Letter to the Editor

In Regard to Trotter et al.

Loic Ah-Thiane, MD,^{a,*} Caroline Rousseau, MD, PhD,^{b,c} and Stéphane Supiot, MD, PhD^{d,e}

^aDepartment of Radiotherapy, ICO René Gauducheau, Boulevard Jacques Monod, St-Herblain, France; ^bDepartment of Nuclear Medicine, ICO René Gauducheau, St. Herblain, France; ^cCRCIbNA, UMR ac07 Inserm—UMR 6075 CNRS, Nantes University, Nantes, France; ^dDepartment of Radiotherapy, ICO René Gauducheau, Boulevard Jacques Monod, St. Herblain, France; and ^eCRCI2NA, Inserm UMR1232, CNRS ERL 6001, Nantes University, Nantes, France

Received 24 August 2023; accepted 30 August 2023

One of the first steps in the radiation therapy process is delineating targets to treat. However, this task may lack precision and be hardly reproducible. That is why Trotter et al wrote a review that suggested to include positron emission tomography (PET) imaging for treatment planning in radiotherapy.¹ Delineated volumes could then be modified compared with those based on the scanner alone. However, several issues persist and must be stressed: larger volumes may mean a better coverage of microscopic spreading, but also a wider margin overflowing healthy tissues; interobserver variations may decrease but not lead to changes in outcomes; different PET segmentation algorithms may lead to different volumes; PET benefits may vary according to treated locations. But this review has the merit to highlight the role of PET that could go beyond cancer diagnosis and staging. Image-based phenotypic precision medicine intends to offer therapeutic options for the majority of patients and diseases.² In prostate cancer, Prostate specific membrane antigen (PSMA)-PET was shown to easily identify the dominant lesion,³ enabling an accurate focal boost for a potentially better disease control.⁴ Another example would be mapping hypoxic areas in prostate cancer with [18F]Fluoromisonidazole (18F-FMISO) PET-CT to deliver dose-painting radiotherapy.⁵ RTEP5 is phase 2 trial that found improved overall survival due to a radiation therapy boost in hypoxic tumoral lung lesions identified by 18F-FMISO PET-CT.⁶ Based on a prospective study on phenotypic imaging in breast cancer with a theranostic approach,⁷ a pilot study evaluated immuno-PET using antibodies against carcinoembryonic antigen for a precise mapping and delineation of bone metastasis and showed its feasibility,⁸ paving the way for developing the PET-LINAC, a linear accelerator coupled with a PET-CT. Performing PET imaging could also be time-sparing for radiation oncologists. Indeed, 68Ga-DOTATE PET-CT for meningioma contributed to more precise volumes than magnetic resonance imaging⁹ and could then help semiautomatic delineation by using standard uptake value (SUV) thresholds.¹⁰ Because of the rich (and complex) data from PET imaging, more advanced models for automation can be developed, with for examples the GloD-LoATUNet and the mask-Net, which are deep-learning algorithms trained for automatic delineation of esophageal gross tumor volume and segmentation of non-small cell lung cancer in one second, respectively.^{11,12}

In a near future, PET imaging could become essential for treatment planning for all patients undergoing radiation therapy, and this review represents an inducement to design clinical trials evaluating the role of PET imaging for radiation oncologists, and to better define its place in the practice guidelines.

Disclosures

None to be declared regarding the present paper.

References

1. Trotter J, Pantel AR, Teo BK, et al. Positron emission tomography (PET)/computed tomography (CT) imaging in radiation therapy

https://doi.org/10.1016/j.adro.2023.101410

2452-1094/© 2023 The Authors. Published by Elsevier Inc. on behalf of American Society for Radiation Oncology. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).





www.advancesradonc.org

Sources of support: This work had no specific funding.

^{*}Corresponding author: Loic Ah-Thiane, MD; E-mail: Loic.Ah-Thiane@ico.unicancer.fr

treatment planning: A review of PET imaging tracers and methods to incorporate PET/CT. *Adv Radiat Oncol.* 2023;8:101212.

- Hawkey NM, Broderick A, George DJ, et al. The value of phenotypic precision medicine in prostate cancer. *Oncologist.* 2023;28: 93-104.
- Gaudreault M, Chang D, Hardcastle N, et al. Feasibility of biologyguided radiotherapy using PSMA-PET to boost to dominant intraprostatic tumour. *Clin Transl Radiat Oncol.* 2022;35:84-89.
- 4. Kerkmeijer LGW, Groen VH, Pos FJ, et al. Focal boost to the intraprostatic tumor in external beam radiotherapy for patients with localized prostate cancer: Results from the FLAME randomized phase III trial. *J Clin Oncol.* 2021;39:787-796.
- Supiot S, Rousseau C, Dore M, et al. Reoxygenation during radiotherapy in intermediate-risk prostate cancer. *Radiother Oncol.* 2019;133:16-19.
- **6**. Vera P, Mihailescu SD, Lequesne J, et al. Radiotherapy boost in patients with hypoxic lesions identified by 18F-FMISO PET/CT in non-small-cell lung carcinoma: Can we expect a better survival outcome without toxicity? [RTEP5 long-term follow-up]. *Eur J Nucl Med Mol Imaging*. 2019;46:1448-1456.

- Rousseau C, Goldenberg DM, Colombié M, et al. Initial clinical results of a novel immuno-PET theranostic probe in human epidermal growth factor receptor 2-negative breast cancer. J Nucl Med. 2020;61:1205-1211.
- 8. Pichon B, Rousseau C, Blanc-Lapierre A, et al. Targeting stereotactic body radiotherapy on metabolic PET- and immuno-PET-positive vertebral metastases. *Biomedicines*. 2020;8:548.
- **9.** Perlow HK, Siedow M, Gokun Y, et al. 68Ga-DOTATATE PET-based radiation contouring creates more precise radiation volumes for patients with meningioma. *Int J Radiat Oncol Biol Phys.* 2022;113:859-865.
- **10.** Kriwanek F, Ulbrich L, Lechner W, et al. Impact of SSTR PET on inter-observer variability of target delineation of meningioma and the possibility of using threshold-based segmentations in radiation oncology. *Cancers (Basel)*. 2022;14:4435.
- Yue Y, Li N, Zhang G, et al. Automatic segmentation of esophageal gross tumor volume in 18F-FDG PET/CT images via GloD-LoATU-Net. *Comput Methods Programs Biomed*. 2022;229:107266.
- **12.** Lei Y, Wang T, Jeong JJ, et al. Automated lung tumor delineation on positron emission tomography/computed tomography via a hybrid regional network. *Med Phys.* 2023;50:274-283.