

FDG PET or PET/CT in Evaluation of Renal Angiomyolipoma

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Objective: Angiomyolipoma is the most common benign kidney tumor. However, literature describing FDG PET findings on renal angiomyolipoma (AML) is limited. This study reports the FDG PET and PET/CT findings of 21 cases of renal AML.

Materials and Methods: The study reviews FDG PET and PET/CT images of 21 patients diagnosed with renal AML. The diagnosis is based on the classical appearance of an AML on CT scan with active surveillance for 6 months. The study is focused on the observation of clinical and radiographic features.

Results: Six men and 15 women were included in our study. The mean age of the patients was 57.14 ± 9.67 years old. The mean diameter of 21 renal AML on CT scans was 1.76 ± 1.00 cm (Min: 0.6 cm; Max: 4.4 cm). CT scans illustrated renal masses typical of AMLs, and the corresponding FDG PET scans showed minimal FDG activities in the area of the tumors. None of the 21 AMLs showed a maximum standardized uptake value (SUV_{max}) greater than 1.98. No statistically significant correlation was present between SUV_{max} and tumor size.

Conclusion: Renal AMLs demonstrate very low to low uptake on FDG PET and PET/CT imaging in this study. When a fat-containing tumor in the kidney is found on a CT scan, it is critical to differentiate an AML from a malignant tumor including an RCC, liposarcoma, and Wilms tumor. This study suggests that FDG PET or PET/CT imaging is useful for differentiating a renal AML from a fat-containing malignant tumor.

Index terms: Computed tomography; Fluorine-18 fluoro-2-deoxy-D-glucose; Positron emission tomography; Renal angiomyolipoma.

INTRODUCTION

Renal angiomyolipoma (AML) is a relatively rare renal cortical neoplasm that has been identified in less than 0.2% of the general population. AMLs are composed of smooth

muscle cells, dysmorphic blood vessels, and fatty tissue with a known benign nature. AML is usually asymptomatic and is often managed with active surveillance protocols (1). AMLs account for 24% of spontaneous renal hematomas and sudden-onset flank pain is the most common symptom (2).

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Conditions for intervention are suspicion of malignancy, size greater than 4 cm, spontaneous hemorrhage, and pain (3).

The classical appearance of an AML on a CT scan is a predominantly fatty inhomogeneous mass with varying amounts of tissue density interspersed within it. The fat seen in this lesion is similar in appearance to subcutaneous or retroperitoneal fat (4). On a CT, fat is characterized by its negative number of Hounsfield units (HUs). A tissue attenuation of -10 HU is typically a reliable indicator of fat. A well-circumscribed renal mass with intratumoral fat on CT scan is similar to the diagnostic finding of renal angiomyolipoma (5).

Currently, positron emission tomography (PET) forms part of the routine assessment of malignant diseases. The vast majority of PET scans are performed with fluorine-18 fluoro-2-deoxy-D-glucose (FDG), an analogue of glucose. Many cancers correlate with their metabolic activities, as indicated by the degree of FDG accumulation on PET imaging (6).

Angiomyolipoma is the most common benign tumor of the kidney (5). However, literature describing the FDG PET or PET/CT findings on renal AML is limited. This study reports FDG PET and PET/CT findings of 21 cases of renal AML.

MATERIALS AND METHODS

Between January 2006 and December 2010, 473 patients were diagnosed with typical AMLs on CT scan with active surveillance for 6 months in our hospital. Researchers retrospectively extracted clinical and radiographic data from the complete medical records. Twenty-one cases had received FDG PET or PET/CT imaging for other oncologic surveys; these formed the basis of this study. The time gap between the diagnostic CT and the FDG PET (or PET/CT) was within one month. This study was approved by the hospital ethics committee (DMR-99-IRB-010-2).

Computed Tomography Scan

In all patients, triple-phase dynamic CT examination was performed using MDCT scanners (LightSpeed 16; GE Medical Systems Milwaukee, WI, USA). The images were reconstructed at 5-mm intervals to provide contiguous sections. All patients underwent a contrast-enhanced biphasic helical CT examination including arterial and equilibrium phases. Following a bolus-triggered technique, the arterial phase began 20-35 s after a 100-mL injection of nonionic iodinated contrast material with an automatic

power injector, at a rate of 3 mL/s. The venous phase began 180 s after initiating the contrast injection.

