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Name of Journal: *World Journal of Hepatology*

ESPS Manuscript NO: 26075

Manuscript Type: Review

Response to reviewer's comments (highlighted with green color in the manuscript):

- 1- the whole words were written before abbreviation” hepatitis C virus (HCV) related hepatocellular carcinoma (HCC)”
- 2- The sentence “HCV can induce HCC through direct effects of its protein or through indirect way which is inflammation of liver tissue and its complication as cirrhosis which form the background for HCC in most of HCV –HCC patients” was divided into “HCV can induce HCC through direct effects of its protein or through indirect way. The indirect way occurs as inflammation of liver tissue and /or its complication as cirrhosis which form the background for HCC in most of HCV –HCC patients”.
- 3- The sentence “The viral particle is formed by a nucleocapsid, consisting of the core protein and viral genome and an envelope consisting of envelope glycoproteins E1 and E2”is changed into “The viral particle is formed by a nucleocapsid consisting of the core protein ; viral genome and envelope glycoproteins E1 and E2.”.
- 4- The sentence” These pathways are known to be activated in HCC “ was explained as “Moreover, signaling pathways such as Raf/MAPK, [19] Wnt/ β -catenin, 41 and TGF- β can be stimulated by HCV core protein [20, 21]. The inclusion of different pathways is known to be activated in HCC and may help in progression of cirrhosis process; induce mutation for one or more oncogenes or tumor suppressor genes”.
- 5- The sentence “Interestingly, wild-type NS5A genes were up regulated among HCC patients with liver cirrhosis as background Compared with those who did not” was changed into “Interestingly, wild-type NS5A gene was up regulated among HCC

patients with liver cirrhosis as background Compared with those who did not develop HCC”.

- 6- The sentence” The indirect way for hepatocarcinogenesis during HCV infection is inflammation , persistence of chronic hepatitis, liver fibrosis, cirrhosis ending to malignancy transformation “ was changed into “The indirect way for hepatocarcinogenesis during HCV infection is inflammation of hepatocytes, persistence of chronic hepatitis, liver fibrosis and cirrhosis ending to malignancy transformation”.
- 7- The paragraph ” There is a long way to go in controlling liver carcinogenesis. surgical resection and liver transplantation , the important procedures for HCC therapy, are the most curative options to treat HCC-----“ is important for clarification of importance of genetics in early detection and therapeutic intervention in HCC. This was found in the following paragraph: “Finding treatments for advanced stage cancer isn’t easy. Therefore, early detection methods are paramount in the fight against the disease. A major endeavor to develop novel treatments should involve the implementation of genetic profiling to describe tumors and provide accurate predictions as well as potential therapeutic targets during the process of HCC and an overall improvement in targeted therapies”.
- 8- The company of the panel was given as above

Response to of editor`s comments (highlighted with yellow color in the manuscript):

- Name of journal was added
- Type of manuscript was added.
- Affiliation of authors were corrected.
- Address and telephone of corresponding author were corrected.