**Review Article** 

Medicinal Plants with Antimalarial Potentials from Northern Nigeria: A Review

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# Abstract

Malaria is a prevalent infectious disease that is transmitted by parasites through the bite of infected mosquitoes. Despite advancements in medical science, malaria is still a significant global health challenge, especially in regions like northern Nigeria. Medicinal plants have long played a pivotal role in traditional healthcare systems, serving as valuable sources of therapeutic compounds. In the context of malaria, certain plants in Northern Nigeria have been traditionally recognized for their antimalarial properties, offering an alternative or complementary approach to conventional treatments. The significance of these medicinal plants lies in their potential to provide accessible, cost-effective, and culturally relevant solutions for managing malaria. Traditional knowledge about these plants has been passed down through generations, contributing to the resilience of local communities against the disease. This review explores the rich diversity of medicinal plants in Northern Nigeria with antimalarial properties. Through extensive research utilizing various search engines, we identified and examined 30 distinct plant species traditionally used in the treatment of malaria across the region. The documented uses, phytochemical compositions, and therapeutic potentials of these plants contribute valuable insights into the traditional medicinal practices of Northern Nigeria. Our findings underscore the importance of further scientific investigation and documentation of these natural resources, with the potential for developing novel antimalarial drugs and fostering sustainable healthcare practices in the region.

## Keywords: Antiplasmodial, Malaria, Traditional.

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#### **1. Introduction**

There are fewer than 400,000 flowering plants on the planet, 12% of them are employed in conventional medicine. About 10,000 of such plants have already been studied and described scientifically. Around 25% of prescribed medications in the Western medical system are higher plant-derived chemicals, and 74% of the 121 bioactive plant-derived compounds were discovered through research based on leads from traditional medicine. The elder generation passed on their knowledge of medical plant usage to the younger generation through trial and error, but this knowledge and transmission are now in jeopardy because there isn't always a guarantee that it will happen (1). Indigenous people have long used plants as health-restoring products. The effectiveness of plant-based therapies in treating a variety of ailments has been confirmed. According to the WHO, nearly 70% of people worldwide rely on herbal remedies to heal illnesses (2). The development of new antimalarial medications has historically been dominated by the utilization of medicinal plants. These compounds can be taken directly from the

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plant, manufactured, or utilized as structural models for semi-synthetic antimalarials. Early in the 18th century, the antimalarial compound quinine was initially extracted from Cinchona bark. It served as a base for the later synthesis of chloroquine, mefloquine, and other medications with a similar mechanism of action (3).

The five single-celled eukaryotic Plasmodium parasite species that cause malaria—*P*. *falciparum*, *P. ovale*, *P. malariae*, *P. vivax*, and *P. knowlesi*—are all spread by means of female anopheles mosquitoes' bite that are carrying the parasite. Malaria is a significant public health issue with international significance and scientific interest. In 2016, malaria was the sixth largest cause of death globally, it is estimated that 216 million cases and 719,600 deaths take place despite a decline in occurrences of the disease of 25.9% from 2006 to 2016 (4).

P. falciparum is the germ that causes the most dangerous type of malaria. The female anopheles mosquito serves as the insect vector for the complex life cycle of *Plasmodium* sp while second host is the vertebrate human (5). The sporozoites in the female anopheles mosquito's saliva enter the host's body while feeding on blood, starting a period of hepatic malarial infection lasting 5-7 days. Thousands of merozoites are produced during this asymptomatic phase. Clinical signs begin to manifest 48 hours after the merozoites enter the bloodstream. Chills, fever, prostration, cerebral malaria, anemia and multiple organ failure with a fatal outcome are some of the clinical symptoms. In addition to its asexual reproduction cycle, the parasite divides into sexual forms called gametocytes. The female anopheles mosquitoes take the gametocytes while feeding on blood, and the sexual cycle starts in the host mosquito's stomach (5).

There has been a pressing need to locate new antimalarial chemotherapeutic agents from natural sources, mainly medicinal plants, to potentially prevent problems associated with drug resistance. This is because the malaria parasites have a resistance to many of the current treatment regimens. The two main antimalarial lead compounds quinine and artemisinin are produced by medicinal plants, and because they were frequently used in various traditional medical systems to treat malaria (6). Over time, various nations have documented the usage of medicinal herbs to treat the symptoms of malaria, which includes fever paroxysms. This practice is widespread in Nigeria, and research on a few medicinal herbs has shown that they are effective against plasmodial infection (7).

The aim of this review exercise is to compile the most utilized medicinal plants for malaria in Northern Nigeria with a goal of improving their national standards which will consequently be used to establish their herbal monographs.

#### 2. Sources of the Literature

The search for medicinal plants was conducted using various search engines including Google Scholar, PubMed, Institute for Scientific Information, Hinari, Scopus, Scientific Information Database, etc. The keywords such as malaria endemic, antimalarial plants, antiplasmodial, ethnopharmacology, and northern Nigeria regions are used in this approach.

#### **3. Selected Medicinal Plants with Antimalarial Potentials in Northern Nigeria** *3.1 Magnifera Indica*

Mangifera indica, which is commonly referred to as mango is a type of flowering plant belonging to the Anacadiaceae family. There are 69 different kinds of mango found on tropical and subtropical continents around the globe. Mango's significance cannot be overstated. In addition to being consumed, the fruits are also utilized to make juice and wine (8). The leaves, fruit and immature stem of Magnifera indica is used as decoction in the treatment of malaria in northern part of Nigeria (9). The aqueous extract of Magnifera indica exhibited IC<sub>50</sub> of 18.11 µg/ml, 20.08 µg/ml, and 10.23 µg/ml against three strains of P. falciparum, (NF54, CamWT C580Y, and FA08) respectively (10). The Artemisia species have been found to contain flavonoids, which have been shown to have potent anti-malarial effects against P. falciparium. The flavonoids and alkaloids were discovered in Magnifera indica extracts, leading to the suggestion that their presence may be the cause of the plant's therapeutic effects. Meanwhile, the findings partially support assertions made in conventional medicine about the effectiveness of this plant in combating malaria (8).

### 3.2 Cymbopogon citratus

Popular names for *Cymbopogon citratus* include citronella grass and lemongrass. This species is a member of the Gramineae family, which has 8,000 plant species and 500 genera. A tufted perennial grass, lemon grass may reach a height of one meter and possesses a number of stiff, leafy stems which emerge from roots that are short and rhizomatous. It has an estimated five-year economic lifetime (11). The linear, tapered leaf blade may reach a maximum length of 50 cm and a maximum width of 1.5 cm. The leaf-sheath serves as a pseudo stem and has a tubular form. The leaves are long, glaucous, green, and linear which are tapered upwards and along the edges (12).

Herbalists believe that C. citratus has a variety of medical benefits. In Nigeria, concoction preparations of lemon grass have been used to cure illnesses like typhoid, fever, stomach pains, as well as malaria when combined with other herbs. Its folkloric uses as a possible malaria-controlling medication have been validated (13). Alkaloids, carbohydrates, tannins, flavonoids, cardiac glycosides, steroids, saponins, anthraquinones, carotenoids, volatile and non-volatile terpenoids were found in the aqueous leaf and root extracts of C. citratus, in accordance with the phytochemical screening results which were implicated in antiplasmodial activity of some herbal preparations making the extract a potential antiplasmodial candidate (14). Numerous herbal treatments for malaria may have antiparasitic benefits not only by directly combating the infection but also by indirectly enhancing the host's natural and adaptive defensive systems (15). When compared to chloroquine, Cymbopogon citratus extracts and its essential oils were demonstrated to have 86.6% decrease in the development of Plasmodium berghei (considering 100% inhibition by chloroquine as the reference point) (16, 17).

