

January 28,2015

Dear Editor,

Please find enclosed the edited manuscript in word format (file name:15644-review.doc)

Title :Role of biomarkers in prediction and diagnosis of hepatocellular carcinoma (HCC).

Authors: Mahmoud Khattab ,Magdy Fouad , Elham Ahmed .

Name of Journal:World Journal of Hepatology.

ESPS Manuscript NO:15644

The manuscript has been improved according to the suggestions of reviewers:

1-Format has been updated ,in accordance to directions of administration provided at the edited manuscript.

2-Revision has been made according to the suggestions of reviewers :

(1)Reviewer 1: 00503202:

-All presented markers have been revised to include accuracy indices namely : cut off values ,sensitivity ,specificity ,and AUC for each one when available.

-Combinations between more than one of the most robust biomarkers have been added with increased accuracy of prediction of HCC in cirrhotic patients.

-A table including the most relevant biomarkers with their cut off values and accuracy indices has been added.

-To illustrate the role that may be displayed by biomarkers in diagnosis of HCC ,an edited algorithm utilising biomarkers has been suggested .

-Language has been revised .

(2)Reviewer 2: 02440197:

-All presented biomarkers have been revised with addition of their significant accuracy indices in diagnosis of HCC in cirrhotics together with multiple combinations of biomarkers that increased accuracy of prediction and diagnosis of HCC.

-Regarding your suggestion about clinical studies, of course, this can be the target for future clinical trials that validate one or more of these available biomarkers and/or combinations of more than one .

-Language revision has been done.

(3)Reviewer 3: 02444773 :

-Thank you sir for your suggestion .An edited algorithm utilising biomarkers in diagnosis of HCC has been added to the edited manuscript.

-Revision of the the presented biomarkers to include their accuracy indices (sensitivity ,specificity ,and AUC)at their cut off values has been done.

-language revision has been done.

(4)Reviewer 4 : 00068273 :

-The presented markers in this review article have been revised to include only the most robust biomarkers in prediction and diagnosis of HCC in cirrhotic patients . Furthermore ,multiple combinations with higher accuracy indices have been added to provide meaningful awareness of the relevant biomarkers that can be utilised in an algorithm for diagnosis of HCC.

-Still a marker with strong predictive score for HCC even in cirrhotics without evident focal lesion ,is awaited and a subject for future research.

-Language revision has been done.

3-References and typesetting were corrected according to the Journal rules .

Thank you again for publishing our manuscript in the *World Journal of Hepatology*.

Sincerely yours,



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Dear Fang-Fang Ji ,
Science Editor,BPG,

Greetings, Apologise for delay in response.

Please find attached is our 15644 manuscript after being re-written according to comments of Editoreal authority. The manuscript has been edited by American Journal Experts (AJE) and attached is the AJE certificate that indicate the journal editing of our manuscript, hoping that this version of the manuscript goes with your journal requirements for publication.

Best Regards,
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manuscript 15644
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