

## Role of nuclear medicine in breast cancer

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### **Abstract:**

In the world, this breast cancer is the most popular and growing disease especially in present situations. By early detection only we can prevent this cancer. And there are so many cases that are handled by early detection and decrease the rate of death. And also many research works have been done on this breast cancer. In this paper we are discussing about detection of breast cancer through nuclear medicine. Routine nuclear medicine is a major contribution to a complete range of clinical studies such as early lesion detection and stratification; monitoring, guiding, progression in monitoring, recurrence or metastases and predicting response to therapy. Particularly recent developed instrumentation such as high-resolution device coupled with the diagnostic versatility of conventional cameras have reinserted nuclear medicine as a valuable tool in the broader clinical setting. This shows the outline of nuclear medicine, concluding that targeted radiopharmaceuticals and versatile instrumentation position nuclear medicine as a powerful modality for patients with breast cancer.

### **Introduction:**

In the world, this breast cancer is the most popular and growing disease especially in present situations. By early detection only we can prevent this cancer. And there are so many cases that are handled by early detection and decrease the rate of death. And also many research works have been done on this breast cancer. In this paper we are discussing about detection of breast cancer through nuclear medicine. Routine nuclear medicine is a major contribution to a complete range of clinical studies such as early lesion detection and stratification; monitoring, guiding, progression in monitoring, recurrence or metastases and predicting response to therapy. Particularly recent developed instrumentation such as high-resolution device coupled with the diagnostic versatility of conventional cameras have reinserted nuclear medicine as a valuable tool in the broader clinical setting. This shows the outline of nuclear medicine, concluding that targeted radiopharmaceuticals and versatile instrumentation position nuclear medicine as a powerful modality for patients with breast cancer.

While changing our lifestyle factors such as excessive alcohol consumption, obesity and physically inactive have also contributed to increased the rate of death from breast cancer.

Mortality rate which is caused by breast cancer is currently the second leading cause of death for women in developed regions. Since 2008, the support of clinical approaches that decreases the risk and increase early detection and treatment through worldwide, while frequency increases in 20% and rise in mortality by 14%. When incorporating nuclear medicine technology in the diagnostic plan, it can readily achievable by multimodality approach by using tailored management. In instruments the developments has been taking place such as high resolution dedicated breast device and it has coupled with the diagnostic versatility of conventional cameras have reinserted nuclear medicine as a valuable tool in clinical setting in broader range. This shows general nuclear medicine is a critical role in the care of women with breast cancer including detection and stratification, guiding treatment by using sentinel node imaging (which is used to determine whether cancer has spread beyond a primary tumor into your lymphatic system), monitoring cardiotoxicity from therapeutic regimens and evaluating local and global progression. In addition with that, it describes the role of positron emission tomography or computed tomography is discussed.

### **Scintimammography :**

Scintimammography in nuclear medicine, which utilizes a wide range of instrumentation applications . Especially in recent years, conventional planar scintimammography has been enhanced by single-photon has been enhanced by single-photon emission computed tomography (SPECT) and hybrid SPECT/CT . While hybrid SPECT/CT adds clinical benefit by combining physiologic and anatomical data to facilitate non-palpable lesion biopsies, radiotherapy planning, and treatment monitoring. Breast-specific gamma imaging systems, which are specialised to a limited field of view (FOV), have also become popular internationally. Planar orientations (anterior, lateral, oblique) are obtained within 5-15 minutes after injection for both BSGI and scintimammography. The patient is in a prone position with pendent breasts for oblique and lateral pictures, whereas supine stance is supported for oblique, anterior, and SPECT tomography acquisitions. Image acquisition is preferred in both locations. By reducing photon scattering and boosting imaging contrast, prone placement better separates breast tissue from high pharmacological uptake in the heart and liver, allowing for improved observation of breast activity.

