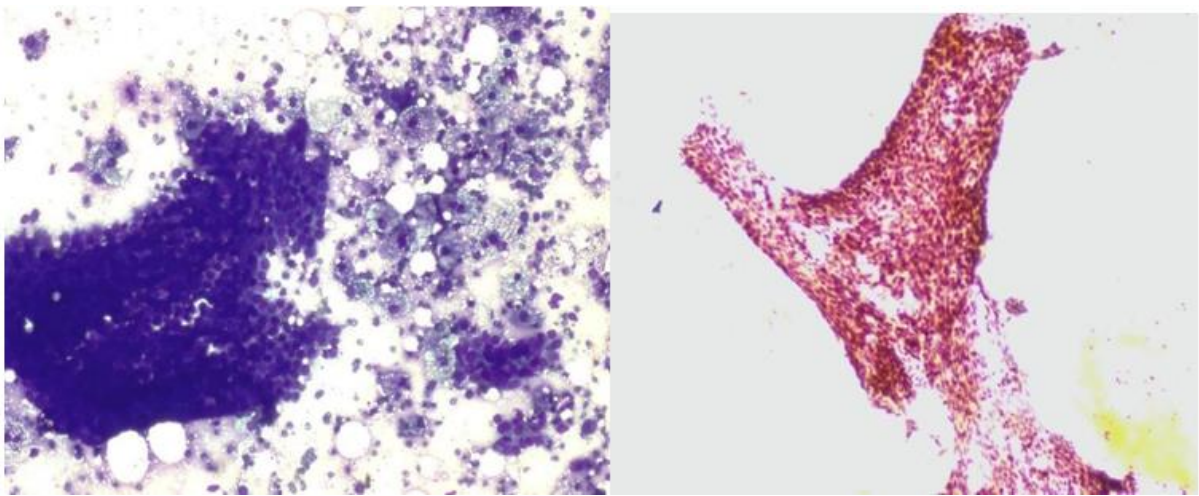


Figure3: MCG and PAP stains smear of Yokohama category 3- Atypical cells

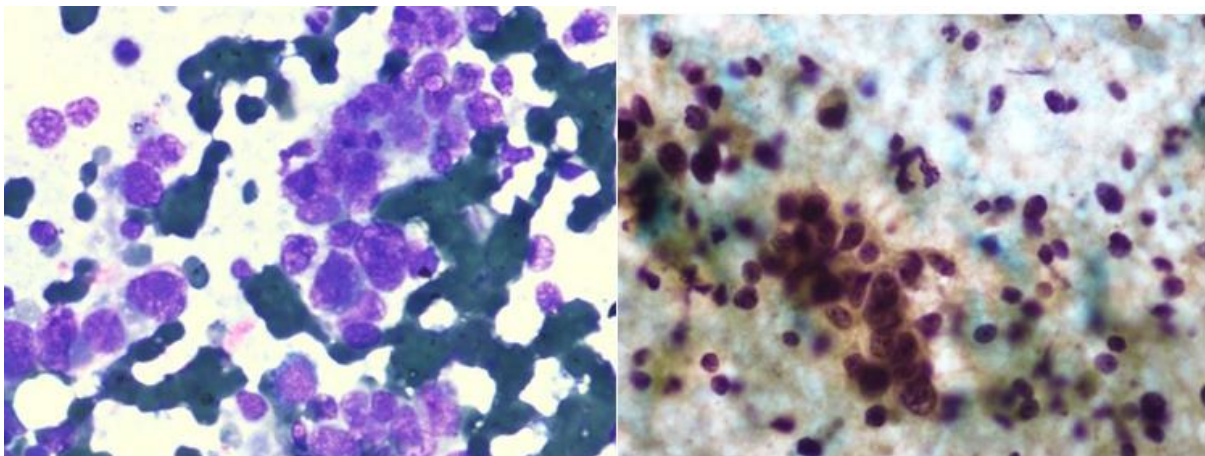


Figure4: MCG and PAP stains smear of Yokohama category 4- Suspicious of malignancy

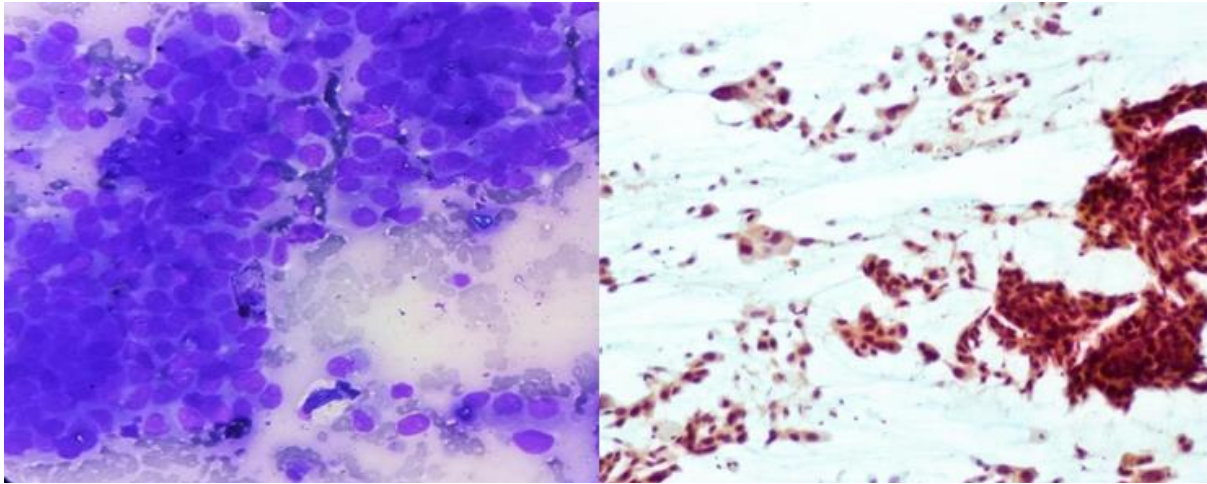


Figure 5: MCG and Papanicolaou (PAP) stain smear of Yokohama category 5 - Duct carcinoma DISCUSSION

