Author’s response to reviews

Title: Physiological and pharmacokinetic effects of oral 1,3-dimethylamylamine administration in men

Authors:

Brian K. Schilling (bschllng@memphis.edu)
Kelley G. Hammond (kghmmond@memphis.edu)
Richard J. Bloomer (rbloomer@memphis.edu)
Chaela S. Presley (cpresle4@uthsc.edu)
Charles R. Yates (cyates4@uthsc.edu)

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The authors reported in the introduction (lines 107-112) that 25 mg is the typical dosage on the dietary supplements containing DMMA. In the conclusions (line 302) the authors also say that 25 mg is the recommended dose. In the previous Revision, a question was made about the characterization of the recommended dose (Question 2). The authors reply was that “The exact doses of DMAA in Jack3d and Oxyelite Pro are not available, as the supplements have a proprietary blend.” As such, there is no indication that 25mg is the recommended dose. In addition, there are no "recommendations" for DMMA in the literature. This type of characterization is not acceptable and should be removed.

Changed as suggested, line 109

The plasma DMMA analysis section was improved since the last revision. However, the chromatograms show a double peak on the DMMA and an additional peak for 2-aminoheptane. I believe that the latter is an internal standard, but that must be described in the method. In addition, the double peak was previously characterized as a mixture of diastereoisomers (see the reference 14). The authors have decided to treat these two diastereoisomers as one single entity, but that remains to be proved, and a thoroughly discussion on this must be included in the manuscript. Eventually, a new PK characterization on each individual diastereoisomers should be considered.

Changed as suggested, lines 169, 176, 321.

Minor Essential Revisions

line 268, the reference "anecdotal reports" must be removed.

Changed to “These data are important to ensure that reports in the lay media of those reportedly taking DMAA suffering heat injury [18] can be contextualized.”, line 277.

line 305, the text "with mean peak values being <70ng•mL-1” must be corrected.
The reported mean Cmax was 76.5 ng/ml.
Changed as suggested, line 314.

The ppb unit should be presented as ng/ml.

Changed as suggested, lines 172-176 and figure legends.