False positive $^{18}$F-FDG PET/CT due to active varicella zoster infection in a Hodgkin’s lymphoma patient

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ABSTRACT

We report a case of Hodgkin lymphoma (classic type) referred for response assessment after two cycles of chemotherapy with ABVD regimen. The F-18 fluorodeoxyglucose positron emission tomography (F-18 FDG PET/CT) showed hypermetabolic cutaneous and subcutaneous lesions with a linear pattern in the left arm with significant F-18 fluorodeoxyglucose positron accumulation in associate with left axillary hypermetabolic lymph nodes. She presented with left arm pruritic rash accomplished by pain from two weeks ago. On clinical examination, painful papulovesicular rash with palpable enlarged axillary lymph node were noted. These findings were compatible with cutaneous herpes zoster infection of the left arm along with axillary reactive lymphadenopathy.

Key words: Active herpes infection; Hodgkin’s lymphoma; Clinical examination; Nuclear medicine

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**INTRODUCTION**

Potential diagnostic errors due to active herpes infection and its associated adenopathy have been reported [1-3]. Herein we report a case of herpes zoster and underlining the possibility of an active viral infection in an immunocompromised patient which may mimic recurrent disease. The importance of awareness of opportunistic infections and precise attention to the patient history and clinical assessment is emphasized (Figure 1).

**CASE PRESENTATION**

A 31 years old lady with a history of classic type Hodgkin’s lymphoma without unfavorable feature, diagnosed with a prevascular anterior mediastinal mass since seven months, was referred to our department for response assessment. She has received two cycles of ABVD regiment without mediastinal radiotherapy, four weeks after the last course of ABVD mid-treatment $^{18}$F-FDG PET/CT was done. $^{18}$F-FDG PET/CT scan revealed hypermetabolic axillary nodes with fatty hilum as well as a subtle linear hypermetabolic cutaneous uptake in the upper left arm in MIP images with no other pathologic $^{18}$F-FDG avid lesion throughout the body. Clinically, erythematous vesicles in the left cervical innervation pathognomonic of herpes zoster infection with enlarged axillary lymph nodes were detected. Hypermetabolic axillary lymph nodes were considered due to a reactive response to viral infection. (Figure 2) Complete response to therapy was reported. Upon clinical examination, the reason of axillary lymphadenopathy and hypermetabolism was obviously due to varicella zoster infection and tissue biopsy was not done. Subsequently, sixth months following antiviral treatment, axillary lymphadenopathy disappeared upon sonographic correlation with no cutaneous lesions. After completion of chemotherapy, the patient was in disease-free state without any evidence of recurrent disease in long term clinical and conventional imaging follow up.

**DISCUSSION**

Negative interim metabolic PET/CT scan results, predict complete remission at the end of treatment and even works better than other prognostic model such as NCCN-IPI (international prognostic Score) [4, 5]. Due to high negative predictive value (NPV) in the early stage of Hodgkin’s lymphoma, negative findings can lead to decrease courses of radiotherapy especially in the good responder patients [6, 7].
According to the literature, the main explained mechanism for 18F-FDG uptake related to high glycolytic activity and overexpressing specific membrane glucose transporter compared to the normal cells. Because 18F-FDG scanning is not a tumor-specific tracer, false positive findings are frequently encountered, atypical physiologic site and benign pathologic lesions such as: infectious, autoimmune process, granulomatous inflammation (tuberculosis, sarcoidosis) [8], recent immunization status [9], postsurgical inflammatory tissue, inflammatory reactions after radiotherapy, nonspecific reactive changes associated with treatment as well as treatment-related change like thymic hyperplasia, Doxil-related condition [10], bleomycin-induced ILD(interstitial lung disease) [11], flare phenomenon, osteonecrosis and insufficiency fracture [12, 13]. Even, technical errors can be a potential cause of misleading interpretation such as suboptimal injection with 18F-FDG uptake in the regional nodes, misregistration artifact and dense materials that may artefactually increase 18F-FDG uptake [14]. It is even reported that within the neoplastic masses, 24% of 18F-FDG uptake was related to the granulation tissue and activated macrophage [15].

To minimize, possible non-tumoral 18F-FDG avid foci the time interval between chemotherapy and scanning should be the least 3-week window [16], through attention to history, clinical examination and observation should be part of nuclear medicine physicians’ responsibilities especially in an immunocompromised patient. Suspected lesions should be further evaluated by delayed images, reviewing non-attenuation correlated images as well as using diagnostic CT or (magnetic resonance imaging) MRI providing anatomical details. Using a consistent guideline for reporting 18F-FDG-PET can also decrease the false positive results as well as a continuous learning activities of reporting physicians [17].

REFERENCES