

Review Article

The Potential of Plants of Family *Fabaceae* with Emphasis on Putri Malu Medicinal Plant 'Mimosa Pudica' (*Fabaceae*) as an Antimalarial & an Insecticide for Malaria Vectors: A Review

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A B S T R A C T

Background: Due to the prevalence of malaria worldwide and cases of resistance to malaria drugs, finding new drug candidates is important in malaria control. Due to its traditional use and phytochemical content, this review was conducted on the medicinal plant *Mimosa pudica* and its family, Fabaceae.

Method: This review collected original articles in online databases using several keywords combined with boolean operators. The articles about the antimalarial and insecticidal effects of *Mimosa pudica* and other *Fabaceae* species were included in the study.

Results: Forty-two articles described 45 species from the *Fabaceae* family exhibiting antimalarial and/ or insecticidal potential including *Mimosa pudica*. The studies showed that crude extract of *M. pudica* showed activity against *P. falciparum* or *P. berghei* and insecticidal activity against *Anopheles subpictus* and *Anopheles stephensi*. More advanced studies were carried out on other *Fabaceae* species, evaluating their activity with crude extracts and fractions, isolated compounds, and silver nanoparticles (AgNPs).

Conclusions: The most promising antiplasmodial activity of *M. pudica* was shown by aqueous, methanol, and water/ methanol extracts from the aerial part against *P. falciparum* FCR-3 strain. In addition, aqueous or ethanolic extracts from the leaves of *M. pudica* revealed their potential against *A. subpictus* and *A. stephensi*.

Keywords: *Anopheles*, Antimalaria, Antiplasmodial, *Fabaceae*, Insecticidal, *Mimosa Pudica*, *Plasmodium*

Introduction

Malaria is one of the most prevalent infectious diseases in the tropical world, with a total of 228 million global cases in 2018, according to World Health Organization (WHO).¹ WHO guidelines explain how to control malaria by controlling the malarial pathogen (*Plasmodium*) and its vector (*Anopheles*) with Artemisin-based Combination Therapy (ACT), Long-Lasting Insecticidal Nets (LLINs) and Indoor Residual Spraying of Insecticides (IRN).^{2,4} However, *Plasmodium* and *Anopheles* developed resistance to drugs used against them. Despite that, ACT, LLINs, and IRS remained to be the first choice of control of WHO as they remained the most effective method, and the resistance towards them has not spread widely.^{4,10} Still, some action is needed to prevent further spreading and malignant resistance.

Finding new drug candidates is vital to overcoming malaria resistance. Medicinal plants are an essential source of new drug candidates. The antimalarial drug to date, artemisinin, is also sourced from medicinal plants (*Artemisia annua*).¹¹ *Mimosa pudica* contained phytochemicals that showed antimalarial or insecticidal activities.^{12,15} This study aims to gain information regarding the potential effects

of this plant from published original research articles. Furthermore, this plant grows quickly, is found abundantly across tropical areas, and is even capable of growing in less fertile soil, making it easy to be cultivated.^{12,16,17} This plant has been used as a traditional medicine in many countries.¹² To the best of our knowledge, a specific review study on this particular topic has not been done before, making this study the first recorded review study on this topic.

Review Method

This study was conducted on the species *Mimosa pudica* and its family, *Fabaceae*. Information on the insecticidal and antiplasmodial effects of this plant has been presented in two tables.

The search strategy used several keywords combined with boolean operators in online databases, as stated in Table 1. This review includes original articles about the effects of *Mimosa pudica* or other *Fabaceae* species towards *Plasmodium* or *Anopheles*. On the other hand, a review article, an article that could not be fully accessed, or an article published before 2016 were excluded from this study, except for the articles about *Mimosa pudica*, since the study about this plant is limited.

Table 1. Keywords and Databases Used

Keywords	Databases
("fabaceae" OR "peas" OR "legumes" OR "mimosa pudica" OR "lajwanti" OR "shameplant") AND ("antimalarial" OR "antiplasmodial" OR "plasmodium" OR "anopheles")	Google Scholar
	Science Direct
	Medline
	Springer Links
	Cochrane
("fabaceae" OR "polong" OR "mimosa pudica" OR "putri malu") AND ("antimalarial" OR "antiplasmodial" OR "plasmodium" OR "anopheles")	Google Scholar Indonesia
	Portal Garuda

Result

Forty-two articles met the inclusion and exclusion criteria (Figure 1). Thirty-four focused on the effects on *plasmodium*, four of which revealed the effects of *Mimosa pudica*, and another thirty showed the effects of *Fabaceae* species. Eight other articles focused on the effect on *Anopheles*, three of which revealed the effects of *Mimosa pudica*, and five articles showed the effects of other *Fabaceae* species. Original articles found in other languages (Bahasa-Indonesian language) were also used as found in Google Scholar. Common papers in different languages were not used as they were considered duplicates.

We reported 45 *Fabaceae* species including *Mimosa pudica*. Of these, 37 species were evaluated for their antimalarial potency, six for their insecticidal activity, and two species, *Cassia occidentalis* and *Mimosa pudica*, for both activities.

