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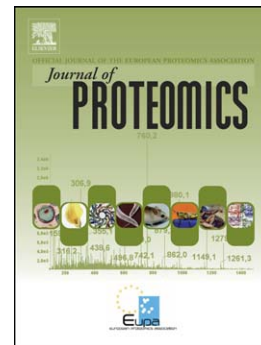
Proteomic approach to nanotoxicity

Magdalena Matysiak, Lucyna Kapka-Skrzypczak, Kamil Brzóska, Arno C. Gutleb, Marcin Kruszewski

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Answers to the reviewers comments

Ms. Ref. No.: JPROT-D-15-00341

Title: Proteomic approach to nanotoxicity

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Reviewers' comments:

Reviewer #1:

The authors present numerous articles on proteomic approaches to nanomaterials from various fields, but these are not adequately covered or reconciled in enough detail. For example, the biological responses and bioaccumulation potential to nanomaterials are clearly different in vivo and in vitro, which should be addressed. The authors should at least add information on living organisms and cells as analyzed by proteomics in Tables 1 and 2. Based on this acknowledgment, they should also mention the relationships between proteins altered by exposure to nanomaterials and the nanomaterial phenomena witnessed in living organisms and cells.

We are a bit surprised with this comment. There is no doubts that in vitro and in vivo response to nanomaterials differ substantially. Thus, our review was already divided into two main sections, ex vivo and in vitro response, and in vivo response. This is reflected by Tab. 1 and Tab 2. Moreover both tables included detailed description of nanomaterials used in the particular study, experimental model and the observed outcome. Nevertheless, a short commentary of the observed results is now added to the summary section. Moreover, an additional graph (Fig. 2) has been added, summarizing subject/specimen that have been studied so far.

A careful reread of the article for English grammar and style is strongly recommended by the reviewer as well.

English is not our mother-tongue language. Despite all efforts we put in English learning, it is clear that we still have a lot to improve. We appreciate all efforts put by both reviewers to improve comprehension of our manuscript. All grammar and style corrections suggested by both reviewers, as well as misplaced references were included, with the exception of Reviewer 2 comment to line 26 [*appliances = applications (?)*]. According to the Webster on-line dictionary '*appliance*' is "an instrument or device designed for a particular use or function", and this was our intention.

Reviewer #2: The manuscript by Matysiak et al. collects available information on a so far neglected topic in toxicology. Data are scattered because many different types of both trigger NPs and target biological systems are used in the exposure tests. With this background, drawing conclusions is certainly not an easy task, however the authors should attempt a summary of the main findings. In its present form the text is mainly a list, whose content, in addition, is duplicated by the Table: a more critical perspective should be taken throughout.

As noticed by the Reviewer, due to the scarcity of data, their sketchiness and inconsistency, drawing reasonable conclusions is not an easy task. Nevertheless, as suggested, a short conclusions summarizing in vitro and in vivo observations were added to the Summary section.

Language and style are uneven; the first section requires a more extensive revision than the rest. I list

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