| No | Tahap Korespondensi Publikasi "Typhonium flagelliforme Lodd. Blume Leaf Extract Inhibits COX-2 Expression on WiDr Colon Cancer Cells" | Tanggal |
|----|---|------------------|
| 1. | Pembuatan akun ScholarOne Manuscript Asian Pacific journal | 25 Juni 2014 |
| | of Tropical Biomedicine | |
| 2. | Manuscript submission | 20 Oktober 2015 |
| 3 | Manuscript lolos review awal | 28 Oktober 2015 |
| 3. | Manuscript decision: Revision needed from two referees | 10 November 2015 |
| 4. | First revision of the manuscript | 16 November 2015 |
| 5. | Provisional Acceptance Letter with Editorial Revision | 23 November 2015 |
| 6 | Publishing Agreement | 26 November 2015 |
| 7 | Proofreading manuscript dari Editor | 29 November 2015 |
| 6. | Online manuscript published | 17 Februari 2016 |

1. Pembuatan akun ScholarOne Manuscript Asian Pacific journal of Tropical Biomedicine

Asian Pacific Journal of Tropical Biomedicine - Account Created in ScholarOne Manuscripts

 $on behalf of + apjtbb + yahoo.com@manuscriptcentral.com \\ < on behalf of + apjtbb + yahoo.com@manuscriptcentral.com \\ > on b$

apjtbb@yahoo.com <apjtbb@yahoo.com>

Wed 6/25/2014 4:53 AM

To:agustinasetiawati85@gmail.com <agustinasetiawati85@gmail.com>;Agustina Setiawati <nina@usd.ac.id>

Dear Mrs. setiawati:

Welcome to the Asian Pacific Journal of Tropical Biomedicine - ScholarOne Manuscripts site for online submission and review.

Your USER ID for your account at http://mc03.manuscriptcentral.com/apjtb is as follows:

USER ID: agustinasetiawati85@gmail.com

If you are unsure of the password that you set when you created your account you should click the link below which will take you directly to the option for setting a new password.

http://mc03.manuscriptcentral.com/apjtb?URL MASK=7c22146f36db43b1b7be28dc2e1e1f07

Thank you for your participation.

Sincerely,

Asian Pacific Journal of Tropical Biomedicine Editorial Office

2. Manuscript submission

Submission APJTB_2015_621 received by Asian Pacific Journal of Tropical Biomedicine





Evise <EviseSupport@elsevier.com>

 \odot



Tue 10/20/2015 3:40 PM

To: Agustina Setiawati

Dear Mrs. Setiawati,

Thank you for submitting your manuscript for consideration for publication in Asian Pacific Journal of Tropical Biomedicine. Your submission was received in good order.

To track the status of your manuscript, please log into EVISE® http://www.evise.com/evise/faces/pages/navigation/NavController.jspx?JRNL_ACR=APJTB and go to 'My Submissions'.

Thank you for submitting your work to this journal.

Kind regards,

Asian Pacific Journal of Tropical Biomedicine

This message was sent automatically. Please do not reply.

3. Manuscript lolos review awal dan butuh proofread.





Dear author,

Your paper has passed the first review, please revise your paper according to the following suggestions and highlight the revision with different color to facilitate our work:

1. Your paper should be read, corrected, and edited with an English native speaker or by someone who is familiar with the English language.

Please send back your revised paper before 5 Nov, so as to publish your paper timely.

Best wishes,

Asian Pacific Journal of Tropical Biomedicine

Asian Pacinic Solution of Implication in the Commission at http://mc03.manuscriptcentral.com/apitb

EVISE® Online submission at https://www.evise.com/evise/faces/pages/navigation/NavController.jspx?JRNL_ACR=APJTB

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http://www.sciencedirect.com/science/journals/22211691

www.apitb.com

Email:apjtbb@yahoo.com

+86-898-66973631, +86-898-36335723 Fax: +86-898-36335723

4. Manuscript decision: Revision needed from two referees.

From: Shunhai Qu <apitbb@yahoo.com>

Sent: Tuesday, November 10, 2015 2:57 PM

To: Agustina Setiawati

Subject: Asian Pacific Journal of Tropical Biomedicine (B5-280) Decision

Dear Agustina Setiawati,

Two referees have assessed your manuscript and have recommended substantial enhancements to be made before it can be considered suitable for publication. Please find the reviewers' comments attached for your reference.

