[IJPPS] Your manuscript IJPPS 9938: Revise and resubmit

From: Editor IJPPS (editor@ijpps.innovareacademics.in)

To: lanhadi@yahoo.com

Date: Saturday, December 5, 2015 at 02:38 PM GMT+7

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938.

Article is quite interesting but it needs major revision Comments-

For revision of your article see the following points

- Format-: Revise article to make it strictly as per format of the Journal. Refer latest issue of the Journal for formatting. Headings and subheading should not be numbered.
- Symbol and units: It should be as per International System of Units (SI). See it in instructions to authors and follow accordingly and strictly.
- Errors: Grammatical and punctuation errors should be rectified. Authors are suggested to use smart tools like 1 checker, ginger, grammarly, white smoke, etc.
- · Headings of figure (s) should be rectified. See latest issue of IJPPS.
- Table: Column headings should be in sentence case and bold. See latest issue of IJPPS.
- Fig: Ensure that titles at x and y axis are in sentence case and bold.
- Ensure that biological names (plants/crude drugs/bacteria/fungus etc) are in italic in whole manuscript including reference also.
- · Sample size and proper foot notes should be mentioned below table(s).
- Discussion could not be found. Only results have been given. Authors should make comparison with previously reported such works to emphasize importance of the presented work.
- References: References are out of format. Uniformity must be ensured in all the references. It should be made strictly as per Instructions to Authors. Journal's title should be non italic, abbreviated without use of full stop.
- Pagination style is incorrect in references. Authors should refer any latest published article in IJPPS. Digit appeared in starting page number should not be repeated in end page number. Ex. 12-5, 25-32, 125-7, 11456-62 etc.
- References: There is no need to mention issue number with volume. It should be provided only if it is supplement issue. Please correct accordingly.

[Few examples of references from journal:

Devi KV, Pai RS. Antiretrovirals: Need for an Effective Drug Delivery. Indian J Pharm Sci 2006;68:1-6. List the first six contributors followed by et al.

Volume with supplement: Shen HM, Zhang QF. Risk assessment of nickel carcinogenicity and occupational lung cancer. Environ Health Perspect 1994;102 Suppl 1:275-82.

Issue with supplement: Payne DK, Sullivan MD, Massie MJ. Women's psychological reactions to breast cancer. Semin Oncol 1996;23(1, Suppl 2):89-97.]

See attachment for more comments and queries.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments:

- 1. Authors are requested to make revision point to point and very strictly. Failure may cause its rejection.
- 2. Authors must give their response to the comments of reviewers at end of the revised copy of manuscript. If authors disagree with any comment they should record response with reason.

Note: Authors must send email to editor@ijppsjournal.com, after submission of revised article compulsorily with subject- "Revised article submitted for Round 2" along with article reference no.

After corrections submit your revised article as follows

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With Regards, Editor IJPPS

editor@ijpps.innovareacademics.in

International Journal of Pharmacy and Pharmaceutical Sciences http://innovareacademics.in/journals/index.php/ijpps



9938-30431-1-SM.docx

343.9kB

REVISED ARTICLE FOR SUBMITTED ROUND 2 - 9938

From: lannie hadi (lanhadi@yahoo.com)

To: editor@ijppsjournal.com

Date: Monday, December 14, 2015 at 12:54 AM GMT+7

Dear The Editor of IJPPS

I have sent a revised article submitted for round 2.

My response to comments:

- 1. The discussion has been included in the part of RESULTS AND DISCUSSION.
- 2. Many grammatically mistake, so I don't give the red color for my revise article.

Thank you for your kind cooperation.

Best Regards, Lannie Hadisoewignyo

[IJPPS] Your manuscript IJPPS 9938: Revise and resubmit

From: Manager (editor@ijpps.innovareacademics.in)

To: lanhadi@yahoo.com

Date: Monday, December 14, 2015 at 05:18 PM GMT+7

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938.

Revision required

Comments-

For further revision of your article see the following points

- Sections Determination of total flavonoid content in the extract and Preparation of tablet formulation development of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts: Rewrite both of it.
 Grmaatically incorret and should be written in past tense third person only.
 Further language should be moodified so it can be unserstood.
- Sample size and proper foot notes should be mentioned below table(s).
- Discussion could not be found. Only results have been given. Authors should make comparison with previously reported such works to emphasize importance of the presented work.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments:

- 1. Authors are requested to make revision point to point and very strictly. Failure may cause its rejection.
- 2. Authors must give their response to the comments of reviewers at end of the revised copy of manuscript. If authors disagree with any comment they should record response with reason.

Note: Authors must send email to editor@ijppsjournal.com, after submission of revised article compulsorily with subject- "Revised article submitted for Round 3" along with article reference no.

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Impact (Cites per doc): 0.55 (SCImago, 2014).

With Regards, Editor IJPPS

editor@ijppsjournal.com

International Journal of Pharmacy and Pharmaceutical Sciences

http://innovareacademics.in/journals/index.php/ijpps

Re: [IJPPS] Submission Acknowledgement: IJPPS 9938

From: lannie hadi (lanhadi@yahoo.com)

To: editor@ijpps.innovareacademics.in

Date: Friday, December 18, 2015 at 11:29 AM GMT+7

Dear The Editor of LIPPS

Thank you for accepting my manuscript, but there are erratum in the manuscript as on a file that I send in this email

There is the addition of the author in the manuscript.

Thank you for your cooperation.

Best regards,

Dr. Lannie Hadisoewignyo, M.Si., Apt. Vice Dean I - Faculty of Pharmacy Widya Mandala Catholic University Surabaya Jl. Raya Kalisari Selatan No. 1, Surabaya - 60112 office phone: +6231 99005291; ext. 10603 fax. +6231 99005288

mobile phone: +6287852942829

On Wednesday, November 18, 2015 3:08 PM, Manager <editor@ijpps.innovareacademics.in> wrote:

Dear Lannie Hadisoewignyo,

Thank you for submitting the manuscript, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN

ANTI-INFLAMMATORY" to International Journal of Pharmacy and Pharmaceutical Sciences.

Manuscript Reference No.: IJPPS 9938

Manuscript URL:

http://innovareacademics.in/journals/index.php/ijpps/author/submission/9938

Username: lanhadi

Decision of manuscript will be communicated with in 15-20 days.

