

Review of: "Methadone In Chronic Non-Oncological Pain: From Disassuefaction Of Painkillers Abuse To The Primary Management Of Opioid Hyperalgesia"

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Potential competing interests: The author(s) declared that no potential competing interests exist.

Foremost, I would like to thank for this opportunity to revise the manuscript titled

"Methadone In Chronic Non-Oncological Pain: From Disassuefaction Of Painkillers Abuse To The Primary Management Of Opioid Hyperalgesia" that was submitted to Qeios.

The article topic presents an interesting concern regarding opioid addiction and therapeutic use of methadone. However, there are several concerns with the manuscript which are as follows-

- 1. Firstly, the language, grammar and writing is not scholarly and fit for an academic literature.
- 2. Many sentences are too lengthy such that the message being delivered gets confusing.
- 3. Many of the factual data are not backed by any references (e.g in Background and Discussion section).
- 4. The material and methods section is not satisfactory- Design, Search timeline, Inclusion, Exclusion criteria, Article flow?
- 5. The Image 3- for molecular mechanisms of OIH appears in other primary article by Roeckel et al.
 - L.A. Roeckel, G.M. LeCoz, C. Gaveriaux-Ruff, F. Simonin.Opioid-induced hyperalgesia: cellular and molecular mechanisms. Neuroscience, 338 (2016), pp. 160-182.

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- 1. Which study population is targeted as the article gives US data and then relates to Italy in between the article?
- 2. The final conclusion adds nothing to the already available literature.

FINAL RECOMMENDATION- REJECT