

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastroenterology*

Manuscript NO: 75757

Title: Duodenal-jejunal bypass reduces serum ceramides via inhibiting intestinal bile acid-farnesoid X receptor pathway

Provenance and peer review: Unsolicited manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05260837

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Italy

Author's Country/Territory: China

Manuscript submission date: 2022-03-11

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2022-04-06 10:23

Reviewer performed review: 2022-04-13 02:39

Review time: 6 Days and 16 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input checked="" type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

Peer-reviewer statements	Peer-Review: [<input checked="" type="radio"/>] Anonymous [<input type="radio"/>] Onymous Conflicts-of-Interest: [<input type="radio"/>] Yes [<input checked="" type="radio"/>] No
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SPECIFIC COMMENTS TO AUTHORS

Bile acids are traditionally known as lipid absorption-facilitating agents. It was not until recent years that the role of bile acids as signaling molecules in modulating metabolism has been unveiled. The intestinal lumina, where bile acid concentrations are high, is the main place for bile acid signaling. The interaction between bile acids and Farnesoid X receptor is more complicated, as different subtypes of bile acids have distinct effect on the downstream pathway of Farnesoid X receptor. Chenodeoxycholic acid represents the most potent Farnesoid X receptor stimulator while ursodeoxycholic acid and β -muricholic acid are Farnesoid X receptor inhibitors. Coincidentally, the changes in lipid metabolism after intestine-selective Farnesoid X receptor inhibition is similar to the changes following duodenal-jejunal bypass that hepatic fat accumulation is alleviated and the key transcriptional regulators and enzymes involved in hepatic de novo lipogenesis are downregulated. In this study, the authors investigate of luminal bile acids following duodenal-jejunal bypass. This study is very well designed and the manuscript is very well written. Methods are described in detail, and the results are interesting. The results are discussed with updated references. The limit of the study is also discussed. The reviewer recommends to accept this interesting study after a minor editing.

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Provenance and peer review: Unsolicited manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05386935

Position: Peer Reviewer

Academic degree: MD

Professional title: Research Associate

Reviewer's Country/Territory: Switzerland

Author's Country/Territory: China

Manuscript submission date: 2022-03-11

Reviewer chosen by: Jin-Lei Wang

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Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

Peer-reviewer statements	Peer-Review: [<input checked="" type="radio"/>] Anonymous [<input type="radio"/>] Onymous Conflicts-of-Interest: [<input type="radio"/>] Yes [<input checked="" type="radio"/>] No
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SPECIFIC COMMENTS TO AUTHORS

This is an interesting study of the duodenal-jejunal bypass and intestinal bile acid-FXR pathway. This study measured the changes of individual luminal bile acid and ceramide concentrations within the enterohepatic circulation after duodenal-jejunal bypass in a high-fat diet and streptozotocin-induced diabetic rat model. The research is well performed, and the results are very interesting. The animals and surgical procedures are reasonable, and RNA analysis results are excellent. Minor comments: 1. The manuscript requires a minor editing. Please take attention about some abbreviations. 2. The figures are too small, please update them with high resolution ratio images.

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Provenance and peer review: Unsolicited manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05429710

Position: Peer Reviewer

Academic degree: MD, PhD

Professional title: Associate Professor

Reviewer's Country/Territory: Spain

Author's Country/Territory: China

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Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No



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Peer-reviewer statements	Peer-Review: [<input checked="" type="checkbox"/>] Anonymous [<input type="checkbox"/>] Onymous Conflicts-of-Interest: [<input type="checkbox"/>] Yes [<input checked="" type="checkbox"/>] No
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SPECIFIC COMMENTS TO AUTHORS

The authors designed this study to investigate of luminal bile acids following duodenal-jejunal bypass. Overall, the study is very interesting. The manuscript is well written. I suggest the author to update the figures with black, white, and gray color. No other comments.