Author’s response to reviews

Title: Determination of Parecoxib and Valdecoxib in Rat Plasma by UPLC-MS/MS and its Application to Pharmacokinetics Studies

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Author’s response to reviews:

Dear Editor and reviewers,

We received the letter with pleasure on the required minor revision to our submitted manuscript (Submission-ID: PHAT-D-19-00201R2) and appreciated very much for offering us an opportunity to improve our work. Following the revision, we tried answering questions point by point. Here we submit the revised clean version for your reference and hope it may have an opportunity to appear in this prestigious journal.

Thanks again for your kindest attention. We look forward to hearing from you soon,

Sincerely yours,
Yijie Chen and Ledan Wang

Reviewer Comments for Manuscript.

Marcia Ratner (Reviewer 1):

Q1: The language in this paper has been improved somewhat from the original submission but numerous language problems remain which make this paper difficult to read.

A1: Thanks for the reviewer’s concern on this issue. A native English speaker has edited the revised manuscript, and I hope it would be more suitable for publication.

Q2: While the paper employs suitable controls including a determination of the stability of the samples used for method validation, the impact of this methodological improvement for determining parecoxib...

A2: Thanks for the reviewer’s comments on this issue. We acknowledge that the method we used in the current work is indeed not groundbreaking, but we achieved some improvements compared to Liu M’s work. For example, Liu M’s work has obtained a great test, yet the employed method always calls for complicated sample preparatory processes including plasma protein precipitation, along with a liquid-liquid extraction strategy and follow by complete evaporation of organic solvent. In this regard, the one-step protein precipitation method was introduced in our study to simplify the process. In the other hand, chromatographic condition settings are prerequisite to acquire reliable results so that optimization of conditions are needed. The mobile phase was supplemented with 0.1 % (v/v) formic acid to obtain symmetrical peak shapes and to improve ionization efficiency in our work. However, Liu M, et al used ammonium acetate to instead of formic acid that indeed can lead to forming distinct peak shapes but compromised ionization efficacy. Taken together, we have made many improvements compare to that of Liu M, et al’s work.

Vijayakumar Sekar, M.Sc.,PGDSP.,Ph.D (Reviewer 2):

Q: Please include all comments for the authors in this box rather than uploading your report as an attachment. Please only upload as attachments annotated versions of manuscripts, graphs, supporting materials or other aspects of your report which cannot be included in a text format. Please overwrite this text when adding your comments to the authors.

A: We appreciate the reviewer’s comments.

Enayatollah Seydi (Reviewer 4):

Q: The article is acceptable.

A: We appreciate the reviewer’s positive comments.