

Point by Point Reviewer Responses

Reviewer comments are listed point by point below. Our response is listed in the bullet below with line numbers provided for you to quickly reference the exact location of the comment. Line numbers will be removed once you are satisfied with our point-by-point reviewer comment responses and the paper is ready for final submission.

REVIEWER 1 COMMENTS

- change “myriad” to variety
 - done
- change “moiety” to “part”
 - done
- delete “very”
 - done
- “Please bear that this is theory. When looking at tumor cell biology, there are multiple factors that influence cell death, for example bystander effect: Molecular mechanism of bystander effects and related abscopal/cohort effects in cancer therapy - PubMed (nih.gov)” with link to article.
 - Added line, “However, a variety of factors mediate tumor killing including bystander effect” with citation of the article provided by the reviewer. Thank you for this clarification.
- “Please mention higher nephrotoxicity with 90Y, see article of Konijnenberg et al. Radiation dose distribution in human kidneys by octreotides in peptide receptor radionuclide therapy - PubMed (nih.gov)” with link to article
 - Response: We are not sure we can definitively conclude, particularly from the article cited by the reviewer, that nephrotoxicity is *actually* higher, but rather that there is evidence the renal dose is higher which *could* lead to greater nephrotoxicity. However, we appreciate the point and feel it is important to include, so we added the line, “Renal dose is also higher than 177Lu which poses a higher risk of nephrotoxicity” with citation of the article provided by the reviewer.
- change “very few” to “rare”, and delete “potential”
 - done

REVIEWER 2 COMMENTS

Minor comments

- Abstract:
 - 1. In the word well differentiated, hyphen should be placed.
 - done
 - 2. In the last sentence, no need of comma after the word development
 - done
- Background:
 - 1. In the second line comma should be placed (“body, which”)
 - done
 - 2. In 2nd paragraph omit the “which is” (which is based on)
 - done
 - 3. In the last paragraph of the background, add comma (“in this paper,”)
 - done
- DOTATATE PET Imaging
 - 1. In first sentence replace the word “which” by “that”.
 - done
 - 2. Replace (The first imaging agents) by (The earlier imaging agents)
 - done
- Technique
 - 1. Replace (prior to) by (Before)
 - done
 - 2. Replace the word “same” by “exact”
 - done

- Normal Biodistribution
 - 1. Use the suitable article before unbound
 - done (replaced “of” with “for”)
 - 2. Remove comma after variable (variable,)
 - done
- Imaging Performance
 - 1. Replace “a very low” by “a deficient”
 - done
 - 2. Replace the sentence “Another cause of false negatives, as seen in all of PET imaging, may arise from small lesions that are below PET resolution” by “Another cause of false negatives, as seen in PET imaging, may arise from small lesions below PET resolution”
 - done
 - 3. Replace “extremely high” by “highly”
 - done – replaced “extremely high” with “highly avid” to make grammatically correct
 - 4. Replace the sentence “Very small lesions can be readily visualized in both phantom and patient clinical studies if the background activity is low” by “Small lesions can be readily visualized in phantom and patient clinical studies if the background activity is low”
 - done
- Role in Evaluation and Value in Management
 - 1. Replace “evaluation of” by “evaluating”
 - done
 - 2. “¹¹¹In pentetreotide and to CT or MR is well established” omit “to”
 - done
- ⁶⁴Cu DOTATATE
 - 1. Replace “compare to” by “than”
 - done
 - 2. Replace “contribute to improved” by “improve”
 - done
 - 3. Replace the sentence “Finally, pre-clinical studies of new investigational agents, such as ⁵⁵Co DOTATATE, are showing the potential for yet further improvements in tumor uptake and image contrast when compared to currently approved agents” by “Finally, preclinical studies of new investigational agents, such as ⁵⁵Co DOTATATE, show the potential for further improvements in tumor uptake and image contrast compared to currently approved agents”
 - done