Tables 2 and 3 present a summary of the latest research on the effect of plant extracts of *M. pudica* and other *Fabaceae* species on *Plasmodium* and *Anopheles* malaria vectors. The widely used parts of this species are the roots, leaves, and bark. *M. pudica* samples were generally used in the form of crude extracts with ethanol, methanol, or water as solvents. Mean while, other *Fabaceae* species were used as crude extracts with water, methanol, hexane, ethanol, and

ethyl acetate as solvents. The most identified metabolites were alkaloids, flavonoids, tannins, and saponins. Other metabolites such as quinone, terpenoid, phenol, chloroform, and glycoside groups were also identified. Of these, 35 active compounds from 10 species were isolated (Table 3).

The studies on *Burkea africana*, *Ormocarpum cochinchinense*, and *Mimosa pudica* are the only studies that explained the mechanism of its metabolites.

However, the pathway stated in their article was gathered from other literature and was not discovered in their investigations. Several studies have used plant-based-synthesised silver nanoparticles (AgNPs), in addition to using natural products to increase drug effectiveness. The materials used were from *Indigofera oblongifolia*, *Ormocarpum cochinchinense*, and *Mimosa pudica* (Tables 2 and 3).

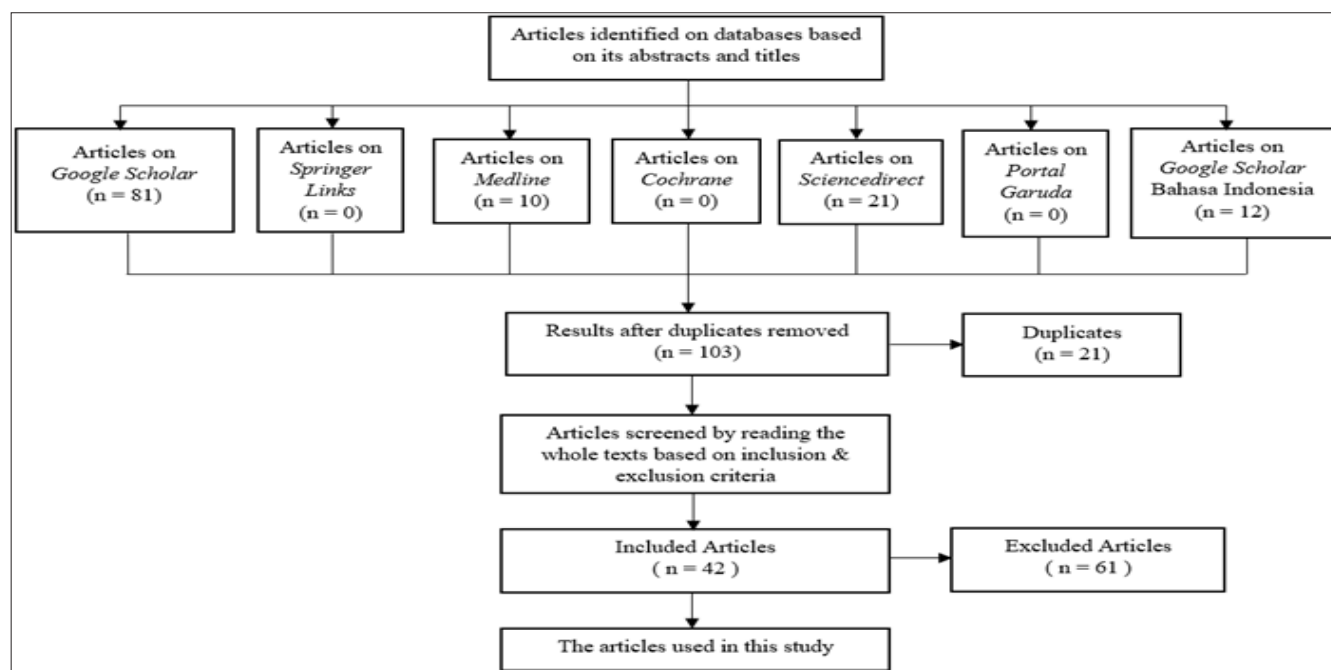


Figure 1. PRISMA Flow Diagram of the Process of Article Selection

Table 2. Results from Studies on *Mimosa Pudica*

S. No.	Authors (Year)	Target Species	Study Design	Sample Used	Results	Active Phytochemicals	Additional Information
1.	N Aarthi & K Murugan (2011) ¹⁸	<i>Plasmodium berghei</i>	In vivo	Ethanol extract from leaves	% Suppression: 38.3% Dose: 800 mg/kg/day	Alkaloids and flavonoids	N/A
2.	N Singh et al. (2015) ¹⁹	CQS <i>Plasmodium falciparum</i> 3D7 strain	Ex vivo	Ethanol extract from whole plant	IC50: 51 µg/mL	N/A	N/A
		CQR <i>Plasmodium falciparum</i> INDO strain	Ex vivo	Ethanol extract from whole plant	N/A	N/A	N/A
3.	OO Ogbole, PA Segun, & PS Fasinu (2018) ²⁰	CQS <i>Plasmodium falciparum</i> D6 strain	In vitro	Methanol extract from leaves	% Suppression: 35% Dose: 15.9 µg/mL	N/A	N/A

