CXCR4 - directed SPECT Imaging using [99mTc]Tc-PentixaTec in Patients with Primary Aldosteronism

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Abstract

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Introduction: Primary aldosteronism (PA) is an underdiagnosed cause of hypertension either caused by bilateral adrenal hyperplasia (BAH) or unilateral aldosterone-producing adenoma (APA). Previous work has shown that C-X-C-motif chemokine receptor 4-directed positron emission tomography/computed tomography (PET/CT) is an important add-on to native abdominal CT and adrenal vein sampling and can aid in therapeutic decision making. The recent development of a first ^{99m}Tc-labeled CXCR4 ligand with suitable in vivo properties, namely [^{99m}Tc]Tc-PentixaTec, now allows CXCR4-targeted conventional scintigraphy and SPECT, reducing costs and improving general availability of this imaging approach. Here, we report on the first clinical experience with [^{99m}Tc]Tc-PentixaTec as diagnostic tool in PA.

Methods: Six patients (4 male, 2 female, 49±15 years) with PA underwent imaging after i.v. injection of 435±50 MBq [^{99m}Tc]Tc-PentixaTec. First, images were visually inspected for asymmetric tracer accumulation in the adrenal glands. Subsequently, lateralization indices using lesion-to-contralateral adrenal ratio were calculated in SPECT/(CT) images. Scintigraphic findings were compared to CT and adrenal vein sampling in all patients.

Results: Four out of six patients showed lateralization of adrenal tracer accumulation with a median lateralization index of 1.53 (range, 1.28 – 1.68); in the remaining two patients no distinct lateralization of $[^{99\text{m}}\text{Tc}]\text{Tc-PentixaTec}$ uptake could be detected. Focally enhanced tracer retention correlated with adrenal adenomas as detected by CT (median size 17 mm; range, 16 - 19 mm) in all four patients, whereas adrenal vein catheterization did not detect any lateralization in 3/4 subjects. In two of the four patients, primary surgery was recommended: histopathologic work-up confirmed APA in one patient, with an adequate decrease in aldosterone/renin ratio (pre- vs. post-therapeutic: 17.27 vs. 3.91), whereas the second patient received mineralocorticoid receptor antagonists instead of surgery upon patient request. The remaining patients started medical therapy with mineralocorticoid receptor antagonists as well. Clinical follow-up is still pending in these patients.

Conclusions: Non-invasive visualization of CXCR4 in patients with PA using [99mTc]Tc-PentixaTec is feasible. Further research is warranted to elucidate its diagnostic performance, especially in comparison to CXCR4-directed PET.