

The answer to Reviewer 1:

1. Title is misleading.

We absolutely agree with the reviewer, and we modify the title name to “Overview of Bile Acid Signaling in the Cardiovascular System”.

2. The description about “cytotoxicity” in Introduction (P.3) and Figure 1 lacks reference.

We look up a large number of literature finds that many reports about the cytotoxicity of hydrophobic bile acids, so we add more precise discussion together with reference.

3. In Table 1, add the explanation of all abbreviations used in footnote.

Yes, we already add the explanation of all abbreviations used in footnote.

4. Again, in Table 1, there is no explanation of LCT which is not found in the text. Is it LCA?

We are so sorry for the confusion brought to the editor, we already add the clear explanation of all abbreviations used in footnote in Table 1.

LCT is lithocholyltaurine, which is different with the lithocholic acid(LCA).

5. Again, in Table 1 footnote, there is no corresponding words of TLCA(taurolithocholic acid),CA(cholic acid),and UDCA(ursodeoxycholic acid)in Table 1.

We are so sorry for the confusion brought to the editor and Thank you very much for your careful reminder, we already add the clear explanation of all abbreviations used in footnote in Table 1.

6. BESP stands for Bile Salt Export Pump, not bile salt output pump.

Yes, we already modify this mistake for all review.

7. There are some irrelevant words in medical article, such as essay, interfacial activity, A 1983 study, ect.

We already checked the whole review and modified the irrelevant words.

The answer to Reviewer 2:

1. There are multiple grammar errors, the author should revise the paper more.

We already checked the whole and modified the grammar errors as far as

possible. In addition, we also find American Journal Experts to help us address the language problems of the manuscript as much as possible.

**The following are my response to the questions and comments raised in the second-round review.**

1. Title is misleading.

We absolutely agree with the reviewer, and we modify the title name to "Overview of Bile Acid Signaling in the Cardiovascular System".

2. The description about "cytotoxicity" in Introduction (P.3) and Figure 1 lacks reference.

We look up a large number of literature finds that many reports about the cytotoxicity of hydrophobic bile acids, so we add more precise discussion together with reference.

3. In Table 1, add the explanation of all abbreviations used in footnote.

Yes, we already add the explanation of all abbreviations used in footnote.

4. Again, in Table 1, there is no explanation of LCT which is not found in the text. Is it LCA?

We are so sorry for the confusion brought to the editor, we already add the clear explanation of all abbreviations used in footnote in Table 1. LCT is lithocholyltaurine, which is different with the lithocholic acid(LCA).

5. Again, in Table 1 footnote, there is no corresponding words of TLCA(taurolithocholic acid),CA(cholic acid),and UDCA(ursodeoxycholic acid)in Table 1.

We are so sorry for the confusion brought to the editor and Thank you very much for your careful reminder, we already add the clear explanation of all abbreviations used in footnote in Table 1.

6. BESP stands for Bile Salt Export Pump, not bile salt output pump.

Yes, we already modify this mistake for all review.

7. There are some irrelevant words in medical article, such as essay, interfacial activity, A 1983 study, ect.

We already checked the whole review and modified the irrelevant words.

8. There are multiple grammar errors, the author should revise the paper more.

We already checked the whole and modified the grammar errors as far as

possible. In addition, we also find American Journal Experts to help us address the language problems of the manuscript as much as possible.

9. We already modify the Figure 1 to Table 1 in our review.

10. We already move the Table 1 footnote under the Table 1.

11. We already modify the Table 1 arrow indication according to editor to make it clear understand.

12. We already modify the Table 1 to Table 2, leave space in each line and add reference in our review.

13. We already modify the Figure 2 to Figure 1 in our review.

14. We already offer the grant application form in PDF format.