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^{203}Pb -VMT- α -NET Scintigraphy of a Patient With Neuroendocrine Tumor

Dirk Müller, PhD,* Hendrik Herrmann, PhD,* Michael K. Schultz, PhD,† Christoph Solbach, PhD,* Thomas Ettrich, MD,‡ and Vikas Prasad, MD*

Abstract: In an end-stage midgut neuroendocrine tumor patient with cardioid heart disease, right ventricular dysfunction, mildly reduced renal function, and refractory to 6 cycles of ^{177}Lu -HA-DOTATATE therapy, planar, and 22 hours SPECT/CT images were acquired after injection of 224 MBq of ^{203}Pb -VMT- α -NET to assess the feasibility of performing ^{212}Pb -VMT- α -NET therapy. A comparison of the 1.5 and 22 hours SPECT/CT images with ^{68}Ga -HA-DOTATATE PET/CT showed high uptake of ^{203}Pb -VMT- α -NET in liver metastases matching with the results of the PET/CT investigation.

Key Words: ^{203}Pb , ^{212}Pb , VMT- α -NET, Tyr3-octreotide, SPECT/CT

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From the *Department of Nuclear Medicine, University of Ulm, Ulm, Germany; †Radiology-Division of Nuclear Medicine, University of Iowa, Iowa, IA; and ‡Clinic of Internal Medicine, University of Ulm, Ulm, Germany.

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Correspondence to: Dirk Müller, PhD, Department of Nuclear Medicine, University of Ulm, Albert-Einstein-Allee 23, 89081 Ulm, Germany. E-mail: dirk.mueller@uniklinik-ulm.de; Vikas Prasad, MD, Department of Nuclear Medicine, University of Ulm, Albert-Einstein-Allee 23, 89081 Ulm, Germany. E-mail: vikas.prasad@uniklinik-ulm.de.

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REFERENCES

- Li M, Zhang X, Quinn TP, et al. Automated cassette-based production of high specific activity [$^{203}\text{Pb}/^{212}\text{Pb}$]peptide-based theranostic radiopharmaceuticals for image-guided radionuclide therapy for cancer. *Appl Radiat Isot.* 2017;127: 52–60. Erratum in: *Appl Radiat Isot.* 2020;156:108733.
- Li M, Sagastume EA, Lee D, et al. $^{203}\text{Pb}/^{212}\text{Pb}$ theranostic radiopharmaceuticals for image-guided radionuclide therapy for cancer. *Curr Med Chem.* 2020;27: 7003–7031.
- Zaid NRR, Kletting P, Beer AJ, et al. Mathematical modeling of in vivo alpha particle generators and chelator stability. *Cancer Biother Radiopharm.* 2021. doi:10.1089/cbr.2020.4112. Epub ahead of print.
- Zaid NRR, Kletting P, Winter G, et al. A physiologically based pharmacokinetic model for in vivo alpha particle generators targeting neuroendocrine tumors in mice. *Pharmaceutics.* 2021;13:2132.
- Li M, Liu D, Lee D, et al. Targeted alpha-particle radiotherapy and immune checkpoint inhibitors induces cooperative inhibition on tumor growth of malignant melanoma. *Cancers (Basel).* 2021;13:3676.
- Dos Santos JC, Schäfer M, Bauder-Wüst U, et al. Development and dosimetry of $^{203}\text{Pb}/^{212}\text{Pb}$ -labelled PSMA ligands: bringing “the lead” into PSMA-targeted alpha therapy? *Eur J Nucl Med Mol Imaging.* 2019;46:1081–1091.
- Schottelius M, Šimeček J, Hoffmann F, et al. Twins in spirit—episode I: comparative preclinical evaluation of [(68)Ga]DOTATATE and [(68)Ga]HA-DOTATATE. *EJNMMI Res.* 2015;5:22.
- Schottelius M, Wester HJ, Reubi JC, et al. Improvement of pharmacokinetics of radioiodinated Tyr(3)-octreotide by conjugation with carbohydrates. *Bioconjug Chem.* 2002;13:1021–1030.