Progress of your submission can be tracked on your account with IJPPS.

If decision is not sent in time frame to you, please remind us with reference no of the manuscript.

Impact: 0.99 (SCImago, 2013)

Important Announcement:

It is my great pleasure to invite you to Innopharm1, 1st international conference on novel frontiers in pharmaceutical and health sciences, which will be held from Oct 10 to Oct 11, 2015 at Bhopal, MP, India. We aim to make this conference one of the most acclaimed meetings in the field of pharmaceutical and health sciences. The meeting is being organized under the auspices of Innovare Academic Sciences in association with International Journal of Pharmacy and Pharmaceutical Sciences as publishing partner.

For more details and registration to submit abstract visit http://innopharm1.innovareacademics.in/

Its extreme pleasure to announce the launching of INNOVARE ACADEMIC SCIENCES (http://www.innovareacademics.in) – Serve to provide quality publication.

Success of International Journal of Pharmacy & Pharmaceutical Sciences and Asian Journal of Pharmaceutical & Clinical Research lead to launch INNOVARE ACADEMIC SCIENCES (http://www.innovareacademics.in) to provide platform for quality publication in various others disciplines such as Medicine, Engineering, Agriculture, Health, Ayurvedic, Education, Social, Business Management, Food, and Life sciences.

Authors/readers interested to be part of editorial team, visit at http://www.innovareacademics.in/Career.php

On facebook, you can join us at http://www.facebook.com/InnovareAcademicSciences https://www.facebook.com/ijpps

Our team sincerely conveys kind regards to authors and editorial members to achieve the impact. You are requested to cite articles which are published in IJPPS in other publications also which will help us to increase the impact factor.

We are happy to inform our authors that IJPPS is now indexed in Scopus, Elsevier, EBSCO, Google Scholar and Directory of Open Access Journal (DOAJ), Index Copernicus, ICAAP, Scientific commons, PSOAR, Open-J-Gate in very short span. Journal has made its presence amongst International community with great impact ICV- 5

For detail visit www.ijppsjournal.com

If you have any questions, please contact us at editor@ijppsjournal.com.

Thanks for your interest in IJPPS.

Manager

International Journal of Pharmacy and Pharmaceutical Sciences International Journal of Pharmacy and Pharmaceutical Sciences http://innovareacademics.in/journals/index.php/ijpps

revised article submitted for round 3

From: lannie hadi (lanhadi@yahoo.com)

To: editor@ijppsjournal.com

Date: Thursday, December 24, 2015 at 11:00 AM GMT+7

Dear The Editor of IJPPS

I have sent a revised article submitted for round 3.

Thank you for your kind cooperation.

Best regards, Dr. Lannie Hadisoewignyo, M.Si., Apt. Vice Dean I - Faculty of Pharmacy Widya Mandala Catholic University Surabaya Jl. Raya Kalisari Selatan No. 1, Surabaya - 60112 office phone: +6231 99005291; ext. 10603 fax. +6231 99005288 mobile phone: +6287852942829

Revised Article

From: lanhadi@yahoo.com (lanhadi@yahoo.com)

To: editor@ijppsjournal.com

Date: Tuesday, January 19, 2016 at 09:19 AM GMT+7

Dear The Editor of IJPPS

I have sent a revised article submitted for round 3 (24 Dec 2015).

Thank you for your kind cooperation.

Sent from my ASUS

Best regards, Dr. Lannie Hadisoewignyo, M.Si., Apt. Vice Dean Faculty of Pharmacy Widya Mandala Surabaya Catholic University Jl. Raya Kalisari Selatan No. 1, Surabaya-60211 Phone. 62-31-99005299 (10603) Hp. 62-87852942829

------- Original Message -------From:Manager <<u>editor@ijpps.innovareacademics.in</u>>
Sent:Mon, 14 Dec 2015 17:18:05 +0700
To:Lannie Hadisoewignyo <<u>lanhadi@yahoo.com</u>>
Subject:[IJPPS] Your manuscript IJPPS 9938: Revise and resubmit

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938.

Revision required

Comments-

For further revision of your article see the following points

- Sections Determination of total flavonoid content in the extract and Preparation of tablet formulation development of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts: Rewrite both of it.
 Grmaatically incorret and should be written in past tense third person only.
 Further language should be moodified so it can be unserstood.
- Sample size and proper foot notes should be mentioned below table(s).
- Discussion could not be found. Only results have been given. Authors should make comparison with previously reported such works to emphasize importance of the presented work.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments:

- 1. Authors are requested to make revision point to point and very strictly. Failure may cause its rejection.
- 2. Authors must give their response to the comments of reviewers at end of the revised copy of manuscript. If authors disagree with any comment they should record response with reason.

Note: Authors must send email to editor@ijppsjournal.com, after submission

about:blank

1/2

of revised article compulsorily with subject- "Revised article submitted for Round 3" along with article reference no.

After corrections submit your revised article as follows

- Log ir
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[IJPPS] Your manuscript IJPPS 9938: Revise and resubmit

From: Editor (editor@ijppsjournal.com)

To: lanhadi@yahoo.com

Date: Thursday, January 21, 2016 at 03:02 PM GMT+7

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938.

Revision required

Comments-

For further revision of your article see the following points

- Heading Determination of total flavonoid content in the extract should be corrected as it gives methods for the ethanlic exract and water fraction also.
- Preparation of tablet formulation development of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts: Rewrite it. Still grammatically incorrect and confusing. Language polishing is must to understand it.
- Tables and figure footnotes: Authors must mention values of n, i.e. number of experiments, for example, 3, 4, 5 or 10 etc.
- Tables and figure footnotes: Please mention that data given in mean±SD or mean±SEM in footnotes of each table/figure.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments:

- 1. Authors are requested to make revision point to point and very strictly. Failure may cause its rejection.
- 2. Authors must give their response to the comments of reviewers at end of the revised copy of manuscript. If authors disagree with any comment they should record response with reason.

Note: Authors must send email to editor@ijppsjournal.com, after submission of revised article compulsorily with subject- "Revised article submitted for Round 4" along with article reference no.

