# <sup>68</sup>Ga-PSMA PET/CT versus <sup>18</sup>F-FDG PET/CT for detecting lesions in a case of fumarate hydratase-deficient renal cell carcinoma

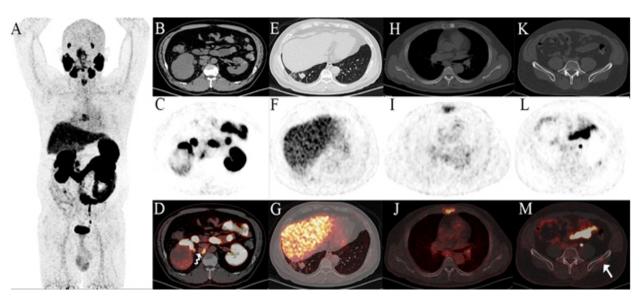
#### Abstract

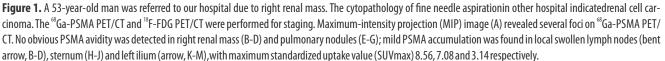
Gallium-68 (<sup>66</sup>Ga)-prostate-specific membrane antigen (PSMA) positron emission tomography/computed tomography (PET/CT) and fluorine-18-fluorodeoxyglucose (<sup>18</sup>F-FDG) PET/CT were performed for staging in a 51-year-old man with renal cell carcinoma. Compared with <sup>18</sup>F-FDG PET/CT, no obvious tracer uptake in right renal mass and less metastatic lesions were found on <sup>68</sup>Ga-PSMA PET/CT. Postoperative pathology demonstrated the diagnosis of fumarate hydratase-deficient renal cell carcinoma (FHRCC).

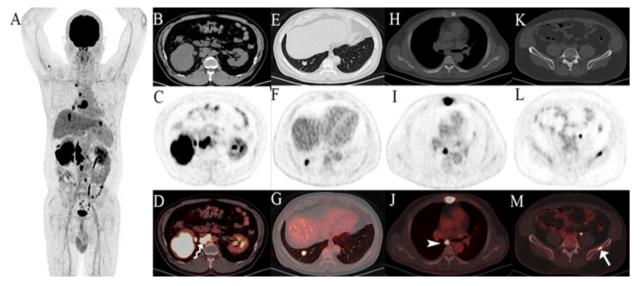
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**Figure 2.** Fluorine-18-FDG PET/CT was further performed next day. Maximum-intensity projection revealed multiple foci on <sup>18</sup>F-FDG PET/CT (A). Axial CT, PET and fused images found intense <sup>18</sup>F-FDG uptake in right renal mass with SUVmax 25.59 (B-D) and local lymph nodes with SUVmax 38.37 (bent arrow, B-D). Besides, pulmonary nodules (E-G), sternum, mediastinal lymph nodes(arrow head, H-J) and left ilium (arrow, K-M) also presented intense <sup>18</sup>F-FDG uptake, with SUVmax 19.18, 22.46, 16.78 and 10.66, respectively.

Combined the imaging results of two tracers, we draw the conclusion that renal cancer with metastases of local lymph nodes, lung, bone, and mediastinal lymph nodes. Subsequent radical right nephrectomy was performed [1], and postoperative pathology revealed the diagnosis of FHRCC. Prostate-specific membrane antigenis a type II integral membrane glycoprotein which is highly expressed in prostate cancer cells and tumor-associated neovasculature of other solid tumors [2]. Baccala et al. (2007) [3] demonstrated PSMA is also expressed in tumor-associated neovasculature of RCC, especially in clear cell renal cell carcinoma (ccRCC). Besides, Spatz et al. (2018) [3, 4] found higher expression of PSMA had worse survival. The first application of PSMA PET/CT on ccRCC was reported in 2014 and promising results were found [5, 6]. Afterwards, PSMA PET/CT was increasingly used in RCC for diagnosis and staging. The SUVmax of primary renal tumors on <sup>66</sup>Ga-PSMA PET/CT can predict aggressive pathological features and the expression of vascular endothelial growth factor receptors and platelet-derived growth factor in RCC patients [7, 8]. With evidence increasing, <sup>68</sup>Ga-PSMA PET/CT was found with high false negative in some nonccRCCs, including papillary RCC, chromophobe RCC, unclassified RCC and Xp11 translocation RCC [9-13]. Fumarate hydratase-deficient renal cell carcinoma is a rare non-ccRCC characterized by the deficiency of fumarate hydratase (FH), which is induced by the germline / somatic gene of FH gene [14-16]. The deficiency of FH would disrupt tricarboxylic acid cycle and activate hypoxia-inducible factor-1, which can increase glucose uptake [17]. To our knowledge, there were no cases reported the imaging of <sup>68</sup>Ga-PSMA PET/CT in FHRCC and compared the ability of <sup>68</sup>Ga-PSMA PET/CT and <sup>18</sup>F-FDG PET/CT in detecting lesions in FHRCC. Based on above imaging results, we thought <sup>18</sup>F-FDG PET/CT may be more suitable than <sup>68</sup>Ga-PSMA PET/CT in FHRCC patients for detecting primary and metastatic lesions.

