

November 15, 2013

Dear Editor,

Please find the attached manuscript in word format (file name : 6520-review)

ESPS Manuscript NO: 6520

Title: Hepato-cardiac disorders

Author Name: YM Fouad, R Yehia

Name of Journal: *World Journal of Hepatology*

The manuscript has been improved according to the suggestions of reviewers:

- 1 Format has been updated
- 2 Revision has been made according to the suggestions of the reviewers
- 3 References and typesetting were corrected
- 4 We thank all reviewers who accepted our review

Please find below the reply to reviewers.

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

Best regards

Sincerely

Dr Yasser M Fouad MD Do

Professor of Gastroenterology and Hepatology

Reviewer # 1 (02444960):

The manuscript by Yasser Mahrous Fouad, is a review focused in classification, management, treatment and overcoming of a group of heart, liver and other diseases causing hepatic, cardiac disorders or both, respectively. Although there are few articles in the recent bibliography exploring these conditions (PMID: 23853073), this paper is of interest in clinical practice. However I have few suggestions regarding some points of the manuscript: 1- To structure the manuscript. Some subsections are not formatted as the others and readers could get confused regarding to which group of disorders they refer. Authors should check also when some sections

should be written in bold (for example, "Chronic HF and congestive hepatopathy" in page 16). 2- To include few schemes or more tables to make feasible the understanding of the manuscript. 3- To check grammar and spelling of manuscript, in order to polish the text, specially words like Algille (change to Alagille in page 21) 4- To modify sentences with no sense like " It is worth noting that....,undergoes a substantial prolongation in the" in page 12 or like that referring to the role of valganciclovir in page 20. 5- To include a section with abbreviations and to write by first time the abbreviation after the group of words they are defining, for example, "primary biliary cirrhosis" (PBC) in page 8. 6- To check there are not missed spacebar after and before "." " ," ":" in the whole manuscript. 7- Bibliography should be formatted following World J Hepatology recommendations, for example page 18 "Roell W, 1997" should be cited with a number, reference 21 (at bibliography section) has the year of the publication between brackets (2013). Authors should unify the format of all the references.

Reply: Thank you for your comments. We made the changes you suggested (highlighted in the text) . we added tables and graph.

Reviewer# 2 (ID: 0003036318):

This is a comprehensive review of the major liver diseases that affect the heart and vice versa. The authors also discuss briefly the main disorders affecting both the heart and the liver. The topic is interesting and provided information is up to date. I believe that the review would benefit from the discussion of two relevant entities, i.e. portopulmonary syndrome and pulmonary hypertension. Moreover, there are several sentences, even whole paragraphs, who lack references and this should be corrected.

Reply: Thank you for your comments. We made the changes you suggested (highlighted in the text) . we add portal hypertension complications to the text.

Reviewer #3 (ID: 00054317):

This manuscript is a very succinct but clinically applicable and very instructive for practicing Cardiologist and Hepatologist alike. The editors need to correct misspellings such as "Alagille" and sentences with poor grammatical closure and syntax can be easily fixed as well as a few typos.

Reply: Thank you for your comments. We tried to make these changes .

Reviewer #4 (ID:00053485)

In this review, Fouad & Yehia. reviewed hepato-cardiac diseases. The authors well described important points from the view of both hepatologist and cardiologist. However, there are several issues in the current manuscript. Major Points 1. There is no Figure in this review. To get further understanding of readers, figures and tables are better to be added. 2. Chronic HCV infection is associated with severe insulin resistance and mild atherosclerosis. One would be interested in the paradoxical effect of HCV and this topic has to be discussed by citing following papers: Atherosclerosis. 2012;221:496-502, J Gastroenterol. 2013;48:93-100., Atherosclerosis. 2013;231:22-6., Aliment Pharmacol Ther. 2013;38:415-23. 3. Budd-Chiari syndrome is a cause of hepatocellular carcinoma. Although authors briefly described this topic, the possible mechanisms are better to be described by citing following papers: Hepatology. 1998;28:1191-8, Hepatogastroenterology. 1999;46:3237-40, Eur J Gastroenterol Hepatol. 2004;16:319-24, Oncol Lett. 2013;6:612-616. 4. Effects of portal hypertension and its treatment on cardiac function would be an interesting topic.

Reply: Thank you for your valuable comments. We made the changes you suggested (highlighted in the text) . we added tables and graph.

Reviewer #5 (00140083):

This review article contains a number of non-evidence based, non referenced or poorly referenced statements. The authors rarely express their opinions and the manuscript is largely a collection of statements from other papers.

The criteria, if any, followed in choosing the manuscripts for this review are also not stated. Most references are outdated. The only table is inappropriate as contains a number of topics not examined in the text. More tables and figures would be needed to illustrate pathophysiologic mechanisms and to catch and focus the attention of the readers. Pages are not numbered.

Reply: Thank you for your comments. The subject is not in intensive research although very important . So no evidence based data are much available. References are added . we also added tables and a graph.