1	
2	Ectopic, intra-thyroid parathyroid adenoma better visualised by deep
3	
4	learning enhanced choline PET/CT.
5	
6	Kevin M Bradley <sup>1</sup> , Daniel R McGowan <sup>2,3,</sup> , Lee Bartley <sup>4</sup> , David Scott-Coombes <sup>5</sup> , John I Rees <sup>4</sup>
7	
8	<sup>1</sup> Wales Research and Diagnostic PET Imaging Centre, Cardiff University, Cardiff, UK
9 10	<sup>2</sup> Department of Medical Physics and Clinical Engineering, Oxford University Hospitals NHS FT, Oxford, UK
10	<sup>3</sup> Department of Oncology, University of Oxford, Oxford, UK
12	<sup>4</sup> Department of Radiology, University Hospital of Wales, Cardiff, UK
12	<sup>5</sup> Department of Endocrine Surgery, University Hospital of Wales, Cardiff, UK
13	Department of Endocrine Surgery, oniversity hospital of Wales, cardin, ok
15	
16	Corresponding author
17	Daniel R McGowan, daniel.mcgowan@oncology.ox.ac.uk
18	Department of Oncology, University of Oxford, Oxford, UK
19	
20	
21	Imaging for the localisation of parathyroid adenomas is commonly
22	performed to permit minimally invasive parathyroidectomy and also to
23	
24 25	detect ectopic adenomas. A plethora of modalities, techniques and
23 26	radiotracers have been advocated, with over four thousand such imaging
27	publications on PubMed. However, there will always be a problem
28	
29	localising very small adenomas, particularly if 'ectopic'. Recently choline
30	PET/CT has emerged as a sensitive technique, with a 2023 meta-analysis
31	
32	of 1716 patients showing a pooled patient-based sensitivity of 93.8% [1].
33	The technique appears to be of particular benefit following failed surgical
34	exploration [2], regarded as the most difficult patients for which to provide
35 36	
30 37	a cure.
38	
39	A (0 year ald man with namiatant mimany hymamanathymaidian fallowing
40	A 60-year-old man with persistent primary hyperparathyroidism following
41	a failed surgical neck exploration, and persistent negative <sup>99m</sup> Tc-
42	SestaMIBI and 4D-CT scans, underwent <sup>18</sup> FluoroEthylCholine-PET/CT
43	
44	with arterial and portal phase iv contrast on a GE Omni Legend PET/CT
45	scanner. This revealed a sub-centimetre focus of increased uptake in the
46 47	right lower pole of the thyroid, more conspicuous on the deep learning
47 48	
49	enhanced PET reconstruction [3,4] (a, coronal PET, axial PET/CT, all
50	images SUV 0-6) than standard BSREM (Block Sequential Regularized
51	
52	Expectation Maximisation) PET reconstruction (b) with associated
53	increase in SUV <sub>max</sub> (maximum Standardised Uptake Value) from 3.1 to
54	4.5 Histopathology from surgical re-exploration with a right
EE	$\pm \beta$ , instruction we contribute that the harden with a term

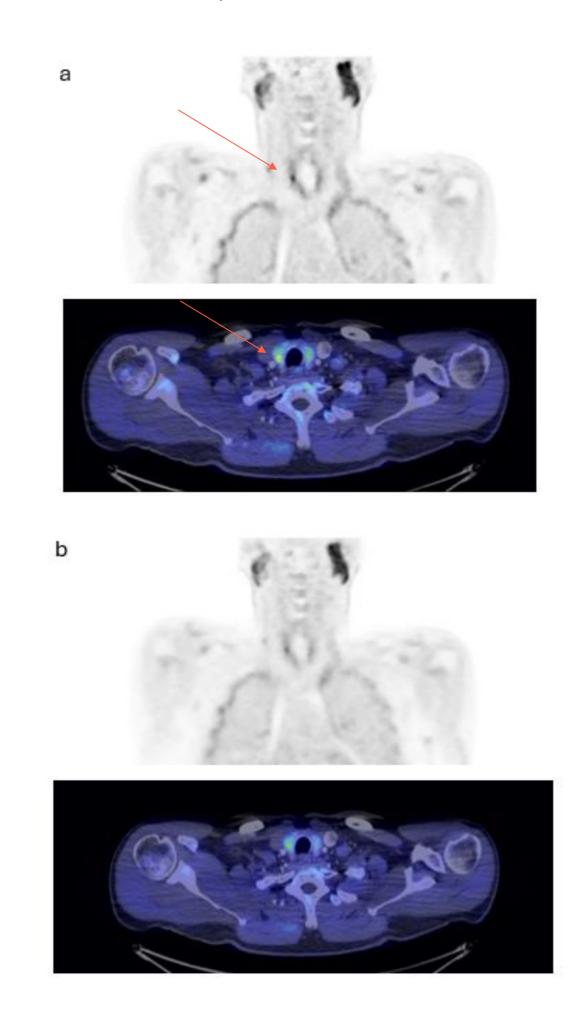
4.5. Histopathology from surgical re-exploration with a right hemithyroidectomy confirmed a 6mm parathyroid adenoma within the excised thyroid. Parathyroid hormone and calcium biochemistry were normalised at day 1, and 3 months, post-operatively.

