



## PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 64869

**Title:** Microbiota shaping – the effects of probiotics, prebiotics and fecal microbiota transplant on cognitive functions: A systematic review

**Reviewer’s code:** 05232520

**Position:** Editorial Board

**Academic degree:** BSc, MSc, PhD

**Professional title:** Professor, Research Scientist

**Reviewer’s Country/Territory:** Brazil

**Author’s Country/Territory:** Italy

**Manuscript submission date:** 2021-02-25

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2021-02-25 11:56

**Reviewer performed review:** 2021-03-11 19:17

**Review time:** 14 Days and 7 Hours

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Re-review</b>	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
<b>Peer-reviewer statements</b>	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No



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## **SPECIFIC COMMENTS TO AUTHORS**

The manuscript submitted by Baldi and cols., aims to review the role of gut microbiota modulation by different approaches and its effects on cognitive functions. However, before a final decision is made, some aspects must be addressed. Major concerns:

- Authors must clarify if the review refers to cognitive disorders (in general) or specifically about dementia.
- In introduction, authors stated that "...the administration in animal models of an adequate posology of multistrain probiotics reduced both Firmicutes/Bacteroidetes ratio and the intestinal permeability...", however, there are some clinical trials addressing the role of such approach in gut microbiota modulation.
- In page 11, please clarify FMT studies T?
- As *Lactococcus* spp may also be used as probiotics in different contexts, the species of microorganisms used in the review must be written with their full names by the first time they are cited in the text
- Authors must describe the putative mechanisms used by probiotics to modulate the activity of central nervous system
- In its current form the manuscript is merely descriptive. The text lacks a more in-depth discussion concerning pros and cons of using microbial modulation to treat or constrain the development of cognitive disorders
- Authors must describe the limitations of each approach highlighting the concerns regarding the use of live microorganisms (probiotics and FMT) in targeted populations, including immunossupressed and the risk of FMT in the transmission of pathogenic multi-resistant bacteria.

Minor concerns:

- There are several typos throughout the text. E.g. "synbiotics" ; "week" – page 11
- Please, replace the word "flora" by "microbiota"
- "mL" rather than "ml"



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**Reviewer’s code:** 03061187

**Position:** Peer Reviewer

**Academic degree:** PhD

**Professional title:** Professor

**Reviewer’s Country/Territory:** Mexico

**Author’s Country/Territory:** Italy

**Manuscript submission date:** 2021-02-25

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2021-03-01 17:03

**Reviewer performed review:** 2021-03-12 07:01

**Review time:** 10 Days and 13 Hours

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Re-review</b>	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No
<b>Peer-reviewer statements</b>	Peer-Review: <input type="checkbox"/> Anonymous <input checked="" type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No



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#### **SPECIFIC COMMENTS TO AUTHORS**

The text entitled “Microbiota shaping: the effects of probiotics, prebiotics and fecal microbiota transplants on cognitive functions. A systematic review” is a nice work prepared by methodic review of the state-of-the-art of important cognitive disorders and its possible treatment by modification of the gut microbiota through the administration of probiotics, prebiotics, synbiotics or even fecal microbiota transplants. The information is clearly presented by the authors, updating the knowledge to the reader in this field of knowledge. Without decreasing the importance of this work, to the eyes of this reviewer, the work lacks interesting information about the use of nutritional interventions used similarly to treat deficiencies in the cognitive functions. There are some suggestions to improve the work, made directly to the reviewed text (attached: ca857959-9f4c-4ea4-a2f0-c597d4071f70\20210312\_64869-Manuscript File-reviewed.pdf) which are highlighted in magenta color.



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**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 64869

**Title:** Microbiota shaping – the effects of probiotics, prebiotics and fecal microbiota transplant on cognitive functions: A systematic review

**Reviewer's code:** 04929451

**Position:** Peer Reviewer

**Academic degree:** MD

**Professional title:** Doctor

**Reviewer's Country/Territory:** China

**Author's Country/Territory:** Italy

**Manuscript submission date:** 2021-02-25

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2021-03-01 07:17

**Reviewer performed review:** 2021-03-19 06:18

**Review time:** 17 Days and 23 Hours

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
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#### **SPECIFIC COMMENTS TO AUTHORS**

Simone Baldi and colleagues inspect the RCT research reports to review the influences of gut microbiota interventions including probiotics, prebiotics, and FMT on cognitive functions. This PRISMA study is very meaningful for the better understanding about gut-brain axis and useful for the pathogenesis study relate to dementia and other brain diseases. This study overall is of good quality and of interest to the journal's audience. Minor comments, The differential effects caused by the different gut microbiota intervention methods needs to be compared strictly in RESULTS. The details of the mechanism in these reports need more in-depth analysis and summary. The uncertainties and risks associated with microbiota intervention, especially FMT, need to be analysed and discussed.



## RE-REVIEW REPORT OF REVISED MANUSCRIPT

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**Reviewer's code:** 05232520

**Position:** Editorial Board

**Academic degree:** BSc, MSc, PhD

**Professional title:** Professor, Research Scientist

**Reviewer's Country/Territory:** Brazil

**Author's Country/Territory:** Italy

**Manuscript submission date:** 2021-02-25

**Reviewer chosen by:** Chen-Chen Gao

**Reviewer accepted review:** 2021-07-19 13:25

**Reviewer performed review:** 2021-07-19 13:30

**Review time:** 1 Hour

<b>Scientific quality</b>	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
<b>Language quality</b>	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
<b>Conclusion</b>	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
<b>Peer-reviewer statements</b>	Peer-Review: <input type="checkbox"/> Anonymous <input checked="" type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

### SPECIFIC COMMENTS TO AUTHORS



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The authors have satisfactorily responded to all my questions