

June 12th, 2021

Editor-in-Chief, Cells

Dear Professor / Doctor,

We wish to submit the manuscript entitled **"Indonesia-based study of the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with allergic rhinitis in private practice"** for consideration of publication in the **Cells: Special Issue "Advances in Immunotherapy of Allergic Diseases"**.

In the field of child allergic diseases, the private sector's role in the Indonesian health care system has grown dramatically over the past decade. There is an overall wide acceptance among Indonesian parents, even among the poorest socio-economic groups, to use private sector providers for specific treatment such as subcutaneous immunotherapy which isn't covered by national health insurance.

In our 6-year analysis looking back in time of children newly diagnosed as having rhinitis allergies, we compared clinical efficacy and health care costs incurred during the 18 months before starting immunotherapy to the costs for these same children that were incurred during the 18 months after completion. If the results are effective to improve quality of life, we would like to advocate our government to implement immunotherapy in the national health-care insurance, so the poorest socio-economic groups could also get the benefit of this treatment.

We believe this study may fulfil scientific holes in the field and hopefully fit the scope of your journal. This study has not been published in part or whole or is not under consideration for publication elsewhere. All authors involved in this study have agreed to be listed and approved the manuscript.

We thank you for considering our work for publication in the **Cells: Special Issue "Advances in Immunotherapy of Allergic Diseases"**.

Yours sincerely,

Anang Endaryanto, M.D., Ph.D.

Corresponding author

Department of Child Health, Soetomo General Hospital, Faculty of Medicine, Universitas Airlangga

Mayjen Prof. Dr. Moestopo Street No.6-8

Surabaya 60286, Indonesia

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Phone: +62-811-327-431

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Journal name: Cells

Manuscript ID: cells-1277684

Type of manuscript: Article

Title: Indonesia-based study of the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with allergic rhinitis in private practice

Authors: Anang Endaryanto *, Ricardo Adrian Nugraha

Received: 12 June 2021

E-mails: anang.endaryanto@fk.unair.ac.id,
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Submitted to section: Cellular Immunology,

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Your manuscript has been assigned to Smina Mukhtar for further processing who will act as a point of contact for any questions related to your paper.

Journal: Cells

Manuscript ID: cells-1277684

Title: Indonesia-based study of the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with allergic rhinitis in private practice

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E-mails: anang.endaryanto@fk.unair.ac.id,
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Journal name: Cells

Manuscript ID: cells-1277684

Type of manuscript: Article

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Dear Dr. Endaryanto,
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Dear Editors, Thank you for excellent comments designed to improve our paper, “Indonesia-based study of the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with allergic rhinitis in private practice” which we have addressed below. We greatly appreciate the time and effort put forth by reviewers and editors to improve our paper. If any responses are unclear or you wish additional changes, please let us know.

Sincerely,

Anang Endaryanto, M.D., Ph.D.

Division of Pediatric Allergy and Immunology, Department of Child Health

Faculty of Medicine, Universitas Airlangga - Dr. Soetomo General Hospital, Surabaya – INDONESIA

No.	Reviewer’s comment	Author response
1	I was really delighted to read this interesting study on the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with AR. The paper is well structured, heavily up-dated, and using appropriate statistics provides the reader with an accurate and up-to-date point of view regarding the importance of subcutaneous allergen immunotherapy in the treatment of children with AR. It is a must for every clinician and health care provider involved in the treatment of children with AR.	Thank you!
2	<p>【General comments】</p> <p>This is a study to evaluate the the clinical and cost-saving benefits of subcutaneous allergen immunotherapy for children with allergic rhinitis in private practice.</p> <p>This is a well-designed and well-described manuscript.</p> <p>However there are some points that must be cleared out.</p> <p>【Specific comments】</p> <p>(1)Table 1</p> <p>As for inclusion criteria, severity of symptom should be normalized as the same ARIA classification. Please inform whether there</p>	<p>We agree with the reviewer’s assessment. Accordingly, throughout the manuscript, we have answered some points</p> <p>(1) Table 1</p> <p>We have added the baseline symptom score, medication score, combination before study enrolment. There are no difference between groups in the symptoms score (base of ARIA classification) before study enrolment.</p>

	<p>existed any symptom severity difference between two groups before study enrollment</p> <p>(2) Table 2</p> <p>In Table2, the effect of SLIT on symptom scores(total) is showed.</p> <p>Can authors show any data about the effect on nasal symptom scores(sneezing, rhinorrhea, and nasal congestion) ?</p> <p>Please let me have a comment .</p> <p>I hope that my comments are useful for the improvement of this manuscript.</p>	<p>(2) Table 2</p> <p>Yes, we can. We have added the data regarding nasal symptom points of score data. Data about the effect on nasal symptom scores (sneezing, rhinorrhea, and nasal congestion) were added to a new table (Table 3).</p>
3	<p>The article is interesting and well written. The argument is not new but the cut linked to the economic aspects in a particular Nation makes it unique. The study has the limits of a retrospective study but the large number of sample population ensures the validity of the results. English is correct and reading is fluent. I approve its publication after specifying the adverse reactions of the drug.</p>	<p>Thank you for your constructive comment and suggestion. Some of the most common adverse reactions associated with immunotherapy treatment is chills, constipation, coughing, decreased appetite, diarrhea, fatigue, fever and flu-like symptoms, headache, infusion-related reaction or injection site pain, itching, localized rashes and/or blisters.</p>



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Manuscript ID: cells-1277684

Type of manuscript: Article

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Authors: Anang Endaryanto *, Ricardo Adrian Nugraha

Received: 12 June 2021

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Dear Dr. Endaryanto,

Congratulations on the acceptance of your manuscript, and thank you for your interest in submitting your work to Cells:

Manuscript ID: cells-1277684

Type of manuscript: Article

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Alexander E. Kalyuzhny, Cord Brakebusch

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Received: 12 June 2021

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Authors: Anang Endaryanto *, Ricardo Adrian Nugraha

Received: 12 June 2021

E-mails: anang.endaryanto@fk.unair.ac.id,
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Type of manuscript: Article

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Authors: Anang Endaryanto *, Ricardo Adrian Nugraha

Received: 12 June 2021

E-mails: anang.endaryanto@fk.unair.ac.id,
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Surabaya 60286, Indonesia
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Corresponding author

Department of Child Health, Soetomo General Hospital, Faculty of Medicine, Universitas Airlangga

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Email: anang.endaryanto@fk.unair.ac.id <<mailto:anang.endaryanto@fk.unair.ac.id>>

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