#### 3.3 Abrus precatorius

*Abrus precatorius* known by other names including rosary pea and jequirity pea. In Hausa language it is known as 'Idon Zakara'. It belongs to Papillionaceae family, it has glabrous internodes and leaves which twines around trees, shrubs, and hedges. Its leaflet length ranges from 1.2 to 1.8 cm. It produces fruit, which is a tiny pod with a deflexed beak and a flat, truncate form. The fruit has three to six ovoid crimson seeds with a black mark at the base that are around 0.6 cm in diameter. It grows worldwide in tropical and subtropical climates (18). Its seeds are interesting since they are generally bright, beautiful crimson with black at the hilum. This unique black and red backdrop resembles the eyes of a chicken (19).

The plant leaves and roots that have a sweet taste have been used for centuries to treat a variety of conditions including fever, stomatitis, bronchitis, asthma, diabetes, chronic nephritis, cancer, sores, scratches, and wounds as well as leucoderma, tetanus, boils, and abscesses. They also have a known hematonic and plasma expander effect (20). The leaves decoction of *A. precatorius* have also been utilized in northern Nigeria for the management of myriad of ailments including malaria, eye infection, typhoid respiratory tract infections, cough, hepatitis and skin diseases (21, 22).

More than 166 chemical constituents have so far been extracted or identified from A. precatorius; these constituents belong to many groups of substances, such as proteins, polysaccharides, alkaloids, esters, flavonoids, phenolics, steroids, terpenoids, organic acids and other constituents (19). Abruquinone, an isoflavanquinone, was discovered in the aerial parts extract and demonstrated anti-malarial activity (23). Significant antiplasmodial efficacy against P. berghei was demonstrated by aqueous leaf extract of A. precatorius at dose levels of 25, 50, and 100 mg/kg, which was equivalent to halofantrine at dose level of 25 mg/kg (24). With an IC<sub>50</sub> value of 12.1 g/ml, the n-hexane and chloroform fractions of Abrus precatorius leaf demonstrated the greatest inhibitory effects against chloroquine and pyrimethamine resistant Plasmodium falciparum (25). The presence of secondary metabolites in the plant may be the cause of the antiplasmodial action that was observed.

#### 3.4. Anogeissus leiocarpus

A deciduous tree from the combretaceae family, *Anogeissus leiocarpus* is often found in tropical and subtropical parts of the world. It is also found in central and west African countries, where it is a common herbal remedy. The plant is known for its scaly, greyish bark, thin, frequently falling branches, and alternating, ovate-lanceletshaped leaves that are 2 to 8 cm long and 1.3 to 5 cm wide (26). *Anogeissus leiocarpus*, often known as the Marke (Hausa), Kojoli (Fulani) and African

birch (English). It is a significant evergreen tree that is common throughout Africa. African and Sudanese traditional medicine utilizes Anogeissus leiocarpus extensively to treat multiple ailments including; toothaches, respiratory illnesses, diarrhea, hepatitis, jaundice, haemorrhoids, headaches, as well as antimalarial. It has potent antibacterial and antifungal action against several pathogenic pathogens (27). Stem bark of Angeissus leiocarpus is boiled and taken 2 times a day for 3 days for the treatment of malaria by the Rumaya people of Kauru Local Government Area of Kaduna State in northern Nigeria (28). In Nupe land, North central Nigeria, malaria is been treated using decoction of the A. leiocarpus. The bark with three ties of its leaves were orally administered. A warmed portion of the decoction was also used with potash or black soap for bathing. Palm kernel oil used to be added to the other portion and the resultant mixture is rubbed on the body (29). While flavonoids were lacking, four distinct secondary metabolites, including saponins, phenols, tannins, and phytosterols, were detected in the phytochemical screening of Anogeissus leiocarpus stem bark extract (30).

An investigation into a chloroquine-resistant strain of *Plasmodium falciparum* found that a methanol extract of the plant's leaves and roots was highly effective in treating malaria. When the stem bark of Anogeissus leiocarpus was examined for in vitro antiplasmodial activity, butanol, ethyl acetate, and methanol extracts were found to have the strongest efficacy. The methanolic extract has strong antimalarial properties and can increase HDL levels in organisms with malaria (31).

#### 3.5 Carica Papaya

The *Carica papaya* plant, also known as pawpaw in English and gwanda in Hausa, is a member of the Caricaceae family and has been grown in most tropical nations. *Carica papaya* is a huge, herbaceous perennial plant that resembles a tree. It has a soft single stem that may reach a height of 5 to 10 meters, sparsely spaced leaves at the top of the trunk, and a scarred lower trunk where leaves and fruits are formed. Despite having a limited lifespan on average, the plants can bear fruit for up to 20 years. The papaya has a convoluted reproductive strategy (32). The fruits are eaten as food and are a reliable source of elements (iron, calcium and potassium), vitamins (vitamin C, riboflavin and thiamine) and other nutrients. Different *Carica papaya* components have a long history of being used for medical purposes (33). In Asia, boiling and eating papaya leaves with spinach has a long tradition. Papaya leaves are one of the many plant parts with therapeutic benefits. The tea extract of *Carica papaya* leaves was displayed to have antispasmodic and antimalarial activities, and a significant effect have been observed on a variety of tumor cell lines. It has been reported to increase appetite, enhance nausea, and decrease menstrual cramps (34).

Carica papaya leaves is used among diverse tribes in northern Nigeria for the management of malaria. Concoction of leaves of Carica papaya is been taken orally and also vapour bath to treat malaria by the people of Madobi town in Kano state north-western Nigeria (35). The fresh leaves of C. papaya, Psidium gujava, and Cymbopogen citratus were combined, heated in water, and then left to cool. For a week, a cupful is consumed three times every day to cure anemia and malaria by the Kanuri tribes of north-eastern Nigeria (36). Also, infusion of the leaves and fruit of Carica papaya is been taken orally to treat malaria by the people of Rukuba town in Bassa Local Government Area of Plateau State north-central Nigeria (37).

Phytochemical components found in papaya leaves include saponins, cardiac glycosides, anthraquinones, alkaloids, flavonoids, and other phenolic chemicals (38). According to a study, C. papaya methanolic leaf extracts at 400 and 600 mg/kg body weight caused a dose-dependent and gradual decline in malaria parasites throughout the course of a curative test (39). According to another study, when compared to the control group, the average daily parasitemia of infected mice treated with chloroquine, 400 mg/kg of papaya leaf extract, and 200 mg/kg of papaya leaf extract each decreased considerably (P<0.05). However, they found no discernible difference between the infected mice treated with 100 mg/kg of Carica papaya leaf extract and the control group in terms of the amount of parasitemia (p > 0.05) (40). The antimalarial properties of C. papaya methanolic leaf extract may be due to flavonoids and alkaloids which have been demonstrated to have strong in vitro antimalarial activity against P. falciparum (41).

