

## Manuscript Number 22661

### Host Nucleotide Polymorphism in HBV-Associated Hepatocellular Carcinoma

#### **Reviewer 1:**

- 1) The review paper addresses an interesting topic. However, the manuscript is poorly written and not well structured. For instance, the introduction is rather shallow and does not introduce the reader to the topic at hand. The three first sections of the manuscript (HBV genotypes and its Global distribution, Common genotyping methods for HBV, HBV gene mutations) are completely disconnected from the title.

**Response:** The three sections of the manuscript (HBV genotypes and its Global distribution, Common genotyping methods for HBV, HBV gene mutations) are removed as suggested.

- 2) The second part of the paper is the main subject for this review paper and it should be the highlight of the whole manuscript.

**Response:** The second part of the manuscript is highlighted and enhanced more as suggested by the reviewer.

- 3) The authors should be encouraged to include few schematic diagrams that will facilitate reading and overall comprehension of their manuscript.

**Response:** A schematic diagram is also included as suggested by the reviewer.

- 4) The conclusions should be rewritten from scratch to capture the real essence of the paper.

**Response:** The discussion and conclusion part is revised as suggested by the reviewer.

- 5) The paper should be proofread by an English native speaker before resubmitting the manuscript to the journal.

**Response:** The revised manuscript is proofread by expertise English speakers.

**Reviewer 2:**

This review deals with not only the relationship between genotypes of HBV and development of hepatocellular carcinoma but also the relationship between HBV infection and host nucleotide polymorphism. This manuscript is well organized so that we easily follow and understand the context. We obtain very useful information without searching further papers in this review. We strongly recommend the review to publication in your journal. However, we found many typo errors. 1. page 10 : line 3 cyclooxygenade-2 -> cyclcoxygenase-2 line 7 in vitro and in vivo -> intalic line 15-16 ; please check that the sentence is correct. “in which “ (?) line 22 : IL-1a and Ib -> IL-1 alpha and 1beta 2. page 14; line 21 : Ebstein -> Epstein 3. page 15; line 5 mRNA[172] -> mRNA [172] 4. page 16 line 8 : (TIM3)-1516 G>T -> delete “1567 G>T” Line12 and line 13 : Tim-> TIM3 Line 16 : TIM 3-> TIM3 because you described “gene name” as “Times New Roman style”. Please unify. 5. page 16 line 21 : (XPC) Codon K939Q -> (XPC) ; delete “Codon K939Q” 6. In Table 1, please delete “ and”.

**Response:** The manuscript is further improved and we corrected many typo errors as suggested by the reviewer.