

**ESPS Peer-review Report**
**Name of Journal:** World Journal of Gastroenterology

**ESPS Manuscript NO:** 9682

**Title:** Potential Mechanism(s) of Hepatitis B virus (HBV) Induced Liver Injury

**Reviewer code:** 02860618

**Science editor:** Yuan Qi

**Date sent for review:** 2014-02-24 13:22

**Date reviewed:** 2014-02-28 19:12

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 checked="" type="checkbox"/> Grade B: minor language polishing	<input type="checkbox"/> Existed	<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"/> Rejection
<input type="checkbox"/> Grade E (Poor)	<input type="checkbox"/> Grade D: rejected	<input type="checkbox"/> Existed	<input checked="" type="checkbox"/> Minor revision
		<input type="checkbox"/> No records	<input type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

The review by Mohd Suhail and co-workers deals with the mechanisms that cause liver injury during Hepatitis B Virus (HBV) infection. In the present manuscript they summarize the role played by the immune system, oxidative and endoplasmic stress, hypoxia and angiogenesis, as well as HBV-DNA integration and individual viral proteins, in HBV-mediated liver injury. Even though this seems to be a complete overview on this topics, I suggest some revisions before publication (please see the attached word file).

1. Please indicate some more keywords.
2. I recommend to change the contents and chapter titles as follows:

***Introduction***

*General Background*

*Various Genotypes and their role in liver disease*

***Pathogenesis of liver injury***

*Role of altered innate and adaptive immunity*

*Role of oxidative and endoplasmic stress*

*Role of Hypoxia and angiogenesis*

*Integration of hepatitis B virus DNA allows persistence of the virus and induces genetic alterations*

*Role of individual viral proteins in liver injury*

*HBx protein*

*Pre core protein*

*Surface protein*

*Polymerase protein*

*Conclusion*

*Acknowledgments*

*References*

3. Summary, line 2: is CAH recognized as a cause of liver injury or liver failure?
4. Summary, line 13: please check for the verb “dictated”. Did you mean “resulted in the alteration of”?
5. Page 7, lines 21-22: please check “In HBV endemic areas, infected patients 80-90% of HCCs of all etiological backgrounds arise in cirrhotic livers”. The meaning of this sentence is not clear.
6. Page 7, line 27: what is the meaning of “tolerable” in this sentence?
7. Page 9, lines 8-9: please check for “bout of hepatitis and regeneration, resulting in outgrowth of HBx-positive hepatocytes.” (incomplete sentence)

8. Page 14, line 6: please check for “...attenuate the host immune response to the”  
(incomplete sentence)
9. Please cite the figure 1 in the manuscript.
10. Page 4, lines 10-13: Reference 2 is not the proper citation; the citation n.5 would be more suitable.
11. Page 4, line 29: please provide a reference for ground squirrels.
12. The references 57, 59, 67 and 68 are about HCV.
13. Page 8, line 1: please convert “Rehermann 2005” in a proper reference.
14. Page 11, lines 25-29 and page 12, lines 1-2: please insert one or more references.
15. Please extend the following abbreviations in the manuscript: PC and BPC (in table 1), PBMCs (page 8, line 17), IRAK4, TRAF3 and IRF7 (page 8, line 19), HIF1a (page 11, line 1).
16. Please check for several grammar and typo's errors along the whole manuscript: for instance, lines 6 and 10 of the summary (“processes” instead of “processed” or “process”), and line 16 of the summary (“hepatocytes”), “Kupffer” on page 7, “CONCLUSION” on page 16.
17. There are lots of double spacing between words; moreover, the Authors do not use hyphenated words (HBV-induced, immune-mediated...) and the choice to use brackets for “mechanism(s)” or “protein(s)” is confusing.

