

Reviewer #1:

Scientific Quality: Grade C (Good)

Language Quality: Grade B (Minor language polishing)

Conclusion: Major revision

Specific Comments to Authors:

In this mini-review, the authors summarized the timely knowledge of intraductal papillary neoplasm of the bile duct (IPNB). The article itself is scientifically sound and potentially useful for both clinicians and basic scientist. The English presentation is readable and understandable. However, the article needs a better organized. Some more information and critical discussion are needed to make the article higher quality. The authors may take the following suggestions to improve the data presentation.

1. The main text is now narrated without any divided subsections. It should be divided into reasonable topics and sub-topics which will be easy to the readers to follow the story.

In response, → appropriate sections and sub-sections have been added along the main text, according to your precious suggestions.

2. When describing the histological classification, microscopic findings, or radiologic findings, it would be better to have the figures demonstrating what the authors are describing. Now it is hard to imagine the characteristics of each IPNB subtypes referring by the authors.

In response, → several figures regarding the histologic and radiologic aspects of IPNB have been added, according to your suggestions.

3. As it is a review article, the critical discussion is important. The authors should add more section to describe the current advancement of the current topic, and their perspective for the future direction of this topic using their expertise. It could be both clinical or research aspects.

In response, → several sections have been added at the end of the manuscript describing the most recent advancements particularly in the field of genetic research with regard to mutations involving different oncogenes and oncosuppressor genes that might lead in future to the development of targeted therapies, particularly addressed at the WNT- β -catenin signaling pathway.

Indeed, this peculiar pathway appears to be specifically involved in the cancerogenesis of IPNB, mainly type 2, and ongoing clinical trials are testing the efficacy of WNT inhibitors in advanced gastro-intestinal neoplasms, including CCA.

Reviewer #2:

Scientific Quality: Grade C (Good)

Language Quality: Grade B (Minor language polishing)

Conclusion: Minor revision

Specific Comments to Authors:

This manuscript is a mini-review that summarize the current concepts regarding IPNB, including classification, diagnosis, management and prognosis. IPNB has not yet well investigated as this disease concept is relatively new. This review article contains interesting and informative knowledges for readers.

The following minor issues require clarification:

Minor 1. (Abstract) Conclusion should be modified based on the contents of Conclusion in the main text.

In response, → In the Abstract, conclusions have been modified according to the Conclusions reported in the main text, as suggested.

2. (Table 1) Please explain an abbreviated word of “n.r”.

In response, → abbreviation “n.r.” has been added below the Table 1.

3. (P13) The authors introduced some non-surgical treatments. I recommend that the authors describe each treatment in more detail as those should be interesting for readers.

In response, → each non-surgical treatment cited has been properly described, according to your suggestions, with particular attention to ablative endoscopic techniques.

4. Subtitles in Main Text, such as classification, diagnosis, management, prognosis and so on, can help readers’ understanding.

In response, → appropriate sections and sub-sections have been added along the main text according to your precious suggestions.