FDG PET and PET/CT Imaging

For the ¹⁸F-FDG PET and PET/CT examinations, patients were asked to fast for at least 4 hours before scanning. Each patient was injected intravenously with 370 megabecquerel (MBq) of 2-¹⁸F-FDG PET and rested supine in a quiet, dimly lit room. Imaging was performed with a positron emission tomography/computed tomography (PET/CT) scanner or PET scanner (PET/CT: Discovery STE, PET: Advance NXi, General Electric Healthcare, Milwaukee, WI, USA). Scanning began approximately 60 min after the FDG injection. When patients were positioned in the PET/CT scanner, a molded headrest and a head-restraining Velcro band were applied to secure their heads firmly, to reduce motion artifact.

Fluorine-18 fluoro-2-deoxy-D-glucose positron emission tomography scanning was performed with an axial field of view of 14.6 cm. Emission scans required a 3-min acquisition time in 2-D mode at every table position, typically requiring 6 or 7 bed positions to cover the entire field of view. After emission scanning a transmission scan was started using ⁶⁸Ge pin sources rotating around the body. Transmission scans were performed from the head to the pelvic floor of the patient, with a 2-min acquisition time at every table position. Image data sets were reconstructed using an iterative algorithm with segmented attenuation correction from the collected data, which had been corrected for scatter, random events, and dead time.

The PET/CT examinations were started by acquiring a CT topogram to define the axial examination range of the PET/CT study. After defining the axial imaging range, a spiral non-contrast-enhanced low-radiation-dose CT scan (0.8-second rotation time, 120 kVp, variable mA with AutomA technique, 3.75-mm slice thickness, and 1.75 : 1 pitch) was acquired for anatomical references and attenuation correction. PET emission images were then acquired after CT scans at 2 min per field of view (FOV) in the 3-D acquisition mode with 11-slice overlap at the FOV borders. The CT images were reconstructed onto a 512 × 512 matrix and converted to a 128 × 128 matrix, with 511-keV-equivalent attenuation factors for attenuation correction of the corresponding PET emission images. The PET images were then reconstructed to attenuation-corrected, 3.27-mm transaxial slice thickness for further interpretation.

Researchers measured the maximum standardized uptake values (SUV_{max}) of each AML on the FDG PET and PET-CT

images.

Statistical Analysis

We expressed the values of age, tumor size on CT scan, and SUV_{max} of tumor on FDG PET and PET/CT as mean \pm standard deviation. The relationship between tumor size on CT scan and SUV_{max} of tumor on FDG PET and PET/CT in renal AML of our study was analyzed using a Spearman rank correlation. All analyses were conducted by STATA 11.0 using a 0.05 level of significance.

RESULTS

Table 1 shows the profile of the renal AML patients, tumor size on CT scan, and SUV_{max} of tumor on FDG PET and PET/CT. Of the 21 patients, 15 were women and 6 were men. The mean age of the patients was 57.14 ± 9.67 years. The mean diameter of 21 renal AMLs on CT scan was 1.76 ± 1.00 cm. The minimum tumor diameter on CT scans was 0.6 cm and maximum diameter was 4.4 cm. The mean SUV_{max} was 1.25 ± 0.34 . None of the 21 AML showed SUV_{max} greater than 1.98 (Figs. 1, 2). Generally, FDG uptake (expressed as $SUV_{max} > 2.5$) is interpreted as being “positive” or PET-avid. No positive FDG PET or PET/CT imaging was found in the renal AML of our study. There was no statistically significant correlation between SUV_{max} and tumor size ($B = 0.045, p = 0.85$).

DISCUSSION

Renal angiomyolipoma is a fairly common benign renal tumor that is often found incidentally. Renal AML is composed of mature adipose tissue, smooth muscle and

blood vessels. AML is not always located in the kidney, but may also be found in the liver, spleen, uterus, and tubes (7).

