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# Ethnomedicinal plants used for malaria treatment in Rukungiri District, Western Uganda



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# Abstract

Background Malaria remains a major global health challenge and a serious cause of morbidity and mortality in sub-Saharan Africa. In Uganda, limited access to medical facilities has perpetuated the reliance of indigenous communities on herbal medicine for the prevention and management of malaria. This study was undertaken to document ethnobotanical knowledge on medicinal plants prescribed for managing malaria in Rukungiri District, a meso-endemic malaria region of Western Uganda.

Methods An ethnobotanical survey was carried out between May 2022 and December 2022 in Bwambara Sub-County, Rukungiri District, Western Uganda using semi-structured guestionnaire. A total of 125 respondents (81 females and 44 males) were randomly selected and seven (7) key informants were engaged in open interviews. In all cases, awareness of herbalists on malaria, treatment-seeking behaviour and herbal treatment practices were obtained. The ethnobotanical data were analyzed using descriptive statistics, informant consensus factor and preference ranking.

Results The study identified 48 medicinal plants belonging to 47 genera and 23 families used in the treatment of malaria and its symptoms in the study area. The most frequently cited species were Vernonia amygdalina, Aloe vera and Azadirachta indica. Leaves (74%) was the most used plant organ, mostly for preparation of decoctions (41.8%) and infusions (23.6%) which are administered orally (89.6%) or used for bathing (10.4%).

**Conclusions** Indigenous knowledge of medicinal plants used as prophylaxis and for treatment of malaria still exist among the local communities of Bwambara Sub-County. However, there is a need to investigate the antimalarial efficacy, phytochemical composition and safety of species (such as Digitaria abyssinica and Berkheya barbata) with high percentage use values to validate their use.

Keywords African traditional medicine, Antimalarial resistance, Ethnobotanical knowledge, Malaria, Medicinal plants

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# Background

Malaria has afflicted humanity for millennia [1]. It is one of the most fatal, preventable and curable parasitic diseases globally, with about 619,000 deaths and 247 million new cases reported in 2021 [2]. In the WHO African Region, 593,000 malaria deaths were reported in 2021 [2]. Recent estimates indicate that almost half of the global population live in 82 malaria-endemic countries [3]. The majority of malaria deaths are common in the tropical and subtropical regions, particularly Central, Western and East Africa, where there is limited healthcare and/or vector (female Anopheles mosquito) control [4].



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According to World Malaria Report 2022 [2], about 96% of the global malaria-related deaths were reported in 29 countries. Uganda has one of the highest recorded malaria transmission rates on the African continent, with over 90% of the population at risk [5]. Malaria has also remained the major cause of morbidity and mortality in Uganda, as evidenced by 30–50% of outpatient visits and 15-20% of hospital admissions. The average economic loss in Uganda due to malaria is more than US\$ 500 million per year [6]. In 2021, the World Health Organization (WHO) estimated that there were 12.4 million malaria cases and over 31,350 malaria deaths in Uganda [2]. In 2022, the Ugandan Ministry of Health indicated that 68 districts recorded malaria upsurge [7]. Rukungiri District recorded 196,070 malaria cases and some deaths between 25th April and 1st May 2022, where Bwambara Sub-County (the focal area of this study) accounted for all the malaria deaths within the whole district [7]. This could be due to the poor public healthcare system in the Sub-County [8].

Inefficient malaria treatment has promoted antimalarial drug resistance and this has been cited to be of considerable concern in East Africa [2]. In Uganda, artemisinin resistance by *Plasmodium falciparum* was identified, where two pathogenic mutations were found in more than 15% of the collected samples [9]. The WHO calls for response to the emergence of antimalarial drug resistance by ensuring that efficacious treatments remain available. This calls for alternative interventions for malaria treatment which include use and exploration of medicinal plants [2].

Medicinal plants contain phytochemicals with intriguing biological activities which provide prospects for new drug discoveries [10]. Plants have remained the focus of many studies aiming at discovering antimalarials, because the current conventional antimalarial drugs such as artemisinin and quinine were discovered from Artemisia annua and Cinchona species [11]. In this context, documentation of medicinal plants used in the treatment of malaria constitutes a vital task in the preservation of indigenous knowledge and biodiversity, as well as improvement of malaria treatment interventions amongst communities. It also contributes to bioprospecting, which lays a foundation for further research on the safety and efficacy of medicinal plants, and identification of bioactive compounds that could act as druggable hit molecules for the development of new antimalarial therapies.

Previous studies in Rukungiri District have documented the ethnomedicinal plants used for managing pediatric diseases [12] and "African" diseases [13]. Based on the latter report, we found that indigenous communities of Bwambara Sub-County (Rukungiri District) cherish medicinal plants and knowledge of their utilization, management and conservation. We identified one of the most used species in this study area (Gouania longispicata Engl.) and reported on its antimicrobial activity [14]. Despite the floral biodiversity and rich ethnocultural heritage of Bwambara Sub-County [15] and Rukungiri District as a whole, there is no report on medicinal plants used in malaria prevention and treatment. Coupled with the rapid loss of vegetation and adoption of foreign cultures by the local societies [16], there is a pressing need to record such indigenous knowledge before it is lost forever. Thus, this study documented the ethnobotanical knowledge on medicinal plants that are used for treatment of malaria in Bwambara Sub-County, Rukungiri District, Western Uganda.

# Materials and methods

# Study area

This study was conducted in Bwambara Sub-County (latitudes: – 0.33778 and – 0.67119, and longitudes: 29.69882 and 29.86939), Rukungiri District, Western Uganda (Fig. 1). The area is dominated by the Bakiga ethnic group, but other ethnic groups such as the Bahororo and Banyabutumbi are also represented. The main occupation is subsistence farming, except in Rwenshama Parish, where fishing is the main economic activity [13]. The major crops grown are coffee, matooke, grapes, pears and peaches, while cattle are majorly kept for milk production [17]. Bwambara Sub-County is bordered by Lake Edward, Queen Elizabeth National Park and Kigezi Game Reserve [18].

The area has forests, forest/savannah mosaic and savannah type of vegetation [19]. It experiences a bimodal rainfall, with a short rainy season from March to May and a long rainy season between September and December. Thus, malaria transmission is perennial with maximum prevalence shortly after the rainy seasons [8, 20] and is, therefore, meso-endemic for malaria with moderateto-high transmission [21]. The public healthcare system in the Sub-County has four health centres that serve its 79 villages with an estimated population of 28,900 people [8]. Two are Public Health Centre II but only limited to outpatient services. The remaining two are Public Health Centre III facilities, which in addition to outpatient services, offer maternity services, has an inpatient ward and microscopy for diagnosing malaria. However, these health centres have inadequate facilities, are poorly stocked and have few health workers [18]. Majority of the population are unable to access good services at the health centres due to long distances, unavailability of medicine, poverty and neglect by medical staff.



Fig. 1 Map of Rukungiri district, Western Uganda showing the location of the study area (Bwambara Sub-County)

## Sample size determination

The calculated sample size (S) was 125 respondents, obtained using the formula suggested by Krejcie and Morgan [22], that is

$$S = X^2 N P (1 - P) \div d^2 (N - 1) + X^2 P (1 - P)$$

where *S* is the required sample size,  $X^2$  = the table value of chi-square for 1 degree of freedom at the desired confidence level (3.841), *N* = the population size, *P* = the population proportion (assumed to be 0.50, since this would provide the maximum sample size) and *d* = the degree of accuracy expressed as a proportion (0.05).

## **Ethnobotanical survey**

A cross-sectional survey using semi-structured questionnaires was undertaken between May 2022 and December 2022. With the help of local council representatives, seven specialized herbalists known to treat patients using herbal medicine were purposively sampled and interviewed as key informants. The study was conducted in *Rukiga/Runyankole*, the commonly used local languages in the area. The questions mainly focused on how informants prevented and diagnosed malaria, and how medicinal plants were prepared and administered for treatment of malaria.

Voucher specimens were prepared for all the collected plant species and deposited at the Department of Biology, Mbarara University of Science and Technology (Uganda), where they were authenticated by a botanist. The scientific plant names were given according to The World Flora Online (http://www.worldfloraonline.org) and International Plant Name Index (www.ipni.org). The family names of the plant species were cross-checked with the Angiosperm Phylogeny Group (www.gbif.org).

