

Medicinal Plants Used in The Management of Epilepsy in Nigeria: A Review of Potential Targets for Drug Discovery

Abubakar Sadiq Wada^{1*}, Abubakar Rabi Abdullahi¹, Abubakar Sule Danbatta¹, Mustapha Mohammed^{2,3}, Mubarak Hussaini Ahmad⁴, Sani Malami¹ and Abdullahi Hamza Yaro¹

¹Department of Pharmacology and Therapeutics, Bayero University, Kano, Nigeria

²School of Pharmaceutical Sciences, Universiti Sains Malaysia, Penang, Pulau Pinang, Malaysia

³Department of Clinical Pharmacy and Pharmacy Practice, Faculty of Pharmaceutical Sciences, Ahmadu Bello University, Zaria, Kaduna, Nigeria

⁴Department of Pharmacology and Therapeutics, Faculty of Pharmaceutical Sciences, Ahmadu Bello University, Zaria, Kaduna, Nigeria

⁵Department of Pharmacology and Therapeutics, Faculty of Pharmaceutical Sciences, Capital City University, Kano, Nigeria

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* Corresponding Author:

pharmwada17@gmail.com
+2347037763700
<https://orcid.org/0000-0001-8221-1159>

ABSTRACT

Background: Medicinal plants are continuously used to manage epilepsy and other neurological disorders. They provide major promising targets in pursuit of new drugs and lead compounds that are affordable, available and accessible to treat the debilitating neurological condition. Several experimental models employed to screen for seizures in laboratory animals provide beneficial information concerning diagnoses, treatment and possible prevention of the disease. This review aims to identify medicinal plants used in the management of epilepsy in Nigeria and to explore pharmacological basis to support their ethnobotanical claims.

Methods: Literature searches of relevant articles in electronic databases including PubMed, African journal online, Google Scholar and ScienceDirect databases were carried out, and information about how these medicinal plants are used traditionally in the management of epilepsy and other diseases in Nigeria were also obtained. Only studies conducted within Nigeria on medicinal plants tested for seizures and epilepsy between 2000 and 2022 were included.

Results: We identified sixty-eight (68) medicinal plants spanning across several families majorly Agavaceae, Amaryllidaceae, Annonaceae, Apocynaceae, Asteraceae, Bignoniaceae, Burseraceae, Compositae, Convolvulaceae, Euphorbiaceae, Lamiaceae, Leguminosae, Lorantheaceae, Moraceae and Rubiaceae, that have been reported to contain bioactive compounds active against seizures using various pharmacological screening models. Plants that have not been fully studied and their main mechanisms of action not ascertained were recorded. We also identified those plants with unknown active constituents responsible for their activity. The review also identified potential medicinal plants for future studies of new as well as alternative therapies for the management of epilepsy and other neurological and neurodegenerative diseases.

Conclusions: This review provided evidence on the use of medicinal plants in the management of epilepsy and possibly rationalized the use of these plant extracts as alternatives in treating seizures and epilepsy.

1. Background

Epilepsy is a chronic neurological disorder characterized by recurrent seizures that occur due to abnormal electrical discharges of groups of neurons in the brain¹⁻³. The commonest causes of epilepsy include severe hypoxia or

birth asphyxia, brain tumors, intracranial trauma during birth, metabolic disturbances, traumatic head injuries, road traffic accidents and congenital malformations of the brain, perinatal insults or infection such as meningitis, cerebral malaria, febrile seizures and encephalitis^{2,4-8}. Remission of

recurrent and unprovoked seizures can be achieved in most cases by treatment with anticonvulsant medication, surgical resection, or nerve-brain electrical stimulation. However, one-third of epileptic patients are still refractory to treatment, this necessitates addressing such challenges⁹⁻¹³. The currently available antiepileptic drugs (AEDs) neither provide a definite clinical cure nor prevent its occurrence or relapse. Moreover, the AEDs are often accompanied by several side effects, including cognitive dysfunction, sedation, blood dyscrasias and teratogenicity. The absence of disease-modifying therapies for epilepsy is also a major concern in the quality of life of these patients¹²⁻¹⁶. About 80% of people with epilepsy living in developing countries receive less or no drug treatment for epilepsy¹⁷⁻¹⁹. Inadequacy, cost of AEDs, lack of access to modern healthcare facilities and stigmatization of patients and their caregivers hinders these patients with epilepsy from seeking proper management¹⁹⁻²⁰. Medicinal plants, however, provide major promising targets in pursuing new drugs and lead compounds that will be affordable and accessible to treat these debilitating neurological disorders²¹⁻²². In Northwestern Nigeria, about 11.9% of patients with epilepsy are reported to take at least a form of an alternative therapy before visiting the health care centres²³. This is perhaps because medicinal plants are seen as natural, harmless to health, relatively cheap compared to orthodox medicines, easily accessible, locally and culturally acceptable and economically affordable, as well as efficacious in curing most ailments²⁴⁻²⁶. Despite appreciable studies on anticonvulsant activities of medicinal plants in Nigeria and other developing countries, paucity of reliable evidences regarding their efficacy and safety as well as inadequate information about their mechanism of action are domineering weaknesses regarding their effectiveness for use in treatments²⁵. This review provides scientific evidence for using various medicinal plants with anticonvulsant activity and portrays the extent of the validated anticonvulsant profile of these plants in Nigeria.

2. Methods

Literature searches of relevant articles in electronic databases including PubMed, African journal online, Google Scholar and Science Direct databases was carried out using 'Epilepsy', 'Convulsion', 'Nigeria', 'Medicinal plants' as keywords for the search. Information about the traditional usage of these medicinal plants in the management of epilepsy and other diseases in Nigeria were also gotten from published papers and texts on

ethnobotanical studies. Only research studies carried out in Nigeria that investigate the effect of medicinal plants on epilepsy and experimental models used to validate such ethnobotanical claims of these plants in Nigeria between January 2000 and November 2022 were included Table 1. This review excludes all articles published before the year 2000. Studies conducted on medicinal plants outside Nigeria were also not included. Several publications obtained from various universities and research institutions that fulfilled the inclusion criteria were also considered. The articles chosen were thoroughly scrutinized for eligibility (Figure 1).

