



**ESPS PEER-REVIEW REPORT**

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

**ESPS manuscript NO:** 20100

**Title:** Naturally derived anti-hepatitis B virus agents and their mechanism of action

**Reviewer’s code:** 00503536

**Reviewer’s country:** Japan

**Science editor:** Ya-Juan Ma

**Date sent for review:** 2015-05-31 16:10

**Date reviewed:** 2015-08-22 14:18

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 checked="" type="checkbox"/> Grade C: Good		<input type="checkbox"/> Duplicate publication	
<input 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 checked="" 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 checked="" type="checkbox"/> No	

**COMMENTS TO AUTHORS**

The review manuscript written by Wu et al. describes various natural anti-HBV agents. Since HBV infection is still a serious problem around the world, the data are interesting and important. However there are some concerns that need to be addressed. Major points 1. The most important problems of current treatment for HBV infection is that any anti-HBV agents has no effect on cccDNA, leading to relapse of HBV replication after discontinuation of the treatment. It is of note that the authors show some natural agents decreased cccDNA in the liver. If that is also true for human HBV infection, chronic HBV infection can be terminated in a drug-free manner. The authors should state more on the effect and mechanism of the action. 2. Emergence of treatment-resistant HBV mutants after long-term drug administration is another problem in HBV treatment. Long term antiviral effect of those natural agents should be shown. 3. Are there dose-dependent effects of each anti-viral agent? 4. Toxicities of each agent should be stated. Minor point 1. HepG2.2.15 cells should have no cccDNA.

## ESPS PEER-REVIEW REPORT

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

**ESPS manuscript NO:** 20100

**Title:** Naturally derived anti-hepatitis B virus agents and their mechanism of action

**Reviewer's code:** 00053556

**Reviewer's country:** Egypt

**Science editor:** Ya-Juan Ma

**Date sent for review:** 2015-05-31 16:10

**Date reviewed:** 2015-09-10 16:05

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 checked="" type="checkbox"/> Grade C: Good		<input type="checkbox"/> Duplicate publication	
<input 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 checked="" 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 checked="" type="checkbox"/> No	

### COMMENTS TO AUTHORS

Comments to the Editor: Thanks for inviting me to review the review article entitled "Naturally derived anti-hepatitis B virus agents and their mechanism of action". Minor Comment: ? Major revision is required concerning language in term of grammar and structure. ? Language evaluation: "B". 1. TITLE reflect the major content of the article. 2. ABSTRACT Fulfil the journal requirements. The aim of the work is clearly identified; however, a conclusion is better to be included 3. TEXT: The section is almost well organized; an overall theoretical analysis concerning the provided data is well covered to great extent, however and in order to satisfy the reader, the authors are better to describe some important data regarding: ? Citations regarding the problem of HBV worldwide (Ref.:1-5) are better to be updated. The provided data are not accurate for ref. No: 3. ? Some important wards have to be fully written when mentioned for the first time ;e.g: DHBV, IC50, EMSA analysis ? The authors discussed the mode of action of several antiviral without identifying the level of studying the antiviral activity; in vitro or in vivo studies. This has to be maintained. ? The antiviral activities of the illustrated agents are better to be mentioned in comparable to positive control. ? An informative descriptive table or a table for each studied group is better to be included to



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show the group, structure, mode of action, experimental level, and IC50 value of all the studied antivirals. 5. REFERENCES: Relevant and sufficient updated references, especially the most current literatures were cited (42/70) references were cited from publications  $\geq$  2010. The journal style for writing this section is well maintained. However, Reference No: 1-5 concerning the problem of HBV have to be updated.