

**Borneo Journal of Pharmacy** Vol 4 Issue 4 November 2021 Page 248 – 259 http://journal.umpalangkaraya.ac.id/index.php/bjop/article/view/2377 DOI: https://doi.org/10.33084/bjop.v4i4.2377 e-ISSN: 2621-4814

Review

# Pharmacological screening of Eryngium foetidum Linn - A Review

Galamula Hewage Tharuka Kanchana Hemachandra <sup>1</sup>

Sinnadurai Thuvaragan 1\*💿

Vinotha Sanmugarajah 20

<sup>1</sup>Department of Pharmacy, Faculty of Allied Health Sciences, University of Jaffna, Jaffna, Northern Province, Sri Lanka

<sup>2</sup>Unit of Siddha Medicine, University of Jaffna, Jaffna, Northern Province, Sri Lanka

\*email: sthuvaragan@univ.jfn.ac.lk

Keywords: Activities Eryngium foetidum Pharmacology Review

### Abstract

Eryngium foetidum L. (Family Apiaceae) is a biennial herb, and it is used as a culinary herb and spice across the different countries of the world, including Sri Lanka, India, Bangladesh, Malaysia, Singapore, etc. due to its high aroma quality. Also, it is used to treat several ailments, such as respiratory diseases, gastrointestinal ailments, and skin diseases among different indigenous populations for its medicinal properties. Based on ethnomedical evidence, many studies have been conducted to identify the phytoconstituents, underlying mechanisms, and related pharmacological effects of different parts of this plant. This study reviewed the current state of findings related to the Pharmacological activities of *E. foetidum*. Based on this review, this plant is widely used for ethnomedical and culinary purposes. Pharmacological screening of the plant revealed that it had different activities such as anti-inflammatory, antioxidant, antimicrobial, anthelminthic, anticonvulsant, anticancer, antidiabetic, antimalarial, larvicidal, and hepatoprotective activities. This review further promised that potential new chemical entities could be elicited from the phytoconstituents of E. foetidum.

Received: June 28<sup>th</sup>, 2021 Accepted: October 24<sup>th</sup>, 2021 Published: November 30<sup>th</sup>, 2021

© 2021 Galamula Hewage Tharuka Kanchana Hemachandra, Sinnadurai Thuvaragan, Vinotha Sanmugarajah. Published by Institute for Research and Community Services Universitas Muhammadiyah Palangkaraya. This is an Open Access article under the CC-BY-SA License (http://creativecommons.org/licenses/by-sa/4.0/). DOI: https://doi.org/10.33084/bjop.v4i4.2377

# INTRODUCTION

Eryngium is the largest and most complex genus in the Apiaceae family<sup>1</sup>, and it includes more than 250 flowering species worldwide. Under this genus, the most studied species have been *Eryngium foetidum* L., cultivated in Asian countries such as Iran, Turkey, Europe, Tropical Africa, and Pacific islands. *Eryngium foetidum* is native to Tropical America and the West Indies. It is used as a culinary herb and spice globally, including in Sri Lanka, India, Bangladesh, Malaysia, Singapore, and others. *Eryngium foetidum* is also considered a general edible food component in Nigeria<sup>2</sup>. *Eryngium foetidum* is commonly known as *'spiny coriander'*<sup>3</sup>, *'spirit weed'* or *'saw-tooth coriander'* and is also called in different countries as *langer koriander* (German);

ketumbar java (Malay); pak chi farang (Thai); ngo gai (Vietnamese); culantro, racao, recao (Spanish); Bahkhawr' (India) and andu kola (Sri Lanka)<sup>4</sup>. The leaves of this plant are frequently used interchangeably with Coriandrum sativum L. due to their similar pungent aroma character<sup>5</sup>. It treats several ailments, such as respiratory diseases, gastrointestinal ailments, and skin diseases, by different indigenous systems<sup>6</sup>.

A review study stated that the technology for mass production is available for this plant, cultivated for commercial purposes<sup>6</sup>. Simultaneously, it is easily propagated by seeds (germinate in 20–25 days) in spring or suckers during monsoon. The seedlings are ready to transplant in 45–50 days after seed sowing<sup>4</sup>. In Bangladesh, a research study provides a baseline of information concerning this plant's cultivation techniques, market potential, and climate resilience<sup>3</sup>.

