INTRODUCTION

Molecular damage in the body can be induced by molecules called free, radicals\(^1\). Free radicals can be formed due to internal and external sources of free radicals\(^2\). Internal sources of free radicals are factors derived from normal metabolite processes in the human body, namely phagocytes, xanthine oxidase, arachidonic pathways, peroxisomes, inflammation and others. External sources of free radicals are factors that originate outside the human body, namely cigarette smoke, environmental pollution, sunlight, chemicals, ozone, several types of drugs, pesticides and others. Excessive levels of free radicals are a trigger for various degenerative diseases and conditions\(^3\). Antioxidants can inactivate the development of oxidation reactions by binding to free radicals and highly reactive molecules so that cell damage can be inhibited\(^4\). The use of natural materials that have biological activity is the motivation for further research, after synthetic compounds that have biological activity such as synthetic antioxidant compounds Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene (BHT) are restricted because they are carcinogenic\(^5\). There is concern about the possible side effects of synthetic antioxidants causing natural antioxidants to become alternatives that need to be developed\(^6\). One of the plants that can be used as medicine is a plant \(Duchesnea indica\) (Jacks.) Focke) known as Arbenan. The community uses this plant as a fever, anti-inflammatory, stop bleeding, destroy blood clots, and reduce swelling\(^6\). Flavonoids are compounds that have antioxidant activity due to the presence of a hydroxy group in their molecular structure so they are called bioflavonoids. Likewise, some tannins have been shown to have antioxidant activity, inhibit tumor growth and inhibit enzymes such as reverse transcriptase and DNA topoisomerase. While some saponins work as antimicrobials\(^6\). However, the use of Arbenan as an

**ABSTRACT**

**Objective:** Arbenan \((Duchesnea indica)\) plants contain saponins, flavonoids, and tannins which have antioxidant activity. The purpose of this research is to perform formulation and evaluation extract ethanol of Arbenan leaves in the form of serum which is pharmaceutically stable.

**Method:** Arbenan leaf powder was macerated with ethanol solvent, and then left for 3-4 days while stirring repeatedly, and then filtering. Furthermore, the liquid ethanol extract that has been obtained is evaporated using a Rotary Vacum Evaporator was used to evaporate the extract. Prepared extract was used to evaluate various parameters like organoleptics, homogeneity, viscosity, and pH.

**Result:** All formulations were having typical smell, light brown color and a little thick consistency. Formulations of leaf extract of Arbenan with four variations bases have shown to have good stability after stress condition. It can be seen from the evaluation result are organoleptics, homogeneity, viscosity, rheology, and pH.

**Conclusion:** Study concludes that a stable leaf extract of Arbenan can be effectively formulated into a serum by the means of various bases.

**Keyword:** Arbenan leaves, ethanol extract, HPMC, serum.
antioxidant for the skin is not widely known by the public. Aesthetic use certainly does not provide comfort. If further research is carried out, Arbenan plants can be formulated to facilitate their use. Preparations, especially cosmetic preparations, have developed into several dosage forms aimed at increasing convenience for their use, one of which is serum. Serum is a preparation that has more bioactive components. Serum has the advantage that it can provide a more comfortable effect and is easier to spread on the skin surface because its viscosity is not too high.

Serum or called concentrate, contains ten times more biologically active substances than cream preparations, so it is faster and more effective. Serum has fast absorption properties and the ability to penetrate into the deeper layers of the skin. The selection of serum preparations is motivated by the form of preparation that is easy to make, practical to use, easily absorbs into the skin and gives a soft and moist feeling after use. Serum works locally on different parts of the body, face, neck, eyelids. This preparation can be used regardless of age. Based on the explanation above, a research will be conducted on the formulation and evaluation of the ethanol extract serum of Arbenan [Duchesnea indica (Jacks.) Focke] leaves which are pharmaceutical stable. This research was conducted to make serum preparations that are practical to use and provide antioxidant activity that can maintain its stability on storage. The use of the percentage of the extract in a formula based on IC50 Arbenan leaf ethanol extract was 30.20 µg/mL. An increase in the extract concentration was carried out up to 100x times that of the IC50 in order to qualify as a serum, namely a highly concentrated skin preparation.

| MATERIALS AND METHODS |
|------------------------|

**Sampling**
The sample used was the leaves of Arbenan [Duchesnea indica (Jacks.) Focke] taken from Mount Bawakaraeng, Gowa Regency, South Sulawesi.

**Preparation Sample**
The collected samples of Arbenan were cleaned of dirt adhering to the leaves using running water and then dried by aerating. After drying the sample is then mashed.

