



**PEER-REVIEW REPORT**

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

**Manuscript NO:** 43097

**Title:** Serum hepatitis B virus RNA is a predictor of HBeAg seroconversion and virological response with entecavir treatment in chronic hepatitis B patients

**Reviewer’s code:** 00071703

**Reviewer’s country:** Turkey

**Science editor:** Ruo-Yu Ma

**Date sent for review:** 2018-12-03

**Date reviewed:** 2018-12-07

**Review time:** 14 Hours, 4 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input checked="" type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good		<input checked="" type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	(General priority)	Peer-reviewer’s expertise on the topic of the manuscript:
<input type="checkbox"/> Grade E: Do not publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Minor revision	<input checked="" type="checkbox"/> Advanced
		<input type="checkbox"/> Major revision	<input type="checkbox"/> General
		<input type="checkbox"/> Rejection	<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

**SPECIFIC COMMENTS TO AUTHORS**

Serum HBV RNA is a predictor of HBeAg seroconversion and virological response with entecavir treatment in CHB patients In this study, the authors aimed to investigate whether HBV RNA can predict virological response or HBeAg seroconversion during



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ETV treatment when HBV DNA is undetectable. Thirty treatment-naive individuals were included in the study. They found that HBeAg seroconversion was more likely in CHB patients with HBV RNA levels below 4.12 log<sub>10</sub> copies/ml before treatment. They have concluded that Serum HBV RNA can predict HBeAg seroconversion and partial virological response during treatment. In the partial virological response group, serum HBV RNA showed an increasing trend. I think this is a good study, and the manuscript is well written; also the tables are all appropriate.

#### **INITIAL REVIEW OF THE MANUSCRIPT**

##### ***Google Search:***

- The same title
- Duplicate publication
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**PEER-REVIEW REPORT**

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

**Manuscript NO:** 43097

**Title:** Serum hepatitis B virus RNA is a predictor of HBeAg seroconversion and virological response with entecavir treatment in chronic hepatitis B patients

**Reviewer's code:** 02445074

**Reviewer's country:** United States

**Science editor:** Ruo-Yu Ma

**Date sent for review:** 2018-12-18

**Date reviewed:** 2018-12-19

**Review time:** 20 Hours, 1 Day

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input checked="" type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
<input type="checkbox"/> Grade E: Do not	language polishing	<input checked="" type="checkbox"/> Minor revision	topic of the manuscript:
publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input checked="" type="checkbox"/> Advanced
		<input type="checkbox"/> Rejection	<input type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

**SPECIFIC COMMENTS TO AUTHORS**

**GENERAL COMMENTS:** Other studies have shown that in HBV patients while HBV DNA is found in serum, there is at levels at least 1,000-fold lower, HBV RNA. And, yet other studies have shown that if HBV DNA levels are lower, then antiviral treatments



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are more likely to succeed. The present studies would like to test/assert that serum HBV RNA, although more difficult to detect, might be a better way to predict/confirm success in antiviral therapies. A major problem of the present study is the limited number of patients with which to test the hypothesis. The accepted patient groups contain only 10 and 15 individuals, with 5 uninfected controls. This major problem remains unsolved even though the authors have provided an heroic attempt at evaluating data for statistical significance. What might be a second problem is the specificity of the RNA assays. The assays might yet be detecting residual HBV DNA. That is, since the HBV DNA is initially in an excess of at least 1,000 fold, the RNA isolation/assay must be much greater than 99.9% greater in removing/detecting DNA. This reviewer did not see control data, obtained in the absence of the reverse transcription step. **SPECIFIC COMMENTS:** Abstract Remove use of first person. Introduction In 1996 Kock first demonstrated that ... HBV RNA levels in serum were ... characteristics of alterations of serum HBV RNA were ... Patients The criteria for inclusion were: (a) patients ... The criteria for exclusion were: ... Informed consent was obtained for all enrolled patients and the study protocol ... by the Ethics Committee ... Results (83%) (37 +/- 6 vs. 33 +/- 12) (3.1 +/- 1.4 vs. 5.5 +/- 1.5) Not convincing that this is a statistically significant difference!! various personal background!! Define flRNA and trRNA Table 3 HBeAg positive

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**PEER-REVIEW REPORT**

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

**Manuscript NO:** 43097

**Title:** Serum hepatitis B virus RNA is a predictor of HBeAg seroconversion and virological response with entecavir treatment in chronic hepatitis B patients

**Reviewer's code:** 02994003

**Reviewer's country:** Egypt

**Science editor:** Ruo-Yu Ma

**Date sent for review:** 2018-12-18

**Date reviewed:** 2018-12-21

**Review time:** 13 Hours, 3 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
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<input checked="" type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
<input type="checkbox"/> Grade E: Do not	language polishing	<input checked="" type="checkbox"/> Minor revision	topic of the manuscript:
publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input type="checkbox"/> Advanced
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			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

**SPECIFIC COMMENTS TO AUTHORS**

Name of Journal: World Journal of Gastroenterology Manuscript Type: OBSERVATIONAL STUDY Serum HBV RNA is a predictor of HBeAg seroconversion and virological response with entecavir treatment in CHB patients Generally, the idea



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of this study is good and the manuscript is well written although it needs minor error editing Title is ok Abstract is simple and please change word discussion to conclusion Key words are enough Introduction, materials and methods are clear. Results contains enough tables and figures but there is several error editing in tables- Please what about relation between HBV DNA and HBsAg Discussion is ok although it is better if it is more longer with more references

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