## Data analysis

Numerical data were captured in Microsoft Excel for Windows (Microsoft Corporation, USA) and thereafter exported to SPSS software (v26, SPSS Inc., USA) for analysis. Qualitative data on malaria prevention, ethnobotanical knowledge and respondents' socio-demographic characteristics were analyzed using descriptive statistics, such as percentages and frequencies. Quantitative ethnobotanical data were used to calculate frequency of citation (FC) and informant consensus factor (ICF). The FC was used to assess the number of informants who were familiar with usage of a particular medicinal plant to treat malaria. Use report (UR) was recorded whenever any informant mentioned use of a medicinal plant in a particular way. The ICF was used to determine the homogeneity of the collected ethnobotanical data using the formula [23]:

$$ICF = (N_{ur} - N_t)/(N_{ur} - 1)$$

where  $N_{ur}$ =total number of use reports for each disease category, and  $N_t$ =total number of species in each use category. The ICF value ranges from 0 to 1. A high ICF value (close to 1) signifies that a high proportion of informants uses a few plant species to treat a particular disease, and this suggests that the community possesses a well-defined mechanism for exchange of indigenous knowledge. Conversely, low ICF values (close to 0) indicate that many plant species are reported to be used by a high proportion of informants to treat a particular disease, implying that there is lack of information exchange within the community or the medicinal plants are randomly selected [24].

## Results

## Socio-demographic characteristics

The study involved 125 informants, of which there were 64.8% females and 35.2% males (Table 1). The largest number of respondents were 31–40 years (27.8%). In general, 53.5% were above 40 years, while only 18.5% were  $\leq$  30 years. The findings revealed that most of the informants had attended primary school (55.6%), while 30.4% did not attain any formal education. The main religion in the area is Anglican (45.6%) and Catholic (31.6%).

Approximately 31.6% of the respondents declared a lack of monthly (source of) income, while 27.8% others earned less than 50,000 Uganda shillings (UgX) (US\$ 13.07) per month. With the exception of a formally employed primary school teacher who receive monthly salary, the rest got their earnings from harvesting crops. The monthly earnings were got from the sum of the block money earned after selling the harvests at the end seasons divided by 12 months. Therefore, the locals hardly had any income outside harvesting periods. Moreover, most locals carried out subsistence farming, where food is simply grown for consumption. The highest monthly income was UgX 560,000 (US\$ 146.33), earned by the primary school teacher.

## Source of knowledge of medicinal plants

Majority of the respondents (62%) acquired knowledge on ethnomedicinal plants for malaria treatment from 
 Table 1
 Socio-demographic characteristics of informants from

 Bwambara Sub-County, Rukungiri District, Uganda
 Page 100 (2000)

Variable	Characteristic	Percentage (%)
Gender	Male	35.2
	Female	64.8
Age group (years)	18–19	0.8
	20–30	17.7
	31–40	27.8
	41-50	21.5
	51-60	15.2
	>60	17
Educational status	None	30.4
	Primary	55.6
	Secondary	13.2
	College	0.8
Religious affiliation	Anglican	45.6
	Catholic	31.6
	Pentecostal	16.5
	Moslem	6.3
Ethnic group	Bakiga	83.5
	Bahororo	5.1
	Banyabutumbi	3.8
	Banyankole	2.5
	Banyarwanda	0.8
	Congolese	0.8
Occupation	Peasant farmer	82.3
	Business	10.1
	Teacher	0.8
Income (UgX)	None	31.6
	< 50,000	27.8
	50,001-80,000	19
	>80,000	12.7

UgX Ugandan shillings, and 1 US\$ = 3827 UgX



Fig. 2 Source of knowledge on plants used for malaria treatment in Bwambara Sub-County, Rukungiri District, Uganda

their parents, while 16.5% got knowledge from older family members, usually grandparents (Fig. 2). Traditional healers contributed 5.1% to the transfer of indigenous knowledge. However, a very small portion of the respondents acquired ethnobotanical knowledge through other means, such as spiritual revelations, radio and television programs, and reading ethnobotanical books.

# Symptoms of malaria and treatment-seeking behaviour of patients

In the study area, malaria is locally known as *Omush-weija gw'ensiri* (*Omushweija* means fever, *ensiri* means mosquito). Thus, this is directly translated to mean "fever of mosquitoes". For this, all informants were aware that malaria is caused by mosquito bites. Some informants reported that drinking unboiled water, exposure to morning dew and rain may cause malaria. Other indirect causes of malaria such as having grown bushes around homesteads, leaving the house doors and windows open in the evening hours and having stagnant water were also cited.

The most important symptoms of malaria were feeling cold (having goosebumps), high body temperature, headache, shivering, vomiting and joint pains (Table 2). It should be noted that all or a combination of the listed symptoms may be present in a malaria patient. Thus, the informants diagnose a certain disease as malaria if the patient presented with at least three of the listed symptoms. The participants were aware that not all patients recover from malaria after taking herbal remedies, which necessitated referral to hospitals. Participants reported that referrals are made basing on severity of the symptoms (i.e., extremely high body temperature and confusion) (Table 3).

# Prevention of malaria

All the informants reported that sleeping under treated mosquito net prevents contracting malaria. Other preventive measures included destruction of mosquito breeding areas such aas stagnant waters and elimination of unwanted vegetation within and around homesteads (Fig. 3).

# Diversity of ethnomedicinal plants used for malaria treatment

A total of 55 plant species were mentioned to be used for treatment of malaria (Table 4). However, only 48 of the voucher specimens collected were taxonomically identified. The other seven species could not be found within the study area during field visits, and only their local

Table 2 Malaria symptoms mentioned by respondents in Bwambara, Rukungiri District, Uganda (n = 125)

General malaria symptoms	Local language (Runyankore/Rukiga)	Percentage
Feeling cold/goosebumps	Okupiipa	82
High temperature	Omuriro	79
Headache	Okuterwa omutwe	77
Shivering	Okutetema	75
Vomiting	Okutanaka	72
Joint pains	Okushaash engyingo	70
Loss of appetite	Okuremwa kurya	58
General body weakness	Okubura amaani	42
Abdominal pain	Okushaasha omunda	37
Wound or burns	Ebironda/amasya	26
Sweating	Okututuuka	23
Sticky or red eyes	Okutukura amaisho	17
Bitter or sour mouth	Okushariira akanwa	15
Anemia-very pale palms, fingernails	Okuhwamu eshagama–okwera engaro n'enono	13
False smell	Okunukirwa ebitariho	12
Flue	Sanyiga	11
Diarrhea	Okwirukana	10
Frequent thirst	Okugyira eiriho buri kanya	9
Teary eyes	Okurira amaisho	8
Severe dehydration indicated by pale eyes	Okuhwamu amaizi–amaisho garikwera	7
Yellow eyes	Amaisho g'akinekye	7
Difficulty in breathing	Okwisya kubi	2
Painful swelling or lumps in the skin	Okuzimba	2

Severe symptoms	Inform	nants						Total	Ranking
	I <sub>1</sub>	I <sub>2</sub>	I <sub>3</sub>	I <sub>4</sub>	I <sub>5</sub>	I <sub>6</sub>	I <sub>7</sub>		
Coma/loss of consciousness	5	4	3	5	4	4	5	30	3rd
Confusion	3	7	5	6	6	5	6	38	2nd
Convulsions	6	3	7	3	3	6	1	29	4th
Diarrhea	1	2	1	4	1	1	2	12	7th
Extremely high body temperature	7	6	6	7	7	7	7	47	1st
Fits in children	4	5	4	3	5	3	4	28	5 <sup>th</sup>
Vomiting—cannot keep any food or drink	2	1	2	1	2	2	3	13	6 <sup>th</sup>

**Table 3** Severe malaria symptoms that warrant hospitalization among indigenous communities in Bwambara Sub-County, Rukungiri District, Western Uganda

/ Informant



Sub-County, Rukungiri District, Western Uganda

names were recorded (Table 4). The identified species belonged to 47 genera and 23 families. Most plant species belonged to family Asteraceae (22.9%), Fabaceae (10.4%), Lamiaceae (8.3%) and Poaceae (8.3%). The calculated ICF value for malaria was 0.81. Of the medicinal plants reported for malaria treatment, a paired comparison indicates that the highest rank was given for *Vernonia amygdalina* (FC=76; UV=0.608), followed by *Aloe vera* (FC=35; UV=0.28) and then *Azadirachta indica* (FC=15; UV=0.12).

# Life forms, plant parts, preparation and administration of herbal remedies for malaria treatment

The identified plant species consisted of mainly shrubs (31.3%), trees (27.1%) and herbs (25.0%) (Fig. 4; Additional file 1). On the other hand, the most used plant organs were leaves (74%), followed by stem bark (9%) and root bark (8%) (Fig. 5). Most of the plants were used in combination with each other (65.5%). All the plant materials were used immediately after collection (while still fresh). Only the stem bark of *Warburgia ugandensis* and

leaves of *Azadirachta indica* were reported to be used either when fresh or in powder form.

