

To the Editors

World Journal of Hepatology

**We thank the editors for considering to accept our manuscript “Mitochondrial Hepatopathy: Respiratory Chain Disorders- ‘breathing in and out of the liver’ ” and giving us an opportunity to submit with revisions. We also thank the esteemed reviewers for their valuable inputs.**

Please find the point to point responses to the reviewers below. Our answers are in italics. **The modifications in the main manuscript are in blue font for the reviewers and editors to track.**

**Reviewer #1:**

**Language quality: Grade A (Priority publishing)**

**Scientific quality: Grade A- Excellent**

**Conclusion: Accept (High priority)**

**Comments to authors & authors’ response :**

1. I was extremely interested in reading the manuscript titled: “Mitochondrial Hepatopathy: Respiratory Chain Disorders- ‘breathing in and out of the liver’”. It is one of the best reviews about mitochondrial hepatopathies I ever read. The authors have made a great effort to present their manuscript in a palatable way, making a tough subject easy to read and understand, i.e. reader-friendly.

*ANSWER: We thank the reviewer for the encouraging comments.*

2. Keywords are lacking

*ANSWER: regret We have rectified the error.. We have now updated in both the main text of the paper and the section provided while submitting the manuscript on the portal*

3. Abbreviations are lacking

*ANSWER: We have scanned the manuscript and detailed all abbreviations in the text for those in main manuscript and in figure legends or table legends for those appearing in the table.*

4. Language modification: “Those surviving this early onslaught experience a respite from hematological symptoms<sup>29</sup>” The aforementioned sentence uses some unusual terms, that add difficulty to a tough subject that the authors are attempting to simplify.

*ANSWER: We thank the reviewer for this pertinent comment. As suggested we have modified the sentence to “If the patient survives this phase of hematological symptoms, its intensity begins to decrease over time<sup>29</sup> and symptoms change from hematological to a phenotype of severe pancreatic insufficiency in late infancy to early childhood, during the same time which villous atrophy is found to appear.”*

5. Also, the expression “simply put” was used repeatedly and can be replaced by an easier and more palatable term “In simple words”

*ANSWER: The suggestion has been incorporated and the phrases have been re-written*

6. In general management: “placement of feeding tubes (oro-nasal)”; I think the authors mean orogastric/nasogastric”

*ANSWER: We have rectified the error and rephrased as “placement of feeding tubes (orogastric or nasogastric)” as pointed out by the reviewer*

7. The Figures are excellent particularly figures 4-8, showing the graphical presentation of each disorder. Similarly, most of the tables are very informative. The conclusions are perfectly written.

*ANSWER: We thank with all humility, the reviewers for these gratifying comments.*

## **Reviewer#2**

**Language quality: Grade A (Priority publishing)**

**Scientific quality: Grade C –Good**

**Conclusion: Minor revision**

**Comments to authors & authors’ response :**

This review discusses the burden of mitochondrial respiratory chain defects, including some uncommon diseases, Mitochondrial DNA deletion syndrome. Genetic evaluation is important for the better understand of the disease, however, the reveiw donot

describe how to do genetic testing. The point should be discussed.

*ANSWER: We thank the reviewer for these encouraging comments and pertinent concerns. As suggested we have added a new section on genetic testing with a succinct paragraph and an additional figure (Figure 13). Hope this change will now be suitable for the journal.*

### **Comments from the scientific editor and authors' reply reply**

5 Issues raised:

(1) The "Author Contributions" section is missing. Please provide the author contributions

*Answer: The author contribution section has been added in the online portal*

(2) The "Key Words" section is missing. Please add the key words

*Answer: The key words section has been added in the online submission portal*

(3) The "Core Tip" section is missing. Please add the core tip

*Answer: The core tip section has been added in the online submission portal*

(4) The authors did not provide original pictures. 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

*Answer: The original pictures (author's own) have been now provided as a power-point file. Each figure is now decomposable/movable/editable and is incorporated on one single slide so that the slide number corresponds to the figure number. Figure legends and abbreviations used have been uploaded in the section available in the online submission portal.*

(5) PMID and DOI numbers are missing in the reference list. Please provide the PubMed numbers and DOI citation numbers to the reference list and list all

authors of the references

***Answer:** PMID and DOI numbers have been added in the references as pointed out by the scientific reviewer*

Kind regards,

Dr Amrit Gopan, and

Dr Moinak Sen Sarma (*Corresponding author*)