Additionally, the prone posture has advantages such as higher spatial resolution, improved evaluation of the chest wall, and better delineation of the breast contour. Supine position allows for better visibility of the main lesion and internal mammary. In the craniocaudal and mediolateral oblique orientations, BSGI obtains 210 micro pictures per breast. Breast lesion size and palpability are extremely important in the diagnosis and prognosis of patients, as small non-palpable lesions suggest early disease. Mammographically non-palpable benign lesions that are metabolically benign could be classified as such, avoiding the need for a laborious biopsy procedure and instead allowing for clinical surveillance. Because of scintimammography's great specificity, a positive scintigraphic finding would support a recommendation for an invasive assessment.

When compared to scintimammography, which had overall sensitivity and specificity of 82 percent and 85 percent, respectively, and showed no improvement over SPECT, which had 86 and 87 percent respectively, multiple studies suggested that dedicated combined with breast positioning during BSGI provided better detection of sub-centimeter and non-palpable lesions. BSGI's unique camera design is highly sensitive for detecting local disease, but it has limitations in the broader clinical situation when compared to planar, SPECT, and SPECT/CT cameras, which can study regional, axial, and global disease.

### **Sentinel Lymph Node Scintigraphy:**

Axial nodal bed status is the most important prognostic factor for newly diagnosed patients with invasive breast cancer, and it's also important for deciding on treatment options. Because imaging methods for axillary staging are rarely sensitive or specific, surgical exploration of nodal involvement is required. Sentinel lymph node status through biopsy (SLNB) was developed as a less intrusive alternative to established staging approaches that often increase morbidity, such as axillary lymph node dissection (ALND). A histopathologically negative sentinel node means that the ipsilateral nodal bed is clear of metastatic illness because it is the first relay receiving lymphatic outflow straight from the tumour. The cost and comorbidities associated with preserving the healthy nodal bed are reduced. The cost and common comorbidities associated with ALND are both reduced when the healthy nodal bed is spared. Negative sentinel node status is related with a 0–2% axillary recurrence rate.

The use of radiolabelled colloids in nuclear medicine planar (dynamic or static) and/or SPECT/CT sentinel node imaging provides surgeons with a visual map to enable correct localization of sentinel nodes and unusual drainage patterns. The identification of the sentinel node is critical to the success of SLNB, and preoperative sentinel node imaging is well adapted for this task, with a detection rate of 94 percent to 100 percent. Sentinel node imaging with SPECT/CT of all tumours may give a more reliable technique to locate and biopsy sentinel nodes for staging in patients with multicentric and multifocal illness whose lymphatic drainage patterns may differ.

Particle size and dose concentration can alter lymphatic transit of radiocolloid particles following injection. Large colloid particles' lymphatic movement is often hampered by their size, resulting in either delayed or non-visualization of axillary sentinel nodes. Small particles, on the other hand, migrate quickly and may not be trapped in the sentinel node. So that lymphatic physiology is adequately depicted, solvent volumes should be adjusted to injection technique. Smaller quantities are better for peritumoural injections, for example, because higher pressure can promote leakage into the extravascular space, which can then migrate to nearby lymphatic channels.

Preoperative sentinel node mapping is consistently provided by planar acquisitions (Fig. 1), and the addition of multimodality SPECT/CT further strengthens this role. Oriented SPECT/CT slices allow surgeons to precisely anatomically localise nodal uptake, decreasing

surgical time and enhancing biopsy accuracy (Fig. 2). Evidence also suggests that SPECT/CT can detect difficult-to-interpret drainage patterns and sentinel nodes (such as parasternal nodes) that aren't seen on planar imaging, limiting blind exploratory surgery. For precise visualisation of lymphatic drainage patterns and labelling sentinel nodes, patient positioning and acquisition settings are critical. Patient photos, patient markers, and an auditory gamma probe all work together to help surgeons identify the radioactive sentinel node or nodes for excision during surgery.

### **Tc-99m Multigated Radionuclide Angiography:**

The use of combination chemotherapy has been well recognised in cancer research to cause cardiac problems, necessitating routine monitoring of left ventricular ejection fraction. Anthracyclines and the monoclonal antibody trastuzumab are two anti-cancer medications with well-known cardiotoxicity among the constantly expanding array of anti-cancer therapies used in breast cancer treatment. Anthracycline treatment causes cumulative dose-dependent cardiotoxicity in myocytes, which can lead to congestive heart failure and even death. When used in combination with anthracyclines, trastuzumab can cause severe cardiotoxicity. Furthermore, pericardial irritants such as radiation therapy and chemotherapy medications enhance the likelihood of pericardial effusion problems, making early diagnosis crucial to reducing the primary or contributing related mortality found in 86 percent of symptomatic cancer patients.