The most common preparation methods were decoctions (41.8%) and cold infusion (23.6%) (Additional file 1). Other methods included bathing (3.6%) and steaming (3.6%). Some herbalists cited continuous use of some medicinal plants such as *Azadirachta indica* as tea or in tea as a prophylaxis for malaria.

# Discussion

This study showed that most of the respondents were females (Table 1). This observation is contrary to a recent report by Tabuti et al. [24] in Eastern Uganda. Previous studies in Uganda have indicated higher participation of females than males in ethnobotanical surveys, since they are the ones who take healthcare needs of their families [25]. Moreover, more females could have participated because of their indigenous knowledge of nutri-medicinal plants as home managers and disease managers amongst children [26, 27]. Most respondents were the Bakiga, because they were the first group of Bantu ethnic group to migrate into this area as early as the 1950s, where the original inhabitants were the minority Banyabutumbi [28]. There were low-income levels among the respondents indicating that most of them depended on subsistence farming. This is typical in Uganda, and may also explain their preference of medicinal plants over conventional antimalarial drugs.

Regarding medicinal plant knowledge dynamics, most respondents hinted that they inherited the indigenous knowledge about medicinal plants for malaria treatment from their parents or grandparents (Fig. 2). This is not surprising, because African traditional medicine recipes are usually a guarded family secret, so that they are only passed orally to children from parents or through

Table 4         Medicinal plants used in the management of malaria in Bwambara Subcounty, Re	}ukungiri	ri District, Uga	anda
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No.	Family Botanical name	Voucher number	Part used	Life form	FC	UV
	Acanthaceae					
1	Justicia anselliana (Nees) T. Anderson	H22-013	L	Shrub	1	0.008
	Apocynaceae					
2	Mondia whitei (Hook.f.) Skeels	H22-030	L	Vine	1	0.008
3	Cascabela thevetia (L.) Lippod	H22-049	L	Tree	1	0.008
	Asphodelaceae					
4	<i>Aloe vera</i> (L.) Burm.f	H22-022	L	Herb	35	0.28
	Asteraceae					
5	Baccharoides lasiopus (O.Hoffm) H.Rob	H22-008	L, RB	Shrub	6	0.048
6	<i>Berkheya barbata</i> (L.f.) Hutch	H22-040	L	Herb	1	0.008
7	Bidens pilosa L	H22-035	FI	Herb	3	0.024
8	<i>Bothriocline longipes</i> (Oliv. & Hiern) N.E.Br	H22-046	L	Herb	1	0.008
9	Crassocephalum vitellinum (Benth.) S. Moore	H22-016	L	Herb	1	0.008
10	Vernonia amygdalina Delile	H22-002	L, RB	Shrub	76	0.608
11	Hoffmannanthus abbotianus (O.Hoffm.) H.Rob., S.C.Keeley & Skvarla	H22-019	L	Herb	4	0.032
12	Laggera alata (D.Don) Sch.Bip. ex Oliv	H22-020	L	Herb	1	0.008
13	Solanecio mannii (Hook.f.) C. Jeffrey	H22-005	L	Shrub	2	0.016
14	Sonchus oleraceus L	H22-011	St	Herb	5	0.04
15	Tithonia diversifolia (Hemsl.) A.Gray	H22-050	L	Shrub	5	0.04
	Bignoniaceae					
16	<i>Kigelia africana</i> (Lam.) Benth	H22-036	L	Tree	1	0.008
17	Markhamia lutea (Benth.) K.Schum	H22-031	RB	Tree	1	0.008
	Canellaceae					
18	Warburgia ugandensis Sprague	H22-014	SB	Tree	3	0.024
	Caricaceae					
19	Carica papaya L	H22-047	SB, L, Fr	Tree	6	0.048
	Cleomaceae					
20	Cleome gynandra L	H22-001	L	Herb	1	0.008
	Cucurbitaceae					
21	Momordica foetida Schumach	H22-025	L, SB	Vine	2	0.016
	Euphorbiaceae					
22	Manihot esculenta Crantz	H22-033	L	Shrub	1	0.008
23	Shirakiopsis elliptica (Hochst.) Esser	H22-009	L	Tree	1	0.008
	Fabaceae					
24	Albizia coriaria Welw. ex Oliv	H22-012	SB	Tree	2	0.016
25	Erythrina abyssinica Lam	H22-010	L	Tree	3	0.024
26	Indigofera arrecta Hochst. ex A.Rich	H22-006	L, RB	Shrub	1	0.008
27	Senna didymobotrya (Fresen.) H.S. Irwin & Barneby	H22-037	L	Shrub	2	0.016
28	Vachellia hockii (De Wild.) Seigler & Ebinger	H22-041	SB	Tree	1	0.008
	Lamiaceae					
29	Clerodendrum capitatum (Wild.) Schumach	H22-021	L	Shrub	5	0.032
30	Ocimum gratissimum	H22-003	L	Shrub	1	0.008
31	Ocimum kilimandscharicum Gürke	H22-018	L	Shrub	2	0.016
32	Plectranthus barbatus Andrews	H22-007	L	Shrub	3	0.024
	Malvaceae					
33	Sida alba L	H22-044	L	Shrub	3	0.024
	Meliaceae					

# Table 4 (continued)

No.	Family Botanical name	Voucher number	Part used	Life form	FC	UV
34	Azadirachta indica A. Juss	H22-048	L	Tree	15	0.12
	Moringaceae					
35	Moringa oleifera Lam	H22-027	Seed, L	Tree	4	0.032
	Musaceae					
36	Musa acuminata Colla	H22-023	L	Herb	1	0.008
	Myrtaceae					
37	Eucalyptus grandis W. Hill ex Maiden	H22-039	L	Tree	2	0.016
	Poaceae					
38	Digitaria abyssinica (Hochst. ex A.Rich.) Stapf	H22-026	L	Grass	6	0.048
39	Imperata cylindrica (L.) P.Beauv	H22-042	L	Grass	1	0.008
40	Pennisetum purpureum Schumach	H22-029	St	Grass	2	0.016
41	Saccharum officinarum L	H22-004	St	Grass	1	0.008
	Rhamnaceae					
42	<i>Gouania longispicata</i> Engl	H22-028	L	Liana	1	0.008
	Rutaceae					
43	Citrus limon (L.) Burm.fil	H22-043	L, SB	Tree	2	0.016
44	Clausena anisata (Willd.) Hook.f	H22-032	L	Shrub	1	0.008
	Solanaceae					
45	Nicotiana tabacum L	H22-015	L	Herb	1	0.008
46	Physalis peruviana L	H22-034	L, RB	Herb	3	0.024
	Verbenaceae					
47	Lantana trifolia L	H22-024	L	Shrub	3	0.024
	Vitaceae					
48	Cyphostemma adenocaule (Steud. ex A.Rich.) Desc	H22-017	L	Liana	1	0.008
49	Unidentified	NA	L	NA	6	0.048
50	Unidentified	NA	L	NA	3	0.024
51	Unidentified	NA	L	NA	1	0.008
52	Unidentified	NA	L	NA	2	0.016
53	Unidentified	NA	L	NA	4	0.032
54	Unidentified	NA	L	NA	1	0.008
55	Unidentified	NA	L	NA	1	0.008

NA not applicable, FC frequency of citation, UV use value, plant parts: Bk bark, Fl flowers, Fr fruits, L leaf, RB root bark, St stem, SB stem bark



 Shrubs • Trees • Herbs • Grass • Liana • Vines
 Fig. 4 Life forms of medicinal plants used in the management of malaria in Bwambara Sub-County, Rukungiri District, Western Uganda

apprenticeships from relatives [16]. It was interesting to note that traditional healers contributed 5.1% to the transfer of indigenous knowledge (Fig. 2). This may be



**Fig. 5** Plant parts used in preparation of remedies for malaria treatment in Bwambara Sub-County, Rukungiri District, Western Uganda

attributed to the cultural acceptability and strong belief in herbal medicines which has driven commercialization of herbal products. Therefore, the impetus behind acquiring herbal medicine knowledge observed in this study may be related to the need to establish a source of income through sale of herbal products [29]. Absence of a respondent who had acquired folk knowledge through formal training is because herbal medicine training centres are rare in Uganda.

