



MELASTOMA MALABATHRICUM L.: A REVIEW OF ITS TRADITIONAL USES, PHYTOCHEMICAL CONSTITUENTS AND BIOACTIVITIES

Jun Sheng Low¹, Kit-Kay Mak^{1,2,3}, Mallikarjuna Rao Pichika^{2,3}, Puvaneswari Marappan¹, Shiming Zhang¹, Apiwat Jiampanichkul¹, Servenna Chua¹, Kavitha Mohandas⁴, Madhu Katyayani Balijepalli^{4,*}

¹School of Postgraduate studies and Research, International Medical University, 126, Jalan Jalil Perkasa 19, Bukit Jalil, 57000, Kuala Lumpur, Malaysia.

²Department of Pharmaceutical Chemistry, School of Pharmacy, International Medical University, 126, Jalan Jalil Perkasa 19, Bukit Jalil, 57000, Kuala Lumpur, Malaysia.

³Centre for Bioactive Molecules & Drug Delivery, Institute for Research, Development and Innovation, International Medical University, 126, Jalan Jalil Perkasa 19, Bukit Jalil, 57000, Kuala Lumpur, Malaysia.

⁴Faculty of Medicine & Biomedical Sciences, MAHSA University, Jln SP 2, Bandar Saujana Putra, 42610 Jenjarom, Selangor

*Corresponding author E-mail: madhukatyayani@gmail.com

ARTICLE INFO

ABSTRACT

Key words:

malabathricum,
Melastomataceae
traditional uses,
bioactivity,
phytochemical,
auranamide

Melastoma malabathricum L. has been used as a traditional medicine in different parts of the world. The aim of this review is to summarize and critically analyze the literature on traditional uses, phytochemistry and bioactivities of *M. malabathricum*. The literature was retrieved from scientific databases. The leaves, flowers, stem and roots are being used for wound healing, treating stomachache, measles, low sperm count, jaundice, pain, leucorrhoea, diarrhea, dysentery, tooth decay, and fever. It contains 71 phytochemicals; flavonoids, terpenoids, tannins, sterols, amides, glycolipids and fatty acids. Extracts demonstrated antimicrobial, gastroprotective, hepatoprotective, anti-inflammatory, antioxidant, cytotoxic, anticancer, wound healing, antinociceptive, antihyperlipidemic, anti-infertile, antiarthritic and antidiabetic activities. This review provides insights on traditional uses, phytochemical constituents, and bioactivities of *M. malabathricum L.* Few papers in the literature were found to be lacking in scientific vigor. Further studies should be carried out on safe use of *M. malabathricum L.* and mechanisms underlying its bioactivities.

INTRODUCTION:

The shrub, *Melastoma malabathricum L.* belongs to the family, *Melastomataceae*, is found in tropical and subtropical regions of the world¹. It has many alternative names; *M. affine*, *M. candidum*, *M. cavaleriei*, *M. esquirolii*, *M. normale*, and *M. polyanthum* (www.theplantlist.org).

One of the major global health concerns is deaths due to infectious diseases and development of bacterial resistance. Plants are reported to have potential to treat infectious diseases and to overcome antimicrobial resistance and therefore many researchers across the world are being investigating the plants to explore their antimicrobial properties especially against

clinically relevant drug-resistant strains². In addition, it is pertinent to note that healing of wounds has always been a challenging task in the clinical practice and there is a growing research interest in exploring the potential of plants for their wound healing properties³. In traditional medicine, *M. malabathricum* L. is reported to be used for the treatment of infectious diseases and wounds. In addition, there are many reports in the literature investigating its antimicrobial and wound healing properties. Among phytochemicals, flavonoids and tannins are reported to possess many interesting biological activities⁴. Many published papers on *M. malabathricum* reported the presence of flavonoids and tannins. In this review, we provide an updated and critical analysis of the literature on *M. malabathricum* regarding phytochemical and biological investigations on extracts and isolated compounds.

Methods

For the collection of the information on reported bioactivities of *M. malabathricum* L., PICOS scheme⁵ was followed: the population are bacteria or cells or tissues or animals; the intervention is treatment with *M. malabathricum* extract or isolated compound; the comparator is no treatment or placebo or standard treatment; the outcome is the effect of *M. malabathricum* extract or isolated compound; the study design is either *in vivo* or *ex vivo* or *in vitro*. Its scientific name is confirmed via www.theplantlist.org. The eligible research papers published in English language found in databases (Scopus, PubMed, SpringerLink, EbscoHost, Google Scholar), as well as textbooks until February 2020 were referred. Specific search terms were used to gather information, such as “*Melastoma malabathricum*”, “*M. malabathricum*”, “ethnomedicinal uses”, “phytochemical constituents”, “pharmacological activities”, and “medicinal uses”. All authors independently screened the titles and abstracts for choosing the eligible papers. The chemical structures of reported phytochemicals were obtained either from the research articles or PubChem

or ChemSpider. The chemical structures were drawn using ChemDraw Professional 18.0.

Distribution

According to <http://www.theplantlist.org>, *Melastoma malabathricum* L. has three synonyms; *Melastoma malabathricum* subsp. *malabathricum*, *Melastoma malabathricum* var. *normale* (D. Don) R.C. Sirvast. and *Melastoma malabathricum* subsp. *normale* (D. Don) Karst. Mey.

M. malabathricum M. malabathricum Geographic distribution and traditional & ethnomedicinal uses

The plant *M. malabathricum* is widely distributed in Asian region and cultivated as an ornamental plant. Various communities and tribes have been using *M. malabathricum* as traditional medicine for protecting their health and treatment of ailments. Its geographic distribution, vernacular names and ethnomedicinal uses are listed in Table 1.

Phytochemical constituents

The major phytochemicals present in *M. malabathricum* L. are flavonoids and tannins. Many very recent literature reviews highlighted the biological activities and therapeutic potential of flavonoids, flavonoid glycosides and tannins^{6,7}. Therefore, the reported traditional uses of *M. malabathricum* L. could be attributed to the presence of flavonoids and tannins. The flavonoids and tannins are widely distributed in leaves compared to other parts which could explain why leaves are the most widely used part of the plant in traditional medicine. However, there were no studies on the amounts and percentage of these compounds in *M. malabathricum* L., a major research gap in the published literature. The chemical structures of important bioactive phytochemicals were shown in Table 2.

Many age-old-well-known phytochemical constituents have been well documented in the 80's and 90's. These include tannins such as Alienanin, Brevifolin-carboxylic acid, Casuarinin/Stachyurin, Dimethyl hexamethoxydiphenate, Malabathrin A, B, C, D, E, F, Nobotanin B, D, G, H, J, Pedunculagin, Procyanidin B2, B5, Pterocarbin C, Strictinin Stenophyllanin A, B;

flavonoids such as Isoquercetin 6''-O-gallate, Kaempferol-3-O- β -D-xyloside, Quercetin-3-O- α -L-rhamnosyl-(1 \rightarrow 2)- α -D-galactoside and Malvidin-3,5-diglucoside; Pentacyclic triterpenoids such as Ursolic acid; sterols such as β -Sitosterol; polyphenols such as (-)-Epicatechin, (-)-Epicatechin gallate, Gallic acid; organic acids such as Hexacosanoic acid, 2,5,6-Trihydroxynaphthoic carbonic acid, *p*-Hydroxybenzoic acid, 1-Octyl decanoate A; and 11-Methyl-1-triacontanol and 32-Methyl-1-triacontanol⁸⁻¹¹. The list detailing the bioactive phytochemical constituents documented in the past 20 years is concisely tabulated in Table 3. This include information regarding their source of origin from part of plants, solvent used for extraction, isolation method as well as identification method.