4. Results

The extracts obtained from the references were prepared either from the whole plant, leaves, stem barks, and root barks extracted in different polar solvents; aqueous, methanolic, ethanolic, hydroalcoholic, butanol and ethyl acetate. Rodents such as mice and rats and day-old chicks were used for the screening for anticonvulsant activity. The medicinal plants reported in this review span across several families such as: Agavaceae, Amaryllidaceae, Anacardiaceae, Annonaceae, Apocynaceae, Asteraceae, Bignoniaceae, Burseraceae, Cochlospermaceae, Compositae, Convolvulaceae, Crassulaceae, Cucurbitaceae, Cyperaceae, Ebanaceae, Euphorbiaceae, Fabaceae, Icacinaceae, Lamiaceae, Leguminosae, lilaceae, Loranthaceae, Meliaceae, Mimosaceae, Moraceae, Moringaceae, Ochnaceae, Olacaceae, Piperaceae, Poaceae, Polygalaceae, Rubiaceae, Rutaceae, Sapindaceae, Sapotaceae, Scrophulariaceae, Solanaceae, Ulmaceae, Vitaceae²⁻⁵

Several medicinal plants in Nigeria have shown appreciable activity against MEST in this review. Notably, at doses of 25 and 50 mg/kg body weight, the methanolic extract of *Chrysanthellum Indicum* protected the chicks (80%) against MEST²⁶, as compared to the saline control group. Likewise, methanolic extract of *Olox subscorpioidea* at doses of 100 and 200 mg/kg provided 30 and 70% protection against MEST in chicks respectively. At all doses tested (100, 200, and 400 mg/kg), the extract significantly ($p < 0.05$) increased the latency of seizures induced by MEST compared to the normal saline control group²⁷. A seizure protection (100%) against MEST-induced seizures at 400 mg/kg of *Evolvulus alsinoides* (in chicks) and *Leucas martinicensis* (in rats), respectively were also reported²⁸⁻²⁹. Other medicinal plants reported to have activity against MEST include *Laggera Aurita*³⁰, *Paullinia pinnata*³¹, *Randia nilotica*³², *Solanum nigrum*³³,

*Afrormosia laxiflora*³⁴, *Securidaca longipedunculata*³⁵ and *Allium cepa*³⁶.

The extract of *Evolvulus alsinoides*, at all doses tested, significantly increased the latency of PTZ-induced seizure and protected 100% of the mice against seizure at the highest dose of 400 mg/kg²⁸. The extract of *A. chevalieri* at 300 mg/kg significantly ($P < 0.01$) increased the mean onset of seizures induced by PTZ³⁷. The aqueous stem bark extract of *Securidaca longipedunculata* at 50 and 100 mg/kg body weight protected (80%) of animals and significantly ($p < 0.05$) prolonged the onset of convulsion³⁵. Dose-dependent protection against PTZ-induced seizure in mice, with complete protection (100%) against seizure at the dose of 30 mg/kg, was observed with aqueous extract of *Solanum nigrum*³³. Other medicinal plants such as *Carissa edulis*³⁸, *Cissus cornifolia*³⁹, *Cochlospermum tinctorium*³¹, *Diospyros mespiliformis*⁴⁰, *Ficus platyphylla*¹⁶, *Globimetula braunii*⁴¹, *Hymenocardia acida*⁴²⁻⁴³, *Laggetera aurita*³⁰, *Paullinia pinnata*³¹, *Albizia glaberrima*⁴⁴, *Moringa oleifera*⁴⁵, *Lannea barteri*⁴⁶, *Piper guineense*⁴⁷, *Securinega virosa*⁴⁸, *Afrormosia laxiflora*³⁴, *Olex subscorpioidea*²⁷ and *Celtis integrifolia*⁴⁹ have demonstrated activity against PTZ induced seizures.

Extract of *Leucas martinicensis* at 200 mg/kg and 400 mg/kg offers protection (100%) to rats against STR-induced seizures⁵⁰. The methanolic extract of *Olex subscorpioidea* offers 50% protection against strychnine-induced seizures in mice at a dose of 100 mg/kg. The extract significantly ($p < 0.01$) prolonged the mean onset of seizures²⁷. Other medicinal plants identified in this review that offers protection to mice against strychnine-induced seizures include *Carissa edulis*³⁸, *Cissus cornifolia*³⁹, *Ficus platyphylla*¹⁶, *Hymenocardia acida*⁴²⁻⁴³, *Sansevieria liberica*⁵¹, *Mitragyna africana*⁵².

*Cissus cornifolia*³⁹, *Hymenocardia acida*⁴²⁻⁴³, *Celtis integrifolia*⁴⁹, *A. chevalieri*³⁷ have been reported to show activity against 4-aminopyridine induced seizures. Medicinally active plants such as *Diospyros mespiliformis*⁴⁰, *Laggetera aurita*³⁰, *Solanum nigrum*³³, *Sansevieria liberica*⁵¹, *Russelia equisetiformis*⁵³ and *A. chevalieri*³⁷ have been found to be active against picrotoxin induced seizures in this review.

*Sansevieria liberica*⁵¹ offer protection to mice against bicuculline-induced seizures. *Carissa edulis* have been reported to offer protection to mice against NMDA-induced seizures and aminophylline-induced seizures³⁸.

