JURNAL INTERNATIONAL BERPUTASI TERAKREDITASI Q3

	CTLA-4 CT-60 A/G and CTLA-4 1822 C/T Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus
JURNAL	The Application of Clinical Genetics - Dove Medical Press Ltd.
	Nur Rochmah ; Muhammad Faizi ; Suhasta Nova ; Retno Asih Setyoningrum ; Sukmawati Basuki ; Anang Endaryanto

No	Perihal
1	BUKTI SUBMIT
2	BUKTIPROSES REVIEW
3	BUKTI KOMENTAR REVIEWER
4	BUKTI RESPON DARI AUTHOR
5	BUKTI ACCEPTED, PUBLISH DAN PAYMENT

Surabaya, 7 Agustus 2023

Dr. NUR ROCHMAH dr., Sp.A(K)

1. BUKTI SUBMIT

Dear Dr Rochmah

I am just contacting you regarding your revised manuscript which was recently resubmitted titled 'CTLA-4 Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus' (ID: 359158)

Upon beginning to process your revised manuscript, we have noticed some issues that prevent us from forwarding your submission to the Editor for final editorial review. Please use the link below to find a list of your files which need further correction. All the required corrections can be found in the included manuscript file as comments. https://www.dovepress.com/upload_files_new.php?submission_id=359158&l=A9YDypACsLaPnI7OrubimGgK1466517

Please perform or respond to each of the requested corrections, then upload the corrected files through the "Upload corrected files' section at the bottom of the page

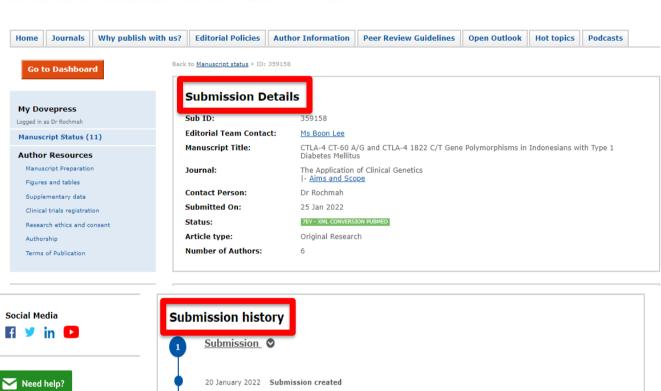
Once we receive the corrected files, we will continue processing your submission, and provided we do not encounter any further issues which need correcting, will notify you once it has been sent to the Editor for final decision.

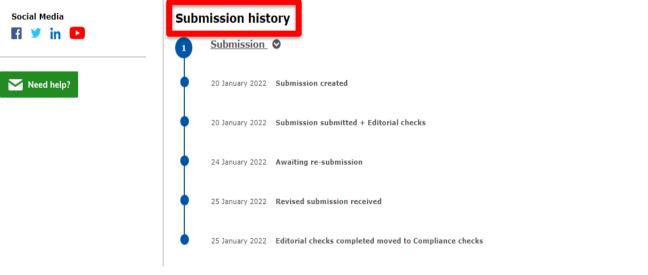
Please don't hesitate to contact me if you have queries about any of the requested corrections.

Regards, Marina Ralph Revised Manuscript Co-ordinator Dove Medical Press

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Manuscript submitted to Dove Medical Press

2. BUKTI PROSES REVIEW



Dear Dr Rochmah

The author proofs for your paper "CTLA-4 CT-60 A/G and CTLA-4 1822 C/T Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus" [ID 359158] are now available by clicking this link:

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Please make your corrections in the online correction tool and ensure you complete the process by using the green or red button in the top right hand corner.

Ensure you show your co-authors the PDF proofs which can be downloaded from the link above, and use the online correction tool to mark any final corrections on their behalf

Please submit your proof within the next 5 working days (due 27 Apr 2022) to ensure your paper is published in a timely manner.