#### 3.6 Citrus aurantifolia

A tiny shrubby tree with a height of around 5 m, C. aurantifolia known as lime (English) and lemun tsami (Hausa). It is a tree that is both evergreen and everbearing, has thickly and erratically branching leaves, and has short, stiff spines (thorns). The leaves are alternate, elliptical to oblong-oviate in form, and have a crenulate edge (4-8 cm by 2-5 cm). A subtle purple hue can be seen on the border of the 1-inch-diameter, yellowish-white blooms. The fruits are globose to ovoid berries that range in size from 3 to 6 cm in diameter and may feature an apical papilla (42). Although it is commonly grown in green for commercial purposes, it turns yellow when fully mature. Fruits and blooms are present all year in the Northern Hemisphere, but are most abundant from May to September. The fruit skins have segments that are heavily glandular and contain pulp vesicles that are yellow-green in color. Fruit juice is sour like lemon juice but more aromatically acidic and fragrant. It is typically prized for having a distinct flavor from other limes. The seeds have a white embryo and are tiny, plump, oval, pale, and smooth (42). Initial analysis of the fruit and other parts of C. aurantifolia revealed the presence of reducing sugars, alkaloids, flavonoids, tannins, saponins, steroids, cardiac glycosides, carbohydrates, and phenols (43).

A decoction of leaves of C. aurantifolia is taken orally and also vapor bath to treat malaria by the people of Madobi town in Kano state northwestern Nigeria (35). The leaves of C. aurantifolia is boiled in water together with the leaves of Magnifera indica and Psidium gwajava, a cupful of the decoction is administered orally three times daily for five days by the Kanuri tribes of north-eastern Nigeria to treat malaria (36). Also, decoction from the root, bark, fruit, stem-twigs and leaves of the plant is taken orally to treat malaria by the people of Rukuba, Bassa Local Government Area of Plateau State north central Nigeria (37). When compared to the standard Chloroquine, C. aurantifolia leaf extracts of various solvents (aqueous, hexane, chloroform, petroleum ether, and ethanol) had a considerable inhibitory impact on the schizont development of P. Falciparum parasites (44). A study found that C. aurantifolia leaf extract has

potent *in vivo* antiplasmodial activity that is dose independent. In another study it was reported that the plant extract tested positive for alkaloids, saponins, flavonoids, cardiac glycosides, and tannins during phytochemical screening. These components may be responsible for the plant's antiplasmodial properties. This study provides a foundation for the ethnomedical use of herbs in Nigeria to treat malaria (45).

#### 3.7 Psidium guajava

The Psidium guajava tree which is normally known as gwava in English and gwaiba in Hausa is around 10m-tall with thin, smooth, uneven, and flaking bark. The oval, 5-15 cm long leaves are opposite, short-petiolate, and have noticeable pinnate veins. The spectacular flowers have many stamens and white petals up to 2 cm long. It thrives in tropical and subtropical regions around the world (46). It belongs to the Myrtaceae family, which has more than 3,800 species and 133 genera. The leaves and bark of the P. guajava tree have been used medicinally for a long time and are being used up to today (47). All parts of Psidium guajava, including the bark, fruit, leaves, and roots, have a long history of therapeutic value in the treatment of a number of illnesses, including trypanosomiasis, malaria infection, wound healing, diabetes, ulcers, hypertension, stomach aches, cholera, obesity, diarrhea, and ulcerative colitis (48). Traditional herbalists also recognize Psidium guajava as a key ingredient in antimalarial polyherbal formulations (48).

The plant is used to cure malaria, according to an ethnobotanical study of P. guajava in the town of Mubi in the Adamawa state, north-eastern Nigeria (49). Decoction of the leaves and bark is taken orally with little potash in it to treat malaria according to ethnobotanical studies carried out in some northern part of Nigeria (35). Using chloroquine and quinine as the reference drugs, Psidium guajava extract was tested for its in vitro antimalarial action against P. falciparum strain. The IC<sub>50</sub> values for methanolic and ethanolic extracts against the P. falciparum strain were found to vary from 0.048 µM to 0.965 µM. Compared to quinine (IC<sub>50</sub>= 0.832  $\mu$ M) and chloroquine (IC<sub>50</sub>= 0.065 µM), these substances demonstrated superior action against the P. falciparum strain (50). In mice infected with Plasmodium berghei, the crude extract of *Psidium guajava* effectively (p <0.05) suppressed parasite multiplication, avoided body weight loss, and reduced packed cell volume (51).

#### 3.8 Anacardium occidentale

The Anacardium occidentale tree which is known as cashew in English and kashu in Hausa has the potential to reach a height of 12 meters and is spreading, evergreen, and heavily branched. It seldom grows taller than 6 meters when planted in lateritic, gravelly, or coastal sandy regions. When produced from seed, the mature root system consists of a very noticeable taproot and a wellestablished and vast network of lateral and sinker roots. Young leaves are pale green or reddish; mature leaves are dark green. The genuine fruit, which is the nut, dries up and does not crack open. The edible cashew nut is a huge, curving seed that is approximately 2.5 cm long. The fleshy, fruitlike structure, which is widest at the apex and is commonly referred to as the fruit, quickly grows from the stalk (receptacle) at the base of the nut as it matures (52). Worldwide ethnomedical traditions have demonstrated the therapeutic potential of extracts from various sections of the A. occidentale tree for the treatment of malaria, dyspepsia, bronchitis, eczema, psoriasis, syphilis, nasal congestion and urinary insufficiency (53). Decoction from the stem bark and leaves of the plant is taken orally with honey added as an adative for 5 days to treat malaria in many parts of northern Nigeria (35). According to a research, 48 hours after starting therapy, chloroquine was completely curative while at 96 hours of therapy, A. occidentale 600 mg/kg and 800 mg/kg produced percentage curatives of 80.66% and 80.69%, respectively (54). Another finding revealed that ethanol crude extract of Anacardium occidentale (Cashew) exhibited significant antiplasmodial property (IC50  $0.577\mu$ g/ml) (55). This has established the fundamental idea behind A. occidentale's historical usage in the treatment of malaria.

#### 3.9 Momordica charantia

*Momordica charantia*, sometimes referred to as bitter melon, bitter guard, and garahunu in Hausa, is a plant that is consumed as food and utilized as a natural remedy. All plant components, including the fruits, contain the very bitter chemical momordicinso. The plants are able to grow in tropical areas (56). M. charantia, a slender, climbing monoecious plant with distinct male and female flowers that are yellow in color and appear in the leaf axils and have long stalks. It belongs to the Cucurbitaceae family (56). In folk medicine, the herb is used to cure a wide range of ailments (57). In some parts of northern Nigeria, a tablespoon of Momordica charantia's fresh leaf and fruit decoction is taken three times per day to cure malaria (58). The M. charantia leaf extract has shown an anti-malarial action, according to Akanji et al. (59), despite the fact that the mechanism remains unclear. The chemosuppression of parasitaemia and lengthened life span in infected mice treated with the extract and solvent fractions of the plant demonstrate the considerable antiplasmodial activity of the crude extract and solvent fractions of Momordica charantia stem (60). Fractionating the crude extract of the plant improved its ability to combat malaria, with the ethyl acetate fraction exhibiting the best antiplasmodial efficacy (60). Compared to the conventional antimalarial drugs chloroquine and arteether, Momordica charantia was shown to exhibit antimalarial activity; this activity may be attributable to the terpenes, alkaloids, and flavonoids or to a combination of more than one metabolite. This study supports the folktale that suggests Momordica charantia's many parts may treat malaria (60).

