



Increased Bone Marrow ¹⁸F-Choline Uptake in a Patient with Hepatocellular Carcinoma and Thalassemia Intermedia

Hepatosellüler Karsinom ve Talasemi Intermedia Tanılı Bir Hastada Artmış Kemik İliği ¹⁸F-Kolin Tutulumu

© Luca Filippi

Santa Maria Goretti Hospital, Clinic of Nuclear Medicine, Latina, Italy

Abstract

A 57-year-old male with history of thalassemia intermedia and hepatocellular carcinoma underwent a positron emission tomography/computed tomography (PET/CT) scan with ¹⁸F-choline before radioembolization procedure with ⁹⁰Y-microspheres. The PET/CT scan with ¹⁸F-choline demonstrated highly increased tracer incorporation within a gross lesion in the hepatic dome coupled with diffuse activity in bone marrow, this latter aspect was probably due to the compensatory hematopoiesis stimulation induced by chronic hemolysis. This pattern of skeletal ¹⁸F-choline uptake should be considered as a peculiar PET/CT finding in thalassemic patients.

Keywords: ¹⁸F-choline, positron emission tomography/computed tomography, hepatocellular carcinoma, thalassemia

Öz

Talasemi intermedia ve hepatosellüler karsinom tanılı bir hastada ⁹⁰Y-mikrosfer ile radyoembolizasyon prosedürü öncesi ¹⁸F-kolin ile pozitron emisyon tomografisi/bilgisayarlı tomografi (PET/BT) taraması yapıldı. ¹⁸F-kolin ile PET/BT taraması karaciğer kubbesindeki büyük bir lezyon içerisinde artmış tracer tutulumu ve muhtemelen kronik hemoliz ile indüklenen kompensatuvar hematopoezise kemik iliğinde artmış aktivite gösterdi. Bu skeletal patterned ¹⁸F-kolin tutulumu talasemik hastalara özgü bir PET/BT bulgusu olarak değerlendirilmelidir.

Anahtar kelimeler: ¹⁸F-kolin, pozitron emisyon tomografisi/bilgisayarlı tomografi, hepatosellüler karsinom, talasemi

Address for Correspondence: Luca Filippi MD, Santa Maria Goretti Hospital, Clinic of Nuclear Medicine, Latina, Italy

Phone: +393921247921 **E-mail:** l.filippi@ausl.latina.it **ORCID ID:** orcid.org/0000-0003-4423-5496

Received: 09.08.2019 **Accepted:** 30.09.2019

©Copyright 2020 by Turkish Society of Nuclear Medicine
Molecular Imaging and Radionuclide Therapy published by Galenos Yayınevi.

