

New radiopharmaceutical shows promise for improved detection of neuroendocrine tumors

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Representative maximum-intensity projection from PET/CT scans comparing [55Co]Co-DOTATATE (A–C), [64Cu]Cu-DOTATATE (D–F), and [68Ga]Ga-DOTATATE (G). Arrows indicates tumor in each animal. Decrease in background was observed for [55Co]Co-DOTATATE for late imaging. Higher relative liver uptake with time was observed for [64Cu]Cu-DOTATATE, whereas high bladder uptake was observed for [68Ga]Ga-DOTATATE. All images are scaled to maximum uptake in tumor. Credit: T.L. Andersen, Dept. of Nuclear Medicine, Odense University Hospital, Odense, Denmark.

The newly developed [55Co]55Co-DOTATATE imaging agent has emerged as a more accurate and sensitive radiopharmaceutical to aid in the diagnosis of neuroendocrine tumor metastases. According to research published in the February issue of the *Journal of Nuclear Medicine*, positron emission tomography/computed tomography (PET/CT) imaging with [55Co]Co-DOTATATE resulted in superior image contrast and enhanced detection of metastases as compared with other commonly used radiopharmaceuticals.

"Currently, [68Ga]Ga-DOTATATE and [64Cu]Cu-DOTATATE are the two imaging agents that are most widely used in patients with neuroendocrine tumors," said Helge Thisgaard, Ph.D., associate professor in the Department of Nuclear Medicine at Odense University Hospital in Odense, Denmark. "However, there are limitations associated with both agents; particularly the short half-life of [68Ga]Ga-DOTATATE and the low positron yield of [64Cu]Cu-DOTATATE."

To address the shortcomings of these common imaging agents, researchers produced the non-conventional radiometal, 55Co, which has a relatively long half-life and high positron yield. After radiolabeling 55Co with DOTATATE, researchers evaluated the new PET-imaging



agent in comparison to [68Ga]Ga-DOTATATE and [64Cu]Cu-DOTATATE.

In the study, ten male mice were inoculated with neuroendocrine tumor cells. After the growth of a five millimeter tumor, the mice underwent PET/CT scanning with the different radiopharmaceuticals—four mice with [68Ga]Ga-DOTATATE, three with [64Cu]Cu-DOTATATE and three with [55Co]Co-DOTATATE. All mice were dynamically scanned within one hour after injection. Due to their longer half-lives, additional imaging was also performed at four hours and 24 hours after injection for the [64Cu]Cu-DOTATATE and [55Co]Co-DOTATATE and [55Co]Co-DOTATATE and [55Co]Co-DOTATATE mice.







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Data analysis of the PET/CT <u>images</u> revealed a rapid increase in tumor uptake within the first hour for all imaging agents. Normal-tissue ratios as represented by tumor-to-liver, tumor-to-kidney and tumor-to-muscle increased significantly over time, with [55Co]Co-DOTATATE showing the highest contrast. Most notably, the tumor-to-liver ratio of [55Co]Co-DOTATATE was 15 times and 30 times higher than the [64Cu]Cu-DOTATATE ratio at four hours and 24 hours, respectively, and was five times higher than [68Ga]Ga-DOTATATE at one hour. Furthermore, the tumor-to-kidney ratio at 24 hours for [55Co]Co-DOTATATE was four times higher than [64Cu]Cu-DOTATATE and eight times higher than [68Ga]Ga-DOTATATE at one hour.

"With [55Co]Co-DOTATATE, we found a highly increased <u>tumor</u> uptake relative to the uptake in healthy tissues, including the liver, which is a major site of metastasis of <u>neuroendocrine tumors</u>," said Thomas Lund Andersen, assistant professor in the Department of Nuclear Medicine at Odense University Hospital in Odense, Denmark. "These results warrant further translation into clinical practice and could lead to an improved detection of metastases, giving increased diagnostic confidence in ensuring the best possible patient care."



Looking toward the future of nuclear medicine and molecular imaging, Thisgaard believes this research has meaningful ramifications. "This study highlights the importance of optimizing not only the biological vector, but also selecting the most appropriate radionuclide to obtain the best imaging agent for a given indication. It illustrates that the use of promising, non-conventional PET isotopes may significantly improve the kinetics of the imaging agent, warranting further studies with novel combinations of biological vectors and new isotopes."

More information: Thomas L. Andersen et al, Improving Contrast and Detectability: Imaging with [55Co]Co-DOTATATE in Comparison with [64Cu]Cu-DOTATATE and [68Ga]Ga-DOTATATE, *Journal of Nuclear Medicine* (2019). DOI: 10.2967/jnumed.119.233015

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