Do contact me if you have any questions:

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Kind regards

Boon Lee
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3. Bukti Review dan Balasan

Dari editor: kotak biru

Dari author : kotak merah

CTLA-4 Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus

Editorial Corrections:

 Abstract Structure: Original research papers require a structured abstract. A structured abstract is an abstract with distinct, labelled sections (e.g., Introduction, Methods, Results, Discussion) for rapid comprehension. Please amend accordingly prior to re-submission.

Author response:

Thank you for your suggestion, we already revised the manuscript. Please see on abstract section page 1-2 line 24-45.

• Figure Files: Please supply all figures in high quality .jpg, .tif or .pdf format, one file for each figure (eg, Fig 1, Fig 2, etc). This should include the figure artwork but not the figure legends. If the figures have also been placed in your manuscript, please remove these. Please note that the figure legends must not be placed within the figure artwork and should instead be listed at the end of the manuscript file. See the figure page on our website for further details (https://www.dovepress.com/author-guidelines/figures-tables).

Author response:

Thank you for your suggestion, we already revised the quality of the figure, please see the figure file 1 and 2, We also uploaded the figures in pdf format.

 Affiliations/Corresponding Author: The presentation of department/institution details differs between author affiliation list and correspondence details. They both need to align. Kindly check and correct.

· Author response:

Thank you, we have edited as suggested, please see page 1 line 18-20.

Reviewer Comments:

Reviewer 1 Title &

Abstract

1. Do the title and abstract cover the main aspect of the work?

please see comments below

Author response:

Thank you for your suggestion, we already revised the manuscript. Please see on abstract section page 1-2 line 3-45.We also changes the tittle of the manuscript and our affiliation.

2. Does the introduction provide background and information relevant to the study?

please see comments below

Author response:

- Thank you for your suggestion, we already add the new reference. Please see on introduction section page 2-3
 line 53-54
- Thank you for your suggestion, we already revised as your suggestion. We have added some paragraphs in the introduction section. Please see pages 3 lines 55-60 in the revised manuscript.
- Thank you, we have added the control sample in our study in our method, please see page 3 line 71 and on result section page 13 in lines 321-323 tables 1 and 2 of the revised manuscript.

Material and Methods

3. Are the methods clear and replicable? Do all the results presented match the methods described?

No, this manuscript is limited to describe how they got the genotypes for two polymorphisms at gene CTLA4. This is not enough for this study. Authors should also have added some information about the patients, their inclusion/exclusion criteria. Importantly, no ethics aproval or statistical approach is described in this section.

Variants should be named after their rs_id.

Author response:

Thank you for your suggestion, We have edited the above suggestion. Please see page 4 line 85-89 of the revised manuscript. We also added some paragraphs in the study participant and statistical analysis subsection. Please see page 3-4 line 72-82 and page 5 line 108-113 in the revised manuscript. Meanwhile, the variant name in our study is based on another study about CTLA-4 gene polymorphisms by Pastuzak-Lewandoska et al., 2011.

5. Do the findings described by the author correlate with the results? Are the findings relevant?

These findings are, not relevant. This study needs a thorough design by geneticist or statistician. Also, authors should include controls, in addition to expanding the sample size of cases.

Author response:

Thank you for your suggestion. We already added the control sample in our study. Please see on result section on page 13 and line 321-323, table 1 and 2 of the revised manuscript. We also revised our discussion section, please see page 6 lines 135-136, page 2 line 161-162, page 8 187-188 and 192-193.

7. If the author has provided figures and tables are the figures and tables clear and legible? Are the figures free fromunnecessary modification?

Yes, tables are clear. However, the information contained in them is insufficient for reaching the aim stated in the introduction

Author response:

Thank you, We have edited as suggested. We choose the table 1 and 2 to be presented and delete table 3 in the result section, please see page 13 and line 321-323, of the revised manuscript.

8. Does the paper raise any concerns?

I believe this study requieres more thorough statistical design and analyisis.