Biological activities

Numerous researchers have elucidated a myriad of biological activities of *M. malabathricum*. The biological activities exhibited by *M. malabathricum* extract discussed in this paper include antimicrobial activity against gram positive & negative bacteria and fungi, gastroprotective activity, hepatoprotective activity, anti-inflammatory activity, antioxidant activity, anticancer activity, wound healing activity and antinociceptive activity. It is believed that the presence of various bioactive such as high flavonoids and tannin contents in the *M. malabathricum* extracts contribute to the synergistic effects in antioxidant, antimicrobial and anti-inflammatory activity. The biological activities of compounds studied between year 2015 to-date are summarized in Table 4.

Antimicrobial activity

The underlying issue with the surge of antibiotic resistance has become one of the main concerns shared among researchers. Apart from research focusing on small molecule organic synthesis, effort is also given to the natural products expertise in order to discover bioactive compound that these microbes can succumb to¹². Thus, extracts of *M. malabathricum* were

evaluated for their antibacterial activity. Most of the up-to date literature extensively investigated the extraction of different parts of *M. malabathricum* plant using methanol for their antimicrobial activity. It was reported in various research that some of the microbes succumbing to the extracts of leaves, flowers and fruit of *M. malabathricum* include *M. malabathricum* *C. capsica*¹³, *M. malabathricum* gram positive bacteria such as *Listeria monocytogenes*¹⁴, *Staphylococcus aureus* and *Bacillus subtilis*; and Gram-negative bacteria namely *Klebsiella pneumoniae*, *Escherichia coli*, *Salmonella typhi*, and *Shigella flexneri*¹⁵. Unfortunately, the extract fall short against as no activities were detected against gram-negative bacteria such as *Escherichia coli* and *Salmonella typhimurium*¹⁴. With this, Che Omar *et al.* and Alwash *et al.* reported that Gram-positive bacteria were more susceptible to the plant extracts than that of the Gram-negative species^{14,16}. There were however no studies reported on antimicrobial activity of *M. malabathricum* roots, in which further studies could potentially be performed.

Gastroprotective activity

It was documented that *M. malabathricum* plant was traditionally used to relieve ulcers. It is interesting to note that *M. malabathricum* leaves showed dose-dependent gastroprotective activity in male adult Sprague-Dawley rats. Literature to date reported the investigation of *M. malabathricum* leaves chloroform extract¹⁷. It was previously reported that the gastroprotective effects of *M. malabathricum* extract was performed using established ethanol-induced gastric ulcer, and ethanol- and indomethacin-induced gastric ulcers¹⁸ models. *M. malabathricum* (500 mg/kg) exhibited equipotent activity of that of Omeprazole (reference drug;) ¹⁹. In a different study, *M. malabathricum* (500 mg/kg) exhibited improved the inhibition of ulcer than that of Rantidine ¹⁷ and carbenoxolone^{20,21}.

Hepatoprotective activity

Not much experimentation was performed to investigate the hepatoprotective properties of *M.*

malabathricum, thus, further research could be performed regarding its effectiveness. Thus far, only methanol extract of leaves was tested for their hepatoprotective activity in paracetamol-induced hepatotoxicity in male adult Sprague-Dawley rats. It was noted by Mamatet *al.* that the main contributors towards its hepatoprotective activity of *M. malabathricum* methanol extract could be due to the presence of flavonoids, tannins, and saponins²². Kamisanet *al.* also reported that *M. malabathricum* methanol extract was found to be effective for hepatoprotective activity where significant improvement in the levels of aspartate aminotransferase (AST) and alanine transaminase (ALT) in both paracetamol-induced liver toxicity and carbon tetrachloride-induced liver toxicity compared to that of their respective control group²³.

Anti-inflammatory activity

Similarly, not many animal studies were performed to elucidate the mechanisms involved in reducing inflammation. Balamurugan *et al.* reported dose-dependent anti-inflammatory activity of leaves ethanol extract in carrageenan induced paw edema adult Wistar albino rats where ultimately, the percent inhibition was equipotent compared to that of the positive control group²⁴. Kumar *et al.* reported methanol extracts of leaves attenuated Complete Freund's adjuvant (CFA)-induced arthritis in a dose-dependent manner where *M. malabathricum* leaves extract at 500 mg/kg showed equipotent effect with that of the reference drug, Indomethacin (10 mg/kg) in modulating the proinflammatory mediators namely tumor necrosis factor- alpha (TNF- α), Interleukin (IL)-6, IL-1 β and COX-2²⁵.

Antioxidant

The antioxidant activity of *M. malabathricum* extracts from different extraction solvent including ethyl acetate²⁶ and ethanol²⁷ were previously reported to exhibit antioxidant activity in a β -carotene bleaching assay²⁶. Sari *et al.* reported that the leaves and fruit of the plant having antioxidant activity as potential herbal tea material where *M. malabathricum* leaves and flower extract respectively inhibited

90% and 88% of DPPH radical compared to that of commercial black tea and green tea (85-90%)²⁸. Danladi *et al.* reported that among leaves, fruits, flower and stem extracts, the flowers extract showed equipotent (IC₅₀: 48 μ g/mL) DPPH free radical scavenging activity to that of Quercetin (IC₅₀: 48 μ g/mL)²⁹. Verma *et al.* described that the *M. malabathricum* extract showed an inhibition of ABTS (99.7%; IC₅₀: 38 μ g/mL) and DPPH (98.8%; IC₅₀: 27 μ g/mL) free radicals³⁰; while Baruah *et al.* demonstrated that ABTS and DPPH values of *M. malabathricum* extract were found to be 55.58 ± 1.09 μ M/g and 0.9271 ± 0.22 μ M/g respectively³¹.

Antitumor activity: Numerous published articles had investigated both *in-vitro* and *in-vivo* for antitumor activities of different extracts from different parts of *M. malabathricum*. Roslenet *al.* reported that the leaves methanol extract showed the highest percentage of MCF-7 cells inhibition³². To date, the anticancer activity possessed by *M. malabathricum* extract was also tested on multiple cell lines, which includes human breast cancer (MCF-7)³³, Hepatoma G2 (HepG2)²⁹ and lung cancer (A549) cell lines³³. In summary, the leaves and the flowers of the plant possess higher antitumor activity compared to the stem bark, but the activity exhibited by the stem bark was still significant.

Wound healing activity

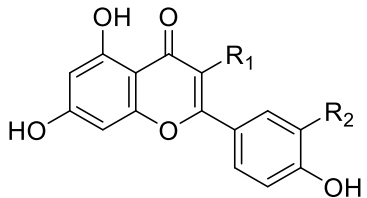
Very limited publications were reported on the wound healing activity of *M. malabathricum*. *In-vivo* wound healing studies were carried out in healthy male adult Sprague-Dawley rats demonstrated that *M. malabathricum* leaves extract was the better treatment for wound healing compared to the then reported conventional drugs, Povidone-Iodine and Acriflavine³⁴. While *in-vitro* studies were performed using human gingival fibroblasts by Ab Rahman *et al.* However, the effectiveness of wound healing activity reported by both publications was only subtle but was significant when compared to untreated groups regarding its wound closure activity³⁵.

