

Answering reviewers (62962)

Reviewer #1:

Scientific Quality: Grade C (Good)

Language Quality: Grade A (Priority publishing)

Conclusion: Major revision

Specific Comments to Authors: It is a good presentation of two cases of nasal metastasis from neuroblastoma, a rare entity. However, there are some issues Abstract In case summary 7-18: It should be included that both NB express PHOX2B 18. CASE presentation 24-25: In case 2. It should be included that she has a history of NB Figure and figure legends 2-3. Fig. 1: For a better exposure of them. It is necessary to include CT scan of both cases. 12-13. Fig. 3: it should be included that: similar findings has been observed in case 2 (image or figure not shown)

Response: Thank you for your assessment. Case summary 7-18: The "PHOX2B" was added. Case presentation 24-25: The sentence about the past history of case 2 was added. Figures and figure legends: the images of CT scan, histological, IHC and FISH of both cases were included in Fig.1-3 in the revised manuscript.

Reviewer #2:

Scientific Quality: Grade C (Good)

Language Quality: Grade A (Priority publishing)

Conclusion: Accept (General priority)

Specific Comments to Authors: In the manuscript entitled "Nasal Metastasis from Neuroblastoma-a rare entity: Report of Two Pediatric Cases and a Review of the Literature", the authors showed two cases of NB metastasis at which the authors found that PHOX2B, a protein participating the development of the peripheral nervous system, is expressed in most cases of NB, but not those of olfactory neuroblastoma. Therefore, I suggest that PHOX2B might be the helpful tool to differentiate NB from olfactory neuroblastoma although only two cases were examined in this study. In conclusion, I suggest that this article should be considered for publication on BPG journal after carefully reviewing.

Response: Thank you for your assessment.

Reviewer #3:

Scientific Quality: Grade D (Fair)

Language Quality: Grade B (Minor language polishing)

Conclusion: Major revision

Specific Comments to Authors: Dear Editor, Thank you for the opportunity to revise this interesting paper. The authors presented two pediatric case report of nasal metastases from Neuroblastoma, and they included a sufficiently focused review of the current literature on this issue. My suggestions are reported below: - To me it seems confusing to present the two cases in a consecutive way. Think to describe each cases separately. In fact the reader can have a view of both cases in Table 1 that contains sufficient informations to distinguish the characteristics of the two patients. - The plural of metastasis is metastases so the title should be modified into Nasal "metastases..." - Line 9, page 3: please modify into "The radiologic imaging results..." - Line 14, page 3: It should be better "...in both cases" - Line 15, page 3: It should be better "Finally,.." - Line

16, page 3: Instead of "with this disease" it should be better "The two patients affected by NB.." - Linea 20, page 3: Instead of "differ", it should be better "to differentiate..." - Line 27, page 3: Please change this sentence "...Nasal metastasis from neuroblastoma (NB) is rare and prone to be misdiagnosed when this disease initially presents with a nasal mass without a history of NB" into " ..Nasal metastasis from neuroblastoma (NB) is rare disease that can be misdiagnosed in cases of initial presentation with a nasal mass without a history of NB" - Line 2, page 4: the sentence "It is difficult to differ metastatic.." should be reformulated into "it can be difficult to differentiate between NB..." - Line 10, page 4: "and only one case has been reported", please add "...and only one ... to date" - In this sentence, the authors did not explain properly concerning the NB localization, when and for how long the patient was undergoing chemotherapy, please clarify this sentence "...without any obvious cause in November 2017 when she still received chemotherapy because of NB". Maybe it should be deleted from this section and better explained in the section of history of past illness. - In the section History of past illness, please provide more details about surgical resection (R0?), chemotherapy, radiotherapy and stem cell transplantation chemotherapy (which schemes?)... - The authors provided only radiological and histological images only for case 1; they should provide separately the images for both the two cases. - It should have been also more impressive if the authors could have provided further two non radiological photos of the facial masses. - Line 25-26, page 6: the sentence "All the pathological results were examined by at least two pathologists", please specify the exact number of pathologists involved in the histopathological analysis. - Line 1, page 7: "The tumor was...But the diagnosis" please add the "final diagnosis" - Line 11-12, page 7 please provide specific details about the therapeutic schemes for chemotherapy and RT. - Line 1, page 8 please change automatic into spontaneous. - In table 1 legend it should be specified "....from adrenal NB" - The authors could briefly propose a schematic diagnostic final solution - The paper requires also a further moderate language editing for diverse grammar, and punctuation errors.

Response: Thank you for your assessment.

-Case presentation: Because of BPG journal online submission system, in which the revised manuscript is automatically generated, this study will automatically present the two cases in a consecutive way. If we could, we would like to describe each case separately.

-Title: The word "metastasis" in the title was modified into "metastases".

- Line 9, page 3: The words "their imaging results" were modified into "the radiologic imaging result".

- Line 14, page 3: The words of "in them" were modified into "in both cases".

- Line 15, page 3: The word "therefore" was modified into "Finally".

- Line 16, page 3: The words "with this disease" were modified into "affected by NB".

- Linea 20, page 3: The word "differ" was modified into "differentiate".

- Line 27, page 3: The sentence "Nasal metastasis from neuroblastoma (NB) is rare and prone to be misdiagnosed when this disease initially presents with a nasal mass without a history of NB" was modified into "Nasal metastasis from neuroblastoma (NB) is a rare disease that can be misdiagnosed in cases of initial presentation with a nasal mass without a history of NB".

- Line 2, page 4: the sentence "It is difficult to differ metastatic NB in the nose from olfactory neuroblastoma in the absence of a history of NB" was reformulated into "It can be difficult to differentiate between metastatic NB in the nose and olfactory neuroblastoma in the absence of a history of NB".

- Line 10, page 4: The word "to date" were added.
- In this sentence, the authors did not explain properly concerning the NB localization, when and for how long the patient was undergoing chemotherapy, please clarify this sentence "...without any obvious cause in November 2017 when she still received chemotherapy because of NB". Maybe it should be deleted from this section and better explained in the section of history of past illness (Line 4-6, page 4): The words "when she still received chemotherapy because of NB" were deleted.
- In the section History of past illness, please provide more details about surgical resection (R0?), chemotherapy, radiotherapy and stem cell transplantation chemotherapy (which schemes?)(Line 4-6, page 4): The more details about the treatments in case 2 were added in the section of history of past illness of case 2.
- The authors provided only radiological and histological images only for case 1; they should provide separately the images for both the two cases: The radiological, histological, IHC and FISH images were provided in Fig. 1-3. The images of the two cases were shown separately.
- It should have been also more impressive if the authors could have provided further two non radiological photos of the facial masses: It is a pity that the non-radiological photos were not taken and could not be provided in the revised manuscript.
- Line 25-26, page 6: The number of pathologists was three and the word "three" was added.
- Line 1, page 7: The word "final" was added.
- Line 11-12, page 7: the specific details about the therapeutic schemes for chemotherapy and RT were added.
- Line 1, page 8: The word "automatic" was changed into "spontaneous".
- In table 1 legend: The words "from NB" was modified into "from adrenal NB".
- The authors could briefly propose a schematic diagnostic final solution: A Schematic diagram of diagnosis procedure was provided in Fig. 4.
- The manuscript has been edited for its language and style by native speakers.