Coronary artery disease, valve disease, constructive pericarditis, and myocardial dysfunction, including congestive heart failure, are all examples of radiation-induced heart disease that can severely affect a patient's quality of life. As a result, determining the amount of a patient's risk of a cardiac episode is critical. Obtaining ejection fraction measures prior to and after chemotherapy treatment in breast cancer patients is the usual way for detecting cardiotoxicity and pericardial effusions. Serial examination allows clinicians to track a patient's cardiac response to treatment, lowering the likelihood of comorbidities caused by chemotherapy.

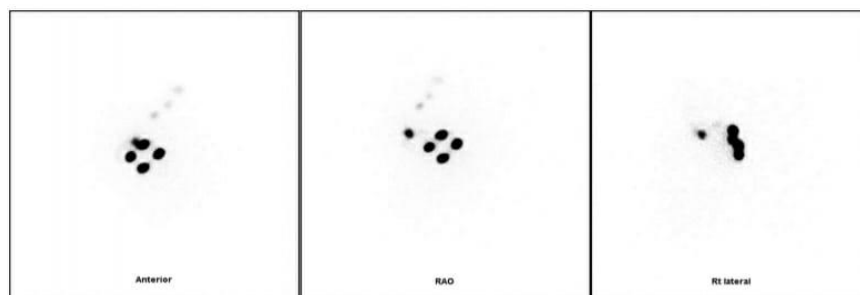


Figure 1. Sentinel node localisation study of the right breast showing four periareolar radiopharmaceutical injections oriented on anterior, right anterior oblique (RAO) and right lateral planar images. An intense focal uptake lateral to the injection sites is identified as well as faint uptake in three axillary lymphatic chain nodes. Image courtesy of Regional Imaging, a member of I-MED Network Radiology, Wagga Wagga, NSW.

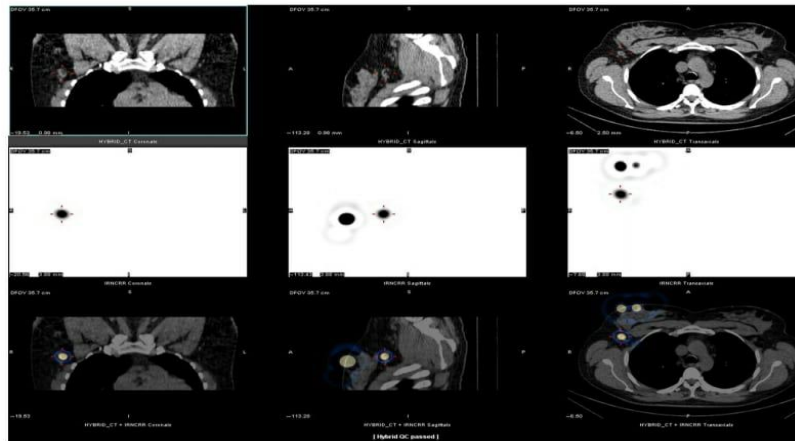


Figure 2. Fused SPECT/CT images provide surgeons with a more accurate anatomical visualisation of the sentinel node with regard to the breast and axilla by depicting the precise location in the axillary area adjacent to the first and second rib saving valuable intraoperative time and increasing surgical confidence. SPECT, single-photon emission computed tomography. Image courtesy of Regional Imaging, a member of I-MED Network Radiology, Wagga Wagga, NSW

In patients undergoing chemotherapy, multigated radionuclide angiography (RNA) or equilibrium radionuclide angiocardiology (ERNA) is considered the gold standard for measuring heart function with good repeatability and low inter-observer variability. RNA is a noninvasive technology that measures regional and global wall motion, ventricular systolic and diastolic function (both right and left ventricular ejection fractions), and ventricular volumes using Tc-99m pertechnetate erythrocyte labelling. In vivo, in vitro, or modified in vivo procedures are used to label red blood cells with Tc-99m pertechnetate, as documented in the literature. The acceptable level of picture quality, patient throughput needs, and technical staff knowledge will all influence the choice of a blood pool agent in any given clinical circumstance.