The Residual aqueous fraction of *Carissa edulis* at the dose of 300 mg/kg reduced both the behavioral seizure scores and the mean seizure duration³⁸. The extract of *A. chevalieri*

also, at all doses tested, decreased the severity of seizures on the Racine scale in PTZ kindled rats, suggestive of antiepileptogenic potentials³⁷. However, there was no protection against seizure nor significant increase or decrease in seizure threshold to animals in which aqueous extract of *Solanum nigrum* was administered for 28 days and later challenged after 24-hours to electrically-induced, picrotoxin (5 mg/kg) and pentylenetetrazol (85 mg/kg) groups³³.

In an attempt to identify, isolate and characterize the active constituents responsible for anticonvulsant activity in the reported medicinal plants, the n-butanol fraction of *Securinega virosa* rich in flavonoids and saponins, protect mice against PTZ-induced seizures⁴⁸. Flavonoids and saponins are implicated to be responsible for anticonvulsant activity⁵⁴⁻⁵⁵. Several other authors, have also reportedly attributed the anticonvulsant activity to bioactive constituents such as flavonoids, saponins, tannins and alkaloids⁵⁵. Methysticin; a pyrone from the rhizomes of *Piper methysticum*. a triterpenoid glycoside from *Tetrapleura tetraptera* and *Spathodea campanulata* and Linalool; monoterpene from *Aeolanthus suaveolens*⁵⁶⁻⁵⁷ have been isolated from medicinal plants with anticonvulsant activity. Alkaloids with anticonvulsant activity have also been isolated from *Capparis baduca*, *Pithecellobium saman*, *Picnomon acarna* while cannabinoids and flavonoids from *Cannabis sativa*⁵⁶.

This review also identified a few authors that isolated active constituents and assayed for anticonvulsant activity. Bioactive-guided fractionation of methanol-methylene chloride root bark extract of *Annona senegalensis* isolated kaurenoic acid which protected mice against PTZ-induced seizures⁵⁶. Lupeol acetate, an ursolic acid also isolated from the leaf of *Milicia excelsa*⁵⁸. Ursolic acid; a triterpenoid carboxylic acid was hitherto shown to possess anticonvulsant effects⁵⁹⁻⁶⁰. Two compounds isolated and characterized from leaf extract of *Pyrenacantha staudtii* are bis(8-hydroxyl-2-methylnonyl) phthalate and bis(8-methylnonyl) phthalate. However, an assay of their anticonvulsant activity only reveals slight activity with the former while the latter does not⁵⁷.

5. Discussions

5.1 Experimental evidence in Epilepsy

Experimental models of epilepsy and seizures remain essential in understanding the mechanisms underlying ictogenesis and epileptogenesis that are instrumental in the drug discovery and development of novel antiepileptic drugs^{15,61}. Several experimental models are employed to

screen for seizures in laboratory animals either in acute or chronic models of seizures and epilepsy. Among the commonly used acute models, the Maximal electroshock seizure test (MEST) and Pentylentetrazol (PTZ) models remain the two-goal standard models of acute seizure screening. They have clearly defined endpoints foretelling equitable effects in humans and require only basic technical expertise¹⁹.

The MEST is a standard antiepileptic drugs (AEDs) test that assesses the testing material's ability to protect against the hind limb tonic extension (HLTE) phase of the seizure⁶². The MEST model induces seizures by sending electrical signals that hyper excite the neurons. MEST recognizes activity against generalized tonic-clonic/ grand mal seizures, and such activity epitomizes action on the seizure focus⁶³. Experimentally, the testing animals are observed for seizures manifesting as hind limb tonic extension (HLTE). The ability of extracts to prevent HLTE and prolong the latency and/or onset of HLTE is considered an indication of anticonvulsant activity⁶⁴⁻⁶⁶. Seizures induced by MEST can be prevented by agents inhibiting sodium channels such as valproate, phenytoin, lamotrigine and felbamate and those blocking glutamatergic excitation mediated by N-Methyl-D-Aspartate (NMDA) receptors.

PTZ exerts its convulsive effect by inhibiting the activity of gamma-amino-butyric acid (GABA) thus a suitable model to primarily identify compounds that raise the seizure threshold. GABA is the major inhibitory neurotransmitter implicated in epilepsy at GABA_A receptors, enhancement and inhibition of GABA neurotransmission do result in attenuation and enhancing convulsion, respectively⁶⁷⁻⁶⁸. Studies also suggested the possible involvement of the glutamatergic mechanism through activation of the NMDA receptor system in the initiation and propagation of PTZ-induced convulsions⁶⁹⁻⁷⁰.

Experimental animals are observed with the experimental timeline for the absence of an episode of clonic spasm for at least 5s duration, which is indicative of an extract's ability to abolish the effect of PTZ⁷¹. AEDs effective against this type of seizure are also effective in treating absence and myoclonic seizures⁷².

Strychnine is a selective and competitive antagonist that acts by blocking the inhibitory effects of glycine at the glycine receptors, extracts that are able to protect mice against lethality within 30 minutes of observation are considered to have anticonvulsant activity⁷³⁻⁷⁵.

4-aminopyridine (4-AP) is a potassium channel antagonist and a potent chemoconvulsant that induces tonic-clonic-convulsion and lethality⁷⁶. It interferes with neuronal

excitability both at resting membrane potential, responsiveness to synaptic inputs, frequency adaptation and neurotransmitters release⁷⁷. The ability of extracts to protect mice from lethality within a 30-minute observational period is considered an indication of anticonvulsant activity⁷⁶.

Picrotoxin also antagonizes the central action of GABA raising the seizure threshold, therefore drugs effective in suppressing seizures induced by picrotoxin in rodents are also beneficial in the absence seizures⁷⁸. They abolish the HLTE or prolong the onset of HLTE induced by picrotoxin⁷⁹⁻⁸⁰.

Bicuculline also acts by blocking the action of GABA at GABA_A receptors⁸¹⁻⁸². In experimental animals, the animals that did not elicit the characteristic turning behavior of two consecutive 360 cycles within 30 min observational period are considered protected⁸³⁻⁸⁴.

