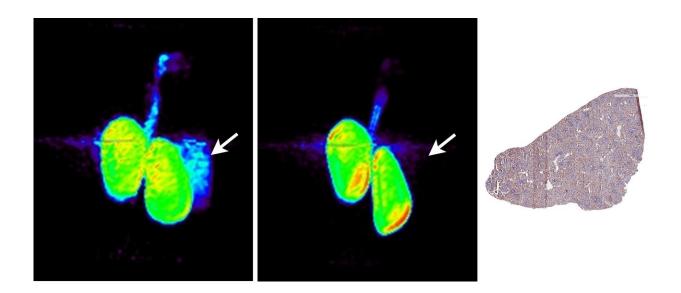


Researchers develop new approach that uses single PET scan to personalize cancer treatment

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Maximal intensity projection images of kidneys and spleen 90 minutes post injection of 18F-BMS-986192 only (left) and of 18F-BMS-986192 and coadministration of 1 mg/kg BMS-986192 in the same monkey (center). Representative anti-PD-L1 immunohistochemistry of healthy monkey spleen tissue (right) Credit: D Donnelly et al., Bristol-Myers Squibb Pharmaceutical Research and Development, Princeton, N.J.

Researchers have developed a same-day, noninvasive positron emission tomography (PET)-based imaging approach to assess PD-L1 positive tumors, and the study is presented in the featured article of *The Journal*



of Nuclear Medicine's March issue.

A healthy immune system strikes a delicate balance between eradicating infections and cancers and not overreacting to damage one's own tissue. Immune checkpoints help control the immune response, but tumors exploit these checkpoint pathways by expressing special proteins that evade antitumor immune responses. One major checkpoint inhibitor pathway is the PD-1 pathway, and its ligand is PD-L1.

In this study, the PD-L1 ligand, which enables cancer to evade a person's immune system, has been successfully targeted for the first time with a fluorine-18 (¹⁸F)-labeled PD-L1 radioligand. Until now, efforts to predict response to treatments targeting PD-1 or PD-L1 have typically been limited to evaluation of a single patient biopsy sample.

"This approach represents an opportunity for physicians to noninvasively assess all of a patient's tumors for PD-L1 expression with a single PET scan and timely readout," explains David J. Donnelly, PhD, at Bristol-Myers Squibb Research and Development in Princeton, New Jersey. "This may help guide treatment decisions and assess treatment response, to help identify the right <u>treatment</u> for the right patient at the right time and right dose."

For the study, an anti-PD-L1 adnectin (an engineered, target-binding protein) was labeled with ¹⁸F to generate ¹⁸F-BMS-986192, which was then evaluated in mice bearing bilateral PD-L1(-) and PD-L1(+) subcutaneous tumors. ¹⁸F-BMS-986192 was also evaluated for distribution, binding and radiation dosimetry in healthy cynomolgus monkey. The results of the study demonstrate the feasibility of the approach, and the radiation dosimetry estimates indicate that the tracer is safe to administer in human studies. Clinical studies are now underway to measure PD-L1 expression in human tumors.



More information: David J. Donnelly et al, Synthesis and Biologic Evaluation of a Novel18F-Labeled Adnectin as a PET Radioligand for Imaging PD-L1 Expression, *Journal of Nuclear Medicine* (2017). DOI: 10.2967/jnumed.117.199596

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