After corrections submit your revised article as follows

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With Regards,

Editor IJPPS

editor@ijpps.innovareacademics.in
International Journal of Pharmacy and Pharmaceutical Sciences
http://innovareacademics.in/journals/index.php/ijpps

revised article submitted for round 4

From: lannie hadi (lanhadi@yahoo.com)

To: editor@ijppsjournal.com

Date: Wednesday, January 27, 2016 at 09:25 AM GMT+7

Dear The Editor of IJPPS

I have sent a revised article submitted for round 4 (27 Jan 2016).

Thank you for your kind cooperation.

Best regards,,

Dr. Lannie Hadisoewignyo, M.Si., Apt. Vice Dean I - Faculty of Pharmacy Widya Mandala Catholic University Surabaya Jl. Raya Kalisari Selatan No. 1, Surabaya - 60112 office phone: +6231 99005291; ext. 10603

fax. +6231 99005288

mobile phone: +6287852942829



9938-30431-1-SM-REVISED ARTICLE SUBMITTED FOR ROUND 4.docx

[IJPPS] Your manuscript IJPPS 9938: Revise and resubmit

From: Editor (editor@ijppsjournal.com)

lanhadi@yahoo.com To:

Date: Wednesday, January 27, 2016 at 02:51 PM GMT+7

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences. "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938.

Revision required

Comments-

For further revision of your article see the following points

Preparation of tablet formulation development of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts: Rewrite it. Still grammatically incorrect and confusing. Language polishing is must to understand it.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments:

- 1. Authors are requested to make revision point to point and very strictly. Failure may cause its rejection.
- 2. Authors must give their response to the comments of reviewers at end of the revised copy of manuscript. If authors disagree with any comment they should record response with reason.

Note: Authors must send email to editor@ijppsjournal.com, after submission of revised article compulsorily with subject- "Revised article submitted for Round 5" along with article reference no.

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With Regards, **Editor IJPPS**

editor@ijpps.innovareacademics.in

International Journal of Pharmacy and Pharmaceutical Sciences

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revised article submitted for round 5

From: lannie hadi (lanhadi@yahoo.com)

To: editor@ijppsjournal.com

Date: Wednesday, January 27, 2016 at 04:53 PM GMT+7

Dear The Editor of IJPPS

I have sent a revised article submitted for round 5 (27 Jan 2016).

Thank you for your kind cooperation.

Best regards,

Dr. Lannie Hadisoewignyo, M.Si., Apt. Vice Dean I - Faculty of Pharmacy Widya Mandala Catholic University Surabaya Jl. Raya Kalisari Selatan No. 1, Surabaya - 60112 office phone: +6231 99005291; ext. 10603

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mobile phone: +6287852942829



9938-30431-1-SM-REVISED ARTICLE SUBMITTED FOR ROUND 5.docx

Fwd: [IJPPS] Your manuscript IJPPS9938: Decision

From: lanhadi@yahoo.com (lanhadi@yahoo.com)

To: raymond@dexa-medica.com; raytjan@yahoo.com

Date: Friday, January 29, 2016 at 01:53 PM GMT+7

Sent from my ASUS

------ Original Message ------From:Editor <<u>editor@ijppsjournal.com</u>>
Sent:Fri, 29 Jan 2016 13:04:35 +0700
To:Lannie Hadisoewignyo <<u>lanhadi@yahoo.com</u>>
Subject:[IJPPS] Your manuscript IJPPS9938: Decision

Dear Lannie Hadisoewignyo,

We have reached a decision regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" with reference no. IJPPS 9938

Your manuscript has been recommended (Provisionally Accepted) for publication after peer review.

Acceptance will be sent on receipt of the registration fee and adequate revision (Revised article).

For publication, all the authors are required to send registration charges (US\$ 50.00 per author). You are requested to inform regarding your acceptance for registration within 3-4 days.

For revision of your article see the following points

· Fig should be written as Fig.

(All the changes made must be highlighted with RED coloured fonts or it should be done in track change mode).

Response to comments: Authors should give their response to the comments of reviewers at end of the revised copy of the manuscript. If authors disagree with any comment, they should record response with reason.

(All the changes made must be highlighted with RED coloured fonts. After corrections submit your revised article as follows

- 1. Log in
- 2. Click on Active
- In section- Click on provisionally accepted- New page
- 4. Scroll down- See Editor Decision
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You can transfer the required amount [US\$ 479 {including US\$ 20.0 for bank charges}+ all other charges] online to following account.

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International Journal of Pharmacy and Pharmaceutical Sciences C/o- Dr Avijeet Jain Associate Editor 6/1, Sanjeevani Medical Stores Shastri Chowk, Sadar Bazar, Sagar (Saugor), MP PIN-472002 India

With Regards,
Editor IJPPS
editor@ijpps.innovareacademics.in
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[IJPPS] Views and downloads of your publication

From: Editor (member@ijcpr.org)

To: lanhadi@yahoo.com

Date: Tuesday, February 2, 2016 at 12:33 PM GMT+7

Dear,

Delight to announce downloads/views of your manuscript published in International Journal of Pharmacy and Pharmaceutical Sciences. Your quality contribution in the IJPPS has given it new heights in the field of research publication and helps to make IJPPS indexed in leading abstracting and indexing agencies like Scopus, Elsevier, Ebsco, Embase and so on. IJPPS and team personally convey vote of thanks to you for your constant support and contribution. IJPPS would always be happy to serve you and demand to keep on your support and share with your colleagues/students about quality publication in our journal, that could be helpful for their research work and publication.

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Thanks once again for your precious time and publication of research outcomes in IJPPS and trust IJPPS will flourish with you and your research community/colleagues/students.

International Journal of Pharmacy and Pharmaceutical Sciences (ISSN- 0975-1491) Rapid Peer review: 15-20 days

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Scopus Cites per doc, SJR (2014)0.55, SCImago www.scimagojr.com and see option Journal Search

For any query, feel free to contact at editor@ijppsjournal.com www.ijppsjournal.com

For Table of content-Click here

With Best Wishes

Editorial Team

[IJPPS] Your manuscript IJPPS 9938: Acceptance

From: Editor (editor@ijppsjournal.com)

To: lanhadi@yahoo.com

Date: Tuesday, February 2, 2016 at 12:52 PM GMT+7

Dear Lannie Hadisoewignyo,

I am happy to inform you regarding your submission to International Journal of Pharmacy and Pharmaceutical Sciences, "FORMULA HERBAL TABLET CONTAINING ETHANOL EXTRACT OF SALAM LEAVES AND ETHANOL EXTRACT OF SAMBILOTO HERBS AS AN ANTI-INFLAMMATORY" that it has been recommended for publication after peer review.

