

6b

Lymphoscintigraphy as the standard of imaging for lymphatic mapping in melanoma and breast cancer

S. Vidal-Sicart (Barcelona)

A renewed interest has been posed on lymphoscintigraphy since its widespread use in the sentinel node (SN) localization technique. Its main indications are melanoma and breast cancer, although it is used in other type of tumours.

An essential prerequisite for a successful SN biopsy is an accurate map of the lymphatic drainage pattern from the tumour site. The role of lymphoscintigraphy is to provide such a map in every patient. Lymphoscintigraphy indicates not only the location of sentinel nodes but also the number of SNs at each location. This mapping is especially important in malignant melanoma, since lymphoscintigraphy can identify the lymphatic vessels draining to the SNs.

Reliable clinical prediction of lymphatic drainage from the skin or breast is not possible. Patterns of lymphatic drainage from the skin are highly variable from patient to patient, even from the same area of the skin.

In breast cancer, lymphoscintigraphy permits accurately detection of SNs outside the axilla, which occurs in about 40 % of cases. These nodes can be visualized in the internal mammary chain, the supra-clavicular region, the intrapectoral region as well as intramammary location. The location of the breast tumour is not a reliable guide to lymphatic drainage, since lymph flow often crosses the center line of the breast.

There are some aspects to take in account in order to reach the best quality in lymphoscintigraphic images. In summary, radiopharmaceutical, site and route of administration and gammacamera positioning and imaging are important issues.

Radiopharmaceuticals

There are numerous radiopharmaceuticals for SN imaging (suitably labelled with ^{99m}Tc). These include dextran, human serum albumin and colloids. Antimony colloid is largely used in Australia, sulfur colloid (filtered or unfiltered) is mostly used in North America and nanocolloidal albumin is widely used in Europe. Their main differences are based on particle size (ranging from 3 to 600 nm), the speed of migration from injection site and the visualization of lymphatic channels and one or more lymphatic nodes. Then, the lesser particle size the faster radiotracer migration to lymph nodes. The particle concentration and the dosage of radiopharmaceutical are important too.

Tracer administration

The injection of the tracer has evolved since the beginning of SN procedure. In the first procedures (melanoma, palpable breast) the injection was performed by the nuclear medicine physician. However, as the technique evolved the injection site can be guided by ultrasound and stereotaxis in non-palpable breast cancers.

In melanoma patients the tracer is injected intradermally, raising a wheal on the skin, around the primary lesion or biopsy scar. The best approach is performing the injection with a 5 mm distance from the lesion in order to gain the best drainage reproducibility.

In breast cancer there is no consensus about the best way to inject the radiotracer. Thus, intradermal, subdermal, periareolar, peritumoral, subtumoral and intratumoral routes have been described. We can categorize them in to groups, skin-related (intradermal, subdermal, periareolar) injections and tumor-related (all the others) injections. All of them get a suitable SN identification and localization in the axilla. However, there is a remarkable variability between them as skin-related injections almost always

depict an axillary node and on the other hand, the tumor-related injections show a significant percentage of extraaxillary drainage.

Gammacamera positioning

Lymphoscintigraphy is mandatory in SN procedure for many reasons:

- To ascertain the lymph node basins at risk for metastatic spread
- To indicate the location and number of sentinel nodes
- To distinguish the sentinel nodes and non-sentinel nodes
- To localize the sentinel nodes in aberrant, in transit or unpredictable locations
- To mark the sentinel node position on the skin

Lymphoscintigraphy has to be performed sequentially.

Dynamic scintigraphy: Normally we use a 30 sec images during the first 10-15 min after injection. This approach facilitates the identification of lymphatic ducts from a lesion with a rapid drainage expected.

Static images: They last at least 180 s in both anterior and lateral projections. Oblique and other special views can be performed in order to clarify the SN location. The time scheduled to obtain these images are 20-30 min, and 2 hours for cutaneous lymphoscintigraphy and until 4 hour or more when a slow migration of tracer is expected.

The body contour can be outlined with a ^{57}Co or $^{99\text{m}}\text{Tc}$ marker or, preferably, with a flood source of ^{57}Co or $^{99\text{m}}\text{Tc}$. The aim of these procedures is to mark the SN location on the skin in the same manner that patient will be subsequently operated.

Clinical results in melanoma and breast cancer

After an adequate lymphoscintigraphic acquisition and skin marking, the surgical SN retrieval is successfully accomplished in the majority of cases. However there are different aspects depending on the tumour to be treated.

Thus, in melanoma patients lymphoscintigraphic SN visualization and surgical harvesting reaches almost 100 % of cases. Important aspects in melanoma are the aberrant or intransit SNs, the unpredictability of some anatomical regions (trunk, head & neck) and, specially, the false negative rate (i. e. the presence of a positive lymph node in a patient with a negative SN).

In breast cancer patients, the identification of SN in lymphoscintigraphy ranges from 85-97 % depending on the way of administration. Important aspects in this clinical scenario are the extraaxillary SN identification (inner mammary chain, supraclavicular, intramammary nodes) that represents as much as 35 % of cases (with intratumoral injection). The false negative rate is currently below 5 % in the majority of studies.

References

1. Tanis PJ, Nieweg OE, Valdés Olmos RA, Kroon BBR. Anatomy and physiology of lymphatic drainage of the breast from the perspective of sentinel node biopsy. *J Am Coll Surg* 2001; 192: 399-409
2. Mariani G, Moresco L, Viale G, Villa G, Bagnasco M, Canavese G et al. Radioguided sentinel lymph node biopsy in breast cancer surgery. *J Nucl Med* 2001; 42: 1198-1215.
3. Uren RF, Howman-Giles R, Thompson JF. Patterns of lymphatic drainage from the skin in patients with melanoma. *J Nucl Med* 2003; 44: 570-582.
4. Valdés-Olmos RA, Tanis PJ, Hoefnagel CA, Nieweg OE, Muller SH, Rutgers EJT. Improved sentinel node visualization in breast cancer by optimizing the colloid particle concentration and tracer dosage. *Nucl Med Commun* 2001; 22: 579-586.
5. Vidal-Sicart S, Pons F, Fuertes S, Vilalta A, Rull R, Puig S, Palou JM, Ortega M, Castel T. Is the identification of in-transit sentinel lymph nodes in malignant melanoma patients really necessary? *Eur J Nucl Med Mol Imaging* 2004; 31: 945-949.