

ANSWERING REVIEWERS

June 5, 2018

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 38929-revised.docx).

Title: The emerging role of long non-coding RNA in the development of gastric cancer

Author: Hang Yu, Long Rong

Name of Journal: World Journal of Gastroenterology oncology

Manuscript NO: 38929

The manuscript has been improved according to the suggestions of reviewers:

1. Format has been updated and spelling mistakes been corrected.
2. Revision has been made according to the suggestions of the reviewers

(1) Reviewer1 (03009243)

1) *This review does not contain any figures or tables. The deficit makes readers difficult to understand the overall concepts or information provided by the authors.* 2) *The author must explain the molecular mechanism of action of lncRNA in more detail. The authors just mentioned chromatin modification, transcriptional regulation, and post-transcriptional regulation as the molecular mechanism. However, other mechanism still exists, such as absorption of micro RNA (sponge effect), precursor for micro RNA, or regulation of splicing. The authors must include additional mechanism.* 3) *The authors cited HOTAIR, H19, GAS5, MEG3, LINC00152, UCA1, and MALAT1 as representative lncRNAs contributing pathogenesis of gastric cancer. However, the function of each lncRNA has not explained in the same fashion. For some, detailed molecular mechanisms have been described, but not for others.* 4) *The description in the literature is not accurate. There are sentences describing contents without citing the literature. The first sentence in the section of 'H19', "LncRNA H19, discovered in 1991 by Bartolomei, was the first imprinted lncRNA gene identified." must cite 'Bartolomei MS et al. Nature, 351: 153-5, 1991.'* 5) *There is no uniformity in the use of abbreviations. For instance, RNA polymerase II was abbreviated to RNAP II,*

but the word again appeared without abbreviation. There is also a mistake in description of the name of the author. The first author of the reference number 48 is Endo H., but not Hiroyuki E.

Authors respond:

1) We added a table that summarized the lncRNAs associated with gastric cancer mentioned in this review and the main biological roles of them. We hoped that this table would make it easier for readers to understand the deregulation of lncRNAs in gastric cancer. 2) Besides chromatin modification, transcriptional regulation and post-transcriptional regulation as the molecular mechanisms, we added other mechanisms including sponge effect, regulation of splicing and precursor for micro RNA, which could be seen in the highlighted paragraphs. 3) In this review, we analyzed some of the well-studied lncRNAs deregulated in gastric cancer. However, as the specific function mechanisms of lncRNAs had not been fully elucidated, it was hard to explain the mechanisms of each lncRNA in exactly the same fashion. We hoped that in the future, more related studies would be held and explore the function mechanisms in more details. 4) The citing literature of H19 had been added in the reference. The first author of some reference had been corrected. 5) The abbreviation of words had been unified.

(2) Reviewer 2 (02446277)

*May be improved with additional data as follows: (1) Because in the literature there are mentioned more long non-coding RNA than described in the manuscript (see *Oncotarget* 2017, 8(46): 81572–81582; *Frontiers in Laboratory Medicine* 2017, 1(2): 59-68), I suggest the authors to include one table to list them, specifying the function, up/down-regulation in gastric cancer, citation, etc. (2) Another suggestion is to include data about the lncRNA association with prognosis (*Frontiers in Laboratory Medicine* 2017, 1(2): 59-68). (3) A minor observation: in *Category of lncRNAs* section, page 3, the letter “s” is missing in the construct: “only a mall proportion”*

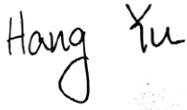
Authors respond:

1) We added a table to list the lncRNAs mentioned in this review and their up/down-regulation, biological role, citations as the reviewer suggested. 2) In the description of some of the lncRNAs, we included the prognosis data. However, the prognosis data of some other lncRNAs were not available. 3) We corrected the spelling mistakes mentioned.

3. References and typesetting were corrected.

Thank you again for publishing our manuscript in the World Journal of Gastroenterology
oncology.

Sincerely yours,

A handwritten signature in black ink that reads "Hang Yu". The signature is written in a cursive style with a loop at the end of the "Y".

Hang Yu

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