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### ESPS PEER REVIEW REPORT

**Name of journal:** World Journal of Hepatology

**ESPS manuscript NO:** 13290

**Title:** THE ROLE OF AUTOPHAGY IN DIFFERENTIAL SENSITIVITY OF HEPATOCARCINOMA CELLS TO SORAFENIB

**Reviewer code:** 02538244

**Science editor:** Ling-Ling Wen

**Date sent for review:** 2013-12-31 20:43

**Date reviewed:** 2014-01-23 08:27

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	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"/> Existing	<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"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> Existing	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

### COMMENTS TO AUTHORS

The authors have an idea to find Sorafenib(SFN) effects in HCC cell lines. To this end, they analyzed the changes of autophagy-related proteins in the cells, and found the different responsiveness to SFN involves autophagy signaling pathway. The paper is interesting, and I think would be of interest to readers of World Journal of Hepatology.

## ESPS PEER REVIEW REPORT

**Name of journal:** World Journal of Hepatology

**ESPS manuscript NO:** 13290

**Title:** THE ROLE OF AUTOPHAGY IN DIFFERENTIAL SENSITIVITY OF HEPATOCARCINOMA CELLS TO SORAFENIB

**Reviewer code:** 00680710

**Science editor:** Ling-Ling Wen

**Date sent for review:** 2013-12-31 20:43

**Date reviewed:** 2014-01-28 18:01

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> Existing	<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"/> No records	<input checked="" type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair		BPG Search:	<input type="checkbox"/> Minor revision
<input checked="" type="checkbox"/> Grade E: Poor	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Existing	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

## COMMENTS TO AUTHORS

The results presented in the manuscript seem interesting though very preliminary and controversial. The major problem is that the authors completely ignored the contradictory effect of sorafenib on Hep3B and Huh7 cell lines as described by Aditi Pandya Martin in Molecular Pharmacology (76:327-341, 2009). This is really very disturbing and makes it impossible to accept the manuscript at this stage. Moreover, the assessment of cellular viability based solely on MTT test seems insufficient, especially in context of contradictory results published by other authors. It would be strongly advisable to do additional tests evaluating apoptosis and necrosis observed in the population of Hep3B and Huh7 cell lines.

## ESPS PEER REVIEW REPORT

**Name of journal:** World Journal of Hepatology

**ESPS manuscript NO:** 13290

**Title:** THE ROLE OF AUTOPHAGY IN DIFFERENTIAL SENSITIVITY OF HEPATOCARCINOMA CELLS TO SORAFENIB

**Reviewer code:** 02861035

**Science editor:** Ling-Ling Wen

**Date sent for review:** 2013-12-31 20:43

**Date reviewed:** 2014-03-02 19:30

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> Existing	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair		BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Existing	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

## COMMENTS TO AUTHORS

The topic the authors trying to investigate is interesting and could potentially leads to future therapy for different type of cancer. However, the experimental design was too brief and simple and the detailed mechanisms not investigated further with an unclear conclusion. I would suggest the authors to make several additions to their experiments prior to publish. 1. The authors showed differential p-AMPK expression between Hep3B and Huh7. This is an interesting observation, the authors should try and knockdown/knockout LKB to investigate whether there will be a change in p-AMPK level in the Huh7 and does that affect sensitivity towards SFN and autophagy; or to induce p-AMPK in Hep3B to investigate the changes in autophagy

## ESPS PEER REVIEW REPORT

**Name of journal:** World Journal of Hepatology

**ESPS manuscript NO:** 13290

**Title:** THE ROLE OF AUTOPHAGY IN DIFFERENTIAL SENSITIVITY OF HEPATOCARCINOMA CELLS TO SORAFENIB

**Reviewer code:** 02860966

**Science editor:** Ling-Ling Wen

**Date sent for review:** 2013-12-31 20:43

**Date reviewed:** 2014-03-12 21:40

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	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 type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> Existing	<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"/> No records	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair		BPG Search:	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Existing	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

## COMMENTS TO AUTHORS

Thank you for the opportunity to review this paper.