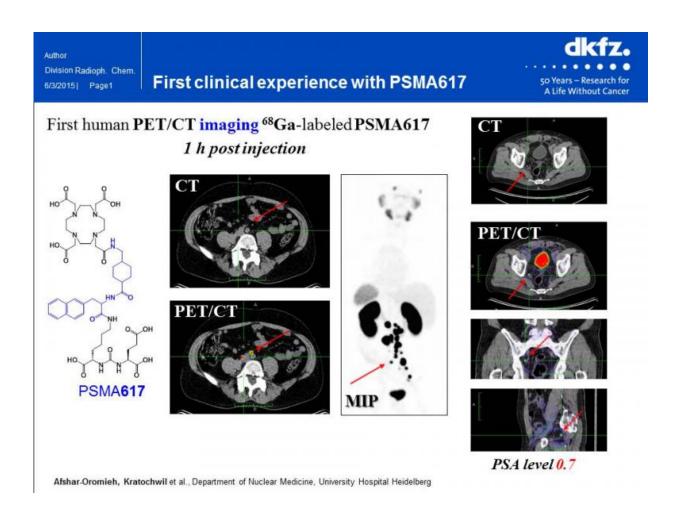


Theranostic drug personalizes prostate cancer imaging and therapy

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First human PET/CT imaging,GA-68-labeled PSMA-617,1 hour post injection. Credit: Afshar-Oromieh, Kratochwil, et al., Department of Nuclear Medicine, University Hospital Heidelberg



A novel radionuclide drug tackles the challenge of prostate cancer imaging and takes a turn as a cancer-killing therapy for tumors in and out of the prostate, according to research presented during the 2015 Annual Meeting of the Society of Nuclear Medicine and Molecular Imaging (SNMMI).

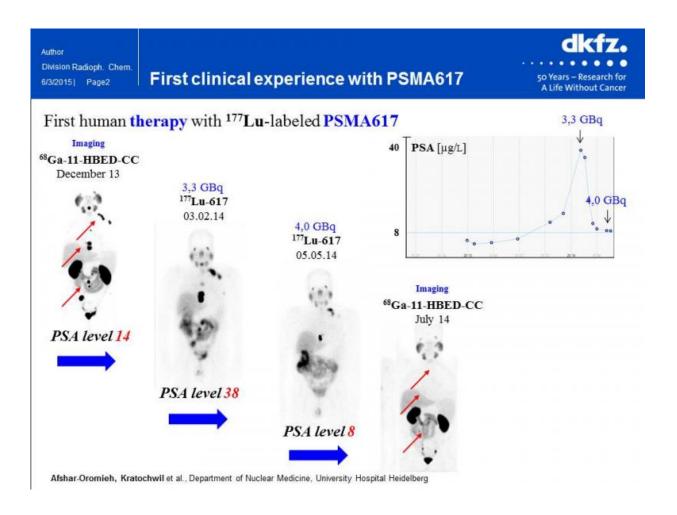
The drug works by delivering diagnostic- or therapy-grade radionuclides to cells that express a protein called prostate-specific membrane antigen (PSMA), found on the surface of prostate cancer cells and their metastatic counterparts throughout the body. Clinicians can diagnose or stage disease and monitor therapy with the aid of a special hybrid scanner used to perform minimally invasive positron emission tomography (PET).

"Prostate cancer still represents one of the main causes for cancerrelated deaths among men," said Matthias Eder, PhD, co-author of the
study and a researcher in the division of radiopharmaceutical chemistry
at the German Cancer Research Center in Heidelberg, Germany. "The
diagnosis and therapy of metastatic prostate cancer is still challenging.
The current clinical methods are not sensitive enough for detecting
disease beyond the prostate, but we are convinced that this novel
theranostic radiotracer represents a significant step forward that could
have a major impact on the future of prostate cancer care."

The PSMA-inhibiting theranostic agent, called PSMA-617, is still in its initial stages, but it could be ideal for the treatment of patients with hormone-refractory prostate cancers, which are notoriously difficult to control and linked to poor prognosis. Options for these patients are few, and they come with substantial adverse effects. Diagnosis and therapy with the theranostic agent PSMA-617 could offer more effective and sensitive visualization, better staging and significantly higher therapeutic potential. To be clear, other PSMA-based theranostics have reached the research bench, but previous contenders had too many limitations in



terms of instability in live subjects, a lack of imaging contrast between targeted tissue and background signal and increased PSMA-associated binding to normal organs such as the kidneys. This new radionuclide drug shows strong binding to the protein PSMA and is readily and safely taken up by malignant PSMA-positive tumors. PSMA-617 could represent a watershed moment for prostate cancer theranostics.



First human therapy with Lu-177-labeled PSMA-617. Credit: Afshar-Oromieh, Kratochwil, et al., Department of Nuclear Medicine, University Hospital Heidelberg



For this research, scientists first imaged mice with an imaging-grade radionuclide, gallium-68, to assess the diagnostic value of PSMA-617. Diagnostic imaging was followed by a therapy-grade radionuclide, lutetium-177, which delivers a more powerful dose of radiation that penetrates and destroys the cells and tissues of tumors when combined with PSMA-617. This phase was followed by a first-in-human clinical trial of both imaging and therapy in a single person.

Results of the human study showed that imaging was effective for the evaluation of metastatic prostate cancer, and subsequent therapy resulted in a drop in prostate-specific antigen levels from 38.0 to 4.6 nanograms per milliliter. Positive response to therapy was verified via combined PET and computed tomography (PET/CT), a hybrid imaging system that shows both functional and structural aspects of the body.

Approximately one out of seven men will be diagnosed with prostate cancer in their lifetime, according to 2015 statistics from the American Cancer Society. About 220,800 new prostate cancer diagnoses and 27,540 prostate-cancer-related deaths are expected to occur in the U.S. this year.

More information: Scientific Paper 63: "PSMA-617 - a novel theranostic PSMA inhibitor for both diagnosis and endoradiotherapy of prostate cancer"

Provided by Society of Nuclear Medicine

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