I acknowledge you receipt of registration fee by NEFT for IJPPS 9938.

Your article is now accepted for publication and your article is scheduled to be published in Vol 8 Issue 3, Mar 2016.

All the author(s)/coauthor(s) are also requested to like our facebook page to get latest notification about latest issues, conferences, and other happenings. (https://fb.me/InnovareAcademicSciences) and (https://www.facebook.com/ijpps)

Impact: 0.55 (SCImago, 2014)

Its extreme pleasure to announce the launching of INNOVARE ACADEMIC SCIENCES (http://www.innovareacademics.in) – Serve to provide quality publication.

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Success of International Journal of Pharmacy & Pharmaceutical Sciences and Asian Journal of Pharmaceutical & Clinical Research lead to launch INNOVARE ACADEMIC SCIENCES (http://www.innovareacademics.in) to provide platform for quality publication in various others disciplines such as Medicine, Engineering, Agriculture, Health, Ayurvedic, Education, Social, Business Management, Food, and Life sciences.

We are happy to inform our authors that IJPPS is now indexed in Scopus, Elsevier.

EBSCO, Google Scholar and Directory of Open Access Journal (DOAJ), Index Copernicus, ICAAP, Scientific commons, PSOAR, Open-J-Gate in very short span. Journal has made its presence amongst International community with great impact 0.55 (SCImago, SJR 2014).

Our team sincerely conveys kind regards to authors and editorial members to achieve the impact factor. You are requested to cite articles which are published in IJPPS in other publications also which will help us to increase its impact.

For detail visit www.ijppsjournal.com

Editor IJPPS

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Galley proof IJPPS 99-38/2016- For any correction- Time duration 72 h only.

From: Galleyproof IJPPS (galleyproof@ijppsjournal.com)

To: lanhadi@yahoo.com

Date: Friday, February 19, 2016 at 12:06 PM GMT+7

Dear Sir/Madam

Your article is scheduled to be published in Vol. 8, Issue 3, March 2016- First week.

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Original Article

FORMULATION DEVELOPMENT AND OPTIMIZATION OF TABLET CONTAINING COMBINATION OF SALAM (SYZYGIUM POLYANTHUM) AND SAMBILOTO (ANDROGRAPHIS PANICULATA) ETHANOLIC EXTRACTS

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ABSTRACT

Objective: The present study was aimed at finding the appropriate type of a binder and a disintegrant which can be used formulate tablets containing a combination of two ethanol extracts of *Syzygium polyanthum* leaves and *Andrographis paniculata* herbs, as well as to find their optimum concentrations which would produce the tablets with desired hardness and disintegration time.

Methods: Tablet manufacturing was done in two methods with the use of PVP K30 as a binder using the wet granulation method, and gelatin as a binder using the direct compression method. The design optimization used was the 2² full factorial design with two factors and two levels, with the factors used being the binder concentration and the disintegrant.

Results: Tablets produced using gelatin as a binder had a hardness level similar to those using PVP K30 as a binder. However, their disintegration time was strongly affected by the choice of the binder. The result suggested that different binder (PVP K30 and gelatin) affected different disintegration time despite producing tablet hardness values that were almost identical.

Conclusion: The herbal tablet development of combined ethanolic extracts of *Syzygium polyanthum* and *Andrographis paniculata* herbs could be made using gelatin as a binder and crospovidone as a disintegrating agent using the direct compression method. The optimal formula was be obtained using gelatin 3.61% and crospovidone 3.45%, which had theoretical results in the hardness of 6.50 kp, the friability of 0.35%, and the disintegration time of 12.76 min.

Keywords: Syzygium polyanthum, Andrographis paniculata, Gelatin, Crospovidone, Optimization

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INTRODUCTION

The Indonesian-origined Salam leaves (*Syzygium polyanthum* folium) contain flavonoids, tannins, essential oils including sitral and eugenol, saponins, alkaloids, flavonoids, and polyphenols [1]. It has been characterized as having a function as an anti-inflammatory, antioxidant, antibacterial, and antifungal agent. Another plant that can be used in the therapy of inflammatory condition is the herb Sambiloto (*Andrographis paniculata* herbs). *Andrographis paniculata* contains resin, kalmeghin, andrographolide, diterpene, neo andrographolide, deoxy andrographolide apigenin, flavonoids and phenols with the bitter taste. A previous study [2] suggested that the optimum dose for a 10:1 combination of water fractions of *Syzygium polyanthum* and *Andrographis paniculata*, respectively, as an anti-inflammatory agent is 250 mg. At this dose, the combination gave a better anti-inflammatory effect (45.68%) than that of ibuprofen at a dose of 36 mg/kg BB (40.63%). The ethanolic extract combination was selected as the active ingredients in this research with doses determined based on comparisons between total flavonoid levels of ethanol extracts and the water fractions.

Polyvinyl pyrrolidone (PVP K-30) is a binder which can be used in a dry state or in a solution [3], and has a high binding strength which is suitable for wet granulation method and direct compression. Gelatin is a natural binder polymer widely used in the tablet formulation process. Gelatin is generally dissolved beforehand in a solvent to be used in a wet granulation process, but gelatin can also be used in the form of good solid powder and granule for direct compression method, with a concentration of 0.1%-99.9% of tablet weight. Gelatin has been used for a variety of uses such as binders, fillers, even as disintegrants. The usual gelatin concentration user for the binder is approximately 5% [4]. The metabolism of gelatin in the body is assisted by the enzyme protease. This enzyme also plays a role in the disintegration of tablets [5].

