

**SYSTEMATIC REVIEW: UJI AKTIVITAS ANTI BAKTERI EKSTRAK
BUAH ANDALIMAN (*ZANTHOXYLUM sp*)**

SKRIPSI

Oleh

AMIRAH NAHDIA BATUBARA

71190811051



FAKULTAS KEDOKTERAN

UNIVERSITAS ISLAM SUMATERA UTARA

MEDAN

2023

**SYSTEMATIC REVIEW: UJI AKTIVITAS ANTI BAKTERI EKSTRAK
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SKRIPSI

Diajukan Sebagai Salah Satu Syarat Kelulusan Menjadi Sarjana Kedokteran

Oleh

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71190811051



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LEMBAR PERSETUJUAN

Hasil Penelitian dengan judul:

**SYSTEMATIC REVIEW: UJI AKTIVITAS ANTI BAKTERI EKSTRAK
BUAH ANDALIMAN (*ZANTHOXYLUM sp*)**

Yang dipersiapkan oleh:

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Judul Skripsi : *Systematic Review: Uji Aktivitas Anti Bakteri*

Ekstrak Buah Andaliman (*Zanthoxylum sp*)

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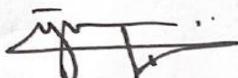
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KATA PENGANTAR

Assalamu'alaikum Wr. Wb

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dan bersedia memberikan masukan dan kritikan yang membangun penyusunan skripsi ini.

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Penulis menyadari bahwa skripsi ini masih jauh dari kesempurnaan. Oleh karena itu, saran dan kritik yang bersifat membangun demi perbaikan dimasa yang akan datang sangat penulis harapkan. Tidak ada kata yang pantas diberikan, selain balasan dari Tuhan Yang Maha Esa untuk kemajuan kita bersama. Akhirnya penulis mengharapkan skripsi ini dapat bermanfaat bagi perkembangan ilmu pengetahuan di masa yang akan datang.

Medan, 15 Januari 2023

Penulis

Amirah Nahdia Batubara

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DC	: de Candolle
WHO	: <i>World Health Organization</i>
DNA	: <i>Deoxyribonucleic Acid</i>
RNA	: <i>Ribonucleic Acid</i>
KHM	: Kadar Hambat Minimum
MIC	: <i>Minimum Inhibitory Concentration</i>
DDH	: Diameter Daya Hambat

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LAMPIRAN

Lampiran 1 Daftar Riwayat Hidup



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LEMBAR PENGESAHAN
JUDUL SKRIPSI

Judul : Studi literatur Uji Aktivitas Anti Bakteri
Ekstrak Buah Andaliman (Zanthoxylum
acanthopodium DC.)

Tujuan Umum : Untuk mengetahui gambaran umum
tertentu Publikasi efek Anti Bakteri Ekstrak Buah
Andaliman (Zanthoxylum acanthopodium DC.)

Tujuan Khusus :
 1. Mengidentifikasi jumlah artikel publikasi efek
Anti Bakteri Ekstrak andaliman.
 2. Mengidentifikasi metode dan hasil penelitian
efek Anti Bakteri Ekstrak buah Andaliman.

Nama : Amirah Nahdya Batubara
NIM : 71190811051

Pembimbing
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(dr. Syaiful Barubara)

Ka. Prodi S.Ked

(dr. Irma Yanti Rangkuti, M.Si., M.Biomed)

NB : Mohon dikembalikan ke Bag. Unit Penelitian kurang dari 2 minggu sejak melapor ke dosen pembimbing

Lampiran 3 Surat Keabsahan Daftar Pustaka

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SURAT PERNYATAAN KEABSAHAN DAFTAR PUSTAKA

Saya yang bertanda tangan dibawah ini telah memeriksa kebenaran daftar pustaka yang digunakan oleh :

Nama Mahasiswa : Amirah Nahdiah Bahubara
NPM : 71100011051
Judul Skripsi : Systematic Review Uji Aktivitas Anti Bakteri Ekstrak Buah Abdalom (Zanthoxylum sp)

Demikian pernyataan ini disampaikan, atas kerjasamanya diucapkan terima kasih.

Dosen Pembimbing Skripsi

Dr. Dafue Bahubara, M.Psi

Lampiran 4 Lembar Persetujuan Revisi Proposal

LEMBAR PERSETUJUAN

Proposal Penelitian dengan judul:

SYSTEMATIC REVIEW UJI AKTIVITAS ANTI BAKTERI EKSTRAK
BUAH ANDALIMAN (*ZANTHOXYLUM sp*)

Yang dipersiapkan oleh :

AMIRAH NAHDIA BATUBARA

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Propasal Penelitian ini telah diperiksa dan disetujui untuk

dilanjutkan ke lahan penelitian

Medan, 27 Juli 2022

Disetujui,

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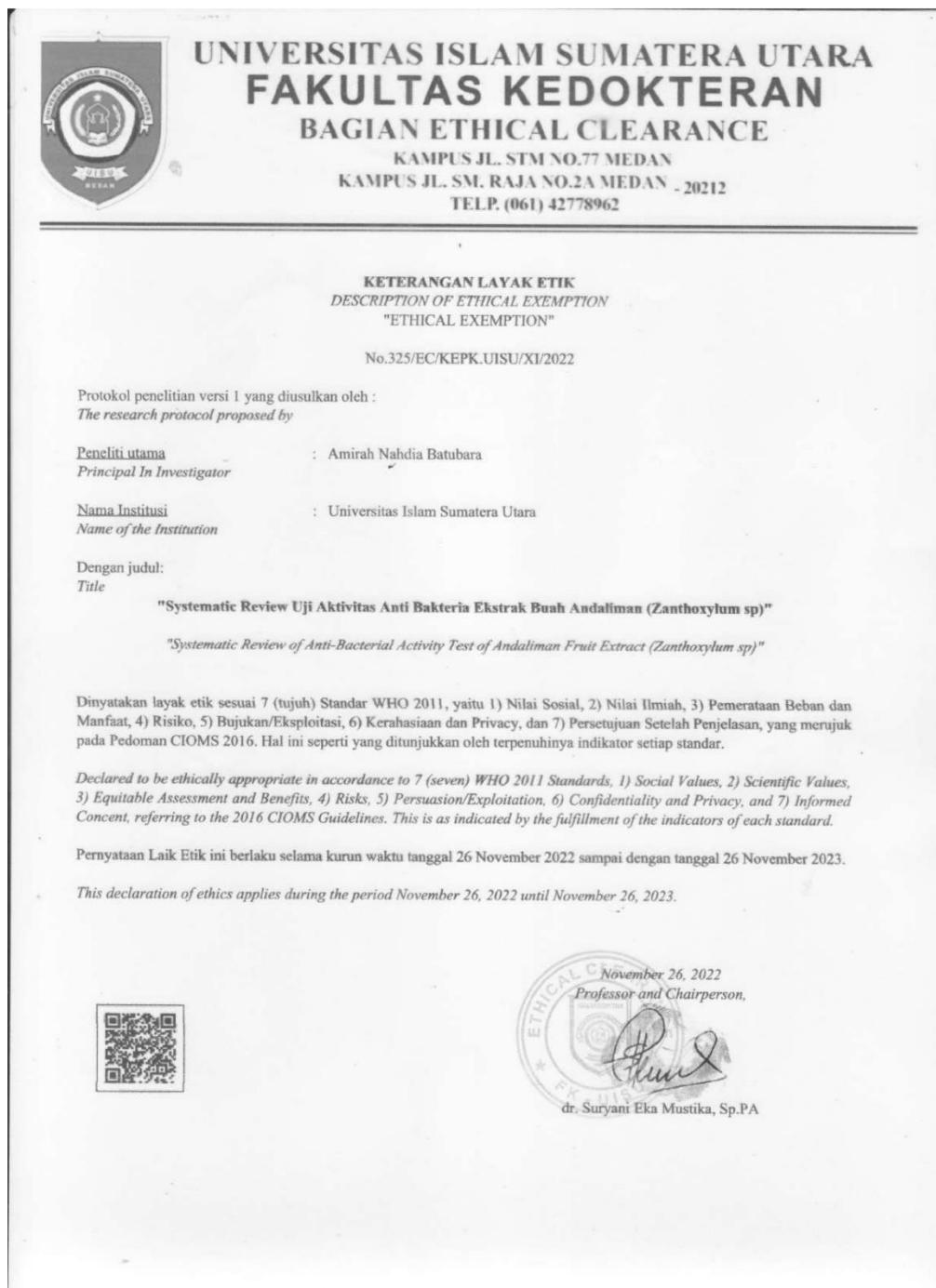
Pembanding I