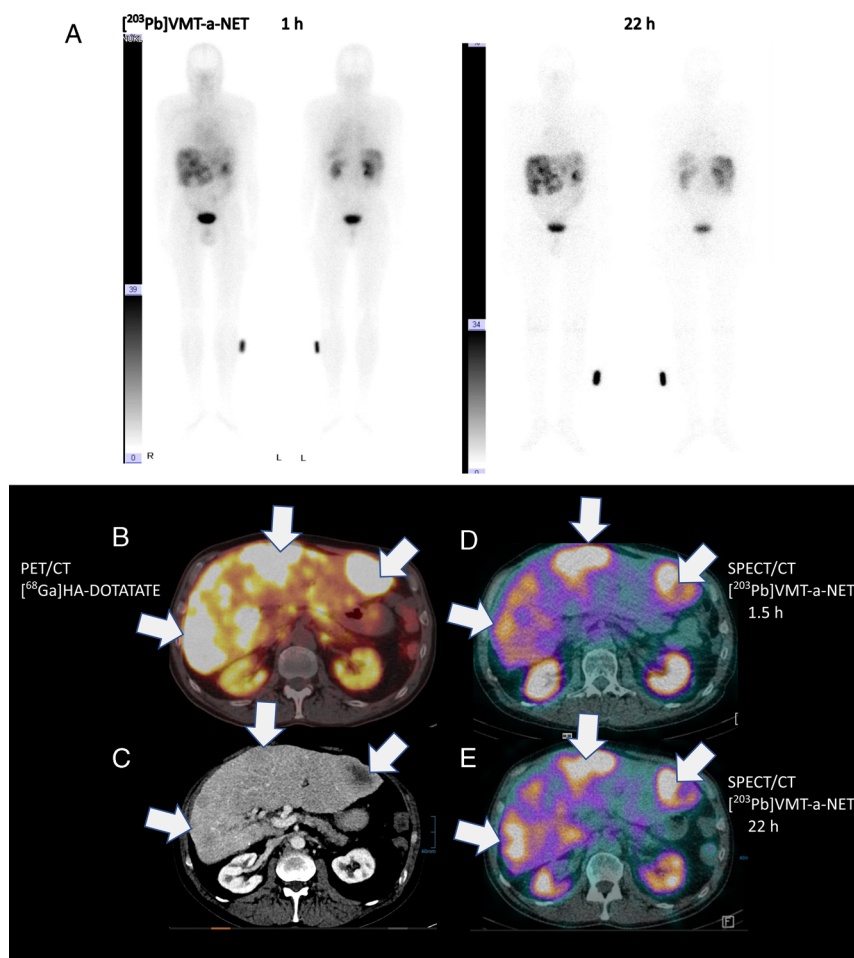


FIGURE 1. Tyr3-octreotide (TOC) variant VMT- α -NET is a novel and ideal ligand for $^{203}\text{Pb}/^{212}\text{Pb}$ that shows lower renal excretion, higher neuroendocrine tumor uptake, and high chelation properties for retention of daughter α -particle emitting radionuclide ^{212}Bi . In an end-stage midgut neuroendocrine tumor patient with carcinoid heart disease, right ventricular dysfunction, mildly reduced renal function, and refractory to 6 cycles of ^{177}Lu -HA-DOTATATE therapy, planar, and 22 hours ^{203}Pb -VMT- α -NET SPECT/CT (low dose) images were acquired after injection of 224 MBq of ^{203}Pb -VMT- α -NET to assess the feasibility of performing ^{212}Pb -VMT- α -NET therapy. The patient tolerated the injection without any significant alteration in the vital parameters. There was rapid renal clearance of the tracer within the first hour itself as was evident from tracer excretion in the urinary bladder and in the renal pelvicalyceal system. Because of known right ventricular dysfunction, there was evidence of blood pool activity in the heart, which however decreased significantly after 21 hours. A comparison of the 1.5 and 22 hours SPECT/CT images with ^{68}Ga -HA-DOTATATE PET/CT images (contrast-enhanced diagnostic CT) showed high uptake of ^{203}Pb -VMT- α -NET in liver metastases matching with the results of the PET/CT. **A**, Anterior and posterior planar images acquired at 1 and 22 hours. **B**, Fused ^{68}Ga -HA-DOTATATE PET/CT transverse slice. **C**, Transverse contrast-enhanced CT image. **D** and **E**, SPECT/CT (low-dose) fused transverse slice. Arrows are showing the liver metastases.¹⁻⁸