## **Bibliography**

- 1. Barata PC, Rini Bl. Treatment of renal cell carcinoma: Current status and future directions. CA Cancer J Clin 2017;67:507-24.
- 2. Uijen MJM, Derks YHW, Merkx RIJ et al. PSMA radioligand therapy for solid tumors other than prostate cancer: background, opportunities, challenges, and first clinical reports. *Eur J Nucl Med Mol Imaging* 2021;48:4350-68.
- 3. Baccala A, Sercia L, Li J et al. Expression of prostate-specific membrane antigen in tumor-associated neovasculature of renal neoplasms. *Urology* 2007; 70: 385-90.
- 4. Spatz S, Tolkach Y, Jung K et al. Comprehensive evaluation of prostate specific membrane antigen expression in the vasculature of renal tumors: Implications for imaging studies and prognostic role. *J Urol* 2018; 199: 370-7.
- 5. Demirci E, Ocak M, Kabasakal L et al. <sup>68</sup>Ga-PSMA PET/CT imaging of metastatic clear cell renal cell carcinoma. *Eur J Nucl Med Mol Imaging* 2014;41:1461-2.
- 6. Gühne F, Seifert P, Theis B et al. PSMA-PET/CT in patients with recurrent clear cell renal cell carcinoma: Histopathological correlations of imaging findings. *Diagnostics (Basel)* 2021; 11:1142.
- 7. Gao J, Meng L, Xu Q et al. <sup>66</sup>Ga-PSMA-11 PET/CT parameter correlates with pathological VEGFR-2/PDGFR-β expression in renal cell carcinoma patients. *Mol Imaging Biol* 2022; 24: 759-68.
- 8. Gao J, Xu Q, Fu Y et al. Comprehensive evaluation of <sup>68</sup>Ga-PSMA-11 PET/CT parameters for discriminating pathological characteristics in primary clearcell renal cell carcinoma. *Eur J Nucl Med Mol Imaging* 2021;48:561-9.
- 9. Siva S, Callahan J, Pryor D et al. Utility of <sup>66</sup>Ga prostate specific membrane antigen positron emission tomography in diagnosis and response assessment of recurrent renal cell carcinoma. *JMed Imaging Radiat Oncol* 2017;61:372-8.
- 10. Sawicki LM, Buchbender C, Boos J et al. Diagnostic potential of PET/CT using a <sup>68</sup>Ga-labelled prostate-specific membrane antigen ligand in wholebody staging of renal cell carcinoma: initial experience. *Eur J Nucl Med Mol Imaging* 2017;44: 102-7.
- 11. Mittlmeier LM, Unterrainer M, Rodler S et al. <sup>18</sup>F-PSMA-1007 PET/CT for response assessment in patients with metastatic renal cell carcinoma undergoing tyrosine kinase or checkpoint inhibitor therapy: preliminary results. *Eur J Nucl Med Mol Imaging* 2021;48:2031-7.
- 12. Muselaers S, Erdem S, Bertolo R et al. PSMA PET/CT in renal cell carcinoma: An overview of current literature. J Clin Med 2022; 11: 1829.
- 13. Yin Y, Campbell SP, Markowski MC et al. Inconsistent detection of sites of metastatic non-clear cell renal cell carcinoma with PSMA-Targeted <sup>18</sup>F-DCFPyL PET/CT. *Mol Imaging Biol* 2019; 21:567-73.
- 14. Lau HD, Chan E, Fan AC et al. A clinicopathologic and molecular analysis of fumarate hydratase-deficient renal cell carcinoma in 32 patients. *Am J Surg Pathol* 2020; 44: 98-110.
- 15. Smith SC, Trpkov K, Chen YB et al. Tubulocystic carcinoma of the kidney with poorly differentiated foci: A frequent morphologic pattern of fumarate hydratase-deficient renal cell carcinoma. *Am J Surg Pathol* 2016; 40: 1457-72.
- 16. Sun G, Zhang X, Liang J et al. Integrated molecular characterization of fumarate hydratase-deficient renal cell carcinoma. *Clin Cancer Res* 2021; 27: 1734-43.
- 17. Schmidt C, Sciacovelli M, Frezza C. Fumarate hydratase in cancer: A multifaceted tumour suppressor. Semin Cell Dev Biol 2020; 98: 15-25.

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