

Format for ANSWERING REVIEWERS



August 5, 2013

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 4217-review.doc).

Title: The quality of ulcer healing (QOUH) of gastric ulcer; Natural product beyond acid suppression.

Author: Napapan Kangwan, Jong-Min Park, Eun-Hee Kim, and Ki Baik Hahm

Name of Journal: *World Journal of Gastroenterology*

ESPS Manuscript NO: 4217

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

Reviewer 02446765

- (1) This is a wonderful review on peptic ulcer and the effect of natural product.

[Answer]

Thank you so much.

- (2) Since this is a review article on natural product for peptic ulcer, other natural product should be commented (*Lafoensia pacari*, *Arctium lappa*, *Luffa acutangula*, etc).

[Answer]

As you commented, we have added other natural products in this revised manuscript. (Lines 324-357)

- (3) The authors described intracellular pathways on ulcer healing. It will be very helpful for the readers when the authors draw a picture for this pathway.

[Answer]

In accordance with the reviewer's suggestion, we have added a schematic representation of intracellular pathway on ulcer healing as shown in Figure 1

- (4) One of the most important reasons for Hp eradication is to prevent the recurrence of peptic ulcer. It was omitted in Abstract.

[Answer]

As you commented, we have added the sentence in Abstract (Lines 43-44) and more information in Lines 268-271.

- (5) In Introduction, should we consider antacid as a gastroprotectant?

[Answer]

Yes it is. In this revised manuscript, we added the following description in the text of Introduction (Lines 101-105); "Antacids talcid activates EGF and EGF-R expression in normal and ulcerated gastric mucosa. Since EGF and EGF-R are crucial for cell proliferation, migration,

re-epithelialization and gland reconstruction within the scar. In addition, antacids, especially hydrotalcite are due to activation of prostaglandin synthesis; binding to and inactivation of pepsin, lysolecithin, bile acids and *H. pylori* toxins; activation of heat shock proteins and, activation of the genes encoding EGF, bFGF and their receptors."

- (6) Spelling errors: 1) Abstract: "gastroprotectant" -> "gastroprotectant" 2) References: bold type letters in Reference 2

[Answer]

As you commented, we have corrected.

Reviewer 00053723

- (1) Authors must include at least 4 figures or schemes 1) about ulcer development (and the cascade of inflammation, loss of mucosa integrity, etc) 2) traditional therapy of ulcer 3) healing process (including the cascade of events that can be modulated by therapeutic agents). 4) protective effect of natural product with emphasis in their antioxidant and anti-inflammatory effects the literature about other plants protective effects must be included (preferentially a n extensive revision of the last 5-10 years must be included). Otherwise, the interest in this microreview will be small.

[Answer]

In accordance with the reviewer's suggestion, we have added a figure including ulcer development, traditional therapy of ulcer and ulcer healing process as shown in Figure 1. In addition, we have added the effects of other natural products on the ulcer healing in this revised manuscript in Lines 324-357.

Reviewer 00504722

- (1) An extensive review of experimental (animal) data of ulcer formation and healing, and even natural products may be beneficial in enhancing quality of ulcer healing. The manuscript needs careful proof reading to exclude typographical errors (QOUH, QUOH), and other writing issues, etc ..

[Answer]

Thank you so much.

As per reviewer's suggestion, we have modified the manuscript very carefully and added more information about the animal data of ulcer formation and healing in Lines 190-205.

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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