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Dear Editor,

We are grateful to the reviewers for their valuable comments and recognition of the relevance of our Editorial for the readership of *the World Journal of Hepatology*. We have updated the manuscript in line with the requests from the Science Editor and Reviewers. A detailed description of the changes is given below.

The manuscript has been reviewed for grammatical clarity and appropriate vocabulary by a colleague whose native language is English.

We are convinced that our Editorial meets the high standards of *the World Journal of Hepatology*.

Sincerely,

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Requests from Reviewer#1 (02462252)

COMMENTS TO AUTHORS

Good article will be of interest to general readership.

Response: We thank the reviewer for the kind words.

Requests from Reviewer#2 (01560464)

COMMENTS TO AUTHORS

1)The review is too simple to describe the role of Wnt signaling in molecular pathogenesis and targeted therapy of CCA. It need to be further supplemented according by the two parts of Wnt signaling in molecular pathogenesis and targeted therapy of CCA. 2) I suggest that the review can be reviewed again after supplementaion .

Response: We would like to point out that we received an invitation to write an Editorial about an original study that is of interest to the readership of *World Journal of Hepatology*. Thus, our manuscript **is not a review** aiming to give an extensive overview of the state of the art knowlegde on cholangiocarcinogenesis, within or outside the context of Wnt signaling. We have extensively discussed, and provided context, for the experimental findings of Boulter and colleagues. We were not entirely sure what the reviewer exactly meant with remark 1, but interpreted it as being outside the scope of **an Editorial**.

Requests from Reviewer#3 (02441405)

COMMENTS TO AUTHORS

This editorial nicely outlines the role of Wnt signaling in the development of cholangiocarcinoma based upon the human tissue and animal studies. The manuscript is well written. Only a minor comment: Page 2, the second last line, "Molecular-targeted therapies hold potential for (personalized) treatment of malignancies including CCA". Can it be clearly stated as "Molecular-targeted therapies hold potential for PERSONALIZED treatment of malignancies including CCA"?

Response: We thank the reviewer for the nice words. We have now clearly stated that "Molecular-targeted therapies hold potential for PERSONALIZED treatment of malignancies including CCA"?

Requests from Reviewer#4 (00182833)

COMMENTS TO AUTHORS

Thank you to invited me to review this article (Cholangiocarcinoma, gone without the Wnt?) ?These review addressed the prospect for understanding of the molecular pathogenesis of CCA . ?Cholangiocarcinoma is not frequent in cirrhotic Liver ?Wnt signaling cascade raise interest of other recherches for more study in human to decrease the incidence ?In patient with CCA. In the future if we succeeded , Wnt inhibition with or without other line of treatment tiis good achievement

Response: The manuscript has been reviewed for grammatical clarity and appropriate vocabulary by a colleague whose native language is English.