

ROLE OF IMPRINT CYTOLOGY IN DIAGNOSIS OF BREAST LUMPS-A REVIEW

Dr Vinod k	Department of General surgery Mahatma Gandhi medical college and Research institute Sri Balaji Vidyapeeth University.
Dr Tirou Aroul S	Department of General surgery Mahatma Gandhi medical college and Research institute Sri Balaji Vidyapeeth University.
Dr Anandraj Vaithi	Department of General surgery Mahatma Gandhi medical college and Research institute Sri Balaji Vidyapeeth University.
Dr Saravana Kumar S	Department of General surgery Mahatma Gandhi medical college and Research institute Sri Balaji Vidyapeeth University.

ABSTRACT
In yet another study conducted by Khanna et all Fine needle aspiration cytology, Imprint cytology and Tru-cut biopsy were done in a total of 86 patients with breast lump and the results were finally interpretated and the data compared FNAC, IC and Tru cut biopsy and it was found that the sensitivity and specificity of FNAC was 96.8% and 100% whilst imprint cytology had a sensitivity and specificity 98.4%, 100%, Tru cut 100% sensitivity and specificity. Of the 86 patients taken up for the study 15 people were rejected due to inadequate sample



KEYWORD-

histopathological, imprint cytology, squash smears

*Corresponding Author Dr Tirou Aroul S

Department of General surgery Mahatma Gandhi medical college and Research institute Sri Balaji Vidyapeeth University.

INTRODUCTION:

Breast diseases are commonly encountered clinical entities in surgery outpatient department. Fine needle aspiration cytology is a well established diagnostic modality in detecting breast lumps with an increased sensitivity, core needle biopsy has even more accuracy than fine needle aspiration cytology in diagnosing breast lumps .Imprint cytology is touch preparation of the intra-operative specimen in the form of cells in glass slide followed by examination (1)

Intraoperative evaluation of histopathological type during surgery might help the surgeon decide the appropriate surgery on table. Intraoperative imprint cytology remains still a topic of controversy(2)

The most common malignancy amongst woman is breast carcinoma. Despite adequate recognition of histological variant by needle biopsies and also evaluation of lumpectomy margins and operative nodal status by frozen sections it has an added advantage of delivering the results within minutes that might influence therapeutic decisions.(3). The incidence of breast carcinoma in Indian population is 25.8 per one lakh population and it's been estimated to be around 112 cases per 100000 patients being screened for breast cancer (4,5)

Imprint cytology has a better visualization of morphological features of cells (8)It has a great impetus to histological diagnosis due to its rapidity, simplicity and cost effectiveness (9). Apart from diagnosing breast lesions it is also being employed in the diagnosis of a wide variety of other lesions including thyroid, sentinel nodes, endoscopic biopsy and prostate etc(10,11)The sensitivity and specificity of imprint cytology in breast lesions was found to be 100% and 96.43%(3)

METHODS:

Relevant publications collected from Pub Med , Google scholar, EMBASE, MEDLINE , Cochrane database of systematic reviews were included in the review. The key search words used were imprint cytology, Histopathology and breast lesions/lumps .No restrictions were made based on date of publication. Duplicate copies of the publications were eliminated. Data compiled based on accuracy of imprint cytology in the diagnosis of breast lumps.

DISCUSSION: IMPRINT CYTOLOGY

The intraoperative accuracy of tumor is essential for the patient's treatment. Though intraoperative accuracy of a tumor can be achieved with the help of frozen section and cytologic examination. Various methods have been established using FNAC, imprint cytology and squash smears to establish the tumor and surgical margins.

Origin of cytology dates back for more than a century in diagnostic pathology. In imprint cytology tissue is touched onto the slide and it leaves behind its imprint in the form of cells on a glass slide .Diagnostic cytology is the science of interpretation of cells on epithelial surface or from the ones derived from various sources by artificial means(25). Imprint cytology has been a useful alternative to frozen section in underdeveloped countries(26). Imprint cytology is a form of touch preparation wherein tissues are touched onto a slide only to create imprint on the glass slides (cell forms)(27). Indeed a second surgery could be avoided if a well documented imprint report is obtained (28). Similar imprint studies have been conducted even in the setting of endoscopic cytology by Hughes et al to aid in the diagnosis of carcinoma stomach(29). Certain studies have disparities in efficacy of imprint cytology (30). Similar studies by Tew et al to

evaluate the evidence of metastasis to sentinel node have also been conducted and were proven to have increased efficacy(31).

