

PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05516254 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Iran

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-02-10 06:16

Reviewer performed review: 2021-03-05 10:09

Review time: 23 Days and 3 Hours

Scientific quality	[] Grade A: Excellent [Y] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [Y] Accept (General priority) [] Minor revision [] Major revision [] Rejection
Re-review	[]Yes [Y]No
Peer-reviewer	Peer-Review: [Y] Anonymous [] Onymous
statements	Conflicts-of-Interest: [] Yes [Y] No



SPECIFIC COMMENTS TO AUTHORS

This is a well designed and very useful manuscript. It just needs a little editing polishing.



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05933767 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: China

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-03-24 05:21

Reviewer performed review: 2021-03-25 02:52

Review time: 21 Hours

Scientific quality	[] Grade A: Excellent [Y] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [Y] Accept (General priority) [] Minor revision [] Major revision [] Rejection
Re-review	[]Yes [Y]No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

Specific Comments To Authors: 6MWT is often used to evaluate the prognosis of patients with heart disease and chronic obstructive pulmonary disease. There are few studies on the correlation between 6MWT and cirrhosis. This study has certain clinical value. However, the manuscript needed to be revised. 1 The manuscript needs to be revised in the format required by the magazine. 2 The participants were all from a liver transplantation center; however, the manuscript does not indicate whether the patients had ever undergone a liver transplant. "Whether patients who have previously received a liver transplant have been excluded? " should be included in the exclusion criteria. 3 The manuscript mentions that "The study has been performed in accordance with The Declaration of Helsinki (2000) and approved by The Ethics Committee of our Institution". But no ethics statement was submitted.



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05769843 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: France

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-03-24 09:26

Reviewer performed review: 2021-03-29 13:37

Review time: 5 Days and 4 Hours

Scientific quality	[] Grade A: Excellent [] Grade B: Very good [Y] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [] Accept (General priority) [Y] Minor revision [] Major revision [] Rejection
Re-review	[Y] Yes [] No
Peer-reviewer	Peer-Review: [Y] Anonymous [] Onymous
statements	Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

World Journal of Hepatology MANUSCRIT NO: 64130 Comments to the Author General's comments: The manuscript is an original work and the topic is important. This study retrospectively evaluated the six -minute walk test (6MWT) in 106 outpatients with liver cirrhosis with a short follow up of one year. They reported that a decreased 6MWT is associated with CHILD PUGH score and that 6MWT was an independent predictor of death. And tolerated in selected patient (pas trop grave) Comments: 1) Methods: a. Can the authors specify why patients should have stopped alcohol for 6 months before enrollment and why do they exclude recent decompensations. b. Do they exclude all patients with alcohol peripheral neuropathy, arterial obliterating disease, or cardiac dysfunction (TTechocardiography results)? c.

Did some patients have pulmonary complication of cirrhosis? d. Do all patients were included for survival analysis (even the 3 who had liver transplantation?), censured or not at liver transplantation? e. How was cirrhosis diagnosed (histologically for all patients? Morphological criteria?) f. Proportion of others cause of cirrhosis high, could the author's precise causes, do some patients had several cause of cirrhosis? 2)

Results: a. Did 6MWD differ according to cardiovascular features? Pulmonary arterial pressure? Existence of hepatopulmonary shunt? b. Does 6MWD predict survival independently of MELD score (multivariate logistic regression)? If yes, what about a survival analyses (log rank) according to low MELD and 6MWD > 387 vs high MELD and 6MWD < 387 c. Do the authors could specify specificity and specificity of two cut off they observed?



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05311389 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor, Research Fellow

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-03-25 11:50

Reviewer performed review: 2021-03-30 11:43

Review time: 4 Days and 23 Hours

Scientific quality	[] Grade A: Excellent [] Grade B: Very good [Y] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [] Accept (General priority) [] Minor revision [Y] Major revision [] Rejection
Re-review	[Y]Yes []No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

