Author's response to reviews

Title: Pyrrolo[1,2-B][1,2,5]Benzothiadiazepines (PBTDs) induces apoptosis in K562 cells

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Author's response to reviews: see over
Dear Editor-in-Chief

We have revised the manuscript. It is now resubmitted. We followed all the specific comments of the two reviewers, as outlined below.

We hope that the new manuscript version is satisfactory.
Yours sincerely,

Dr. PhD  Gabriella Marfe
Reviewer #:1 Yoo Hong Min

Pyrrolo[1,2-b][1,2,5]benzothiadiazepines (PBTDs) induce apoptosis in k562 cells

In this manuscript, experimental data results indicate that some of the tested analogs induce cell death activating a complex cascade of intracellular biochemical events which, even though partly understood, are different and somehow peculiar for each molecule. These events could contribute to some of pharmacological effects associated with PTBDs treatment, including the previously described antitumor activity in CML cell lines. Future studies will be devoted to analyzing the possible role of compounds-induced apoptosis in other experimental systems, including in vivo antitumor activity focus together with ATP competitor TK inhibitors.

- **Reply:** In the new version we have performed others experiments in order to understand the role of Bcl-2 in PBTDs-induced apoptosis.
- **Reply:** In the new version we improved grammatical English
Reviewer #2: Cecil Pace-Asciak

Pyrrolo[1,2-b][1,2,5]benzothiadiazepines (PBTDs) induce apoptosis in k562 cells

Thank for your suggestions which have been helpful to us.

- **Reply:** In the new version we have performed others experiments in order to understand the role of Bcl-2 in PBTDs-induced apoptosis.

- **Reply:** In the new version we improved nomenclature of compounds.

- **Reply:** Thank for your suggestions. In our lab, we are testing new compounds (~30) in order to improve the chemical structure for modulating the apoptotic activity. After these experiments, we will start the experiments in vivo.