#### 3.10 Sida acuta

Sida acuta, the common wireweed belongs to the family malvaceae is known as tsanya in Hausa. The plant is a little perennial herb or small shrub with erect, branching stems that reach a height of around 1.5 meters. The root of this tree is long, narrow, cylindrical, and very rough. Its bark is smooth and greenish. It has lance-shaped, almost glabrous leaves, and peduncles that are the same length as petioles. Its seeds are smooth and black, with yellow blooms that can bloom alone or in pairs. It may be found in large quantities in cultivated fields, waste areas, and roadside verges. This plant may be propagated successfully using both seeds and stem cuttings (61). The Sida acuta species was phytochemically screened, and it was found to contain tannins, phenolic compounds, steroids (ecdysterone, -sistosterol, stig1naterol, ampesterol), saponosides, and alkaloids including vasicine, ephedrine, and cryptolepine (the plant's main alkaloid) (62). Sida acuta has been established to have a range of pharmacological potentials in traditional medicine, including antipyretic, anti-inflammatory, antibacterial, anti-ulcer, and antimalarial properties (63). The herb is commonly used in Nupe land in north-central Nigeria in the management of fever associated to malaria. The decoction of the leaves is taken orally to treat malaria in northern Nigeria (58), Sida acuta leaf crude ethanol extract was safe, and no overtly acute side effects were noticed. With an LD<sub>50</sub> of 2154 mg/kg bodyweight, the crude alkaloid extract from S. acuta leaves exhibited antiplasmodial action against the P. berghei malaria parasite, which was comparable to chloroquine (standard drug) (64). According to study carried out by Adesina et al (65), S. acuta methanolic extract dem-

onstrated effective in vitro antimalarial property against Plasmodium falciparum with IC50 value of  $50 \text{ g mL}^{-1}$ . In another study (66) Sida acuta leaf alkaloid, flavonoid, and phenolic extracts have demonstrated antiplasmodial activity. Phenolic extract at the dose of 600 mg/kg bw) indicated the highest level of plasmodial with 64.64% suppression level among the other extracts and was near to the positive control (chloroquine), that caused plasmodial suppression value of 76.79%. The various qualities of the plant and its historical use in medication are a result of bioactive constituents such as alkaloids, cardiac glycosides, coumarins, sesquiterpenes, saponins, steroids, phenolic compounds, and flavonoids in a significant amount in the plant extract.

Other medicinal plants with antimalarial

S/N	Scientific Name	Part of the plant	Pharmacologically active component(s)	Antimalarial/Antiplasmodial Activity	Ref
1	Tithonia diversifolia	Leaves	Orizabin and tagitinin C a type of sesquiter- pene lactones.	Its dichloromethane extract exhibited high antiplasmodial activity ( $IC_{50} < 10 \mu g/ml$ ) <i>in vitro</i> .	(67)
2	Ficus vallis- choudae	Leaves	Wighteone, maslinic acid, and genistein.	It showed high activity in <i>in vitro</i> antiplasmo- dial and <i>in vivo</i> antimalarial assays.	(68)
3	Zea mays	Husk	Quinine and also tan- nin a polyphenol	It has demonstrated a significant <i>in vivo</i> antimalarial activity toward <i>P. berghei</i> infec- tion in mice It also exhibited antiplasmodial activity against <i>P. falciparum</i> , chloroquine sensitive and resistant strains <i>in vitro</i> .	(69)
4	Picralima nitida	Seed	Akuammine an indole alkaloid	The ethanol seed extract demonstrated <i>in vivo</i> antiplasmodial activity in addition to the <i>in vitro</i> action.	(70)
5	Azadirachta indica	Leaves and Stem bark	β-sitosterol, nimbo- lide, gedunin and other limonoids	It exhibited a good antiplasmodial activity against <i>P. falciparum</i> sensitive (3D7) and resistant (RKL9) strains.	(71)
6	Chromolae- na odorata	Leaves	Flavonoids	The antimalarial activity of its ethanol frac- tion was comparable to that of the common medication chloroquine and artermether- lumenfantrine.	(72)
7	Khaya sen- egalensis	Stem bark	Fissinolide	The crude extract exhibited high antiplasmo- dial activity against chloroquine-resistant <i>P.</i> <i>falciparum</i> .	(73)
8	Ocimum gratissimum	Leaves	Flavonoids and an- thraquinones	It has a significant effect in reducing <i>P. ber-</i> <i>ghei</i> growth <i>in vivo</i> .	(74)

Table 1. Medicinal Plants with Antimalarial Potentials

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Cont	inued Table 1				
9	Vernonia	Whole	Sesquiterpene lac-	It has a very promising antiplasmodial	(75)
	cinereal	Plant	tones	activity.	
10	Ananas	Peel	Linoleic acid and	It has shown some modest antimalarial	(76)
	comosus		palmitic acid	action.	
11	Afzelia afri- cana	Stem bark	Alkaloids, flavo- noids, phenolics and anthraquinones	It exhibits antiplasmodial action against <i>P. falciparum</i> strain 3D7.	(77)
12	Senna oc- cidentalis	Root	Flavonoids and an- thraquinnes	It has remarkable antimalarial activity, <i>in vitro</i> and in mice.	(78)
13	Pteridum aquilinum	Fond	Haemanthamine	It possesses significant antimalarial activity against <i>P. berghei</i> .	(79)
14	Phyllanthus amarus	Whole plant	1-O-galloyl-6-O- luteoyl-R-D-glucose	<i>P. amarus</i> has good antimalarial activity in both ethanol and aqueous extracts.	(80)
15	Acacia mel-	Leaves and	3-(Z)-trans coumar-	Strong antioxidant and antimalarial proper-	(81)
	lifera	Stem bark	oylbetulin and 3-(E)-	ties were demonstrated by extracts from the	
			cis coumaroylbetulin	leaves and stem bark.	
16	Moringa	Leaves	Apigenin, kaemp-	<i>M. oleifera</i> leaves have high activity against	(82).
	oleifera		ferol, rutin, and quercetin	P. berghei.	
17	Trema orien-	Leaves and	Xanthone, secoiri-	Its aqueous extracts have strong antiplasmo-	(83).
	talis	stem bark	doid, triterpenes and phytosteroids	dial properties.	
18	Vitex doni-	Leaves	Abieta-11(12)-ene-	It has the potential to be a good medication	(84).
	ana		9α,13α-endoperoxide, Abieta-11(12)-ene-	option for treating malarial infection.	
			9β,13β-endoperoxide		
			and 9αH-manoyl oxide		
19	Burkea afri-	Stem bark	Rutin and caffeic acid	It has antiplasmodial potential and its anti-	(85).
	cana			plasmodial constituents are concentrated in its dichloromethane fraction.	
20	Guiera sen-	Leaves	Harman and tetrahy-	It exhibits strong antiplasmodial activity.	(86).
	egalensis		droharman		
				tential as sources of novel antimalarial c	hemicals

potentials from Northern Nigeria are summarized in table 1:

#### 4. Conclusion

The numerous ethnobotanical studies conducted in northern Nigeria have made it possible to describe a significant variety of plants used by the local communities to treat malaria in the areas where the disease is endemic, as discussed herein. This article discusses the species that have been tested for antimalarial activity as well as their potential as sources of novel antimalarial chemicals and therapeutic leads. This will serve as a solid foundation for future pharmacological studies and serve as a clear instruction on how to extract the key bioactive components for the manufacture of potential antimalarial medicines. One expects that the identification of active plant components may lead to the development of more potent medications that are both inexpensive and accessible to rural populations most at risk for disease morbidity.