4.	QL Tran et al. (2003) ²¹	<i>Plasmodium falciparum</i> FCR-3 strain	In vitro	Aerial part-derived: 1. Aqueous extract 2. Methanol extract 3. Aqueous: Methanol extract (1:1)	EC50: 4.4 µg/mL 6.2 µg/mL 4.0 µg/mL	N/A	This is the first recorded study on the antimalarial potency of <i>M. pudica</i>
5.	S Marimuthu et al. (2011) ²²	<i>Anopheles subpictus</i> larvae	In vivo	Leaves-derived: 1. Aqueous extract 2. AgNPs	LC50: 45.82 µg/mL 13.9 µg/mL	Silver AgNPs, terpenoids, flavonoids	Using AgNPs rather than extraction, fractionation, or isolation of active compounds
6.	N Aarthi et al. (2011) ²³	<i>Anopheles stephensi</i> larvae Instar I	In vivo	Ethanol extract from leaves	LC50: 0.723 µg/mL	Alkaloids, flavonoids, phenols, steroids	N/A
		<i>Anopheles stephensi</i> larvae Instar II	In vivo	Ethanol extract from leaves	LC50: 1.150 µg/mL	Alkaloids, flavonoids, phenols, steroids	N/A
		<i>Anopheles stephensi</i> larvae Instar III	In vivo	Ethanol extract from leaves	LC50: 1.540 µg/mL	Alkaloids, flavonoids, phenols, steroids	N/A
		<i>Anopheles stephensi</i> larvae Instar IV	In vivo	Ethanol extract from leaves	LC50: 2.073 µg/mL	Alkaloids, flavonoids, phenols, steroids	N/A
		<i>Anopheles stephensi</i> pupa	In vivo	Ethanol extract from leaves	LC50: 2.835 µg/mL	Alkaloids, flavonoids, phenols, steroids	N/A
7.	S Amilah & E Fitria (2015) ²⁴	<i>Anopheles sp.</i> larvae Instar III	In vivo	Ethanol extract from leaves	LC50: 1.88 g/L	Alkaloids, saponins, flavonoids	Mechanism of action stated, derived from other literatures

EC50: Half Maximal Effective Concentration, IC50: Half Maximal Inhibitory Concentration, LD50: Half Maximal Lethal Dose, N/A: Not Available

ED50: Half Maximal Effective Dose, IC50: Half Maximal Inhibitory Concentration, LD50: Half Maximal Lethal Dose, N/A: Not Available

Table 3. Results from Studies on other Fabaceae Species

S. No.	Fabaceae Species	Target Species	Study Design	Sample Used	Results	Active Phytochemicals	Additional Information	References
1.	Acacia karroo	<i>CQS Plasmodium falciparum</i> 3D7 strain	In vitro	Leaves-derived: 1. Methanol extract 2. Fraction 4	IC50: 9 µg/mL 8.2 µg/mL	Phenol, 2,4-bis(1,1-dimethylethyl)-; benzenepropanoic acid, 3,5-bis(1,1-dimethylethyl)-4-hydroxy-; mome inositol; and 3',5'-dimethoxyacetophenone	N/A	C Sachdeva et al. (2020) ²⁵
		CQR I falciparum INDO strain	In vitro	Leaves-derived: 1. Methanol extract 2. Fraction 4	IC50: 14 µg/mL 4.8 µg/mL			
		<i>Plasmodium berghei</i> ANKA strain	In vivo	Methanol extract from leaves	% Suppression: 57%			
2.	Acacia nilotica	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from bark	IC50 = 208.33 µg/mL	Tannins, alkaloids, saponins	N/A	M Ohashi et al. (2018) ²⁶
		<i>Plasmodium berghei</i>	In vivo	Ethanol extract from bark	% Suppression: 26.35%	N/A	The method used was different from other in vivo studies (not 4-day suppressive test)	NT Dabo et al. (2016) ²⁷
		Artesunate sensitive <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from: 1. Bark 2. Pods 3. Leaves	IC50: 4.28 µg/mL IC50: 4.16 µg/mL IC50: 1.29 µg/mL	N/A	N/A	MB Sadiq et al. (2017) ²⁸
3.	Afzelia africana	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from bark	IC50: 222.36 µg/mL	Alkaloids, tannins, flavonoids, saponins	N/A	M Ohashi et al. (2018) ²⁶

4.	Baphia nitida	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from bark	IC50: > 1000 µg/mL	Tannins, flavonoids, saponin glycosides	N/A	M Ohashi et al. (2018) ²⁶
5.	Bauhinia rufescens	<i>Plasmodium berghei</i> ANKA strain	In vivo	Extract from leaves	% Suppression: 50.6%	Flavonoids, tannins, triterpenes, saponins, and alkaloids	Means of extraction not mentioned	LN Bonkian et al. (2018) ²⁹
6.	Burkea africana	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Bark-derived: 1. Ethanol extract 2. Dichloro-methane fraction	IC50: 28.69 µg/mL 6.44 µg/mL	N/A	The fraction showed a promising effect on the resistant strain, which means that it has a different mechanism of action from the drugs.	IC Ezenyi et al. (2021) ³⁰
		Multidrug resistant <i>Plasmodium falciparum</i> W2 mef strain	In vitro	Bark-derived: 1. Ethanol extract 2. Dichloro-methane fraction	IC50: 25.19 µg/mL 6.3 µg/mL			
7.	Caesalpenia bonducella	<i>Plasmodium berghei</i> ANKA strain	In vivo	Dichloro-methane extract from roots	% Suppression: 55.96%	Less to medium polar compounds		RS Nondo et al. (2016) ³¹
8.	Cassia alata	CQS <i>Plasmodium falciparum</i> D10 strain	In vitro	Extract CH ₂ Cl ₂ / MeOH (1:1) from leaves	IC50: 7.02 µg/mL	N/A	N/A	O Da et al. (2016) ³²
		<i>Plasmodium berghei</i> ANKA strain	In vivo	Extract CH ₂ Cl ₂ / MeOH (1:1) from leaves	% Suppression: 45.2%			
		<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from leaves	IC50: 57.60 µg/mL	Flavonoids and glycosides	N/A	M Ohashi et al. (2018) ²⁶