Early diagnosis of cardiotoxic abnormalities on serial imaging allows for prompt intervention, reducing the risk of related patient morbidity or mortality from trastuzumab and anthracycline therapy. The use of RNA to assess global LV systolic function and diastolic performance indexes has been shown to be effective for early detection of functional changes after chemotherapy when compared to baseline, highlighting the importance of serial imaging at all stages of treatment to assess patient prognosis. A decrease in peak fill rate as measured by multigated RNA implies poor diastolic function, which occurs before systolic function declines in anthracycline-induced cardiotoxicity and, more critically, is an early indicator of compromised heart function. Pericardial effusions in small or moderate amounts may not impair LVEF, although they can cause comorbidities such as difficulty breathing and caused chest discomfort.

## Positron Emission Tomography/ Computed Tomography :

Oncologic investigations accounted for 94 percent of the projected 1.5 million PET/CT procedures performed in the United States in 2011. F-18 fluoro-2-deoxyglucose (F-18 FDG), a tumor-avid glucose analogue, has acquired general acceptance as a marker of cellular metabolism, providing molecular insight into cancer physiology. PET/ability CT's to precisely stage WB metastatic disease as well as quantify and evaluate therapy response has had a huge impact on how breast cancer patients are treated.

Because the method may not sufficiently resolve small primary breast tumours or axillary nodal areas, a PET/ CT with high sensitivity and specificity is optimal for detecting distant metastases. Dedicated positron emission mammography (PEM) devices were created to increase detection of small primary lesions by maximising system spatial resolution, and are currently being investigated to address this problem. PET/CT is comparable to and may surpass whole body bone scan (WBBS) in the diagnosis of breast cancer metastases to bone, with similar sensitivity but higher specificity due to superior metabolic and morphologic characterization of osseous lesions.

In many treatment contexts, F-18 FDG can also give an earlier and more objective prediction and assessment of response. Serial standard uptake value (SUV) measurements are produced directly from tumour metabolic avidity using a standardised technique, and so provide a semi-quantitative depiction of cancer physiology across time. A greater lesion SUVmax, including values acquired from osseous lesions, also has a favourable connection with prognostically poor aggressive illness. PET/CT has also been used to investigate new positron-emitting radiopharmaceuticals that target oestrogen, progesterone, oestrogen growth factor, and somatostatin receptors found on breast tumours to improve specificity.

Receptor-mediated molecular imaging aids tumour characterization, customised therapeutic research, disease burden extent, and disease monitoring through serial imaging.

