PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Pathophysiology*

Manuscript NO: 82233

Title: Polymorphism of genes encoding drug-metabolizing and inflammation-related enzymes for susceptibility to cholangiocarcinoma in Thailand

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 05453202

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Reviewer_Country

Author's Country/Territory: Japan

Manuscript submission date: 2022-12-11

Reviewer chosen by: AI Technique

Reviewer accepted review: 2023-01-12 01:58

Reviewer performed review: 2023-01-22 06:21

Review time: 10 Days and 4 Hours

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No creativity or innovation

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Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

The manuscript has the potential to publish, as the author presented valid observations of positive OV individuals closely associated with common risk factors such as the smoking status and alcohol consumption were delivered in the manuscript. However, there are still rooms for improvement to enhance the visibility and quality of the work. The following are comments from my point of view for a manuscript entitled, "Review manuscript: Polymorphism of genes encoding drug-metabolizing and inflammation-related enzymes for susceptibility to cholangiocarcinoma in Thailand".   1. The methodology was comprehensively described in the manuscript, however, the associated with each of the inflammation genes and the well-known metabolizing enzymes were highly due to the small sample size in the study. 2. If the statement or inclusion criteria of normal alpha fetoprotein are not the main contributor variable in the study, please remove. Else, there is no point mentioning in the manuscript. 3. Were the blood samples stored at -80C before DNA extraction? Why was it not extracted immediately after? Or was the DNA that was stored at -80C? 4.   Please emphasis the basis of assessing the inflammation genes polymorphisms with CCA. And the link

between the carcinogen infection was unclear to correlate it with CCA and the inflammation markers. Need to justify the purpose.

Scientific quality	[ ] Grade A: Excellent [Y] Grade B: Very good [ ] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[Y] Grade A: Excellent [ ] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No creativity or innovation

Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [Y] Grade B: Good [ ] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[Y] Grade A: Priority publishing [ ] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[Y] Yes [ ] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

Dear Authors, I believe you and your team have put tremendous work into this collaboration, studying and collecting data overseas. Your study is interesting and worth to be studied further. The possibility of CCA could be caused by OV and external factors (e.g. carcinogens or chemicals). Authors found no significant association of CCA with polymorphisms of drug metabolizing enzyme and inflammation-related genes. Perhaps, authors should state why CYP2E1 was tested, as there are other human CYPs that could be carcinogen-activating enzyme e.g. CYP4B1.   Based on your manuscript, understand that this is a continuous study, the cases of CCA collected were dated back to 1999-2005, perhaps authors could compare data with the current population? I am not well-versed with case study manuscript, but it would be good if authors could provide the conclusion after the discussion section, limitations of your study and future work. All the best, keep up the good work.   Thank you