

February 28, 2015



Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 16486-review.doc).

Title: Mechanisms of Givinostat inhibiting hepatic stellate cells proliferation and activation of lysine-acetylated proteins

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Name of Journal: *World Journal of Gastroenterology*

ESPS Manuscript NO: 16486

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 reviewer.

We are very grateful to your comments for the manuscript. According with your advice, we amended the relevant part in manuscript. Some of your questions were answered below.

(1) *The authors should avoid using acronyms in the abstract, if they are not repeated. For instance HDAC. Other acronyms are not disclosed (e.g., ROS). Please review this.*

The acronyms in the abstract have been checked and corrected.

(2) *Introduction: Line 21, page 3. Hepatic sinusoid capillarization occurs without activation of HSC. It occurs with aging, without proliferation or myofibroblastic transformation of HSC. Please delete this.*

Corrected accordingly.

(3) *Materials and methods Line 10, page 9. Collagen fibers accumulate but they do not proliferate. Please change "proliferation of collagen" to collagen deposition. It is not needed to state which are the colours of the Masson's Trichrome. The authors could have quantified in the control, CCl4 and givinostat model. This would increase the value of the manuscript.*

These sentences such as "proliferation of collagen" in page 9 have redirected according to the suggestions of reviewer. And we have quantified in the control, CCl4 and givinostat model according to the suggestions of reviewer.

(4) *Figures Figure 5 The images should be reviewed. The images at 40x do not have good quality (bubbles in two images) and should be changed. The Legend should be reviewed (Hyperplasia does not apply to collagen).*

Figure 5 has been replaced. The Legend has be reviewed according to the suggestions of reviewer.

(5) *In the Methods section, it is important to quantify the amount of collagen and/or grade of fibrosis in the control and treatment groups in order to statistically verify the presence of significant changes. See for example, PLoS ONE 9(11): e112384.*

we have quantified in the control, CCl4 and givinostat model according to the suggestions of reviewer and referenced in this article.

(6) In the Methods section, the authors did not describe in details the animal model of liver fibrosis due to incomplete sentence in page 9, line 1. The sentence “four portions of CCl₄ and six of....” is incomplete and the duration of the treatment was not stated.

The sentence “four portions of CCl₄ and six of....” should be “four portions of CCl₄ and six of olive oil”.

(7) Page 4, line 6, the word “imbalanced” should read “imbalance” without “d”

Corrected accordingly.

(8) Page 4, line 9, the word “avalproic acid” at the beginning of the line should be deleted. It is both incorrect and redundant.

Corrected accordingly.

(9) Page 5, 3rd line of last section, in the expression (1×10^6 cells/well) the exponent is misplaced.

The word “ 1×10^6 cells/well” should be “ 1×10^6 cells/well”, Superscript.

(10) In the Results section; page 12, line 7; the word “by givinostat” should be “by CCl₄”

Corrected accordingly.

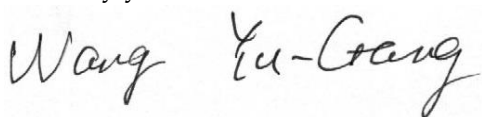
(11) Page 23, the figure legend “Fig. 5 Pathological changes of liver tissue in mouse models of liver fibrosis induced by givinostat treatment” should read “Fig. 5 Pathological changes of liver tissue in mouse models of liver fibrosis induced by CCl₄ treatment” because it is the CCl₄ that induced liver fibrosis but givinostat treatment reduces the fibrotic process.

Corrected accordingly.

3 References and typesetting were corrected

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

Sincerely yours,



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