**Extraction Method**
Arbenan leaf powder is weighed as much as 300 grams, put in a maceration container then 2700 mL of 70% ethanol solvent is added until the sample is submerged, then left for 3–4 days while stirring repeatedly, then filtering and obtaining residual and liquid ethanol extract. Furthermore, the liquid ethanol extract that has been obtained is evaporated using a Rotary Vacum Evaporator to obtain a thick ethanol extract. The use of extract percentage in formulas based on IC50 Arbenan leaf ethanol extract is 30.20 µg/mL. Increase the concentration of the extract to 100x times that of IC50 in order to qualify as a serum with high concentrated skin preparations.

**Preparation of serum ethanol extract of Arbenan leaf**
Prepared tools and materials. Weigh all ingredients to be used. The suitable basic formulation is selected from the optimization results. The base is dispersed in heated aquadest and added with propyl paraben and methyl paraben which have been dissolved in propylene glycol, added tocopherol then homogenized.

**Characterization of serum extract of Arbenan Leaf**

**Organoleptic**
Organoleptic tests are performed visually on serum preparations which include shape, color, and smell.

**Homogeneity**
The preparation is placed between two glass slides and then the presence of coarse particles or inhomogeneity under the light is observed.

**Measurement of Viscosity and Flow Properties**
Viscosity measurements were carried out using a Brookfield viscometer. The preparation is put into a measuring cup then the appropriate spindle is lowered until the spindle limit is immersed into the preparation. Then the motor and spindle are started. Player speed is set 0.5 successively; 2; 5; 10; and 20 rpm is then reversed from 20; 10; 5; 2; and 0.5 rpm. The viscosity number indicated by the red needle is noted. Then it is multiplied by the correction factor in the table on the tool brochure.

**Stability Test**
Evaluation of the stability of the preparation is carried out before and after the conditions are imposed. The condition was enforced by storing the preparation as much as ±100.00 mL at a temperature of 5°C and 35°C alternately for 12 hours each for 10 cycles.

**Deployment Ability**
A total of 0.5 mL of the preparation was placed on a diameter of 15 cm round glass, another glass was placed on it and allowed to stand for 1 minute. Then, a 50 gram load is added and allowed to stand for 1

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**Table 1: Different formulations of serum extract of Arbenan leaves.**

| Materials      | Formula 1 (%b/v) | Formula 2 (%b/v) | Formula 3 (%b/v) | Formula 4 (%b/v) |
|----------------|------------------|------------------|------------------|------------------|
| Extract Arbenan| 0.302            | 0.302            | 0.302            | 0.302            |
| Leaves         | 0.5              | 1                | -                | -                |
| HPMC           | -                | -                | 0.5              | 1                |
| Na CMC         | 0.02             | 0.02             | 0.02             | 0.02             |
| Propyl paraben | 0.02             | 0.02             | 0.02             | 0.02             |
| Methyl paraben | 0.02             | 0.02             | 0.02             | 0.02             |
| Propylen Glycol| 5                | 5                | 5                | 5                |
| α-tocopherol   | 0.03             | 0.03             | 0.03             | 0.03             |
| Aquadest add   | 100              | 100              | 100              | 100              |
minute and then a constant diameter of 5-7 cm is measured, showing a semisolid consistency which is very comfortable to use. The pH examination of serum preparations was carried out before and after the stress condition using a pH meter. The pH meter is immersed into the serum preparation to the limit of the mark and the pH value of the serum preparation will be read.

| Tabel 2: The results of the measurement of the different parameters of the Arabenan leaf extract serum formula before and after the conditions were imposed. 

| Formula | Average Viscosity (Poise) | Diameter (Average) | pH (Average) |
|---------|--------------------------|-------------------|--------------|
|         | Before | Before | After | After | After | After |
| 1       | 127.6  | 9.50   | 6.02  | 6.07  | 9.38  | 126.3 |
| 2       | 754.3  | 9.25   | 5.6   | 5.5   | 9.63  | 752   |
| 3       | 33.6   | 14.88  | 6.17  | 6.02  | 14.50 | 33.3  |
| 4       | 144.6  | 12.75  | 6.03  | 6.12  | 13.25 | 143.3 |