Literature on the role of FDG PET or PET/CT in the diagnosis AMLs is limited. Lhommel et al. (8) reported a case where a liver AML did not have increased FDG uptake but had increased C11 acetate uptake. Ho et al. (9) reported increased C11 acetate uptake but no abnormal FDG uptake

Table 1. Clinical Features of Patients, Size on CT Scan and SUV_{max} on FDG PET or PET/CT of Renal AML

| Patient | Gender | Age | Tumor Size (cm) | SUV_{max} |
|---------|--------|-----|-----------------|-------------|
| 1 | Male | 51 | 1.7 | 1.45 |
| 2 | Female | 60 | 0.7 | 1.07 |
| 3 | Female | 34 | 2 | 1.54 |
| 4 | Male | 60 | 1.1 | 0.96 |
| 5 | Female | 58 | 2.2 | 1.35 |
| 6 | Female | 53 | 0.8 | 1.45 |
| 7 | Female | 56 | 1.5 | 1.98 |
| 8 | Female | 56 | 1 | 0.87 |
| 9 | Female | 76 | 1.3 | 0.97 |
| 10 | Male | 58 | 2.2 | 0.94 |
| 11 | Female | 53 | 0.6 | 1.15 |
| 12 | Male | 55 | 0.8 | 1.23 |
| 13 | Female | 57 | 2.4 | 1.51 |
| 14 | Female | 55 | 1 | 1.07 |
| 15 | Female | 47 | 2.6 | 0.88 |
| 16 | Female | 71 | 4.4 | 0.92 |
| 17 | Female | 48 | 3.5 | 1.76 |
| 18 | Male | 77 | 3 | 1.37 |
| 19 | Female | 59 | 1.1 | 1.83 |
| 20 | Female | 65 | 2 | 0.81 |
| 21 | Male | 51 | 1.2 | 1.04 |

Note.— SUV_{max} = maximum standardized uptake values, AML = angiomyolipoma

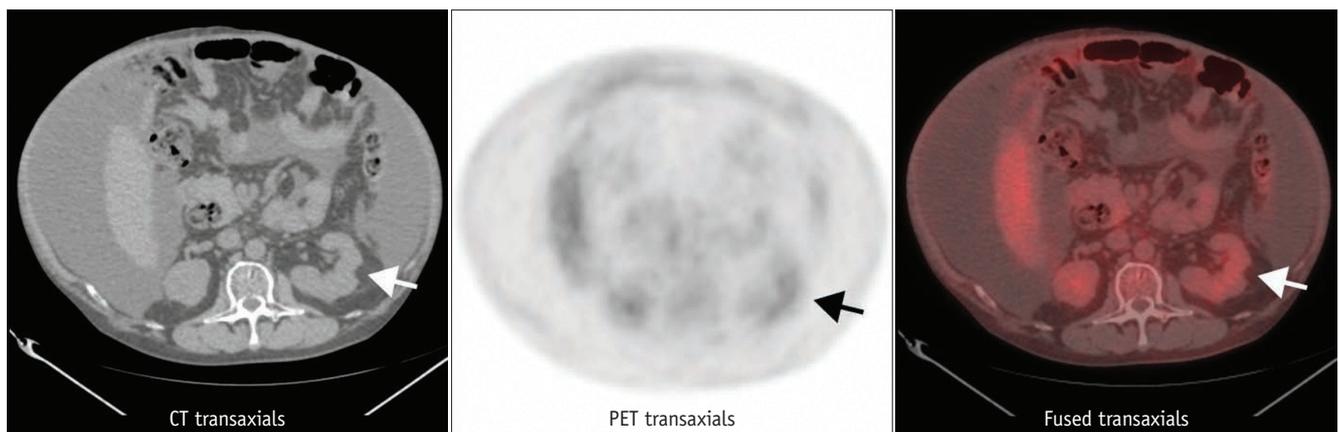


Fig. 1. 58-year-old hepatoma male patient with liver cirrhosis with massive ascites and angiomyolipoma (AML) in left kidney received whole body FDG PET/CT for detecting extrahepatic metastases (patient no. 10 in Table 1). Uptake of ^{18}F -FDG in renal AML was lower than that of spine (arrows).

in a renal AML. Arnold and Myers (10) documented renal AML as metabolically active on FDG PET and falsely positive for renal cell carcinoma (RCC).