Based on ethnomedical evidence, many studies have been explored phytoconstituents, underlying mechanisms, and related pharmacological effects of the different parts of this plant<sup>7</sup>. Still, many compounds actively responsible for different pharmacological effects of this plant are yet to be detected. Even though some therapeutic effects have been elicited using *in vitro* models, further studies will be needed to correlate these activities using *in vivo* models. This study reviews the current state of findings related to the Pharmacological screening of *E. foetidum*.

# PLANT CLASSIFICATION

Classification and local names of *E. foetidum* are presented in **Table I**. This plant also has several synonyms, with many local names from various parts of the world. However, in general, this plant is better known as long coriander.

Table I.	Classification of I	E. foetidum <sup>8</sup>

Classification	Identity
Domain	Eukaryote
Kingdom	Plantae
Phylum	Tracheophyta
Class	Magnoliopsida
Order	Apiales
Family	Apiaceae
Genus	Eryngium
Species	Eryngium foetidum L
Synonyms	Eryngium antihystericum Rottler, Eryngium molleri
	Gand
Common	Long coriander, wild coriander, fitweed, culantro,
name	Mexican coriander
English name	Long coriander/Sea Holly

### **BOTANICAL DESCRIPTION**

*Eryngium foetidum* is a leafy herb and distributed in the tropical zone of the world. It is indigenous to tropical zones such as America and Caribbean islands, from southern Mexico to Panama via Brazil and from Cuba to Trinidad. It is generally grown in tropical areas like Africa, South Asia, Southern Europe, and the Pacific

islands. It is extensively cultivated in Costa Rica and Puerto Rico for its usage and exporting to the US<sup>4</sup>.

A biennial pungently smelling herb and *E. foetidum* (Figure 1) consists of fleshy waxy in nature, oblanceolate shape, 30 cm long and 4 cm broad serrated or dentate margin with dark green leaves; each tooth of the margin has a slight yellow spine spirally arranged around a short thick stem from a basal rosette. Flowers white, sessile and bisexual, white, narrow, oblonged and notched petals; corolla-creamy white with green calyx, sepals are tubular, acute, persistent, longer than petals and forming an umbel inflorescence on a long stalk, which arises from the leaf rosette at the bottom; fruits are globose to ovoid and evenly branched fibrous roots<sup>6</sup>.



Figure 1. Eryngium foetidum plant

### TRADITIONAL-ETHNOMEDICAL USE

*Eryngium foetidum* has been widely used in Southeast Asian countries, Caribbean islands, and Latin America, where its harvested leaves are commonly used in foods<sup>7</sup>. In traditional medicine, the leaves and roots have been used to treat insanity, skin diseases, mucosal diseases, pulmonary ailments, diabetes mellitus, stomach disorders, and conditions related to the nervous system, such as convulsions, paralysis, spasms, and pains<sup>9</sup>. Moreover, it is used as a vermifuge and febrifuge<sup>10</sup>. *Eryngium foetidum* has been used to treat colds and fever, burns, earache<sup>11</sup>, pneumonia, flu, convulsions in children, malaria<sup>12</sup>, hypertension, constipation, worm infections, and infertility complications<sup>13</sup>. The roots have stomachic, sudorific, and diuretic effects<sup>13,14</sup>, where leaf juice/decoction stimulates the gut as a laxative<sup>7</sup>. It was also reported to treat several poisons, including the treatment of snake bites and scorpion-sting venom<sup>4,13</sup>. In West Africa, aerial parts of this plant were used to treat

respiratory diseases (asthma, cough, cold, and sinusitis), gastrointestinal ailments such as diarrhea, and crushed leaves in heated palm oil to treat rheumatism<sup>15</sup>. Tea or infusions prepared from the leaves of this plant were used as an inhalation to alleviate respiratory disorders<sup>12</sup>. In traditional Chinese medicine, *E. foetidum* is widely used for the treatment of inflammation<sup>2</sup>.