9.	Cassia occidentalis	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from leaves, seeds, and whole plant	IC50: > 1000 µg/mL (for all parts)	Anthraquinone, flavonoids	N/A	M Ohashi et al. (2018) ²⁶
		Anopheles stephensi Liston eggs	In vivo	Leaves-derived: 1. Hexane extract 2. Ethyl acetate extract 3. Methanol extract	Egg mortality rate: 85% 96.1% 94.9%	N/A	N/A	V Raja et al. (2016) ³³
10.	Cassia podocarpa	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from leaves	IC50: > 1000 µg/mL	Anthraquinone	N/A	M Ohashi et al. (2018) ²⁶
11.	Cassia siamea	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from leaves	IC50: > 1000 µg/mL	Anthraquinones, flavonoids	N/A	M Ohashi et al. (2018) ²⁶
12.	Cassia sieberiana	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from roots & leaves	IC50: 432.48 µg/mL (roots) IC50: > 1000 µg/mL (leaves)	Galactosides, flavonoids Flavonoids, alkaloids	N/A	M Ohashi et al. (2018) ²⁶
13.	Cassia tora	Anopheles gambiae larvae Instar III-IV	In vivo	Seeds-derived: 1. Ethyl acetate extract 2. Fraction 3 3. Fraction 4 4. Aurantio-obtusin isolated compound 5. Obtusin-isolated compound	LD50: 2.5 µg/mL LD50: 5 µg/mL LD50: 7 µg/mL LD50: 10 µg/mL LD50: 10.2 µg/mL	Aurantio-obtusin and obtusin	Obtusin may not be the compound responsible for the effect in the extracts	VC Mbatchou et al. (2017) ³⁴

14.	Copaifera reticulata	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Oleoresin	IC50: 2.54 µg/mL	β-caryophyllene &	Using oleoresin rather than extraction, fractionation, or isolation of active compounds	GA de Souza et al. (2016) ³⁵
		CQR <i>Plasmodium falciparum</i> W2 strain	In vitro	Oleoresin	IC50: 1.66 µg/mL	β-bisabolene		
		<i>Plasmodium berghei</i> ANKA strain	In vitro	Oleoresin	% Suppression: 96%			
15.	Cylicodiscus gabunensis	<i>Plasmodium falciparum</i> Dd2 strain	In vitro	Bark-derived: 1. Ethanol extract 2. Hexane extract 3. Ethyl acetate fraction 4. Butanol fraction 5. Aqueous fraction 6. CGEBU-F10 fraction 7. CGEBU-F10-7 fraction	IC50: 20.8 µg/mL IC50: 32.1 µg/mL IC50: 16.1 µg/mL IC50: 10.4 µg/mL IC50: 25.8 µg/mL IC50: 6.5 µg/mL IC50: 4.7 µg/mL	Gallic acid, ethyl gallate, benzoic acids	Fractionation and isolation of active compound has been done	O Aldulaimi et al. (2017) ³⁶
16.	Dalbergia katangensis	<i>Plasmodium berghei</i> ANKA MRA 311 strain	In vivo	Leaves-derived: 1. Methanol extract 2. Aqueous extract	% Suppression: 74.81% 73.38%	Terpenoids, steroids, polyphenols, flavonoids, anthraquinones	N/A	BC Valentin et al. (2020) ³⁷
17.	Detarium microcarpum	CQS <i>Plasmodium berghei</i>	In vivo	Methanol extract from leaves	% Suppression: 80.92%	Alkaloids, saponins, flavonoids, triterpenes, tannins	N/A	AR Abdullahi et al. (2020) ³⁸
18.	Dialium angolense	<i>Plasmodium berghei</i> ANKA MRA 311 strain	In vivo	Leaves-derived: 1. Methanol extract 2. Aqueous extract	% Suppression: 70.81% 70.38%	Anthraquinones, flavonoids, polyphenols, terpenoids	N/A	BC Valentin et al. (2020) ³⁹

