

Reviewer #1:

1. In the whole manuscript, please replace flora with microbiota.

Answer: Yes, I have already replaced all.

2. Abstract: Methods: please insert that you used forty mice.

Answer: Yes, I have inserted.

3. Key words are missing. Please insert relevant words.

Answer: Thank you for your warm reminder. I have inserted the key words.

4. Core Tip: This does not exist. It is mandatory. Please write it and insert it.

Answer: Thank you for your warm reminder. I have inserted the Core Tips.

5. Introduction: Clearly states the importance of this research; just please define (by the end of Introduction) the aim of your study, not what you performed during your research.

Answer: Thank you for your advice. I revised the last part of introduction, as following. Based on the above information and our previous study, we aimed to explore the mechanism of *B. infantis* on PD-1/PD-L1 signaling and the differentiation and function of Tregs.

6. Materials and Methods should be one paragraph, with multiple subparagraphs. Also, in "Materials": Please mention in a sentence that you used those materials, instead of a list.

Answer: I have corrected it according to your request.

7. Title of Table 1 is missing.

Answer: I have added the title.

8. Figure 5: *Bifidobacterium infantis* appears abbreviated as B9. Please mention it.

Answer: Thank you for your warm reminder. I have made the correct changes of Figure 5.

9. Paragraph Discussion should be expanded, with references from 2021. Please insert limitations of your study.

Answer: I accept your opinion with pleasure. I added three references from 2021 and inserted limitations in the discussion.

10. Please provide definite perspectives for future research.

Answer: I have added the definite perspectives in the end of the article.

11. References – there is none from 2021. Please update.

Answer: I add three references from 2021.

12. Please provide a list of abbreviations.

Answer: I have provided in the Supplementary Material.

13. There are no « Conflict-of-Interest Disclosure Form » and « Copyright License Agreement ». Please add.

Answer: I am sorry! I have uploaded « Conflict-of-Interest Disclosure Form » and « Copyright License Agreement ».

14. The overall format of the WJG is not respected, including references. Please revise.

Answer: I have revised the format as required.

15. Please also add authors' ORCID numbers.

Answer: I have added all the authors' ORCID numbers.

Reviewer #2:

Specific Comments to Authors: The title reflects well the main subject of the manuscript.

The abstract is well written and summarizes the work described in the manuscript. The manuscript adequately describes the background, present status and significance of the study. The manuscript describes methods in adequate detail. The research objectives were achieved by the experiments used in this study. And I think that this study contributed to increase our knowledge about IBD and the role of the microenvironment. The manuscript interprets the findings adequately and appropriately. The findings are important and help to understand the interaction between the immune function of the intestinal mucosa and the intestinal flora of an affected animal. The figures, diagrams and tables are sufficient and gave good quality. The manuscript is coherently organized and presented. But the authors forgot the Key words.

Answer: Thank you for your warm reminder. I have inserted the key words.

Reviewer #3:

Specific Comments to Authors: Thank you for valuable researching, I only suppose to add more references about Bifidobacterium's beneficial effects on immune system in discussion part [39]Bozkurt, H.S.; Quigley, E.M.M. Bifidobacteria and Mucosal-Associated Invariant T (MAIT) Cells: A New Approach to Colorectal Cancer Prevention? *Gastrointest. Disord.* 2019, 1, 266-272. [40]Bozkurt HS, Kara B. A new treatment for ulcerative colitis: Intracolonic Bifidobacterium and xyloglucan application. *European Journal of Inflammation*. January 2020 [41]Bozkurt HS, Quigley EM. The probiotic Bifidobacterium in the management of Coronavirus: A theoretical basis. *Int J Immunopathol Pharmacol.* 2020 Jan-Dec. [42]Bozkurt HS, Bilen Ö. Oral booster probiotic bifidobacteria in SARS-COV-2 patients. *International Journal of Immunopathology and Pharmacology*. January 2021 Again, congratulation for this working

Answer: I have inserted « A new treatment for ulcerative colitis: Intracolonic Bifidobacterium and xyloglucan application » in the discussion. The other references were about cancer and coronavirus.