Region	Vernacular name(s)	Part of plant	Mode of preparation	Traditional use(s)	Ref
Malaysia					
Shah Alam Kelantan Negeri sembilan Pahang	Senduduk, Sekedudok, Sikadudok, Kendudok, Kedudok, Sedudok, Lingangadi, Gosing-Gosing, Gagabang, Ngongodo, Gata-Gata	Leaves	Aqueous extract	Given to postpartum women for healing & womb strengthening.	40
		Fruit	Juice	Relieve dry lips.	
		Shoots	N/A	Puerperal infections, blood pressure, diabetes, toothache, leucorrhea.	41
		Roots	Decoction	Epilepsy, rheumatoid arthritis, tenderness in legs	42
				Diarrhea.	43
		Leaves & roots	Poultice	Aid healing process & reduce haemorrhoid discomfort.	44
Leaves & flowers	N/A	Cholera, diarrhea, prolonged fever, dysentery, leucorrhea, wounds, various skin diseases.			
India					
Assam, Manipur, Tripura, Mizoram, Odisha	Shapti, Bobuchunmei, Rongmei, Rindha, Palore, Nekkarike, Ankerki, Kinkerika, Gongoi, Koroti,	Leaves	Decoction, juice	Reduce smallpox scars, Dysentery, diarrhea, hemorrhoids	45
			Eaten raw	Dysentery.	46
			Paste	Cuts & wounds, stomach disorders, fever.	47
		Stem barks	N/A	Skin diseases.	47

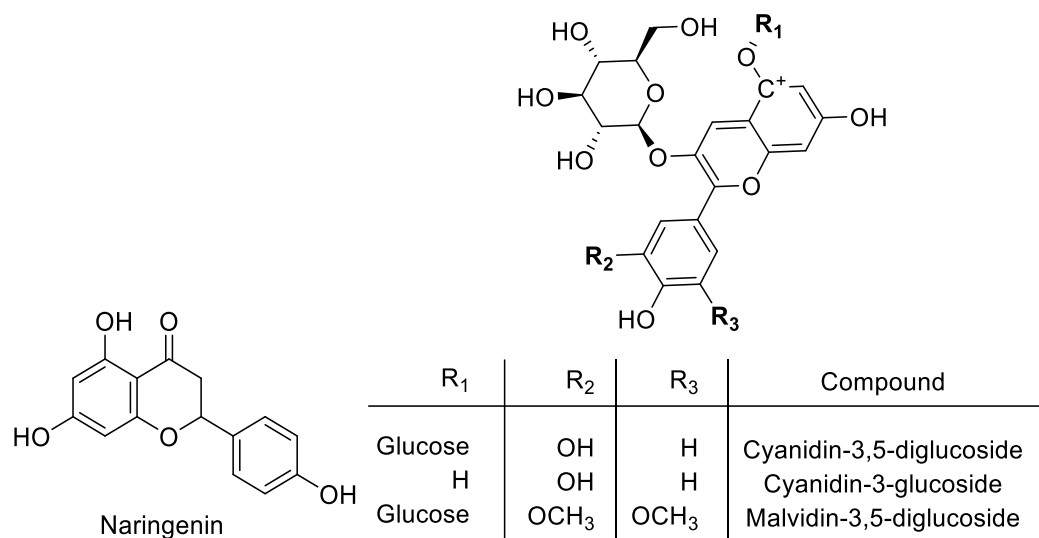
	Myetpyai, Phutki, Phutkala, Koroli, Karali	Bark, leaves and seeds	Decoction	Skin diseases, leucorrhea, diarrhea, dysentery.	48
Thailand					
N/A	Khlong Khlong, KheeNok, Mang Khre , Mang Re, Bre, Kadu-Da, Chuk Naaree	Roots	Decoction	Fever	49
Indonesia					
Sumatra West Java	Harendong, Senggani, Kemanden, Kluruk, Kendudu, Pucuk, Kendudu, DaunKhalendon g, Mua e bong, Sendudok	Leaves	N/A	Diarrhea, dysentery, gastric ulcers, haemorrhoids.	50
			Ingested raw	Toothache, Given to postpartum women for healing & womb strengthening	51
		Roots	N/A	Rheumatoid arthritis, toothache, epilepsy	50
		Flowers	Cold infusion	Anemia due to gastrointestinal bleeding, Epigastric pain.	52
Philippines					
N/A	Malatungau, Bubtoi, Yagomyum	Leaves	Powder	Dysentery	53
		Leaves, roots and flowers	Juice	Sedative, haemorrhoids, indigestion, leucorrhea, chronic diarrhoea	

Bangladesh					
Bandarban, Netrakona, Rangamati	Koiam-Pay-Bang, Kakkhu, Aksio, Koaim-pay-bang, Kakkhu	Roots	Juice	Jaundice, Leucorrhea	54
		Leaves	Juice	Diuretic, Urinary tract infections.	
Pacific Islands					
Tahiti	N/A	N/A	N/A	Diarrhoea, dysentery	53

Table 2. Chemical structures of isolated compounds from *M. malabathricum*

Flavonoids		
		
Name	R1	R2
Quercetin	OH	OH
Quercetrin	O-β-D-glu	OH
Kaempferol	OH	H
Kaempferol-3-O-β-D-glucopyranoside	O-β-D-glu	H
Kaempferol-3-O-(2'',6''-di-O-p-trans-courmaoyl)-β-glucoside	O-2,6-di-O-p-trans-courmaroyl)-β-D-glucoside	H
Quercetin-3-O-Dgalactoside	O-D-gal	OH
Kaempferol 3-O-α-L-rhamnopyranoside	O-α-L-rha	H
Kaempferol-3-O-β-D-galctopyarnoside	O-β-D-gal	H
Kaempferol 3-O-(2'',6''-di-O-E-p-	O-(2'',6''-di-O-E-p-coumaryl)-β-	H

coumaryl)- β -D-galactopyranoside	D-gal	
Kaempferol-3-O- β -D-xyloside	O- β -D-xylose	
Isoquercetin-6''-O-gallate	O- β -D-(6-O-gallyl)glu	OH
Quercetin-3-O- α -L-rhamnosyl-(1 \rightarrow 2)- α -D-galactoside	O- α -L-rha-(1 \rightarrow 2)- α -D-gal	OH
Rutin	O- β -D-(6-O- α -L-rhamnosyl)glucose	OH



Antinociceptive activity

Only a few published articles reported the investigation of antinociceptive activity of *M. malabathricum*. The reported literature only investigated methanol and ethanol extracts from the leaves and the stem bark in *in-vivo*, using male Balb/C mice³⁶ or male Sprague-Dawley rats^{37,38}. Sulaiman *et. al.* reported an inhibition of 84.4 % in acetic acid-induced abdominal writhing test with of *M. malabathricum* (300 mg/kg *i.p.*) ethanol extract³⁶. Zakaria *et. al.* reported that acetic-acid induced nociception was significantly ($p < 0.05$) attenuated by *M. malabathricum* petroleum ether (EC_{50} : 119.5 mg/kg) extract³⁷. Besides these methods latency of discomfort also showed positive results^{37,38}. Despite the different test animals and methods, it was reported that *M. malabathricum* exhibited antinociceptive activity.