19.	Dichrostachys cinerea	CQS <i>Plasmodium berghei</i> NK-65 strain	In vivo	Bark-derived: 1. Ethanol extract 2. Chloroform fraction 3. Ethyl acetate fraction 4. Buthanol fraction	% Suppression: 59.66% 74.71% 23.41% 70.38%	Phenols, flavonoids, tannins	N/A	LA Fadipe et al. (2020) ⁴⁰
		CQS <i>Plasmodium falciparum</i> D6 strain	In vitro	Whole stem-derived: 1. Dichloromethane extract 2. Methanol extract Stem bark-derived: 1. Dichloromethane extract 2. Methanol extract	IC50: 11.47 µg/mL 2.96 µg/mL IC50: 2.37 µg/mL > 1000 µg/mL	Steroids, flavonoids, saponins, cardiac glycosides, flavonoids, tannins, triterpenoids, saponins	N/A	PA Kweyamba et al. (2019) ⁴¹
		CQR <i>Plasmodium falciparum</i> Dd2 strain	In vitro	Whole stem-derived: 1. Dichloromethane extract 2. Methanol extract Stem bark-derived: 1. Dichloromethane extract 2. Methanol extract	IC50: > 1000 µg/mL > 1000 µg/mL IC50: 11.92 µg/mL > 1000 µg/mL	Steroids, flavonoids, saponins, cardiac glycosides, flavonoids, tannins, triterpenoids, saponins	N/A	PA Kweyamba et al. (2019) ⁴¹
		<i>Plasmodium berghei</i>	In vivo	Dichloromethane extract from bark	% Suppression: 53.12 %	Steroids, flavonoids, saponin	N/A	PA Kweyamba et al. (2019) ⁴¹
20.	Dipteryx Lacunifera	CQR <i>Plasmodium falciparum</i> W2 strain	In vitro	Seed-derived: 1. Isolated isoliquiritigenin 2. Isolated 6,3',4'-trihydroxyflavone 3. Diethyl ether fraction	% Suppression: 81.25% 88.98% 89.46%	Isoliquiritigenin 6,3',4'-trihydroxyflavone	Suspected active compounds have been isolated	LS Alexandre et al. (2020) ⁴²

21.	Distemonanthus benthamianus	<i>Plasmodium berghei</i> ANKA strain	In vivo	Ethanol extract from bark	% Suppression: 56.43%	Tannins, glycosides, saponins, alkaloids, terpenes, flavonoids, coumarins	N/A	F Ayisi et al. (2021) ⁴³
22.	Eriosema montanum	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Root-derived: 1. Extract CH ₂ Cl ₂ / MeOH Root-derived isolation of: 2. Eucomic acid 3. 7-O -glucopyranosyl-isopruneitin 4. Genistin 5. Malonyl genistin 6. Isopruneitin 7. Isoluteolin 8. Genistein	IC50: 17.68 µg/mL IC50: 0.057 µg/mL 0.113 µg/mL 7.867 µg/mL > 10 µg/mL 0.042 µg/mL 0.121 µg/mL 7.736 µg/mL	Eucomic acid, isoflavonoid derivates (7-O -glucopyranosyl-isopruneitin, genistin, malonyl genistin, isopruneitin, isoluteolin, genistein)	Suspected active compounds have been isolated	JC Tomani et al. (2021) ⁴⁴
23.	Erythrina fusca	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Bark-derived: 1. Methanol extract 2. nHexane fraction 3. CHCl ₃ fraction 4. Aqueous fraction	IC50: 13.64 µg/mL 21.44 µg/mL 22.55 µg/mL 4.94 µg/mL	Phaseolin, phytol, β-amyrin, lupeol, and stigmaterol	Fractionation has been done	SA Sazed et al. (2021) ⁴⁵
		CQR <i>Plasmodium falciparum</i> Dd2 strain	In vitro	Bark-derived: 1. Methanol extract 2. nHexane fraction 3. CHCl ₃ fraction 4. Aqueous fraction	IC50: 8.22 µg/mL 4.88 µg/mL 13.77 µg/mL 18.77 µg/mL	Phaseolin, phytol, β-amyrin, lupeol, and stigmaterol	Fractionation has been done	SA Sazed et al. (2021) ⁴⁵
24.	Erythrina schliebenii	<i>Plasmodium berghei</i> ANKA strain	In vivo	Aqueous extract from bark	% Suppression: 28.64%	N/A	N/A	RS Nondo et al. (2016) ³¹
25.	Indigofera amoxyllum	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethyl acetate extract from: 1. Leaves 2. Bark	IC50: > 50 µg/mL > 50 µg/mL	N/A	N/A	A Ledoux et al. (2018) ⁴⁶

26.	Indigofera oblongifolia	<i>Plasmodium chabaudi</i>	In vivo	AgNP from methanol extract derived from leaves	% Suppression 99%	N/A	Outstanding result near the control's value from AgNP	S Al-Quraishy et al. (2020) ⁴⁷
27.	Indigofera spicata	<i>Plasmodium berghei</i> ANKA strain	In vivo	Methanol extract from roots	% Suppression: 53.42%	Alkaloids, flavonoids, tannins	N/A	EM Birru et al. (2017) ⁴⁸
28.	Kotschya speciosa	Anopheles gambiae larvae	In vivo	Ethanol extract from: 1. Leaves 2. Roots	LC50: 75.83 µg/mL 252.03 µg/mL	Phenols	N/A	I Daniel et al. (2020) ⁴⁹
29.	Kotschya strigosa	Anopheles gambiae larvae	In vivo	Ethanol extract from: 1. Leaves 2. Roots	LC50: 37.08 µg/mL 237.31 µg/mL	Phenols	N/A	I Daniel et al. (2020) ⁴⁹
30.	Kotschya thymodora	Anopheles gambiae larvae	In vivo	Ethanol extract from: 1. Leaves 2. Roots	LC50: 16.35 µg/mL 53.35 µg/mL	Phenols	N/A	I Daniel et al. (2020) ⁴⁹
31.	Kotschya uguenensis	Anopheles gambiae larvae	In vivo	Leaves-derived: 1. Ethanol extract 2. Isolated ent-halim-1(10)-ene-15-oic acid 3. Isolated 3-O-methyl-D-chiro-inositol	LC50: 94.01 µg/mL 30.05 µg/mL 80.73 µg/mL	ent-halim-1(10)-ene-15-oic acid 3-O-methyl-D-chiro-inositol	Suspected active compounds have been isolated	B Samwel et al. (2019) ⁵⁰
32.	Mezoneuron benthamianum	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Leaves-derived: 1. Hydroethanolic extract 2. Precipitate 3. Isolated ethyl gallate 4. Isolated quercetin 5. Isolated 13b-OH-pheophorbide a	IC50: 32.6 µg/mL 6.4 µg/mL 6.2 µg/mL 9.5 µg/mL 5.1 µg/mL	Ethyl gallate, quercetin, 13b-OH-pheophorbide A	Suspected active compounds have been isolated	O Jansen et al. (2017) ⁵¹

