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OPEN PEER REVIEW

Effects of Adding Tizanidine on Stimulants in Treatment of ADHD: A Randomized, Double-Blind, Placebo-Controlled Trial

Afsaneh Karbasi Amel, Mostafa Haghshenas, Leila Sahraian

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1 Original submission

2 comments · 15 Dec 2024 – 15 Dec 2024

Associate Editor

MAJOR REVISION 15 Dec 2024

Dear EIC,
Dear EIC,
Thanks to the authors and appreciation to the researchers. The manuscript is a valuable piece of research. But some revisions are needed. The authors should revise it based on the reviewers' comments. Also, the abstract should be summarized, especially in the Background section.

Editor-in-Chief

MAJOR REVISION 15 Dec 2024

Dear Author,
Thank you for submitting the article. It needs some revision. Please revise it based on the reviewers' and AE's comments.

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Revision 1

AUTHORS' RESPONSE TO REVIEWERS

Authors' reply to reviewers

The next pages contain the authors' point-by-point reply to the comments of this revision round, as submitted with the revised manuscript. It is published as the authors wrote it.

Dear EIC.

First First, I would like to appreciate you for spending your valuable time in reviewing our manuscript.

Based on you and your colleagues' valuable comments, we have edited the whole manuscript. Still, we are fully ready for any potential further amendments or revisions.

Reviewer 2:

1. *Limited Patient Diversity:*
   The study exclusively focuses on patients unresponsive to Lisdexamfetamine. Including patients who use other common medications, such as Ritalin (Methylphenidate), would enhance the generalizability of the findings

Answer: because comply the homogeneity of participants(due to small sample size) we just select Vyas group

2. *Small Sample Size:*
   The sample size of 40 patients is relatively small and limits the statistical power of the study. Larger-scale trials are recommended to detect more subtle effects and validate the results.

Answer: Sample size was according the previous similar studies but as you noted should be increased enough in future studies for detect more subtle effects.

3. *Short Study Duration:*
   The 8-week study period might be too brief to observe the long-term effects of Tizanidine. Extending the follow-up to 12 or 16 weeks could provide more comprehensive data.

Answer : due to previous similar studies the duration of study was 8 week and this was a limitation for this study.

4. *Low Dosage of Tizanidine:*
   The maximum dose of 4 mg/day might not be sufficient to observe significant therapeutic effects. Future studies could explore the impact of higher doses with appropriate safety precautions

Answer :because this was the first time that Tizanidine was used in pediatric age group, the dose of tizanidin was in lower limit. In future studies the dose could be rise.

5. *Focus on Limited Outcomes:*
   The study primarily examines ADHD symptoms without considering broader

outcomes such as quality of life or psychological well-being. Including these measures in future research would provide a more holistic view of the drug's impact.

Answer: whereas considering quality of life and psychological wellbeing is optimal in these type of studies,but time limitation (8week for quality of life) and small sample size (for assessing psychological well beeng) are abstacle.

6. *Underexplored Positive Results:*
   The finding that Tizanidine reduces conduct problems is significant but underanalyzed. Further exploration of this effect and its clinical implications is highly encouraged

Answer: it was corrected

2 Revision 1

2 comments · 12 Jan 2025 – 12 Jan 2025

Associate Editor

ACCEPT 12 Jan 2025

Dear EIC,
I read the manuscript, and the authors corrected the requested items. The article could be accepted at this stage.
Sincerely;

Editor-in-Chief

ACCEPT 12 Jan 2025

Dear Author,
I read the manuscript, and the authors corrected the requested items. The article could be accepted at this stage.
Sincerely;