Author response:

Thank you, we have edited it as suggested. We added the statistical analysis subsection the in material and methods section. Please see page 5 lines 109-113 in the revised manuscript.

English editing

10. Do you think the manuscript requires English editing to correct the grammar or flow?

Author response:

Thank you for your response. This manuscript has been edited to ensure language and grammar accuracy by "editage". We also uploaded the certificate of editing.

Reviewer 2 Title &

Abstract

1. Do the title and abstract cover the main aspect of the work?

Here is the ORIGINAL RESEARCH from Rochman et al on entitle CTLA-4 Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus.

This study aimed to investigate the association of CTLA-4 gene polymorphism in childhood T1DM in the Indonesian population using PCR-RFLP.

The title and abstract cover the main aspect of the work but not completed

The abstract is incorrect in line 26 mention 31 subjects (including 12 males and 27 Javanese). Should be male/female? Please conclude the findings what the significant impact of CTLA-4 gene polymorphism study.

Author response:

Thank you for your suggestion, we already revised the manuscript. Please see on abstract section page 1-2 line 3-45. We also changes the tittle of the manuscript and our affiliation.

2. Does the introduction provide background and information relevant to the study?

The manuscript is technically sound. However, there are some points to be considered in the revision of the manuscript: 1)Classification of diabetes should be more updated than ref. 1-2.

2)The rationale of gene and SNP selection would be elaborated. 3)The cross section was chosen instead of case-control.

Author response:

- Thank you for your suggestion, we already add the new reference. Please see on introduction section page 2-3 line 53-54.
- Thank you for your suggestion, we already revised as your suggestion. We have added some paragraphs in the introduction section. Please see pages 3 lines 55-60 in the revised manuscript.
- Thank you, we have added the control sample in our study in our method, please see page 3 line 71 and on result section page 13 in lines 321-323 tables 1 and 2 of the revised manuscript.

4. Are the methods clear and replicable? Do all the results presented match the methods described?

1) The rationale of SNP selection would be elaborated.



2) Inclusion criteria of study participants especially T1DM should be more detail which guideline.

- 3) The rationale of selection just regularly visiting the hospital during the study not T1DM patients hospitalized in the pediatric intensive care should be explained.
- 4) Define the ethnic in section subject as in table 1. More define why divide age of subject as table 3 (0-4 years / 5-11 years

/12-18 years)

5) No statistic method is noted.

Author response:

- Thank you for your suggestion, we already revised our suggestion. We have added some paragraphs in the material and methods section. Please see page 4 line 85-89 in the genetic analysis subsection
- Thank you for your suggestion, we are revising as your suggestion. We have added some paragraphs in the study participant subsection. Please see page 3-4 line 72-82 in the revised manuscript.
- Thank you, we exclude children in the pediatric intensive care unit because those children need to be stabilized first
- Thank you for your suggestion, we have already deleted table 3 in the revised manuscript. We divide the children's age based on the WHO
- Thank you for your suggestion, we already added the statistical analysis subsection to the on in material and methods section. Please see pages 5 line 109-113 in the revised manuscript.
 - 5. If relevant are the results novel? Does the study provide an advance in the field? Is the data plausible?

The result is not so now

- 1) More consider for typo in table 2, Polymorphisme.
- 2) The analysis of polymorphism and clinical characteristic would be more interested.

Author response:

- Thank you for your suggestion, we already revised table 2 with the result of the case-control statistic. Please see page 13 line 323 in the revised manuscript.
- Thank you for your suggestion, we already revised the table 1 and 2 with the result of the case-control statistic.
 Please see page 13 in line 321-323 tables 1 and 2 in the revised manuscript.

6. Do the findings described by the author correlate with the results? Are the findings relevant?

1) It would be more interesting if the authors mention the genotype/haplotype comparing other populations especially in the Asian population or between ethnic.

