

Answering Company editor-in-chief

English Language Certificate:

Dear Editor-in-Chief, Prof. Li Ma,

According to the journal's guidelines, when one of the authors is a native English speaker, there is no need to have a certificate of English editing.

We confirm that Olivia K. Heintz is a native American English speaker. She is currently affiliated to the University of Massachusetts – School of Medicine (USA). At the time of the first submission, she was moving from Harvard University (USA) to University of Massachusetts – School of Medicine (USA). She was born in the USA and resides at Boston, MA, USA. Considering the guidelines of the World Journal of Stem Cells, our understanding is that there is no need of English editing certificate.

Reference Citation Analysis:

By using the suggested tool "Reference Citation Analysis", the most relevant articles cited in the tool were already included in the first version of the manuscript.

Abbreviations

We checked all abbreviations and adjusted the manuscript.

Other comments

We adjusted the order of topics, format and subtitles as requested.

Copyright license agreement

The Copyright license agreement has been replied and attached.

All co-authors received the e-mail from WJSC requesting to fill a on-line agreement form with the exception of our co-authors Rubia Casagrande (rubiaca@uel.br) and Olivia K. Heintz (olivia.heintz@umassmed.edu). If the on-line agreement form must be completed by all authors, we would appreciate if you can send it again to these two co-authors (Rubia Casagrande [rubiaca@uel.br] and Olivia K. Heintz [olivia.heintz@umassmed.edu]).

Conflict-of-interest disclosure form

The Conflict-of-interest disclosure form has been replied and attached.

Answers to Reviewers

Answer to Reviewer #1

Question 1. I do not find the necessity of this review paper in the Introduction.

Answer 1: Thank for evaluating our manuscript and helpful input. We have added a justification for this review article in the introduction section.

Question 2. I believe it would be better if a figure was provided to interpret the pain mechanisms.

Answer 2: We added a new figure to illustrate the pain mechanisms as suggested. Please, see novel figure1.

Question 3. I do not understand the necessity of How can pain be quantified in animal tests section.

Answer 3: Our aim was to provide a review article that is complete instead of a review article that the reader must necessarily search for many other sources to fully understand it. In this sense, in our experience, not only readers that work in the pain field read the review articles on pain. Therefore, a brief explanation is helpful for the reader to fully understand all the nuances of the article. We prefer to maintain this section as explained above, however, if the Reviewer sees it as prejudicial to the manuscript, we can delete it.

Question 4. Please design a paragraph to summarize administration routes and dosages of stem cell.

Answer 4: We added a paragraph as suggested. Please, see the novel conclusion section. Updated Tables 2 and 3 as well as Figure 3 also bring this information.

Question 5. Please provide Non-Native Speakers of English Editing Certificate.

Answer 5: Thank you for the comment. One of the Authors of this manuscript is a native English speaker. According to the journal guidelines, there is no need of a certificate of English editing in this case. We confirm that the latest version of this manuscript was corrected by Olivia K. Heintz who is a native English speaker.

Answers to Reviewer #2:

Question 1: The review manuscript is excellent, containing a wide information. The contents may provide readers and investigators with good information. I recommend the manuscript should be published in WJSC, pending on checking the format of Reference list, and update some (unpublished) references.

Answer 1: Thank you for evaluating our manuscript and approval. We have checked the reference formatting and new references that were unpublished at the time of original submission were included in the R1 version of the manuscript.