In a study by Kashiwagi et al imprint of core needle biopsies have been done and it was proven to be a effective rapid reliable method for diagnosing breast lesions(32). Imprints have also been used in assessing the surgical margins in operative cases of breast lesions(33).

In a study conducted by Narendranath Swain, Bharat Kumar Behera, Sukanti Majhi, Manindra Nayak et al ,in about 64 patients 60 patients were female and 4 patients are male with female to male ratio being 15:1 with average age range between 11-70 years with mean age in females being 40.5 years and male being 40.5 years .Amongst 64 people the chief complaints were breast lump in 100 percent of patients related to mastlgia in 24 ,menstrual cycle in 12,nipple discharge in 4 and skin ulceration in 8, anatomically it was in upper outer quadrant in 30,12 in upper inner quadrant,11 in lower outer quadrant,8 in lower inner quadrant and 3 in central region. On cytological examination the cause was inflammatory in 3 people, benign in 32 numbers and malignant in 29 people. FNAC had a sensitivity of 93.3% and a accuracy of 95.3%.tru cut needle biopsy had a sensitivity of 100% and specificity of 100% and a accuracy of 98.4%.Imprint cytology had a sensitivity of 96.4% and a overall accuracy of 95.3% (43).

In yet another study conducted by Khanna et all Fine needle aspiration cytology, Imprint cytology and Tru-cut biopsy were done in a total of 86 patients with breast lump and the results were finally interpretated and the data compared FNAC,IC and Tru cut biopsy and it was found that the sensitivity and specificity of FNAC was 96.8% and 100% whilst imprint cytology had a sensitivity and specificity 98.4%, 100%, Tru cut 100% sensitivity and specificity. Of the 86 patients taken up for the study 15 people were rejected due to inadequate sample(44).

In a study done by Dr Ravindran Chirukandathl, Dr Venita Juliet Noronha, Dr RemaniK, Dr Sarath Krishnan imprint cytology was done on a total of 110 freshly excised surgical specimens and the results were found to be astounding with an impeccable accuracy rate. It was found that the imprint cytology ad a sensitivity of 95.6% and a specificity of 100%. although the draw backs were the lymphoma detection was not accurate. Hence the imprint cytology has been established to be a quick alternative for intraoperative specimen histopathological examination (28).

In yet another study by Asha Mahadevappa, Thattamparambil Gopalakrishnan Nisha, GubbannaV Manjunath. A total number of 62 patients were included in the study one was excluded as the sample was inadequate hence only 61 patients were included in the study and compared with histopathological examination. In imprint cytology and frozen section one was false positive due to error in interpretation. Imprint correlation with histopathological examination and showed a total of 93.1 percent correleation. Sensitivity and specificity of imprint cytology was 100% and 96.4%. accuracy was 98.6%. frozen section had a sensitivity and specificity of 100 and 96.55%(3).

In a study conducted by Ramraje et al with a main objective to prove the accuracy of imprint cytology in comparison with histopathological examination the study was conducted in a total of 90 patients over a period of 2 years and it was found that imprint cytology diagnosed 81 cases of 90 accurately which is approximately 90%(45).

In a study conducted by Ronald enrique Delgado bocanegra et althe mean age was found to be 51 years and the most

common histological variant was ductal carcinoma .Imprint cytology had a sensitivity of 61.8% and an accuracy of 86.3%(46).

In a study conducted by Motomura k et al imprint cytology results were compared with frozen section and imprint cytology had a sensitivity and specificity of 96% and 90% while frozen section had a sensitivity and specificity of 90% and 98.5% and thus again proving the diagnostic accuracy of imprint cytology amongst other diagnostic modalities (47). In a study conducted by Petroupoluet al ,a total of 60 patients were studied and sensitivity and specificity of sentinel lymph node was 90% and 100% and 80% and 100% for frozen section (48).

Intraoperative diagnosis of breast cancer deposits in axillary nodes is essential to prevent a second surgery. Imprint cytology is an effective technique aiding in the diagnosis with a high specificity and sensitivity. In a study conducted by K Tew et al which was a metanalysis. Thirty one studies were included and they all showed a pooled sensitivity of imprint cytology to be 63 percent and specificity of 99 percent .and frozen section had more sensitivity and specificity than imprint cytology(31).

In another study conducted by Karla Esbona et al intraoperative imprint cytology in assessment of surgical margins the final results were interpretated in terms of reexcision and was found to be 35% for histopathology,11% for imprint cytology and 10% for imprint cytology and the sensitivity was 83% for imprint cytology and 72% for frozen section(50).

