**Author’s response to reviews**

**Title:** The effects of oral administration of Cola nitida on the pharmacokinetic profile of metoclopramide in rabbits

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**POINT BY POINT RESPONSE TO REVIEWERS' COMMENTS**

**REVIEWER 3**

**COMMENT 1:** "Cola nitida produced a reduction in the oral exposure/bioavailability of metoclopramide. This reduction strongly suggests an increased catalytic activity of CYP2D6 and SULT2A1 and/or efflux function of P-gp in the intestine and/or liver by Cola nitida."

The data presented do not suggest anything regarding CYP activity changes. It should be emphasized, instead, that these could be potential mechanisms that were NOT investigated in the present work.

Also, a similar conclusion should be removed from the Abstract. The abstract conclusion should just contain the text in black print, indicating a possible interaction.

**RESPONSE:** Conclusion has been rephrased to include reviewer’s suggestions and also, a similar conclusion in abstract has been removed. Abstract page 1, lines 18; Conclusion section, page 9, lines 3-4.

**COMMENT 2:** In addition, I do not know why the authors say in their response to Reviewer 4 that the animals received paracetamol - it was never mentioned in previous versions of the manuscript?

**RESPONSE:** So sorry for the typing error. The animals did NOT receive paracetamol in the study. I was writing a different manuscript on paracetamol in the course of making corrections for this work. So, I must have typed in paracetamol in error.

**REVIEWER 4**

**COMMENT 1:** Please be more explicit in your discussion about the weaknesses of the experimental design with respect to controls and small sample size

**RESPONSE:** This has been done in the discussion section, lines 1-4

**COMMENT 2:** previous criticism of the figure plotting appears to have been misunderstood. My
criticism was of the meandering lines joining points which do not have any basis in the data (see attached file). I do not think the references you supplied justify this approach.

RESPONSE: Graph has been replotted to remove meandering lines joining data points (Figure 2).