**RESULT AND DISCUSSION**

Molecular damage in the body can be induced by molecules called free radicals. Free radicals can be formed due to internal and external sources of free radicals. Excessive levels of free radicals are a trigger for various degenerative diseases and conditions. Antioxidant can be prevents or prevents oxidation, or natural or synthetic substances. Antioxidants can inactivate the development of oxidation reactions by binding to free radicals and highly reactive molecules so that cell damage can be inhibited. Synthetic compounds that have biological activity such as synthetic antioxidant compounds Butylated Hydroxyanisole (BHA) and Butylated Hydroxytoluene (BHT) because of their use because they are carcinogenic. Therefore, natural antioxidants are an alternative that needs to be developed. One of the uses of natural materials that have biological activity is Arabenan plants. All parts of the Arabenan plant contain saponins, flavonoids, and tannins. Flavonoids are compounds that have antioxidant activity due to the presence of a hydroxy group in their molecular structure so they are called bioflavonoids. However, the use of Arabenan as an antioxidant for skin is not widely known by the public. Aesthetic use certainly has typical smell, light brown color and a little thick consistency. The homogeneity test was carried out to see a homogeneous serum composition. The composition of the serum is said to be homogeneous if there is an even color equation and no different particles are found. Homogeneity observations were carried out visually with a glass object, where smearing the serum sample on a glass object was observed. From the results of testing the homogeneity of F1, F2, F3 and F4 before and after the forced conditions on the serum formula of Arabenan leaf extract showed that the formula was homogeneous which was marked by the absence of coarse particles in the preparation. Viscosity testing aims to determine the consistency of preparations that affect the skin. The higher the viscosity value, the more difficult it is to apply to the skin. The factors that affect viscosity are pressure, temperature, size and molecular weight. The results obtained that the serum using HPMC had a higher viscosity than that of Na CMC. In measuring the viscosity of Arabenan leaf extract serum, a Brookfield Viscometer was used. The viscosity of the preparation was measured using a spindle number 62 with the rotating speed adjusted to 0.5 successively; 2; 5; 10; and 20 rpm is then reversed from 20; 10; 5; 2; and 0.5 rpm for four replications. The results obtained can be seen in Table 2. The viscosity of the preparations before and after the stress condition there was a change in the decrease in the mean viscosity at F1, F2, F3 and F4. This may be because in the formulation there are variations in the concentration of the base used to improve the appearance of the preparation. The viscosity data obtained were analyzed statistically using the One-Way ANOVA method. The results of the analysis can be seen in Table 2, which shows that for serum preparations, the viscosity of all formulas. There was slight significant change in the conditions. This suggests that the existence of a forced condition greatly affects the viscosity of all norms.
In determining the flow type of Arabenan leaf extract serum, a Brookfield Viscometer was also used. The type of flow can be seen from the rogram and the yield value of the preparation. The yield value is the price that must be met in order for the preparation to flow. The yield value is obtained from measuring the viscosity of the preparation at several rpm, then the data obtained can be determined by shearing stress and shear speed (rate of shear). After creating a rheogram linking the shearing stress and the shear speed (rate of share), the newton flow type is obtained which is formed from the four formulas, namely plastic flow. It is said to be plastic flow because the results of the rogram show that the rising and falling curves cut the yield value in the absence of a hysteresis loop. Flow types in F1, F2, F3 and F4 did not change the flow both before and after the accelerated condition. The results obtained can be seen in the rheogram.

The spreadability test is carried out to determine how much the spreadability of the serum is, because it is a good preparation and is preferable if it can spread easily and is comfortable to use. The greater the dispersibility value, the easier the serum will spread on the skin. From the test results, the spreadability is inversely proportional to the viscosity of a preparation, the thicker the consistency, the smaller the dispersion power produced. The value of the scattering power whose consistency is very comfortable to use is 5-7 cm. In the Mappa study it was also said that the dispersibility of the gel was 8-15 cm. In the Mappa study it was also said that the dispersibility was very good because the viscosity of Na CMC is too high. When Na CMC is put into water, Na + is released and replaced with H + ions and forms HCMC which will increase the viscosity and Na CMC also determines the viscosity stability and spreadability of the gel preparation so that further research is needed regarding the effect of differences in the concentration of additional strength of the gel on physical stability, with this study it is necessary to adjust the formula. After the spreadability test, pH measurements were carried out which aims to see whether the pH on the preparation matches the pH on the skin. The pH measurement of the preparation was carried out before and after the conditions were imposed. This is related to the problem of stability and safety of using preparations to avoid irritation of the skin for its users, the pH of skin preparations should have a pH that is approximately the same as the pH of the skin, which is between 5.6 to 6.17. A stable pH indicates that the components in the preparation are still in the pH range category and are not affected by temperature so that the preparation remains stable during storage. So that the results of pH measurements for each formula 1 and 2 on the basis of HPMC and formulas 3 and 4 on the basis of Na CMC are concluded to remain stable and safe to use. Even though the pH has decreased and increased after the conditions are imposed, the indicated pH changes are very small and still acceptable because they still meet the pH range of the preparation for the skin.
CONCLUSION

Results of the research that have been carried out can be concluded that leaf extract of Arabenan can be formulated into a serum with various bases of HPMC and Na CMC. The serum formula for leaf extract of Arabenan is stable and complies with the required parameters.

AUTHORS CONTRIBUTION

All authors have worked equally for the literature survey, lab work and writing of the manuscript.

CONFLICT OF INTEREST

No conflict of interest, associated with this work.