(dr. Surya Akbar, MMed)

Pembanding II

(dr. Ichwan Alamsyah Lubis, M. Biomed)

Lampiran 5 Surat Keterangan Layak Etik



Lampiran 6 Penelusuran Jurnal

The screenshot shows a Google Scholar search results page for the query "aktivitas antibakteri andaliman". The search results page displays several academic papers related to the topic. An advanced search overlay window is overlaid on the main search results. The overlay has the title "Penelusuran lanjutan" and contains fields for "Temukan artikel" (Search for article) with the query "Zanthoxylum sp" "antibakteri", "dengan frasa yang sebenarnya" (with exact phrase), "dengan minimal satu kata" (with at least one word), and "tanpa kata" (without words). It also includes checkboxes for "di manapun dalam artikel" (anywhere in the article) and "dalam judul artikel" (in the title). Below these, there are fields for "Kembalikan artikel yang ditulis oleh" (Return articles written by) and "Kembalikan artikel yang dipublikasikan di" (Return articles published in), both with dropdown menus. There are also fields for "Kembalikan artikel tertanggal antara" (Return articles from date to date) with the values "2012 — 2022" and "misalnya, 1996". The overlay is semi-transparent, allowing the underlying search results to be seen.

Lampiran 7 Jurnal 1

PLOS ONE

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Structure-Activity Relationship of Benzophenanthridine Alkaloids from *Zanthoxylum rhoifolium* Having Antimicrobial Activity

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Abstract

Zanthoxylum rhoifolium (Rutaceae) is a plant alkaloid that grows in South America and has been used in Brazilian traditional medicine for the treatment of different health problems. The present study was designed to evaluate the antimicrobial activity of the stem bark crude methanol extract, fractions, and pure alkaloids of *Z. rhoifolium*. Its stem bark extracts exhibited a broad spectrum of antimicrobial activity, ranging from 12.5 to 100 µg/mL using bioautography method, and from 125 to 500 µg/mL in the microdilution bioassay. From the dichloromethane basic fraction, three furouquinoline alkaloids (1–3), and nine benzophenanthridine alkaloids (4–12) were isolated and the antimicrobial activity of the benzophenanthridine alkaloids is discussed in terms of structure-activity relationships. The alkaloid with the widest spectrum of activity was chelerythrine (10), followed by avicine (12) and dillidydrochelerythrine (4). The minimal inhibitory concentration of chelerythrine of 1.50 µg/mL for all bacteria tested, and between 3.12 and 6.25 µg/mL for the yeast tested, show this compound to be a more powerful antimicrobial agent when compared with the other active alkaloids isolated from *Z. rhoifolium*. To verify the potential importance of the methylendioxy group (ring A) of these alkaloids, chelerythrine was selected to represent the remainder of the benzophenanthridine alkaloids isolated in this work and was subjected to a demethylation reaction giving derivative 14. Compared to chelerythrine, the derivative (14) was less active against the tested bacteria and fungi. Kinetic measurements of the bacteriolytic activities of chelerythrine against the bacteria *Bacillus subtilis* (Gram-positive) and *Escherichia coli* (Gram-negative) were determined by optical density based on real time assay, suggesting that its mechanism of action is not bacteriolytic. The present study did not detect hemolytic effects of chelerythrine on erythrocytes and found a protective effect considering the decrease in TBARS and AOPP (advanced oxidized protein products) levels when compared to the control group.

Chelerythrine Tavares LSC, Zanon G, Weber AD, Neto AT, Mostardeiro CP, et al. (2014) Structure-Activity Relationship of Benzophenanthridine Alkaloids from *Zanthoxylum rhoifolium* Having Antimicrobial Activity. PLoS ONE 9(5): e97000. doi:10.1371/journal.pone.0097000

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Introduction

Many studies have underlined the importance and use of plants as a source of antimicrobial agents [1]. In this context, plants of the genus *Zanthoxylum* (Rutaceae), which encompasses about 250 species distributed throughout the world, have been of great importance. Among these, *Zanthoxylum rhoifolium* (syn. *Zigaria rhoifolium*), native to South America (Brazil, Uruguay, Paraguay and Argentina) has been used in Brazilian traditional medicine for the treatment of many different health problems [2–4]. In French Guiana, *Z. rhoifolium* has been used as an antimarial treatment [5]. These plants are known to be a rich source of natural products, mainly alkaloids and lignans. Some of these alkaloids, including the benzophenanthridines and furouquinolines, display a variety of biological activities, such as antitumor [6–11], antimicrobial [12], anti-inflammatory [13–15] and antiparasitic [16–18] activity.