Crospovidone is one of the super disintegrants which does not form a complex with the active ingredient because it is a nonionic polymer. The disintegration of the tablet by crospovidone is done through a combination mechanism of wicking, swelling, and deformation. Crospovidone rapidly expands when it is in contact with the water without forming a gel, and it has a higher compression and disintegrating activity which is not affected by the pH of the medium [6]. Starch is a polysaccharide found in plants. Starch has many functions. One pharmaceutical use of starch is for tablet disintegration. In this study corn starch was used. It has the property of a white powder, tasteless, odorless and insoluble in water. Starch undergoes plastic deformation during compression, but this property depends on the size, size distribution, and particle shape. Starch is composed by two kinds of main components of the polymer of D-glucopyranose is 15-30% amylose (linear polymers), 70-85% amylopectin (branched polymer), and 5-10% of other material (protein and fat) [7]. Starch can serve as a disintegrating agent on the tablet by increasing the amount of water that goes into a tablet since starch powder has pores that cause capillary action. An increase in the amount of water that goes into the tablet causes the tablet to disintegrate into granules. The granules will then disintegrate into powders which will dissolve in bodily fluid and give their pharmacological effects [8].

The aim of this research was therefore to find an optimal concentration and specification of a binder and a disintegrant which were suitable for making tablet dosage form containing *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extracts.

MATERIALS AND METHODS

Materials

The Syzygium polyanthum and Andrographis paniculata ethanolic extracts were obtained from PT. Javaplant, Solo. Chemicals used in

this research were purchased as follows: PVP K-30 and crospovidone were from PT. BASF Indonesia), lactose monohydrate was from Leprino Foods, US. Talc and magnesium stearate were sourced from Peter Greven, Venlo, the Netherlands). AlCl₃, FeCl₃, hydrochloric acid, anhydrous acetic acid, sulfuric acid, chloroform, mayer reagents, dragendroff reagent, quercetin were purchased from Sigma, Switzerland. Ethanol 96% distilled water were sourced locally from Surabaya, Indonesia.

Standardization of dry ethanolic extracts

Standardization of dry ethanol extracts was done with the determination of the organoleptic, determination of total ash, determination of water soluble ash, determination of acid insoluble ash, and lost on drying was performed according to standard parameters that fulfill the BPOM Indonesia and the Indonesian Materia Medika requirements.

Phytochemical screening

Phytochemical screening test for the *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extracts included were standardized alkaloids, steroids or triterpenoids, flavonoids, tannins, saponins, and glycosides amounts.

Determination of total flavonoid content in the ethanolic extracts and the water fraction

Determination of total flavonoid content in the *Syzygium polyanthum* leaves and *Andrographis paniculata* ethanolic extracts and also in the water fraction is performed by a modification of the method by Humadi and Istudor [9]. The standard solution was prepared by weighing 25 mg quercetin in 25 ml of volumetric flask and dissolved in 15 ml of ethanol 96% and volume was made up with ethanol 96% (1000 µg/ml). The stock solution was further diluted with ethanol 96% to obtain working solution in the range 6.5–100 µg/ml. 1 ml of 96% to obtain working solution was withdrawn and added a mixture of 3 ml of ethanol 96%, 0.2 ml of 10% AlCl3, 0.2 ml of 1M sodium acetate, and 5.6 ml of distilled water. The absorbance of the solution was measured at 440.8 nm after incubation for 30 min at room temperature.

The ethanolic extract or the water fraction of *Syzygium polyanthum* leaves-*Andrographis paniculata* 10: 1 (1 g) was weighed accurately and dissolved in 25 ml of ethanol 96%, then hydrolyzed with 4N HCl (1:1) for 30 min. After that, 25 ml of distilled water was added. The flavonoids were extracted by using ethyl acetate. The extract solution was evaporated to remove the solvent. The residue was dissolved in ethanol 96% to made up 25 ml volume. 1 ml of the solution was withdrawn and added a mixture of 3 ml of ethanol

 $96\%,\,0.2$ ml of 10% AlCl3, 0.2 ml of 1M sodium acetate, and 5.6 ml of distilled water. The flavonoid was evaluated by spectrophotometer uv-vis 440.8 nm after incubation for 30 min at room temperature.

Preparation of tablet formulation development of *Syzygium* polyanthum leaves and *Andrographis paniculata* ethanolic extracts

Tablet manufacturing was done in two methods, wet granulation, and direct compression. The tablets weight was fixed to 700 mg. Wet granulation method of tablet was made by mixing dry extract (1 tablet contains 450 mg with the comparison between *Syzygium polyanthum* leaves, and *Andrographis paniculata* are 10: 1), lactose monohydrate, disintegrating agent (crospovidone or corn starch) and PVP K-30 solution to form a damp coherent mass which was screened through a sieve no 16 and was dried at 50 °C. The compositions of the batches are shown in table 1 (F1-8).

Direct compression method was used to prepare tablet with gelatin as a binder. Dry extracts were taken and then mixed with Pharmatose 80M, gelatin, and cornstarch. Magnesium stearate and talc were mixed and followed by compression of the blend. The compositions of the batches are shown in table 1 (F9-16).

Physical properties evaluation of powder

The physical properties of powder were evaluated by their bulk density, tapped density, Carr's index, and Hausner ratio.

Physical properties evaluation of tablets

Tablet hardness test was done by taking 20 tablets at random, and then placed one by one in the clamp on the hardness tester. Tablet hardness values can be seen on the screen. The terms of good tablet hardness were 4-8 kp.

Twenty tablets were cleaned from dust, weighed (W_1) , and then put in friability tester, 4 min of rotating at 25 RPM, and then the whole tablet removed, cleaned of dust and reweighing (W_2) . Loosing weight in percentage can be calculated by the formula in equation 1. Terms friability good is less than 0.8%.

% Friability tablet =
$$(W_1-W_2)/W_1 \times 100\%$$
.....(Eq.1)

Tablet disintegration time was tested by inserting 6 tablets into the basket, and the basket fluctuates in the immersion fluid at a constant frequency rate between 29-32 cycles. If there was no parts of the tablets are left on the screen, except fragments of the coating agent, the tablet could be declared as disintegrated. Disintegration time for the uncoated tablet was no more than 15 min [13].