If you have any questions or technical issues, please contact the journal's editorial office.

Yours sincerely,

Shunhai Qu Editor-in-Chief

Asian Pacific Journal of Tropical Biomedicine EVISE® Online submission at https://www.evise.com/evise/faces/pages/navigation/NavController.jspx?JRNL_ACR=APJTB

http://www.ncbi.nlm.nih.gov/pmc/journals/2020/ http://www.sciencedirect.com/science/journal/22211691

Email:apjtbb@yahoo.com Tel: +86-898-36335723 Fax: +86-898-36335723

#Reviewer 1

Reviewers' comments:

Authors have studied "Typhonium flagelliforme Lodd. Blume Leaf Extract Inhibits COX-2 Expression on WiDr Colon Cancer Cells" a study on. The following points are strongly

- 1. In the method section, extract preparation, what kind of extract was evaporated? It should be clear.
- 2. I think in immunocytochemistry method assay, the cells must be washed after first antibody incubation.
- 3. How do you quantify COX-2 expressing cells?

#Reviewer 2

Reviewer #2: The manuscript of Setiawati et al. reports "Typhonium flagelliforme Lodd. Blume Leaf Extract Inhibits COX-2 Expression on WiDr Colon Cancer Cells" The results are novel and interesting; however, they need to be improved. It is necessary to have a minor revision.

- 1. In the result section, the first paragraph must be revised.
- 2. The double staining method must be clarifying the difference between apoptotic and necrotic cells.
- 3. The number of cells expressing COX-2 must be notified by statistical analysis different.
- 4. The content found in the initial paragraph of the 'Discussion' section appears more fitting for an introduction rather than a discussion of the results. The author's presentation serves as an introduction to the topic rather than an in-depth analysis of the obtained results.
- 5. The conclusion needs to be improved.

5. First revision of the manuscript

From: Agustina Setiawati <nina@usd.ac.id>
Sent: Monday, November 16, 2015 5:01 PM
To: Shunhai Qu <apjtbb@yahoo.com>

Subject: Re: Asian Pacific Journal of Tropical Biomedicine (B5-280) Decision



Dear Editor in-Chief,

I am writing to you to submit our revised manuscript encoded B5-280.

Please find the following attachment in this email. Here is our responses to Reviewer's comment:

eviewei #1

1. In the method section, extract preparation, what kind of extract was evaporated? It should be clear.

Response:

We appreciated your comment. The evaporated extract was liquid ethanolic extract. Therefore, we revised our manuscript as mentioned below:

"The liquid extract was slowly evaporated to discard the residual solvent, until viscous extract was obtained".

2. I think in immunocytochemistry method assay, the cells must be washed after first antibody incubation.

Response:

Thank you for your critical comment. We washed our fixed cells after antibody incubation. Hence, we revised our manuscript:

"The cells were washed three times with PBS and added with secondary antibody, incubated at room temperature for ten minutes, and washed four times with PBS"

3. How do you quantify COX-2 expressing cells?

Responses:

We mentioned the quantification method of COX-2 expressing cells in data analysis.

Reviewer#2

1. In the result section, the first paragraph must be revised.

Responses

We revised our paragraph accordingly.

2. The double staining method must be clarifying the difference between early and late apoptotic cells.

Responses

Thank you for your critical comment. In early apoptotic cells, the nuclei showed condensed yellow-green fluorescence by acridine orange. On the other hand, the nuclei of late apoptotic cells showed condensed in orange fluorescence by ethidium bromide.

We mentioned it in the discussion section.

3. The content found in the initial paragraph of the 'Discussion' section appears more fitting for an introduction rather than a discussion of the results. The author's presentation serves as an introduction to the topic rather than an in-depth analysis of the obtained results.

Responses:

We revised our paragraph accordingly.

4. The conclusion needs to be improved.

Responses

We revised our conclusion accordingly.

I am looking forward to hearing from you.