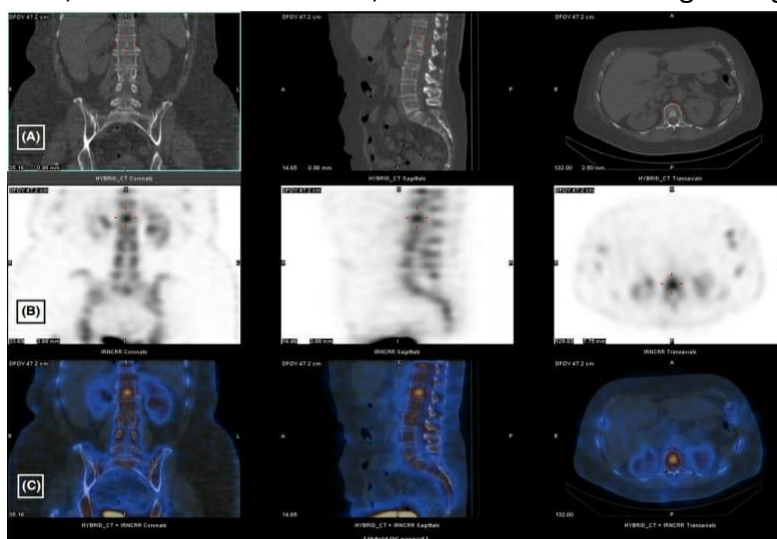


Figure: 3 SPECT/CT images of the same patient revealing sclerosis in the vertebral body of L1 on CT images (A) which correspond to an area of increased focal uptake on SPECT (B) and fused SPECT/CT (C) most consistent with metastatic disease. This lesion was not apparent on WBBS. Additionally, degenerative changes (which affect intervertebral discs or facet joints) and metastatic disease (which primarily affect the vertebral body) are both common pathologies occurring in the lumbar region and thus require anatomic information for an accurate diagnosis. These images clearly highlight the significance of hybrid imaging in evaluating vertebral metastatic involvement and increasing diagnostic confidence. SPECT, single-photon emission computed tomography; WBBS, whole body bone scan. Image courtesy of Regional Imaging, a member of I-MED Network Radiology, Wagga Wagga, NSW

## **Conclusion :**

Nuclear medicine is a vital component of illness diagnosis, therapy, and prognosis because it fills a physiologic gap in medical imaging for cancer patients, delivering information that is representational of function.

General nuclear medicine, in particular for breast cancer care, has diagnostic value at every stage of the disease and is considered the first-line modality in a variety of clinical settings. The adaptability of gamma cameras with SPECT and/or SPECT/CT is an appropriate tool for patients in a broader clinical setting, providing a very detailed view into the current physiologic state of bodily structures and their functions. PET/CT has established itself as a powerful oncologic tool, and with the introduction of tailored radiopharmaceuticals, it is expected to see a significant increase in its use in the treatment of breast cancer.

Future research should look at multi-acquisition (planar and SPECT) and multi-modality (SPECT/CT) imaging in the full scope of physiologic breast cancer imaging to better understand the added utility of instrument hybridisation methods and processes in nuclear medicine imaging.

## **References:**

1. International Agency for Research on Cancer. Breast cancer estimated incidence, mortality and prevalence worldwide in 2012. World Health Organization [updated 2012]. Available from: [http://www.globocan.iarc.fr/Pages/fact\\_sheets\\_cancer.aspx](http://www.globocan.iarc.fr/Pages/fact_sheets_cancer.aspx) (accessed 8 January 2015).
2. Shah R, Rosso K, Nathanson SD. Pathogenesis, prevention, diagnosis and treatment of breast cancer. *World J Clin Oncol* 2014; 5: 283–98.
3. Brem R, Rechtman L. Nuclear medicine imaging of the breast: a novel, physiological approach to breast cancer detection and diagnosis. *Radiol Clin North Am* 2010; 48: 1055–74.
4. Munnink TH, Nagengast WB, Brouwers AH, et al. Molecular imaging of breast cancer. *Breast* 2009; 18: S66–73.
5. Ferrara A. Nuclear imaging in breast cancer. *Radiol Technol* 2010; 81: 233–46.