2) Limitation of the study should be mentioned

Author response:

- 1. Thank you for your suggestion, we already revised the discussion, please see page 6 lines 139-150.
- Thank you for your suggestion, we already added the statement in the discussion section. Please see page 8 lines 196-200 in the revised manuscript.

7. Do the conclusions correlate to the results found?

Please conclude the findings what the significant impact of CTLA-4 gene polymorphism study.

Author response:

Thank you for your suggestion, we already revised the conclusion, please see page 8 lines 202-205.

Figures & Tables

8. If the author has provided figures and tables are the figures and tables clear and legible? Are the figures free from unnecessary modification?

1)The Figure legend should be modified please add more detail what is lane 1-8 and what method used in the experiment. 2)The figures of agarose gels are not good.

Author response:

- 1. Thank you, we already revised it as your suggestion. We also uploaded the figures in pdf format.
- 2. Thank you, we already revised the quality of the figure, please see the figure file 1 and 2



Here is the ORIGINAL RESEARCH from Rochman et al on entitle CTLA-4 Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus.

This study aimed to investigate the 2 CTLA-4 gene polymorphisms in childhood T1DM in the Indonesian population using PCR-RFLP. 1)No statistic method is noted. It is not the new knowledge, it will be great if mention especially ethnic group but the limitation of number of subjects were considered the power of test.

Author response:

- Thank you for your suggestion, we added the statistical analysis subsection the material and methods section
 and already changes the statistical analysis from cross-sectional to case-control. Please see page 3 line 72 and
 page 4 line 108—113 in the revised manuscript.
- 3. Do you think the manuscript requires English editing to correct the grammar or flow?

Yes

Author response:

Thank you for your response. This manuscript has been edited to ensure language and grammar accuracy by "editage". We also uploaded the certificate of editing.



Editing Certificate

This document certifies that the manuscript listed below has been edited to ensure language and grammar accuracy and is error free in these aspects. The logical presentation of ideas and the structure of the paper were also checked during the editing process. The edit was performed by professional editors at Editage, a division of Cactus Communications. The author's core research ideas were not altered in any way during the editing process. The quality of the edit has been guaranteed, with the assumption that our suggested changes have been accepted and the text has not been further altered without the knowledge of our editors.

MANUSCRIPT TITLE

CTLA-4 GENE POLYMORPHISMS IN INDONESIANS WITH TYPE 1 DIABETES MELLITUS

AUTHORS

Nur Rochmah

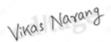
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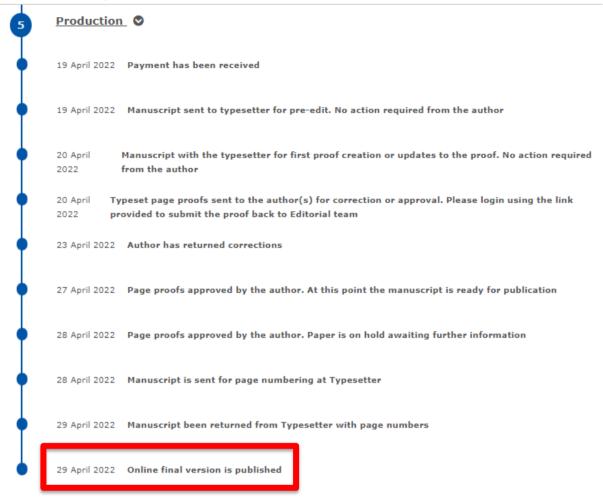
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4. BUKTI ACCEPTED, PUBLISH DAN PAYMENT



Dear Dr Rochmah

Thank you, we have received approval for your paper "CTLA-4 CT-60 A/G and CTLA-4 1822 C/T Gene Polymorphisms in Indonesians with Type 1 Diabetes Mellitus".

We will arrange to have the paper prepared for publication. An email will be sent to you when your paper has been published on our website.

Do contact me if you have any questions: Live Support https://www.dovepress.com/live_help.t Email: boonlee@dovepress.com Phone: +64-9-443 3060

Kind regards

Boon Lee
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