In previous studies of our research team on the crude extract of *Z. rhoifolium*, the isolation and structural elucidation of a number of alkaloids have been reported, including zanthoxylamine, nitrarine, oxyntidine, and skimmianamine [19]. We have also reported the antibacterial activity of the crude MeOH extract, aqueous extract, acid and basic fractions of *Z. rhoifolium* by disc diffusion method [20] and of some isolated alkaloids by the TLC bioassay (bioautography method) [21]. More recently, our research team reported the presence of several benzophenanthridine alkaloids with interesting antimicrobial activities from the same source [22]. Because many diseases that are widespread among underprivileged and indigenous populations, such as fever, stomach upset and respiratory diseases, among others, can be caused by fungi and bacteria, this study investigated the antimicrobial activities of the crude methanol extract, fractions obtained from this extract and purified compounds from the stem bark of *Z. rhoifolium* collected in Rio Grande do Sul, Brazil, were initially screened by

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Unit Penelitian dan Pengabdian Masyarakat
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UJI AKTIVITAS ANTIBIOTIK EKSTRAK BUAH ANDALIMAN (*Zanthoxylum acanthopodium DC*) TERHADAP PERTUMBUHAN BAKTERI *Staphylococcus aureus* SECARA IN VITRO

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Abstrak

Latar belakang. *Staphylococcus aureus* merupakan flora normal yang diperkirakan 20-75% ditemukan pada saluran pernapasan atas, muka, tangan, rambut dan vagina. Infeksi bakteri ini dapat menimbulkan penyakit dengan tanda-tanda yang khas, yaitu peradangan dan tampak sebagai jerawat, infeksi dan inflamasi folikel rambut dan pembentukan abses. Di antara organ yang sering diserang oleh bakteri *Staphylococcus aureus* adalah kulit yang mengalami luka. Buah Andaliman (*Zanthoxylum acanthopodium DC*) memiliki efek antibiotik terhadap bakteri gram positif. Flavonoid pada buah Andaliman diketahui dapat menghambat bakteri *Staphylococcus aureus*. **Tujuan.** Penelitian ini bertujuan untuk mengetahui aktivitas antibiotik ekstrak buah Andaliman terhadap pertumbuhan bakteri *Staphylococcus aureus*. **Metode.** Penelitian ini menggunakan metode eksperimental. Teknik yang digunakan dalam mengukur aktivitas antibiotik adalah metode difusi cakram. **Hasil.** Dari hasil penelitian menunjukkan bahwa ekstrak buah Andaliman (*Zanthoxylum acanthopodium DC*) dengan konsentrasi 8%, 6%, 4% dan 2% menghasilkan rata-rata diameter zona bening masing-masing yaitu 13.2 mm, 11.30 mm, 10.24 mm, 8.29 mm sedangkan diameter zona bening sefotaksim yaitu 27.67 mm dan akuedas tidak diperoleh zona bening. **Kesimpulan.** Ekstrak buah Andaliman dengan konsentrasi 8% memiliki zona bening tertinggi terhadap pertumbuhan bakteri *S.aureus*. Efek antibiotik ekstrak buah Andaliman seluruh konsentrasi tidak berbeda nyata sedangkan sefotaksim dengan ekstrak buah Andaliman seluruh konsentrasi memiliki daya hambat yang nyata.

Kata kunci: *Staphylococcus aureus*, Ekstrak buah Andaliman

Abstract

Background. *Staphylococcus aureus* is a normal flora that estimated 20-75% found in the upper respiratory tract, face, hands, hair and vaginal. This bacterial infection can cause diseases with distinctive signs, namely inflammation, infection and abscess formation. Among the organs which are often attacked by the bacteria *Staphylococcus aureus* are skin injuries. Andaliman fruit (*Zanthoxylum acanthopodium DC*) has antibiotic effect against gram positive bacteria. Flavonoid in Andaliman fruit is known to inhibit *Staphylococcus aureus*. **Purpose.** The purpose of this research is to determine the antibiotic activity of Andaliman fruit extract towards the growth of *Staphylococcus aureus*. **Method.** This research used experimental method. The technique used to measure antibiotic activity was diffusion disk method. **Result.** The result showed that the average diameter of clear zone of 8%, 6%, 4% and 2% concentration of Andaliman fruit extract were 13.2 mm, 11.30 mm, 10.24 mm, 8.29 mm respectively and the diameter of sefotaksim was 27.67 mm and no clear zone was obtained. **Conclusion.** Andaliman fruit extract with 8% concentration has the highest clear zone compared to other concentrations. Antibiotic effect of Andaliman fruit extract at all concentrations did not significantly different from each other while sefotaksim with Andaliman fruit extract did not significantly different from each other.

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medicines

Article

Chemical Composition and In Vitro Cytotoxic and Antimicrobial Activities of the Essential Oil from Leaves of *Zanthoxylum monogynum* St. Hill (Rutaceae)

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Abstract: **Background:** The *Zanthoxylum monogynum* species belongs to the family Rutaceae and is found in Southeast, Midwest, and Northeast Brazil. For this genus several biological activities have been described. **Methods:** The essential oil (EO) was obtained from the leaves of *Zanthoxylum monogynum* by hydro-distillation and was analyzed by gas chromatograph and gas chromatograph/mass spectrometry (GC and GC/MS). Also the EO of *Z. monogynum* was evaluated for in vitro cytotoxic activity against six tumor cell lines and for antimicrobial activity, performing disk diffusion and MIC assays with yeast and bacterial strains. **Results:** The chemical analysis afforded the identification of 18 components (99.0% of the EO). The major components were found to be citronellol (43.0%) and farnesol (32.0%). The *in vitro* cytotoxic activity against tumor cell lines, resulted in IC₅₀ values ranging from 11–65 µg/mL against all tested cell lines. Antimicrobial activity of the essential oil was also tested and oil was effective, especially against *Cryptococcus* sp. yeast. All the tested yeast strains showed at least 90% growth inhibition. **Conclusions:** the essential oil from leaves of *Z. monogynum* has a different qualitative and quantitative composition when compared to the composition previously described. Also this EO has significant cytotoxic activity and moderate activity against *Cryptococcus* sp. and *Saccharomyces cerevisiae* yeasts.

Keywords: *Zanthoxylum monogynum*; Rutaceae; essential oil; cytotoxic activity; antimicrobial activity

1. Introduction

The Rutaceae family has approximately 150 genera distributed worldwide in tropical, subtropical, and temperate regions. The *Zanthoxylum* genus comprises more than 200 species, being the most abundant genus in Rutaceae regarding to the number of species. In *Zanthoxylum* genus, it is common to find secondary metabolites such as alkaloids, lignans, coumarins, amides, flavonoids, and triterpenes [1,2].

In the *Zanthoxylum* genus, several biological activities—lарvicide [3], trypanocidal [4], anti-tumor [5], anti-psoriasis [6], and anti-inflammatory [7], among others—have been described. In Brazil, people of the state of Rondonia use the tea from leaves and roots of *Zanthoxylum* sp. to relieve toothache [8]. Additionally, the species *Z. simulans* and *Z. chalybeum* are used for the treatment of malaria [9,10].