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1. Awadh Ali NA, Al Sokari SS, Gushash A, Anwar S, Al-Karani K, Al-Khulaidi A. Ethnopharmacological Survey of Medicinal Plants in Albaha Region, Saudi Arabia. Pharmacognosy Res. 2017 Oct-Dec;9(4):401-407. doi: 10.4103/pr.pr 11 17. PMID: 29263636; PMCID: PMC5717795.

Laryea MK, Borquaye LS. Antimalarial 2. Efficacy and Toxicological Assessment of Extracts of Some Ghanaian Medicinal Plants. J Parasitol Res. 2019 Aug 1;2019:1630405. doi: 10.1155/2019/1630405. PMID: 31467688; PM-CID: PMC6699259.

Clement, O. A., Anthony, A. E., Félicien, 3. M. K., Mercy, G. T., Hedmon, O., Anke, W., . . . Patrick, E. O. A review for selecting medicinal plants commonly used for malaria in Uganda. African Journal of Pharmacy and Pharmacology, 2020, 14(9), 347-361. https://doi.org/10.5897/ ajpp2020.5182

4. YYu T, Fu Y, Kong X, Liu X, Yan G, Wang Y. Epidemiological characteristics of imported malaria in Shandong Province, China, from 2012 to 2017. Sci Rep. 2020 May 5;10(1):7568. doi: 10.1038/s41598-020-64593-1. PMID: 32371895; PMCID: PMC7200687.

Osman, C. P., and Ismail, N. H. An-5. tiplasmodial Anthraquinones from Medicinal Plants: The Chemistry and Possible Mode of Actions. Natural Product Communications, 2018, 13(12),1934578X1801301.https://doi. org/10.1177/1934578x1801301207.

Uzor PF. Alkaloids from Plants with An-6. timalarial Activity: A Review of Recent Studies. Evid Based Complement Alternat Med. 2020 Feb 12;2020:8749083. doi: 10.1155/2020/8749083. PMID: 32104196; PMCID: PMC7037883.

7. Ezenyi IC, Verma V, Singh S, Okhale SE, Adzu B. Ethnopharmacology-aided antiplasmodial evaluation of six selected plants used for malaria treatment in Nigeria. JEthnopharmacol. 2020 May 23;254:112694. doi: 10.1016/j.jep.2020.112694. Epub 2020 Feb 21. PMID: 32092499.

Awoibi, K. and Amah, H. Antiplasmodial 8. and Toxicological Effects of Ethanolic Extracts of literatures.

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#### **Conflict of Interest**

The authors declare no conflict of interest.

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Mango (Mangifera indica) Leaves and Bitter Cola (Garcinia kola) Seeds in Albino Rats. International Journal of Research and Scientific Innovation (IJRSI) 2019, Volume VI, Issue VIII, | ISSN 2321-

9. Dogara, A. M., Labaran, I., and Yunusa, A. Ethnobotany of medicinal plants with antimalarial potential in Northern Nigeria. Ethnobotany Research and Applications, 2020, 19. https://doi. org/10.32859/era.19.32.1-8

10. Jibira Y, Cudjoe E, Tei-Maya FM, Ayensu B, Amoah LE. The Effectiveness of Varying Combination Ratios of A. cordifolia and M. indica against Field and Laboratory Strains of P. falciparum In Vitro. J Parasitol Res. 2020 Nov 14:2020:8836771. doi: 10.1155/2020/8836771. PMID: 33294217; PMCID: PMC7691008.

Mukarram M, Choudhary S, Khan MA, 11. Poltronieri P, Khan MMA, Ali J, Kurjak D, Shahid M. Lemongrass Essential Oil Components with Antimicrobial and Anticancer Activities. Antioxidants (Basel). 2021 Dec 22;11(1):20. doi: 10.3390/antiox11010020. PMID: 35052524; PM-CID: PMC8773226.

12. Available from ANR. California Agriculture, 2014, 68(4), 160-160. https://doi. org/10.3733/ca.v068n04p160

Arome, D., Chinedu, E., Ameh, S., and 13. Sunday, A. Comparative antiplasmodial evaluation of Cymbopogon citratus extracts in Plasmodium berghei-infected mice. Journal of Current Research in Scientific Medicine, 2016, 2(1), 29. https://doi.org/10.4103/2455-3069.184126.

Avoseh O, Oyedeji O, Rungqu P, Nkeh-14. Chungag B, Oyedeji A. Cymbopogon species; ethnopharmacology, phytochemistry and the pharmacological importance. Molecules. 2015 Apr 23;20(5):7438-53. doi: 10.3390/molecules20057438. PMID: 25915460; PMCID: PMC6272507.

Raymond, B. B., Germain S. T., David, 15. D. F. S., Orelien Sylvain, M. B., Hart Mann, A. Y., Seraphin, N. O. M., Liliane, L. T., and Helen, K. K. Prophylactic antimalarial effects of Cymbopogon citratus (DC.) Stapf (Poaceae) in a mouse model of Plasmodium berghei ANKA infection: normalisation of haematological and serum biochemical status. GSC Biological and Pharmaceutical Sciences, 2021, 15(1), 05–017. https://doi. org/10.30574/gscbps.2021.15.1.0084

16. Shah G, Shri R, Panchal V, Sharma N, Singh B, Mann AS. Scientific basis for the therapeutic use of Cymbopogon citratus, stapf (Lemon grass). *J Adv Pharm Technol Res.* 2011 Jan;2(1):3-8. doi: 10.4103/2231-4040.79796. PMID: 22171285; PMCID: PMC3217679.

17. Magotra, S., Singh, A. P., and Singh, A. P. A review on pharmacological activities of cymbopogon citratus. International Journal of Pharmaceutics and Drug Analysis, 2021, 151–157. https://doi.org/10.47957/ijpda.v9i2.475

18. Kuete, V. Toxicological Survey of African Medicinal Plants, book chapter, 2014, 635-657. Available at https://www.sciencedirect.com/science/article/pii/B9780128000182000224

19. Qian H, Wang L, Li Y, Wang B, Li C, Fang L, Tang L. The traditional uses, phytochemistry and pharmacology of Abrus precatorius L.: A comprehensive review. *J Ethnopharmacol.* 2022 Oct 5;296:115463. doi: 10.1016/j.jep.2022.115463. Epub 2022 Jun 14. PMID: 35714881.