The indigenous communities surveyed were wellinformed that malaria is caused by mosquito bites, though some believed that taking unboiled water, exposure to morning dew as well as rain are other potential causes. Other indirect causes of malaria such as having grown bushes around homesteads, leaving the house doors and windows open in the evening hours and having stagnant water were also cited. A previous study in Uganda by Adia et al. [25] reported that 96% of the respondents understood that malaria was caused by infected mosquito bites. In similar study in Zimbabwe, all respondents recognized mosquito bites as the main cause of malaria [30] but traditional healers still held it that exposure to morning dew and drinking water from unprotected wells caused malaria. In Kenya, it was cited that malaria is believed to be a direct cause of mosquito bites, exposure to cold weather [31] or consumption of fresh unboiled milk (Cheko che makiyo in the Tugen dialect), dirty water, *ikwek* (i.e., vegetables, such as Gynadropis gynadra and Solanum nigrum) [32], witchcraft (sorcery attacks) and supernatural forces [33, 34]. In Eastern Uganda, some people believe that keeping a dirty homestead with dense bushes or pools of stagnant water caused malaria [35]. Though bushes and stagnant water may not be the direct causes of malaria, they are breeding sites for mosquitoes which transmit malaria. In this case, some communities would close windows and doors in the evening to guard against mosquito bites [35].

The most important symptoms of malaria were feeling cold/goosebumps, high body temperature, headache, shivering, vomiting and joint pains. These findings are consistent with Stangeland et al. [36] who reported high body temperature, shivering and headache as the most frequent symptoms of malaria in Nyakayojo Sub-County, Western Uganda. Another study in Central Uganda mentioned high body temperature, red eyes and vomiting as the most common malaria symptoms [25]. In the malaria-endemic Eastern Uganda, fever was mentioned as the major malaria symptom, along with vomiting, body weakness, headache, diarrhea, convulsions, inappetence and body chills [24]. Outside Uganda, malaria has been associated with periodic fever, sweating, backache, chills, headache, vomiting, diarrhea and joint pains [37, 38], coldness, goose pimples and headaches [30], which aligns well with our findings.

As indicated by the herbalists, some severe malaria symptoms such as very high body temperatures and confusion (probably due to cerebral malaria) were indicative that the patient needed to be referred for further treatment using conventional medicine (Table 3). This suggests that most people seek for conventional medical treatment when malaria is in severe stages. The listed symptoms relate with those of cerebral malaria, which may signify its prevalence in the study area. A study by Stangeland et al. [36] in Western Uganda found that hospital referrals were being recommended only in cases, where herbal remedies did not relieve malaria symptoms within 2-3 days of treatment or if the patient presented severe symptoms. The current findings agree with previous studies among traditional healers which reported few hospital referrals in exceptional cases when patients showed severe malaria symptoms or herbal remedies failed completely [24, 30, 37]. In a study by Ngarivhume et al. [30], lack of a patient's improvement was attributed to evil spirits, where the patient needed cleansing/placating the spirits and then repeat or be given a different prescription thereafter. Such practices have been cited in Kenya, where malaria which is thought to be caused by supernatural forces required the intervention of diviners (such as Oloiboni among the Maasai and Orgoiyon among the Tugen) [33]. Such practices and beliefs, coupled with improper diagnosis and wrong choice of appropriate herbal remedies may result in death. This requires sensitization of the population about proper diagnosis of malaria and the dangers associated with late treatment.

All the informants reported that sleeping under treated mosquito net prevents contracting malaria (Fig. 3). This could be because since November 2018, Uganda as part of the eleven (high burden to high impact) countries distributed insecticide-treated mosquito nets (ITN) across the country [39]. This countrywide campaign could be the reason as to why each informant knew about ITN. Other preventive measures included the destruction of mosquito breeding areas, such as stagnant waters, and the reduction of unwanted vegetation within and around the homesteads. In Uganda, Tabuti [35] and Adia et al. [25] reported similar preventive measures used to guard against malaria. These findings are comparable to that in Zimbabwe where the use of ITN, destruction of mosquito breeding and resting areas, and reduction of unwanted undergrowth and puddles within and around homesteads, avoiding dew and residual spraying as malaria preventive measures are used to prevent malaria [30].

We identified 48 species, out of the 55 species mentioned by the respondents. The other species which could not be found during the field visits suggest that some plants are rare in the study area, or may have become extinct due to over exploitation or habitat destruction [18]. These raises need for instituting conservation measures. The identified plant species were majorly members of family Asteraceae, Fabaceae, Lamiaceae and Poaceae, which is in agreement with studies in Uganda [24, 25, 36, 40] and elsewhere [37, 41]. The prevalence of these families may be attributed to the frequent distribution in the area, diverse habitat, availability and presence of diverse secondary metabolites which increase the effectiveness of herbal remedies from them [42].

Of the 48 plants mentioned by herbalists, most species were listed for use in the management of malaria in Uganda and other countries. For instance, *Carica papaya*, *Albizia coriaria*, *Warburgia ugandensis*, *Azadirachta indica*, *Aloe vera* and *Vernonia amygdalina* in Uganda [24, 25, 40, 43–45], Zimbabwe [30], Kenya [38], Cameroon [46], Indonesia [47] and Ethiopia [48]. The calculated ICF for malaria was 0.81, implying that there is substantial agreement amongst the local population about the medicinal plants used in malaria treatment.

Leaves of shrubs, trees and herbs are the commonly used parts for preparing malaria remedies, corroborating some ethnobotanical reports in other parts the country [24, 25, 35, 36]. This could translate into a more sustainable use of the plants, as compared to roots. Leaves regenerate under favorable conditions to preserve the occurrence of the plants but their high use frequency in this study may plausibly be related to their year-round availability [49]. The observed use of different plants or plant parts could be to exploit their synergistic therapeutic effects [36, 50, 51], mask toxicity of efficacious plants or as a trick of keeping the secrecy of herbal formulae [52].

Oral administration (89.6%) was the major route used, but use of prepared decoctions for bathing was also practiced (10.4%). This is expected, because oral dosage forms are easy to prepare and administer, especially because malaria is an internal disease [38, 53]. One striking feature was that posology of the remedies were based on cups or glasses, a phenomenon that has been cited among herbalists throughout the East African botanical plate [24, 38, 40].

An in-depth search of literature indicated that 22 species recorded in this ethnobotanical study have been evaluated for their antimalarial/antiplasmodial activities (Table 5). Most species possess acceptable preclinical efficacy and safety, with the exception of Vernonia amygdalina [54] and Senna Didymobotrya [55] which showed high toxicity. It is worth noting that Azadirachta indica and Vernonia amygdalina has been subjected to clinical trials. A homeopathic Azadirachta indica preparation was shown to be effective for reducing malaria attacks in a highly endemic area for Plasmodium falciparum [56]. Similarly, Challand and Willcox [57] performed a clinical trial utilizing a remedy of Vernonia amygdalina leaves for the treatment of uncomplicated malaria. They found that there was an adequate clinical response at day 14 in 67% of cases. Total parasite clearance was observed only in 32% of those with adequate clinical response, where recrudescence occurred in 71% and this hampers complete remission of the parasite from the body [57]. In conclusion, these reports support that the inventoried medicinal plants possess antimalarial properties which can be further investigated for development of new antimalarial drug candidates. Specifically, clinical trials should be done on some of the species that have exhibited promising antimalarial efficacy. However, these maybe impeded by the strict regulatory requirements for clinical studies, as well as the financial muscle required [38]. In addition, formulations rich in known bioactive compounds from these species should be specifically targeted.

## Conclusion

The local communities in Bwambara sub-county of Rukungiri District possess rich and diverse indigenous knowledge of plant-based medicine used to manage malaria. This could indicate high malaria cases, and/ or accessibility, availability and potency of herbal remedies. More than 30% of the plants recorded in this study have been reported elsewhere for treatment of malaria and have experimentally recorded antimalarial and antiplasmodial activities. No study has reported any pharmacological properties, safety and efficacy of *Digitaria abyssinica* and *Berkheya barbata*. There is need to validate the toxicity and efficacy of other unstudied species to foster the discovery of efficacious, safe and effective standardized antimalarial drugs.

