

Review of: "Targeting Alzheimer's disease hallmarks with the Nrf2 activator Isoeugenol"

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Potential competing interests: No potential competing interests to declare.

Dear Editor,

On manuscript entitled "Targeting Alzheimer's disease hallmarks with the Nrf2 activator Isoeugenol"

Comments:

- 1- Text should be edited. There are various typo and grammatical errors.
- 2- Abstract need to rewrite, methods and results should be added in brief and clear way.
- 3- introduction is too long. You should concise the introduction. in the current form, AD explained more. I suggest that introduction write in 4 separate paragraph (1-AD and its complications, pathophysiology and ...2- Role of nrf2, neuroinflammation and oxidative stress and ... in the pathophysiology of AD, 3- Isoeugenol and its pharmacological and pharmacokinetic effects and 4- aim of study)
- 4-Results should be deleted from end of introduction section.
- 5-all methods need appropriate references.
- 6-How many mice were used?
- 7- What is your rational for dose and time of agent administrations?
- 8- Why number of mice in groups is different?
- 9- A schematic for study design in mice is needed to help better understanding.
- 10- methods used for sacrificing are different for Pharmacokinetic and pharmacodynamic studies. Please clarify. Doses of ketamine and xylazine need references
- 11- gene expressions were measured in which samples? please clarify.
- 12-rational for performing hot plate test should be clarified.
- 13- did each mouse subjected to one trial? if not, please clarify timeline with



intervals between tests.

14- If possible please concise methods. primers include in table.

15-please show histopathological changes in figure.

16-Discussion is too long with unnecessary topic in some points. strongly suggested that brief this section. limitation of study should be added.

Qeios ID: JM0LG2 · https://doi.org/10.32388/JM0LG2