In a study done by Chandrakar et all the results showed out of 110 cases, 85 were diagnosed to be malignant and 25 were benign whilst 11 cases were reported to be false negative. Of these 85 malignant cases 78 were correctly diagnosed, 7 were negative. Accuracy of imprint was 83.63%(27).

CONCLUSION:

Both FNAC and imprint cytology have got their own merits and demerits. In concordance with the evidences in literature our study has shown that imprint cytology has high specificity and sensitivity in diagnosing breast lumps. However histopathology has got a better sensitivity, specificity in diagnosing breast lump. Larger study population and multi centric study may be required in establishing further efficacy of imprint cytology.

REFERENCES:

- H C, Das S. Diagnostic efficacy of imprint cytology and frozen section of breast lesions. Indian J Pathol Oncol. 2019 Feb 15;6(1):128–36.
- 2. Khalid A, Haque AU. Touch Impression Cytology Versus Frozen Section as

 $Intra operative\ Consultation\ Diagnosis\ International\ Journal\ of\ Pathology; 2004;$

2(2):63-70.

- Mahadevappa A, Nisha TG, Manjunath GV. Intra-operative Diagnosis of Breast Lesions by Imprint Cytology and Frozen Section with Histopathological Correlation. J Clin Diagn Res JCDR. 2017 Mar; 11(3)
- 4. Shreshtha M,Sarangadhara A B,Uma D ,Sunita S.Epidemiology of breast cancer
- in Indian women Malvia 2017 Asia-Pacific Journal of Clinical Oncology, volume 13(4)
- Bleyer A, Welch HG. Effect of Three Decades of Screening Mammography on Breast-Cancer Incidence. N Engl J Med. 2012 Nov 22;367(21):1998–2005.
- Brem RF, Floerke AC, Rapelyea JA, Teal C, Kelly T, Mathur V.
 Breast-specific Gamma Imaging as an Adjunct Imaging Modality for the Diagnosis of Breast Cancer. Radiology.

2008 Jun 1:247(3):651-7.

- Farokhzad OC, Langer R. Nanomedicine: Developing smarter therapeutic and diagnostic modalities. Adv Drug Deliv Rev. 2006 Dec 1;58(14):1456-9.
- 8.Kamatchi V,Aravindha babu N,Leena sankari S,Rajesh E Imprint cytology
- Journal of pharmacy and bioallied science2015 Apr; 7(Suppl 1):S207–S208.
- 9. Gore CR, Singh BK, Chandanwale SS, Gurwale SG, Kumar H, Bawikar R, et al. Imprint cytology: A boon in tissue diagnosis. Med J Dr Patil Univ. 2017 Jan 1;10(1):58.
- Suen KC, Wood WS, Syed AA, Quenville NF, Clement PB. Role of imprint cytology in intraoperative diagnosis: value and limitations. J Clin Pathol. 1978 Apr 1;31(4):328–37.
- Gentry JF. Pelvic lymph node metastases in prostatic carcinoma. The value of touch imprint cytology. Am J Surg Pathol. 1986 Oct;10(10):718–27.
- 12. John B,Stan lee K,Farhad K,Tacin O The History of the Breast ,Basic Human anatomy ,2012 Jan;1(2)213-215
- Brunicardi FC, editor. Schwartz's principles of surgery. Eleventh edition. New York: McGraw-Hill; 2018;500-525.
- Nguyen C, Kettler MD, Swirsky ME, Miller VI, Scott C, Krause R, et al. Male Breast Disease: Pictorial Review with Radiologic-Pathologic Correlation. RadioGraphics. 2013 May 1;33(3):763–79.
- 15. Hughes L Classification of benign breast disorders. The ANDI classification based on physiological processes within the normal breast. British medical bulletin 1991 Apr;47(2)251-7.
- Lee M, Soltanian HT. Breast fibroadenomas in adolescents: current perspectives. Adolesc Health Med Ther. 2015 Sep 2:6:159–63.
- 17. Chen Y-Y, Fang W-H, Wang C-C, Kao T-W, Chang Y-W, Yang H-F, et al. Examining the Associations among Fibrocystic Breast Change, Total Lean Mass, and Percent Body Fat. Scientific Reports 2018;8;9180 Jun 15
- Boufelli G, Giannotti MA, Ruiz CA, de Barros N, Chala LF, Maesaka JY, et al. Papillomas of the breast: factors associated with underestimation. Eur J Cancer Prev. 2018 Jul;27(4):310–4.
- Parker SJ, Harries SA. Phyllodes tumours. Postgraduate Medical Journal. 2001 Jul 1;77(909):428–35.
- London SJ, Connolly JL, Schnitt SJ, Colditz GA. A Prospective Study of Benign Breast Disease and the Risk of Breast Cancer. JAMA. 1992 Feb 19;267(7):941–4.
- 21. Crooke PS, Justenhoven C, Brauch H, Dawling S, Roodi N, Higginbotham KS, et al. Estrogen metabolism and exposure in a genotypic-phenotypic model for breast cancer risk prediction. Cancer Epidemiol Biomark Prev Publ Am Assoc Cancer Res Cosponsored Am Soc Prev Oncol. 2011 Jul;20(7):1502-15.
- Ford D, Easton DF, Bishop DT, Narod SA, Goldgar DE. Risks of cancer in BRCA1-mutation carriers. The Lancet. 1994 Mar 19;343(8899):692–5.
- Chen S, Parmigiani G. Meta-Analysis of BRCA1 and BRCA2 Penetrance. J Clin Oncol Off J Am Soc Clin Oncol. 2007 Apr 10;25(11):1329–33.
- 24. Mehta R,Kunal T,Goyal N,Basak U,Gupta A. Triple approach for diagnosing breast lesions-experience at a Tertiary Care Hospital Journal of marine medical society2017;vol19(2)123-127
- Verma R, Kumar M, Khan SS, Sachdev AS. Imprint cytology: an appraisal. J Oral Med Oral Surg Oral Pathol Oral Radiol. 2016;2(2):110.
- 26. Oncology Overview: Selected Abstracts, a Service of the International Cancer Research Data Bank. U.S. Department of Health and Human Services, Public Health Service, National Institutes of Health, National Cancer Institute.; 1981.148 p.
- Chandrakar J, Srivastava S. Evaluation of the relevance of touch imprint cytology in the diagnosis of various neoplastic lesions. Int J Res Med Sci. 2017 Jan

