A case of hepatic focal nodular hyperplasia mimicking hepatocellular carcinoma identified on gallium-68-prostate-specific membrane antigen positron emission tomography/computed tomography

ABSTRACT
Prostate-specific membrane antigen (PSMA) is a glycosylated type-II transmembrane protein highly expressed in individual tumor cells. Lesions with PSMA expression in the liver are commonly reported as prostate cancer metastasis or hepatocellular cancer previously. This is the first case reported as hepatic focal nodular hyperplasia, mimicking hepatocellular carcinoma with imaging features. This patient, having a lesion that has been enlarged from 2.0 cm to 2.5 cm in 3 months, was referred to our department for restaging by gallium-68 PSMA (68Ga-PSMA) positron emission tomography/computed tomography (PET/CT). The CT scan showed a focal segment VI hypodensity, which was significantly PSMA-avid. Consequently, its biopsy resulted as focal nodular hyperplasia in liver. His follow-up 68Ga-PSMA PET/CT ultimately revealed a mass lesion of 8 cm of axial diameter.

Keywords: Focal nodular hyperplasia, gallium-68-prostate-specific membrane antigen positron emission tomography/computed tomography, hepatocellular cancer, liver, metastasis, pitfall, prostate cancer

INTRODUCTION
Focal nodular hyperplasia (FNH) is the second most common benign tumor of the liver, with an incidence of approximately 3%–5% in general population. We present a male case which has been followed conservatively with FNH in which the growth of liver lesion is thought to be associated with the use of androgen deprivation therapy (ADT) for primary prostate cancer.

CASE REPORT
A 66-year-old man who had been diagnosed as prostate cancer pT2cN0M1b, with Gleason score 8 (4 + 4) and had received chemotherapy in 2015 underwent restaging by gallium-68-prostate-specific membrane antigen (68Ga-PSMA) positron emission tomography/computed tomography (PET/CT) in June 2017 after a prostate-specific antigen (PSA) spike of 8.0 ng/ml. among his previous fluctuating PSA levels without a steady increase, after having his written consent. This PET/CT revealed increased levels of radiopharmaceutical in prostate gland with metastatic bone and lymph node lesions, with a subtle liver lesion in segment VI [Figure 1a-d] without any prominent lesion on noncontrast CT.

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Submission: 03-Aug-2020, Revised: 16-Oct-2020, Accepted: 17-Aug-2020, Published: 12-Feb-2021

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How to cite this article: Bilgic S, Sayman HB, Sager MS, Sonmezoglu K. A case of hepatic focal nodular hyperplasia mimicking hepatocellular carcinoma identified on gallium-68-prostate-specific membrane antigen positron emission tomography/computed tomography. World J Nucl Med 2021;20:192-4.
The heterogeneous uptake of $^{68}$Ga-PSMA in liver segment VI was interpreted as hepatocellular cancer (HCC) or an organ metastasis from his primary prostate cancer. Although follow-up magnetic resonance imaging (MRI) was in favor of metastasis [Figure 2], the double checked histopathological findings ended-up as ductular proliferation and fibrosis of liver containing some vascular structures, that is, typical for FNH.[1]

During his follow-up, another $^{68}$Ga-PSMA PET/CT ordered in January 2020 showed multiple lesions with intense uptakes of the tracer in prostate, multiple infra-diaphragmatic lymph nodes, left of acetabulum posteriorly, and a markedly progressive liver lesion in segment VI [Figure 3b].

DISCUSSION

Metastatic prostate cancer has potential to disseminate in body organs especially after it changes to castration resistant form. Main sites of metastasis were known as lymph nodes, bone and lungs but liver metastasis are not so infrequent in these patients.[2] $^{68}$Ga-PSMA hybrid imaging is becoming modality of choice in the current follow-up of this kind of patients selectively wherever it is available and found more useful than biochemical markers. But it has some drawbacks due to its false-positive results: HCC,[3] renal cell cancer,[4] colorectal cancer,[5] follicular thyroid carcinoma,[6] anthracosis,[7] mediastinal sarcoidosis,[8] hepatic hemangioma,[9] vertebral hemangioma,[10] and glioblastoma.[11]

In this case report, we presented an intriguing liver uptake of $^{68}$Ga-PSMA, which could be interpreted as HCC or a metastatic lesion from prostate cancer. Further work-up even with MRI was not confirmative unless a histopathological examination was done.

The continued clinical survey of this patient suggested the primary prostate cancer had favorable course in pace with the PET/CT images showing stable disease in metastatic lymph node and bone lesions [Figure 3]. However, the relatively increased size of segment VI lesion in liver was noticed in his serial follow-up MRIs. This phenomena was hypothetically defined as a lesion progression probably due to the increased levels of estrogen as a result of ADT for primary prostate cancer and should be considered as a pitfall in the interpretation of liver lesions in those prostate cancer patients scanned with $^{68}$Ga-PSMA PET/CT.[12]

FINANCIAL SUPPORT AND SPONSORSHIP
Nil.

CONFLICTS OF INTEREST
There are no conflicts of interest.
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Figure 3: The patient treated with gonadotropin releasing hormone analogs and restaged in January 2020 with a decrease in prostate specific antigen levels to 3.54 ng/ml from previous records. Current scans (a-d) showed increase in liver lesion, from 2.0 cm to 8.0 cm, diagnosed as focal nodular hyperplasia with stable metastatic lymph node and bone lesions from prostate cancer.