

Dear Editors and Reviewers:

Thank you for your letter and for the reviewers' comments concerning our manuscript entitled "Electroconvulsive therapy plays an irreplaceable role in the treatment of major depressive disorder" (Manuscript NO: 75999). Those comments are all valuable and very helpful for revising and improving our paper, as well as the important guiding significance to our researches. We have studied comments carefully and have made correction which we hope meet with approval.

Reviewer #1: This paper is a careful letter to the editor, which further expands on the role of electro-convulsive therapy and the mechanisms which underpin it on neuro-biological level. The authors may Also consider to comment on the whole-brain voxel-based estimation of the quantitative changes in the brain volume over the time course

followingECT:<https://www.sciencedirect.com/science/article/pii/S1053811921001725>

Response: Thank you very much for your reference. I will review the paper you recommend at my leisure.

Reviewer #2: In this letter to the editor the authors have discussed the physiological and neurobiological role/impact of ECT in various psychiatric disorders including major depressive disorder. They are highlighting that ECT should not be considered as the second line of management or last resort of treatment for major depressive disorder. Apart from several grammatical errors (one of the big ones being use of present and past tense in consecutive sentences) the letter covers the salient point of not overlooking the relevance of ECT in management of major depressive disorder among other psychiatric disorders.

Response: Thank you very much for your valuable advice. We have carefully revised the grammar and sentences of the whole paper.

We tried our best to improve the manuscript and made some changes in the manuscript. These changes will not influence the content and framework of the paper. And here we did not list the changes but marked in red in revised paper.

We appreciate for Editors/Reviewers' warm work earnestly, and hope that the correction will meet with approval.

Once again, thank you very much for your comments and suggestions.