Diagnosis of renal AMLs relies on demonstrating macroscopic fat within a lesion on a CT scan. Some studies report that renal AML may be distinguished from RCC on

images by identifying macroscopic fat (11). Davenport et al. reported that a threshold of -10 HU or lower enables accurate differentiation of renal AML from RCC on non-enhanced phase (12). Although demonstrable fat on a CT scan is highly suggestive of an AML, with differential diagnosis of a fat-containing tumor in the kidney, the

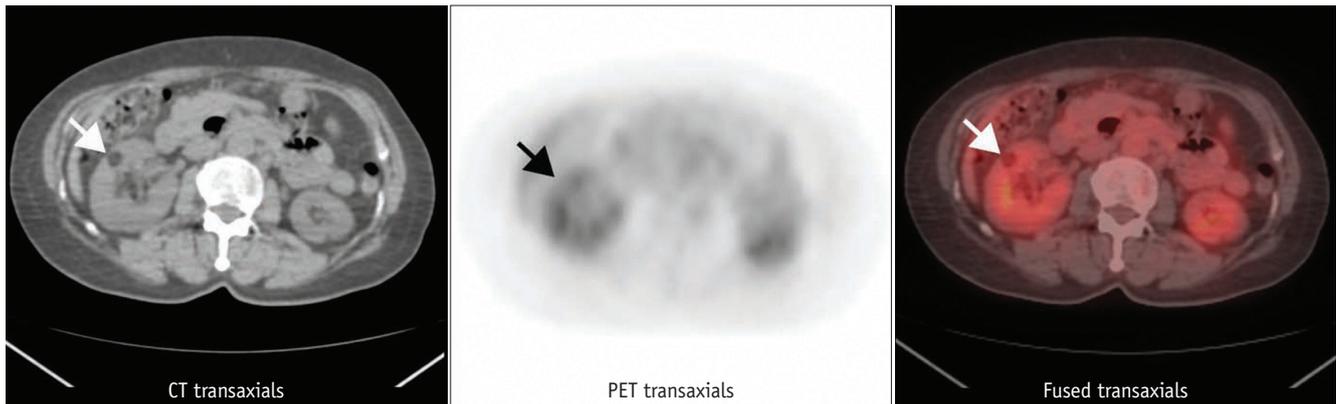


Fig. 2. 59-year-old female patient with angiomyolipoma (AML) in right kidney (patient no. 19 in Table 1). Uptake of ^{18}F -FDG in renal AML was higher than that of spine but lower than hepatic activity (arrows).

