## RESPONSE TO THE REVIEWER'S COMMENTS

The Reviewer text has been copied and left in Black. The answers to questions, and the actions taken to adapt the text to the Reviewer comments have been written in blue in the next page.

Name of journal: World Journal of Diabetes

Manuscript NO: 62581

Title: Estrogens and the regulation of glucose metabolism

Reviewer's code: 03072151

**Position:** Editorial Board **Academic degree:** MD, MSc

**Professional title:** Associate Professor, Attending Doctor, Surgeon

Reviewer's Country/Territory: Taiwan

Author's Country/Territory: Spain

Manuscript submission date: 2021-01-22

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-01-27 13:49

Reviewer performed review: 2021-02-01 10:15

**Review time:** 4 Days and 20 Hours

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [ Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Language quality	[ ] Grade A: Priority publishing [ Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [ Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[Y]Yes []No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous  Conflicts-of-Interest: [] Yes [Y] No

## SPECIFIC COMMENTS TO AUTHORS

Dear editor and authors, The authors presented a detailed, balanced, and comprehensive narrative review to clarify Estrogens and its regulation of glucose metabolism. This is a very thorough review and has brought a lot of knowledge from this area into one review article. The paper is generally well-written and the overview given is fairly informative.

Thank you.

In addition to the changes suggested by the Reviewer, and following the instructions of the Science Editor, the text has been thoroughly revised, and a number of references from my research group (all signed by me) have been eliminated because the number of autocites was considered excessive. This has forced the slight modification of parts of the initial redaction and the use in some cases of more general (albeit more imprecise) references to support the text.

The major strength of the article is that the authors also review the mechanisms of action of Estrogens. Here are my suggestions to improve the quality of the manuscript and make this even a stronger paper.

Thank you

All titles are numbered. However, title 5 is missing in the manuscript. Please recheck if this is a numbering error or some paragraphs are missing.

No text is missing. This was only a careless numbering error for which I apologize.

The manuscript has a clear focus and is written well. In my opinion, it adds valuable information on the topic that can help readers a further knowledge of the hormone. I would suggest adding a "Preface" section in the beginning to be more comprehensive.

Thanks for the idea. The text has been extended, as suggested, with a more general introduction of what are and represent the estrogens. Thus, a new initial section has been appended to the manuscript:

1-The complex and growing implication of steroid hormones in homeostasis

The initial paragraphs of the first draft text (now section 2, retaining the initial name), and Boxes 1 and 4 have been slightly adapted to incorporate this expansion of the previous introduction.

Furthermore, it would be very interesting if the review continue some information about "Future applications and perspectives" before the "General considerations and conclusions" section.

Thanks, again for this suggestion. In fact, initially I did not dare to go along this path, ending the text with a final summary and conclusions. I thought along the lines you suggested and decided that this coda could be better presented, without breaking the linear discourse, by simply adding this text in a separate new "Box" (number 5).

## Box 5 Perspectives: the need for further advance of our knowledge of estrogens

This way, I maintained the linear text as initially designed, but I took the opportunity to extend the question of perspectives, a possible insight in the future of estrogens (essentially on metabolic control) and the needed clarification of the multiple roles of the known estrogens. I am happy you presented this idea, I was aware that something was lacking, but now I believe that this addition to the manuscript completed the vision.

## I cannot see Table 1 neither Boxes 1-4 in the word file submitted.

I am sorry the Reviewer had not access to Table 1, since —in my opinion—this is a summary of the final classification-discussion already discussed in the text.

I hope that you will be able to read all in the re-review process.

Thank you very much for your comments and insight; your ideas helped me to improve (I hope) the text, irrespective of its final acceptance for publication.