To,

The Editors,

World Journal of Clinical Pediatrics.

Subject: Corrections and queries in the manuscript

Respected Editors,

I am extremely thankful for your invaluable suggestions and queries on the manuscript. In this letter I have tried to address all the queries which were raised.

In addition, as advised, I have tried to improve upon the grammar of the write up and the quality of the sentences.

Title:

1. Changing the title of the manuscript to "a case report and metanalysis of literature"

Although we agree to the point raised by the review with regards to difficulty in identifying cases of DS with PH due to lack of a screening protocol, we have tried to use the definition of PH as the screening criteria while considering studies for inclusion, hence feel that calling it a case report might narrow the scope of the study hence would request to keep the title same.

Abstract:

- 1. Increased prevalence
 - As advised the phrase "increased prevalence" of pulmonary hypertension, has been edited appropriately.
- 2. Incongruence between statements in abstract and core tip
 - With the help of core tip we had highlighted the need for this study i.e to highlight lack of a screening protocol for PH in DS, this we wanted to undertake by showcasing the prevalence of PH in said patients and. As advised, objective of the study has been added to the coretip to remove any incongruency.
 - 3. The language of the subject matter has been greatly improved upon.

Introduction:

- As advised, we have talked about the higher interferon-related inflammatory profile typical of the DS and its role in causing interferon mediated pulmonary hypertension (Ref. 21 and 22).
- The term trisomy 21 has been changed to Down's syndrome wherever present.
- In line 22 DiGeorge syndrome spelling has been corrected.
- "Children DS" has been corrected to "children with DS"

Methods:

• PROSPERO trial registration number has also been added.

Discussion:

- In the discussion, it is not clear how "the prevalence ranges from various countries range from 42% to 54%"", when the overall prevalence is 25.5%.
- We would like to edit this part as the statistics were wrongly quoted from unsuitable studies hence, this discrepancy of prevalence has been appropriately edited.

Conclusion:

- A need for multidisciplinary approach for diagnosis and management of PH in patients with DS has been emphasized.
- "This article highlights" section has been added as advised by the editor.

References:

• Requested changes have been made in the references with the addition of DOI and PMID of all the possible available references.

I am extremely grateful for your consideration. I welcome any further comments or criticism on the manuscript.

Hoping for a positive response.

Thanking you.

With warm regards,

Dr. Amar Taksande

Dr. Divya Pujari

Dr. Patel Zeeshan Jameel

Dr. Bharati Taksande

Dr. Revat Meshram