33.	Mucuna pruriens	CQS <i>Plasmodium berghei</i> NK - 65 strain	In vitro	Ethanol extract from leaves	Significant parasitaemia decrease	n-hexadecanoic acid	Not using the same results as others	OE Ezim et al. (2021) ⁵²
		<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Seeds-derived: 1. Ethanol extract 2. Aqueous extract	%Viability: 63.80 % 65.86 %	Phenolic compounds	Not using the same results as others	MA Jimoh et al. (2020) ⁵³
34.	Mundulea sericea	CQR <i>Plasmodium falciparum</i> W2 strain	In vitro	CH ₂ Cl ₂ / MeOH extract from roots. Leaves-derived isolated: 1. Lupinifolinol 2. Lupinifolin 3. Mundulinol	IC50: 0.6 µg/mL IC50: 2.0 µM 12.1 µM 5.9 µM	Lupinifolinol Lupinifolin Mundulinol	Isolation of suspected active compounds has been done.	C Chepkirui et al. (2021) ⁵⁴
		CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	CH ₂ Cl ₂ / MeOH extract from roots. Leaves-derived isolated: 1. Lupinifolinol 2. Lupinifolin 3. Mundulinol	IC50: 1.8 µg/mL IC50: 6.6 µM 3.6 µM 2.4 µM	Lupinifolinol Lupinifolin Mundulinol	Isolation of suspected active compounds has been done.	C Chepkirui et al. (2021) ⁵⁴
35.	Ormocarpum cochinchinense	Anopheles stephensi	In vivo	Aqueous extract AgNP from aqueous extract	LC50:164.72 µg/mL LC50:10.43 µg/mL	AgNPs	Mechanism of action of AgNPs is stated	M Govindarajan & G Benelli (2016) ⁵⁵
36.	Parkia clappertoniana	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from: 1. Leaves 2. Bark	IC50: 501.23 µg/mL > 1000 µg/mL	Saponin, flavonoid, tannins	N/A	M Ohashi et al. (2018) ²⁶
37.	Piliostigma thonningii	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from: 1. Leaves 2. Bark	IC50: > 1000 µg/mL 514.63 µg/mL	Tannins, alkaloids, flavonoids	N/A	M Ohashi et al. (2018) ²⁶

38.	Prosopis juliflora	<i>Plasmodium berghei</i> NK-65 strain	In vivo	Chloroform extract from: 1. Leaves 2. Pods	% Suppression: 35% 35%	Julifloridine & juliprosopine	Suspected active compounds identified	R Batista et al. (2018) ⁵⁶
39.	Pterocarpus erinaceus	<i>Plasmodium berghei</i> ANKA strain	In vivo	Methanol extract from leaves	% Suppression: 75%	N/A	N/A	O Noufou et al. (2016) ⁵⁷
40.	Pterocarpus santalinoides	<i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from bark	IC50: 907.52 µg/mL	Alkaloids, flavonoids, tannins	N/A	M Ohashi et al. (2018) ²⁶
41.	Senna occidentalis	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Leaves-derived: 1. Hexane extract 2. Methanol extract 3. Aqueous extract	IC50 & % Suppression: 3.47 µg/mL & 75.52% 3,79 µg/mL & 71.60 % 4.03 µg/mL & 56.94 %	Anthraquinones phenols, tannins, alkaloids, flavonoids	N/A	AM Daskum et al. (2019) ⁵⁸
42.	Senna singueana	CQS <i>Plasmodium berghei</i> ANKA strain	In vivo	Ethanol extract from leaves	% Suppression: 47.32%	N/A	Showed great adjuvant properties when tested alongside chloroquine	MG Hiben et al. (2016) ⁵⁹
43.	Sophora denudata	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethyl acetate Extract from: 1. Leaves 2. Barks	IC50: > 50 µg/mL 17.88 µg/mL	N/A	N/A	A Ledoux et al. (2018) ⁴⁶
44.	Tamarindus indica	CQS <i>Plasmodium falciparum</i> 3D7 strain	In vitro	Ethanol extract from: 1. Leaves 2. Barks	IC50: > 1000 µg/mL > 1000 µg/mL	Saponins, tannins, glycoside, phenols, flavonoids	N/A	M Ohashi et al. (2018) ²⁶

Discussion

This review presents information in 2 groups: (1) studies related to the plasmodial and insecticidal activity of *Mimosa pudica*, and (2) from other species of the *Fabaceae* family. The information obtained includes the part of the plant used, the type of extract/ fraction/ isolated compound used, the type of study conducted (in vitro, in vivo, or ex vivo), and the toxicity parameters (IC50, EC50, LC50, LD50, or percentage suppression) against *Plasmodium* sp. or *Anopheles* sp. Isolation and further purification of the extract are needed to study a specific effect of a medicinal plant because of its phytochemical properties.⁶⁰

