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PEER-REVIEW REPORT

Name of journal: *World Journal of Clinical Oncology*

Manuscript NO: 86498

Title: Advances in drug resistance of triple negative breast cancer caused by pregnane X receptor

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 00069774
Position: Editorial Board
Academic degree: PhD
Professional title: Associate Professor
Reviewer's Country/Territory: Thailand
Author's Country/Territory: China

Manuscript submission date: 2023-06-21

Reviewer chosen by: Geng-Long Liu

Reviewer accepted review: 2023-07-21 04:22

Reviewer performed review: 2023-07-30 04:31

Review time: 9 Days

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Novelty of this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No novelty
Creativity or innovation of this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No creativity or innovation

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Scientific significance of the conclusion in this manuscript	[ ] Grade A: Excellent [ ] Grade B: Good [Y] Grade C: Fair [ ] Grade D: No scientific significance
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [ ] Minor revision [Y] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No
Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

Manuscript describes the classical role of PXR in regulation of drug metabolizing enzymes phase 1, 2, and phase 3 drug transporters and the association with the metabolism and transport of chemotherapeutic drugs. The major drug metabolizing enzymes and transporters with sufficient evidences linked to chemotherapeutic resistance should be tabulated for a summary to shorten the narrative. Many important roles of PXR were not included in the manuscript as it highly relates to the manuscript title. This includes regulation of PXR and function of PXR independence of drug metabolism. Regulation of PXR is very important in that several chemicals and inflammatory mediators which likely occur in many cancers can regulate expression of PXR and PXR-regulated genes. PXR also acts independence of drug metabolism in regulation of various processes for instances, apoptosis, cell cycle progression and inflammation via p53, p21, NF-kB, STAT3. Moreover, many above effects can be tissue specific in that PXR acts differently in various tissues. These two aspects should be included in the manuscript.

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Reviewer's code: 00505473
Position: Peer Reviewer
Academic degree: PhD
Professional title: Head
Reviewer's Country/Territory: India
Author's Country/Territory: China

Manuscript submission date: 2023-06-21
Reviewer chosen by: Geng-Long Liu
Reviewer accepted review: 2023-07-06 13:54
Reviewer performed review: 2023-08-02 20:23
Review time: 27 Days and 6 Hours

Scientific quality	[ ] Grade A: Excellent [ ] Grade B: Very good [Y] Grade C: Good [ ] Grade D: Fair [ ] Grade E: Do not publish
Language quality	[ ] Grade A: Priority publishing [Y] Grade B: Minor language polishing [ ] Grade C: A great deal of language polishing [ ] Grade D: Rejection
Conclusion	[ ] Accept (High priority) [ ] Accept (General priority) [Y] Minor revision [ ] Major revision [ ] Rejection
Re-review	[ ] Yes [Y] No

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Peer-reviewer statements	Peer-Review: [Y] Anonymous [ ] Onymous
	Conflicts-of-Interest: [ ] Yes [Y] No

SPECIFIC COMMENTS TO AUTHORS

In this review, the therapeutic importance of pregnane X receptor (PXR) in TNBC was discussed. Following comments can be made: 1.The authors have discussed the three phases of the chemotherapy drug resistance mechanism caused by PXR in general (Figure-1). However, authors have superficially discussed the alterations of PXR regulated genes associated with the three phases of drug metabolism in TNBC. The discussion should be made in detail with diagrammatic representation for better understanding of the role of PXR regulating genes in drug resistance in TNBC. This should be in concordance of title of the review. 2.The conclusion of the review should be made in accordance of the discussion above.