There is no clear explanation of the mechanism of Aminophylline-induced seizure⁸⁵. However, theophylline used in therapeutics often induces intractable seizures and mortality. The possibility of free radicals and oxidative stress involvement in AMI-induced seizures has also been demonstrated in various Studies⁸⁶⁻⁸⁸. The testing animals are observed for the onset of myoclonic seizures, THLE, and mortality within the observable period of 30 minutes⁸⁹.

The chronic model of epilepsy provides more information on the processes of epileptogenesis occurring from an initial epileptogenic insult to the brain to the occurrence of spontaneous seizures^{14,61,90}. To mimic spontaneous seizures occurring in an epileptic brain, the kindling model has been beneficial both as a tool for understanding chronic epileptogenesis as it relates to epilepsy in humans and as a model for testing AEDs with the potentials for treating complex partial seizures^{15,90}. Seizures beget seizures, through kindling, a normal functioning brain can be epileptic through repeated focal stimulation. It induces progressive intensification of seizures and may be useful in the identification of promising agents necessary for the prevention of seizures, seizure modification as well as a possible correction^{15,91}.

4.3 Experimental evidence in Safety

Toxicological evaluations of medicinal plants are critical to drug development and safety. These toxicities to plants are usually dependent on the plant part and amount consumed, the species and stage of development, as well as the susceptibility of the victim⁹². Noting that the cumulative effects of plants ingested over time are not well understood in traditional medicine, toxicological evaluations such as

acute toxicity, sub-acute toxicity, sub-chronic toxicity, and chronic toxicity studies are commonly employed in predicting the safety of medicinal plants.

Knowledge of the traditional method of preparation and use of such plants is necessary to anticipate the ingestion of such toxic plants and their parts or other finished herbal products. Although the safety of herbal products use is still a major concern among healthcare practitioners in promoting their integration into healthcare systems⁹³, the trend of traditional medicines (TM) use is increasing even among healthcare personnel as a reflection of its increasing acceptance in the health system⁹⁴.

This review identified that the median lethal dose (LD50) of these plants were reported, further toxicological studies should be carried out on these plants. A comprehensive analysis of plant's extracts and phytochemical constituents presents, identification, isolation and characterization of the individual bioactive constituents in the medicinal plants, as well as determination of their exact mechanisms of action, will ensure a critical assessment of their therapeutic potential. This will also be beneficial in the development of potential inexpensive remedies to reduce the epilepsy treatment gap in developing countries and also could provide new treatments for drug-resistant seizures.

5. Conclusions

This review provided evidence for the use of medicinal plants in the management of epilepsy and possibly rationalized the use of plant extracts as alternatives in treating seizures and epilepsy. This review also identifies the screening models carried out on the medicinal plants and thus identifies those that have not been fully studied and their mechanisms of action not ascertained. It also identified those whose active constituents are responsible for their activity remain unknown. The review also identifies potential target plants for future studies of new and alternative therapies for managing epilepsy and other neurological and neurodegenerative diseases.

List of abbreviations

4-AP: 4-aminopyridine; AEDs: Antiepileptic drugs; AMI: Aminophylline-induced; BRU: brucine; ELT KIND: Electrically induced kindling; GABA: Gamma amino butyric acid; HLTE: Hind limb tonic extension; INH: Isoniazid; LD50: median lethal dose; MEST: Maximal electroshock seizure test; NMDA: N-Methyl-D-Aspartate; PIC: Picrotoxin; PIL: Pilocarpine; PTZ KIND: PTZ induced kindling; PTZ: Pentylentetrazol; STR: Strychnine; STR: Strychnine; WHO: World health

organization;

Declarations

Ethics Approval and consent to participate

Not applicable

Consent for Publication

Not applicable

Availability of data and materials

Not applicable

Competing interests

The authors declare no conflict of interest.

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Authors' contributions

ASW conceptualized the original idea. *ASW* and *ARA* developed the study methods and co-wrote the manuscript. *ASD*, *MM* and *MHA* performed the literature review and co-wrote the manuscript. *ASW* and *SM* edited the final draft and critically reviewed the manuscript for intellectual content. *AHY* reviewed and approved the final version of the manuscript.