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The extraction of antimicrobials component of andaliman (*Zanthoxylum acanthopodium* DC.) and its application on catfish (*Pangasius sutchi*) fillet

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Abstract. Andaliman (*Zanthoxylum acanthopodium* DC.) is a well known wild species in North Sumatra and used for seasoning in Batak's traditional cuisine. This study was aimed to examine the phytochemical constituents of andaliman fruit extracts after simple macerated in water, methanol, ethyl acetate and hexane using qualitative phytochemical analysis, and to determine its potential antimicrobial activity against *Staphylococcus aureus*, *Escherichia coli* and *Salmonella* sp by using agar well diffusion method and minimum inhibitory concentration (MIC). Phytochemicals such as alkaloids, flavonoid, glycosides, saponins, tannins, triterpene/steroid and glycoside anthroquinones were detected in the methanol extracts, but steroids and glycoside antraquinones were absent in the ethyl acetate extract. The ethyl acetate extracts showed maximum zone of inhibition and minimum inhibitory concentrations against all the experimental microorganisms. The minimum zone of inhibition was determined in hexane extracts showing less antimicrobial activity against all the experimental microorganisms. The MIC of the ethyl acetate extracts was 0,5% w/v for all tested bacteria. Application of ethyl acetate extracts of andaliman fruits showed effective for catfish (*Pangasius Sutchi*) fillet stored in refrigerator (5 °C) for 3 days.

1. Introduction

Andaliman (*Zanthoxylum acanthopodium* DC.) fruits is known as the Batak society spicy and used to eliminate the smell of fish and raw meat. They are wildly grown in Tapanuli, North Sumatera at 1500 meters above sea level [1] at a temperature of 15-18 °C [2,3]. The fruits shape are like pepper, small round, green, but becomes black when they are dry. The fruits contain aromatic compounds with a bitter taste and produce the effect of thrilling sensation on taste buds and cause the tongue felt numb. Wijaya [4] found that citronellan and limonen were the most impacting compounds on the aroma of andaliman, but β-myrcene, (z)-β-ocimene, linalool, β-citronellol, nerol, geraniol, geranial, geranyl acetate, unknown compound, and a sesquiterpene were also contributing to andaliman's fresh citrus and warm sweet-peppery aroma. It had been reported to has an anti-inflammatory activity [5] and antioxidant activity [6]. Based on Cahyana and Mardiana [7], andaliman has an antimicrobial activity.

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DAYA HAMBAT EKSTRAK BUAH ANDALIMAN (*Zanthoxylum acanthopodium DC*) DALAM ETIL ASETAT TERHADAP PERTUMBUHAN *Escherichia coli* Inhibition of Andaliman Fruit Extract (*Zanthoxylum acanthopodium DC*) in Ethyl Acetate Against The Growth of *Escherichia coli*.

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Abstract

This study aims to determine the Minimum Inhibitory Concentration (MIC) of andaliman fruit extract (*Zanthoxylum acanthopodium DC*) in ethyl acetate against the growth of *Escherichia coli* bacteria. The design of this study used an experimental method with 10 extract concentrations for MIC testing, namely 10%, 20%, 30%, 40%, 50%, 60%, 70%, 80%, 90% and 100% with 2 repetitions so that they were obtained 20 experimental units. The variables observed were the formation of clear zones, namely the inhibition produced by the presence of antimicrobial activity from andaliman fruit extract in ethyl acetate against the growth of *Escherichia coli*. Data from research results are presented in the form of tables, figure and discussed descriptively. Andaliman fruit extracts in ethyl acetate have varying inhibitory forces, namely at the concentrations of concentrations of 10% to 20% was 0.65 mm to 3.15 mm with weak inhibition categories, concentrations of 30% to 50% was 5.25 mm to 6, 60 mm in the moderate inhibition category and concentrations of 60% to 100% showed strong inhibition categories with inhibition diameters of 7.20 mm to 9.60 mm. Minimum Inhibitory Concentration (MIC) for the growth of *Escherichia coli* was at the concentration of 60% with a clear zone diameter of 7.2 mm with a strong category.

Keywords : Andaliman fruit, *Escherichia coli*, antibacterial.

PENDAHULUAN

Andaliman merupakan rempah asli dari Sumatera Utara yang disebut sebagai *the golden spice from North Sumatera*. Tanaman ini ditemukan tumbuh liar di daerah Tapanuli dan dimanfaatkan sebagai rempah pada masakan adat Batak Angkola dan Batak Mandailing. Buah andaliman dipakai sebagai bumbu penyedap masakan yang memberikan rasa pedas dan aroma yang khas (Katzer, 2004). Andaliman telah lama dipergunakan oleh suku batak sebagai campuran masakan untuk berbagai jenis makanan, seperti ikan mas arsik, natinombur, dan saksang. Masakan yang menggunakan buah andaliman

dengan cara ekstraksi. Parhusip (2006) mengekstrak buah andaliman menggunakan metode maserasi dengan pelarut etil asetat mampu melarutkan banyak kelompok senyawa seperti alkaloid, flavonoid, terpenoid, saponin, dan steroid. Ekstrak etil asetat juga menghasilkan komponen berbau tajam, komponen itu diduga golongan triterpenoid.

Triterpenoid merupakan golongan terpenoid yang mempunyai aktivitas antioksidan yang sangat bermanfaat bagi kesehatan dan berperan penting untuk mempertahankan mutu produk pangan dari berbagai kerusakan seperti ketengikan,

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UJI AKTIVITAS ANTIMIKROBA EKSTRAK ANDALIMAN (*Zanthoxylum acanthopodium* DC.) PADA *Staphylococcus aureus*

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ABSTRAK

Indonesia merupakan Negara yang kaya akan rempah-rempah. Rempah-rempah mengandung senyawa-senyawa bioaktif yang dapat berfungsi sebagai antioksidan dan, antimikroba, sehingga dapat dimanfaatkan sebagai antimikroba alami dan bias diaplikasikan pada berbagai produk pangan. Andaliman (*Zanthoxylum acanthopodium* DC) merupakan salah satu tumbuhan rempah yang banyak terdapat di daerah Kabupaten Toba Samosir dan Tapanuli Utara, Sumatera Utara. Ekstrak andaliman digunakan sebagai antimikroba. Proses ekstraksi komponen antimikroba pada andaliman dengan menggunakan metode maserasi dengan 4 jenis pelarut (Air, Metanol, Etil-Asetat, Heksana). Ekstrakan daliman yang diperoleh diuji fitokimia lalu diuji pada Bakteri *Staphylococcus aureus*. Senyawa aktif seperti flavanoid, saponin dan tanin yang terdapat pada ekstrak andaliman mampu menghamabat pertumbuhan *Staphylococcus aureus*.

Keyword: Andaliman, Antimikroba, *Staphylococcus aureus*

PENDAHULUAN

Indonesia Negara yang kaya akan rempah-rempah. Rempah-rempah merupakan sebagai bahan yang dikeringkan, dan merupakan bagian dari tanaman baik dalam bentuk utuh atau potongan, serta lebih berfungsi sebagai bahan penyedap rasa dibandingkan untuk meningkatkan nilai gizi suatu pangan (Mawadah, 2009). Andaliman

selain itu kulit, akar dan daun secara tradisional dimanfaatkan sebagai obat untuk menyembuhkan sakit perut, sakit gigi, batuk, rematik dan sakit pinggang. Andaliman memiliki beberapa aktivitas biologis seperti larvasida, anti inflamasi, analgesik, antimikroba, antioksidan dan antijamur (Negi dkk, 2011).

