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PEER-REVIEW REPORT

Name of journal: World Journal of Psychiatry

Manuscript NO: 64964

Title: G-protein coupled receptors and synaptic plasticity in sleep deprivation.

Reviewer's code: 05945693 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Canada

Author's Country/Territory: Canada

Manuscript submission date: 2021-02-25

Reviewer chosen by: Jin-Lei Wang

Reviewer accepted review: 2021-03-29 13:59

Reviewer performed review: 2021-04-06 18:55

Review time: 8 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Scientific quality	[] Grade A: Excellent [] Grade B: Very good [Y] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish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) [] 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

Your review article entitled "G-protein coupled receptors and synaptic plasticity in sleep deprivation" by Parmar et al. attempted to summarize the recent findings on GPCRs and their role in the formation of long-term memories which occurs while you sleep. In the event you suffer from inadequate sleep or sleep deprivation, these memory consolidation processes (synaptic plasticity via LTP and LTD) are affected which can explain memory loss and other psychoneurological processes that are associated with sleep deprivation. Essentially the authors concluded that this is an issue for drugs used to treat neurological conditions since they act on or are GPCRs and that this information is essential when it comes to drug development. In general, I found the article to be well written, organized in a logical way and fairly easy to read. I did identify a few spelling and syntax errors and well as some inconsistencies with the spelling of some terms. I also identified a few instances where more detail could be added to improve the review article, particularly in the conclusion and with regards to the figures. sleep is so fundamental to this review, I feel the last paragraph on page 5 of section I. a. Sleep and Its significance could be expanded. You describe what normally occurs in the brain when one sleeps however with regards to sleep deprivation you don't explain what changes or differs with regards to the REM and SWS. What are some typical and atypical scenarios where people would suffer from sleep deprivation? Do people who work nights or shift work suffer from more memory impairment than those who work during the day? 2. In the last sentence on Page 6/first sentence of Page 7 you define LTP (high frequency stimulation = high Ca2+ influx) but you don't define LTD (low frequency stimulation = low Ca2+ influx). As I continue to read the article you frequently talk about "LTP and LTD" and so describing the processes for each of these synaptic mechanisms would help the reader understand both the difference and importance of each of these processes and what happens (i.e. Ca2+ increases, AMPA



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receptors, etc). I strongly suggest a table or figure be created to represent this. 3. In the last paragraph on page 10 you mention a hypothesis by Lynch. I felt you did not summarize the findings of the article adequately. Something is missing. What caused the increase in the number of glutamate receptors and what did it have to do with calcium proteinases? 4.Midway through page 11 you talk about now ionotropic and metabotropic receptors are included in synaptic plasticity and have different mechanisms of action. It would be great to give the reader an example of each type of receptor (in brackets) and the different mechanism of action it has with regards to synaptic plasticity like the ones you talk about later in the article. 5. Within the last few lines on page 14, you say "Considering that NO has a role in so many other important physiological functions, " I feel you should give an example of an important physiological function. 6. On page 15, section 4 and specifically In line number 8 you write "high in the hippocampus (300) and about 20..." what does the (300) refer to? Is it a reference? Or are you referring to how many transcripts? Please clarify 7. see several instances where you write "post synaptic", "post-synaptic" and "postsynaptic". Please make it consistent throughout the manuscript. The same for "pre-synaptic". 8.

Regarding the last 2 sentences in section "a." on page 21, you summarize the results from reference 131. However, something is missing. Please review the 2 sentences and possibly add more detail about the study. How were the rats sleep deprived? What part of the brain did the authors study? Which method was used to measure the expression levels of the receptors? 9. In both Figure 1 and in the concluding paragraph on page 24-25 you talk about psychiatric therapeutics, however you don't list any examples. I strongly suggest you highlight (and give examples) of a few drugs that are used to treat the disorders you talk about in your manuscript particularly those that are or act upon GPCRs. Also what is the mechanism of action of these drugs? Do they increase the availability of neurotransmitters which in turn will cause an increase the transcription of



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their respective receptors? Are they GPCRs themselves? 10. I believe a much stronger link needs to be made with regards to sleep deprivation and psychiatric disorders. I can see where you are trying to go with regards to the conclusion of this review, that being that changes are occurring to GPCRs as a result of sleep deprivation and it is important to study this because it can affect the formation of memories as well as the ability of drugs that target these receptors to perform their action when it comes to treating neuropsychiatric disorders. One angle that you could take to make the link stronger is maybe highlight how one of the side effects of taking drugs that treat neuropsychiatric disorders is (likely) problems with sleeping (which you kind of touch on in the future direction paragraph). I think it would also be a good idea to briefly describe drugs that treat sleeping problems as well unless wordcount/space is an issue.



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RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Psychiatry

Manuscript NO: 64964

Title: G-protein coupled receptors and synaptic plasticity in sleep deprivation.

Reviewer's code: 05945693 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Canada

Author's Country/Territory: Canada

Manuscript submission date: 2021-02-25

Reviewer chosen by: Man Liu

Reviewer accepted review: 2021-07-28 14:37

Reviewer performed review: 2021-08-02 15:08

Review time: 5 Days

Scientific quality	[] Grade A: Excellent [Y] Grade B: Very good [] 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	[Y] Accept (High priority) [] Accept (General priority) [] Minor revision [] Major revision [] Rejection
Peer-reviewer statements	Peer-Review: [Y] Anonymous [] Onymous Conflicts-of-Interest: [] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

The revised version of the manuscript is adequate as well as the responses to each



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suggested revision.