

## Reply to Reviewer 1

Thank you for your valuable comments and suggestions, which were helpful for improving our manuscript.

### Comments and answers

Comment 1: How was 5-HT released after intravenous administration of cisplatin? The authors should discuss in detail.

Answer 1: Following your suggestion, we added the following sentence in the discussion.

Anticancer drugs distribute on the rapidly dividing cells in the GI tract. The action of these cytotoxic drugs initiates within the GI tract, resulting in histopathological changes, activation of the biosynthesis of 5-HT, and increase in the concentration of 5-HT.

Comment 2: The results after phentolamine, propranolol, atropine and metoclopramide administration should be shown in figures.

Answer 2: Following your suggestion, we did not show Figures after phentolamine (regitin<sup>®</sup>), propranolol (inalderal<sup>®</sup>), atropine and metoclopramide administration because these Figures were similar to Figure 3 after dexamethasone administration.

Comment 3: In Figure 3, decadron is the trade name. It should be replaced by the common name such as dexamethasone. In Figure 4, 5B, 6 and 7, kytril is also the trade name.

Answer 3: Following your suggestion, we re-wrote in Figures 3, 4, 5B, 6 and 7.

Comment 4: In Figure 8A, 8B, 9A, 9B and 9C, the time points of 5-HT administration should be indicated.

Answer 4: Following your suggestion, we re-wrote in Figures 8A, 8B, 9A, 9B and 9C.

Reply to Reviewer 2

Thank you for your valuable comments and suggestions, which were helpful for improving our future study.