Response to Editor

In the submission of our first revision, the journal provided only two reviewers comments as Peer Review Report. That is why we were not able to respond to all reviewer's comments. In this version, we have responded to all comments of reviewers.

1. As per editor's suggestion, we have revised the manuscript according to the Cross-Check report. They are marked in RED.
2. We have provided the editable figures in power point as 53130-image files
3. We have modified the multiple institutions for authors and are marked in RED.
4. We included only one institution for corresponding author.
5. Abbreviations and acronyms were defined the first time when they were used in all sections.
6. Corrected all cited references number like [number], there are no spaces before [number] but our Endnote did not allow us to avoid the space between two references like [1, 2].
7. We arranged the references as per your suggestion.

Response to Reviewers

We thank the Reviewers for being so positive regarding this Manuscript and their comments on high significance of the effects of alcohol on HBV-infection pathogenesis and treatment options. We would like to address the suggestion raised by Reviewer 1:

Reviewer id: 03021264

The article discussed the combined effects of alcohol and HBV-infection in the progression of liver diseases. Authors emphasized the synergistic effect of alcohol and Hepatitis B virus infection on progression of end-stage liver diseases and expounded the possible mechanisms. The review can improve the treatment options for the HBV-alcoholic patients.

"*But the article content is too much, especially the first part of the introduction of HBV epidemiology and other content can be considered appropriate briefly*".

Response:

Following the Reviewer's suggestion, we removed the information about the vertical transmission of HBV-infection since in this review, we are not analyzing the role of alcohol on the routes of HBV transmission. We removed the second paragraph on page 4 of initial version.

Reviewer id: 03011757

The review submitted by Genesan et al summarizes the current literature on how alcohol abuse promotes the progression of HBV-associated liver disease. The review is well written, and the content is thoroughly and clearly arranged.

There are some minor points the authors should address prior to publishing:

1. *Figure 1 and Figure 2: Detailed description is missing in the figure legend.*

Response: We included the description of figures

2) The paragraph on HBV innate immunity is very extensive and would benefit from shortening – e.g. remove the basic description of DC etc.

Response: As per reviewer's suggestion we removed few sentences (shortened) in the revised manuscript.

3) Core tip: Worldwide, 1.5 billion people had chronic liver disease: please provide the year this number was imposed.

Response: As per reviewer's suggestion, we included this.

4) Introduction: Current treatment for chronic HBV patients….: I suggest to separately described the medical treatment form the surgical (liver transplantation).

Response: We respect this suggestion, but we are planning a separate review for Treatment options for alcohol abusing HBV patients. Other reviewers suggested shortening the review, and we feel that to expand on some items will be excessive for this review.

5) Page 5: please revise: chronic hepatitis B are ranged from…; HBV infection is one of the major risk factors for the outcome to HCC.

Response: We modified these sentences and marked in red.

6) Page 10, last paragraph: please revise: The first phase is the immune-tolerant…; In the second phase, the immune clearance phase, elevated serum ALT….

Response: We modified these sentences and marked in red.

7) Page 13: …pro- and anti-inflammatory cytokine levels

Response: We modified these sentences and marked in red.

8) Page 14, second paragraph: hydrodynamic injection model of acute HBV (Yang et …)

Response: We apologize for this mistake and we have corrected it.