Physical fitness and functional status in cirrhotic patients are two main features related to survival in this cohort of patients. It is very interesting the attention payed on a simple way (eg. 6MWT) to estimate the functional impairment. Previous studies focused on more complex methods, such as the psoas muscle volume on CT scan, but this approach is not widely applied in clinical practice. I have few suggestions: ENGLISH (it should generally be implemented): Line 10: "RECENT studies" Line 22: "Chronic obstructive PULMONARY disease" Line 26: "There ARE also evidenceS...." Line 31-32: "The aim of this study was to analyze the association between 6MWT and long-term mortality IN A COHORT OF CIRRHOTIC PATIENTS" Line 35-36: A total of 106 outpatients with liver cirrhosis (57 male) WAS included in the present study" Line 61: Patients were stratified according TO their ability...." Line 213: "follow-up" Line 213-214: "Poor performance during 6MWT may warrant that the at-risk patients SHOULD be followed more closely..." ERRORS IN FIGURES AND TABLES: Table 1: Hepatocellular carcinoma (with two L) Table 1: electolytes should all be written with or without apex (Na, K, Mg, Ca or Na+, K+, Mg2+, Ca2+) Table 1: I suppose that all continuous variable are shown as mean +/- DS... this must be specified Table 2: This table should be implemented with %6MWD according to Child-Pugh classes and according to ascites, encephalopathy, previous decompensation, hospital admission and survival Table 3: there are layout problems (i don't kown if it is a file download problem). If not, layout must be improved Table 3: Confidential intervals must be added Figure 4: there are lines in portuguese, they should be traduced STATISTICAL AND TEXT ISSUES: 1) In the methods chapter you say that the main outcomes are death and liver transplantation (line 64), but then in results chapter you do not consider patient underwent transplantation in survival analysis. In my opinion "liver transplantation"



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should be deleted among outcomes 2) As you said, the progressively reduction of 6MWD among Child classes in not unexpected. But quantifying the difference of 6MWD among Child classes remain interesting. In the result chapter and in Figure 2 you show the statistical difference between CP-A vs CP-C and CP-A vs CP-B/C, while no statistical difference came out between CP-A vs CP-B and CP-B and CP-C. Are the correlation between the three classes corrected by a multiplicity test? If yes, it should be clarified in the text; if not, it should be made. Another option is presenting the results as differences in 6MWD between compensated (CP-A) and decompensated (CP-B/C) patients. 3) I have some doubts between 6MWD and %6MWD. If you consider the data "6MWD", in all further analysis (first of all univariate and multivariate) you have to correlate this parameter with others (sex, age, weight, height). So all data presented in Table 3, Figure 3a and Figure 4a are not statistically correct. Otherwise, the data "%6MWD" is already adjusted by age/sex/weight/height so is a more correct parameter for all further evaluations. In my opinion you should focus all your attention and discussion on the correlation of %6MWD and outcome 4) Based on ROC analysis, you found a cut-off of 0.82 as the most accurate in predicting mortality. You should produce a table with all the main characteristics (sex, age, BMI, MELD, Child-Pugh, previous decompensation, ascites, encephalopaty, HCC) of the two cohort of patients (%6MWD <0.82 vs >0.82) and the respective p values. If there are differences between the two groups, they should be added in univariate and multivariate analysis.



PEER-REVIEW REPORT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05477048

Position: Editorial Board

Academic degree: PhD

Professional title: Assistant Professor

Reviewer's Country/Territory: Romania

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-03-29 20:13

Reviewer performed review: 2021-04-05 21:03

Review time: 7 Days

Scientific quality	[] Grade A: Excellent [Y] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [] Accept (General priority) [Y] Minor revision [] Major revision [] Rejection
Re-review	[]Yes [Y]No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

The topic is very important and the manuscript highlights a good research activity. Comments: a. the authors specify that "Cirrhosis was defined by clinical history, physical examination, laboratory analysis and at least one radiology data" – would be better imaging data??? b. the exclusion criteria need to be reviewed: it is not clear whether or not patients with previous/current cardiovascular disease were included; why patients with hemochromatosis were excluded? More, "Whether patients who have previously received a liver transplant have been excluded?" should be included in the exclusion criteria.



RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Hepatology

Manuscript NO: 64130

Title: Six-minute walking test performance is associated with survival in cirrhotic

patients.

Reviewer's code: 05311389 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor, Research Fellow

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2021-02-09

Reviewer chosen by: Man Liu

Reviewer accepted review: 2021-05-22 06:30

Reviewer performed review: 2021-05-23 16:07

Review time: 1 Day and 9 Hours

Scientific quality	[] Grade A: Excellent [] Grade B: Very good [Y] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[Y] Grade A: Priority publishing [] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [Y] Accept (General priority) [] Minor revision [] Major revision [] Rejection
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous Conflicts-of-Interest: [] Yes [Y] No
statements	Commercial pres [1] No



Thanks for the answers. I have nothing more to suggest