- 16;3(11):3046–50.
- Chirukandath D R. Accuracy of intra-operative imprint cytology in breast tumours and lymph node swellings: Can it be a useful alternate to Frozen section. Journal of Medical Science Clinical Residence. 2019 Jun5;vol(2):46-47.
- Young JA, Hughes HE. Three year trial of endoscopic cytology of the stomach and duodenum. Gut. 1980 Mar 1;21(3):241-6.
- Hiregoudar AD, Godhi AS, Malur PR, Gogeri BV, Metgud SC. Accuracy of intra-operative imprint smears in breast tumours: A study of 40 cases with review of literature. Indian J Surg. 2006;68(6):4.
- 31. Tew K, Irwig L, Matthews A, Crowe P, Macaskill P. Metaanalysis of sentinel node imprint cytology in breast cancer.BJS.2005;92(9):1068–80.
- 32. Kashiwagi S, Onoda N, Asano Y, Noda S, Kawajiri H, Takashima T, et al. Adjunctive imprint cytology of core needle biopsy specimens improved diagnostic accuracy for breast cancer. SpringerPlus 2013 Aug 6(2):12.
- Bakhshandeh M, Tutuncuoglu SO, Fischer G, Masood S. Use of imprint cytology for assessment of surgical margins in lumpectomy specimens of breast cancer patients. Diagn Cytopathol. 2007 Oct; 35(10):656-9.
- 34. Britton PD. Fine needle aspiration or core biopsy. The Breast. 1999 Feb 1;8(1):1-4.
- 35. Lever JV, Trott PA, Webb AJ. Fine needle aspiration cytology. J Clin Pathol. 1985 Jan 1;38(1):1-11.
- 36. Ciatto S, Brancato B, Risso G, Ambrogetti D, Bulgaresi P, Maddau C, et al. Accuracy of fine needle aspiration cytology (FNAC) of axillary lymph nodes as a triage test in breast cancer staging. Breast Cancer Res Treat. 2007 May 1;103(1):85–91.
- Das MK, Koirala A. Cytological evaluation of breast lesion and its histopathological correlation in a tertiary care center. J Nobel Med Coll. 2017;6(2):66–71.
- Khanam KF, Akter N, Tabashum T, Raza AKMM, Hosna AU, Rahman F, et al. A Clinicopathologic Study of Various Breast Lesions by Fine Needle Aspiration Cytology (FNAC). J Curr Surg. 2018 Oct 17;8(3-4):27-31-31.
- Badge SA, Ovhal AG, Azad K, Meshram AT. Study of fineneedle aspiration cytology of breast lumps in rural area of Bastar district, Chhattisgarh. Med J Dr Patil Univ. 2017 Jul 1:10(4):339.
- 40. Chandanwale SS, Rajpal M, Jadhav PS, Sood SK, Gupta K, Gupta N. pattern of benign breast lesions on fnac in consecutive 100 cases : a study at tertiary care hospital in India. Annals of International Medical and Dental Research 2013vol 3(3).
- Zimmermann CJ, Sheffield KM, Duncan CB, Han Y, Cooksley CD, Townsend CM, et al. Time Trends and Geographic Variation in Use of Minimally Invasive Breast Biopsy. J Am Coll Surg. 2013 Apr;216(4):814–24.
- Sun W, Li A, Abreo F, Turbat-Herrera E, Grafton WD. Comparison of fine-needle aspiration cytology and core biopsy for diagnosis of breast cancer. Diagn Cytopathol. 2001;24(6):421-5.
- 43. Swain N, Behera BK, Majhi S, Nayak M. A Comparative Evaluation of Fine Needle Aspiration Cytology, Trucut Needle Biopsy & Imprint Cytology and Histopathology in Breast lumps. Ann Int Med Dent res vol3Issue3:77
- 44. Khanna AK, Singh MR, Khanna S, Khanna NN. Fine Needle Aspiration Cytology, Imprint Cytology and Tru-Cut Needle Biopsy in Breast Lumps: A Comparative Evaluation. J Indian Med Assoc. 1991 Jul;89(7):192–5.
- 45. Ramraje SN, Bharambe BM, Tote VD. Imprint smear cytology and histopathology of breast lesions - a comparative evaluation with review of literature. International Journal of Pharmacy and Biological science2012 issue 4;129-138
- Delgado-Bocanegra RE, Millen EC, Nascimento CM do, Bruno K de A, Delgado-Bocanegra RE, Millen EC, et al. Intraoperative imprint cytology versus histological

