

Santiago, May 25th, 2017

Dr.
Yuan Qi
Science Editor
World Journal of Gastroenterology

Dear Dr. Qi:

Thank you for all the comments that the reviewers made to the document, which enrich and facilitate the understanding of this review. I would like to explain and comment regarding the incorporation of the changes suggested by the reviewers of the review Glucocorticoid Therapy in Inflammatory Bowel Diseases: from Clinical Practice to Molecular Biology, Manuscript N° 33686.

Reviewer's code: 03413692

"I would include some epidemiological data to confirm the lesser rate of side-effects that 2nd generation glucocorticoids have vs. 1st generation"

We included a relative risk of adverse effects of 2dn generation of GCs compared to conventional corticosteroid therapy that was published in a pooled analysis that used the Cochrane database. Please find this change in the section titled **Budesonide, similar effectiveness with less adverse effects.**

"When comparing BDP with PP, last sentence is confusing because the article in which BDP is less effective is not mentioned at all"

We apologize for this mistake, because according to the reference, BDP has similar effect to PP treatment. We corrected the phrase:

This evidence leads to the conclusion that BDP similar or less effective than traditional and other second generation GCs.

To:

This evidence leads to the conclusion that BDP has similar effect to the traditional GCs treatment.

"Before conclusion it would be great if you could include some thoughts about the role of epigenetics in GC dependence (see Harris RA et al. Epigenetics. 2013 Feb;8(2):157-63)"

We thanks for this suggestion because this improves our review considerably. Before the conclusion, we incorporated paragraph about epigenetics in GC resistance and dependence in IBD, which includes the suggested paper.

Reviewer's code 3413692

"The English of the text should be revised"

We appreciate the suggestion of the reviewer. The English language was reviewed and revised exhaustively by Payton Ottum to make improvements throughout the manuscript. We hope that you find these changes sufficient to approve the language evaluation.

"The authors should include a table gathering the main findings from literature"

We want to thank the suggestion of the reviewer because this table helps to summarize and increase the consolidation of the evidence described. We include this information as Table N° 2 Molecular highlights on response to steroidal treatment.

Please consider and discuss the paper from Principi M et al. J Crohns Colitis. 2013 Nov;7(10):e427-33.

We included the paper suggested in the section titled *First generation GCs: more effective than 5-asa and immunomodulatory treatment with significant secondary effects.*

We hope that these incorporations improve the quality of our review and are favorably accepted by the reviewers.

Regards,



Marcela Hermoso Ramello
Corresponding Author