

Response to the Reviewer

We thank the editors for their constructive comments. We have made detailed modifications according to the editor's comments. We believe that these changes have improved the paper and we appreciate the efforts of the editors in this behalf. Specific point-by-point responses are below.

List of issues that need to be addressed by authors in a conditionally-accepted manuscript

Issues raised:

- (1) please give brief reasoning and explanation about determining XHP dose for mice. he title is too long, and it should be no more than 18 words;

According to the pharmacological dosage regulations of XHP, the daily dose of XHP for adults is 6 g/d, and the body weight of each mice is about 20g. The dose of XHP was determined by dose extrapolation based on dose-body surface area normalization, this dosage was converted to 78 mg/kg[17].

Reference: Nair A, Morsy MA, Jacob S. Dose translation between laboratory animals and human in preclinical and clinical phases of drug development. *Drug Dev Res* 2018; **79**: 373-382 [PMID: 30343496 DOI: 10.1002/ddr.21461]

- (2) Discussion: since XHP consists of many compounds, please give reasoning about this in relation with your study and results. Please point out the limitations of your study.

We added the following:

In this study, the high resolution mass spectrometry was used to analyze the compound composition of xihuang pill. Twelve compounds in xihuang pill were identified, which were valine, 4-gingerol, myrrhone, ricinoleic acid, glycocholic acid, curzerenone, 11-keto- β -boswellic acid, oleic acid, germacrone, 3-acetyl-9,11-dehydro- β -boswellic acid, 5 β -androstane-3,17-dione, and 3-acetyl-11-keto- β -boswellic acid. Among them, Curcuzederone, 11-keto- α -Boswellic Acid, Oleic Acid and 3-acetyl-11-keto-beta-Boswellic Acid have been reported to have good anticancer activity. They inhibit tumor growth by promoting apoptosis of cancer cells[34-39]. We speculate that the anti-tumor effect of XHP may be through these active ingredients, and we will further clarify the role of the active ingredients in future studies.

(3) In figure 5 there are 8 mice, while your sample is 10 mice, please add the missing 2 mice for better viewing.

We have supplemented and modified the picture

(4) There is figure 6 at the end of your manuscript. Where is Fig. 6 located in the text?

Figure 6 is a schematic of the experimental mechanism, which we have deleted.

We have revised the language of the manuscript and submitted the Non-Native Speakers of English Editing Certificate. Finally, we have further revised the manuscript according to the guidelines for revising the content.