

Dear Editor Yuan-Qi:

Thank you for offering me the opportunity to revising my manuscript submitted in World Journal of Gastroenterology (manuscript No. 21909).

I am very grateful for your patience, cautious and conscientious.

I have read the guidelines for several times but can't understand the details of the tile page fully. I change the format of the title page and wish that could meet your requirements. If there are some other mistakes, could you please help me revise.

Thank you very much for your attention and consideration.

Sincerely yours,

Zhiming Huang.

Here are the modifications of my manuscript.

1. In the third paragraph of "Introduction" part, I modify the cites number "[10-12]" to "[9-11]";
2. In "statistical analysis" part, I add "All the statistical review of the study was performed by a biomedical statistician." at the of this part to meet the requirement;
3. In legend of "Figure 2", I add "(39kDa)" and "(23kDa)" following "Phosphorylated" and "nonphosphorylated" separately. And also I have replaced Fig. 2.
4. In legend of "Figure 2", I modify the sentence "and protein expression level of AQP-2 showed significant differences after day 2 and successive days." to "and protein expression level of AQP-2 showed significant differences since day 2 (3mg/kg: 100%±22.9% vs 54.7%±11.7, P<0.05; 5mg/kg: 100%±22.9% vs 53.0%±9.4%, P<0.05) and successive days."
5. In legend of "Figure 3", I modify the sentence "A significant difference was detected after day 1 and successive days." to "A significant difference was detected since day 1 (3mg/kg: 100%±16.3 vs 68.50±19.20, P<0.05; 5mg/kg: 100%±16.3 vs 34.16±15.14, P<0.01) and successive days."
6. In legend of "Figure 4", I modify the sentence "The fecal water content increased from day 1 in the 5 mg/kg tolvaptan group, and from day 2 in the 3 mg/kg group." to "The fecal water content increased from day 1 in the 5 mg/kg tolvaptan group (41.4%±12.8% vs 66.8%±10.3%, P<0.05), and from day 2 in the 3 mg/kg group (40.0%±13.3% vs 59.2±15.8, P<0.05)."

All the modification has been noted in the revision manuscript.

Here are my responses to reviewers:

03261947

Response: thank you very much for reviewing my manuscript and we are very grateful for your acceptance.

03251423

Response: thank you very much for reviewing my manuscript and we are very grateful for your acceptance.

02445715

Response: thank you very much for reviewing my manuscript and we are grateful for your opinion to improve the qualification of my manuscript.

Here are my responses for your comments:

1) In Fig. 2, which are phosphorylated or non-phosphorylated protein bands? Please show me the real bands by marking arrows.

Thank you for your comment and I have added “(39kDa)” and “(23kDa)” following “Phosphorylated” and “nonphosphorylated” separately. I also have replaced Fig. 2.

2) In Figs. 2 and 3, I cannot see a positive control. You should show the data.

Thank you for your comment. Positive and negative control should be the most basic part of the study to verify the impact of the drug and in our study, valid drug which could down regulate aquaporin-2 (AQP-2) in distal colon in cirrhotic model would be the proper positive control. However, as the description of the available cites, AQP-2 is reported mainly to be blocked in kidney in cirrhotic model and in distal colon of healthy entities by tolvaptan. Other methods, such as low sodium diet, have not been confirmed its effect in distal colon in cirrhotic model. So we believe there is no proper positive control group in this study. What's more, we performed this experiment to explore tolvaptan's effect on AQP-2 expression and fecal water content, and after the administration of tolvaptan, sensible results had been showed. So we believe this protocol might be proper to illustrate this issue.

3) In Fig. 3, you should show Q-PCR data.

Thank you for your comment and I have added Q-PCR data in the Fig. 3 legend to present a better understanding of the picture. In legend of “Figure 3”, I modify the sentence “A significant difference was detected after day 1 and successive days.” to “A significant difference was detected since day 1 (3mg/kg: 100%±16.3 vs 68.50±19.20, P<0.05; 5mg/kg: 100%±16.3 vs 34.16±15.14, P<0.01) and successive days.”