Andaliman mengandung senyawa flavonoid yang mempunyai aktivitas

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Antimicrobial activity of Mamica de Porca *(Zanthoxylum rhoifolium Lam)* Extract against Gram-positive and Negative bacteria

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Abstract— *Mamica de porca* (*Zanthoxylum rhoifolium Lam.*, Rutaceae) is a native medicinal plant of occurrence in the Brazilian Cerrado Biome, the most diverse and rich savanna worldwide. It's present in the states such as Tocantins, Pará, Amazonas, Acre, and Rondônia. The leaves are used to treatment of microbial infections and parasitic diseases, conferring alternative for health promotion and recovery. However, little is known about the action against Gram-positive and negative bacteria, expanding the knowledge about the potential of the antibacterial. To analyze the antimicrobial activity of leaf/crude extract of *Z. rhoifolium* L. front the standard strains Gram-positive *Staphylococcus aureus* and Gram-negative *Escherichia coli*. The ethanolic extract was obtained in the Natural Products Laboratory (UAP) of the University UnirG, Gurupi-TO, Brazil allotted to Muller Hinton Agar plates, in triplicate, inoculated with bacterial strains Standard American Type Culture Collection (ATCC). Gram-Positive *S. aureus* ATCC® 29213 and Gram-Negative *E. coli* ATCC® 25922, with the technique of wells for depositing 50 µL saline (negative control) and extract with Gentamicin® on disk 10µg for positive control. The plates were incubated in a greenhouse at 35 ± 1 °C for 24 hours, microbiology laboratory, measuring the diameter of the growth inhibition halos. The crude leaf extract of *Z. rhoifolium* showed biological activity against Gram-positive bacteria *S. aureus* and Gram-negative *E. coli*, with moderately sensitive response and growth inhibition halos ranging from 14 to 16mm and 10 to 10.3mm, respectively. The antimicrobial potential of the crude leaf

extract *Z. rhoifolium* was checked against microorganisms of different cellular structures, expanding the possibilities of antimicrobial action, especially Gram-negative bacteria, being incipient studies in this perspective and not commonly identified for other medicinal species.

Keywords— Medicinal Plants, Antibacterial Agents, Natural Products.

I. INTRODUCTION

The medicinal plants traditionally used confer an alternative for health promotion and recovery, recommended the safe and effective use according to the World Health Organization and National Policy of Medicinal Plants and Phytotherapy. Considering the chemical and biological diversity of Brazil, this review highlights the Brazilian natural products that were successfully used to develop new products and the value of secondary metabolites from Brazilian biodiversity with potential application for new products and technologies [1]. The therapeutic properties are attributed to the presence of bioactive constituents, many of which are employed in the development of drug production [2]. Although research in the field of complementary and alternative medicine has revealed the mechanisms of action and efficacy of this type of treatment, many extracts vegetables yet comply with research to be properly used for therapeutic purposes [3]. This is one of the ways to the rational use of this drug alternative antimicrobial therapy and combating bacterial multiplying resistance, a worldwide problem that persists in the area of hospital health.

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Antibacterial, cytotoxicity and metabolite profiling of crude methanolic extract from andaliman (*Zanthoxylum acanthopodium*) fruit

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Abstract. Sibero MT, Stewart AP, Frederick EH, Wijaya AP, Syafitri E, Farabi K, Murwani R, Saito S, Igurasu Y. 2020. Antibacterial, cytotoxicity and metabolite profiling of crude methanolic extract from andaliman (*Zanthoxylum acanthopodium*) fruit. *Biodiversitas* 21: 4147-4154. The local community in North Sumatra has utilized andaliman fruit (*Zanthoxylum acanthopodium*) as spices for traditional cuisines because it has a unique flavor. Information on the antimicrobial activity of *Z. acanthopodium* fruit against aquaculture pathogens and its bioactivity against leukemia cell lines are limited. The purposes of this study were to evaluate the antimicrobial activity of *Z. acanthopodium* fruit against *Tenacibaculum marinum*, *Vibrio alginolyticus*, *V. angillarum*, *V. harveyi* that are known as pathogens in aquaculture; to determine cytotoxic property against murine P388 leukemia cells; and to characterize its metabolites profile. The sample was extracted using methanol by the maceration method. Antibacterial assay was conducted by Kirby-Bauer disc diffusion method; while cytotoxicity assay using the XTT method. Proximate analysis showed that *Z. acanthopodium* fruit contained 63.41% of moisture, 24.73% of crude fiber, 9.81% of crude protein, 6.90% of ash, and 2.55% of crude fat. Several phytochemical components were detected, such as alkaloid, flavonoid, tannin, triterpenoid, and steroid. The GC/MS analysis indicated the presence of various compounds from terpenoid and terpenes derivatives. This study indicated that *Z. acanthopodium* fruit was not potential as antibacterial agents against the aquaculture pathogens; however, the methanol extract showed cytotoxic potential with IC₅₀ 19.14 µg/mL against murine P388 leukemia cells.

Keywords: Andaliman, cytotoxicity, leukemia, terpenes, *Zanthoxylum*

INTRODUCTION

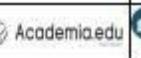
The genus *Zanthoxylum* is a member of the Rutaceae family, with 225 species found in pantropic countries (Appelhans et al. 2018). In Indonesia, *Zanthoxylum acanthopodium*, known as *andaliman* or *tuba* (in Karoese language), is a species that grows in North Sumatra, Indonesia. This plant was reported to grow in Dairi, North Tapanuli, Sidikalang, Simalungun, Sumbul, and several districts around Lake Toba (Siregar 2003; Sinaga et al. 2015; Asbur and Khairunnisyah 2018; Junaidi and Nurlaeli 2019; Saragih and Arsita 2019).