20. Tion, M., Fotina, H., and Saganuwan, S. Phytochemical screening, proximate analysis, median lethal dose (LD50), hematological and biochemical effects of various extracts of Abrus precatorius seeds in Mus musculus. Journal of Advanced Veterinary and Animal Research, 2018, 5(3), 354. https://doi.org/10.5455/javar.2018.e286 21. Paul E. D., Sangodare R. S. A., Uroko R. I., Agbaji A.S. and Dakare M. A. Chemical analysis of leaves of Abrus precatorius. International Journal of Plant Physiology and Biochemistry, 2013, 5(5), 65-67. https://doi.org/10.5897/IJP-PB2013.0193

22. Iyamah PC, Idu M. Ethnomedicinal survey of plants used in the treatment of malaria in Southern Nigeria. *J Ethnopharmacol.* 2015 Sep 15;173:287-302. doi: 10.1016/j.jep.2015.07.008. Epub 2015 Jul 14. PMID: 26187278.

23. Oladimeji, A. V., and Valan, M. F. The Potential Therapeutic Advantage of Abrus precatorius Linn. an Alternative to Glycyrrhiza glabra: A Review. Journal of Pharmaceutical Research International, 2021, 79–94. https://doi.org/10.9734/

#### jpri/2020/v32i4031035

24. Saganuwan, S. A., Onyeyili, P. A., Ameh, E. G., and Udok Etuk, E. In vivo antiplasmodial activity by aqueous extract of Abrus precatorius in mice. Revista latinoamericana de química, 2011, 39(2), 32-44.

25. Saganuwan, S. A., Patrick, A. O., Igoche, G. A., Ngozi, J. N., and Reto, B. In vitro antiplasmodial, antitrypanosomal, antileishmanial and cytotoxic activities of various fractions of Abrus precatorius leaf. International Journal of Tropical Disease and Health, 2015, 5(3), 221-229.

26. Ahmad, H. A., Review on Anogeissus leiocarpus: A Potent African Traditional Drug. International Journal of Research in Pharmacy and Chemistry , 2014, 4(3):496500

27. Bello, A. and Jimoh, A. Some physical and mechanical properties of African birch (Anogeissus leiocarpus) timber. Journal of Applied Sciences and Environmental Management, 2018, 22(1), 79. https://doi.org/10.4314/jasem.v22i1.14

28. Madara, A. A., Iliya, C. B., Azare, B. A. and Elkanah, O.S. Ethnobotanical Survey of plants used in the treatment of malaria by the Rumaya people of Kauru Local Government Area of Kaduna State. Direct Research Journal of Public Health and Technology, 2018, 3 (5)58-62. https:// doi.org/10.26765/DRJPHET.2018.3027

29. Nda-Umar U, Gbate M, Umar AN, Mann A. Ethnobotanical Study of Medicinal Plants Used for The Treatment of Malaria in Nupe land, North Central Nigeria. *Global J Res Med Plants Indigenous Med.* 2014 04;3(4):112-126.

30. Hussaini, Y., Bello, R. Y., and Mustapha, T. Preliminary phytochemical screening and GC-MS analysis of Anogeissus leiocarpus stem bark extract. The Pharma Innovation Journal. 2022, 11(11): 113-117 ISSN (E): 2277-7695.

31. Shuaibu MN, Wuyep PT, Yanagi T, Hirayama K, Ichinose A, Tanaka T, Kouno I. Trypanocidal activity of extracts and compounds from the stem bark of Anogeissus leiocarpus and Terminalia avicennoides. *Parasitol Res.* 2008 Mar;102(4):697-703. doi: 10.1007/s00436-007-0815-1. Epub 2007 Dec 9. PMID: 18066599.

32. Basalingappa, K. M. Medicinal Uses of Carica Papaya. Journal of Natural and Ayurvedic Medicine, 2018, 2(6). https://doi.org/10.23880/jo-nam-16000144

33. Oraebosi MI, Good GM. Carica papaya

augments anti-malarial efficacy of artesunate in Plasmodium berghei parasitized mice. *Ann Parasitol.* 2021;67(2):295-303. doi: 10.17420/ ap6702.342. PMID: 34598401.

34. Akhila, S. and Vijayalakshmi, N. G. Phytochemical studies on Carica papaya leaf juice. International Journal of Pharmaceutical Sciences and Research, 2015, 6. 880- 883.

35. Mukhtar, Y., Adam, A. I., Abdulkadir, A. I., Yakudima, I. I., Galalain, A. M. Ethno Botanical Survey of Medicinal Flora Used for the Treatment of Malaria in Madobi Town, Kano State – Nigeria. Iconic Research and Engineering Journals, 2019, 3 (2): 400-409.

36. Ene, A. C. and Atawodi, S. E. Ethnomedicinal survey of plants used by the Kanuris of North-eastern Nigeria. Indian Journal of Traditional Knowledge

, 2012, 11 (4),640-645.

37. Kunle, O. F., Ali, A. A. and Egharevba, H. O. Medicinal Plants Used for the Treatment of Malaria in Rukuba, Bassa Local Government Area of Plateau State, Nigeria. International Journal of Basic and Applied Sciences, 2013, 2(4) 134-138 ISSN: 2277-1921

38. Dwivedi, M. K., Sonter, S., Mishra, S., Patel, D. K., and Singh, P. K. Antioxidant, antibacterial activity, and phytochemical characterization of Carica papaya flowers. Beni-Suef University Journal of Basic and Applied Sciences, 2020, 9(1). https://doi.org/10.1186/s43088-020-00048-w

39. Momoh, J. O., Damazio, O. A., and Oyegbami, O. M. GC–MS Analysis and Antimalarial Activity of Methanolic Leaf Extract of Carica papaya against Plasmodium berghei NK65 Infection in Swiss Mice. Annual Research and Review in Biology, 2020, 183–197. https://doi.org/10.9734/ arrb/2020/v35i1230323

40. Longdet, I., and Adoga, E. Effect of Methanolic Leaf Extract of Carica papaya on Plasmodium berghei Infection in Albino Mice. European Journal of Medicinal Plants, 2017, 20(1), 1–7. https://doi.org/10.9734/ejmp/2017/34698

41. Omagha, R., Idowu, E., Alimba, C., Otubanjo, A., Agbaje, E., and Ajaegbu, H. Physicochemical and phytochemical screening of six plants commonly used in the treatment of malaria in Nigeria. Journal of Phytomedicine and Therapeutics, 2021, 19(2), 483–501. https://doi.org/10.4314/jopat.v19i2.6

42. Enejoh, O. S., Ogunyemi, I. O., Bala, M. S., Oruene, I. S., Suleiman, M. M., and Ambali, S. F. Ethnomedical Importance of Citrus Aurantifolia (Christm) Swingle. The Pharma Innovation Journal, 2015, 4(8): 01-06 ISSN: 2277-7695.

43. Enejoh, S. O., Suleiman, M. M., Ajanusi, J. O., and Ambali, S. F. In vitro anthelmintic efficacy of extracts of Citrus aurantifolia (Christm) Swingle fruit peels against Heligmosomoides bakeri ova and larvae. International Journal of Current Pharmaceutical Research, 2015, 7(2):92-96.

44. Oderinde, O., James, O., Shagari, A., Bashar, K. and Bilbis, F. Phytochemical Analysis and In-Vitro Screening of Citrus Aurantifolia Leaf Extracts for Schizonticidal Activity on Clinical Isolates of Plasmodium falciparum. The Beam journal of arts and science, 2016, 9. 81.