Deferences	Tovicity profile	Deported bioactive	Antinlacmodial/	Colyon+ <sup>1</sup>	nepro tuelo	dicinal nlant
					anda	trict, Western Ug;
ara Sub-County, Rukungiri	a phytotherapy of Bwamb	rom plants reported in malari	of extracts and compounds isolated fr	d toxicity studies	odial, antimalarial and	ole 5 Antiplasmo

Medicinal plant	Plant organ	Solvent <sup>1</sup>	Antiplasmodial/ antimalarial ( <i>Plasmodium</i> spp) efficacy <sup>2</sup>	Reported bioactive phytochemicals	Toxicity profile	References
Aloe vera	Leaves	Water	EC <sub>50</sub> values for extract: 0.289 to 1056 µg/mL (MRC-2), 169.76 (3D7) for aloin	Aloin	Considered to be safe	[58, 59]
Albizia coriaria	Stem bark	MeOH	15.2 (D6); 16.8 (W2)	Triterpenoids, lupeol, lupenone	Cytotoxic to the human glioblastoma cell line U87 CD4 CXCR4 (CC <sub>50</sub> = 6.4 and 14.9 µg/ml for ethanol and DMSO extracts)	[60–63]
Azadirachta indica	Leaves	Water, MeOH	17.9 (D6); 43.7 (W2)	Terpenoids, isoprenoids, gedunin, limonoids: khayan- thone, meldenin, nimbinin	Cytotoxicity LD <sub>50</sub> of 101.26 and 61.43 µg/ml for water and methanol extracts	[64–68]
Bidens pilosa	Leaves	DCM, water, MeOH	8.5, 5, 11, 70 (D10)	No reports	Hydro and ethanolic extracts are not toxic in mice ( $LD_{50} = 12.3$ and 6.2 g/kg bw, respectively). Safe in humans	[69–71]
Carica papaya	Leaves	Ethyl acetate	2.96 (D10), 3.98 (DD2), 0.2 μM (carpaine)	Carpaine	Carpaine has high selectivity (106), nontoxic to normal red blood cells and rat skeletal myoblast (L6) cells	[72–74]
Clausena anisata	Twigs, Leaves	DCM/MeOH, water	18 (D10),> 100 (D10); 55,> 100 (D10)	No reports	Potentially toxic	[69, 75]
Cleome gynandra	Whole plant	Hexane, ethyl acetate, MeOH	Schizonticidal activity in vitro (NK65)	Apigenin, caffeic, Ferulic, hexadecanoic acids, kaempferol, taraxasterol, pheophytin A, sitosterol, stigmasterol; 5,7,13,15-eico- satetraen-9, 12-diol and 9-hydroxy-5,7,13,15- eicosatetraen-12-one	No report	[76]
Clerodendrum capitatum	Leaves	Ethanol	65.3% chemosuppression at 400 mg/kg	Tannins, flavonoids, alkaloids, triterpenes	LD <sub>50</sub> > 5000 mg/kg	[27]
Erythrina abyssinica	Stem bark	Ethyl acetate	83.6% inhibition of <i>P. falcipa-</i> <i>rum</i> at 10 µg/ml	Chalcones (5-prenylbutein, homobutein), flavanones such as 5-deoxyabyssinin II, abys- sinin III and abyssinone IV	Minimal toxicity reported in animal studies	[54, 78]
Kigelia africana	Bark, fruit	Chloroform/ethyl acetate, MeOH	59.9 (K39), 83.8 (V1/S); fruits had 165.9 (K39)	None reported	Aqueous fruit extract elicited hepatorenal toxic effects	[79, 80]

Table 5 (continued)						
Medicinal plant	Plant organ	Solvent <sup>1</sup>	Antiplasmodial/ antimalarial ( <i>Plasmodium</i> spp) efficacy <sup>2</sup>	Reported bioactive phytochemicals	Toxicity profile	References
Markhamia lutea	Leaves	Ethyl acetate	71% inhibition of <i>P. falciparum</i> at 10 µg/ml	Phenylpropanoid glycosides, cycloartane triterpenoids, musambins A-C, Cand- musambiosides A-C	Aqueous leaf and ethanolic extracts with curative anti- inflammatory activity attenu- ated paclitaxel toxicity in rat's intestine	[54, 81, 82]
Momordica foetida	Shoot	Water	6.16 (NF54); 0.35 (FCR3)	Saponins, alkaloids, cardiac glycosides	No toxicity against human hepatocellular (HepG2) and urinary bladder carci- noma (ECV-304, derivative of T-24) cells	[83-85]
Mondia whitei	Whole root	MeOH	2% parasitemia suppression (ANKA)	Chlorogenic acid	Low toxicity on mice exposed to extract for 90 days	[86, 87]
Moringa oleifera	Flower, leaves	Water, MeOH, acetone, hexane	Parasitemia suppression upto 99.48%	Flavonoids	Not toxic	[88]
Ocimum gratissimum	Leaves/twigs	DCM	8.6 (W2)	Flavonoids	LD <sub>50</sub> of the butanolic and ethyl acetate fraction of MeOH leaf extract were 2154.1 and 3807.9 mg/kg	[64, 89, 90]
Ocimum kilimandscharicum	Leaves, twigs	DCM	0.843 (D6); 1.547 (W2)	No reports	No reports	[89]
Plectranthus barbatus	Leaves	DCM	No activity	No toxicity recorded	No reports	[89, 91]
	Rootbark	Water (hot), chloroform/ MeOH	100 mg/kg/day of extracts had 55.23% and 78.69% para- site chemosuppression	No report	No reports	
Senna didymobotrya	Leaves twigs	MeOH, DCM/MeOH (1:1)	> 100 (K39), 9.5 (D10)	Quinones	Root extracts were toxic $(LD_{50} = 1927 mg/kg)$	[55, 69, 92]
Solanecio mannii	Leaves	MeOH	21.6 (3D7), 26.2 (W2)	Phytosterols, n-alkanes and N-hexacosanol	No reports	[36, 93]
Tithonia diversifolia	Aerial parts, leaves	MeOH, ether	1.2 (3D7); 1.5 (W2), MeOH extract had 74% parasitemia suppression	Tagitinin C, sesquiterpene lactones	Aerial parts reportedly cytotoxic against cells from human foetal lung fibroblast cell line	[94–97]
Vernonia amygdalina	Leaves	MeOH/DCM, ethanol	2.7 (K1), 9.83. In vivo parasite suppression of between 57.2– 72.7% in combination with chloroquine	Vernolepin, vernolin, vernolide, vernodalin and hydroxy vernodalin, steroid glucosides	Petroleum ether extract elic- ited strong cytotoxicity	[36, 54, 57, 85, 95, 98]
Warburgia ugandensis	Stem bark	MeOH, water, DCM	6.4 (D6); 6.9 (W2), 12.9 (D6); 15.6 (W2) 69% parasite suppression	Coloratane sesquiterpenes, e.g., muzigadiolide	Cytotoxic to the human glioblastoma cell line U87 CD4 CXCR4 ( $CC_{50}$ = 7.2 and 2.0 µg/ml for ethanol and DMSO extracts	[60, 61, 95, 99–101]
<sup>1</sup> Solvents used: DCM = dichlore	omethane, MeOH = me	thanol. <sup>2</sup> Antiplasmodial activity are	e reported as ICso (ua/mL). <i>Plasmodi</i>	um falciparum isolates: chloroguine	sensitive strains are 3D7, MRC-2, D	6, D10, FCR3, and NF54;

### Abbreviations

DCM	Dichloromethane
ICF	Informant consensus factor
FC	Frequency of citation
ITN	Insecticide-treated mosquito nets
MeOH	Methanol
WHO	World Health Organization

# **Supplementary Information**

The online version contains supplementary material available at https://doi. org/10.1186/s41182-023-00541-9.

Additional file 1. Details of medicinal plants used in the management of malaria in Bwambara Subcounty, Rukungiri District, Uganda.

#### Acknowledgements

The authors acknowledge the local communities and key informants for sharing this valuable indigenous knowledge. Miss Racheal Akatuhebwa is highly appreciated for proofreading this work.

#### Author contributions

HG, EAO and PM conceived and designed the study. HG collected ethnobotanical data. HG, JBL and TO analyzed and interpreted the collected data. HG and TO wrote the first draft of the manuscript. HG, EAO, PM, JBL and TO revised, read and approved the final manuscript.

#### Funding

This research was supported by the Faculty of Science, Mbarara University of Science and Technology, Mbarara, Uganda (No. PF/Conf-2022/1).

## Availability of data and materials

The raw data supporting the conclusions of this study are available within this article and its supplementary files.

## Declarations

### Ethics approval and consent to participate

This study was approved by the Uganda National Council for Science and Technology (Approval no. NS34ES). The Resident District Commissioner (RDC) of Rukungiri District and the Local Council I (LC I) were consulted and subsequently approved the study to be done in the respective villages. The respondents were first informed about the study objectives and those who accepted to participate gave a written informed consent at the beginning of the study.

### **Consent for publication**

Not applicable.

#### **Competing interests**

The authors declare that there is no conflict of interest regarding the publication of this paper.

Received: 25 June 2023 Accepted: 25 August 2023 Published online: 30 August 2023

#### References

- Yi B, Zhang L, Yin J, Zhou S, Xia Z. 1-3-7 surveillance and response approach in malaria elimination: China's practice and global adaptions. Malar J. 2023;22:152.
- WHO. World Malaria Report 2022. World Health Organization 2022. https://apps.who.int/iris/rest/bitstreams/1484818/retrieve . Accessed 06 May 2023.