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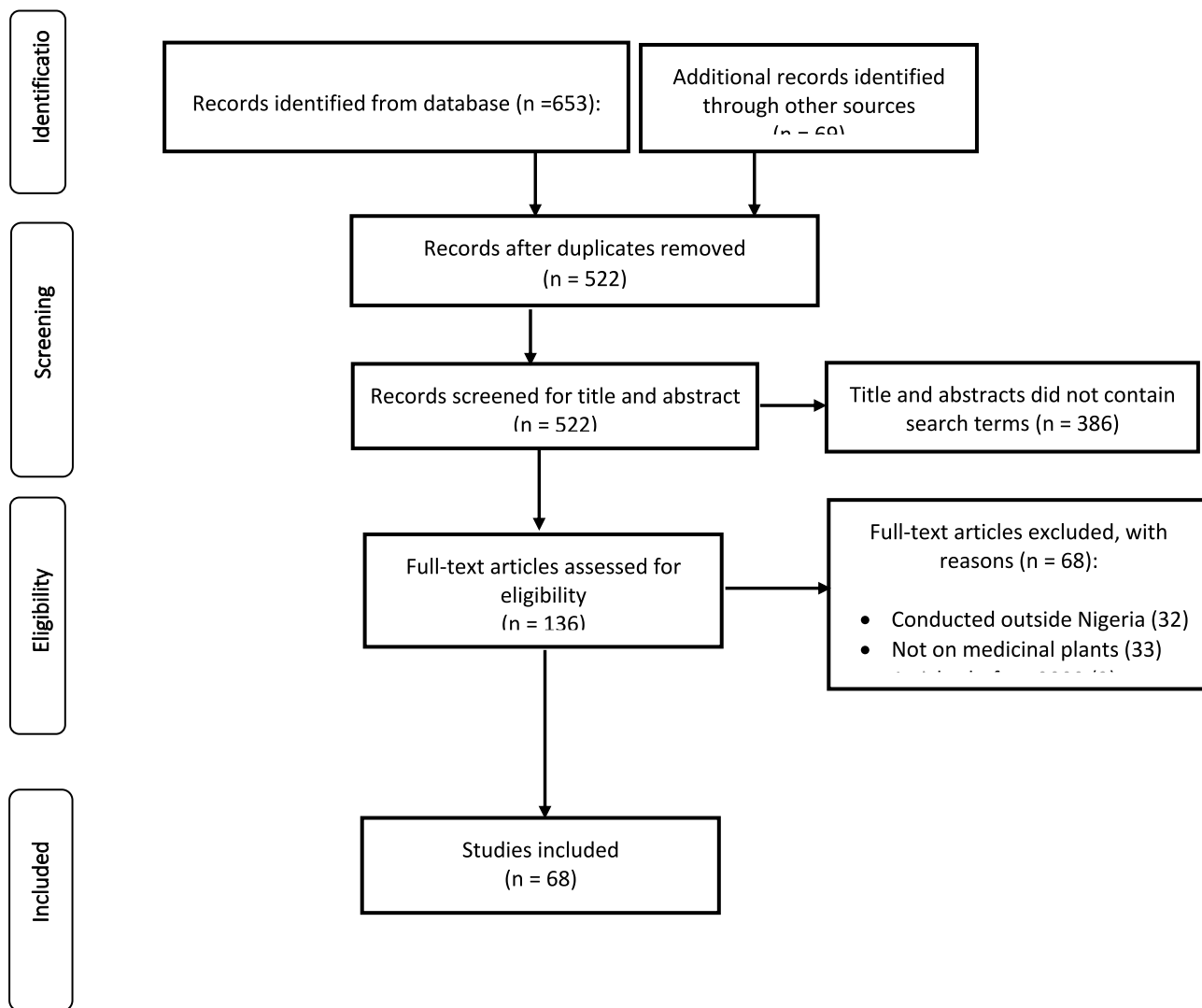


