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All Author List: Kamila A Kwiatek-Średzińska, Kinga Trochimczyk, Jacek Jamiołkowski and Dariusz M Lebensztejn
Corresponding Author: Kamila A Kwiatek-Średzińska, Department of Pediatrics, Gastroenterology, Hepatology, Nutrition, Allergology and Pulmonology, Medical University of Bialystok, Waszyngtona Street 17, Bialystok 15-274, Poland. kamila.kwiatek-sredzinska@umb.edu.pl
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REVIEWERS' COMMENTS:

Reviewer 1:

1) In the Results section, the authors state, "Table 2 shows the diagnostic performance of urinary trypsinogen-2 after 48 hours as a marker for the diagnosis of AP in children." However, no interpretation of these findings is provided. The authors should clearly explain the clinical significance of these results.

Thank you for the comment. We have expanded the description of Table 2 in the Results section and added an appropriate explanation of the clinical meaning of these findings in the Discussion section.

2) A similar issue is noted for Table 5 and Figure 2, where it is stated that "Table 5 and Figure 2 present the assessment of urinary TAP after 48 and 72 hours as predictors of AP severity compared with serum CRP," without sufficient interpretation.

Thank you for the comment. We have expanded the description of Table 5 in the Results section and added an appropriate explanation of the clinical meaning of these findings in the Discussion section.

Reviewer 2:

1) Title and Keywords Consider ensuring that all major keywords (particularly those central to methodology or population characteristics) are reflected in the title for improved indexing and discoverability.

Thank you for the comment. The title has been revised to better reflect the key concepts of the study, including urinary biomarkers, the pediatric population, and acute pancreatitis, in order to improve indexing and discoverability.

2) Abstract Clarification While the abstract follows the IMRaD structure, numerical results (including effect sizes and confidence intervals where applicable) should be presented more explicitly to strengthen scientific clarity.

Thank you for the comment. We have revised the Results section of the abstract to provide more explicit numerical data.

3) Introduction The introduction would benefit from a brief expansion of the most recent literature (last 3–5 years) to better position the study within current research trends.

Thank you for the comment. The introduction has been expanded to include recent literature.

4) Materials and Methods Please clarify the justification for selected parameters (e.g., sample size calculation, inclusion/exclusion criteria thresholds if applicable). If applicable, specify the software used for statistical analysis and version number. Confirm whether ethical approval reference number and approval date are clearly stated.

Thank you for the comment. Due to the relatively low incidence of pediatric acute pancreatitis, the sample size reflects all eligible consecutive patients admitted during the study period. This clarification has been added to the Materials and Methods section. The version number of the statistical software has been specified. We confirm that the ethical approval number was already provided in the manuscript. We have added the approval date.

5) Results Section Ensure consistency in reporting p-values (e.g., exact values vs. p<0.05). If applicable, consider adding confidence intervals to improve interpretability of effect estimates.

Thank you for the comment. The manuscript has been revised to ensure consistent reporting of *P* values throughout the text. We have added confidence intervals, where appropriate.

6) Figures and Tables Minor formatting adjustments may improve readability. Ensure all abbreviations are defined in table/figure footnotes.

Thank you for the comment. Tables and figures have been revised. All abbreviations are defined in table/figure footnotes, and minor formatting adjustments have been implemented.

7) Discussion Strengthen the section addressing study limitations. Even if limitations are minor, clearly articulating them improves transparency and scientific rigor. Briefly discuss potential future research directions.

Thank you for the comment. We have expanded the discussion on the study limitations and potential future research directions.

8) Language and Style The manuscript is generally well written; however, minor grammatical polishing and sentence restructuring in a few sections would enhance clarity and flow.

Thank you for the comment. The manuscript has undergone additional English language editing.

SCIENCE EDITOR'S COMMENTS:

1) The abstract of the manuscript needs to be revised. The authors should provide more details in the methods section. The authors should clearly state the clinical significance of the results. The authors should strengthen the discussion on the limitations of the study. Minor adjustments to the formatting of figures and tables can improve readability. The language of the manuscript should be improved to enhance clarity.

Thank you for the comment. We have revised the abstract of the manuscript, the Materials and Methods, Results and Discussion sections. Tables and figures have been revised. The manuscript has undergone additional English language editing.

2) **Author contributions:** The 'Author contributions' passage describes the specific contribution(s) made by each author. The author's names will be listed in the following format: full family (sur)name, followed by abbreviated first and middle names.

Thank you for the comment. We have corrected Author contributions section**.**

3) The name of the author(s) of a reference is listed in the sentence, the reference number should be placed immediately after the author(s) of the reference.

Thank you for the comment. We have corrected the placement of the reference numbers.

4) To ensure the accuracy of the references, please use "**Edit References by Auto-Analyser**"

Thank you for the comment. We have revised the references using Auto-Analyser.

5) Abbreviations must be defined upon first appearance in the **Figure Legends.** Do not use non-standard abbreviations, unless they appear at least two times in the text preceding the first usage/definition.

Thank you for the comment. We have revised the abbreviations in the figures.

6) **Figure Color:** When using different color markings on the Figures, please avoid using **red or green**, including the **arrow colors** on the Figures.

Thank you for the comment. We have modified the colours used in the figures.

7) Abbreviations are not allowed in table titles.

Thank you for the comment. We have removed abbreviations in table titles.

8) **AIM.** The purpose of the study should be stated clearly, with no or minimal background information, must start with "To", following the format of: "To investigate/study/determine…".

Thank you for the comment. We have revised the aim.

REVIEWER’S COMMENTS: 1) Minor grammatical and typographical errors remain throughout the manuscript and should be corrected through careful proofreading. Thank you for the comment. The manuscript has undergone additional language editing and proofreading. 2) Some figure legends would benefit from slightly more detailed descriptions of statistical significance and abbreviations. Thank you for the comment. Figure legends were expanded to provide clearer descriptions and abbreviation definitions. 3) The discussion section may be strengthened further by citing a few additional recent references related to the investigated pathway and multidrug resistance mechanisms. Thank you for the comment. We have revised the Discussion section and incorporated additional recent references addressing the role of pathological trypsinogen activation and emerging urinary biomarkers in acute pancreatitis. These references provide further context for the investigated biological pathway and support the interpretation of our findings. As multidrug resistance mechanisms were not directly investigated within the scope of the present study, we focused on literature specifically related to acute pancreatitis pathophysiology and biomarker development. 4) The authors should ensure consistency in formatting, terminology, and abbreviation usage throughout the manuscript. Thank you for the comment. We have carefully revised formatting, terminology, and abbreviation usage and standardized them throughout the manuscript. Thank you for your time and consideration.