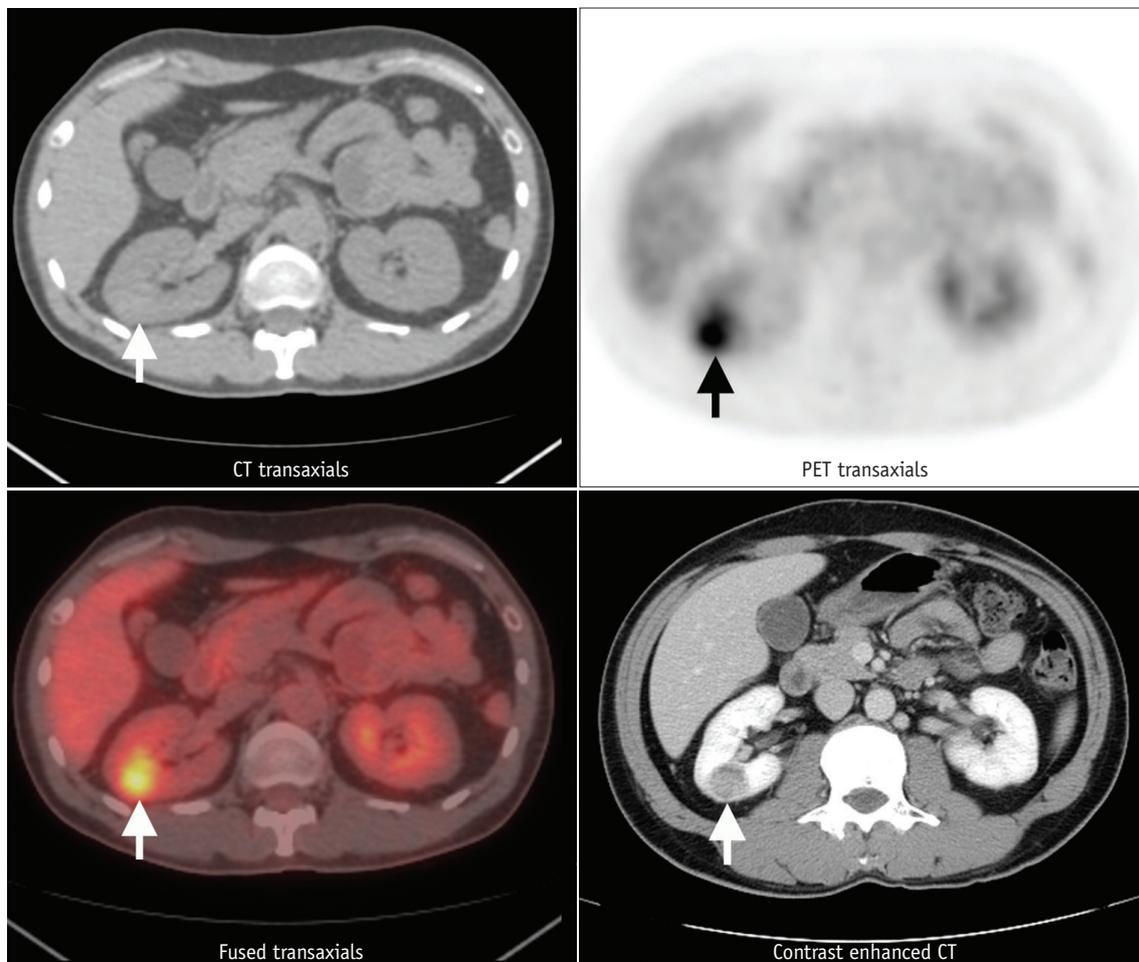


Fig. 3. 30-year-old male with renal cell carcinoma in right kidney (arrows). Maximum standardized uptake value on FDG PET is 5.43.

