

Professor Monjur Ahmed,
Editors-in-Chief,
World Journal of Gastrointestinal Oncology

Dear Prof. Monjur Ahmed:

We are very grateful for giving us the opportunity to submit our revised manuscript (Manuscript ID:64713) for publication in *World Journal of Gastrointestinal Oncology*.

In this revised manuscript (Manuscript ID:64713), we added scheme figures of ncRNA, summary table and clinical section. Moreover, we carefully revised our manuscript to make it more compact and complete. We believe that the review content in this paper is of great interest to the broad audience of *World Journal of Gastrointestinal Oncology*.

The responses to reviewers are shown as below.

Thank you very much for your kind consideration.

Best regards,

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Response to reviewers

World Journal of Gastrointestinal Oncology

Manuscript ID:64713

Title: The role of regulatory RNAs, microRNA, long-non coding RNA and circular RNA in colorectal cancer stem cells

Reviewer #1:

Scientific Quality: Grade C (Good)

Language Quality: Grade B (Minor language polishing)

Conclusion: Minor revisionSpecific

Comments to Authors:

1. This article summarized recent researches on the role of microRNA, long-non coding RNA and

circular RNA in colon cancer stem cells.

2. It would be better if there were some the opinion of author for the three kind of RNA
3. The author should point out whether targeted RNA therapy has limitations in clinical applications

■ **Response:** Thank you for your comment. We added a section to address the clinical applications of ncRNA and our opinions.

Reviewer #2:

Scientific Quality: Grade B (Very good)

Language Quality: Grade A (Priority publishing)

Conclusion: Minor revision

Specific Comments to Authors: In this paper, the authors have comprehensively described in detail the regulation of regulatory RNAs in colorectal carcinoma stem cells and the related mechanisms, including miRNA, lncRNA, and circRNA. The article is well organized, logical, and informative, with some innovation, and no previous review was conducted to summarize this topic, and the language was also embellished. Suggested for acceptance after proper revision. The authors need to address the following issues.

1. I suggested that the authors may add a section on the use of ncRNA in colorectal cancer (treatment) and the related clinical trial, as well as the corresponding Challenges and Prospection.

■ **Response:** Thank you for your suggestion. We added a Challenges and Prospection section to address the clinical applications of ncRNA.

2. It is suggested that the authors could add a Table listing the relevant ncRNAs and their functions, targets, and references, so that readers can quickly identify the ncRNAs of interest and find the corresponding literature.

■ **Response:** Thank you for your comment. We added a Summary Table (Table 1) in our manuscript.

3. In the previous literature, there has been studies on colorectal cancer related miRNA mechanism diagrams, the authors may consider adding lncRNA and circRNA related mechanism diagrams to increase the readability of the article and present the related mechanism to the readers well.

■ **Response:** Thank you for your comment. We added scheme figures of lncRNA and circRNA related mechanisms in our manuscript.

4. The author may need to update the literature while revising the manuscript, including some of the latest related research in each part (2021/1-2021/6) Such as: Identification and Validation of New Cancer Stem Cell-Related Genes and Their Regulatory microRNAs in Colorectal Cancerogenesis. Stem cells as therapeutic targets in colorectal cancer.....

■ **Response:** Thank you for the suggestion. We added the latest research (2021/1-2021/7) of miRNA and lncRNA in our manuscript.

5. “designation of co-first authors and/or co-corresponding authors is not permitted.” This rule of BPG should be considered.

■ **Response:** Thank you for the suggestion. We revised this part.