

Response to comments:

Reviewer #1:

Scientific Quality: Grade C (Good)

Language Quality: Grade B (Minor language polishing)

Conclusion: Minor revision

Specific Comments to Authors: In this review, the author introduce diagnosis, chemotherapy and molecular landscape of HCC, and also Epigenetics and its potential role in HCC. However, part of description requires further improvement. 1.Primary liver cancer is mainly divided into hepatocellular carcinoma (HCC), intrahepatic cholangiocarcinoma (ICC), and HCC-ICC according to pathological type. These three types vary greatly in pathogenesis, biological behaviors, histological morphology, treatment methods, and prognosis. HCC accounts for 85–90% of all cases of primary liver cancer. Thus at the beginning, need to defined that the liver cancer described in this review is HCC. 2.Both lenvatinib and sorafenib, are used as standard-of-care chemotherapeutics in patients with advanced HCC, the study of lenvatinib should be described. 3.In the section of the Molecular landscape of HCC, more recently studies should be summarized. Jiang et.al used quantitative proteomic data stratify the early-stage hepatocellular carcinoma into the subtypes S-I, S-II and S-III, each of which has a different clinical outcome. (PMID: 30814741, DOI: 10.1038/s41586-019-0987-8); Gao et.al performed proteogenomic characterization of HCC identifies three subgroups with distinct features in metabolic reprogramming, microenvironment dysregulation, cell proliferation, and potential therapeutics. (PMID: 31585088, DOI: 10.1016/j.cell.2019.08.052). These two studies should be cited and described.

Response:

We appreciate the comments given by reviewer. We accept all suggestions and have updated the text at line # 58-62, 162-169, and 220-229 in manuscript.

LANGUAGE QUALITY

Please resolve all language issues in the manuscript based on the peer review report. Please be sure to have a native-English speaker edit the manuscript for grammar, sentence structure, word usage, spelling, capitalization, punctuation, format, and general readability, so that the manuscript's language will meet our direct publishing needs.

Response:

Entire manuscript has been subjected to Grammarly professional account to resolve all language issues and certificate is uploaded for the same. Further, we have had the manuscript reviewed by a native English speaker and hope that it is satisfactory.

5 EDITORIAL OFFICE'S COMMENTS

Authors must revise the manuscript according to the Editorial Office's comments and suggestions, which are listed below:

(1) Science editor: 1 Scientific quality: The manuscript describes a Review of the epigenetics and its potential role in HCC. The topic is within the scope of the WJH. (1) Classification: Grade C; (2) Summary of the Peer-Review Report: The part of description requires further improvement. In the section of the Molecular landscape of HCC, more recently studies should be summarized. The

questions raised by the reviewers should be answered; (3) Format: There are 2 tables and 5 figures; (4) References: A total of 103 references are cited, including 19 references published in the last 3 years; (5) Self-cited references: There are 2 self-cited references; and (6) References recommendations (kindly remind): 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. The Editorial Office will close and remove the peer reviewer from the F6Publishing system immediately.

Response:

We have updated the manuscript and references as per suggestion.

2 Language evaluation: Classification: Grade B.

Response:

No comments.

3 Academic norms and rules: No academic misconduct was found in the Bing search.

Response:

No comments.

4 Supplementary comments: This is an invited manuscript. No financial support was obtained for the study. The topic has not previously been published in the WJH.

Response:

No comments.

5 Issues raised: (1) The language classification is Grade B. Please visit the following website for the professional English language editing companies we recommend: <https://www.wjgnet.com/bpg/gerinfo/240>; (2) The authors did not provide original pictures. Please provide the original figure documents. Please prepare and arrange the figures using PowerPoint to ensure that all graphs or arrows or text portions can be reprocessed by the editor; and (3) If an author of a submission is re-using a figure or figures published elsewhere, or that is copyrighted, the author must provide documentation that the previous publisher or copyright holder has given permission for the figure to be re-published; and correctly indicating the reference source and copyrights. For example, "Figure 1 Histopathological examination by hematoxylin-eosin staining (200 ×). A: Control group; B: Model group; C: Pioglitazone hydrochloride group; D: Chinese herbal medicine group. Citation: Yang JM, Sun Y, Wang M, Zhang XL, Zhang SJ, Gao YS, Chen L, Wu MY, Zhou L, Zhou YM, Wang Y, Zheng FJ, Li YH. Regulatory effect of a Chinese herbal medicine formula on non-alcoholic fatty liver disease. *World J Gastroenterol* 2019; 25(34): 5105-5119. Copyright ©The Author(s) 2019. Published by Baishideng Publishing Group Inc[6]". And please cite the reference source in the references list. If the author fails to properly cite the published or copyrighted picture(s) or table(s) as described above, he/she will be subject to withdrawal of the article from BPG publications and may even be held liable.

Response:

The manuscript has been screened using Grammarly professional version. Also, the certificate for the same is uploaded. Further, we have had the manuscript reviewed by a native English speaker and hope that it is satisfactory. All figures given in manuscript are original and generated by the authors. We are providing original figures using PowerPoint Presentation file. However, we have used Biorender tool for figure 5. The output provided by this tool is a PDF file. We have tried our best to decompose the image but only text part is editable.

6 Recommendation: Conditional acceptance.

Response:

No comments.

Company editor-in-chief: I have reviewed the Peer-Review Report, full text of the manuscript, and the relevant ethics documents, all of which have met the basic publishing requirements of the World Journal of Hepatology, 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.

Response:

We thank editor-in-chief for evaluation of manuscript.