Author’s response to reviews

Title: The synergistic antitumor effect of Huaier combined with 5-flourouracil in human cholangiocarcinoma cells

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We really appreciate the hard work of the editors and constructive comments of the reviewers. We have carefully examined our manuscripts and made the following replies.

The answer to the question of reviewer 1

Question 1
The reviewer could download the image directly from the data, where the resolution is higher.

Question 2
We are sorry that we have no conditions for fingerprint of Huaier extract, whose ingredients have been confirmed to be polysaccharides.
Question 3
We have tried our best to correct the linguistic and style errors in our manuscript, please check them.

Question 4
Although Huaier could reduce STAT-3 activation, but in reality it couldn’t completely suppressed the effect of proliferation, cell cycle, apoptosis as well as cellular migration and invasion caused by IL-6. And because of space limitation, we don’t show the results.

Question 5
e we are sorry that we only have one kind of cholangiocarcinoma cell line.

The answer to the question of reviewer 2

It is well known that Huaier has been widely used in the treatment of hepatocellular carcinoma in clinical practice around China. What kinds of agents we use to perform the experiment were aqueous which provided by the pharmaceutical factory of Huaier, it is customary method. Main phytochemicals are polysaccharides, please refer to my citation. We have used Chou-Talalay methods to demonstrated the relationship between 5-Fu and Huaier is synergistic at a certain dose in result of our manuscript. we really wrote that “Huaier has been reported exhibiting a wide range of anti-cancer functions, including……” in introduction, and in reality Huaier has been used in the treatment of liver and breast cancer in clinical practice. The doses of huaier were determined by Chou-Talalay methods, which is commonly used to estimate the effect and dose of the combination of two drugs. Finally, we are sorry that Huh28 cells haven’t subcutaneous tumorigenicity, so we didn’t perform experiments in vivo.