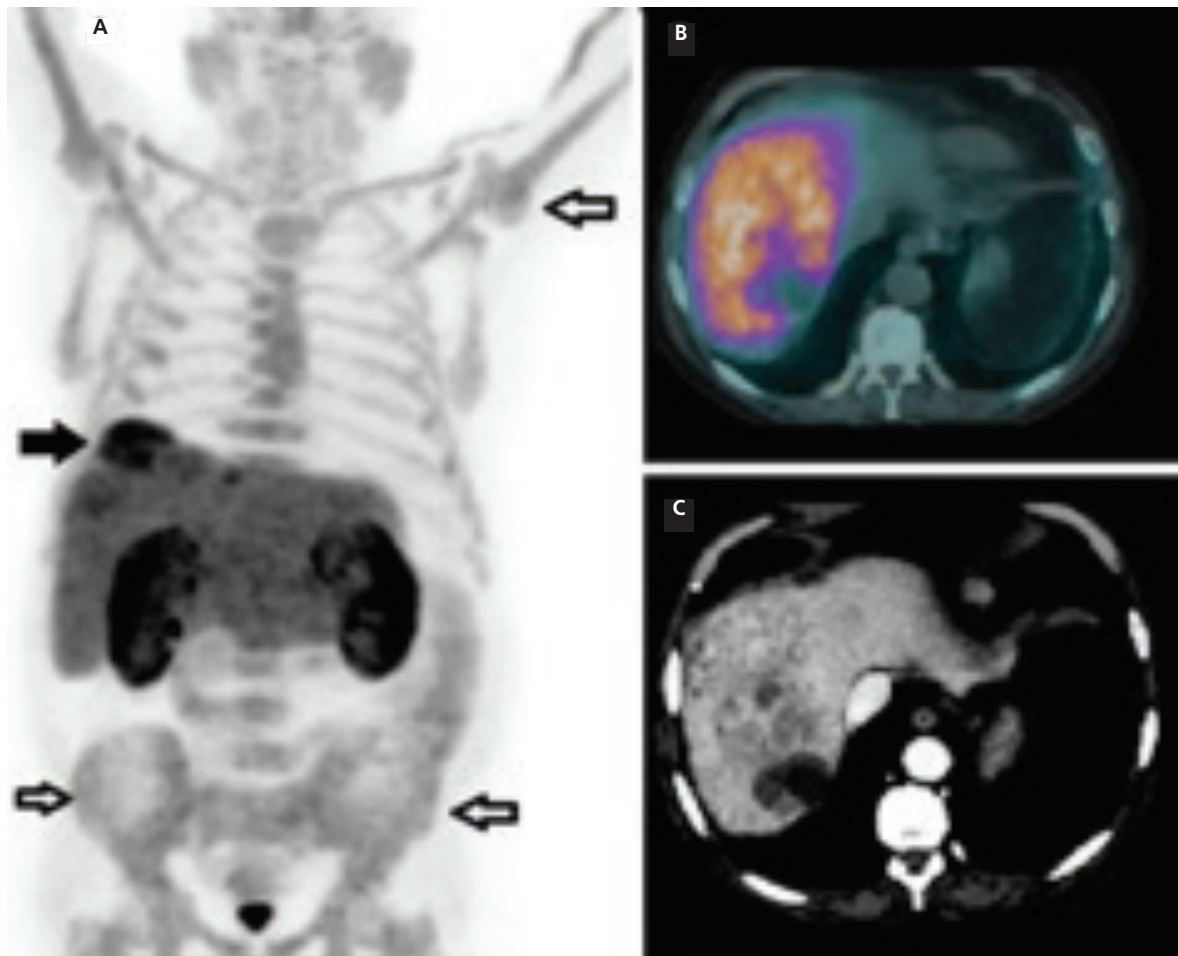


Figure 1. A 57-year-old man was diagnosed as having thalassemia intermedia at the age of 3 years (genotype CD39/IVS 1-6). He received sporadic blood transfusions since childhood and was submitted to splenectomy at the age of 15 years due to giant splenomegaly. Over the years, he developed hemochromatosis secondary to iron overload and was infected by hepatitis C, which was most probably transmitted via blood transfusion before 1990. In April 2018, during a periodical clinical follow-up, an abdominal ultrasound examination revealed multiple lesions in the right hepatic lobe, subsequently confirmed by contrast-enhanced/computed tomography (ce-CT). The patient underwent biopsy which resulted positive for well-differentiated hepatocellular carcinoma (HCC). He received sorafenib until September 2018 when treatment was discontinued due to the onset of cutaneous toxicity and evidence of progressive disease shown by ce-CT. He was enrolled for a loco-regional treatment of the hepatic lesion through radioembolization with ⁹⁰Y-microspheres. Before the radioembolization procedure, he was submitted to positron emission tomography/CT (PET/CT) with ¹⁸F-choline. **(A)** PET maximum intensity projection showed increased tracer uptake in the hepatic dome (black arrow) and diffuse hyperactivity in the axial and appendicular skeleton (black counter arrows). The corresponding fused PET/CT **(B)** and ce-CT **(C)** axial slices demonstrated multiple lesions, with a necrotic peripheral component, in the right hepatic lobe, characterized by intense ¹⁸F-choline incorporation with much higher uptake values (SUV_{max} : 17.0, SUV_{mean} : 6.1) than those calculated in the normal liver parenchyma (SUV_{max} : 7.7, SUV_{mean} : 5.7).