- Global Health Policy. The President's Malaria Initiative and Other U.S. Government Global Malaria Efforts. https://www.kff.org/globalhealth-policy/fact-sheet/the-presidents-malaria-initiative-and-otheru-s-government-global-malaria-efforts/. Accessed 20 Apr 2023.
- WHO. Malaria. https://www.who.int/news-room/fact-sheets/detail/ malaria. Accessed 10 Jun 2023.
- Nuwa A, Oola J, Obonyo SO, Feldman M, Karungi S, Kertho E, Odong DS, Kimera I, Magumba G, Beinomugisha G, Chitty A, Tibenderana J, Opigo J, Abwaimo F. District-led malaria surveillance and response as an effective way to manage malaria upsurges following the withdrawal of indoor residual spraying: a case study from Nwoya District, Northern Uganda. Malar J. 2022;21:55.
- Zalwango JF, Nankabirwa JI, Kitutu FE, Akunzirwe R, Buhuguru R, Rokani JB, Ssendikwanawa E, et al. Malaria diagnostic and treatment practices for febrile children under 5 years at two general hospitals in Karamoja, a high transmission setting in Uganda. Malar J. 2022;21:312.
- MOH. Weekly Malaria Report. Ministry of Health-Uganda. http://libra ry.health.go.ug/sites/default/files/resources/Malaria%20Bulletin-Epi% 20week%2026.pdf. Accessed 20 Jun 2023.
- Lal S, Ndyomugenyi R, Alexander ND, Lagarde M, Paintain L, Magnussen P, et al. Health facility utilisation changes during the introduction of community case management of malaria in South Western Uganda: an interrupted time series approach. PLoS ONE. 2015;10: e0137448.
- Asua V, Conrad MD, Aydemir O, Duvalsaint M, Legac J, Duarte E, Tumwebaze P, Chin DM, Cooper RA, Yeka A, Kamya MR, Dorsey G, Nsobya SL, Bailey J, Rosenthal PJ. Changing prevalence of potential mediators of aminoquinoline, antifolate, and artemisinin resistance across Uganda. J Infect Dis. 2021;223:985–94.
- Nasim N, Sandeep IS, Mohanty S. Plant-derived natural products for drug discovery: current approaches and prospects. Nucleus. 2022;65:399–411.
- Muangphrom P, Seki H, Fukushima EO, Muranaka T. Artemisinin-based antimalarial research: application of biotechnology to the production of artemisinin, its mode of action, and the mechanism of resistance of *Plasmodium* parasites. J Nat Med. 2016;70:318–34.
- Tugume P, Nyakoojo C. Ethno-pharmacological survey of herbal remedies used in the treatment of paediatric diseases in Buhunga parish, Rukungiri District, Uganda. BMC Complement Altern Med. 2019;19:353.
- Gumisiriza H, Sesaazi CD, Olet EA, Kembabazi O, Birungi G. Medicinal plants used to treat "African" diseases by the local communities of Bwambara sub-county in Rukungiri District, Western Uganda. J Ethnopharmacol. 2021;268: 113578.
- Gumisiriza H, Birungi G, Lejju JB, Olet EA, Kembabazi O, Duncan C, Sesaazi CD. Ethnobotany and antimicrobial activity of *Gouania longispicata* Engl. J Complement Med Res. 2020;11:86–94.
- Hartter J, Stampone MD, Ryan SJ, Kirner K, Chapman CA, Goldman A. Patterns and perceptions of climate change in a biodiversity conservation hotspot. PLoS ONE. 2012;7: e32408.
- 16. Omara T. Plants used in antivenom therapy in rural Kenya: ethnobotany and future perspectives. J Toxicol. 2020;2020:1828521.
- Common wealth network. Uganda. Rukungiri. 2020. https://www. commonwealthofnations.org/info/regions-uganda/western-region/ rukungiri/#:~:text=The%20majority%20of%20the%20workforce. Accessed on 02 Aug 2023.
- Gumisiriza H, Birungi G, Olet EA, Sesaazi CD. Medicinal plant species used by local communities around Queen Elizabeth National Park, Maramagambo Central Forest Reserve and Ihimbo Central Forest Reserve, South western Uganda. J Ethnopharmacol. 2019;239: 111926.
- African Development Bank. Road Sector Support Project 5 (RSSP V) upgrading of Rukungiri–Kihihi–Ishasha/Kanungu and Mbale-BumbobiManafwa-Lwakhakha Roads. 2013. https://www.afdb.org/en/docum ents/document/uganda-road-sector-support-project-5-rssp-v-upgra ding-of-rukungiri-kihihi-ishasha-kanungu-and-mbale-bumbobi-manaf wa-lwakhakha-roads-esia-rap-summary-44822. Accessed 16 Aug 2023.
- Monitor. Residents turn mosquito nets into building materials. 2021. https://www.monitor.co.ug/uganda/news/national/residents-turn-mosquito-nets-into-building-materials-3461638. Accessed 02 Aug 2023.

- Kigozi SP, Kigozi RN, Sebuguzi CM, Cano J, Rutazaana D, Opigo J, et al. Spatial-temporal patterns of malaria incidence in Uganda using HMIS data from 2015 to 2019. BMC Public Health. 2020;20:1913.
- 22. Krejcie RV, Morgan DW. Determining sample size for research activities. Educ Psychol Meas. 1970;30:607–10.
- Trotter RJ, Logan MH. Informant consensus. A new approach for identifying potentially effective medicinal plants. In: Etkin NL, editor. Plants in indigenous medicine and diet. Bedford Hills: Newyork: Redgrave; 1986, pp. 22.
- 24. Tabuti JRS, Obakiro SB, Nabatanzi A, Anywar G, Nambejja C, Mutyaba MR, Omara T, Waako P. Medicinal plants used for treatment of malaria by indigenous communities of Tororo District, Eastern Uganda. Trop Med Health. 2023;51:34.
- Adia MM, Anywar G, Byamukama R, Kamatenesi-Mugisha M, Sekagya Y, Kakudidi EK, Kiremire BT. Medicinal plants used in malaria treatment by Prometra herbalists in Uganda. J Ethnopharmacol. 2014;155:580–8.
- Asiimwe S, Namutebi A, Borg-Karlsson A, Kamatenesi-Mugisha M, Oryem-Origa H. Documentation and consensus of indigenous knowledge on medicinal plants used by the local communities of western Uganda. J Nat Prod Plant Resour. 2014;4:34–42.
- Nalumansi P, Kamatenesi-Mugisha M, Anywar G. Medicinal plants used in paediatric health care in namungalwe sub county, Iganga district, Uganda. Nova J Med Biol Sci. 2014;2:1–14.
- Linden I. The History of Kigezi—a history of Kigezi in South-West Uganda. Edited by Donald Denoon. The National Trust. Adult Education Centre, Kampala, Uganda, 1972. pp. 302. J Afr Hist. 1972; 13: 694–695.
- Walusansa A, Asiimwe S, Ssenku JE, Anywar G, Namara M, Nakavuma JL, Kakudidi EK. Herbal medicine used for the treatment of diarrhea and cough in Kampala city, Uganda. Trop Med Health. 2022;50:5.
- Ngarivhume T, Van't Klooster CI, de Jong JT, Van der Westhuizen JH. Medicinal plants used by traditional healers for the treatment of malaria in the Chipinge district in Zimbabwe. J Ethnopharmacol. 2015;159:224–37.
- Olala CN. Identification of plants used for treatment of malaria and factors influencing their use in Boro division, Siaya county, Kenya. MSc Thesis, Kenyatta University, Nairobi, Kenya. 2014.
- Kaendi JM. Coping with Malaria and Visceral Leishmaniasis (Kala-azar) in Baringo District, Kenya: Implications for Disease Control. PhD Thesis, University of California, Los Angeles, CA, USA. 1994.
- Sindiga I. Indigenous medical knowledge of the Maasai. Indig Knowl Dev Monit. 1994;2:16–8.
- Kiringe JW. A survey of traditional health remedies used by the Maasai of Southern Kaijiado district, Kenya. Ethnobot Res Appl. 2006;4:61–73.
- 35. Tabuti J. Herbal medicines used in the treatment of malaria in Budiope county, Uganda. J Ethnopharmacol. 2008;116:33–42.
- Stangeland T, Alele PE, Katuura E, Lye KA. Plants used to treat malaria in Nyakayojo sub-county, western Uganda. J Ethnopharmacol. 2011;137:154–66.
- Mukungu N, Abuga K, Okalebo F, Ingwela R, Mwangi J. Medicinal plants used for management of malaria among the Luhya community of Kakamega East sub-County, Kenya. J Ethnopharmacol. 2016;194:98–107.
- Omara T. Antimalarial plants used across Kenyan communities. Evid Based Complement Altern Med. 2020;2020:4538602.
- WHO. World Malaria Report 2021. World Health Organization 2021. https://www.who.int/teams/global-malaria-programme/reports/world-malaria-report-2021. Accessed 17 Jun 2023.
- Opio D, Andama E, Kureh GT. Ethnobotanical survey of antimalarial plants in areas of: Abukamola, Angeta, Oculokori and Omarari of Alebtong District in Northern Uganda. EJMP. 2018;21:1–14.
- Suleman S, Beyene Tufa T, Kebebe D, Belew S, Mekonnen Y, Gashe F, Mussa S, Wynendaele E, Duchateau L, De Spiegeleer B. Treatment of malaria and related symptoms using traditional herbal medicine in Ethiopia. J Ethnopharmacol. 2018;213:262–79.
- 42. Obakiro SB, Kiprop A, Kowino I, Kigondu E, Odero MP, Omara T, Bunalema L. Ethnobotany, ethnopharmacology, and phytochemistry of traditional medicinal plants used in the management of symptoms of tuberculosis in East Africa: a systematic review. Trop Med Health. 2020;48:68.