Table 1: Formulation of tablets

Formula	Ingredients in each ta	ablet (700 mg per table	et)						
	Dry Extract of	Dry Extract of	PVP	Gelatin	Crospovidone	Corn	Lactose	Mg.	Talk
	Syzygium	Andrographis	K-30		_	Starch	Monohydrate	Stearate	
	polyanthum Leaves	paniculata							
F1	409.09	40.91	21	-	21	-	173	7	28
F2	409.09	40.91	35	-	21	-	159	7	28
F3	409.09	40.91	21	-	35	-	159	7	28
F4	409.09	40.91	35	-	35	-	145	7	28
F5	409.09	40.91	21	-	-	21	173	7	28
F6	409.09	40.91	35	-	-	21	159	7	28
F7	409.09	40.91	21	-	-	35	159	7	28
F8	409.09	40.91	35	-	-	35	145	7	28
F9	409.09	40.91	-	21	21	-	179	7	28
F10	409.09	40.91	-	35	21	-	159	7	28
F11	409.09	40.91	-	21	35	-	159	7	28
F12	409.09	40.91	-	35	35	-	145	7	28
F13	409.09	40.91	-	35	-	35	173	7	28
F14	409.09	40.91	-	35	-	21	159	7	28
F15	409.09	40.91	-	21	-	35	159	7	28
F16	409.09	40.91	-	21	-	21	145	7	28

Determination of total flavonoid content in the tablet

An accurately weighed amount of the powdered tablets (700 mg) was dissolved in 25 ml of ethanol 96%, and then hydrolyzed with 4N

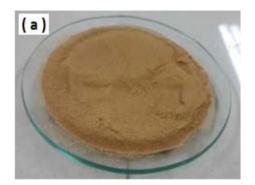
HCl (1:1) for 30 min. After that, 25 ml of distilled water was added. The flavonoid was extracted by using ethyl acetate. The extract solution was evaporated to remove the solvent. The residue was dissolved in ethanol 96% to made up 25 ml volume. 1 ml of the

solution was withdrawn and added a mixture of 3 ml of ethanol 96%, 0.2 ml of 10% AlCl3, 0.2 ml of 1M sodium acetate, and 5.6 ml of distilled water. The flavonoid was evaluated by spectrophotometer uv-vis at 440.8 nm after incubation for 30 min at room temperature.

RESULTS AND DISCUSSION

Standardization of *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extracts

The dry ethanolic extract and the results of standardization of the *Syzygium polyanthum* and *Andrographis paniculata* are shown in fig. 1 and table 2.



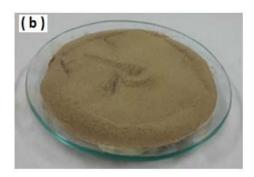


Fig. 1: Ethanolic extract of *Syzygium polyanthum* leaves (a) and ethanolic extract of *Andrographis paniculata* (b)

Phytochemical screening of *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extract

Phytochemical screening was conducted to determine the qualitative secondary metabolite contents of *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extract (table 3). The various phytochemical compounds detected are known to have beneficial importance in medicinal sciences. The flavonoids have been referred to as nature's biological response modifiers, because of their inherent ability to modify the body's reaction to allergies and virus and their antiallergic, anti-inflammatory, antimicrobial and anticancer activities [14].

Table 2: Standardization of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts

Standardization parameters	Ethanol extract of Syzygium	Requirement	Ethanol extract of Andrographis	Requirement
	polyanthum Leaves		paniculata	
Organoleptic:		-		-
-Appearance	powder		powder	
-Color	light brown		greenish brown	
-Taste	chelates	\ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \	very bitter	
-Odor	aromatic		specific	
Total ash content (%)	5.30±0.04	≤ 5.5%	9.66±0.05	≤ 10%
Water soluble ash content (%)	1.61±0.20		7.80±0.10	
Acid insoluble ash content (%)	2.12±0.10	≤ 1.8%	9.16±0.05	≤ 0.1%
Lost on drying (%)	5.73	<6%	5.23	≤ 6%

All values represent mean±SD, n=3. The standardization of dry extract shown to have a parameter that does not meet the requirements is acid insoluble ash this may be due to impurities in the extract.

Table 3: The qualitative analysis of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts

Phytoconstituents	Ethanol extract of Syzygium	Other research	Ethanol Extract of Andrographis	Other research
	polyanthum leaves	[11·12]	paniculata	[13]
Triterpenoid-Steroid	(+)	(+)	(+)	(+)
Flavonoid	(+)	(+)	(+)	(+)
Tannin	(+)	(+)	(+)	(+)
Glycoside	(-)	(-)	(-)	(-)
Alkaloid	(+)	(+)	(+)	(+)

Table 4: Total flavonoids content in the water fraction and the ethanolic extracts

Sample	Flavonoid total (%)	Comparison of ethanol extract and water fraction Syzygium polyanthum leaves-Andrographis paniculata
Ethanol Extract of Syzygium polyanthum Leaves- Andrographis paniculata (10:1)	0.175±0.011	1: 5.25
Water Fraction of Syzygium polyanthum Leaves- Andrographis paniculata (10:1)	0.916±0.011	

All values represent mean±SD, n=3

Determination of total flavonoid content in the ethanol extracts and the water fractions of *Syzygium polyanthum* leaves-*Andrographis paniculata*

The calibration curve was constructed by plotting absorbance against respective concentration of quercetin. The result shows an excellent correlation between absorbance and concentration within the concentration range 6.5–100 $\mu g/ml$. The correlation coefficient was greater than 0.995 (fig. 2).

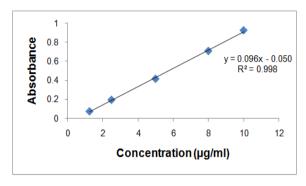


Fig. 2: Linear curve between quercetin absorbance versus concentration ($\mu g/ml$)

The physical properties of the powder and the tablet

The physical properties of the powder and the tablet are shown in table 5. F1, F2, F3, and F4 use PVP K30 as a binder and crospovidone as a disintegrant produce a granular material which has good flow properties indicated by the Carr's index and Hausner ratio which obtained. The tablet has good hardness with low friability. It is also found in F5, F6, F7, and F8 are using PVP K30 as a binder and corn starch as a disintegrant. The use of PVP K30 as a binder in the manufacture of tablets by wet granulation method will produce granules with good flow properties due to the fact that a solution of PVP K30 can spread well during granulation so as to form a bridge of liquid between the particles that will produce a good bonding resulting in excellent tablet hardening process. The use of PVP K30 as a binder produces tablets with good hardness and low friability, but it has a high disintegration time, which does not meet the requirements of the Indonesian Pharmacopoeia. The high disintegration time resulted from the strong bonds between the particles, and hence, they also produce high tablet hardness. Hygroscopic properties of PVP K30 are causing water in the air to be absorbed into the tablet which can reduce its ability of swell when

contact with a liquid medium, resulting in a longer tablet disintegration time.

