Review Article



Biological Potential of Elephantopus scaber Linn.

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ABSTRACT

Elephantopus scaber Linn., belonging to the family Asteraceae, is a small herb found in Asia, Africa, Australia and Europe. The most common name in english for *Elephantopus scaber* is Elephant's Foot, but often it is called more precisely Prickly-leaves Elephant's Foot or Rough-leaved Elephant's Foot. The parts of this plant have been used traditionally for the treatment of number of diseases in many countries. The plant has been extensively screened and proved for anticancer activity, which is mainly for its deoxyelephantopin containing. Many other biological activities such as antimicrobial, hepatoprotective, antioxidant, antidiabetic, anti-inflammatory, analgesic, antiasthamatic, antiplatelet and wound healing ability have been reported in various research articles.

Keywords: Elephantopus scaber, phytochemicals, traditional medicine, pharmacological actions.

INTRODUCTION

n recent decades, research has shown that plants produce a diverse range of bioactive molecules for industrial interest, making them a rich source of different types of medicines and have shown a promising effect in therapeutics¹. The lectotype species of *Elephantopus* genus, family Asteraceae, consists of 32 species of centered in Asia, Africa, Australia and Europe²⁻⁵. The plant vigorously grows in tropical deciduous forest areas. Traditionally the plant is used as folk medicine by tribal communities across India. *Elephantopus scaber* Linn. is one of the medicinally important species of this genus. Hiradeve and Rangari⁶ have recently reviewed the ethnomedical history of *E. scaber*. The whole plant, its various parts and the extracts of *E. scaber* have been used for the treatment of a number of diseases.



Elephantopus scaber

PHYTOCHEMICALS

Flavonoids have been shown to be good taxonomic markers for Asteraceae. More than 800 compounds comprising 4700 flavonoid were occurred with some implications of flavonols, flavones and other types of the family Asteraceae ⁷ apart from that *Elephantopus scaber* has been explored for a large amount of salt such as

potassium chloride and minerals especially calcium, magnesium, iron and zinc. The organic solvent like acetone extract of air-dried powdered seeds is reported to contain terpenoids, flavonoids, steroids, glycosides, alkaloid, quinones, phenols⁸. A number of phytochemical studies have demonstrated the presence of several classes of chemical compounds, the main ones being Phenolic acids, flavonoids, terpenoids, coumarines, quinones, essential oils.

Traditional Uses

E. scaber has been used as traditional medicine in many countries. In Thailand, it has been used as traditional medicine. In Brazil, the decoction of whole plant is used to stimulate diuresis, reduce fever and eliminate bladder stones, febrifuge, and diaphoretic against cough, bronchitis, and asthma ⁹. In Malaysia, decoction of *E*. scaber root has been used to accelerate contraction of abdominal area and prevent inflammation after childbirth. Besides, whole E. scaber was also boiled with red bean to remove flatulence ¹⁰. In Thailand people have used E. scaber to treat cough, as a tonic (root decoction) ¹¹, chapped lips and galactogogue (whole plant decoction) ¹⁰. It has also been used in Madagascar as an antipyretic agent (decoction of aerial part)¹²; in Taiwan (whole plant decoction) to treat hepatitis; in Nigeria (hot water extract of leaves) to cure arthritis and in Mauritius to treat diarrhoea, urinary problems and pimples (root paste)¹³. In India whole plant of E. scaber are used for the treatment of toothache as a toothbrush ¹⁴. Also, the whole plant has been reported to be useful for curing insomnia, diabetes, bronchitis, viral or bacterial infection, leukaemia, rheumatism, snake bite, diuresis, antipyresis, to eliminate bladder stones and for filariasis. The leaf crushed and mixed with salt is used to treat dysentery, while the water extract of the leaves is applied externally



130

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to treat eczema and ulcers ^{15, 16}. In Southern China, Hong Kong and Taiwan, the whole plant of *E. scaber*, is widely used in the treatment of nephritis, edema, dampness, pain in the chest, fever, scabies, arthralgia due to wound and cough of Pneumonia ¹⁷. The Murut people of Sabah Malaysia used the roots of the Elephantopus specie to treat bloody stool ¹⁸.

