

Review Form 1.6

Journal Name:	Journal of Pharmaceutical Research International
Manuscript Number:	Ms_JPRI_78832
Title of the Manuscript:	ANTI-PROLIFERATIVE EFFECT OF HYDROETHANOLIC LEAF EXTRACT OF CITRULLUS COLOCYNTHIS (L) ON RETINOBLASTOMA CELL LINE
Type of the Article	Original Research Article

General guideline for Peer Review process:

This journal's peer review policy states that **NO** manuscript should be rejected only on the basis of '**lack of Novelty**', provided the manuscript is scientifically robust and technically sound. To know the complete guideline for Peer Review process, reviewers are requested to visit this link:

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PART 1: Review Comments

	Reviewer's comment	Author's comment (if agreed with reviewer, correct the manuscript and highlight that part in the manuscript. It is mandatory that authors should write his/her feedback here)
Compulsory REVISION comments	<p>Dear Editor, I found that manuscript is suitable for publication and the reviewer recommended it publication after major revision.</p> <p>The authors contributed the work entitled "ANTI-PROLIFERATIVE EFFECT OF HYDROETHANOLIC LEAF EXTRACT OF CITRULLUS COLOCYNTHIS (L) ON RETINOBLASTOMA CELL LINE". This review covers the knowledge in the field and expands the current understanding of the field.</p> <p>Some of the traditional uses have been validated by phytochemical and modern pharmacological studies but still many needs to be validated. Although increasing interest has prompted more studies on the phytochemistry and pharmacology of Citrullus colocynthis, there are still many areas where the current knowledge could be improved: 1) the detailed quantitative data about the alkaloids and other secondary metabolites are still needed, 2) the Citrullus colocynthis fruit is a rich source of cucurbitacins, many with same skeletal structure. Therefore, it would be interesting to investigate the structure-activity relationships of these compounds and 3) several traditional uses of the Citrullus colocynthis fruit have been validated in the recent pharmacological studies; however, some of these studies were only tested in vitro. Thus the effectiveness of Citrullus colocynthis fruit extracts and isolated compounds needs to be further investigated for their efficacy and safety using in vivo assays. The plant is known to be beneficial in multiple gut disorders such as, indigestion, gastroenteritis, constipation and colic pain, while there is no study available on any gut disorders. The plant has been shown to possess anti-inflammatory activity, but, unlike NSAIDs, it may not cause gastric ulcer, because, it is traditionally used in gastroenteritis (yet to be scientifically validated). Thus, if proved to possess anti-ulcerogenic, this may offer a safer option for inflammatory conditions. The plant has not been studied for its effectiveness in cardiovascular disorders except a couple of preliminary reports showing antilipidemic effect. Thus, this plant with therapeutic potential in metabolic and cardiovascular disorders warrants immediate attention for preclinical and clinical studies. According to what has been mentioned in this paragraph:</p> <ol style="list-style-type: none"> 1. The introduction as well as the discussion should be improved and updated. 2. In addition, I suggest updated the manuscript with this relevant article. https://doi.org/10.1007/s12094-020-02428-1 3. Please discuss about the potential systemic toxicity and safety evaluation of <i>Citrullus colocynthis</i>. 4. Please include the limitations of the study in the discussion. 5. Generally, the writing should be polished to improve the readability. 	
Minor REVISION comments		
Optional/General comments		

PART 2:

	Reviewer's comment	Author's comment (if agreed with reviewer, correct the manuscript and highlight that part in the manuscript. It is mandatory that authors should write his/her feedback here)
Are there ethical issues in this manuscript?	<i>(If yes, Kindly please write down the ethical issues here in details)</i>	

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