Figure 1. PRISMA Flowchart for the study selection process

Table 1. Medicinal plants with reported anticonvulsant activity in Nigeria

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|----------------|---|---|---|--------------------------------------|------------|
| 1. | <i>Afromosia laxiflora</i> Benth. Ex Baker Harms | Leguminosae | Satin wood (E), Shedun, Makarho, Abua ocha | Headache, body pains, edema, dysentery. | Root bark (Aqueous) | MEST, PIC | 34 |
| 2. | <i>Albizia chevalieri</i> Harms | Mimosaceae | Silk tree (E) | Diabetes mellitus, asthma, hemorrhoids, gonorrhoea, leprosy, diarrhoea, bronchitis | Leaf (Methanol) | MEST, PTZ, PIC, 4-AP, PTZ KIND | 37 |
| 3. | <i>Albizia glaberrima</i> Schum & Thonn | Leguminosae | White Nongo (E), Ayunre (Y) | Anemia, liver complaints, bilharzia, and chest pain | Leaf (Aqueous) | PTZ, PIC, STR | 44 |
| 4. | <i>Allium cepa</i> L. | Amaryllidaceae | Onion (E), Albasa (H), Alubosa (Y) | Cough, bronchitis, asthma, dysentery, ulcer wounds, pains | Bulb (Aqueous) | MEST, PTZ | 36 |
| 5. | <i>Annona senegalensis</i> Pers. | Annonaceae | Wild Custard Apple (E), Gwandar daji (H), Abo (Y), Uburu ocha (I) | Sleeping sickness, cancer, chest pain, coughs, anemia, urinary tract infections, stomach ache, diarrhoea, dysentery, arthritis, rheumatism, intestinal and guinea worms, venereal diseases, head and body ache, trypanosomiasis, lice infestation, leishmaniasis, eyelid swelling, and snakebites | Leaf (Methanol); Root bark (Aqueous) Root bark (Kaurenoic acid) | PTZ; MEST, PTZ; MEST, PTZ, PIC | 56, 95-96 |
| 6. | <i>Aspilia Africana</i> Pers. C.D Adams | Asteraceae | Wild sunflower, iodine or hemorrhage plant (E) | Osteoporosis, tuberculosis, cough, measles, diabetes, malaria | Leaf (Ethanol) | MEST, PTZ, STR | 97 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|---|--------------|---|--|--|---|------------|
| 7. | <i>Boswellia dalzielii</i> Hutch | Bursaceae | Frankincense tree (E), Hano (H), Andakehi (F) | Fever, arthritis, rheumatism, gastro-intestinal problems. Emesis, mental derangement | Stem bark (Methanol) | MEST, PTZ, PIC, STR, 4-AP | 98 |
| 8. | <i>Bryophyllum Pinnatum</i> Lam. Oken | Crassulaceae | Never die, Air plant, Life plant (E), Abamoda, Eru-odundun (Y) | Tranquilizer, fever, inflammation, refractory cough, intestinal pains. | Leaf (Aqueous) | PIC, STR | 99 |
| 9. | <i>Carissa edulis</i> Vahl | Apocynaceae | Cizaki, Bagazaki, Lemun tsuntsuu, Uwaa banzaa (H) Arabic num-num (E), Kanboro (F) | Toothache, hernia, fever, sickle cell anemia, edema, cough, ulcer, cancer and worm infestation | Root bark (Aqueous) | MEST, PTZ, PIC, STR, NMDA, INH, AMI, ELT KIND | 38 |
| 10. | <i>Celtis integrifolia</i> Lam | Ulmaceae | African Hackberry (E), Zuwo (H), Aspe (Y), Ngezo (K), Gamki (F) | Leprosy, microbial infections and measles | Leaf (Methanol) | MEST, PTC, 4-AP, STR | 49 |
| 11. | <i>Chrysanthellum indicum</i> Linn | Compositae | Rariyar kasa, dunkufe, Goshin bu'ana (H) Oyigi, Abilere (Y) | Boils, fevers, jaundice, gonorrhea, hepatitis, heart problems. | Whole plant (Methanol) | MEST, PTZ, STR | 26 |
| 12. | <i>Cissus cornifolia</i> Planch | Vitaceae | Rigarbiri or Duwawun biri (H) | Gonorrhea, septic tonsil and pharyngitis and malaria (Burkill, 1985) | Root bark; Leaf (Methanol) | MEST, PTZ, STR, 4AP, PIC | 39, 100 |
| 13. | <i>Clausea anisata</i> (Wild.) Hook.f.ex Benth) | Rutaceae | Agbasa, Atapari obuka (Y). | Antidiabetic, antihypertensive, anti-inflammatory, gastrointestinal disorders, mental disorders, Toothache | Leaves; root bark; stem bark (Ethanol) | PTZ | 101 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|----------------------------|---|--|--|------------------------------|------------|
| 14. | <i>Cochlospermum tinctorium</i> A. Rich | Cochlospermac eae | Rawaya, kyamba (H), Obazi (I), Sewutu (Y). | Jaundice, yellow fever, heart irregularities, diarrhea, dysentery and colic. | Root (Methanol) | MEST, PTZ, STR | 31 |
| 15. | <i>Commiphora Kerstingi</i> Engl. | Burseraceae | Ararabi, Dashi, Dali, Kwaor (H) | Venomous stings, antibacterial, laxative | Leaf (Methanol) | MEST, PTZ, STR, 4-AP | 102 |
| 16. | <i>Crinum jagus</i> L. | lilaceae Amaryllidaceae | Edesuku (Y), Gaadal (F), Ede chukwu, Olodi (I) | Boils, open wounds, chronic cough | Bulb (Aqueous); (Methanol) | MEST, PTZ, STR, PIC; MEST | 103-104 |
| 17. | <i>Cyperus esculentus</i> L. | Cyperaceae | Tiger nut (E), aya (H), ofo (Y), akihausa (I) | Lactogenic | Seeds (Methanol) | PTZ | 105 |
| 18. | <i>Denmettia Tripetala</i> Bak.f. | Annonaceae | Pepper fruit (E) | Source of vitamins | Fruit (ethanol) | ISO | 106 |
| 19. | <i>Detarium senegalense</i> J.F. Gmelin | Fabaceae | Tallow tree (E), Ofo (I) | fever, anemia, diarrhea, cough, ulcer, worm infestation, cancer | Leaves (Ethanol) | PTZ, BRU, INH | 107 |
| 20. | <i>Diospyros mespiliformis</i> Hochst. | Ebanaceae | West African ebony, Monkey guava (E), Kanya, Kaiwa (H), Igi dudu (I) | Malaria, pneumonia, syphilis, leprosy and dermatomycoses, anthelmintic, mild laxative, fever, dysentery, hemostatic agent to wounds, gingivitis and toothache | Leaf (Methanol) | MEST, PTZ, STR, PIC, 4AP | 40 |
| 21. | <i>Emilia sonchifolia</i> (L.), DC. | Compositae Asteraceae | Cupid shaving brush, Sow thistle (E), | Fever, measles, sore throat, rashes, inflammatory diseases, eye and ear ailments, vertigo, regenerating bath therapy anti- inflammatory antidiabetic cytotoxic, antitumor, antioxidant, and properties | Leaf (Ethanol, chloroform and aqueous) | MEST, STR | 108 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|---|----------------|---|---|--|----------------------------------|------------|