Pharmacological Effects

Anticancer & antitumor activity

Cancer is the 2nd leading cause of death worldwide ¹⁹. Deoxyelephantopin (DET) is one of the major sesquiterpene lactone derived from E. scaber. In vitro and have vivo experiments demonstrated that in deoxyelephantopin possesses cytotoxic activity against a variety of cancer cell lines and malignant tumors. Deoxyelephantopin has shown significant cytotoxicity against human breast cancer cell lines Deoxyelephantopin in nasopharyngeal carcinoma (CNE) cells triggred Akt and Mitogen-activated protein kinase (MAPK) signalling pathways²¹ and showed pronounced activity against melanoma derived cell line MEXF 394NL and mammary cancer cell line MEXF 401NL with IC70 value of 1.1 μ g mL⁻¹. ²². Very recently one literature has shown that DET could kill cancer cells selectively inducing apoptosis through multiple signaling pathways ¹⁹. The antitumor activity of the leaves of E. scaber has been evaluated against Dalton's ascitic lymphoma (DAL) in Swiss albino mice ²³. Deoxyelephantopin from elephantopus scaper inhibits HCT116 human colorectal carcinoma cell growth through apoptosis ²⁴. Ethanol extract of E. scaber may be a potential anti-cancer agent for human breast cancer cells by the induction of p53dependent apoptosis ²⁵. Study of E scaber on skin papillomas induced by DMBA and croton oil in mice showed tumor inhibitory activity of the active fraction against chemically induced tumors and an ability to inhibit the development of solid tumors ²⁶.

Antimicrobial activity

There is a vital need to develop new classes of antibiotics as the currently used antibiotics developing a dangerous microbial resistance ²⁷. Different study showed that Elephantopus scaber is a herb with good antimicrobial activity. The methanolic leaf extract of E. scaber showed significant antibacterial activity against S aureus, E coli, P aeruginosa, B subtilis and P vulgaris²⁸. Ethyl acetate extract of the plant showed growth inhibitory effect at 4 mg/ml concentration in all the bacterial isolates tested except Klebsiella pneumonia where it showed ~75% inhibition. Lower concentration of the extract showed concentration-dependent inhibition effect. At 2 mg/ml 50% inhibition in all the cultures ²⁹. Further research into the antimicrobial activity was done by the use of different strains of pathogenic bacteria and fungi. The result was analysed using well diffusion method ³⁰. The acetone fraction of E. scabber demonstrated remarkable antibacterial effect against methicillin resistant

Staphylococcus aureus and methicillin sensitive Staphylococcus. aureus ³¹. A novel terpenoid from Elephantopus scaber showed antibacterial activity against Staphylococcus aureus and suggests that it can act as a drug for bacterial infections ³². Study showed strong in vitro antibacterial activity of terpenoid derivatives against ES beta-lactamase-producing methicillin resistant Staphylococcus aureus ³³.

Anti-diabetic activity

The effect of the crude extract and acetone fractions from Elephantopus scaber (Asteraceae) on glycemia was investigated in streptozotocin (STZ) diabetic rats. Oral administration of crude extract significantly reduced serum glucose levels and increased the lowered insulin concentrations in diabetic animals at 150 mg kg⁻¹. Nevertheless, the hypoglycemic effect of the acetone fractions (mg kg⁻¹) of *Elephantopus scaber* in diabetic rats was evident at 2 h after treatment, where the sugar lowering effect of the most effective fraction was more than 69%. Phytochemical analysis revealed the compound to be a terpenoid ³⁴. Effect of E. scaber leaf extract on STZ-induced diabetic rats showed the methanol extract exhibited better hyperglycemic action than the hexane and ethyl acetate extracts. An acetone extract exhibited significant anti-diabetic activity by reducing blood glucose and restoring the insulin levels in STZ-induced diabetic rats ³⁵. Oral administration of aqueous extract of leaves and roots into alloxan induced diabetic rats significantly reduced serum glucose, glycosylated hemoglobin and the activity of gluconeogenic enzyme glucose-6-phosphatase, but increased serum insulin, liver and skeletal muscle glycogen content and the activity of glycolytic enzssyme glucokinase ³⁶. Report reveals that Terpenoid and 2,6,23 trienolide isolated from whole plant act as anti diabetic agent³⁷.

Anti-inflammatory activity

Search for the novel natural anti-inflammatory substances are still necessary due to intolerable side effects of the synthetic anti-inflammatory drugs 38-40. A compound isolated from the hydroalcoholic extract of aerial part of E. scaber was studied for the *in vivo* anti-inflammatory activity in albino rats and showed that higher dose of the compound is highly effective in inhibiting carragenan induced edema ⁴¹. A study to investigate protective mechanism of E. scaber using lipopolysaccharide (LPS) induced inflammation of BV-2 microglial cells and acute liver injury in Sprague-Dawley rats. E. scaber reduced LPS induced nitric oxide (NO), interleukin (IL)-1, IL-6, reactive oxygen species and prostaglandin (PG) production in BV-2 cells. It significantly decreased serum aspartate aminotransferase (AST) and alanine aminotransferase (ALT) levels in LPS-treated rats ⁴². Ethyl acetate fraction from the leaves of E. scaber exhibited antineuroinflammatory effect in lipopolysaccharide (LPS)induced microglia cells (BV-2) by blocking nuclear factor κ B (NF- κ B) via the significant reduction of NO, inducible nitric oxide synthase (iNOS), cyclooxygenase(COX)-2,



Available online at www.globalresearchonline.net © Copyright protected. Unauthorised republication, reproduction, distribution, dissemination and copying of this document in whole or in part is strictly prohibited. Prostaglandin- E_2 (PGE₂), IL-1b, and Tumor necrosis factor(TNF)- α production ⁴³.