Other pharmacological activities

There were also a few studies reporting interesting pharmacological activities of the plant. One such study is the study ethanolic extract of leaves increased the sperm density and motility in male albino rats³⁹. In this study, it was reported that male albino rats treated with *M. malabathricum* ethanol leaves extract (500mg/kg) daily for 14 resulted in a significant ($P < 0.01$) increase of sperm concentration ($486.30 \pm 14.83 \times 10^6$ mil) compared to that of the control group.

Concluding remarks and future perspectives

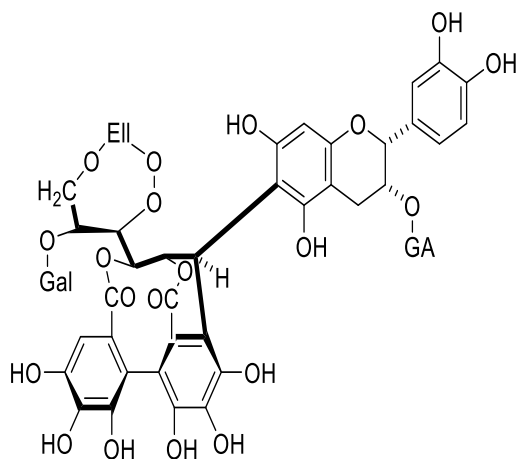
In this review we have presented the critical analysis of the reported literature on traditional uses, phytochemical constituents,

Tannins

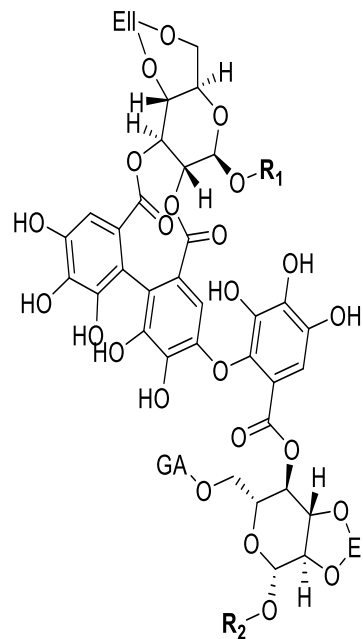
and biological activities of *M. malabathricum* L. Exhaustive literature has shown that, different parts of this plant *M. malabathricum* have been used in ethnomedicine - leaves being the most common one. Biological activity studies have been carried to provide scientific evidence for ethnomedicinal uses.

Studies have reported the phytochemical constituents and biological activities of this plant. The results have indicated the presence of tannins, flavonoids, pentacyclic triterpenoids, sterols, polyphenols, lactones, amides and glycolipids. The tannins and flavonoids are the major components. Studies also proved the biological activities; antimicrobial, antioxidant, gastroprotective, hepatoprotective, anti-inflammatory, antitumor, antinociceptive, antifertile and antidiabetic; of crude extracts.

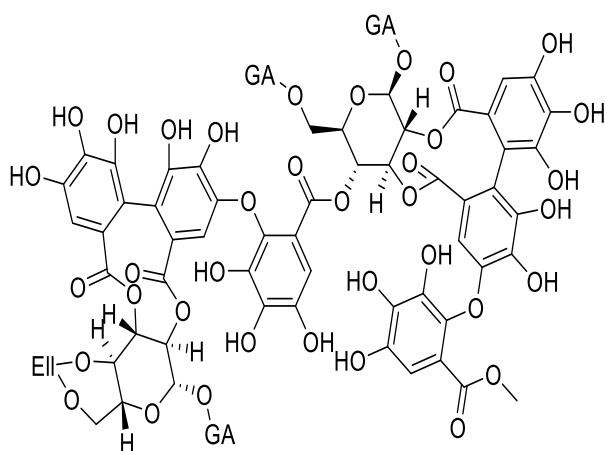
In many studies, the authors did not follow activity guided fractionation to isolate the bioactive compounds. Besides, a few studies did not report the use of adequate spectroscopic techniques to characterize the compounds. Other than few antimicrobial studies, no other studies directly compared the bioactivity of crude extracts and the isolated compounds in addition to the lack of mechanism of activity studies. No reports on quality control, toxicity and safety of crude extracts were found in the literature. The use of disease-relevant animal experiment models was scarce. The published *in vivo* assays did not follow the international standards in terms of the adequate number of animals, dose-range, use of appropriate positive & negative controls and dose-response studies.



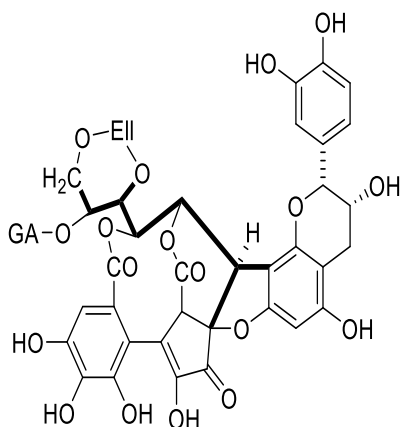
Malabathrin A



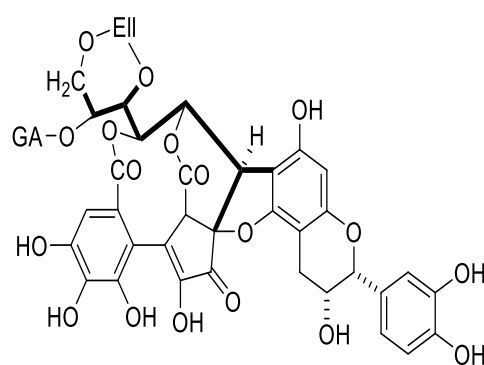
R ₁	R ₂	Compound
Gal	H	Malabathrin B
H	Gal	Malabathrin C



Malabathrin D

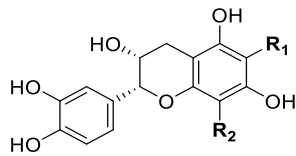
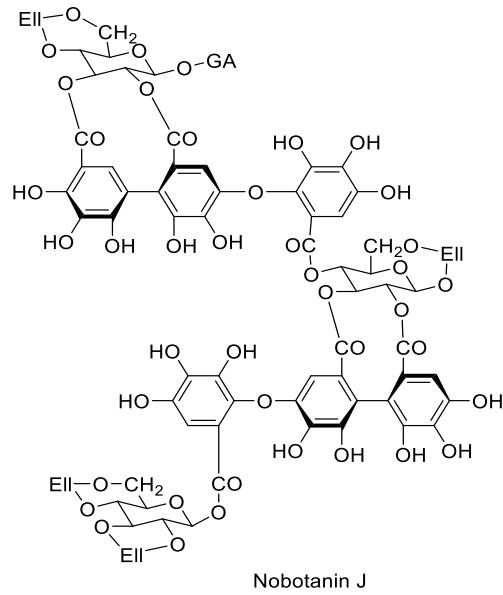
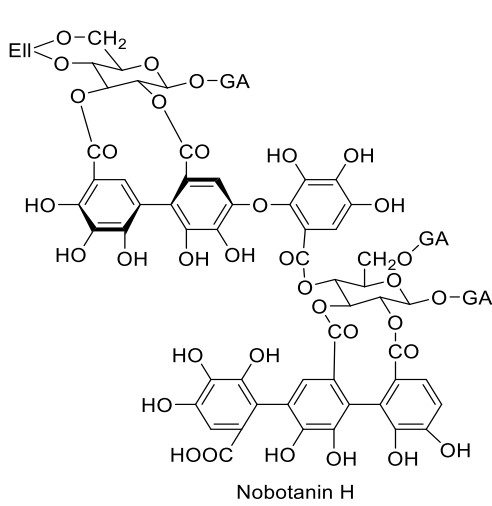