| 22. | <i>Enantia chlorantha</i> Oliv | Annonaceae | African white wood (E), Osopupa, Ikale (Y) | Febrile illnesses, stomach upset, toothache, wounds infections, jaundice, high blood pressure, dysentery, painful and swollen joints. | Stem bark (ethanol) | PTZ, PIC | 109 |
| 23. | <i>Evolvulus alsinoides</i> (Linn.) Linn | Convolvulaceae | kaafi- mallam or matakin kurciya, (I) Efunje, Efunle (Y) | dysentery and depression, asthma mental disturbances fever, loss of memory, syphilis and to promote hair growth | Whole plant (Methanol) | MEST, PTZ | 28 |
| 24. | <i>Ficus platyphylla</i> Del. Holl. | Moraceae | Flake rubber, Red Kano rubber (E), Danko gawi, Dnko, gamjii (H) | Depression, psychoses, pain and inflammation | Stem bark (Methanol) | MEST, PTZ, STR, PIC, INH | 16 |
| 25. | <i>Ficus sycomorua</i> Linn | Moraceae | Farin Baure (H), Tarmu (K), | Diarrhea, dysentery, painful urination and vaginal infections, mental illness, diarrhea, pain relief | Root bark (Methanol); Stem bark (Aqueous) | MEST, PTZ, 4-AP; PTZ, STR | 110-111 |
| 26. | <i>Globimetula braunii</i> (Eng.) Van Tiegh | Loranthaceae | Kauchii (H), Afomoonishano (Y). | Cardiovascular diseases, hepatic illness, malaria. rheumatism, infertility and stomach problems | Leaf (Ethyl acetate) | MEST, PTZ, 4-AP | 41 |
| 27. | <i>Hymenocardia acida</i> Tul | Euphorbiaceae | Heart-fruit (E), jan yaro (H) yawa satoje (F), ikalaga (I), Orumpa (Y). | Abdominal and menstrual pains, tumors, arthritis, chest complaints, small pox, headaches, rheumatic pains, | Leaf, Stem bark (Ethanol) | MEST, PTZ, STR, 4AP | 42-43 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--------------------------------------|----------------|---|--|-----------------------------|----------------------|------------|
| | | | | toothaches, hypotension, sickle cell, schizoprenia, diabetes | | | |
| 28. | <i>Ipomea involucriata</i> P. Beauv. | Convolvulaceae | Duman-kwaadu (H), Fifi lori (I), Apiti, ododo (Y) | Asthma, fever, headache, jaundice, gonorrhoea, dysmenorrhoea | Root (Ethanol) | 4-AP | 112 |
| 29. | <i>Jatropha curcas</i> Linn | Euphorbiaceae | Purging nuts, Barbados nut, Physics nut (E). Butuje (Y) | Cancer, skin rashes, oral candidiasis, scabies, eczema, dermatitis, helmiths, snake bite, rheumatism, dropsy. | leaf (Ethanol & Chloroform) | PTZ, STR | 113 |
| 30. | <i>Jatropha gossypifolia</i> Linn | Euphorbiaceae | Binidazugu (H), Lapalapa (Y), Wluluidi (I) | Anti-inflammatory, anti-diarrheal, analgesic, antipyretic, antidiabetic, antimicrobial & antihemorrhagic activities. | Leaf (Methanol) | MEST, PTZ, STR, 4-AP | 114 |
| 31. | <i>Laggera aurita</i> Linn | Asteraceae | Abanaadene (I), Eru-tabata (Y), taba taba (H) | pediatric malaria, inflammation, fever, rheumatic pain, stomatitis, asthma, bronchitis, dyspepsia, nasal congestion, antibacterial, constipation, dysentery and cancer | Leaf (Methanol) | MEST, PTZ, STR | 30 |
| 32. | <i>Lannea barteri</i> (Oliv.) Engl. | Anacardiaceae | Babban baraa, Faaru, Faarun birri (H), Aka (Y) | Gastritis, childhood convulsions and inflammation | Stem (Ethanol) | MEST PTZ, STR, PIC | 46 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|-------------|--|---|---------------------|----------------------|------------|
| 33. | <i>Leucas martinicensis</i> Jacq. R. Br. | Lamiaceae | Whitewort or mosquito plant (E), Bunsurun fadama (H) | Cough, kidney disorders, rheumatism, diarrhoea, fevers, skin rashes. | Leaves (Aqueous) | MEST, STR | 50 |
| 34. | <i>Lophira lanceolata</i> Tiegh. ex Key | Ochnaceae | Namijin Kadanya (H) | Erectile dysfunction | Stem (Ethanol) | MEST, PTZ, PIC | 115 |
| 35. | <i>Milicia excelsa</i> (Welw.) C.C. Berg | Moraceae | African teak, Iroko (E) | Malaria, anaemia, lactation failure, mental illnesses, sexual dysfunction, rheumatism | Leaf (Ethanol) | PTZ, PIC, STR | 58 |
| 36. | <i>Milletia aboensis</i> Hook. F. | Leguminosae | | Ring worm | Leaf (Ethanol) | 4-AP | 112 |
| 37. | <i>Mitragyna africana</i> Willd | Rubiaceae | Uburu (I), abura (Y), guljeya or gyayya (H), Kawui (K) | Bacterial infections especially gonorrhea, dysentery, mental disorder, treatment of sterility and African sleeping sickness | Stem (Methanol) | STR | 52 |
| 38. | <i>Mitragyna inermis</i> (Willd.) O Kuntze | Rubiaceae | Giyayya (H), | antimicrobial, antidiabetic, antimalarial, antihypertensive | Stem bark (Ethanol) | PTZ, STR | 116 |
| 39. | <i>Mitragyna stipulosa</i> (DC.) Kuntze | Rubiaceae | African linden (E), Opepe (Y). | fever, hypertension, dysentery, gonorrhea, leprosy, ulcers, amenorrhea, colds, cough, chest pain, and stomach ache | Leaf (Ethanol) | PTZ, PIC, STR | 117 |
| 40. | <i>Mondia whitei</i> (Hook f.) Skeel | Apocynaceae | White ginger (E) | Urinary infections, impotence and sexual dysfunction, constipation, gonorrhoea, abdominal pain, inducement of | leaf (Methanol) | PIL | 118 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|-------------|--|--|----------------------------|----------------------------------|------------|
| 41. | <i>Moringa oleifera</i> Lam | Moringaceae | Horseradish or tree of life (E) | labour. stress, paralysis, antimalarial and anthelmintics Gastric ulcers, skin diseases, lowering blood sugar, diabetes, cancer, fatigue, hay fever, impotence, edema, headaches, sore gums | Leaf (Ethanol) | PTZ, PIC, STR | 45 |