### PHYTOCHEMICAL ANALYSIS

Phytochemical screening of this plant found several secondary metabolites such as flavonoids, phenols, tannins, saponins, ascorbic acid, and terpenoids<sup>16,17</sup>. Leaves are an excellent source of vitamin A<sup>18</sup>. Mabeku *et al.*<sup>19</sup> found that the methanol extracts of leaves of *E. foetidum* contained alkaloids, phenols, flavonoids, anthraquinones, and sterol. However, another study explored the presence of flavonoids, tannins, saponin, coumarins, and triterpenoid and the absence of alkaloids in the plant's leaves<sup>1,6,20</sup>. The chloroform extracts of dried aerial parts by soxhlet extraction followed by chromatographic fractionation yielded a pentacyclic triterpenoid saponin and O-glycoside<sup>21</sup>. Some free triterpenoids were too identified from leaves' hexane extract, including stigmasterol-the main phytosterol<sup>22,23</sup>.

*Eryngium foetidum* is a rich source of phenolic compounds<sup>5,24,25</sup>. Leitão *et al.*<sup>26</sup> found that the *E. foetidum* leaves showed major carotenoids as all-trans-lutein and all-trans- $\beta$ -carotene and major phenolic compounds as

chlorogenic acid and ferulic acid. The major constituent (E)-2-dodecenal with 14 compounds was identified in chloroform and methanol leaf extracts of *E. foetidum* via HPLC analysis. Moreover, another study found that the ethanol extracts *E. foetidum* shows higher flavonoid contents<sup>27</sup>.

Dried leaves yielded 0.1 to 0.95% of essential oil<sup>18</sup>. Gas chromatography-mass spectroscopy (GC-MS) analysis of essential oil identified 63 different compounds<sup>24</sup>. The linear unsaturated aldehyde, (E)-2-dodecenal was reported as the main constituent in the aldehydes, which were identified as the major constituents of the essential oil of E. foetidum6,12,14,24-25,28-32. A study showed that the (E)-2-dodecenal was responsible for the fruity, sweet, sour, and characteristic "cilantro" aroma of this plant with the highest flavor dilution factor33. The monoterpene hydrocarbons<sup>5</sup> and positional isomers of trimethylbenzaldehyde29 were present as the highest constituents from the stem and root essential oil of E. foetidum, respectively.

Seeds of *E. foetidum* yielded 0.2% of essential oil in which carotol (19.3%) was identified as the main constituent<sup>34</sup>. Acharya *et al.*<sup>35</sup> investigated the phytochemical analysis of *E. foetidum* found in coastal Odisha, India, and revealed that leaves and branches of *E. foetidum* consisted of chiefly 10-undecenal followed by 2,4,6-trimethylbenzaldehyde, (*Z*)-9-tetradecenal, and (*Z*)-7-tetradecenal. This study further suggested that the chemo profile of *E. foetidum* and seasonal *C. sativum* were the same, and *E. foetidum* can be used as an alternative to seasonal *C. sativum*. Structures of different chemical constituents of the leaves, stem, and root of *E. foetidum* were given in the appendix<sup>5,24,25,34</sup>.

### PHARMACOLOGICAL ACTIVITIES

Some compounds like (E)-2-dodecenal or 'Eryngial,' which present abundantly in *E. foetidum*, revealed to have

multiple bio-activities, and numerous studies revealed that the *E. foetidum* has anti-inflammatory, antioxidant, antimicrobial, antifungal, anti-helminthic, anti-tumor, anti-diabetic, antimalarial larvicidal and anticonvulsant activities. These activities are summarized in **Table II**.

### Anti-inflammatory activity

Several studies explored the anti-inflammatory activity of the plant with pathways of mechanisms responsible for the activity<sup>36</sup>. The hexane extract from the leaves of E. foetidum has shown possible anti-inflammatory activity<sup>3</sup>. Prior treatment with ethanol extract of E. foetidum leaf inhibited the elevation of interleukin 6 (IL-6), tumor necrosis factor-alpha (TNF-a), inducible nitric oxide synthase (iNOS), and cyclooxygenase (COX-2), together with their cognate messenger ribonucleic acid (mRNAs) in a dose-dependent manner. Nitric oxide (NO) and intracellular reactive oxygen species (ROS) contents were similarly reduced. This result suggested that E. foetidum leaf extract possesses suppressive effects on proinflammatory mediators, and the plant has the potential to reduce the risk of cancer associated with inflammation<sup>37</sup>. Leaf extract of E. foetidum consisted of lutein, β-carotene, chlorogenic acid, kaempferol, and caffeic acid, which had bioactive properties<sup>38</sup>.

