

To Reviewers

Aviano (PN-Italy); 6<sup>th</sup> of august 2021

**Objective: response to manuscript comments**

Specific Comments to Authors:

The review was well organized, the topic was highly interesting but several recent published reviews focus on the same topic. The authors need to highlight their recent approach

Issues raised: 1) The manuscript needs to be revised again and to highlight what the authors want and correct the words, as there are many misspellings.

Summary of the Peer-Review Report: The authors need to highlight their recent approach.

Response to comments

Thank you for appreciating the review organization and the scientific topic.

As evidenced, other published reviews were focused on such issue. We admit that the rationale and the slant (i.e. the point of view) of the present manuscript, that distinguish this work from others, was not sufficiently explained in the first version of the text.

The manuscript was edited in response to your comments. On one side, we frequently underlined in the text the selection rationale of scientific information included in the review; approaches and publications were, in fact, chosen if translation to Good Manufacturing Practice compliant protocols was judged as concretely possible. In our opinion, these text amendments improved the overall quality of the work and contributed to differentiate it from previously published reviews.

On the other hand, to improve originality features of the work, we briefly improved chapters regarding our recently published approach aimed to the definition of a clinical grade and effective expansion protocol of mesenchymal stem cells.

We further polished the text from misspellings and typing mistakes; moreover, the manuscript underwent language editing process by a native English speaker (dr. Gonzalo Almanza; University of California San Diego). This finally improved the language quality of the manuscript.

All text modifications were highlighted and tracked in the resubmitted manuscript.

We thank the reviewer and the editor for their efforts in revising the manuscript, to ameliorate its overall scientific quality.

Kind regards.

F. Agostini (on behalf of all authors).