

Point-by-point responses to reviewers' comment

May 21, 2015

Dear Dr. Ma,

Please find attached the edited manuscript in Word format (file name: 18196_revised.doc).

Title: Drug- and Herb-Induced Liver Injury: progress, current challenges and emerging signals of post-marketing risk

Authors: Emanuel Raschi, Fabrizio De Ponti

Name of Journal: *World Journal of Hepatology*

ESPS Manuscript NO: 18196

1. Revision has been made according to the suggestions of the reviewers. A point-by-point outline of responses is provided below. In the current version, we have also added a personal viewpoint in the last paragraph (concluding remarks) on key issues that should be prioritized in terms of research agenda in the next future.

Reviewer # 13180.

The mini review on drug and herb-induced liver injury was already subjected to a previous review before I was asked to review. While the authors have tried to incorporate many of the recent advances in the field including up to date references, the quality of the manuscript will be enhanced by considerably by editing by an English language editor. There are countless words that need to be spaced between them.

We have revised the text for readability, typos and English style, where appropriate.

Although the title mentions drug and Herb-induced DILI, the information/ details on HILI appear to be inadequate. By contrast drug-induced liver injury is more than comprehensively covered.

*We are aware of this issue. The point is that available data and published studies have actually addressed DILI, whereas HILI is not comprehensively covered as it is difficult to capture the use of herbs in the population. Most of the evidence on HILI comes from case reports, which are accruing and systematically collected by different research teams. In the current version of the manuscript, we have emphasized this point, especially in the final section (research agenda on HILI), where a very recent review is now quoted (Teschke R, Eickhoff A. *Front Pharmacol.* 2015 Apr 23;6:72). HILI is also now mentioned as a priority topic for research activities.*

The second sentence under “signals emerging from observational studies” is too long and needs to be broken down to two or more sentences for clarity purposes.

We revised the text according to reviewer's suggestion.

Page 9, end of second paragraph “Follow-up is also.....experienced DILI” needs to be referenced.

We added a relevant reference.

Reviewer # 41858.

The manuscript is well-written and conforms to the accepted standards.

We thank the reviewer for this comment. No answers to specific issues are required.

2. References and typesetting were corrected.

Thank you again for considering our manuscript in the *World Journal of Hepatology*.

Sincerely yours,

A small, square image containing a handwritten signature in black ink on a light blue background. The signature appears to read 'Fabrizio De Ponti'.

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