Hepatoprotective activity

E. scaber was observed for its hepatoprotective effect in mice; the plant extract was able to reverse the liver damage induced by ethanol administration. This action buttressed the traditional use of the plant as a liver tonic⁴⁴. The water extract of E. *scaber* was observed to have a hepatoprotective effect on SD rats (LPS-induced liver damage). The mechanism of *E. scaber* protection involves an antioxidant effect and inhibition of p38 MAP kinase and COX-2 expressions in LPS-stressed acute hepatic injury in SD rats ⁴². The hepatoprotective activity of *E. scaber* was evaluated using the methanolic extract. Administration of the extract to rats in which liver damage was previously induced by carbon tetrachloride was found to reverse the damage ⁴⁵.

Anti-parasitic activity

Twenty three extracts derived from eleven plant species including *Elephantopus scaber* (leaf) showed promising antiplasmodial activities against the blood stage chloroquine resistant P. falciparum (EC50 < 10 μ g/ml) with negligible toxicity effect to MDBK cells *in-vitro* [46]. *In vitro* antitrypanosomal (parasitic) activity of *Elephantopus scaber* was also reported ⁴⁷.

Anti-HIV activity

Potent HIV-1 RT inhibitory actions were obtained from the water extract of *Elephantopus scaber* L. leaves and root. IC50 of leaves and root were 69.9 and 107.57 μ g/ml, respectively [⁴⁸. The hot aqueous extract of *Elephantopus scaber* L. Leaves shown 48% inhibition against HIV-1 protease at the dose of 250 microgram/ml⁴⁹.

Anti-asthmatic activity

Ethanol extract of *Elephantopus scaber* leaves was evaluated for preliminary phytochemical screening and antiasthmatic activity using histamine and acetylcholine-induced bronchospasm, mast cell degranulation and histamine induced constriction on isolated guinea pig tracheal chain at different dose levels ⁵⁰.

Memory power

Ethanolic extract of *Elephantopus scaber* (Linn.) leaves may exhibit memory enhancing activity in aged mice and this enhancement of memory may be due to increase anticholinesterase and caspase activity in mice brain ⁵¹.

Nephroprotective Activity

The nephroprotective effect of *Elephantopus scaber* could be due to flavonoid content and the inherent antioxidant moieties in the extract. The ethanolic extract of Eelphantopus scaber leaves could constitute a lead to discovery of a novel drug for the treatment of drug-induced Nephrotoxicity⁵².

Wound healing activity

Deoxyelephantopin, may be due to the presence of active moiety, α methylene γ lactone showed more significant effect than ethanolic extract towards wound healing activity by increasing cellular proliferation, formation of granulation tissue, synthesis of collagen and increase in the rate of wound contraction ⁵³. Aqueous extract of E. scaber is taken orally to heal wounds ⁵⁴.

Anti-pest activity

Elephantopus scaber leaf extract was found to be toxic against pests like Red Flour Beetle (*Tribolium castaneum*), Cotton Stainer (*Dysdercus cingulatus*) and Maize Weevil (*Sitophilus zeamais*), therefore it can be used as a natural pesticide against those bugs, and the plant is worth further investigation for its anti-pest potential ⁵⁵.

CONCLUSION AND FUTURE PROSPECTIVE

The use of plants as a source of medicine is as old as the existence of man, renewed interest into the use of herbal medicines is not unrelated to the emergence of new diseases, ineffectiveness of existing drugs and adverse effects of synthetic drugs. The present review tried to look at Elephantopus scaber on a broad perspective, with a knowledge that a lot of research has already been done on the plant. This paper tried to expose the different traditional uses of Elephantopus scaber such as its use for hepatitis, insomnia, diabetes, bronchitis, viral or bacterial infection, leukaemia, rheumatism, pain relief, snake bite, diuresis, antipyresis, etc. Different biological activities of the plant have been investigated, such as anticancer, antimicrobial, anti-inflammatory, anti-parasitic, anti-HIV, anti-diabetic, anti-asthmatic, hepatoprotective, nephroprotective, anti-asthmatic wound healing, memory power and anti-pest activities. Further investigations are necessary to validate its many other traditional uses. This plant may be a source to bring the new lead as anticancer and antimicrobial in near future. As evidenced from the barrage of reports regarding the biological activity and chemical composition of Elephantopus scaber the plant could be a potential source of income for the regions where it is found. Also, further synthesis is required to produce drugs from the isolated compound in large scale; this will reduce dependence on the natural source.

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