Breast cancer is the commonest malignant tumor of all female cancers, and there is increasing incidences, morbidity, and mortality globally. There are various diagnostic modalities by which material/tissue samples can be obtained from the breast. They are Percutaneous FNAC, USG guided FNAC and CNB. Martin and Ellis first introduced the application of FNAC for the diagnosis of breast masses in 1930. Since then it has been established as an essential tool for evaluating breast lesions although CNB is gold standard⁸. Even FNAC material can be used for ancillary techniques like Immunohistochemistry and molecular testing with satisfactory results⁹. Cytology material obtained by the FNAC has been shown to provide good quality of DNA and molecular in situ techniques performed on cytology material show an optimal concordance with histology¹⁰. The least invasive with least amount of turnaround time is the FNAC procedure. Percutaneous FNAC using USG is used for lesions. It offers real time monitoring of sample collection. USG guided FNA is used for small lesions. Large and palpable lesions Routine FNAC is used. FNAC has minimal complication. There were no reported complications of FNAC procedure in this study other than mild pain and discomfort.

The Yokohama system, which utilizes the triple assessment approach, followed the same criteria in present study. This approach helps assess breast lesions, which combines clinical, radiological, and pathology information to ensure accurate diagnosis and patient management^{2,3,11,12}. The IAC Yokohama system for reporting breast FNA cytology defines five categories for reporting breast cytology and help to stratify breast lesions by their ROM and give a management algorithm for each category.

The ROM was assessed and compared with the recent studies^{4-6,13,14}. In the present study it was observed that ROM amounted to 33.3% in category 1 and 10.6% in category 2, as compared with the other studies, we had an increased number of cases reported as insufficient category, probably due to lack of training and experience of radiologist and cytology technician,

as the procedure and smearing was done by new trainee students due to which the material obtained was sparse and smearing was suboptimal, however is correlating with 30.3% of Hoda et al meta-analysis.⁴ In study done by Montezuma et al the respective ROM for each category was 4.8% for category 1 (insufficient material), 1.4% for category 2 (benign), 13% for category 3 (atypical), 97.1% for category 4 (suspicious for malignancy), and 100% for category 5 (malignant).⁶

A retrospective study by Wong S et al., over a period of 36 months on 2,696 breast FNAC's, where 579 cases with matched histopathology and 456 cases had Rapid Onset evaluation (ROSE), showed lower ROM for categories 1 and 2 when compared with the present study⁵. Incorporating ROSE helped them in decreasing the percentage of insufficient from 17.1% without ROSE to 4% with ROSE.

A meta-analysis done by Hoda RS and Brachtel EF, included publications between January 1, 1997, and December 31, 2017, by reviewing literature obtained a case-cohort of 33,341 breast FNAC's, which was drawn from 27 studies through a PubMed database⁴. They collected data for the number of total cases and each category when available and calculated ROM for each category. Their ROM for the insufficient category was 30.3%, on the higher side than other mentioned studies, however was similar to present study^{2,6,13}. They concluded that the diagnostic categories of the

new IAC Yokohama System each carry an implied ROM, which increases from the benign to malignant categories as observed in other studies, including the present study.

Another study done by Wai CJ et al depicted physical examination was 92% accurate (95% confidence interval [CI] 0.89-0.96, $p < 0.0001$) at predicting whether a mass was benign or malignant. Imaging was 88% accurate (95% CI 0.84-0.92, $p < 0.0001$) and needle biopsy was 95% accurate (95% CI 0.92-0.98, $p < 0.0001$). The modified triple test was 99% accurate (95% CI 0.98-

1.00, $p < 0.0001$). Each 1-point increment in the mTTS was associated with an increased risk of cancer, with an odds ratio of 9.73 (CI 5.16-18.4, $p < 0.0001$)¹³

In an Indian study by Kamatar PV et al., they analysed a total of 470 cases, obtained between January 2017 and December 2018, which included 453 (96%) female patients with 17 (4%) male patients¹⁴. They retrospectively reviewed breast FNAC cases, and they also observed that ROM was an increase from benign to malignant categories as observed in our study and other studies as well.

Chauhan Vet et al., a study on 468 patients, where in they had a more significant number of category 2 (Benign) cases, 342 (73.07%) and category 4, category 5 amounting to 11 (2.35%) and 85 (18.16%), when compared with present study where we had 52.3% (136) of category five cases, 5.80% (15), 28.8% (75) category four and two respectively¹⁵.

CONCLUSION

FNAC is a reliable, well-tolerated diagnostic modality which can be used in diagnosis of breast lesions. FNAC can be used to arrive at a diagnosis. In situations where the patient is too sick and other more invasive methods cannot be used, FNAC becomes the diagnostic modality of choice for the initial management of such patients. The turnaround time with FNAC is less than helping in the overall management of patients. Following the IAC Yokohama system, FNA is a useful tool and requires specific training & ongoing experience.

Therefore, this newly proposed IAC Yokohama system for reporting breast cytopathology is a simple system that allows greater diagnostic clarity and, consequently, better communication between pathologist and treating clinician, also helps in predicting the ROM.

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