## **Dear editor and reviewer(s)**,

Thanks a lot for your great efforts in reviewing my revised version. I'm very excited that I could have a precious opportunity to revise my manuscript. As soon as I received the comments, I began to revise my manuscript immediately. I have read these comments and revised my manuscript carefully and thoroughly according to the every comment of the reviewer(s). And some necessary explanations are listed in the responses. The point-to-point responses are listed as follows:

Reviewer #1:

Scientific Quality: Grade C (Good) Language Quality: Grade B (Minor language polishing) Conclusion: Minor revision

## Specific Comments to Authors: The paper is well written.

RE: Thanks for your comments and recognition. This review aims to highlight the clinical evaluation and the potential regulatory signaling and the key factors. So we hope that this review could be beneficial to the clinicians and the clinical practice.

However, the authors should highlight the consequence of the mechanism of liver injury in clinical management and how this evidences can chance our clinical practice.

RE: Thanks for your comments. As you said, in the former version, we didn't well in highlighting the consequence of the mechanism of liver injury in clinical management and how this evidences can change our clinical practice. Therefore, we revised our manuscript and made some necessary changes according to your suggestions. The treatment of LIAAP is closely related to the treatment of AP, because the degree of liver damage depends on the severity of the pancreatitis. The clinical management of LIAAP include anti-inflammatory and antioxidant therapies, the improvement of the microcirculation, the inhibition of apoptosis, the promotion of liver cell regeneration and supportive therapy. We added the related therapeutic drugs and mechanisms, including sodium butyrate, antioxidants, recombinant human soluble thrombomodulin and prostaglandin E1, nilotinib, heme oxygenase-1, vascular endothelial cell growth factor ,somatostatin and ethyl. We don't know whether the above responses are satisfactory or not, please email me freely(zuohoud@163.com) if you have any questions. Thank you very much again for your great efforts.

## The title does not match with the content of the paper in my opinion should be modified

RE: Thanks for your comments. I agree with your comment. The title of the manuscript should be changed, which can reflect the current research situation. Because the review comprises two parts: the clinical evaluation and potential mechanism of LIAAP, so we modified the title. And we think the title "Liver injury associated with acute pancreatitis: the current status of clinical evaluation and involved mechanisms" may be more appropriate. We don't know whether the new title is suitable or not. If it's not appropriate, would you like to give us some constructive

suggestions? Thank you very much. Please email me freely (zuohoud@163.com) if you have any questions. Thank you very much again for your great efforts.

Reviewer #2:

Scientific Quality: Grade C (Good)

Language Quality: Grade B (Minor language polishing)

**Conclusion: Accept (General priority)** 

Specific Comments to Authors: The manuscript gives a well-structured overview of the relationship between liver damage and acute pancreatitis. Different facets of this clinical picture are highlighted on the basis of the current literature.

RE: Thanks for your precious comments and recognition of this review. We also want to make some my own contributions to the understandings and the treatment of the liver injury associated with AP.

We have made some changes according to the reviewers comments carefully and thoroughly. The version with marked changes will be uploaded as the supplemental file if necessary.

Finally, I express my sincere appreciation to editor and reviewer(s) again. Although I try my best to revise my manuscript, it still may not be satisfactory completely to all the reviewers. Nevertheless, I'm eager to get a chance to publish this review, so that I can deliver the relative information to the clinicians and make my own contributions. Thank you very much. Please don't hesitate to email me, if you have any questions. I sincerely appreciate your efforts and constructive comments. Have a nice day! Best regards

Houdong Zuo(zuohoud@163.com)