

## PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 34562

**Title:** Dihydromyricetin-mediated inhibition of the Notch1 pathway induces apoptosis in QGY7701 and HepG2 hepatoma Cells

**Reviewer's code:** 02860871

**Reviewer's country:** Indonesia

**Science editor:** Yuan Qi

**Date sent for review:** 2017-05-19

**Date reviewed:** 2017-05-22

| CLASSIFICATION                                    | LANGUAGE EVALUATION   | SCIENTIFIC MISCONDUCT                          | CONCLUSION   |
|---|---|--|--|
| <input type="checkbox"/> Grade A: Excellent       | <input type="checkbox"/> Grade A: Priority publishing                 | Google Search:                                 | <input type="checkbox"/> Accept                        |
| <input type="checkbox"/> Grade B: Very good       | <input checked="" type="checkbox"/> Grade B: Minor language polishing | <input type="checkbox"/> The same title        | <input type="checkbox"/> High priority for publication |
| <input type="checkbox"/> Grade C: Good            |   | <input type="checkbox"/> Duplicate publication |  |
| <input checked="" type="checkbox"/> Grade D: Fair | <input type="checkbox"/> Grade C: A great deal of language polishing  | <input type="checkbox"/> Plagiarism            | <input type="checkbox"/> Rejection                     |
| <input type="checkbox"/> Grade E: Poor            | <input type="checkbox"/> Grade D: Rejected                            | <input type="checkbox"/> No                    | <input type="checkbox"/> Minor revision                |
|   |   | BPG Search:                                    | <input checked="" type="checkbox"/> Major revision     |
|   |   | <input type="checkbox"/> The same title        |  |
|   |   | <input type="checkbox"/> Duplicate publication |  |
|   |   | <input type="checkbox"/> Plagiarism            |  |
|   |   | <input type="checkbox"/> No                    |  |

## COMMENTS TO AUTHORS

This article entitled "Dihydromyricetin-mediated inhibition of the Notch1 pathway induces apoptosis in QGY7701 and HepG2 hepatoma Cells" " by Lu et al is interesting in its field. The authors investigate whether DHM inhibits cell proliferation and promotes apoptosis by downregulating Notch1 expression. This article is well written with few typos inside. Here are some comments : Major comments: 1. Activation of Notch signaling is normally tightly controlled by direct interactions with ligand-expressing cells. It would be better if author also analyze the Notch ligand's activity in their cells or samples. 2. Please provide more result of the profile of Notch receptors and target genes in the samples 3. There is an anomaly in patient number 2, which was Notch 1 is significantly higher in liver than in tumor. What is characteristic of this patient and what might be the possible explanation for this anomaly. 4. Author mentioned that DHM inhibits cell proliferation and promotes apoptosis by downregulating Notch1 expression. Please prove that by silencing the Notch1 genes may attenuate the inhibitory effects of



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DHM on the cells. Minor comments: 1. For figure 1a, please provide also the result of protein analysis for Hes1. 2. In figure 5, please provide the effectiveness of Notch1 siRNA 3. Please provide the patients characteristics for all the 9 samples

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**Title:** Dihydromyricetin-mediated inhibition of the Notch1 pathway induces apoptosis in QGY7701 and HepG2 hepatoma Cells

**Reviewer's code:** 00187828

**Reviewer's country:** Afghanistan

**Science editor:** Yuan Qi

**Date sent for review:** 2017-05-19

**Date reviewed:** 2017-05-24

| CLASSIFICATION                              | LANGUAGE EVALUATION  | SCIENTIFIC MISCONDUCT                          | CONCLUSION   |
|---|--|--|--|
| <input type="checkbox"/> Grade A: Excellent | <input type="checkbox"/> Grade A: Priority publishing                | Google Search:                                 | <input type="checkbox"/> Y] Accept                     |
| <input type="checkbox"/> Grade B: Very good | <input type="checkbox"/> Y] Grade B: Minor language polishing        | <input type="checkbox"/> The same title        | <input type="checkbox"/> High priority for publication |
| <input type="checkbox"/> Grade C: Good      | <input type="checkbox"/> Grade C: A great deal of language polishing | <input type="checkbox"/> Duplicate publication | <input type="checkbox"/> Rejection                     |
| <input type="checkbox"/> Grade D: Fair      | <input type="checkbox"/> Grade D: Rejected                           | <input type="checkbox"/> Plagiarism            | <input type="checkbox"/> Minor revision                |
| <input type="checkbox"/> Grade E: Poor      |  | <input type="checkbox"/> No                    | <input type="checkbox"/> Major revision                |
|   |  | BPG Search:                                    |  |
|   |  | <input type="checkbox"/> The same title        |  |
|   |  | <input type="checkbox"/> Duplicate publication |  |
|   |  | <input type="checkbox"/> Plagiarism            |  |
|   |  | <input type="checkbox"/> No                    |  |

## COMMENTS TO AUTHORS

The manuscript entitled "Dihydromyricetin-mediated inhibition of the Notch1 pathway induces apoptosis in QGY7701 and HepG2 hepatoma Cells" by Lu et al., was evaluated as follows; This is an interesting study in which authors demonstrated that DHM downregulates the expression of Notch1, Hes1 and Bcl-2 in QGY7701 and HepG2 cells. However, Bax was upregulated after the depletion of Notch1 via siRNA. Consequently down-regulation of Notch1 activates the mitochondrial apoptotic pathway. These results show that the anti-tumor activity of DHM in the QGY7701 and HepG2 hepatocarcinoma cell lines is partially exerted through inhibition of the Notch1 pathway. In summary, our experiments confirmed that Notch1 is involved in the development of HCC and may serve as a potential diagnostic marker for HCC and an indicator of malignancy. DHM inhibits cell proliferation and promotes apoptosis by down-regulating the expression of Notch1, indicating that it can be used as a candidate drug for the treatment of HCC. This is the first study to demonstrate that DHM inhibits cell proliferation and promotes



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apoptosis by down-regulating the expression of Notch1. It is well-written and presented. However it needs language polishing.

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**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 34562

**Title:** Dihydromyricetin-mediated inhibition of the Notch1 pathway induces apoptosis in QGY7701 and HepG2 hepatoma Cells

**Reviewer's code:** 02861252

**Reviewer's country:** Turkey

**Science editor:** Yuan Qi

**Date sent for review:** 2017-05-19

**Date reviewed:** 2017-06-01

| CLASSIFICATION   | LANGUAGE EVALUATION  | SCIENTIFIC MISCONDUCT                          | CONCLUSION   |
|--|--|--|--|
| <input type="checkbox"/> Grade A: Excellent            | <input checked="" type="checkbox"/> Grade A: Priority publishing     | Google Search:                                 | <input checked="" type="checkbox"/> Accept             |
| <input checked="" type="checkbox"/> Grade B: Very good | <input type="checkbox"/> Grade B: Minor language polishing           | <input type="checkbox"/> The same title        | <input type="checkbox"/> High priority for publication |
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|  |  | <input type="checkbox"/> No                    |  |

## COMMENTS TO AUTHORS

Good work...