- Okello D, Kang Y. Exploring antimalarial herbal plants across communities in Uganda based on electronic data. Evid Based Complement Alternat Med. 2019;2019:3057180.
- 44. Lee YJ, Adusumilli G, Kazungu R, Anywar G, Kyakulaga F, Katuura E, Parikh S, Willcox M. Treatment-seeking behavior and practices among caregivers of children aged ≤5 y with presumed malaria in rural Uganda. Trans R Soc Trop Med Hyg. 2019;131:525–33.
- 45. Anywar G, Byamukama R, vant Klooster CIEA, Wilcox M, Nalumansi P, de Jong J, Rwaburindori P, Kiremire BT. Medicinal plants used in the treatment and prevention of malaria in Cegere sub-county, Northern Uganda. J Ethnobot Appl Res. 2016;14:505–16.
- Pierre S, Toua V, Tchobsala, Tchuenguem FF, Alexandre-Michel NN, Jean M. Medicinal plants used in traditional treatment of malaria in Cameroon. J Ecol Nat Environ. 2011; 3: 104–117.
- Taek MM, Bambang PEW, Mangestuti A. Plants used in traditional medicine for treatment of malaria by Tetun ethnic people in West Timor Indonesia. Asian Pac J Trop Med. 2018;11:630–7.
- Alebie G, Urga B, Worku A. Systematic review on traditional medicinal plants used for the treatment of malaria in Ethiopia: trends and perspectives. Malaria J. 2017;16:307.
- Ssenku JE, Okurut SA, Namuli A, Kudamba A, Tugume P, Matovu P, Wasige G, Kafeero HM, Walusansa A. Medicinal plant use, conservation, and the associated traditional knowledge in rural communities in Eastern Uganda. Trop Med Health. 2022;50:39.
- Namukobe J, Kasenene JM, Kiremire BT, Byamukama R, Kamatenesi-Mugisha M, Krief S, Dumontet V, Kabasa JD. Traditional plants used for medicinal purposes by local communities around the Northern sector of Kibale National Park, Uganda. J Ethnopharmacol. 2011;136:236–45.
- Dossou AJ, Fandohan AB, Omara T, Gbenou J. Traditional knowledge and phytochemical screening of plants used in snakebite prevention in Benin. Bull Natl Res Cent. 2022;46:160.
- 52. Kuria KA, De Coster S, Muriuki G, Masengo W, Kibwage I, Hoogmartens J, Laekeman GM. Antimalarial activity of *Ajuga remota* Benth (Labiatae) and *Caesalpinia volkensii* Harms (Caesalpiniaceae): *in vitro* confirmation of ethnopharmacological use. J Ethnopharmacol. 2001;74:141–8.
- Murphy MW, Dunton RF, Perich MJ, Rowley WA. Attraction of *Anopheles* (Diptera: culicidae) to volatile chemicals in Western Kenya. J Med Entomol. 2001;38:242–4.
- Lacroix D, Prado S, Kamoga D, Kasenene J, Namukobe J, Krief S, Dumontet V, Mouray E, Bodo B, Brunois F. Antiplasmodial and cytotoxic activities of medicinal plants traditionally used in the village of Kiohima, Uganda. J Ethnopharmacol. 2011;133:850–5.
- Nyamwamu LB, Ngeiywa M, Mulaa M, Lelo AE, Ingonga J, Kimutai A. Acute toxicity of Senna didymobotrya Fresen Irwin roots used as a traditional medicinal plant in Kenya. Am Int J Contemp Res. 2015;5:74–7.
- Barlow-Benschop NM, Gamba C, Barlow SP, Blasco TM. The effect of a homeopathic neem preparation for the prophylaxis of malaria. An exploratory trial in an at-home setting in Tanzania, pp. 1–12. https:// pdfs.semanticscholar.org/177a/390381a2a7c7a19566a20f57330f50 751101.pdf.
- Challand S, Willcox M. A clinical trial of the traditional medicine Vernonia amygdalina in the treatment of uncomplicated malaria. J Alternat Compl Med. 2009;15:1231–7.
- Kumar S, Yadav M, Yadav A, Rohilla P, Yadav JP. Antiplasmodial potential and quantification of aloin and aloe-emodin in *Aloe vera* collected from different climatic regions of India. BMC Complement Altern Med. 2017;17:369.
- 59. Van Zyl R, Viljoen A. In vitro activity of *Aloe* extracts against *Plasmodium falciparum*. South Afr J Bot. 2002;68:106–10.
- Muthaura CN, Keriko JM, Mutai C, Yenesew A, Gathirwa JW, Irungu BN, Nyangacha R, Mungai GM, Derese S. Antiplasmodial potential of traditional antimalarial phytotherapy remedies used by the Kwale community of the Kenyan Coast. J Ethnopharmacol. 2015;170:148–57.
- Anywar GU, Kakudidi E, Oryem-Origa H, Schubert A, Jassoy C. Cytotoxicity of medicinal plant species used by traditional healers in treating people suffering from HIV/AIDS in Uganda. Front Toxicol. 2022;4: 832780.
- 62. Omara T, Kiprop AK, Kosgei VJ. Isolation and characterization of compounds in ethanolic extract of *Albizia coriaria* (Welw ex. Oliver) leaves:

a further evidence of its ethnomedicinal diversity. Bull Natl Res Cent. 2022;46:30.