Z. acanthopodium essential oils due to volatile compounds such as β-myrcene, limonene, and citronellal (Wijaya et al. 2019). Previous studies showed the potency of *Z. acanthopodium* as an antioxidant, antibacterial against human pathogens, anti-inflammatory, and antiae (Julistiomo et al. 2018; Wijaya et al. 2019). The potency of *Z. acanthopodium* against breast (T47D and 4T1), and cervical cancer cell (HeLa) lines were reported previously (Rosidah et al. 2019; Satria et al. 2019; Syari et al. 2019). However, its potency against leukemia cancer cells has not been determined yet. Leukemia is one of the five types of cancer that causes the highest mortality in Indonesia

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	POTENSI EKSTRAK TUMBUHAN ANDALIMAN (<i>Zanthoxylum acanthopodium DC.</i>) SEBAGAI ANTIBAKTERI <i>Staphylococcus aureus</i> Oma Sepriani ^{1*} , Nurhamidah ² , Dewi Handayani ³ ^{1,2,3} Pendidikan Kimia, Jurusan PMIPA FKIP, Universitas Bengkulu *email : omasepriani@yahoo.com	
	     	

ABSTRACT

The aims of this study is to determine the secondary metabolite compounds in Roots, stem bark, leaves and fruit of Andaliman (*Zanthoxylum acanthopodium DC.*), and determine the most potent part of Andaliman plant extract in inhibiting the growth of *Staphylococcus aureus* bacteria. Andaliman plant taken in Simarpingan, Sipaholon village sub district of North Sumatra. Part of the Andaliman plantar are cleaned, cut into small pieces and then dried and mashed, samples that have been finely phytochemically tested and extracted. Testing of secondary metabolite compound by doing phytochemical screening of roots, stem bark, leaves and fruit of Andaliman, alkaloid test using meyers reagent, terpenoid test and steroid using TLC plate with eluent n-hexane : ethyl acetate with ratio 6:4, flavonoid test using solution HCl and Mg band, saponin test using soap test, tannin test using ferro (III) chloride solution. Test of antibacterial activity using disc diffusion method. Phytochemical results showed that all parts of the plants andaliman contain secondary metabolite compounds namely alkaloids, flavonoids, terpenoids, tannins and also saponins. The result of antibacterial activity test of leaf extract of leaf and leaf which has the most potential as antibacterial of *S.aureus* is bark. Because it able to inhibit *S.aureus* growth with 11 mm inhibition zone at concentration 5×10^4 ppm, which is included in the strong category. In addition to skin of andaliman leaf stem is also potential as an antibacterial with a 10 mm inhibition zone at concentration 5×10^4 ppm, which is included in the medium category.

Keywords: Andaliman, Profile of Phytochemicals, Antibacterial, *Staphylococcus aureus*

ABSTRAK

Penelitian ini bertujuan untuk mensentukan senyawa metabolit sekunder pada akar, kulit batang, daun dan buah Andaliman (*Zanthoxylum acanthopodium DC.*), dan menentukan bagian dari ekstrak tumbuhan Andaliman yang paling berpotensi dalam menghambat pertumbuhan bakteri *Staphylococcus aureus*. Tumbuhan Andaliman diambil di desa Simarpingan kecamatan Sipaholon Sumatera Utara. Bagian dari tumbuhan Andaliman dibersihkan, dipotong kecil-kecil kemudian dikering dan dihaluskan, sebagian sampel yang telah halus dilakukan uji fitokimia dan diekstraksi. Pengujian senyawa metabolit sekunder dengan melakukan skrining fitokimia terhadap akar, kulit batang, daun dan buah Andaliman, uji alkaloid menggunakan pereaksi meyers, uji terpenoid dan steroid menggunakan plat KLT dengan eluen n-heksana : etil asetat dengan perbandingan 6 : 4, uji flavonoid menggunakan larutan HCl dan pita Mg, uji saponin menggunakan uji sabun, uji tannin menggunakan larutan besi (III) klorida. Uji aktivitas antibakteri menggunakan metode difusi cakram. Hasil uji fitokimia menunjukkan bahwa seluruh bagian dari tumbuhan andaliman mengandung senyawa metabolit sekunder yaitu alkaloid, flavonoid, terpenoid, tannin dan juga saponin. Hasil uji aktivitas antibakteri ekstrak kulit batang dan daun yang paling berpotensi sebagai antibakteri *S.aureus* adalah kulit batang karena mampu menghambat pertumbuhan bakteri *S.aureus* dengan zona hambat 11 mm pada konsentrasi 5×10^4 ppm, yang termasuk kategori kuat. Selain kulit batang daun andaliman juga berpotensi sebagai antibakteri dengan zona hambat sebesar 10 mm pada konsentrasi 5×10^4 yang termasuk kategori sedang.

Kata Kunci : Andaliman, Profil Fitokimia, Antibakteri, *Staphylococcus aureus*

PENDAHULUAN

Indonesia adalah negara yang memiliki sumber daya alam yang melimpah, dimana sumber daya alam itu terdiri dari tumbuhan, hewan, mineral

spesies dari 30.000 spesies yang berasal dari hutan Indonesia. Salah satu tumbuhan yang terdapat di hutan Indonesia berpotensi sebagai obat adalah Andaliman. Tumbuhan Andaliman (*Zanthoxylum acanthopodium*

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Hindawi
Journal of Tropical Medicine
Volume 2020, Article ID 2803063, 7 pages
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Research Article

In Vitro Antibacterial Activities of Methanolic Extracts of Fruits, Seeds, and Bark of *Zanthoxylum armatum* DC

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and Sangeeta Rajbhandary¹

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Crude methanol extracts of fruits, seeds, and bark of *Zanthoxylum armatum* were investigated in vitro for antimicrobial activities against 9 different bacterial strains: *Bacillus subtilis*, *Enterococcus faecalis*, *Proteus vulgaris*, *Pseudomonas aeruginosa*, *Salmonella typhi*, *Shigella dysenteriae*, *Staphylococcus aureus*, methicillin-resistant *Staphylococcus aureus* (MRSA), and *Staphylococcus epidermidis* using agar well diffusion method, and the MBC values were determined. Only 5 bacteria, i.e., *Bacillus subtilis*, *Enterococcus faecalis*, MRSA, *Staphylococcus aureus*, and *Staphylococcus epidermidis* exhibited antibacterial properties against the different extracts. The fruit and seed extracts showed activities against 5 bacteria, while the bark extract was active against 2 bacteria only (MRSA and *Staphylococcus aureus*). *Staphylococcus aureus* was found to be more susceptible for all the extracts compared to other strains. The maximum ZOI of 20.72 mm was produced by fruits (wild) and 18.10 mm (cultivated) against *Staphylococcus aureus*. Likewise, the least ZOI of 11.73 mm was produced by seeds (wild) and 11.29 mm (cultivated) against *Escherichia coli*. Similarly, the lowest MBC value of 0.78 mg/mL was obtained for fruit extracts against MRSA, 1.56 mg/mL for fruits, seeds, and bark extracts against *Bacillus subtilis*, MRSA, and *Staphylococcus aureus*, and highest value of 50 mg/mL for fruits and seeds extracts against *S. epidermidis*. The fruits, seeds, and bark extracts of *Z. armatum* exhibited remarkable antibacterial properties against different pathogenic bacteria causing several diseases, which suggests the potential use of this plant for treating different bacterial diseases such as skin infection, urinary tract infection, dental problems, diarrhea, and dysentery.