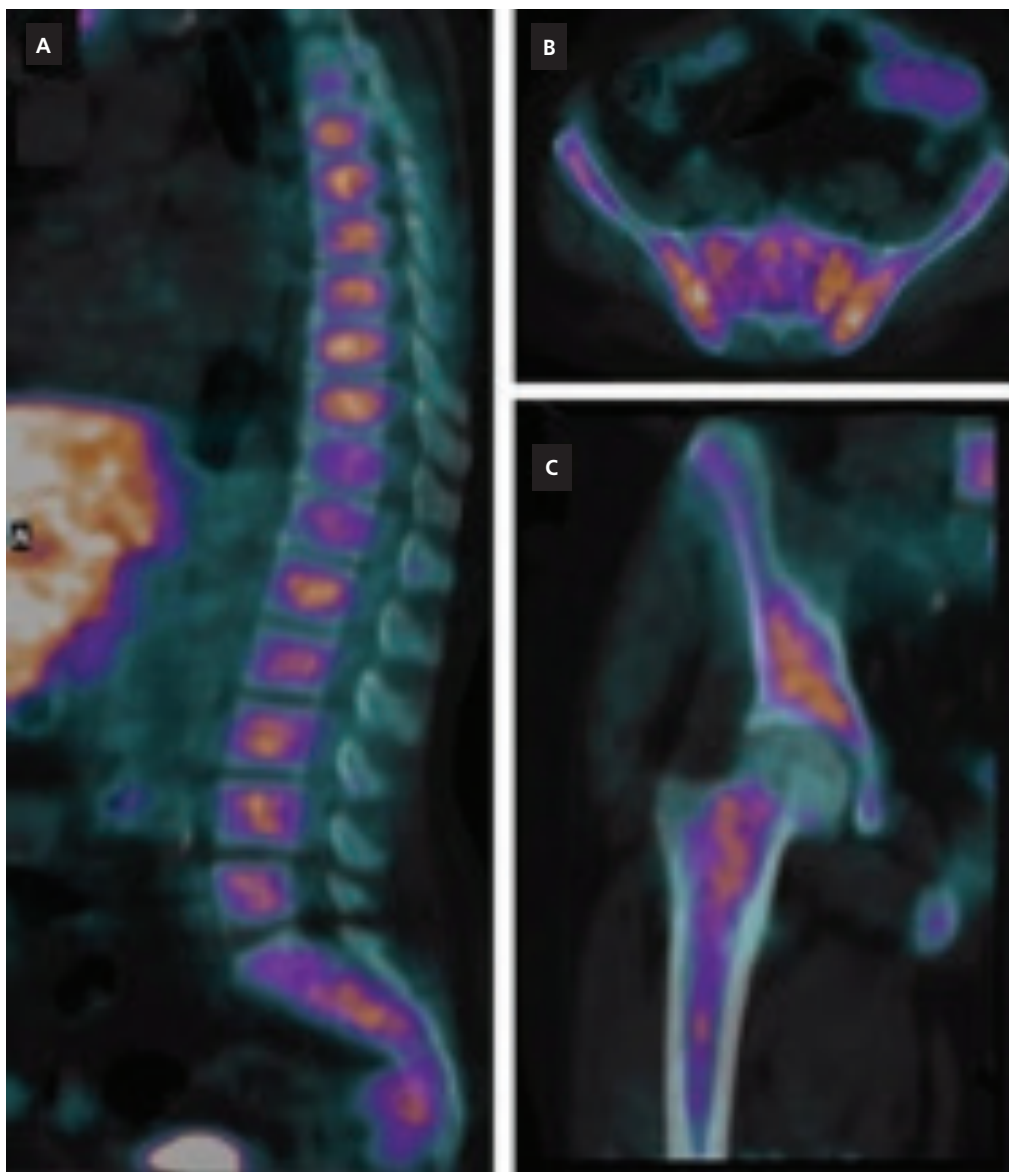


Figure 2. Fused PET/CT well documented tracer incorporation in the endomedullary compartment of the bones, as evident in the sagittal view of vertebrae (**A**), in the axial slice of the pelvic bone (**B**) and in the detailed coronal view of the right femur (**C**). Semiquantitative indices measured in bone marrow, specifically in the pelvic bones, showed significantly increased uptake value (SUV_{max} : 6.2, SUV_{mean} : 4.8) compared with the value reported by Schillaci et al. (1). In a cohort of 80 patients evaluated for assessing the physiological ^{18}F -choline biodistribution (i.e. bone marrow SUV_{mean} : 2.8). Thalassemia intermedia is a rare inherited genetic disease, characterized by a wide spectrum of clinical manifestations (2). Iron overload due to the chronic hemolysis and periodic blood transfusion leads to severe complications, especially at cardiac and hepatic level. Since recent improvements in treatment of thalassemia have led to a significantly prolonged survival, HCC, most probable related to the frequent association of hepatitis C virus-infection and hemochromatosis in thalassemic patients, has emerged as a relatively new complication in long-term survivors (3). Although conventional radiological imaging through CT and magnetic resonance imaging represents the first-line approach for HCC diagnosis, PET/CT with ^{18}F -choline has been introduced as a useful tool for the imaging of HCC, especially before and after loco-regional treatments (4,5). To the best of our knowledge, this is the first report describing the pattern of ^{18}F -choline uptake in a thalassemic patient. It has to be pointed out that diffuse skeletal uptake of ^{18}F -fluciclovine has been recently described in a thalassemic patient affected by prostate cancer with suspicion of bone metastasis (6). Although ^{18}F -fluciclovine and ^{18}F -choline represent different molecular probes in oncology, since the former reflects the upregulation of transmembrane amino-acids transport (7) while the latter is a biomarker of phospholipid synthesis (8), increased uptake of both these 2 tracers in the bone marrow of thalassemic patients are most probably due to hematopoiesis stimulation induced by chronic hemolysis.