- 63. Omara T, Kiprop AK, Kosgei VJ. Two new pentacyclic triterpenoids, an alkaloid and a long-chain fatty acid from *Albizia coriaria* (Welw ex. Oliver). Fr-Uk J Chem. 2022;10:128.
- 64. Asase A, Akwetey GA, Achel DG. Ethnopharmacological use of herbal remedies for the treatment of malaria in the Dangme West District of Ghana. J Ethnopharmacol. 2010;129:367–76.
- Bray DH, Warhurst DC, Connolly JD, O'neill MJ, Phillipson JD. Plants as sources of antimalarial drugs. Part 7. Activity of some species of Meliaceae plants and their constituent limonoids. Phytother Res. 1990;4:29–35.
- 66. Khalid SA. Isolation and characterization of antimalarial agents of the neem tree *Azadirachta indica*. J Nat Prod. 1989;52:922–7.
- Kirira PG, Rukunga GM, Wanyonyi AW, Muregi FM, Gathirwa JW, Muthaura CN, Omar SA, Tolo F, Mungai GM, Ndiege IO. Anti-plasmodial activity and toxicity of extracts of plants used in traditional malaria therapy in Meru and Kilifi Districts of Kenya. J Ethnopharmacol. 2006;106:403–7.
- Nanyingi MO, Kipsengeret KB, Wagate CG, Langat BK, Asaava LL, Midiwo JO. In vitro and in vivo antiplasmodial activity of Kenyan medicinal plants. In: J. O. Midiwo and J. Clough, editors. Aspects of African biodiversity: proceedings of the Pan-Africa chemistry network, RCS Publishing, Cambridge, UK; 2010. pp. 20–28.
- Clarkson C, Maharaj VJ, Crouch NR, Grace OM, Pillay P, Matsabisa MG, Bhagwandin N, Smith PJ, Folb PI. In vitro antiplasmodial activity of medicinal plants native to or naturalised in South Africa. J Ethnopharmacol. 2004;92:177–91.
- Frida L, Rakotonirina S, Rakotonirina A, Savineau JP. In vivo and in vitro effects of *Bidens pilosa* L. (Asteraceae) leaf aqueous and ethanol extracts on primed-oestrogenized rat uterine muscle. Afr J Tradit Complement Altern Med. 2007;5:79–91.
- Lai BY, Chen TY, Huang SH, Kuo TF, Chang TH, Chiang CK, Yang MT, Chang CL. *Bidens pilosa* formulation improves blood homeostasis and β-cell function in men: a pilot study. Evid Based Complement Alternat Med. 2015;2015: 832314.
- Teng WC, Chan R, Suwanarusk W, Ong A, Ho HK, Russell B, Rénia L, Koh HL. In vitro antimalarial evaluations and cytotoxicity investigations of *Carica papaya* leaves and carpaine. Nat Prod Comm. 2019;14:33–6.
- Melariri P, Campbell W, Etusim P, Smith P. Antiplasmodial properties and bioassay-guided fractionation of ethyl acetate extracts from *Carica* papaya Leaves. J Parasitol Res. 2011;2011: 104954.
- Julianti T, De Mieri M, Zimmermann S, Ebrahimi SN, Kaiser M, Neuburger M, Raith M, Brun R, Hamburger M. HPLC-based activity profiling for antiplasmodial compounds in the traditional Indonesian medicinal plant *Carica papaya* L. J Ethnopharmacol. 2014;155:426–34.
- Omara T, Kiprop AK, Kosgei VJ, Kagoya S. Clausena anisata (Willd.) Hook.f. ex. Benth. (Rutaceae): ethnomedicinal uses, phytochemistry, pharmacological activities, toxicity, and clinical application. Tradit Med Res. 2022;7:51.
- Igoli JO, Masao CA, Orkpeh U, Nakamatte E, Yakumbur DT, Nnadozie T, et al. In vivo antimalarial activity of *Cleome gynandra* extracts. J Nat Prod Res Updates. 2016;2:19–29.
- Busola OA, Iwuji IG, Nasir HA, Abdulrahman A. Antimalarial activity of *Clerodendrum capitatum* (Willd) ethanol leaf extract in *Plasmodium berghei* infected mice. Afr J Pharmaceut Res Dev. 2020;12:248–56.
- Obakiro SB, Kiprop A, Kigondu E, K'Owino I, Odero MP, Manyim S, Omara T, Namukobe J, Owor RO, Gavamukulya Y, Bunalema L. Traditional medicinal uses, phytoconstituents, bioactivities, and toxicities of *Erythrina abyssinica* Lam. ex DC. (Fabaceae): a systematic review. Evid Based Complement Alternat Med. 2021;2021:5513484.
- 79. Oketch-Rabah HA, Dossaji SF, Mberu EK. Antimalarial activity of some Kenyan medicinal plants. Pharmaceut Biol. 1999;37:329–34.
- Farah H, El Hussein MA, Khalid EH, Osman HM. Toxicity of *Kigelia africana* fruit in rats. Adv Res. 2018;12:1–9.
- Lacroix D, Prado S, Deville A, Krief S, Dumontet V, Kasenene J, Mouray E, Bories C, Bodo B. Hydroperoxy-cycloartane triterpenoids from the leaves of *Markhamia lutea*, a plant ingested by wild chimpanzees. Phytochemistry. 2009;70:1239–45.

- Azanze NE, Mbiantcha M, Madjo KYK, Yousseu NW, Fagni Njoya ZL, Adjouzem CF, Marthe VM, Ateufack G. *Markhamia lutea* leaves aqueous and ethanolic extract with curative anti-inflammatory activity attenuates paclitaxel toxicity in rat's intestine. J Complement Integr Med. 2023. https://doi.org/10.1515/jcim-2023-0017.
- Adia MM, Emami SN, Byamukama R, Faye I, Borg-Karlson AK. Antiplasmodial activity and phytochemical analysis of extracts from selected Ugandan medicinal plants. J Ethnopharmacol. 2016;186:14–9.
- Froelich S, Onegi B, Kakooko A, Siems K, Schubert C, Jenett-Siems K. Plants traditionally used against malaria: phytochemical and pharmacological investigation of *Momordica foetida*. Rev Bras Farmacogn. 2007;17:1–17.
- Obbo CJD, Kariuki ST, Gathirwa JW, Olaho-Mukani W, Cheplogoi PK, Mwangi EM. In vitro antiplasmodial, antitrypanosomal and antileishmanial activities of selected medicinal plants from Ugandan flora: refocusing into multi-component potentials. J Ethnopharmacol. 2019;229:127–36.
- Olanlokun JO, Bodede O, Prinsloo G, Olorunsogo OO. Comparative antimalarial, toxicity and mito-protective effects of *Diospyros mespiliformis* Hochst. ex A. DC. and Mondia whitei (Hook.f.) Skeels on *Plasmodium berghei* infection in mice. J Ethnopharmacol. 2021;268: 113585.
- Oloro J, Tanayen J, Barbra K, Lawrence I, Waako P, Francis B, Amon A. Toxicity of four herbs used in erectile dysfunction: *Mondia whiteii, Cola acuminata, Urtica massaica,* and *Tarenna graveolensin* male rats. Afr J Pharm Pharmacol. 2015;9:756–63.
- Bezerra JJL, Pinheiro AAV, Dourado D. Antimalarial potential of *Moringa* oleifera Lam. (Moringaceae): a review of the ethnomedicinal, pharmacological, toxicological, and phytochemical evidence. J Venom Anim Toxins Incl Trop Dis. 2023;29: e20220079.
- Owuor BO, Ochanda JO, Kokwaro JO, Cheruiyot AC, Yeda RA, Okudo CA, Akala HM. In vitro antiplasmodial activity of selected Luo and Kuria medicinal plants. J Ethnopharmacol. 2012;144:779–81.
- Njan AA, Olaoye SO, Afolabi SO, Ejimkonye BC, Soje A, Olorundare OE, Iwalewa EO. Safety effect of fractions from methanolic leaf extract of *Ocimum gratissimum* on reproduction in male Wistar rats. Toxicol Rep. 2019;6:496–504.
- Kiraithe MN, Nguta JM, Mbaria JM, Kiama SG. Evaluation of the use of Ocimum suave Willd (Lamiaceae), Plectranthus barbatus Andrews (Lamiaceae) and Zanthoxylum chalybeum Engl (Rutaceae) as antimalarial remedies in Kenyan folk medicine. J Ethnopharmacol. 2016;178:266–71.
- Muregi FW, Chhabra SC, Njagi EN, Lang'at-Thoruwa CC, Njue WM, Orago AS, Omar SA, Ndiege IO. In vitro antiplasmodial activity of some plants used in Kisii, Kenya against malaria and their chloroquine potentiation effects. J Ethnopharmacol. 2003;84:235–9.
- Kigondu EV, Rukunga GM, Keriko JM, Tonui WK, Gathirwa JW, Kirira PG, Irungu B, Ingonga JM, Ndiege IO. Anti-parasitic activity and cytotoxicity of selected medicinal plants from Kenya. J Ethnopharmacol. 2009;123:504–9.
- 94. Muganga R, Angenot L, Tits M, Frédérich M. Antiplasmodial and cytotoxic activities of Rwandan medicinal plants used in the treatment of malaria. J Ethnopharmacol. 2010;128:52–7.
- Onguéné P, Ntie-Kang F, Lifongo LL, Ndom JC, Sippl W, Mbaze LM. The potential of anti-malarial compounds derived from African medicinal plants, part I: a pharmacological evaluation of alkaloids and terpenoids. Malar J. 2013;12:449.
- Goffin E, Ziemons E, De Mol P, de Madureira MC, Martins AP, da Cunha AP, Philippe G, Tits M, Angenot L, Frederich M. In vitro antiplasmodial activity of *Tithonia diversifolia* and identification of its main active constituent: tagitinin C. Planta Med. 2002;68:543–5.
- Oyewole IO, Ibidapo CA, Moronkola DO, Oduola AO, Adeoye GO, Anyasor GN, Obansa JA. Anti-malarial and repellent activities of *Tithonia diversifolia* (Hemsl.) leaf extracts. J Med Plants Res. 2008;2:171–5.
- Omoregie ES, Pal A, Sisodia B. In vitro antimalarial and cytotoxic activities of leaf extracts of *Vernonia amygdalina* (Del.). Niger J Basic Appl Sci. 2011;19:121–6.
- 99. Wube A, Bucar F, Gibbons S, Asres K, Rattray L, Croft SL. Anti-protozoal activity of sesquiterpenes from *Warburgia ugandensis* towards *Trypanosoma bruceirhodesiense* and *Plasmodium falciparum* in vitro. Planta Med. 2008;74:PA222.
- 100. Were PS, Kinyanjui P, Gicheru MM, Mwangi E, Ozwara HS. Prophylactic and curative activities of extracts from *Warburgia ugandensis* Sprague

(Canellaceae) and Zanthoxylum usambarense (Engl) Kokwaro (Rutaceae) against Plasmodium knowlesi and Plasmodium berghei. J Ethnopharmacol. 2010;130:158–62.

101. Okello D, Komakech R, Matsabisa MG, Kang YM. A review on the botanical aspects, phytochemical contents and pharmacological activities of *Warburgia ugandensis*. J Med Plants Res. 2018;12:448–55.

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