Dear Editorial Office, Company Editor-in-Chief, Jin-Lei Wang:

We are grateful to the reviewers for his/her careful reading and helpful comments. We have revised the manuscript, according to the comments and suggestions of the reviewer, and responded point by point to the comments as listed below. And the detailed changes were made in the corresponding full-text.

Replies to Reviewer

First of all, we thank the reviewer for his/her positive and constructive comments and suggestions.

Reviewer #1:

Specific Comments to Authors: This report presented a 6-year-old boy diagnosed with Crouzon syndrome caused by a de novo mutation in the fibroblast growth factor receptor 2 (FGFR2) gene - c.1026C>G (p.Cys342Trp). Since Crouzon syndrome is rare genetic disorder, this case is worthy of reporting. However, there are a few minor issues in the current manuscript.

Comment (1) To clarify boy/girl twins, "fraternal twins" may be used.

Answer: Thank you for your kind comment, we have changed the term "twins" by "fraternal twins", changed the term "twin boy or twin girl" by "fraternal twin boy or fraternal twin girl" in this paper.

Comment (2) In Core tip, you mentioned "There are no case reports showing the Crouzon syndrome occurs in the twins." But, this statement is NOT correct according to the following reference, which should be rephrased. Lloyd MS, Trost JG, Khechoyan DY, Hollier LH Jr, Buchanan EP. Identical Twins with Crouzon Syndrome: Eight-Year Follow-up, Genetic Considerations, and Operative Management. Craniomaxillofac Trauma Reconstr. 2017 Dec;10(4):286-291. doi: 10.1055/s-0036-1592091. Epub 2016 Sep

2. PMID: 29109840; PMCID: PMC5669987.

Answer: Thank you for your constructive suggestion, we have read the reference "Lloyd MS, Trost JG, Khechoyan DY, Hollier LH Jr, Buchanan EP. Identical Twins with Crouzon Syndrome: Eight-Year Follow-up, Genetic Considerations, and Operative Management. Craniomaxillofac Trauma Reconstr. 2017 Dec;10(4):286-291. doi: 10.1055/s-0036-1592091. Epub 2016 Sep 2. PMID: 29109840; PMCID: PMC5669987" and realized that our statement was wrong. So we corrected that incorrect statement as follows: Rarely has this syndrome been seen and evaluated in fraternal twins, only one of whom has the Crouzon syndrome.

Comment (3) Please correct grammatical and spelling errors. For example: In the Title, "…. resulting from a novo mutation,.." "The most common feartures include…" "Since the twin sister and the parents did not present gene mutations and ta similar presentation,"

Answer: Thank you for your kind suggestion, we have corrected grammatical and spelling errors in this paper. We have corrected "…. resulting from a novo mutation,.." by "in a fraternal twin", "The most common feartures include…" by "The most common features include…", "Since the twin sister and the parents did not present gene mutations and ta similar presentation," by "Since the fraternal twin sister and the parents did not present gene mutations and similar presentation". And we have checked the grammatical and spelling issues in the full text.

Comment (4) Please remove "case report" from Key Words.

Answer: Thank you for your kind suggestion, we have removed "case report" from Key Words.

Comment (5) Please rephrase the Conclusion parts to emphasize the patient presented, rather than providing general statements regarding Crouzon

syndrome.

Answer: Thank you for your positive and constructive suggestions, we have rephrased the Conclusion parts as follows:

"CONCLUSION: In summary, Crouzon syndrome could occur in a fraternal twin caused by a de novo mutation of the *FGFR2* gene, characterized by craniosynostosis, a prominent forehead, midface hypoplasia, and prortosis. Oral hygiene instruction and preventive programs on oral hygiene should be performed regularly. A multidisciplinary approach involving oral and maxillofacial surgeons, and orthodontists were required for the treatment of midface hypoplasia. "

Reviewer #2:

Specific Comments to Authors: Dear author, It is well written rare case, quality is fair and as basis try to gain more cases.

Answer: Thank you for your positive and constructive suggestions. Since Crouzon Syndrome is a rare genetic disorder, we will try to gain more cases of Crouzon Syndrome.

Replies to editorial office's comments:

First of all, we thank for your kind and constructive comments and suggestions.

(1) Science editor:

1. To clarify boy/girl twins, "fraternal twins" may be used. 2. In Core tip, you mentioned "There are no case reports showing the Crouzon syndrome occurs in the twins." But, this statement is NOT correct according to the following reference, which should be rephrased. Lloyd MS, Trost JG, Khechoyan DY, Hollier LH Jr, Buchanan EP. Identical Twins with Crouzon Syndrome:

Eight-Year Follow-up, Genetic Considerations, and Operative Management. Craniomaxillofac Trauma Reconstr. 2017 Dec;10(4):286-291. doi: 10.1055/s-0036-1592091. Epub 2016 Sep 2. PMID: 29109840; PMCID: PMC5669987. 3. Please correct grammatical and spelling errors. For example: In the Title, "…. resulting from a novo mutation,.." "The most common feartures include…" "Since the twin sister and the parents did not present gene mutations and ta similar presentation," 4. Please remove "case report" from Key Words. 5. Please rephrase the Conclusion parts to emphasize the patient presented, rather than providing general statements regarding Crouzon syndrome.

Answer: Thank you for your constructive suggestions, we have corrected the issues in the manuscript.

(2) Company editor-in-chief:

I have reviewed the Peer-Review Report, the full text of the manuscript, and the relevant ethics documents, all of which have met the basic publishing requirements of the World Journal of Clinical Cases, and the manuscript is conditionally accepted. I have sent the manuscript to the author(s) for its revision according to the Peer-Review Report, Editorial Office's comments and the Criteria for Manuscript Revision by Authors. Before its final acceptance, the author(s) must provide the Signed Consent for Treatment Form(s) or Document(s). For example, authors from China should upload the Chinese version of the document, authors from Italy should upload the Italian version of the document, authors from Germany should upload the Deutsch version of the document, and authors from the United States and the United Kingdom should upload the English version of the document, etc. Please provide the original figure documents. Please prepare and arrange the figures using PowerPoint to ensure that all graphs or arrows or text portions can be

reprocessed by the editor. Authors are required to provide standard three-line tables, that is, only the top line, bottom line, and column line are displayed, while other table lines are hidden. The contents of each cell in the table should conform to the editing specifications, and the lines of each row or column of the table should be aligned. Do not use carriage returns or spaces to replace lines or vertical lines and do not segment cell content.

Answer: Thank you for your kind suggestions, we have provided the Signed Consent for Treatment Form(s) or Document(s), the original figure documents, and the original figure documents as required.

Best regards

Ji-Mei Su, MD, Chief Doctor，Professor

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