Antimalarial Potency of *Mimosa Pudica*

Initial studies on *Mimosa pudica* were carried out using aqueous, methanol, and aqueous/ methanol extracts of aerial part-derived for antiplasmodial effect against *Plasmodium falciparum* FCR-3 strain. The methanol extract from leaves of *M. pudica* was also investigated against *P. falciparum* strain D6. The ethanol extract has an antiplasmodial effect against *P. berghei* and *P. falciparum* 3D7 or INDO strains. Among all the tests, aqueous, methanolic, and water/ methanol extracts of aerial part-derived showed the most promising results against *P. falciparum* FCR-3 strain, considering its IC50 value was less than 10 µg/mL.^{30,59} The phytochemical screening of ethanolic extract revealed the presence of flavonoids and alkaloids. Phytochemical screening of other extracts or further purification of active compounds has not been carried out (Table 2).

Other *Fabaceae* families used leaves, bark, pods, roots, stems, and whole plants to investigate their antiplasmodial activity against *P. falciparum*, *P. chabaudi*, or *P. berghei*. The methanol extracts of *Acacia karroo*, *Dalbergia katangensis*, *Detarium microcarpum*, *Dialium angolense*, *Dichrostachys cinerea*, *Erythrina fusca*, *Indigofera spicata*, *Pterocarpus erinaceus*, and *Senna occidentalis* have shown promising results against *P. falciparum* or *P. berghei* with IC50 less than 10 µg/mL or suppressing percentage of more than 50%.^{3,59} The most promising result is methanol extract from leaves of *Dichrostachys cinerea* against *P. falciparum* D6 strain with IC50 2.37 µg/mL and *Detarium microcarpum* that suppressed 80.92% *P. berghei* (Table 3). The ethanol extracts from *Acacia nilotica*, *Dichrostachys cinerea*, and *Distemonanthus benthamianus* have shown promising results against *P. falciparum* and *P. berghei*. The most promising result has been shown by ethanol extract from leaves of *Acacia nilotica* against Artesunate sensitive *Plasmodium falciparum* strain 3D7 with IC50 value of 1.29 µg/mL (Table 3).

Aqueous extract from leaves of *Dalbergia* or *Dialium angolense* showed plasmodial activity against *P. berghei*

strain ANKA MRA 311 with 70% suppression. Still, the extract from the bark of *Erythrina schliebenii* is less active against *P. berghei* strain ANKA MRA 311 with 28.64% suppression. Aqueous leaf extract also has good activity against *P. falciparum* 3D7 strain on *Senna occidentalis*. A few other extracts have also revealed good antiplasmodial performance, such as CH₂Cl₂/ MeOH extract from leaves of *Cassia alata* against *P. falciparum* strain D10, dichloromethane extract from the bark of *Dichrostachys cinerea* against *P. berghei*, CH₂Cl₂/ MeOH extract from the roots of *Mundulea sericea* against *P. falciparum* 3D7 strain, and hexane extract from the leaves of *Senna occidentalis* against *P. falciparum* 3D7 strain with IC50 under 10µg/mL (Table 3).

Further purification increased the plasmodial activity many folds, such as the dichloromethane fraction from the bark of *Burkea africana* and the isolated compound from *Eriosema montanum* or *Mezoneuron benthamianum*. The expected results were also shown by the aqueous and n-hexane fraction of *Erythrina fusca* against *P. falciparum* 3D7 and Dd2 strains, respectively. However, further purification did not show any increased activity in the methanol fraction of *Acacia karroo*. It even showed decreased activity in fractions 3-4 and in compounds isolated from the ethyl acetate extract of *Cassia tora*, although the IC50 value was still promising (Table 3).

Isoliquiritigenin and 6,3',4-trihydroxyflavone-isolated compounds were reported to have antiplasmodial activity against *P. falciparum* W2 strain with 80% suppression. In addition, researchers have also succeeded in isolating eucomic acid, 7-O-glucopyranosyl-isoprunitin, isoprunitin, isoluteolin, genistin, genistein, ethyl gallate, quercetin, 13b-OH-pheophorbide a, lupinifolinol, lupinifolin, and mundulinol to fight *P. falciparum* 3D7 or W2 strains with IC50 less than 10 µg/mL (Table 3).

In summary, the methanol and aqueous extract showed good antiplasmodial activity of *M. pudica*. Studies on other *Fabaceae* suggest that methanol extract of *M. pudica* could be evaluated against another *Plasmodium*, as indicated by *Indigofera oblongifolia* against *P. chabaudi* or by *Acacia karroo* against *P. falciparum* 3D7 and INDO strains.^{25,47} Further studies against *P. falciparum* FCR-3 strain can be carried out using fractions or isolated compounds. Unfortunately, antiplasmodial potential against *P. falciparum* has only been reported by *M. pudica* and not by other *Fabaceae* species. On the other hand, an aqueous extract of *M. pudica* could also be tested against *P. berghei* ANKA MRA 311 strain as shown by *Dalbergia katangensis* or *Dialium angolense*.^{37,39} Furthermore, aqueous extract and its fraction of *M. pudica* can be evaluated against *P. falciparum* 3D7 strain as in *Senna occidentalis*.⁵⁸

