Peer Review File

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<mark>Reviewer A</mark>

In this manuscript, the Authors present a retrospective study of 23 SCLC (small cell lung cancer) patients with SIADH (syndrome of inappropriate antidiuretic hormone hypersecretion) focusing on the effect of hyponatremia correction on prognosis and survival.

The topic posed by the Authors can be considered of interest since the paucity of data available in the Literature because SIADH is an uncommon and its pathogenesis remains unknown. Furthermore, growing evidences seem to suggest that SIADH might play a concrete role on patient's prognosis.

Nevertheless, the paper shows several flaws that ultimately do not make it suitable for publication in the present form. Among the others, please consider the following points of discussion:

• English language should be reviewed (some grammatical errors)

Reply 1: Thanks to the reviewer's suggestions, the reviewer's suggestions will help improve the quality of this article. We invited Alvin Han, undergraduate student, Department of Biochemistry and Biomedical Sciences, McMaster University to help us review the grammatical problem, and we corrected the wrong expression.

Change in the text: (see Page 4, line 22)(see Page 5, line 2-12)(see Page 6, line 1-4, line 6-7)(see Page8, line 3-4)(see Page 9, line 2, 7, 12)(see Page 10, line 22)(see Page 11, line 12-13, 17) (see Page12, line 19-20) (see Page13, line 4, 19) (see Page14, line22) (see Page 18, line 2) (see Page 19, line 6)

Change in the Table 2 : we have modified our text as "after 3 days of tolvaptan"

• The Authors should spell check the text (several orthographic mistakes)

Reply 2: Thank you very much for the suggestions of the reviewers, which helped us avoid mistakes, the language correctness was reviewed.

Change in the Table 1: we have modified the word "total, develop", Change in the Fig 1B, we have modified the word "treatment".

• In the Introduction section the Authors should explain the difference between SIADH and the modern acronym SIAD

Reply 3: Thank you very much for the suggestions of the reviewers, the suggestions of the reviewers have enriched our content, we explained the difference between SIAD and SIADH based on the reviewer's suggestions.

Change in the text: (see Page 4, line 9-13)



In their paper the Authors present study on the role of Tolvaptan in the treatment of cancer related SIADH. The retrospective and monocentric study enrolled 23 patients with small cell lung cancer affected by SIAD and treated with Tolvaptan. They showed that a lower dosage could be equally efficacy than dosage indicated by the data sheet, with lower risk of side effects.

The question posed by the Authors can be considered of interest since hyponatremia and SIADH represent important negative predictive and prognostic parameter in several tumors and a prompt correction of this electrolyte disorder might improve patients' outcome. However, there is lack of data in the literature on the real management of SIAD treatment with Tolvaptan and this paper might contribute in improve knowledge about this setting. The article in question is properly structured; the rationale and the hypothesis are clearly expressed. Data and results, including figures, are well explained and properly interpreted in the discussion. After careful evaluation, I would suggest that the manuscript may be suitable for publication after minor revisions.

Reply 1: Thank you for the encouraging comments, we have made detailed changes to the article.

<mark>Reviewer C</mark>

Major comments:

This paper retrospectively analyzed tolvaptan treatment for SIADH in SCLC. It mainly shows that the effect of tolvaptan is not dose-dependent. Well written overall.

Minor comments:

TLV is abbreviation for tolvaptan, but it is not used in the text, so "(TLV)" in Abstract, Keywords, and Introduction are unnecessary.

Reply 1: We would like to thank you for the constructive comments, we fully accepted your suggestion, we have deleted the "TLV" in the text.

Change in the text: (see Page 2, line 4) (see Page 3, line 16) (see Page 5, line 11) In Methods, 23 patients were repeated 3 times.

Reply 2: Thanks for the reviewer's suggestion, the reviewer's suggestions make the article's expression more reasonable, we have deleted "23 patients" in unnecessary sentences.

Change in the text: (see Page 7, line 9)

In Demographic characteristics "body mass index (BMI)" is "BMI"

Reply 3: Thanks for the reviewer's suggestion, we have modified our text as "BMI". Change in the text: (see Page 9, line 5)