| 42. | <i>Olax subscorpioidea</i> Oliv. | Olacaceae | Ifon (Y) | Pains, mental illness, asthma, rheumatism and articular pains, Cancer, typhoid fever, microbial diseases, yellow fever, jaundice, venereal diseases and guinea worm infestation. | Leaf (Ethanol); (Methanol) | PTZ, PIC, STR; MEST, PTZ, STR | 27, 119 |
| 43. | <i>Paullinia pinnata</i> Linn | Sapindaceae | Goron dorina, Yatsu biyar (H) Edefina, Aliligo (I) Ogbe-okiyè (Y). | Yellow fever, Jaundice, and heart irregularities, diarrhea, dysentery and colic | Root bark (Methanol) | MEST, PTZ, STR | 31 |
| 44. | <i>Pennisetum glaucum</i> (L.) R.Br. | Poaceae | Pearl millet (E), gero (H), oka (Y) | | Seeds (Aqueous) | MEST, PTZ, PTZ KIND | 55 |
| 45. | <i>Piper guineense</i> Schumacher & Thonn. | Piperaceae | West African Black Pepper (E) Uziza (I) and Iyere (Y) | Cough, stomach disorder, rheumatism and bronchitis, intestinal diseases, gonorrhoea, aphrodisiac, mental illness | Fruit (hydrodistillation) | PTZ | 47 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|------------------|--|---|--|---------------------------|------------|
| 46. | <i>Plectranthus aegyptiacus</i> (Forssk.) C.Chr. | Lamiaceae | Efinrin-Oyinbo (Y) | Pain, cough, fever, sore throat, ear ache, sensory diseases, respiratory system infections, & abdominal disorders | Leaf (hydro-distillation) | MEST, PTZ, STR | 120 |
| 47. | <i>Pyrenacantha staudtii</i> Engl. | Icacinaceae | | Hypertension, ulcer, inflammation, intestinal pain, blenorhoea, hernia, and insomnia | Leaf (ethanol) | DMSO | 121 |
| 48. | <i>Randia nilotica</i> Stapf. | Rubiaceae | Tsibra, barbaji (H), Gialgoti (F) | | Stem (Ethanol) | MEST, PTZ, STR | 32 |
| 49. | <i>Rauvolfia vomitoria</i> (Afzel) | Apocynaceae | Asofeyeje (Y), Akanta (I) | Blood pressure, antimalarial, antipyretic, analgesic, haematonic | Leaf (Aqueous) | PTZ, STR, PIC | 122 |
| 50. | <i>Russelia equisetiformis</i> Scgdl. & Cham. | Scrophulariaceae | Firecracker, coral and fountain plant (E) | Diabetes and leukemia | Whole plant (Methanol) | PIC, STR | 53, 123 |
| 51. | <i>Rauvolfia vomitoria</i> Afzel | Apocynaceae | Poison devils pepper (E) | Cancer, diabetes, fever, high blood pressure | Root bark (Ethanol) | 4-AP | 112 |
| 52. | <i>Sansevieria liberica</i> Jerome & Labroy | Agavaceae | African Bowstring or Leopard Lily (E) | Headache, fever, cold, analgesic, anti-inflammatory and antibiotic | Root bark (Aqueous) | STR, PIC, BIC, PTZ | 51 |
| 53. | <i>Securidaca longepedunculata</i> Fresen | Polygalaceae | Violet tree (E), <i>Sanya</i> or <i>Uwarmagunguna</i> (H), <i>Ezeogwu</i> (I) and <i>Alali</i> (F) | Pain and inflammation, | Root bark (Aqueous); Stem bark (Aqueous) | STR, PIC; MEST, PTZ, 4-AP | 21, 35 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|---------------|--|---|-----------------------|---------------------------|------------|
| 54. | <i>Securinega virosa</i> Roxb (ex Willd) Baill. | Euphorbiaceae | Tsuwaawun karee (H), Iranje (Y), Njisi nta (I). | Dysentery, ulcer, menstrual problems, gerna, pain, infertility. | Root (methanol) | MEST, PTZ, PIC, STR, 4AP. | 48 |
| 55. | <i>Solanum nigrum</i> Linn | Solanaceae | Black shade (E) | Inflammation, liver problems, antiproliferative | Leaf (Aqueous) | MEST, PTZ, PIC | 33 |
| 56. | <i>Spathodea campanulata</i> P. Beauv | Bignoniaceae | African Tulip (E), Imiewu (I), Oruru (Y) | Analgesic, anti-inflammatory and anti-plasmodial, fungal infections, impetigo, herpes, scabies, other skin infections | Leaf (Ethanol) | MEST, PTZ | 57 |
| 57. | <i>Stereospermum Kunthianum</i> Cham. | Bignoniaceae | Pink jacaranda (E) | Indigestion, hiccups, diarrhea, vomiting, fever, asthma, diabetes | Stem (Aqueous) | MEST, PTZ | 124 |
| 58. | <i>Synsepalum dulcificum</i> Schumach. & Thonn.) Daniell | Sapotaceae | Miracle fruit (E) | Taste modifying | Seed (Aqueous) | MEST, PTZ, STR | 125 |
| 59. | <i>Tapinanthus globiferus</i> (A Rich.) van Tiegh | Loranthaceae | mistletoe (E), Kauchin (H), afomo (Y), Osisi or Okwuma osa (I) | Rheumatism, syphilis, fever, hypertension, cancer, diabetes, diuretic agent, body pain, ulcers. | Whole plant (Aqueous) | MEST, PTZ | 126 |
| 60. | <i>Telfairia occidentalis</i> Hook.fil. | Cucurbitaceae | Fluted gourd (E), | Fish and human poisons, blood tonic, anemia, malaria gastrointestinal disorders, | Leaf (Aqueous) | MEST, STR, PTZ | 127 |
| 61. | <i>Trichilia roka</i> Forskal | Meliaceae | Cape or Natal mahogany (E), Goron Talaka (H), | Antihelminthic, aphrodisiac, antiplasmodial, pains, inflammation | Stem bark (Ethanol) | MEST, PTZ, STR, PIC | 128 |

| S/N | Plants Name | (Family) | Local Names | Other Ethnobotanical Uses | Part used (Solvent) | Methods of Screening | References |
|-----|--|-----------|-------------|--|------------------------|----------------------|------------|
| 62. | <i>Xeromphis nilotica</i> Stapf. Keay | Rubiaceae | Barbaji (E) | Fever, asthma, Stomach pain, dropsy, abdominal pain and to induce labour | Root (aqueous) bark | PTZ | 129 |

KEY: E-English, H- Hausa, F- Fulfulde, Y- Yoruba, I- Igbo, K- Kanuri, MEST- Maximal electroshock test, PTZ- Pentylenetetrazol, PIC- Picrotoxin, PIL- Pilocarpine, STR- Strychnine, NMDA- N-Methyl-D-aspartate (NMDA), INH- Isoniazid, AMI- Aminophylline, BRU- brucine; ELT KIND- Electrically induced kindling, PTZ KIND- PTZ induced kindling