1. Introduction

Among the 8 species of *Zanthoxylum* (family Rutaceae) found in Nepal, *Zanthoxylum armatum* DC. is one of the most common spice plants with many medicinal values. Commonly known as timur, it is a subdeciduous aromatic shrub or small tree up to 6 m high. It is found in hot valleys of subtropical to temperate Himalayas (Kashmir to Bhutan), northeast India and Pakistan, Lao, Myanmar, Thailand

diseases, which may result in undesirable side effects and serious medical problems [3, 4]. The indiscriminate use of commercial antimicrobials has also resulted in the development of multiple drug resistance. This has urged researchers to identify or extract natural antimicrobials from natural sources with less health severity [5]. Plants are the potential source of novel antimicrobial agents [6, 7]. Herbal medicinal practice, which uses plant sources to cure various infectious diseases, is also getting popularity these days. A

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IOP Publishing

Effectiveness of The Antibacterial Activity of n-Hexane Andaliman (*Zanthoxylum Acanthopodium DC*) Extract Against *Bacillus subtilis*, *Salmonella typhi*, and *Staphylococcus aureus*

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 Department of Chemistry, Faculty of Mathematic and Natural Sciences, Universitas Negeri Medan, Jl. Willem Iskandar, Pasar V, Medan, 20221, North Sumatera, Indonesia

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Abstract Infection caused by bacteria is a problem found in various countries, including Indonesia. Different pathogenic bacteria become resistant to some antibiotics. One way to inhibit the activity of these bacteria is to study natural materials that have antibacterial activity. One of the natural ingredients that have these activities is andaliman. It evidenced by the results of the analysis that the diffusion test of Andaliman extract discs uses two types of solvents: n-hexane and ethyl acetate. In the inhibition zone diameter of three types of bacteria, namely *Salmonella typhi*, *Bacillus subtilis* and *Staphylococcus aureus* and for the antibacterial test several variations of concentration of andaliman extract were used, namely: 12.5, 25, 50, and 75%, the best results for n extract -hexane at a level of 50%, namely: 26.7 mm for the best ethyl acetate extract at a concentration of 75%, 21.5 mm, both of the *Staphylococcus aureus* bacteria. These results indicate that andaliman extract is useful as an antibacterial.

1. Introduction
 Nowadays, more and more infections found that Indonesian attack people[1]. The disease is a common health problem suffered by the community due to the growth of microorganisms, one of which is pathogenic bacteria. The bacteria that cause the most cases of infection in the city are pathogenic bacteria of the species *Staphylococcus aureus* (*S. aureus*) and *Escherichia coli*(*E. coli*)[2].
 Indonesia has an extraordinary biodiversity, which is 40,000 plant species of that number, 1,300 of which used as traditional medicine. Based on this potential, conventional Indonesian medicinal products can be widely developed, for example, like herbal medicine, standardized herbal medicine or phytopharma. One of the typical plants in Indonesia that have medicinal properties is andaliman which grows in the Tapanuli region[3].
 Traditionally, andaliman fruit widely used as an aromatic ingredient, tonic, appetite stimulant, stomach-ache medicine, and diarrhoea[4]. Indian society uses andaliman fruit to treat paralysis and various skin diseases, such as boils and leprosy. Andaliman fruit also used as a spice in North Sumatra, especially North Tapanuli. Plants of the genus *Zanthoxylum* contain many phenol hydroquinone, flavonoids, steroids / triterpenoids, tannins, glycosides, essential oils, alkaloids, coumarin, lignans, amides and terpenes[5].
 Based on the research of A Muzafr, et al. in 2018 about "The extraction of antimicrobials component of andaliman (*Zanthoxylum acanthopodium DC*). Moreover, its application on catfish

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DOI: 10.30829/jumantik.v6i3.8866

Research Article

Potensi Bakteri Endofit Andaliman (*Zanthoxylum acanthopodium DC.*) dalam Menghambat Bakteri Penyebab Infeksi pada Manusia

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⁵Departemen Farmakologi Fakultas Kedokteran dan Ilmu Kesehatan Universitas Bengkulu

Abstract

*Endophytic bacteria are a beneficial bacteria that live in plant tissue. Several studies have shown that certain endophytic bacteria can produce secondary metabolites that have health effects. The andaliman (*Zanthoxylum acanthopodium DC.*) have several properties, one of which is as antibacterial against pathogenic bacteria. The most common bacterial infections in humans are caused by commensal bacteria in the body such as *Escherichia coli* and *Staphylococcus aureus*. This study used a qualitative data collection method with the type of experimental laboratory research to isolate the endophytic bacteria andaliman which had been tested against the pathogenic bacteria *Escherichia coli* and *Staphylococcus aureus*. The initial stage were isolation of endophytic bacteria and observation of colony characteristics, after obtaining isolates with different colony characteristics, a Gram stain test and a antagonist test for pathogenic bacteria were carried out. In this study, 252 colonies grew and there were 85 isolates with different colony characteristics. From all the isolates, there were two isolates that could inhibit the growth of *Escherichia coli* and seventeen isolates that could inhibit the growth of *Staphylococcus aureus*.*

Keywords: Andaliman, Endophytic bacteria, *Escherichia coli*, *Staphylococcus aureus*