Ethics

Informed Consent: This article does not contain any studies with human participants or animals performed by any of the authors.

Peer-review: Externally and internally peer-reviewed.

Financial Disclosure: The author declared that this study received no financial support.

References

1. Schillaci O, Calabria F, Tavolozza M, Ciccio C, Carlini M, Caracciolo CR, Danieli R, Orlacchio A, Simonetti G. ¹⁸F-choline PET/CT physiological distribution and pitfalls in image interpretation: experience in 80 patients with prostate cancer. *Nucl Med Commun* 2010;31:39-45.
2. Aessopos A, Kati M, Farmakis D. Heart disease in thalassemia intermedia: a review of the underlying pathophysiology. *Haematologica* 2007;92:658-665.
3. Mancuso A. Hepatocellular carcinoma in thalassemia: A critical review. *World J Hepatol* 2010;2:171-174.
4. Hartenbach M, Weber S, Albert NL, Hartenbach S, Hirtl A, Zacherl MJ, Paprottka PM, Tiling R, Bartenstein P, Hacker M, Haug AR. Evaluating Treatment Response of Radioembolization in Intermediate-Stage Hepatocellular Carcinoma Patients Using ¹⁸F-Fluoroethylcholine PET/CT. *J Nucl Med* 2015;56:1661-1666.
5. Filippi L, Schillaci O, Bagni O. Recent advances in PET probes for hepatocellular carcinoma characterization. *Expert Rev Med Devices* 2019;16:341-350.
6. Schmitt CR, Schmitt CT, Hinds PR, Sawyer KJ. Use of ¹⁸F-Fluciclovine to Diagnose Recurrent Prostate Carcinoma in a Patient With Beta-Thalassemia. *Clin Nucl Med* 2019;44:544-545.
7. Savir-Baruch B, Zaroni L, Schuster DM. Imaging of Prostate Cancer Using Fluciclovine. *Urol Clin North Am* 2018;45:489-502.
8. Bagni O, Filippi L, Schillaci O. Incidental detection of colorectal cancer via ¹⁸F-choline PET/CT in a patient with recurrent prostate cancer: usefulness of early images. *Clin Nucl Med* 2015;40:328-330.