

PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Pharmacology and Therapeutics

Manuscript NO: 60773

Title: Overview of Drug Induced Liver Injury in Brazil: What is the role of public health policy on the evidence?

Reviewer's code: 04054541

Position: Peer Reviewer

Academic degree: PhD

Professional title: Postdoc, Research Fellow

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2020-11-25

Reviewer chosen by: Le Zhang

Reviewer accepted review: 2020-12-28 14:44

Reviewer performed review: 2020-12-30 13:14

Review time: 1 Day and 22 Hours

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|---------------------------------|---|
| Scientific quality | <input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" 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 Grade E: Do not publish |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 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: <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

Here the authors provide an overview of the current situation in Brazil of drug-induced liver injury (DILI). After a careful analysis of literature using PubMed, Scielo, Science Direct, and the Brazilian thesis bank, they considered different variables as well as: age, gender, comorbidities, suspected drugs, immunodeficiency virus (HIV), tuberculosis (TB) or hepatitis C (HCV) infection, use of an algorithm to support diagnosis, classification criteria for hepatotoxicity, outcomes related to DILI, and so on. The authors reported that the drugs more implicated in DILI in Brazil are Isoniazid, Rifampicin, and Pyrazinamide, Nevirapine, in addition to methotrexate, propylthiouracil, azathioprine. However, the diversity of methods and criteria for the definition of hepatotoxicity did not allow obtaining frequency estimates. So, as the authors suggest, this review could be useful to make standardized criteria for the identification of DILI, leading to the creation of a wide national DILI monitoring network. In my opinion, the paper is generally well written and structured. The introduction is clear and also the other sections are adequately described. I suggest only a minor revision and the correction of some oversights and misspellings in the text that I have listed below. - In the Abstract, in line 27, there is a double space within the word “non-prescription”, please remove it. - In the Abstract, in line 36, the verb “lead” sounds incorrect, please replace it with “led”. - The same error occurs in line 55, please replace “lead” with “led”. - The keywords are left-aligned, please justify them. - Sometimes the reference number appears after space, while other times it is attached to the word. Please standardize the references. - In line 118, a full stop is missing at the end of the sentence, please add it. - From lines 122 to 125 and from lines 148 to 150, please always use the comma as a punctuation in the list, removing the semicolon. - In line 139, the word “cross-sectional” shows two different fonts, please correct it. - Please revise the English language.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Pharmacology and Therapeutics

Manuscript NO: 60773

Title: Overview of Drug Induced Liver Injury in Brazil: What is the role of public health policy on the evidence?

Reviewer's code: 04737076

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Consultant Physician-Scientist, Doctor, Lecturer, Pediatric Gastroenterology Fellow, Professor, Senior Postdoctoral Fellow, Senior Researcher, Senior Scientist

Reviewer's Country/Territory: Russia

Author's Country/Territory: Brazil

Manuscript submission date: 2020-11-25

Reviewer chosen by: Le Zhang

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Reviewer performed review: 2021-01-02 09:13

Review time: 4 Days and 11 Hours

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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 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 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 |

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| Peer-reviewer statements | Peer-Review: [<input type="checkbox"/>] Anonymous [<input checked="" type="checkbox"/>] Onymous Conflicts-of-Interest: [<input type="checkbox"/>] Yes [<input checked="" type="checkbox"/>] No |
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SPECIFIC COMMENTS TO AUTHORS

The title of the manuscript corresponds to the content. The abstract reflects the content of the manuscript. Key words reflect the essence of the manuscript. Research methods are described in full. A review is made of a large number of publications devoted to medicinal liver damage. It has been revealed that a significant part of medicinal liver damage, including severe liver damage, is caused by drugs such as Isoniazid, Rifampicin and Pyrazinamide, Nevirapine, as well as methotrexate, propylthiouracil, azathioprine. The assessment of the criteria for hepatotoxicity of drugs was carried out. Twenty-two different strategies in the definition of liver injury were found, and also made it difficult to compare the findings. It is shown that the Roussel Uclaf Causality Assessment Method (RUCAM) algorithm for the diagnosis of medicinal liver lesions is rarely used. Shown in the Brazilian population, the frequency of clinical trials with data on hepatotoxicity of drugs is low. A cohort study and pharmacovigilance activity has been found to be the ideal plan for identifying drug side effects. It has been shown that the frequency of liver damage associated with certain drugs can vary depending on the region, on the characteristics of the studied populations and the profile of drug use. An important factor in potential drug damage to the liver is the use of large amounts of drugs. Attention is focused on such widely used drugs, including in pediatric practice, as Amoxicillin with clavulanate and other antimicrobials, non-steroidal anti-inflammatory drugs, as well as antineoplastic drugs.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Pharmacology and Therapeutics

Manuscript NO: 60773

Title: Overview of Drug Induced Liver Injury in Brazil: What is the role of public health policy on the evidence?

Reviewer's code: 01213276

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Professor

Reviewer's Country/Territory: Serbia

Author's Country/Territory: Brazil

Manuscript submission date: 2020-11-25

Reviewer chosen by: Le Zhang

Reviewer accepted review: 2020-12-30 04:34

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Review time: 3 Days and 11 Hours

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| Scientific quality | <input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input checked="" 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 type="checkbox"/> Minor revision <input checked="" type="checkbox"/> Major revision <input type="checkbox"/> Rejection |
| Re-review | <input checked="" type="checkbox"/> Yes <input 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

In this article authors analyzed Brazilian studies reporting the drug-induced liver injury. „Hepatotoxicity can be induced both by n on-prescription drugs and by those used for chronic diseases,, That sentence haven't sense - prescription or non-prescription, acute or chronic disease, not non-prescription or chronic. DILI's-we haven't the full term before abbreviation. Finally, the main criticism is too extensive discussion. I have a impression that authors wonted to analyze all tiny aspects of the problem. Some parts like „Clinical trials are not the best study design to assess drug safety...,, and „great cultural diversity make it difficult to carry out a single representative study in the country,, and „For this reason, it is important to encourage further regional studies,, are well knowing and didn't improve quality of the paper. By my opinion, first four or five passage are redundant. „Are researcher's unfamiliarity regarding algorithms?... to .. which would make possible to classify the liver injury and improve the knowledge of the outcomes...,, is really nor necessary. However this is a good try, authors made a grate effort and with some abbreviations and modifications this article can by a suitable for publishing.

RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Gastrointestinal Pharmacology and Therapeutics

Manuscript NO: 60773

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Reviewer's code: 04054541

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Academic degree: PhD

Professional title: Postdoc, Research Fellow

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2020-11-25

Reviewer chosen by: Le Zhang

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|---------------------------------|---|
| Scientific quality | <input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" 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 |
| 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



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The authors fulfilled all my requests but key words are still left aligned, please justify them. Moreover in line 150 there is a semicolon, please use the comma. I have no other requests.