Hendrati et al. (2011) used a water extract of Syzygium polyanthum Leaves-Andrographis paniculate, with a dose of 400 mg per tablet (Syzygium polyanthus Leaves: Andrographis paniculata = 2: 1), stating that the use of PVP K-30 4.45%. crospovidone 4.45% and a combination of Avicel and emcompress as a filler would produce a tablet with hardness of 6.6 kp, friability of 0.15%, and the disintegration time of 6.6 min [15]. In contrast to previous studies, in this research, the extract which used is ethanolic extract at a dose of 450 mg per tablet (Syzygium polyanthum Leaves: Andrographis paniculata = 10: 1) and the filler which used is lactose monohydrate. The use of ethanol in the extraction process produced more active substances which extracted, wherein flavonoids more easily soluble in ethanol. In this research, lactose monohydrate was used as a filler which it was easily soluble in water, so when the manufacture of tablets with wet granulation method, lactose monohydrate can be functioning as the self-binder, therefore increasing the disintegration time. This problem can be overcome by using gelatin as a binder and selected direct compression method in the manufacture of tablets. The reason to selected lactose monohydrate as filler was to increase the density of the extract that was known to have a small density.

The use of gelatin as a binder with crospovidone or corn starch as a disintegrant obtained granule with poorer flow properties compared to PVP K30 as a binder. This can be caused by a solution of gelatin in water, while at the same concentration with a solution of PVP K30, has a high viscosity. It needed a better mixing upon the addition of the binder to the powder mixture. The liquid binder can spread more evenly throughout the particle to form liquid bridges between particles that will generate strong granules and reduce the occurrence of fines which can worsen the flow properties of powders. Tablets produced using gelatin as a binder has a hardness that is no different from the tablet hardness that uses PVP K30 as a binder. This result suggests that the compressibility and compatibility of granules produced good enough to be compressed into a tablet because the bonding between the particles has good bond strength. Disintegration time of a tablet that use gelatin as a binder is faster than that using PVP K30, although both have a tablet hardness value that is nearly equal, an effect related to the porosity of the tablet, wherein the tablet which uses gelatin will have a porosity greater than a tablet that uses PVP K30 as a binder. Differences in values of bulk density and tapped density in granules that use gelatin as a binder is greater than granules using PVP K30, resulting in a tablet that is more porous, which can facilitate the entry of water in order to obtain disintegration time faster than the tablet using a PVP K30.

Table 5: Physical properties of the powder and the tablet

Formula	Bulk Density	Tapped density	Carr's Index	Hausner ratio	Hardness	Friability	Disintegration time
	(g/ml)	(g/ml)	(%)		(kp)	(%)	(min)
F1	0.36±0.04	0.42±0.04	15.00±1.41	1.18±0.02	5.72±0.11	0.46±0.00	41.03±1.97
F2	0.35±0.03	0.41±0.03	15.50±0.71	1.19±0.01	6.16±0.09	0.39 ± 0.05	48.85±0.14
F3	0.35±0.03	0.41±0.01	14.50±0.71	1.17±0.01	6.18±0.22	0.44 ± 0.00	39.37±1.42
F4	0.33±0.00	0.39±0.00	15.75±0.35	1.19±0.01	7.21±0.03	0.39 ± 0.00	42.77±0.08
F5	0.40 ± 0.02	0.47 ± 0.03	14.76±0.35	1.17±0.00	6.15±0.08	0.63 ± 0.02	47.55±0.78
F6	0.37 ± 0.01	0.44 ± 0.01	14.75±0.35	1.17±0.01	7.02±0.35	0.52±0.14	44.96±1.68
F7	0.38±0.01	0.44 ± 0.01	15.00±0.00	1.18±0.00	6.02 ± 0.11	0.59 ± 0.05	42.27±0.46
F8	0.37 ± 0.00	0.44 ± 0.01	15.49±0.69	1.18±0.01	6.67 ± 0.10	0.48 ± 0.11	45.08±0.01
F9	0.72 ± 0.02	0.44 ± 0.01	18.50±0.71	1.23±0.01	6.55±0.38	0.36 ± 0.00	12.50±0.33
F10	0.72 ± 0.01	0.57±0.00	18.50±0.71	1.23±0.01	7.03±0.31	0.33 ± 0.13	13.54±0.06
F11	0.73 ± 0.01	0.59±0.01	18.50±0.71	1.23±0.01	7.32±0.26	0.39 ± 0.01	12.73±0.15
F12	0.72 ± 0.00	0.59±0.00	18.00±1.41	1.22±0.02	6.96±0.23	0.32 ± 0.04	13.68±0.21
F13	0.59±0.00	0.73 ± 0.01	18.74±0.61	1.23±0.01	6.17 ± 0.01	0.52±0.05	13.35±0.11
F14	0.59±0.01	0.73±0.01	18.62±0.80	1.23±0.01	6.24±0.01	0.56±0.01	13.54±0.06
F15	0.58±0.01	0.71±0.01	19.01±0.62	1.23±0.01	5.84±0.02	0.48 ± 0.01	13.42±0.12
F16	0.59±0.01	0.72±0.01	18.18±0.18	1.22±0.00	5.72±0.05	0.40±0.04	13.58±0.25

All values represent mean±SD, n=3

Tablet disintegration time is strongly influenced by the binder used. The different binder (PVP K30 and gelatin) poses a very different disintegration time, despite producing tablet hardness value which is almost identical.

The use of PVP K30 as a binder with crospovidone or corn starch as a disintegrant will produce a tablet with disintegration time nearly the same time as the tablet that uses gelatin as a binder. A tablet that uses gelatin as a binder gives a friability value higher than that which uses PVP K30 as a binder. This is due to the fact that gelatin usage as a binder produces granules with a higher content of fines which can lead to increased tablet friability.

This research also conducted the optimization of the tablet formula which uses gelatin as a binder and as a disintegrant used crospovidone and corn starch.