Best regards, Agustina



Faculty of Pharmacy, Sanata Dharma University

Paingan, Maguwoharjo, Depok, Sleman, Yogyakarta, Indonesia 55281

November 16, 2015

Dear Prof Shunhai Qu Editor in Chief Asian Pacific Journal of Tropical Biomedicine

We hereby resubmit our manuscript, entitled "Typhonium flagelliforme Lodd. Blume Leaf Extract Inhibits COX-2 Expression on WiDr Colon Cancer Cells" (Research Article) to the Asian Pacific Journal of Tropical Biomedicine that you handled as the responsible editor in a prior submission.

We were delighted that our work is accepted to be published in this journal. We have taken editor comments seriously and have responded to all recommendations, as detailed below. We appreciate your response to our original submission and we hope that the revisions made to the manuscript raise it to the high standard expected of submissions to Asian Pacific Journal of Tropical Biomedicine

I will continue to serve as the corresponding author for this manuscript. Please do not hesitate to contact me if you have further questions or concerns.

Yours sincerely,

Agustina Setiawati, Assistant Professor

Faculty of Pharmacy, Sanata Dharma University, Paingan,

Maguwoharjo, Depok, Sleman, Yogyakarta, Indonesia 55281

Reviewer #1

| Comments | Responses |
|---|--|
| In the method section, extract preparation, what kind of extract was evaporated? It should be clear. | We appreciated your comment. The evaporated extract was liquid ethanolic extract. Therefore, we revised our manuscript as mentioned below: "The liquid extract was slowly evaporated to discard the residual solvent, until viscous extract was obtained". |
| I think in immunocytochemistry method assay, the cells must be washed after first antibody incubation | Thank you for your critical comment. We washed our fixed cells after antibody incubation. Without washing, the result will reveal lots of noise and unclear between false positive or noise signal of the colour changing. Hence, we revised our manuscript: "The cells were washed three times with PBS and added with secondary antibody, incubated at room temperature for ten minutes, and washed four times with PBS" |
| How do you quantify COX-2 expressing cells? | Thank you for confirming the quantification method for COX-2. In our study we adopt the previous study by MIze (1994) by quantifying brown colour cells under the light microscope. The number of COX-2 expressing cells was then analyzed by t-test using Microsoft Excel 2013. Reference: R.R. Mize. 1994. Quantitative image analysis for immunocytochemistry and in situ hybridization. https://doi.org/10.1016/0165-0270(94)90195-3. |

Reviewer #2

| In the result section, the first paragraph must be revised. Responses: | We appreciated your comment. We revised our paragraph accordingly. |
|--|--|
| | |

| The double staining method must be clarifying the difference between early and late apoptotic cells. | Thank you for your critical comment. Our study employed dual AO/EB fluorescent staining which can detect basic morphological changes in apoptotic cells. In addition, it allows for the distinction between normal cells, early and late apoptotic cells, and necrotic cells. In early apoptotic cells, the nuclei showed condensed yellow-green fluorescence by acridine orange. On the other hand, the nuclei of late apoptotic cells showed condensed in orange fluorescence by ethidium bromide. We mentioned it in the discussion section. Reference: Liu et al. 2015. Dual AO/EB Staining to Detect Apoptosis in Osteosarcoma Cells Compared with Flow Cytometry. Med Sci Monit Basic Res. 2015; 21: 15–20. |
|---|--|
| The content found in the initial paragraph of the 'Discussion' section appears more fitting for an introduction rather than a discussion of the results. The author's presentation serves as an introduction to the topic rather than an in-depth analysis of the obtained results. | We revised our paragraph accordingly |
| The conclusion needs to be improved. | We revised our conclusion accordingly. |

Provisional Acceptance Letter with Editorial Revision & Publishing Agreement

From: Shunhai Qu <apjtbb@yahoo.com>

Sent: Monday, November 23, 2015 7:53 PM

To: Agustina Setiawati

Subject: Asian Pacific Journal of Tropical Biomedicine (B5-280)--Acceptance Letter & Publishing agreement

Dear Dr. Agustina Setiawati

After the preliminary review, your article "Typhonium flagelliforme Lodd. Blume Leaf Extract Inhibits COX-2 Expression on WiDr Colon Cancer Cells" (B5-280) was agreed to be published. Here attached the publishing agreement and the acceptance letter. Please reply us within 5 days.