45. Ettebong, E., Ubulom, P., and Etuk, A. Antiplasmodial activity of methanol leaf extract of Citrus aurantifolia (Christm) Swingle. Journal of Herbmed Pharmacology, 2019, 8(4), 274–280. https://doi.org/10.15171/jhp.2019.40

46. Dange, S. S., Rao, P. S. and Jadhav, R. S. Traditional Uses of Guava: A Review. World Journal of Pharmaceutical Research, 2020, 9(5), 452-464. DOI: 10.20959/wjpr20205-17297.

47. Bijauliya, R.K., Alok, S., Kumar, M., Chanchal, D.K., Sabharwal, M., and Yadav, R.D. An Update of Pharmacological Activity of Psidium Guajava In the Treatment of Various Diseases. International Journal of Pharmaceutical Sciences and Research, 2018, 9(3): 883-893 ISSN: 2320-5148.

48. Joseph, B., and MiniPriya, R. Review on nutritional, medicinal and pharmacological properties of guava (Psidium guajava Linn.). International journal of pharma and bio sciences, 2011, 3(2): 56-73.

49. Yusuf, C. S., Zakawa, N. N., Tizhe, T. D., Timon, D., Andrew, A. D. and Musa, I. F. Ethnobotanical Survey and Phytochemical Analysis of Guava (Psidium guajava L.) Leaves in Some Communities of Mubi North, Adamawa State, Nigeria. Asian Journal of Research in Botany, 2021, 5(4): 26-33, https://doi.org/10.53294/ijflsr.2021.1.1.0035

50. Akshay, R.Y., Pravin, P. H., Manisha, D. R., Vidya, N. D., and Shrinivas, K. M., Antimalarial Activity of Psidium guajava Leaf Extracts, International Journal of Scientific Research in Chemistry, 2020, 5(6)63-68. http://ijsrch.com/IJS-RCH205612

51. Alozieuwa, U.B., Mann, A., Kabiru, A.Y., and Ogbadoyi, E.O. In vivo antimalarial efficacy of Psidium guajava leaf crude extract and fractions in Plasmodium berghei infected mice. AROC in Natural Products Research, 2022, 2(1);28-37, https://doi.org/10.53858/arocnpr02012837

52. Orwa C., Mutua A., Kindt R., Jamnadass R., and Simons A. Agroforestree Database: a tree reference and selection guide version 4.0. World Agroforestry Centre, Kenya, 2009. https://www.worldagroforestry.org/output/agroforestree-database accessed on 12-12-2022.

53. Belonwu, D. C., Ibegbulem, C. O., and Chikezie, P. C. Systemic evaluation of antibacterial activity of Anacardium occidentale. Journal of Phytopharmacology, 2014, 3(3), 193-199.

54. Joseph, A. O., Samson, O. T. Antiplasmodial Efficacy of Anacardium occidentale in Albino Mice Infected with Plasmodium berghei. Journal of Family Medicine and Disease Prevention, 2020, 6:123. doi.org/10.23937/24695793/1510123

55. Gimenez, V. M. M., Alvarenga, T. A., Groppo, M., Silva, M. L. A. E., Cunha, W. R., Januário, A. H., Smilkstein, M. J., Riscoe, M. K., and Pauletti, P. M. Antiplasmodial evaluation of Anacardium occidentale and alkyl-phenols. Revista Brasileira de Farmacognosia, 2019, 29(1), 36-39. https://doi.org/10.1016/j.bjp.2018.11.003

56. Chekka, S. and Mantipelly, N. Momordica charantia: A natural medicinal plant. GSC Biological and Pharmaceutical Sciences, 2020, 12(02), 129-135. https://doi.org/10.30574/gscbps.2020.12.2.0251

57. Aljohi Matou-Nasri А, S, Ahmed Proper-N. Antiglycation and Antioxidant ties of Momordica charantia. PLoS One. 2016 11;11(8):e0159985. doi: Aug 10.1371/journal.pone.0159985. PMID: 27513747; PMCID: PMC4981456.

58. Abdulrahman, M. D., Bradosty, S. W., Hamad, S. W., Ibrahim, M. T., Lema, A. A., Sunusi, N., Usman, M., Ashiru, I., Ahmad, N. B., Wada, N., and Bussmann, R. W. Traditional Methods for Treatment and Management of Measles in Northern Nigeria: Medicinal plants and their molecular docking. Ethnobotany Research and Applications, 2022, 23, 1–18. Retrieved from https://ethnobotanyjournal.org/index.php/era/article/view/359 59. Akanji C, Mojisola C, Taiwo E and Ola O. The antimalaria effect of Momordica charantia L. and Mirabilis jalapa leaf extracts using animal model. Journal of Medicinal Plants Research, 2016, 10: 344–350.

60. Akintola, A.O., Kehinde, B. D., Ayoola, P. B., Ibikunle, G. J., Oyewande, E. A., Arotayo, R. A., Akwu B. P., and Bello, M. O. Antimalarial Activity of the Crude Extract and Solvent Fractions of the Stem of Momordica Charantia in Plasmodium Berghei Infected Mice. Journal of Communicable Diseases, 2022, 54(3): 34-47.

61. Murali, S. and Deepa, N. A Comprehensive Review of Sida Acuta: Potential Plant of Medical Interest. European Journal of Molecular and Clinical Medicine, 2022, 9(3): 10629-10638.

62. Nwankpa P., Chukwuemeka, O. G., Uloneme, G. C., Etteh, C. C., Ugwuezumba, P., Nwosu D. Phyto-nutrient composition and oxidative potential of ethanolic leaf extracts of Sida acuta in wistar albino rats. African Journal of Biotechnology, 2015, 14(49):3264- 3269. https://doi. org/10.5897/AJB2015.14897

63. Dinda B, Das N, Dinda S, Dinda M, Sil-Sarma I. The genus Sida L. - A traditional medicine: Its ethnopharmacological, phytochemical and pharmacological data for commercial exploitation in herbal drugs industry. *J Ethnopharmacol.* 2015 Dec 24;176:135-76. doi: 10.1016/j. jep.2015.10.027. Epub 2015 Oct 21. PMID: 26497766.

64. Zakariyya, Y. M., Adefolalu, F.S. and Abubakar. A. Antiplasmodial Effect of Crude Ethanol and Alkaloidal Extracts of Sida acuta Leaf in mice. Journal of Science, Technology, Mathematics and Education, 2019, 4(2):65-68.

65. Adesina, D. A., Adefolalu, S. F., Jigam, A. A., and Lawal, B. Antiplasmodial effect and subacute toxicity of alkaloid, flavonoid and phenolic extracts of Sida acuta leaf on Plasmodium bergheiinfected animals. Journal of Taibah University for Science, 2020, 14(1), 943–953. https://doi.org/10. 1080/16583655.2020.1790912

66. Nakkliang, K., Chaichareonkul, W., Kuesap, J., Rungsihirunrat, K. Evaluation of in vitro antimalarial activity of Sida acuta Burm.f. crude extract. SNRU Journal of Science and Technology, 2020), 12(1), 130–136.

67. Waiganjo B, Moriasi G, Onyancha J, Elias N, Muregi F. Antiplasmodial and Cytotoxic Ac-

tivities of Extracts of Selected Medicinal Plants Used to Treat Malaria in Embu County, Kenya. *J Parasitol Res.* 2020 Jul 7;2020:8871375. doi: 10.1155/2020/8871375. PMID: 32724666; PM-CID: PMC7364238.