Malabathrin E

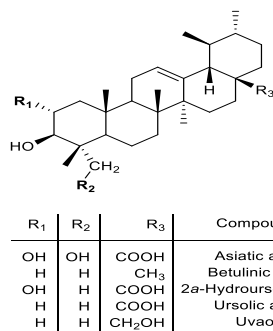
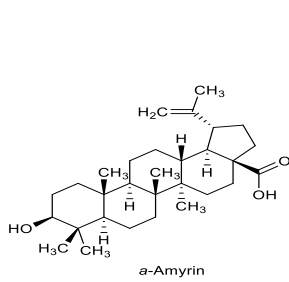
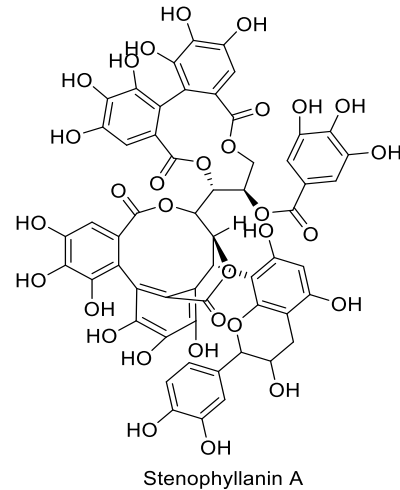
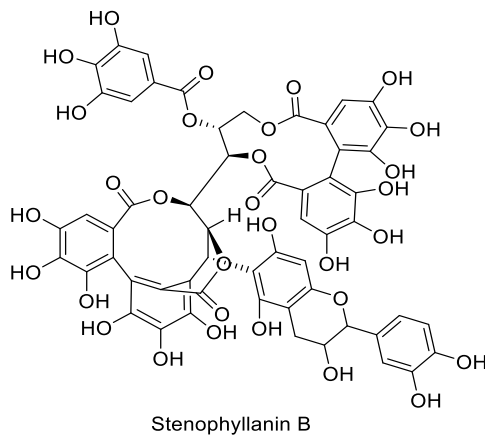


Malabathrin F

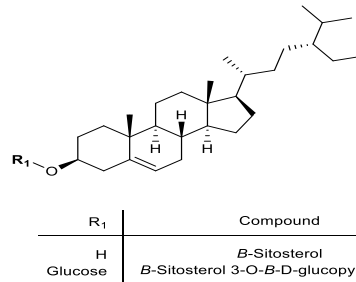
Tannins (continued)



R ₁	R ₂	Compound
H	(-)-Epicatechin	Procyanidin B2
(-)-Epicatechin	H	Procyanidin B5



R ₁	R ₂	R ₃	Compound
OH	OH	COOH	Asiatic acid
H	H	CH ₃	Betulinic acid
OH	H	COOH	2 α -Hydroursolic acid
H	H	COOH	Ursolic acid
H	H	CH ₂ OH	Uvaol

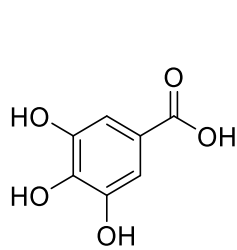


R ₁	Compound
H	<i>B</i> -Sitosterol
Glucose	<i>B</i> -Sitosterol 3-O- <i>B</i> -D-glucopyranoside

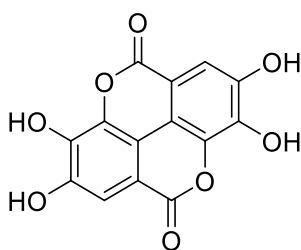
ic triterpenoids

Pentacycl

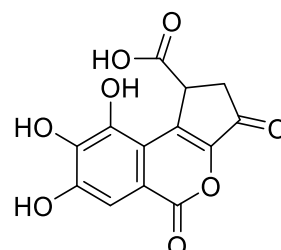
Polyphenols



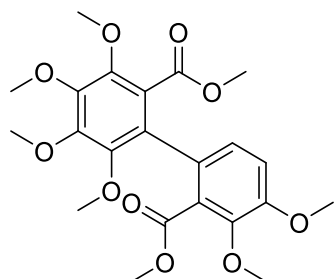
Gallic acid



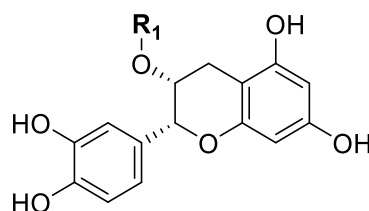
Ellagic acid



Brevifolincarboxylic acid

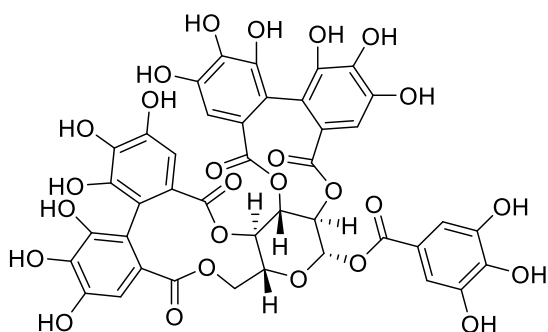


Dimethyl hexamethoxydiphenate



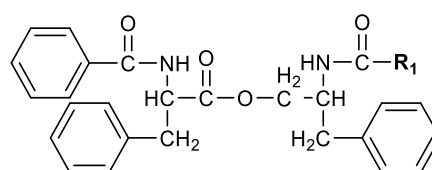
R ₁	Compound
H	(-)-Epicatechin
Gal	(-)-Epicatechin gallate