Please revise your paper according to the following suggestions and highlight the revision with different color to facilitate our work:

- 1. The language still needs to be improved. Your paper should be read, corrected, and edited with an English native speaker or by someone who is familiar with the English language.
- 2. Please add more content to the part of Results.
- 3. Please provide Foundation Name and Grant No. (Don't forget to provide the Grant No.).
- 4. Update the references and make sure half of the references are in recent 3 years!!!!! (2012-2015), another one-fourth references are in recent 5 years (i.e. 2010-2015). Old references are better replaced with updated ones. References must meet these requirements!!!! Citations should be labeled with square bracket and should appear in front of the period or comma at the end of the sentence/clause. All references should be cited. It is better to include citation from Pubmed, the more the better.

You could search the key words on PMC et al. index and add new references to your paper.

- 5. Please provide the URL for reference no. 2 and 5 for validation.
- References should be standard.

Examples:

Journal:

[1] Kihara K, Sato K, Ando M, Oshima H. Ability of lumbar splanchnic nerves and inability of thoracic nerves to generate seminal emission in dogs. J Urol 1992; 147: 260-3. Book:

[1] Armitage P. Statistical methods in medical research. Oxford: Blackwell; 1971, p.102-11.

Chapter in a book:

- [1] Mettam GR, Adams LB. How to prepare an electronic version of your article. In: Jones BS, Smith RZ.(eds.) Introduction to the electronic age. New York: E-Publishing Inc; 1999, p. 281-
- [2] Brinton L, Lacey J, Devesa SS. Epidemiology of breast cancer. In: Donegan WL, Spratt JS. (eds.) Cancer of the Breast. 5th ed. Lodon: Saunders Elsevier Science; 2002, p. 111-32.
- [3] In the references, the author's first name should be abbreviated with the first letter uppercase, and the family name is remained and should be in front of abbreviation of the first name.
- [4] World Health Organization. Malaria elimination. A Field manual for Low and Moderate Endemic Countries. Geneva: World Health Organization; 2007. [Online] Available from: http://www.who.int/malaria/docs/elimination/MalariaElimination_BD.pdf. [Accessed on 27 December, 2011].

Wishes,

Lynn

Asian Pacific Journal of Tropical Biomedicine

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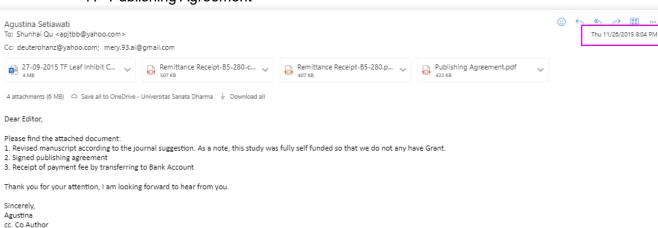
http://www.ncbi.nlm.nih.gov/pmc/journals/2020/

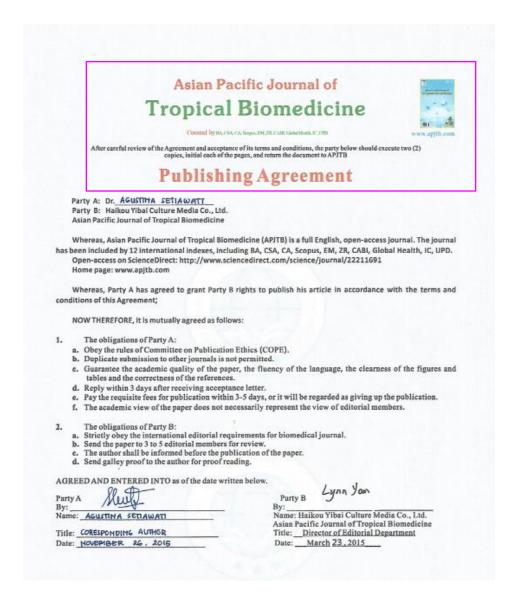
http://www.sciencedirect.com/science/journal/22211691

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7. Publishing Agreement





8. Manuscript Proofreading before Published



9. Manuscript was Online Published