6. Tiling R, Kebler M, Untch M, Sommer H, Linke R, Hahn K. Initial evaluation of breast cancer using Tc-99m sestamibi scintimammography. *Eur J Radiol* 2005; 53: 206–12.
7. de Cesare A, Giuseppe DV, Stefano G, et al. Single photon emission computed tomography (SPECT) with Technetium-99m sestamibi in the diagnosis of small breast cancer and axillary lymph node involvement. *World J Surg* 2011; 35: 2668–72.
8. Mettler F, Guiberteau M. *Essentials of Nuclear Medicine Imaging*, 6th edn. Mosby Elsevier, St. Louis, MO, 2012.
9. Specht JM, Mankoff DA. Advances in molecular imaging for breast cancer detection and characterization. *Breast Cancer Res* 2012; 14: 206–17.
10. Hendrick RE. Radiation doses and cancer risks from breast imaging studies. *Radiology* 2010; 257: 246–53.
11. Taillefer R. Clinical applications of 99mTc-sestamibi scintimammography. *Semin Nucl Med* 2005; 35: 100–15.
12. Jacobsson H. Single-photon-emission computed tomography (SPECT) with 99mTechnetium sestamibi in the diagnosis of small breast cancer and axillary node involvement. *World J Surg* 2011; 35: 2673–4.
13. Kong FL, Kim E, Yang D. Targeted nuclear imaging of breast cancer: status of radiotracer development and clinical applications. *Cancer Biother Radiopharm* 2012; 27: 105–12.
14. Lee J, Rosen E, Mankoff D. The role of radiotracer imaging in the diagnosis and management of patients with breast cancer: part 1 – overview, detection and staging. *J Nucl Med* 2009; 50: 569–81.
15. Nguyen BD, Roarke MC, Karstaedt PJ, Ingui CJ, Ram PC. Practical applications of nuclear medicine in imaging breast cancer. *Curr Probl Diagn Radiol* 2009; 38: 68–83.
16. Kim SJ, Kim IJ, Bae YT, Kim YK, Kim DS. Comparison of quantitative and visual analysis of Tc-99m MIBI scintimammography for detection of primary breast cancer. *Eur J Radiol* 2005; 53: 192–8.
17. Prekeges J. Breast imaging devices for nuclear medicine. *J Nucl Med Technol* 2012; 40: 71–8.
18. Waxman A. The role of 99mTc methoxyisobutylisonitrile in imaging breast cancer. *Semin Nucl Med* 1997; 27: 40–54.
19. Silvera S, Rohan T. Benign proliferative epithelial disorders of the breast: a review of the epidemiologic evidence. *Breast Cancer Res Treat* 2008; 110: 397–409.
20. Sergieva S, Alexandrova E, Baitchev G, Parvanova V. SPECT-CT in breast cancer. *Arch Oncol* 2012; 20: 127–31.
21. Whitman G, Strom E. Workup and staging of locally advanced breast cancer. *Semin Radiat Oncol* 2009; 19: 211–21.
22. Glendenning J, Cook G. Imaging breast cancer bone metastases: current status and future directions. *Semin Nucl Med* 2013; 317–23.
23. Iqbal B, Currie G, Wheat J, Raza H, Ahmed B, Kiat H. Incremental value of SPECT/CT in characterizing solitary spine lesions. *J Nucl Med Technol* 2011; 39: 201–7.
24. Fink C, Hasan B, Deleu S, Pallis A, Baas P, O'Brien M. High prevalence of osteoblastic bone reaction in computed tomography scans of an European Organisation for Research and Treatment of Cancer prospective randomised phase II trial in extensive stage small cell lung cancer. *Eur J Cancer* 2012; 48: 3157–60.
25. IMVinfo.com. [homepage on the Internet]. IMV 2012 PET imaging market summary report [updated 2012 Aug 7]. Available from: <http://www.imvinfo>.



- com/user/documents/content\_documents/def\_dis/2012\_08\_07\_13\_51\_48\_43\_IMV\_PET2012\_report\_datasheet.pdf (accessed 7 January 2015).
26. Gallamini A, Zwarthoed C, Borra A. Positron emission tomography (PET) in oncology. *Cancers* 2014; 6: 1821–89.
  27. Bourgeois A, Warren L, Chang T, Embry S, Hudson K, Bradley Y. Role of positron emission tomography/ computed tomography in breast cancer. *Radiol Clin North Am* 2013; 51: 781–98.
  28. Hong S, Li J, Wang S. 18FDG PET-CT for diagnosis of distant metastases in breast cancer patients. A meta-analysis. *Surg Oncol* 2013; 22: 139–43.
  29. Bensch F, van Kruchten M, Lamberts L, Schroder C, Hospers G, Brouwers A, van Vugt M, de Vries E. Molecular imaging for monitoring treatment response in breast cancer patients. *Eur J Pharmacol* 2013; 171: 2–11. *Nuclear Medicine and Breast Cancer* L. R. Greene et al. 64<sup>a</sup> 2015 The Authors. *Journal of Medical Radiation Sciences* published by Wiley Publishing Asia Pty Ltd on behalf of Australian Institute of Radiography and New Zealand Institute of Medical Radiation Technology
  30. Kalles V, Zografos GC, Provatopoulou X, Koulocheri D, Gounaris A. The current status of positron emission mammography in breast cancer diagnosis. *Breast Cancer* 2013; 20: 123–30.