Lactone



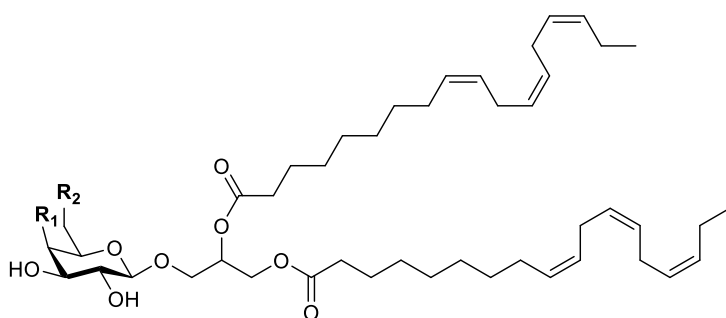
Casuarictin

Amides

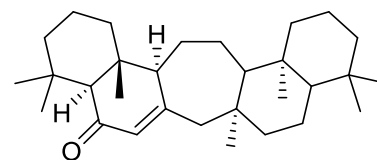
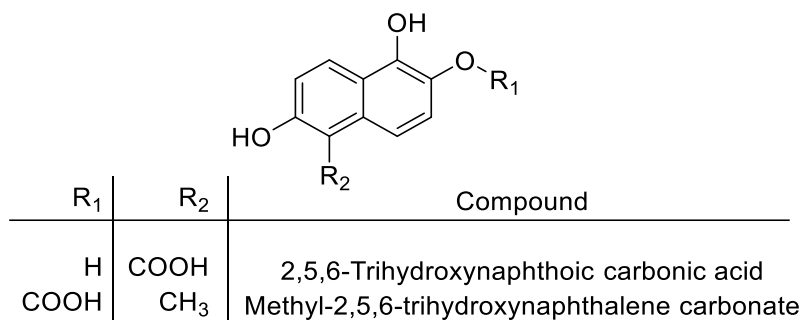


R ₁	Compound
Phe	Auranamide
CH ₃	Patriscabratine

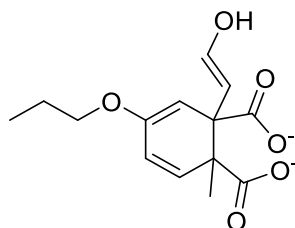
Miscellaneous compounds



R ₁	R ₂	Compound
OH	OH	Glycerol-1,2-dilinolenyl-3-O- <i>b</i> -D-galactopyranoside
2,2-dimethyl-dioxane		Glycerol-1,2-dilinolenyl-3-O-(4,6-di-O-isopropylidene)- <i>b</i> -D-galactopyranoside



Serrat-14-en-16-one



2-(2'-Hydroxyvinyl)-1-methyl-4-propoxyphthalate

Neither pharmacokinetic nor metabolic profiling of phytochemicals present in the extracts were found in the literature. The rationale of the preparing the extract in relation to how it is being prepared in traditional medicine is not reported elsewhere. None of the published reports indicated the recommended human dose. The above-said findings exposed the research gaps in the existing knowledge of *M. malabathricum L.* Therefore, this review recommends few future scientific studies to be considered for providing an evidence to support traditional uses and safe-use of *M. malabathricum L.* So far, the antimicrobial and antioxidant efficacy of the extracts were proven only using *in vitro* models, therefore further studies should be warranted to test the safety & efficacy in physiologically relevant animal modes. Since the extracts showed promising antimicrobial activities against standards strains the studies could be extended further to test their activity against clinical and drug-resistant strains. Only preliminary studies on gastroprotective, hepatoprotective, anti-inflammatory, antitumor, antinociceptive, antifertile and antidiabetic activities of the extracts were reported suggesting plenty of opportunities for future studies to elucidate the mechanisms of action and safety, metabolic & pharmacokinetic profile of the extracts. In addition, many reported traditional uses of

M. malabathricum L. such as rheumatoid arthritis, epilepsy, leucorrhoea, diarrhoea, dysentery, skin diseases, haemorrhoid and as a diuretic; were not investigated thus providing an opportunity to design future studies to prove these claims. A special attention should be paid in future studies to mimic the preparation of extracts to that of traditional preparation (decoction or juice). Lastly, all the good practices in phytopharmacology research; such as identification & authentication of the plant material, identification & characterisation of bioactive compounds, use of physiological & disease relevant experimental models, ethical standards in animal experiments and etc., must be followed in future studies.

Acknowledgement: The authors would like to thank International Medical University, Malaysia and MAHSA University, Malaysia for the library facilities.

Funding:

This work was supported by the Ministry of Education, Malaysia under Grant [FRGS/1/2018/WAB13/MAHSA/02/1].

Conflict of interest:

The authors declare that they have no conflict of interest.

REFERENCES

- Meyer K. Revision of the Southeast Asian genus *Melastoma* (Melastomataceae). *Blumea J Plant Taxon Plant Geogr.* 2001;46(2):351-398.
- Khameneh B, Iranshahy M, Soheili V, Fazly Bazzaz BS. Review on plant antimicrobials: A mechanistic viewpoint. *Antimicrob Resist Infect Control.* 2019;8(1). doi:10.1186/s13756-019-0559-6
- Bahramsoltani R, Farzaei MH, Rahimi R. Medicinal plants and their natural components as future drugs for the treatment of burn wounds: An integrative review. *Arch Dermatol Res.* 2014;306(7):601-617. doi:10.1007/s00403-014-1474-6
- Valverde Malaver CL, Colmenares Dulcey AJ, Rial C, et al. Hydrolysable Tannins and Biological Activities of *Meriania hernandoi* and *Meriania nobilis* (Melastomataceae). *Molecules.* 2019;24(4). doi:10.3390/molecules24040746
- Hajizadeh-Sharafabad F, Ghoreishi Z, Maleki V, Tarighat-Esfanjani A. Mechanistic insights into the effect of lutein on atherosclerosis, vascular dysfunction, and related risk factors: A systematic review of in vivo, ex vivo and in vitro studies. *Pharmacol Res.* 2019;149:104477. doi:10.1016/j.phrs.2019.104477
- Jucá MM, Cysne Filho FMS, de Almeida JC, et al. Flavonoids: biological activities and therapeutic potential. *Nat Prod Res.* 2020;34(5):692-705. doi:10.1080/14786419.2018.1493588
- Ajebli M, Eddouks M. The Promising Role of Plant Tannins as Bioactive Antidiabetic Agents. *Curr Med Chem.* 2019;26(25):4852-4884. doi:10.2174/0929867325666180605124256
- Yoshida T, Nakata F, Hosotani K, Nitta A, Okuda T. Tannins and Related Polyphenols of Melastomataceous Plants: V: 1) Three New Complex Tannins from *Melastoma malabathricum* L. *Chem Pharm Bull.* 1992;40(7):1727-1732. doi:10.1248/cpb.40.1727
- Dinda B, Saha MK. A flavonol-diglycoside from *Melastoma malabathricum*. *J Indian Chem Soc.* 1988;65(3):209-211.
- Das KK, Kotoky J. A new aliphatic constituent of *Melastoma malabathricum* Linn. *J Indian Chem Soc.* 1988;65(5):385-386.
- Lowry JB. The distribution and potential taxonomic value of alkylated ellagic acids. *Phytochemistry.* 1968;7(10):1803-1813. doi:10.1016/S0031-9422(00)86652-2
- Ventola CL. The antibiotic resistance crisis: causes and threats. *P T J.* 2015;40(4):277-283. doi:Article
- Johnny L, Yusuf UK, Nulit R. Antifungal activity of selected plant leaves crude extracts against a pepper anthracnose fungus, *Colletotrichum capsici* (Sydow) butler and bisby (Ascomycota: Phyllachorales). *African J Biotechnol.* 2011;10(20):4157-4165.
- Che Omar SN, Ong Abdullah J, Khairoji KA, Chin Chin S, Hamid M. Effects of flower and fruit extracts of *Melastoma malabathricum* Linn. on growth of pathogenic bacteria: *Listeria monocytogenes*, *Staphylococcus aureus*, *Escherichia coli*, and *salmonella typhimurium*. *Evidence-based Complement Altern Med.* 2013;2013. doi:10.1155/2013/459089
- Shailasree S, Sampathkumara KK, Niranjana SR, Prakash HS. Bioactive potential of medicinal plants from Western ghats region, India. *J Herbs, Spices Med Plants.* 2014;20(3):221-234. doi:10.1080/10496475.2013.860070
- Alwash MSA, Ibrahim N, Yaacob WA, Din L Bin. Antibacterial, antioxidant and cytotoxicity properties of traditionally used *Melastoma malabathricum* linn