68. Chouna HSD, Dize D, Kagho DUK, Bankeu JJK, Fongang YSF, Tali MBT, et al. Constituents from ripe figs of Ficus vallis-choudae Delile (Moraceae) with antiplasmodial activity. *Parasitol Res.* 2022 Jul;121(7):2121-2127. doi: 10.1007/s00436-022-07540-5. Epub 2022 May 17. PMID: 35578036; PMCID: PMC9110216.

69. Okokon JE, Antia BS, Mohanakrishnan D, Sahal D. Antimalarial and antiplasmodial activity of husk extract and fractions of Zea mays. *Pharm Biol.* 2017 Dec;55(1):1394-1400. doi: 10.1080/13880209.2017.1302966. PMID: 28320254; PMCID: PMC6130627.

70. Okokon JE, Antia BS, Igboasoiyi AC, Essien EE, Mbagwu HO. Evaluation of antiplasmodial activity of ethanolic seed extract of Picralima nitida. *J Ethnopharmacol.* 2007 May 22;111(3):464-7. doi: 10.1016/j.jep.2006.12.016. Epub 2006 Dec 20. PMID: 17234375.

71. Hawadak J, Kojom Foko LP, Pande V, Singh V. In vitro antiplasmodial activity, hemocompatibility and temporal stability of Azadirachta indica silver nanoparticles. *Artif Cells Nanomed Biotechnol.* 2022 Dec;50(1):286-300. doi: 10.1080/21691401.2022.2126979. PMID: 36214490.

72. Elebiyo, T. C., Oluba, O. M., and Adeyemi, O. S. Anti-malarial and haematological evaluation of the ethanolic, ethyl acetate and aqueous fractions of Chromolaena odorata. BMC Complementary Medicine and Therapies, 2023, 23(1). https://doi.org/10.1186/s12906-023-04200-8

73. Amang À Ngnoung GA, Nganso Ditchou YO, Leutcha PB, Dize D, Tatsimo SJN, Tchokouaha LRY, et al. Antiplasmodial and Antileishmanial Activities of a New Limonoid and Other Constituents from the Stem Bark of Khaya senegalensis. *Molecules*. 2023 Oct 23;28(20):7227. doi: 10.3390/molecules28207227. PMID: 37894704; PMCID: PMC10609173.

74. Tchoumbougnang F, Zollo PH, Dagne E, Mekonnen Y. In vivo antimalarial activity of essential oils from Cymbopogon citratus and Ocimum gratissimum on mice infected with Plasmodium berghei. *Planta Med.* 2005 Jan;71(1):20-3. doi: 10.1055/s-2005-837745. PMID: 15678368.

75. Aboubakar, S., Souleymane, S., Adama, G., Lamoussa, P. O., Noufou, O., Jean-Baptiste, N., and Sodiomon, B. S. Antiplasmodial activity of Vernonia cinerea Less (Asteraceae), a plant used in traditional medicine in Burkina Faso to treat malaria. African Journal of Pharmacy and Pharmacology, 2017, 11(5), 87–93. https://doi.org/10.5897/ajpp2016.4703

76. Ajayi AM, Coker AI, Oyebanjo OT, Adebanjo IM, Ademowo OG. Ananas comosus (L) Merrill (pineapple) fruit peel extract demonstrates antimalarial, anti-nociceptive and anti-in-flammatory activities in experimental models. *J Ethnopharmacol.* 2022 Jan 10;282:114576. doi: 10.1016/j.jep.2021.114576. Epub 2021 Aug 27. PMID: 34461191.

77. Vigbedor BY, Osei-Owusu J, Kwakye R, Neglo D. Bioassay-Guided Fractionation, ESI-MS Scan, Phytochemical Screening, and Antiplasmodial Activity of Afzelia africana. *Biochem Res Int.* 2022 Apr 13;2022:6895560. doi: 10.1155/2022/6895560. PMID: 35465443; PM-CID: PMC9020990.

78. Mogaka, S., Molu, H., Kagasi, E., Ogila, K., Waihenya, R., Onditi, F., and Ozwara, H. Senna occidentalis (L.) Link root extract inhibits Plasmodium growth in vitro and in mice. BMC Complementary Medicine and Therapies, 2023, 23(1). https://doi.org/10.1186/s12906-023-03854-8

79. Barine, N. I., Augustine, U., and Comfort, M. In vivo Antimalarial Activity of Methanolic Extract of Young Fronds of Teridium aquilinum I. Kuhn in Mice Infected with Plasmodium berghei. Pharmacologyonline, 2014, 4 (1): 114-120

80. Aliyu K, Mohammed Y, Abdullahi IN, Umar AA, Bashir F, Sani MN, Kabuga AI, Adamu AY, Akande AO. In vitro antiplasmodial activity of Phyllanthus amarus against Plasmodium falciparum and evaluation of its acute toxicity effect in mouse model. *Trop Parasitol.* 2021 Jan-Jun;11(1):31-37. doi: 10.4103/tp.TP\_78\_20. Epub 2021 May 14. PMID: 34195058; PMCID: PMC8213120.

81. Mutai C, Rukunga G, Vagias C, Roussis V. In vivo screening of antimalarial activity of Acacia mellifera (Benth) (Leguminosae) on Plasmodium berghei in mice. *Afr J Tradit Complement Altern Med.* 2007 Oct 27;5(1):46-50. PMID: 20162054; Mahmoud Suleiman Jada et al.

PMCID: PMC2816603.

82. Olasehinde, G., Ayanda, O., Egwari, L., Ajayi, A., and Awofeso, T. In vivo Antiplasmodial Activity of Crude Ethanolic and N-hexane Extracts of Moringa oleifera Leaves. International Journal of Agriculture and Biology, 2016, 906– 910. https://doi.org/10.17957/ijab/15.0161

83. Oyebola OE, Morenikeji OA, Ademola IO. In-vivo antimalarial activity of aqueous leaf and bark extracts of Trema orientalis against Plasmodium berghei in mice. *J Parasit Dis.* 2017 Jun;41(2):398-404. doi: 10.1007/s12639-016-0815-0. Epub 2016 Jul 7. PMID: 28615849; PM-CID: PMC5447592.

84. Okpe, O., Elijah J, P., Obi, B. C., and Fred, C. N. O. Vitex doniana, In-Vitro Antioxidant, Membrane Stabilization Potential and Protective Impact against Plasmodium berghei-Passaged Mice. Research Journal of Pharmacognosy, 2023, 10(3), 15-23.

85. Ezenyi IC, Okpoko CK, Ufondu CA, Okhale SE, Adzu B. Antiplasmodial, antinociceptive and antipyretic potential of the stem bark extract of Burkea africana and identification of its antiplasmodial-active fraction. *J Tradit Complement Med.* 2021 Jan 6;11(4):311-317. doi: 10.1016/j. jtcme.2020.12.004. PMID: 34195025; PMCID: PMC8240106.

86. Bonkian LN, Yerbanga RS, Koama B, Soma A, Cisse M, Valea I, et al. In Vivo Antiplasmodial Activity of Two Sahelian Plant Extracts on Plasmodium berghei ANKA Infected NMRI Mice. *Evid Based Complement Alternat Med.* 2018 May 24;2018:6859632. doi: 10.1155/2018/6859632. PMID: 29977316; PMCID: PMC5994278.