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REFERENCES

1. Andarwulan N, Faradila, Fitri RH. Phenolic compounds in some indigenous vegetables from Indonesia. SEAFAST Publisher IPB: West Java, 2012. https://doi.org/10.1016%2Fj.foodchem.2010.01.033
2. Andana M, Aeyni V, Ibrahim A. Formulation and optimization of HPMC (Hydroxy Propyl Methyl Cellulose) gel base with various concentrations. J Trop Pharmacy Chemical 2015; 3(2). https://doi.org/10.25026/jtpc.v3i2.95
3. Artanti N, Jamilah, Hartati S. Technical report of the development of potential anticancer potential compounds from Taxus sumatrana and Benalu, LIPI Chemical Research Center, Serpong, 2003, 2 (3). https://doi.org/10.22146/jpc.v3i2.95
4. Baumann L, Allemann IB. Cosmetic Dermatology, mcGraw-Hill eBook, University of Miami, 2009. https://fliphtml5.com/xzqr/jtph/basic
5. Dalimartha S. Atlas of Indonesian Medicinal Plants, Pupsapwara, Jakarta, 2004; 3(117), https://doi.org/10.2305/IJUCN.UK.2019-3.3RTX.T117309548A124281670.en
6. Farmawati N, Azirahwati, Anwar E. Formulation of tyrosinase inhibiting serum containing phytosomes of longan seed extract (Dimocarpus longan Lour) using casein-xanthan gum coprocessing excipients. Faculty of Pharmacy, University of Indonesia, Depok, 2014.
7. Hanindyo RB. Antioxidant activity test on robusta coffee bean extract (Coffea canephora) DPPH method, faculty of medicine and health sciences. UIN Syarif Hidayatullah, Jakarta, 2014. https://doi.org/10.3390/nu6020466
8. Mardiyyanti S, Anwar E, Saputri FC. Serum formulation as a burn healer made from main raw material of cork fish (Channa strias) concentrate powder, faculty of pharmacy, University of Indonesia, Depok, Indonesia 2016; 14(2). ISSN 2614-6495. https://doi.org/10.1155%2F2018%2F3032790
9. Nuraziza, Seniwati, Waris R. Activity test of ethanol extract of Arbenan leaves (Duchesnea indica (Andr)) flocked with the DPPH method, As-Syifaa 2017; 09(02), ISSN: 2085-4714.
10. Sayuti NA. Formulation and physical stability test for gel extract of Chinese Keputing leaves (Cassia alata L.), Indonesian Pharm J 2015; 5 (2): 74-82. https://doi.org/10.22159/ijap.2017.v9i5.20073
11. Sinko PJ, Martin’s Physical and Pharmacy and Pharmaceutical Science 6th edition, Lippincott Williams & Wilkins, New York, London, 2011.
12. Swastika et al., Optimization of combination of carbopol 940 and Hydroxyppyphil Methylcellulose (HPMC) on the effectiveness of antiseptic gel for ethyl acetate fraction of Kesum leaves with simple design method. J Pharmacy Student Faculty of Med UNTAN Indonesia 2013:1 (1). https://doi.org/10.22159/ajpcr.2019.v12i3.31504
13. Sasidharan S, Joseph P, Junise. Formulation and evaluation of fairness serum using polyherbal extracts, Kerala India. Int J Pharm 2014; 4(3):105-112.
14. Sumarni T, Antioxidant activity of free radical capture of some sprouts from plant seeds of some sprouts from plant seeds of Papilionaceae family. Indonesian Pharm J 2005; 2 (2):53-61. https://doi.org/10.1016/j.foodchem.2013.07.064
15. Waris R, Dahlia A, Mursyid A. Antioxidant activity and phytochemical screening content for ethanolic extract and water extract of Arbenan leaf (Duchesnea indica (Andr.) Focke.). Proceedings of 1st International Conference On Health Science In Developing Country, Pascasarjana Universitas Muslim Indonesia, 2019:33. https://doi.org/10.1155/2020/3865139
16. Winarsi, H., Natural Antioxidants and Free Radicals, Kanisius, Yogyakarta, 2007.
17. Wyatt EL., Sutter SH, Drake LA, Dermatology Pharmacology. In: Hardman, J. G., Limbird, L.E., & Gilman, A. G. (eds), Goodman & Gilman’s the Pharmacological Basis of Therapeutics. 10th edition 1763. McGraw-Hill, New York. 2008. https://doi.org/10.1097/00000539-200205000-00085
18. Zhu M, Dong X, Guo M. Phenolic profiling of Duchesnea indica combining Macroporous Resin Chromatogr. (MRC) with HPLC-ESI-MS/MS and ESIT-IT-MS, Molecules, 2015; 20. https://doi.org/10.3390/molecules201219859