Reviewer #1: The case-report, 'Two novel mutation in the VPS33B gene in a Chinese patient with ARCS1: A case report', presented here has identified two novel mutations in the gene responsible for the ARCS1 disease. The case-report presents here two new mutations with a potential for employed in genetic counselling and testing. This is an interesting work of investigation and few comments for the clarity of work presented are: 1) The manuscript file attached is not the final formatted version and still incorporates the highlighted changes. 2) The manuscript could benefit from minor language editing, for example, the word 'mutation' in the title should be 'mutations' to show the plural nature of the work described 3) Introduction section has not detailed the role of genes mentioned and just alluded the names as VPS33B and VPS33B. They should be elaborated with their role in the disease pathogenesis. 4) There are no details about the whole-exome sequencing and CNV analysis being referred to, that should also be clearly explained with the methodology adopted. 5) The manuscript has not detailed the role of reported mutations in the pathogenicity of the disease and how they concluded their role in this particular phenotype. 6) Figure 2 and Figure 3 have not detailed the methodology used to obtain those chromatograms shown.

Thanks for your comments, we have made the following adjustments based on your advice:
1) We revised the content of the article based on the opinions of the reviewers to ensure the accuracy of the manuscript;
2) We replaced part of the mutation with mutations to ensure the accuracy of the grammar;
3) We briefly describe some clinical manifestations of abnormal VPS33B gene expression. Unfortunately, we have not yet conducted any molecular studies.
4) We provide a supplementary description of the details of whole-exome sequencing and CNV analysis.
5) According to ACMG guidelines, the clinical significance of the variation was pathogenic, and other gene mutations associated with the patient’s phenotype were not detected.
6) We have added the method to the main text (it could be seen in 4).

Reviewer #3: This is an interesting read of a child with cholestasis and hypothyroidism that turned out to have 2 compound heterozygous in the VPS33B gene. Their finding adds on to the understanding of genetic mutations involved in ARCS1. Nevertheless, quite a number of sentences are unclear of their meaning in the case description. Sentence structure needs some revision. Perhaps, this could be relook into by the authors so that the content remains valid and not affected by translation work. Some examples: "Breath sounds in both lungs were rough, and dry and moist rales were not heard., Embrace reflex and sucking reflex could elicit., The ends of the limbs were warm." and many more.

Thanks for your comments, we have made adjustments to the relevant description to avoid unclear expressions.
RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Clinical Cases
Manuscript NO: 77064
Title: Two novel mutations in the VPS33B gene in a Chinese patient with ARCS1: A case report

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind
Reviewer’s code: 06079635
Position: Peer Reviewer
Academic degree: BSc, MSc, PhD
Professional title: Assistant Professor
Reviewer’s Country/Territory: Pakistan
Author’s Country/Territory: China
Manuscript submission date: 2022-04-13
Reviewer chosen by: Han Zhang
Reviewer accepted review: 2022-07-27 09:34
Reviewer performed review: 2022-07-27 09:56
Review time: 1 Hour

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<th>Grade B: Very good</th>
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Peer-reviewer: [Y] Anonymous [ ] Onymous
SPECIFIC COMMENTS TO AUTHORS
The manuscript has been revised and authors have made the suggested changes. The manuscript still needs a thorough editing for grammar and typos particularly the section on physical examination. Also, the primer sequences should be added that were used to sequence the region of the gene mentioned in the manuscript. It will also be good to incorporate the images to show the predicted structural variation upon analysis by REVEL.

Answer:
Thanks for your suggestions. To ensure the integrity and accuracy of the article, we deleted the relevant expressions of the molecular research of genes, and considered the detailed discussion in the new article after the completion of the experiment.
**RE-REVIEW REPORT OF REVISED MANUSCRIPT**

**Name of journal:** *World Journal of Clinical Cases*

**Manuscript NO:** 77064

**Title:** Two novel mutations in the VP 3B gene in a Chinese patient with ARC : A case report

**Provenance and peer review:** Unsolicited Manuscript; Externally peer reviewed

**Peer-review model:** Single blind

**Reviewer’s code:** 05086048

**Position:** Editorial Board

**Academic degree:** PhD

**Professional title:** Postdoctoral Fellow

**Reviewer’s Country/Territory:** Germany

**Author’s Country/Territory:** China

**Manuscript submission date:** 2022-04-13

**Reviewer chosen by:** Han Zhang

**Reviewer accepted review:** 2022-07-27 19:54

**Reviewer performed review:** 2022-07-27 19:57

**Review time:** 1 Hour

### Scientific quality

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### Conclusion

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**Peer-reviewer**

Peer-Review: **Anonymous**
SPECIFIC COMMENTS TO AUTHORS

Manuscript may be accepted in the present form

Answer:

Thank you for your review and help!
RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Clinical Cases

Manuscript NO: 77064

Title: Two novel mutations in the VP 3B gene in a Chinese patient with ARC: A case report

Provenance and peer review: Unsolicited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer’s code: 05204241

Position: Editorial Board

Academic degree: FRCP (C)

Professional title: Staff Physician

Reviewer’s Country/Territory: Malaysia

Author’s Country/Territory: China

Manuscript submission date: 2022-04-13

Reviewer chosen by: Han Zhang

Reviewer accepted review: 2022-07-27 13:29

Reviewer performed review: 2022-08-03 00:05

Review time: 6 Days and 10 Hours

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SPECIFIC COMMENTS TO AUTHORS

There are major typo errors in this version. "There were no obvious abnormalities in routine blood examination. Liver function showed: total bilirubin (TBiL) 229.7 +ACY-mu+ADs-mol/L, direct bilirubin (DBil) 152.4 +ACY-mu+ADs-mol/L, and total bile acid (TBA) 50.6 +ACY-mu+ADs-mol/L. Neonatal jaundice was diagnosed. i am not sure if this errors occurred due to the incompatible format but this makes the whole manuscript difficult to read.

otherwise, the scientific content is acceptable

Answer:

Thank you for your suggestions. After our testing and communication, we found that this is because the characters µ contained in the common units of the laboratory inspection can not be identified in the submission system. We have sent the word and PDF version of the manuscript to the editor in charge to more accurately describe our content.