

PEER-REVIEW REPORT

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Title: Amiodarone-induced hepatotoxicity: quantitative measurement of iodine density in the liver using dual-energy CT: a report of three cases and literature review

Reviewer's code: 02451459

Position: Peer Reviewer

Academic degree: PhD

Professional title: Associate Professor

Reviewer's Country/Territory: Singapore

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Reviewer chosen by: Xiao-Quan Yu

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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 Grade E: Do not publish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
Conclusion	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No
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

SPECIFIC COMMENTS TO AUTHORS

This is an original case report and research about the use of dual-energy CT to measure liver iodine concentration. The goal is to associate this iodine level to the potential for hepatotoxicity due to amiodarone exposure. Using 3 separate case studies, evaluation of both liver density and iodine concentration revealed the significant exposure of the liver to amiodarone. However associating this exposure to the onset of hepatotoxicity will require larger sample size as well as control samples of patients on amiodarone without hepatotoxicity. Nonetheless, this report sets the stage for such future development. Specific questions are as follow: 1. Amiodarone-induced liver toxicity often comes in the form of steatohepatitis. In this case, accumulation of fats, cholestatic injury markers (e.g. GGT and bilirubin) would be more suitable indicators of injury as compared to ALT and AST. Such markers were not consistently reported in all the 3 cases included here. Will be good to also include that information. 2. What is the reference range for iodine concentration in the liver? Do we know if the 1.4-2.9 mg/mL range detected is considered higher than normal? Also, is there is a difference between patients taking amiodarone with or without hepatotoxicity? This information will help in the conclusion for this report.