Author’s response to reviews

Title: Optic glioma and precocious puberty in a girl with neurofibromatosis type 1 carrying an R681X mutation of NF1. Case report and review of the literature.

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Author’s response to reviews:

Dear Editor,

We thank the reviewers for most useful comments and suggestions. We believe that the revised manuscript will be suitable for publication in BMC Endocrine Disorders.

Please find our answers about the changes that we introduced in the revised manuscript as suggested by reviewers

Reviewer #1

We changed the Tanner stage for breast development to B1…B2 in the text and in the Table 1

We have also included in the case report the results of GH values on stimulation test and on OGTT, as well as the IGF1 values

Reviewer #2
We are thankful for the very useful criticism and suggestions. We have made major revisions to the manuscript as follows:

1. We reviewed the literature carefully, and we agree about the frequency of optic glioma and precocious puberty in NF1 children. The important study of Gan HW was published online (26 July) after the submission of our manuscript (7 July), and at that time we were not aware of his important findings. We revised the wording in the abstract, introduction and the discussion accordingly. We also cited the suggested authors.

2. The criticism of the sentence “Some authors also suggest that even children without OPG are at risk of developing CPP as in non NF1 population (Zacharin et al)”: We were aware of the letters to the editor by Bertelloni and Habiby, but also of the answer by Zacharin. Since it is not crucial for this manuscript, we have omitted the sentence and the author from the citations.

3. We agree that the results of our literature search do not support strongly the idea of testing NF1 children with CPP and optic glioma for the R681X mutation at this time. Therefore we have changed the conclusion.

Editorial requests
We reviewed again the instructions of the journal and prepared revised manuscript accordingly.

Consent
All procedures were performed according to the Declaration of Helsinki and approved by the Ethics Committee at the University Pediatric Clinic. Written informed consent was obtained from the patient’s parent for publication of this Case report and any accompanying images. A copy of the written consent is available for review by the Editor of this journal.

Availability of data
All patient data, including laboratory analyses, imaging scans, and genotyping results are part of the electronic and paper patient record system at the University Pediatric Clinic of Skopje, available upon request.

Competing interests
The authors declare that they have no competing interests.

Authors’ contributions

The patient was under the care of MK, who conceived the idea of the report and drafted the manuscript. EK participated in the writing, review of the literature, text editing and finalization of the manuscript. ESA carried out the molecular genetic studies and participated in the review of the literature. All authors read and approved the final manuscript.