- diagnosis for the detection of sentinel lymph nodes in breast cancer treated with neoadjuvant chemotherapy. Clinics2018:73:e363
- 47. Motomura K, Inaji H, Komoike Y, Kasugai T, Nagumo S, Noguchi S, et al. Intraoperative sentinel lymph node examination by imprint cytology and frozen sectioning during breast surgery. BJS. 2000;87(5):597-601.
- Petropoulou T, Kapoula A, Mastoraki A, Politi A, Spanidou-Karvouni E, Psychogios I, et al. Imprint cytology versus frozen section analysis for intraoperative assessment of sentinel lymph node in breast cancer. Breast Cancer Targets Ther. 2017 May 5;9:325–30.
- 49. Celebioglu F, Sylvan M, Perbeck L, Bergkvist L, Frisell J. Intraoperative sentinel lymph node examination by frozen section, immunohistochemistry and imprint cytology during breast surgery – A prospective study. Eur J Cancer. 2006 Mar 1;42(5):617–20.
- Esbona K, Li Z, Wilke LG. Intraoperative Imprint Cytology and Frozen Section Pathology for Margin Assessment in Breast Conservation Surgery: A Systematic Review. Ann Surg Oncol. 2012 Oct 1;19(10):3236–45.
- Santen RJ. Benign Breast Disease in Women. In: Feingold KR, Anawalt B, Boyce A, Chrousos G, Dungan K, Grossman A, et al., editors. Endotext 2018 May 25vol18:(2);23.
- Samoli E, Trichopoulos D, Lagiou A, Zourna P, Georgila C, Minaki P, et al. The hormonal profile of benign breast disease. Br J Cancer. 2013 Jan 15;108(1):199–204.
- 53. Forae GD, Nwachokor FN, Igbe AP, Odokuma EI, Ijomone EA. Benign breast diseases in Warri Southern Nigeria: A spectrum of histopathological analysis. Ann Niger Med. 2014 Jan 1;8(1):28.
- 54. Francis IM, Das DK. Role of Fine Needle Aspiration, Intraoperative Imprint Cytology and Frozen Section in the Diagnosis of Breast Lumps and Thyroid Lesions. Med Princ Pract. 1999;8(3):173–82.
- M R S, Ahmed Z, N U. the diagnostic accuracy of imprint cytology in breast lesions. J Evol Med Dent Sci. 2015 Mar 25;4(25):4299–307.
- Akhtar Z.M.,1 Qureshi S.A.,2 Aziz N.,3 Niazi S.,4 Qureshi S.S.,5 Bukhari M.H. 6 evaluation of accuracy of intra operative imprint annals vol 16. no.1 jan. - mar. 2010.:7.