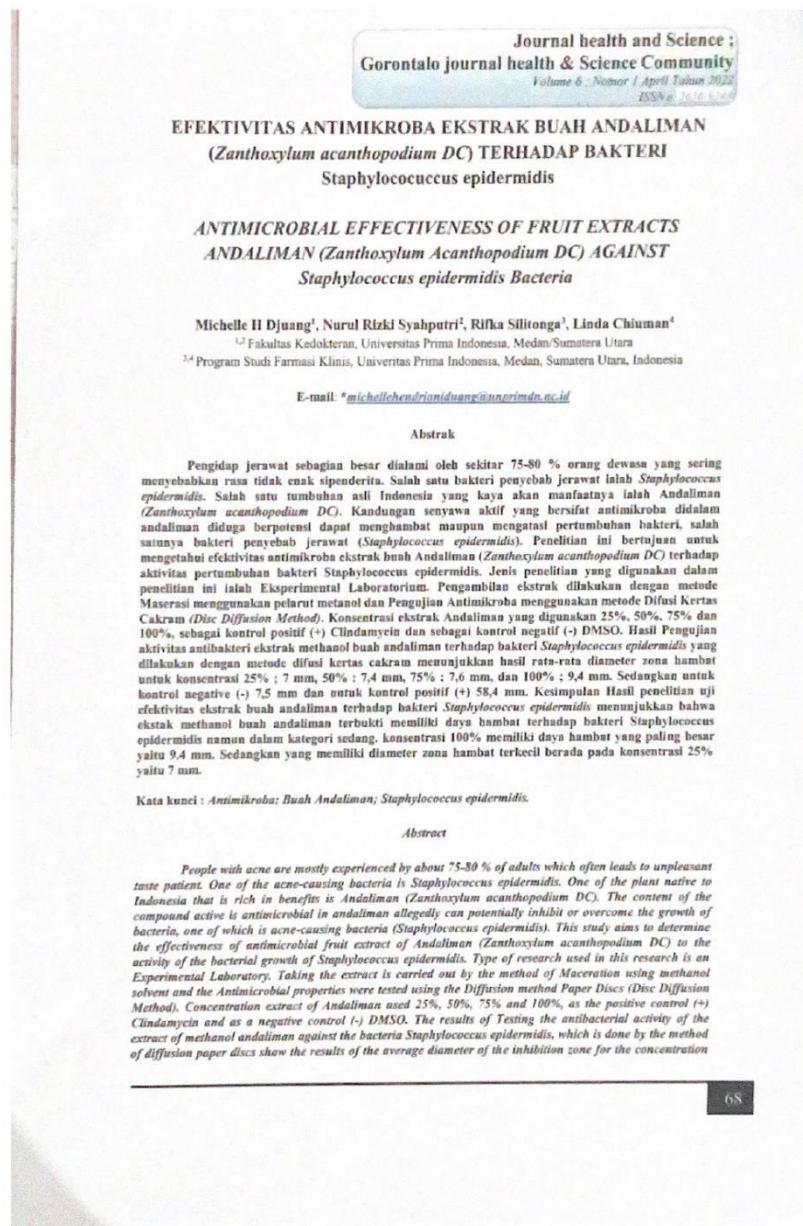
Pendahuluan

Penyakit infeksi masih menjadi salah satu masalah kesehatan yang utama. Menurut Badan Kesehatan Dunia (*World Health Organization - WHO*), sekitar 83% kematian disebabkan oleh penyakit infeksi, kelahiran dan kondisi gizi yang didapatkan oleh anak-anak (WHO, 2015). Berdasarkan hasil Riset Kesehatan Dasar (2018) oleh Kemenkes RI prevalensi untuk penyakit infeksi di Indonesia dapat dilihat dari beberapa data yang terlapor yaitu Infeksi Saluran Pernapasan Atas (ISPA) sebesar 4,4%, malaria sebesar 0,4%, pneumonia sebesar 2,0%, diare pada balita sebesar 12,3% dan diare yang terdiagnosa temage kesehatan sebesar 6,8%. Sebagian besar

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 Web : <http://kesans.nifa institute.com/index.php/kesans/index>

KESANS
International Journal of Health and Science

Testing of Andaliman Extract (*Zanthoxylum Acanthopodium* DC) With 4 Types of Solutions (Ethyl Acetate, Aquades, Methanol, And Hexane) on Growth of Bacteria Escherichia Coli

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 Indonesia
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Article Information

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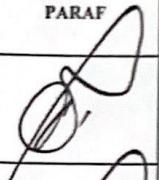
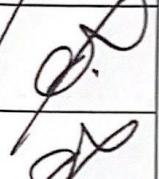
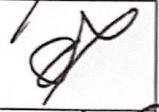
Abstract
*In general, this study aims to see whether Andaliman extract is able to inhibit the growth of *Escherichia coli* microorganisms. the first step in this research is the extraction of Andaliman with 4 types of solvents (equates solvents, methanol, ethyl acetate n-hexane). In the second stage, phyochemical tests were carried out on the andaliman extract to see the content of the active compounds in the andaliman extract, in the andaliman extract the active compounds were found to be alkaloids, steroids, and free-form triterpenes, saponins, tannins, flavonoids, and glycosides. The third stage of this study was the zone activity test inhibit andaliman extract on *Escherichia Coli* assay on bay leaf extract using blank disc paper. The extract solution that has been diluted with a concentration of 25%, 50%, 75%, 100%. Andaliman extract with an ethyl-acetate solvent with a concentration of 100% has the greatest inhibition on the growth of *Escherichia coli* with dimensions of 19.15 mm, while the has the smallest inhibition of hexane solvent with a concentration of 25% at 5.80 mm.*

Keywords: *Andaliman; Escherichia Coli; Ethyl Acetate; Aquades; Methanol; Hexane;*

How to Cite

Al Muzafri, Ria Karno Testing Of Andaliman Extract (*Zanthoxylum Acanthopodium* DC) With 4 Types of Solutions (Ethyl Acetate, Aquades, Methanol, And Hexane) On Growth of Bacteria *Escherichia Coli* Vol. 1, No. 4, January 2022
<https://doi.org/10.54043/kesans.v1i4.59>
 2808-7178
 Ria Institute

Lampiran 21 Lembar Kegiatan Bimbingan Proposal Penelitian

TANGGAL	MATERI DISKUSI	KETERANGAN	PARAF
22 Maret 2022	Judul Penelitian	Konsultasi Awal	
25 Maret 2022	Revisi Judul	-	
13 April 2022	Konsultasi Bab I		
20 April 2022	Konsultasi Bab II		
17 Mei 2022	Konsultasi Bab III		
6 Juni 2022	Penyajikan literatur		

TANGGAL	MATERI DISKUSI	KETERANGAN	PARAF
14 Juni 2022	Acara Seminar	-	

Lampiran 22 Lembar Mengikuti Seminar Proposal

MENGIKUTI SEMINAR PROPOSAL

NO	TANGGAL	PEMAKALAH	JUDUL	PARAF PEMBIMBING
1.	9/8-22	Nurjannah Br Ritonga	Hubungan Posisi duduk Pada aktivitas sehari-hari dengan terjadinya LBP pada Mahasiswa	(dr. Nondang P. Siregar, M.Sc)
2	10-22	Nadhlila Umarah syamdra	Hubungan Posisi duduk Mahasiswa FK UIN pada kegiatan sehari-hari dengan duraan terjadinya kiposis Pacival	(dr. Nondang P. Siregar, M.Sc)
3	22/8-22	Shandy	Hubungan Pengibukan Pem MPASI dan status obesitas G-2A bila di pul. malam blurn	
4.	24/8-22	Syahnillah Thalha	Hubungan Rasa lemas gadis dengan gangguan virus pada mahasiswa FK UINU angkatan 2019	

Lampiran 23 Lembar Kegiatan Bimbingan Hasil Penelitian

LEMBAR KEGIATAN BIMBINGAN HASIL PENELITIAN			
TANGGAL	MATERI DISKUSI	KETERANGAN	PARAF
24 agustus 2022	Rensi Proposal	<i>Dju</i>	<i>Dju</i>
8 September 2022	Rensi Proposal		<i>Dju</i>
26 September 2022	Konsul BAB 4		<i>Dju</i>
27 Oktober 2022	Konsul BAB 4		<i>Dju</i>
10 November 2022	Rensi BAB 4		<i>Dju</i>
4 Desember 2022	Konsul BAB 5		<i>Dju</i>

TANGGAL	MATERI DISKUSI	KETERANGAN	PARAF
7 Desember 2022	Konsul BAB 5		<i>Dju</i>
9 Desember 2022	Acc. Seminar Hasil		<i>Dju</i>

Lampiran 24 Lembar Mengikuti Seminar Hasil Penelitian

MENGIKUTI SEMINAR HASIL PENELITIAN

NO	TANGGAL	PEMAKALAH	JUDUL	PARAF PEMBIMBING
	10/ 12 2022	Rizkia Annisa	Hubungan dukungan keluarga dengan kepuasan minum obat	