

December 19, 2013

Dear editor,

I really appreciate several important comments of two reviewers on my manuscript (Manuscript revision NO: 6715). My responses to reviewers' comments are as follows. I highlighted the changes in my manuscript. Please find enclosed the edited manuscript in Word format. (file name : WJG Review 6715-edited (I.Nakamura) (2013-12).docx).

Title : Impairment of Innate Immune Responses in Cirrhotic Patients and Treatment by Branched-Chain Amino Acids

Author: Ikuo Nakamura

Name of Journal : *World Journal of Gastroenterology*

ESPA Manuscript NO: 6715

<Reviewer 00506513>

- 1. Description of chemical formula of BCAA was added on page 9.**
- 2. Figure 1 was deleted. Alternatively, Table 1 (Impairment of innate immune responses in cirrhotic patients) and Table 2 (Impairment of adaptive immune responses in cirrhotic patients) were produced**
- 3. Several descriptions were revised. (p4, p5, p6, p7, p8, p9 and p13)**

<Reviewer 01560505>

1. The following descriptions were inserted on page 7. <Reduction of NK cell activity might occur partially due to lower frequency of NK cell in peripheral blood^[13]. However, the mechanisms of diminished NK cell activity was not clarified so far. >

2. Other speculated mechanisms of reduction of the risk of HCC development by BCAA are (1) suppression of vascular endothelial growth factor (VEGF)

expression, (2) Induction of apoptosis of tumor cells through mTORC1 and mTORC2-dependent mechanisms. Details were already described in page 12 and 13.

3. On page 6, the pointed description was revised. <Polymorphonuclear cells (PMNs)>

Thank you again for publishing my manuscript in the *World Journal of Gastroenterology*

Sincerely yours,



Dec. 19, 2013

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