Insecticidal Potency of *Mimosa Pudica* towards Malarial Vectors

Researchers have reported the insecticidal activity of *Mimosa pudica* against *Anopheles subpictus* and *Anopheles stephensi*. The action of the aqueous extract of *M. pudica* against *A. subpictus* is lower than their ethanol extract against *A. stephensi* (LC50 45,82 µg/mL and 0.723 – 2.835 µg/mL, respectively) (Table 2). The aqueous extract of *O. cochinchinense* (*Fabaceae* family) also showed their activity against *A. stephensi*, but the action is lower than *M. pudica* (LC50 value of 164.72 µg/mL) (Table 3). These data indicate that the ethanol extract of *M. pudica* is more promising against *A. stephensi* than the other family members (8-45 times higher). Another report revealed that ethanol extract from leaves of other family members (*K. speciosa*, *K. strigosa*, *K. thymodora*, and *K. uguenensis*) had good insecticidal activity against *A. gambiae* with LC50 value of 16.35-94.01 µg/mL. Therefore, it is interesting to know the effect of ethanol extract from leaves of *M. pudica* against other vectors such as *A. gambiae*.

Further purification of the ethanol extract of *M. pudica* is needed to improve drug effectiveness. Another study reported that the insecticidal activity of ent-halim-1(10)-ene-15-oic acid from the ethanolic extract of *Kotschya uguenensis* leaves could increase its insecticidal activity three times against *A. gambiae*.⁵⁰ Insecticidal activity against *A. gambiae* was also shown by the ethyl acetate extract from seeds of another member of *Fabaceae* family, *Cassia tora*. The study used a crude extract, isolated ethyl acetate fractions, and isolated compounds aurantio-obtusin and obtusin. Interestingly, a crude extract obtained the most promising LD50 value rather than its fractions or isolated compounds (LD50 of crude extract: 2.5 µg/mL; fraction 5-7 µg/mL; isolated compounds 10-10.2 µg/mL) (Table 3). The result of this study revealed that a combination of compounds works more effectively than one active compound. Thus, ethyl acetate extract from the seeds of *M. pudica* against *A. gambiae* can also be used for future studies.

Plant-Synthesised Silver Nanoparticles

The green nanoparticle approach is promising in increasing drug sensitivity, one of which is using plant extracts with silver nanoparticles (Ag-NPs). This nanoparticle synthesis method is interesting considering its low cost, ability to accommodate large-scale production, use of simple techniques, and it being environmentally friendly.^{61,62} In this review, we showed that the toxicity of extracts of *Mimosa pudica* or its family with nanoparticles was higher than without nanoparticles (Tables 2 and 3). The aqueous extract of *Mimosa pudica* with silver nanoparticles was more effective against *Anopheles subpictus* larvae than without silver nanoparticles (Table 2). The same thing

was also shown by *Ormocarpum cochinchinense*, where aqueous extracts with silver AgNPs were more effective against *Anopheles stephensi* than without AgNPs (Table 3). As an antiplasmodial, AgNPs with methanol extract of *Indigofera oblongifolia* can suppress 99% of *P. chabaudi* (Table 3). AgNPs with aqueous extracts of plant species from other families (such as *Azadirachta indica* and *Ocimum sanctum*) also showed good results against *P. falciparum* 3D7 strain.⁶² In addition, AgNPs are also known to exert a protective effect on liver tissue injury induced by *P. chabaudi* through regulating the iron regulatory genes or reducing parasitaemia in *Plasmodium chabaudi*-infected mice.^{63,64} The unique metabolites of plant extracts can affect the properties of AgNPs formed, including nano size, high conductivity, and optical properties, which make them essential in increasing drug sensitivity. The nano size might make AgNPs penetrate more easily against *Plasmodium* sp. or *Anopheles* sp. larvae. Thus, biosynthesis of AgNPs with aqueous, methanol, and ethanol extracts of *Mimosa pudica* against *P. falciparum* strains D6 or *P. berghei* could be carried out in future studies. For insecticidal effect, biosynthesis of AgNPs with aqueous or ethanolic extract of *M. pudica* can also be carried out to increase its ability to fight *A. subpictus* and *A. stephensi* larvae.

Conclusion

This review summarises the antiplasmodial and insecticidal potential of *Mimosa pudica*. The most promising antiplasmodial activity was shown by aqueous, methanol, and aqueous/ methanol extract from the aerial part of *M. pudica* against *P. falciparum* FCR-3 strain. Meanwhile, aqueous or ethanolic extracts from leaves of *M. pudica* revealed the ability against *A. subpictus* and *A. stephensi* larvae. Future studies may aim to investigate the activity of the methanol extract of *M. pudica* against *P. chabaudi* or *P. falciparum* 3D7 and INDO strains. The conducted research can describe the action of methanol fraction or isolated compounds against *P. falciparum* FCR-3 strain. The aqueous extract of *M. pudica* can be evaluated against *P. berghei* ANKA MRA 311 strain or *P. falciparum* 3D7 strain. Extraction with other solvents can also be carried out, such as n-hexane or butanol extract to fight *P. falciparum* Dd2 strain, and the extract and ethyl acetate fraction to fight against *Anopheles gambiae* larvae. The use of plant-based-synthesised silver nanoparticles (AgNPs) needs to be intensified for further studies to increase the effectiveness of extracts against *Plasmodium* sp. or *Anopheles* sp.

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