

## ***Response to Review's comments:***

### ***(1) Science editor:***

Scientific quality: The manuscript was a good review that presents the usual risk factors for colorectal cancer in the context of early-onset disease. The topic is within the World Journal of Gastrointestinal Oncology. (1) Classification: one Grade B. (2) Summary of the Peer-Review Reports: The author gathers new insights on the development of early-onset colorectal cancer (EOCRC) based on current evidence. Quality of scientific data included in the paper is very good. Key messages are concisely explained without delving much into details of each study cited. With the exception of genetic features, all other risk factors are proposed to be related to the increased incidence of EOCRC but the discussion cites the lack of a clear causative link between them and EOCRC. The paper recognizes the limitations of the studies mentioned, and suggests the need for further research on these topics. Despite several gaps in research on EOCRC risk factors, the review mentions ways to mitigate these risks in young individuals in order to stem the rise in numbers of patients with EOCRC based on current knowledge.

Response: We would like to thank the Reviewer for the constructive comments on our review article.

But the one minor correction would just involve the names of bacterial organisms in the section on the gut microbiome. Genus name should be capitalized and the species name should be in small letters; the whole scientific name should be italicized from a few studies, some hold the opinion that severity of liver cirrhosis should be highlighted in clinical practice.

Response: All the bacterial organisms scientific names have now been amended on the manuscript.

The questions raised by the reviewers should be answered. (3) Format: There is 1 figure. (4) References: A total of 96 references are cited, including 47 references published in the last 3 years; Self-cited references: There is 1 self-cited references. The self-referencing rate should be less than 10%. Please keep the reasonable self-citations (i.e., those which are most closely related to the topic of the manuscript) and remove all other improper self-citations. If the authors fail to address the critical issue of self-citation, the editing process of this manuscript will be terminated; (6) References recommendations: 1) The authors have the right to refuse to cite improper references recommended by the peer reviewer(s), especially references published by the peer reviewer(s) him/herself (themselves). If the authors find the peer reviewer(s) request for the authors to cite improper references published by him/herself (themselves), please send the peer reviewer's ID number to [editorialoffice@wjgnet.com](mailto:editorialoffice@wjgnet.com). The Editorial Office will close and remove the peer reviewer from the F6Publishing system immediately.

Response: Only 1 self-cited reference is in the review article, which is less than 10% of total references cited.

2) Language evaluation: Classification: One Grade B. 3) Academic norms and rules: No academic misconduct was found in the search. No academic misconduct was found in the Google/Bing search. 4) Supplementary comments: This is an invited Manuscript. The study was not supported by any foundation. The topic has not previously been published in the World Journal of Gastrointestinal Oncology. 5) Issues raised: (1) The sections of abstract, keywords, Author Contributions and the statement of conflict of interest are missing in main text. Please provide them. (2) The formats of references and cited in main text should be modified according to the guideline for authors.

Response: Abstract, keywords and statement of conflict of interest are now added to the manuscript. Formats of references have now been modified according to guideline.

(3) The context of the file named Non-Native Speakers of English Editing Certificate is not the certificate. 6) Recommendation: Conditional Acceptance.

Response: Both authors of this article are native English speakers.

***(2) Company editor-in-chief:***

I have reviewed the Peer-Review Report, the full text of the manuscript, and the relevant ethics documents, all of which have met the basic publishing requirements of the World Journal of Gastrointestinal Oncology, and the manuscript is conditionally accepted. I have sent the manuscript to the author(s) for its revision according to the Peer-Review Report, Editorial Office's comments and the Criteria for Manuscript Revision by Authors. Before final acceptance, uniform presentation should be used for figures showing the same or similar contents; for example, "Figure 1 Pathological changes of atrophic gastritis after treatment. A: ...; B: ...; C: ...; D: ...; E: ...; F: ...; G: ...".