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## ESPS PEER REVIEW REPORT

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

**ESPS manuscript NO:** 13216

**Title:** Involvement of the TAGE-RAGE system in NASH: A novel therapeutic strategy

**Reviewer code:** 00002726

**Science editor:** Fang-Fang Ji

**Date sent for review:** 2014-08-12 18:49

**Date reviewed:** 2014-08-29 03:12

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	<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 work is a timely review of the TAGE involvement in NAFLD. Specific comments 1. Fig. 1 is very busy and therefore difficult to understand. It should be simplified by removing the chemical formulas. 2. The review is long, and there is a lot of information related to diseases and cell types, particularly from that of the authors' work, that is not directly relevant to the liver. These sections should be cut and the review focused more on the liver. 3. The review is not well organized and quickly switches topics. For example, on page 12 the first paragraph starts about endothelial cells, then within the same paragraph hepatic stellate cells are discussed for a few sentences. The next paragraph discusses hepatocytes. The material should be better organized into more coherent subthemes. 4. There are a number of typos, for example, just on page 5: line 6 should read "more significantly;" line 13 should read "discuss;" line 19, "residues." The manuscript requires extensive editing. 5. There are a large number of abbreviations, some nonstandard (e.g., OS) and a list of abbreviations would be helpful.



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### ESPS PEER REVIEW REPORT

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

**ESPS manuscript NO:** 13216

**Title:** Involvement of the TAGE-RAGE system in NASH: A novel therapeutic strategy

**Reviewer code:** 00227368

**Science editor:** Fang-Fang Ji

**Date sent for review:** 2014-08-12 18:49

**Date reviewed:** 2014-08-13 03:06

CLASSIFICATION	LANGUAGE EVALUATION	RECOMMENDATION	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input checked="" 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 checked="" 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

None



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## ESPS PEER REVIEW REPORT

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

**ESPS manuscript NO:** 13216

**Title:** Involvement of the TAGE-RAGE system in NASH: A novel therapeutic strategy

**Reviewer code:** 00227509

**Science editor:** Fang-Fang Ji

**Date sent for review:** 2014-08-12 18:49

**Date reviewed:** 2014-08-16 13:05

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	<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

In this manuscript, Takeuchi and the colleagues summarized the involvement of TAGE-RAGE system in NASH using recent findings. The authors have been investigating about TAGE-RAGE system in various metabolic diseases, and advocating the vicious roles of TAGE-RAGE system as a new concept in metabolic syndrome. In this review article, the authors well summarized about TAGE-RAGE system in NASH. However, there are some concerns remain as followings.

Comments 1. In this manuscript, the authors represented that liver function inversely correlated with serum TAGE levels in chronic liver diseases, and also represented that serum TAGE levels decreased in NASH patients by using anti-DM drugs. Which is more important in each liver disease (virus associated liver diseases, NAFLD)? Liver functions or glucose tolerance? Please explain precisely. 2. The authors showed that HMG-CoA reductase inhibitor, statins, decreased serum TAGE levels. What is the putative mechanism that decrease TAGE levels by statins? Is this secondary effects of statins? 3. In conclusion, the authors described serum TAGE levels were significantly higher in NBNC-HCC. How about other HCC patients (HCV, HBV associated HCC)? 4. After (TBARS), “,” is missing (Page 25, line 10).