This demonstrates the benefit of deep learning-based PET reconstruction image enhancement to reveal small foci of uptake. In this example, an elusive ectopic, intra-thyroidal parathyroid adenoma. Many such elusive adenomas are small and therefore challenging for imaging localisation. Intra-thyroidal, ectopic, parathyroid adenomas are particularly difficult to locate, suggested <1% of ectopic adenomas [5], and if completely embedded within the thyroid, are also invisible on surgical exploration, although some may be deep within a cleft or fold from the surface of the thyroid, described as a 'boutonnière' adenoma. This study also supports the European Association of Nuclear Medicine guidelines [6] which recommend that PET/CT localisation should be performed on a 'scanner with the highest system sensitivity and reconstruction protocols optimized for small lesion detection' and provides an insight into the benefits of current developments in PET image reconstruction.

## Learning points for clinicians:

- Choline PET/CT is a sensitive technique for localising parathyroid adenomas, particularly ectopic adenomas
- Ectopic parathyroid adenomas may rarely be intra-thyroid
- PET/CT image reconstruction can now be enhanced by deep learning, an AI (artificial intelligence) technique

https://mc.manuscriptcentral.com/qjm



## References

 1. Quak E, Cavarec M, Ciappuccini R, Girault G, Rouzier R, Lequesne J. *Detection, resection and cure: a systematic review and meta-analysis of 18F-choline PET in primary hyperparathyroidism.* Q J Nucl Med Mol Imaging. 2023 Jun;67(2):122-129.

2. Christakis I, Khan S, Sadler GP, Gleeson FV, Bradley KM, Mihai R. 18Fluorocholine PET/CT scanning with arterial phase-enhanced CT is useful for persistent/recurrent primary hyperparathyroidism: first UK case series results. Ann R Coll Surg Engl. 2019 Sep;101(7):501-507.

3. Mehranian A. Wollenweber SD, Walker MD, Bradley KM, Fielding PA, Huellner M, et al., *Deep learning-based time-of-flight (ToF) image enhancement of non-ToF PET scans*. European Journal of Nuclear Medicine and Molecular Imaging, 2022. **49**(11): p. 3740-3749.

4. Mehranian A. Wollenweber SD, Bradley KM, Fielding PA, Huellner M, Iagaru A, et al., *Deep learning Time-of-Flight (ToF) Enhancement of non-ToF PET scans for Different Radiotracers*. European Journal of Nuclear Medicine and Molecular Imaging, 2025 (in press)

5. Goodman A., Politz D, Lopez J, Norman J. *Intrathyroid Parathyroid Adenoma: Incidence and Location - The Case Against Thyroid Lobectomy*. Otolaryngol Head Neck Surg, 2011. **144**(6): p. 867-871

6. Ovcaricek PP, Giovanella L, Gasset IC, Hindie E, Huellner MW, Luster M., et al., *The EANM practice guidelines for parathyroid imaging*.
European Journal of Nuclear Medicine and Molecular Imaging, 2021. 48: p. 2801-2822.

## Acronyms

AI – artificial intelligence

BSREM – Block Sequential Regularized Expectation Maximisation

- CT computed tomography
- <sup>58</sup> PET positron emission tomography <sup>60</sup> SUV standardized untake solve
  - SUV standardized uptake value

60

