

PEER-REVIEW REPORT

Name of journal: *World Journal of Neurology*

Manuscript NO: 79884

Title: Possible convulsion and electroencephalographic abnormality in a patient taking long-term oral clarithromycin

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03199608

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Chief Physician, Director, Professor

Reviewer's Country/Territory: China

Author's Country/Territory: Japan

Manuscript submission date: 2022-09-10

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-09-13 20:42

Reviewer performed review: 2022-09-19 06:12

Review time: 5 Days and 9 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input checked="" type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

**Peer-reviewer
statements**

Peer-Review: ☒ Anonymous ☐ Onymous

Conflicts-of-Interest: ☐ Yes ☒ No

SPECIFIC COMMENTS TO AUTHORS

Clarithromycin increases neuronal activity, stimulate CA3 pyramidal neurons of the hippocampus by reducing GABAergic signaling, and cause neuronal hyperexcitability. The patient had been prescribed clarithromycin for chronic sinusitis for >10 years. Liver CYP3A2 metabolize clarithromycin. Diltiazem could inhibit CYP3A. In the present case, diltiazem was added as an antihypertensive just before the seizure. Thus, the introduction of diltiazem may have increased the blood concentration of clarithromycin. Clarithromycin also causes QT interval prolongation. Our case showed a QTc time of 448 milliseconds at admission, and after discontinuation of clarithromycin, his QTc time improved to 385 milliseconds. Measurement of QT time may be a valuable method of assessing clarithromycin excess, since QT interval prolongation might induce seizures. One question, CYP3A2 is expressed in animal or human? Or they are isoenzyme with CYP3A4 in human? QT interval prolongation might be a keyword in the paper.

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Reviewer's code: 06185324

Position: Peer Reviewer

Academic degree: MBBS

Professional title: Reader (Associate Professor)

Reviewer's Country/Territory: Nigeria

Author's Country/Territory: Japan

Manuscript submission date: 2022-09-10

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-10-12 09:26

Reviewer performed review: 2022-10-22 16:59

Review time: 10 Days and 7 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
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Peer-reviewer statements	Peer-Review: [<input checked="" type="checkbox"/>] Anonymous [<input type="checkbox"/>] Onymous Conflicts-of-Interest: [<input type="checkbox"/>] Yes [<input checked="" type="checkbox"/>] No
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SPECIFIC COMMENTS TO AUTHORS

I congratulate the authors for this interesting and well written manuscript. Kindly note the following. 1. Make the introduction more robust 2. Introduction should have references 3. In the discussion, quote case reports or studies by other researchers on same subject. 4. Adopt the changes, if acceptable, as suggested the review.