- leaves. *Adv J Food Sci Technol*. 2014;6(1):6-12.
17. Zakaria ZA, Nur Zainol AS, Sahmat A, et al. Gastroprotective activity of chloroform extract of *Muntingia calabura* and *Melastoma malabathricum* leaves. *Pharm Biol*. 2016;54(5):812-826. doi:10.3109/13880209.2015.1085580
 18. Zabidi Z, Wan Zainulddin WN, Mamat SS, et al. Antiulcer activity of methanol extract of *Melastoma malabathricum* leaves in rats. *Med Princ Pract*. 2012;21(5):501-503. doi:10.1159/000337406
 19. Hussain F, Abdulla MA, Noor SM, Ismail S, Ali HM. Gastroprotective effects of *Melastoma malabathricum* aqueous leaf extract against ethanol-induced gastric ulcer in rats. *Am J Biochem Biotechnol*. 2008;4(4):438-441. doi:10.3844/ajbbsp.2008.438.441
 20. Zakaria ZA, Balan T, Mamat SS, Mohtarrudin N, Teh LK, Salleh MZ. Mechanisms of gastroprotection of methanol extract of *Melastoma malabathricum* leaves. *BMC Complement Altern Med*. 2015;15(1). doi:10.1186/s12906-015-0638-z
 21. Ismail Suhaimy NW, Noor Azmi AK, Mohtarrudin N, et al. Semipurified Ethyl Acetate Partition of Methanolic Extract of *Melastoma malabathricum* Leaves Exerts Gastroprotective Activity Partly via Its Antioxidant-Antisecretory-Anti-Inflammatory Action and Synergistic Action of Several Flavonoid-Based Compounds. *Oxid Med Cell Longev*. 2017;2017. doi:10.1155/2017/6542631
 22. Mamat SS, Kamarolzaman MFF, Yahya F, et al. Methanol extract of *Melastoma malabathricum* leaves exerted antioxidant and liver protective activity in rats. *BMC Complement Altern Med*. 2013;13. doi:10.1186/1472-6882-13-326
 23. Kamisan FH, Yahya F, Ismail NA, et al. Hepatoprotective Activity of Methanol Extract of *Melastoma malabathricum* Leaf in Rats. *JAMS J Acupunct Meridian Stud*. 2013;6(1):52-55. doi:10.1016/j.jams.2012.08.002
 24. Balamurugan K, Sakthidevi G, Mohan VR. Anti-inflammatory activity of leaf of *Melastoma malabathricum* L. (Melastomataceae). *Int J Res Ayurveda Pharm*. 2012;3(6):801-802. doi:10.7897/2277-4343.03622
 25. Kumar V, Bhatt PC, Rahman M, et al. *Melastoma malabathricum* Linn attenuates complete Freund's adjuvant-induced chronic inflammation in Wistar rats via inflammation response. *BMC Complement Altern Med*. 2016;16(1). doi:10.1186/s12906-016-1470-9
 26. Karupiah S, Ismail Z. Antioxidative effect of *Melastoma malabathricum* L extract and determination of its bioactive flavonoids from various location in Malaysia by RP-HPLC with diode array detection. *J Appl Pharm Sci*. 2013;3(2):19-24. doi:10.7324/JAPS.2013.30204
 27. Balamurugan K, Nishanthini A, Mohan VR. Anticancer activity of ethanol extract of *Melastoma malabathricum* L. leaf against Dalton's lymphoma. *J Pharm Sci Res*. 2013;5(5):111-114.
 28. Sari NM, Kuspradini H, Amirta R, Kusuma IW. Antioxidant activity of an invasive plant, *Melastoma malabathricum* and its potential as herbal tea product. In: *IOP Conference Series: Earth and Environmental Science*. ; 2018. doi:10.1088/1755-1315/144/1/012029
 29. Danladi S, Wan-Azemin A, Sani YN, et al. Phytochemical screening, antioxidant potential and cytotoxic activity of *Melastoma malabathricum* Linn. from different locations. *Int J Pharm Pharm Sci*. 2015;7(7):408-413.
 30. Verma A, Bhatt PC, Kaithwas G, et al. Chemomodulatory effect of *Melastoma malabathricum* Linn against chemically induced renal

- carcinogenesis rats via attenuation of inflammation, oxidative stress, and early markers of tumor expansion. *Inflammopharmacology*. 2016;24(5):233-251. doi:10.1007/s10787-016-0276-1
31. Baruah S, Brahma D, Upadhyya P. Phytochemical study of some selected medicinal plants and its ethnobotanical importance to the indigenous communities of assam. *Med Plants*. 2018;10(2):145-150. doi:10.5958/0975-6892.2018.00023.0
32. Roslen NA, Alewi NAM, Ahamada H, Rasad MSBA. Cytotoxicity screening of *Melastoma malabathricum* extracts on human breast cancer cell lines in vitro. *Asian Pac J Trop Biomed*. 2014;4(7):545-548. doi:10.12980/APJTB.4.2014C658
33. Idris A, Zulkipli IN, Zulhilmi NR, et al. *Melastoma malabathricum* ethyl acetate fraction induces secondary necrosis in human breast and lung cancer cell lines. *Pharmacogn Mag*. 2017;13(51):S688-S692. doi:10.4103/pm.pm_465_15
34. Nurdiana S, Marziana N. Wound healing activities of *Melastoma malabathricum* leaves extract in Sprague dawley rats. *Int J Pharm Sci Rev Res*. 2013;20(2):20-23.
35. Ab Rahman MR, Abdul Razak F, Mohd Bakri M. Evaluation of wound closure activity of *nigella sativa*, *Melastoma malabathricum*, *Pluchea indica*, and *Piper sarmentosum* extracts on scratched monolayer of human gingival fibroblasts. *Evidence-based Complement Altern Med*. 2014;2014. doi:10.1155/2014/190342
36. Sulaiman MR, Somchit MN, Israf DA, Ahmad Z, Moin S. Antinociceptive effect of *Melastoma malabathricum* ethanolic extract in mice. *Fitoterapia*. 2004;75(7-8):667-672. doi:10.1016/j.fitote.2004.07.002
37. Zakaria ZA, Jaios ES, Omar MH, et al. Antinociception of petroleum ether fraction derived from crude methanol extract of *Melastoma malabathricum* leaves and its possible mechanisms of action in animal models. *BMC Complement Altern Med*. 2016;16(1). doi:10.1186/s12906-016-1478-1
38. Jaios ES, Abdul Rahman S, Siew MC, Kadir AA, Desa MNM, Zakaria ZA. Possible mechanisms of antinociception of methanol extract of *Melastoma malabathricum* leaves. *Brazilian J Pharmacogn*. 2016;26(5):586-594. doi:10.1016/j.bjp.2016.01.011
39. Balamurugan K, Sakthidevi G, Mohan VR. Stimulatory effect of the ethanol extract of *Melastoma malabathricum* L. (*Melastomataceae*) leaf on the reproductive system of male albino rats. *J Appl Pharm Sci*. 2013;3(2):160-165. doi:10.7324/JAPS.2013.30228
40. Khare CP. *Melastoma malabathricum* Linn. In: *Indian Medicinal Plants*. ; 2007:1-1. doi:10.1007/978-0-387-70638-2_988
41. Zakaria ZAA, Raden Mohd. Nor RNSNS, Hanan Kumar G, et al. Antinociceptive, anti-inflammatory and antipyretic properties of *Melastoma malabathricum* leaves aqueous extract in experimental animals. *Can J Physiol Pharmacol*. 2006;84(12):1291-1299. doi:10.1139/Y06-083
42. Jaganath IB, Ng LT. *Herbs: The Green Pharmacy of Malaysia*. Vinpress; 2000.
43. Lin KW. Ethnobotanical study of medicinal plants used by the Jah Hut peoples in Malaysia. *Indian J Med Sci*. 2005;59(4):156-161. doi:10.4103/0019-5359.16121
44. Burkill IH. A Dictionary of the Economic Products of the Malay Peninsula. *Nature*. 1936;137(3459):255. doi:10.1038/137255c0
45. Sharma HK, Chhange L, Dolui AK. Traditional medicinal plants in Mizoram, India. *Fitoterapia*. 2001;72(2):146-161. doi:10.1016/S0367-326X(00)00278-1

