1 REPLY TO LETTER TO THE EDITOR: The importance of an adequate surgical 2 template during salvage lymph node dissection for node-recurrent prostate cancer 3 4 Authors: Wolfgang Peter Fendler¹, Andrea Farolfi^{1,2} 5 6 Institutions: 7 (1) Department of Nuclear Medicine, University Hospital Essen, Essen, Germany 8 (2) Nuclear Medicine Unit, IRCCS Azienda Ospedaliero-Universitaria di Bologna, Bologna, 9 Italy 10 11 Address for correspondence: 12 Prof. Dr. Wolfgang Fendler, Klinik für Nuklearmedizin, Universitätsklinikum Essen, 13 Hufelandstr. 55, 45147 Essen, Germany. Phone: +49 201 723 2033. Fax +49 201 723 5964. 14 Email address: wolfgang.fendler@uk-essen.de 15 16 Reply: 17 In line with the response by Bravi et al, PSMA PET demonstrated low sensitivity but high 18 specificity and negative predictive value for the detection of pelvic nodal metastases 19 compared to histopathology in a multicenter prospective phase III imaging study employing 20 blinded independent central reads (1). PSMA PET detection inversely correlates with size of 21 tumor deposits and thus PET is prone to miss micro-metastatic disease (2). Despite 22 underestimation on a single-lesion level (3), PSMA PET positivity raises a red flag for 23 diseased regions (4). As nodal spread follows lymph drainage anatomy, PSMA PET 24 guidance towards regions at risk seems feasible. The ongoing prospective ProsTone trial 25 (NCT04271579) will assess whether unilateral pelvic lymph node dissection on the PSMA 26 PET-positive side will lead to improved efficacy toxicity trade-off by sparing potentially 27 undiseased contralateral nodal regions. We agree with Bravi and colleagues suggesting the 28 adoption of an adequate template will be key to maximize the benefit for patients undergoing 29 local salvage therapy. In addition, standardized reporting of PSMA PET using a rationale 30 anatomic framework together with implementation into clinical trials on local therapy will be 31 key to define the future role of PSMA PET for treatment guidance (5). 32

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