Optimization tablet is using 22 full factorial design

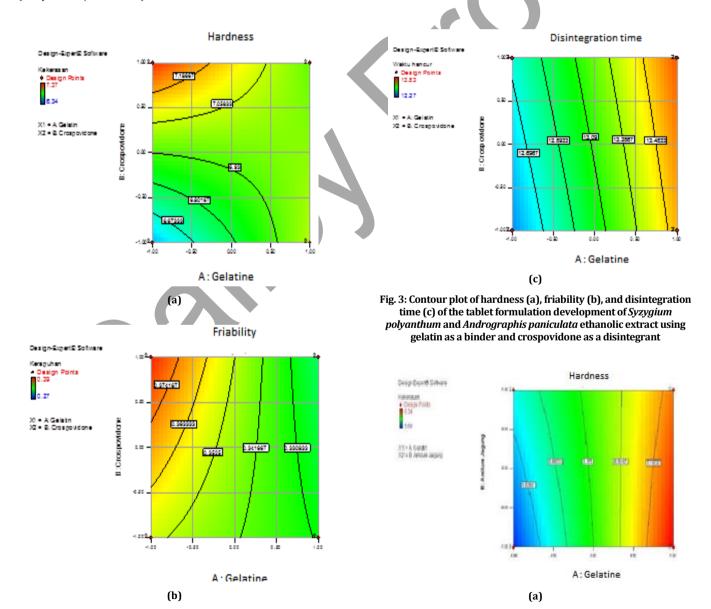
Based on data from the tablet hardness, tablet friability, and the disintegration time can be obtained the polynomial equations that are shown in table 6.

Based on the polynomial equation depicted in table 6, it appears that the concentration of gelatin has a positive correlation to the tablet hardness response, which means that increasing the concentration of gelatin will increase the hardness of tablet on either combination, with the corn starch or with crospovidone. On the response of friability, if gelatin is combined with crospovidone, it reduces brittleness compared to when combined with corn starch which can increase friability. This is due to the nature of crospovidone which is more compressible than the other super disintegrant, so it will form a tablet with a higher hardness and low friability. Based on the polynomial equations as in table 6 can be obtained contour plot as shown in fig. 3 and fig. 4.

Table 6: Polynomial equations

Response	Gelatin-crospovidone	Gelatin-corn starch
Hardness (kp)	$Y = 6.9625 + 0.0325X_A + 0.175X_B - 0.21X_AX_B$	$Y = 5.99 + 0.21X_A + 0.014X_B - 0.046X_{AB}$
Friability (%)	$Y = 0.35 - 0.023 X_A + 5.10^{-3} X_B - 0.01 X_A X_B$	$Y = 0.49 + 0.051X_A + 0.00875X_B - 0.031X_{AB}$
Disintegration tme (min)	$Y = 13.11 + 0.49875 X_A + 0.091 X_B - 0.021 X_A X_B$	$Y = 13.47 - 0.026X_A - 0.089X_B - 0.00875X_{AB}$

Y is the response, X_A is the value of the level of concentration of gelatin, X_B is the value of the level of concentration of the disintegrant (crospovidone/corn starch).



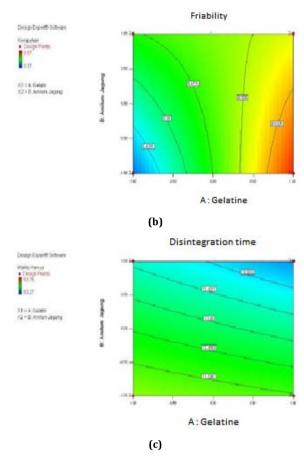


Fig. 4: Contour plot of hardness (a), friability (b), and disintegration time (c) of the tablet formulation development of Syzygium polyanthum leaves and Andrographis paniculata ethanolic extracts use gelatin as a binder and corn starch as a disintegrant

In this research combination of gelatin as a binder and crospovidone as a disintegrant was selected because it produces a tablet with a

higher hardness and a low disintegration time. The determination of the optimum formula using design expert program and obtained contour plots of superimposed as shown in fig. 5. The lower and upper limits are determined to get the optimum area can be seen in table 7.

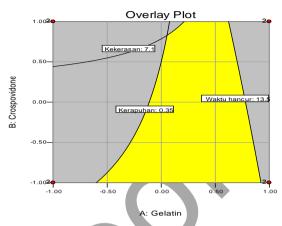


Fig. 5: Superimposed counter plot

Yellow areas depict regions predicted optimum tablet-ethanol extract of $Syzygium\ polyanthum\ leaves-Andrographis\ paniculata$ with the desired response. One composition that is used as the optimum formula by using gelatin concentration of 3.61% and 3.45% crospovidone concentration.

Physical properties of granules and tablets on a optimum tablet formula

The test results theoretically and testing of hardness, friability, and disintegration time can be seen in table 8. Based on one test sample t-test, it was found that the test results in the hardness, the friability, and the disintegration time are not significantly different with theoretical results. It can be seen from the value of-11.00<t(0.025,1) (12.706), the value of t friability (0.195)<t (0.025,1) (12.706), and the disintegration time t value (-0.404)<t (0.025,1) (12.706), which shows that a polynomial equation for a response of hardness, friability, and disintegration time is valid.

Table 7: The lower and upper limits to determined the optimum formula

Response	Lower limit	Upper limit	
Hardness (kp)	6.0	7.0	
Friability (%)	0.25	0.35	
Disintegration time (min)	10	13	

Table 8: Comparison between experiment and theoretical

Tablet properties	Theoretical	Experiment	
Hardness (kp)	6.77	6.50±0,.3	
Friability (%)	0.35	0.35±0.02	
Disintegration time (min)	12.85	12,.76±0,.32	

All values represent mean±SD, n=3

Determination of total flavonoid content in the tablet

The maximum absorption wavelength used is 440.8 nm with a standard curve equation Y =-0.049+9,610 x 10^{-2} X. The assay of total flavonoids in the tablet formulation development of *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extract is 0.112% with 98.44% recovery.

CONCLUSION

The herbal tablet formulation development of *Syzygium polyanthum* and *Andrographis paniculata* ethanolic extract can be prepared using

gelatin as a binder and crospovidone as a disintegrant using the direct compression method. The optimum formula derived from the design optimization using 2^2 full factorial design is the use of gelatin (3.61%) and crospovidone (3.45%), which will generate hardness of 6.50 kp, friability of 0.35%, and disintegration time of 12.76 min.

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CONFLICT OF INTERESTS

Declared none

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