46. Sajem AL, Gosai K. Traditional use of medicinal plants by the Jaintia tribes in North Cachar Hills district of Assam, northeast India. *J Ethnobiol Ethnomed.* 2006;2. doi:10.1186/1746-4269-2-33
47. Ringmichon CL, Shimpi SN, Gopalkrishnan B. *Ethnomedicinal Investigation on Melastoma Malabathricum Linn. from Manipur.*; 2010.
48. H.N. Thatoi, S.K. Panda, S.K. Rath SKD, Thatoi HN, Panda SK, Rath SK, Dutta SK. Antimicrobial activity and ethnomedicinal uses of some medicinal plants from Similipal Biosphere Reserve, Orissa. *Asian J Plant Sci.* 2008;7(3):260-267. doi:10.3923/ajps.2008.260.267
49. Chuakul W. Medicinal Plants in the Khok Pho District, Pattani Province (Thailand). *Thai J Phytopharm.* 2005;12(2):23-45.
50. Mohd. Joffry S, Yob NJ, Rofiee MS, et al. Melastoma malabathricum (L.) smith ethnomedicinal uses, chemical constituents, and pharmacological properties: A review. *Evidence-based Complement Altern Med.* 2012;2012. doi:10.1155/2012/258434
51. Roosita K, Kusharto CM, Sekiyama M, Fachrurozi Y, Ohtsuka R. Medicinal plants used by the villagers of a Sundanese community in West Java, Indonesia. *J Ethnopharmacol.* 2008;115(1):72-81.
52. Jung JM, Kwon SH, Noh H, Han DC, Jeon JS, Jin SY. Acute interstitial nephritis following kudzu root juice ingestion. *Clin Nephrol.* 2013;80(4):298-300. doi:10.5414/cn107850
53. Umali-Stuart G, Stiuart-Santiago A. *Phillippine Medicinal Plants: Family Melastomaceae.* 2010.
54. Rahmatullah M, Hossan S, Hanif A, et al. *Ethnomedicinal Applications of Plants by the Traditional Healers of the Marma Tribe of Naikhongchhari, Bandarban District, Bangladesh.*; 2009.
55. Vankar PS, Tiwari V, Singh LW, Potsangbam L. Sonicator dyeing of cotton fabric and chemical characterization of the colorant from Melastoma malabathricum. *Pigment Resin Technol.* 2009;38(1):38-42. doi:10.1108/03699420910923562
56. Wong K-C, Ali DMH, Boey P-L. Chemical constituents and antibacterial activity of Melastoma malabathricum L. *Nat Prod Res.* 2012;26(7):609-618. doi:10.1080/14786419.2010.538395
57. Ibrahim N, S. N, A.W. NZ, I.B. A. Cytotoxicity and antiviral activity of Melastoma malabathricum extracts. *Malaysian Appl Biol.* 2008;37(2):53-55.
58. Ali DMH, Keng CW, Boey PL. *Triterpenoids, Glycolipids and Flavonoids of Melastoma Malabathricum: Isolation, Spectrometric Characterization and Antibacterial Activity.* VDM Publishing; 2010.
59. Susanti D, Sirat HM, Ahmad F, Ali RM, Aimi N, Kitajima M. Antioxidant and cytotoxic flavonoids from the flowers of Melastoma malabathricum L. *Food Chem.* 2007;103(3):710-716. doi:10.1016/j.foodchem.2006.09.011
60. Mazura MP, Susanti D, Rasadah MA. Anti-inflammatory action of components from Melastoma malabathricum. *Pharm Biol.* 2007;45(5):372-375. doi:10.1080/13880200701214797
61. Sirat HM, Susanti D, Ahmad F, Takayama H, Kitajima M. Amides, triterpene and flavonoids from the leaves of Melastoma malabathricum L. *J Nat Med.* 2010;64(4):492-495. doi:10.1007/s11418-010-0431-8
62. Nuresti. Chemical components of Melastoma malabathricum. *ACGC Chem Res Commun.* 2003;16:28.
63. Pongprom N, Khanapan C, Tosert P. Chemical constituents of Melastoma malabathricum Linn. In: *The 29th Congress on Science and Technology of Thailand.* ; 2003:130.
64. Faravani M. The population biology

- of Straits Rhododendron (*Melastoma malabathricum* L.). 2008.
65. Koay SS. *Establishment of Cell Suspension Culture of Melastoma Malabathricum L. for the Production of Anthocyanin.*; 2008.
 66. Halim SZ, Zakaria ZA, Omar MH, Mohtarrudin N, Wahab IRA, Abdullah MNH. Synergistic gastroprotective activity of methanolic extract of a mixture of *Melastoma malabathricum* and *Muntingia calabura* leaves in rats. *BMC Complement Altern Med.* 2017;17(1). doi:10.1186/s12906-017-1992-9
 67. Hamid HA, Ramli ANM, Zamri N, Yusoff MM. UPLC-QTOF/MS-based phenolic profiling of Melastomaceae, their antioxidant activity and cytotoxic effects against human breast cancer cell MDA-MB-231. *Food Chem.* 2018;265:253-259. doi:10.1016/j.foodchem.2018.05.033
 68. Yusriandi R, Fauzul ZO, Winata NP, et al. Effect of karamunting fruit juice (*Melastoma malabathricum* L.) to advanced glycation end-products (AGEs) and lipid profile as advanced complications of diabetes mellitus. *J Eng Appl Sci.* 2017;12(2):186-194. doi:10.3923/jeasci.2017.171.175