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

**ESPS Manuscript NO:** 9682

**Title:** Potential Mechanism(s) of Hepatitis B virus (HBV) Induced Liver Injury

**Reviewer code:** 02861134

**Science editor:** Yuan Qi

**Date sent for review:** 2014-02-24 13:22

**Date reviewed:** 2014-03-02 22:50

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"/> Existed	<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	<input type="checkbox"/> Existed	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E (Poor)		<input type="checkbox"/> No records	<input type="checkbox"/> Major revision

**COMMENTS TO AUTHORS**

Report of Reviewer D-r Mariana Penkova Radicheva, MD,PhD Manuscript Number 9682  
 Manuscript Title Potential Mechanism(s) of Hepatitis B virus (HBV) Induced Liver Injury The paper makes original contribution. The paper is well organized. Author Guidelines has been followed properly in preparing the manuscript. The paper is based on sound methodology. Literature review is adequate. Analysis and findings support objectives of the paper. Decision regarding the paper - Accept the paper in its current format. Starting part of this paper is excellent, specially the abstract. It is concise and organized. The study is a timely research. Objectives are consistent with literature review and analysis. The paper can be recommended to publish with the prior approval of the editorial board. 02.03.2014

# ESPS Peer-review Report

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

**ESPS Manuscript NO:** 9682

**Title:** Potential Mechanism(s) of Hepatitis B virus (HBV) Induced Liver Injury

**Reviewer code:** 02860590

**Science editor:** Yuan Qi

**Date sent for review:** 2014-02-24 13:22

**Date reviewed:** 2014-03-06 07:49

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"/> Existed	<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"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

## COMMENTS TO AUTHORS

In this review the authors described the mechanisms involved in the hepatic injury caused by hepatitis B virus (HBV). Although the theme is important there are some points that have to be reviewed regarding this manuscript. Specific comments: 1. Title: There are not spaces between words. In the title page, e.g., Potential mechanism (s). 2. Contents: The author(s) use several forms to write liver injury (e.g. liver injury, Liver injury, Liver Injury) please use one of this forms in the manuscript. Line 11: Liver injury should be reviewed due to grammatical error. 3. Summary: Line 12: Please, the verb "dictated" should be reviewed due to inappropriate use. 4. Introduction: Page 4, Line17-18, Please, clarification should be provided for this phrase. Page 6, table 1, Table 1 title should be adjusted to the table width. Legends should contain sufficient information to provide an adequate understanding of the table by the reader without reference to the text. Please extend the following abbreviations in the inferior legend: HCC, PC, BCP. "outcome Unknown" should be reviewed. Page 7, line 16, "In HBV endemic areas,.... Please, the sentence should be reviewed due to poor grammatical construction. Page 7, line 25, Please, change "IFN remedy" to "IFN therapy". Page 8, line 17, "in the all infected HBV patients", Did you consider only the chronic infection? Page 9, line 2, Please, the sentence should be reviewed due to grammatical error. Page 9, line 5, Please, the sentence should be reviewed due to poor grammatical construction (incomplete phrase). Page 9, line 25, Please, the sentence should be reviewed due to poor grammatical construction. Page 14, line 2, Please, the sentence should be reviewed due to poor grammatical construction (incomplete phrase). Page 14, line 28, There is not space between words "mitogen activated". Page 15, line 18-20, Please, the sentence should be reviewed due to poor grammatical construction. 5. Conclusion: Please, the word



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“conclusion” should be reviewed due to grammatical error. Line 11: Please, the word “insult” should be reviewed due to inappropriate use. 6. Reference: Please, the references should be reviewed. Some references are not in accordance with the subject described (references: 57, 59, 67 and 68). Reviewer conclusion: Accept but needs revision (minor).



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### ESPS Peer-review Report

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

**ESPS Manuscript NO:** 9682

**Title:** Potential Mechanism(s) of Hepatitis B virus (HBV) Induced Liver Injury

**Reviewer code:** 02860705

**Science editor:** Yuan Qi

**Date sent for review:** 2014-02-24 13:22

**Date reviewed:** 2014-03-07 18:22

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"/> Existed	<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"/> Existed	<input type="checkbox"/> Major revision
		<input type="checkbox"/> No records	

### COMMENTS TO AUTHORS

The review is well written and clear.