

Dear Editors and Reviewers:

Thank you for your letter and for the reviewers' comments concerning our manuscript entitled "Paper Title" (NO: 45121). Those comments are all valuable and very helpful for revising and improving our paper, as well as the important guiding significance to our researches. We have studied comments carefully and have made correction which we hope meet with approval. Revised portions are marked in red in the paper. The main corrections in the paper and the responses to the reviewer's comments are as follows:

Responses to the reviewer's comments:

Reviewer #1:

[Comment 1] Of interest, but very invasive methods: CCl₄ poisoning to induce steatosis and cutting open the mice for the blood flow measurements. MRI methods offer the same information and are superior and noninvasive. Relevant studies involving MRI should be cited and discussed, also the US study of Topal et al. J Clin US, 2015;43:26-33. The abstract must give numbers (mice groups, flow and diameter changes). In introduction the last paragraph must also come with references. In results the headings should be shortened to Weight changes, Lipid deposition etc. The value (clinical relevance) of the sinusoids measurements presented here, compared with what the others did, must be made clear.

[Answer 1] Thank you very much for your pertinent advice, and I have benefited a lot from it. According to your suggestions, I made the following changes:

- I added the discussion between clinical image techniques and two-photon fluorescence microscopy imaging technique.
- I added relevant numbers/values to abstract, and result values were presented as the form of Mean \pm SD.
- I traced back to the original papers have been cited by the introduction last paragraph, and added them into paper.
- I shortened the headings in the results part.
- I discussed the value of our study.

Thanks again for your sincere comment, if there are any questions please contact me, and I'll try my best to give you a satisfactory feedback. Best wishes!

Reviewer #2:

[Comment 2] Hemodynamic changes in hepatic sinusoids of hepatic steatosis mice
Corresponding author: Wei-Guang Zhang, MD, PhD, Professor, Zhang et al. conducted an animal study with the aim to evaluate histological changes and modification of intrahepatic hemodynamic in CCl₄ induced steatotic mouse livers. The authors used 2 different strain, CCl₄ for lipid accumulation and H.E. for hepatic sinusoidal vessels. A 2-fluorescence photon microscope was used to assess the flow pattern of the red blood cells. The authors found that as modeling time increase the plasma flow gradually decreased and the diameter of the sinusoidal vessel became smaller. The introduction provides a summary of the studies which

were done so far. The aim of the study is clear stated at the end of the introduction. The used material and methods are listed understandable. In the discussion their own results are discussed with the result from other authors. Most studies which were done with focus on the microcirculation. The conducted study has a different approach and is the first one, which direct indicates visual the changes in fatty livers. From that point of view, the study should be published in the present form.

[Answer 2] Thank you very much for your recognition, and I'll try my best to modify the manuscript to make it as perfect as possible. Best wishes!

We appreciate for Editors/Reviewers' warm work earnestly, and hope that the correction will meet with approval.

Once again, thank you very much for your comments and suggestions.

Yours sincerely,

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