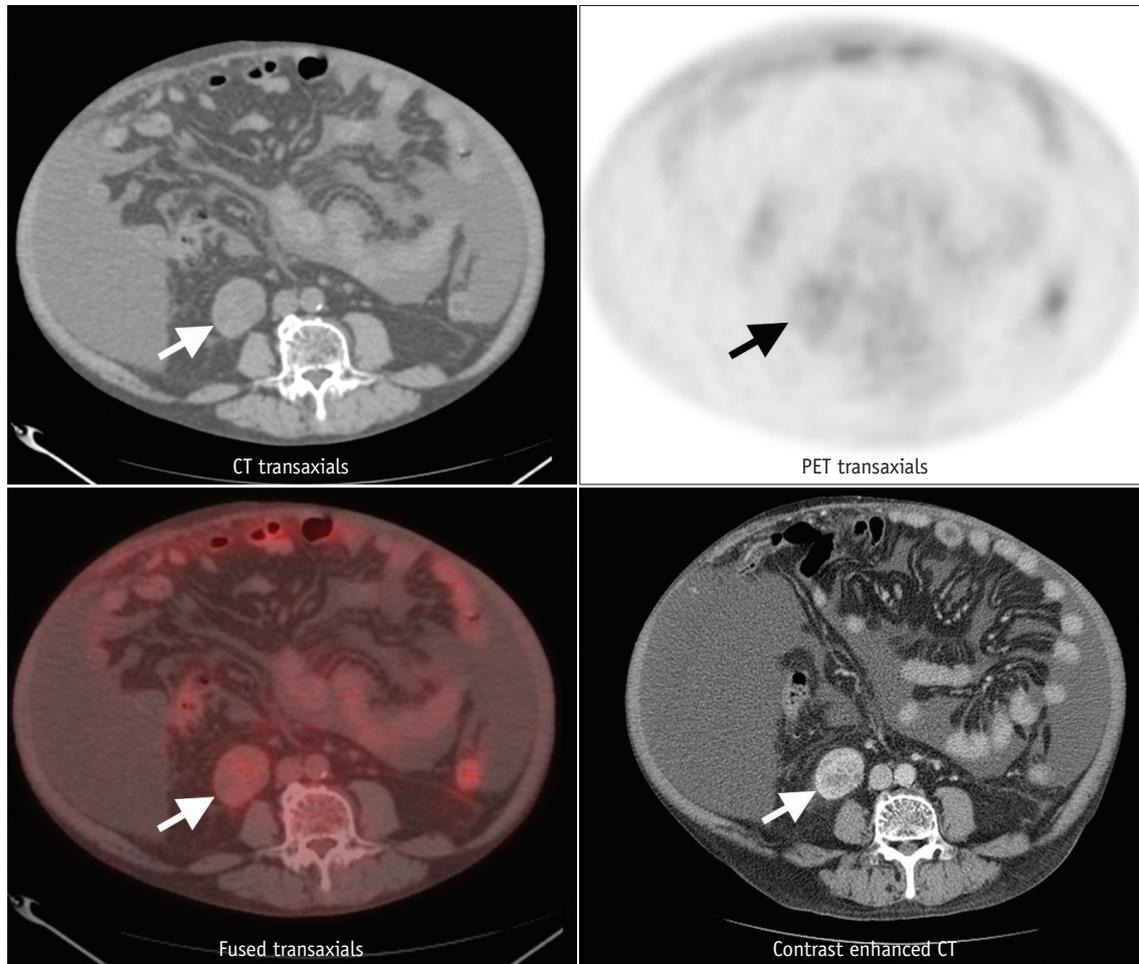


Fig. 4. 58-year-old male with renal cell carcinoma in right kidney (arrows). Maximum standardized uptake value on FDG PET is 1.88.

possibility that it may be an RCC, lipoma, liposarcoma, and Wilms tumor needs to be considered. The most important differential is that of an RCC (5). The mean SUV_{max} value of pathologic confirmed primary RCC in Namura et al. (13) was 10.62 (range, 5.2 - 16.6), which was much higher than the SUV_{max} of renal AML in our study (mean, 1.25; range, 0.81 - 1.98).

Renal cell carcinoma accounts for 3.5% of all adult malignancies and 25% of patients with RCC will not survive (14). Bachor et al. (15) reported that the true positive rate of solid renal masses with confirmed RCC on PET is 77%. A pathologic confirmed RCC case with high SUV_{max} is shown in Figure 3. Kang et al. (16) found that on clinical use of PET in detection of RCC, FDG PET exhibited a sensitivity of 60% and specificity of 100% for primary RCC. Aide et al. (17) reported that high false negative rates of diagnosis of RCC on FDG PET resulted in a sensitivity, specificity, and accuracy of 47%, 80%, and 51%, respectively. A false negative case of pathologic confirmed RCC is shown in Figure 4. One meta-analysis study suggested that FDG PET

can be useful in detecting recurrent or metastatic RCC, with 85.7% accuracy (18). The significant variability in the uptake of FDG by RCC is partly due to the excretion of FDG via the urinary tract.

Determinants of FDG accumulation include tumor blood flow, glucose transport, and glycolytic rate. All renal AMLs in this study demonstrated very low to low uptake on FDG PET and PET/CT imaging, suggesting that these tumors were not FDG-avid. AMLs are known to be hyper-vascular with an associated risk of spontaneous hemorrhage; however, there was no significant correlation between tumor size and SUV_{max} in this study. Neither tumor size nor blood flow explains the relative paucity of FDG activity in renal AML. Studies report glycolytic enzyme expression to be under the influence of mammalian target of rapamycin (mTOR), and that treatment with rapamycin suppressed the rate of glycolysis (19). Jiang et al. (20) reported that mammalian rapamycin complex (mTORC1) is insufficient for increased glycolysis in tumors and that constitutive mTORC1 activity with negatively regulated glucose transporter trafficking.

Their *in vivo* data indicated that constitutive activation of mTORC1 negatively regulates hexose transporter trafficking and glucose uptake. Their study's 3 renal AML cases confirm the relevance of their findings in humans (20). The minimal FDG uptake in 21 renal AML patients in the present study may be explained by mTORC1 activity with negatively regulated glucose transporter trafficking and resulting in a reduction in glucose uptake in our 21 renal AMLs.

Conclusion

Renal AMLs demonstrated very low to low uptake on FDG PET and PET/CT imaging in this study. When a fat-containing tumor in the kidney is present on a CT scan, it is critical to differentiate an AML from a malignant tumor. Based on the results of this study, the authors suggest that FDG PET or PET/CT imaging is a useful tool to differentiate a renal AML from a fat-containing malignant tumor.

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