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Clinically unexpected lymph node: An old enemy to remember in the cancer diagnosis and treatment

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Introduction

Tuberculosis is a common infection especially in the developing countries and it can be detected coincidentally on imaging done for other purposes, such as cancer detection and staging. Treatment of tuberculosis is challenging among cancer patients, due to drug interactions caused by anti-tuberculosis treatment.

Objectives

To describe a patient with tuberculosis lymphadenitis detected incidentally on PET-CT during a staging of newly diagnosed early stage breast cancer and its effect on our treatment decision.

Case report

A 53-year old female patient admitted to clinic with a mass on her right breast. Her past medical history was unremarkable. The breast ultrasound revealed a 29x11 mm solid lesion located 4 cm away from nipple on 7 o'clock position and there was no axillary lymphadenopathy.

A tru-cut biopsy was performed and showed grade 2 invasive ductal carcinoma. A staging PET-CT scan revealed a right supraclavicular lymph node in size of 11x10 mm (SUVm:7.9) and there was also a nodular lesion measured 7x8 mm on the thyroid gland. There was no axillary lymph node involvement (Figure 1).

In the absence of axillary lymph node involvement, the thyroid nodule and cervical lymph nodes made us concern about a second primary malignancy, most probably a thyroid neoplasm. Thyroid fine needle aspiration biopsy showed normal thyroid glands and tru-cut biopsy of the right supraclavicular lymph node showed granulomatous lymphadenitis which is consistent with tuberculosis lymphadenitis (Figure 2A). The PPD test measured 15 mm and serum quantiferon were positive. The patient was consulted to the infectious disease department and an initial intensive-phase therapy with four antibiotics (INH, RIF, PZA, EMB) and continuation-phase therapy with 2 antibiotics (INH, RFA) planned.

The patient underwent lumpectomy and axillary lymph node dissection by surgical team. The pathology of the tumor revealed, 2.8 cm invasive neoplasm with intact surgical margins. The axillary lymph nodes were tumor-free. The tumor showed 80% ER, PR positivity, cErbB2 was negative and Ki-67 proliferation index was %1 (Figure 2B).

Adjuvant endocrine therapy with tamoxifene was planned for perimenopausal early stage breast cancer. However, the usage of rifampin for tuberculosis treatment concerned us about drug

interactions between tamoxifene and rifampin. We started letrozole, an aromatase inhibitor and has no known drug interaction with rifampin, to our patient.

Anti-tuberculosis treatment completed six months after. Ten months after letrozole treatment we made another PET-CT scan and it showed persisting of the thyroid nodule and complete regression of cervical lymphadenopathies (Figure 1B). There were no recurrence findings on examination and imaging.

Discussion

Breast cancer is the most common cancer among women worldwide [1]. It is very unusual to detect distant lymph node metastases of breast

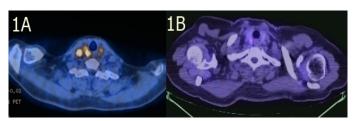


Figure 1. A is showing right cervical lymph node and thyroid involvement seen on PET-CTdetected initially. **B** is showing the loss of FDG involvement in the right cervical lymph node in follow-up.

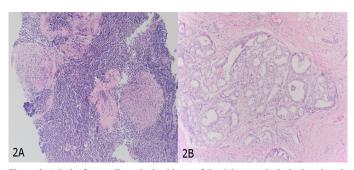


Figure 2. A is the fine needle aspiration biopsy of the right supraclavicular lymph node showing granulomatosis lymphadenitis **B** is the mastectomy material showing invasive ductal carcinoma of the breast.

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cancer without involvement of axillary lymph nodes [2]. In such cases, clinicians should concern about another reason of lymphadenopathy other than breast cancer metastasis. The tuberculosis might be an important cause of lymphadenitis misdiagnosed as cancer metastasis, so patient's risk factors should be considered especially in endemic areas. There would be a devastating result if we assume cervical lymph nodes as metastasis and start chemotherapy without confirmation.

Another challenge is anti-tuberculosis treatment in cancer patients due to both side effects and drug interactions. Tamoxifene, a selective estrogen receptor modulator drug, is frequently used for treatment of breast cancer. However, tamoxifen is metabolized by the CYP2D6 enzyme, so that is greatly influenced by cytochrome induction. Rifampin is a main drug in the treatment of tuberculosis and is a very potent inductor of the cytochrome p450 levels. Binkhorst et al. compared serum levels of tamoxifene and its active metabolite, endoxifene, in four patients tamoxifen administration alone and with rifampin. The results showed that rifampin administration declined serum tamoxifene leves in all patients ranging by %28 to 85. The study strongly indicated that concurrent use of rifampin and tamoxifene should be avoided [3] .

Evaluation of treatment response among patients with tuberculosis lymphadenitis remains challenging. Active tuberculomas have increased macrophage and neutrophil activity. Therefore, 18F-FDG PET can be used in diagnosis, detection and most importantly treatment response of tuberculosis. Various studies showed the promising role of using 18F-FDG PET in the disease course [4]. In our case, the FDG involvement on the cervical region lost after six-month treatment of the tuberculosis and directed us about treatment response.

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