The aqueous bioaccessible fraction of leaves of *E. foetidum*, prepared from simulated digestion, has inhibited IL-8 and MCP-1 levels in the human intestinal Caco-2 cells, which were stimulated with interleukin-1 $\beta$  (IL-1 $\beta$ ). Here, lipophilic constituents such as lutein and beta-carotene had a major role in the anti-inflammatory activity. Further study suggested that the consumption of *E. foetidum* could prevent intestinal inflammation<sup>38,39</sup>.

The topical anti-inflammatory activity of *E. foetidum* was studied in the TPA-induced mice; it was evident with the inhibition of myeloperoxidase (MPO) enzyme activity by hexane extract of *E. foetidum*<sup>40</sup>. Oral administration of decoction of *E. foetidum* inhibited carrageenan-induced

edema in rat paws and TPA-induced edema in the ears of mice. Further, this extract potently inhibited the abdominal writhing induced by acetic acid<sup>41</sup>. In India, the North East Institute of Science and Technology developed a drug formulation for treating arthritis and skin disease in which the essential oil of *E. foetidum* is one of the main components<sup>4</sup>.

### Antioxidant activity

Numerous studies have been conducted to evaluate the antioxidant activity of the plant. A study revealed that pretreatment with *E. foetidum* leaf extract (35-140 µg/ml) inhibited ROS generation mediated by NADPH oxidase in lipopolysaccharide (LPS)-induced mice<sup>37</sup>. Ethanol extract of the *E. foetidum* leaves exhibited 51.44% radical scavenging activity, correlated with a high level of phenolic and flavonoid contents<sup>42</sup>. Amazonian *E. foetidum* has shown that the scavenging capacity of methanol leaves extract against ABTS and DPPH<sup>26</sup>. Another study also proved the remarkable scavenging activity of methanol leaf extract against DPPH and FRAP assay<sup>27</sup>.

The oxygenated compounds like 2,4,6-trimethylphenol, linear saturated aliphatic alcohols, unsaturated aliphatic alcohols, and aliphatic aldehyde compounds of essential oil from the stem were identified as enhancers of antioxidant activity43, and 78.08% of radical scavenging activity was observed with the essential oil extracted from the stem of *E. foetidum*, which were comparable with the activity of standard ascorbic acid<sup>5</sup>. The active polar principles such as chlorogenic acid, caffeic acid, and kaempferol-phenolics in a bioaccessible fraction of E. foetidum extract have reduced intracellular ROS accumulation in the IL-1ß stimulated Caco-2 cells<sup>38</sup>. Methanol extract of E. foetidum shows the highest antioxidant activity compare to aqueous and chloroform extracts<sup>44</sup>. The DPPH assay revealed that the methanol extract has higher antioxidant activity than saponin and essential oils from leaves of *E. foetidum*, and also, the saponin of this plant has been shown to have antioxidant activity<sup>45</sup>. According to these results, the leaf and stem of *E. foetidum* have significant antioxidant properties, which vary due to the different methods of the extraction process<sup>46</sup>. The Nigerian *E. foetidum* volatile oils have shown that the potential source of natural antioxidants is the high acyclic aldehydes and aromatic compounds<sup>47</sup>. Moreover, a study found that these essential oil compounds from Colombian *E. foetidum* have antioxidant capacity against DPPH assay<sup>25</sup>. The methanolic extract of leaves of *E. foetidum* showed high antimicrobial activity, which could be related to the high concentration of polyphenols and flavonoids<sup>48</sup>.

#### Antibacterial activity

Methanol and chloroform extracts from E. foetidum leaves highly inhibited Streptococcus pneumoniae, Staphylococcus aureus, and Listeria monocytogenes (Gram positives bacteria), while aqueous extract highly inhibited Salmonella typhimurium (Gram-negative bacteria)<sup>44</sup>. Methanol extract showed a dose-dependent, mild to moderate antibacterial activity against S. aureus, while it was ineffective against any tested Gram-negative organisms (Escherichia coli and Pseudomonas aeruginosa)<sup>49</sup>. Another study was done with methanol, ethanol, and aqueous extracts from leaves of E. foetidum, S. aureus, Pseudomonas oleovorans, Klebsiella pneumoniae, Salmonella enteric, and E. coli were sensitive to at least one extract. At the same time, Bacillus megaterium, Bacillus subtilis, and Bacillus flexus were sensitive to all three extracts<sup>50</sup>. Another study reported that the blanched *E. foetidum* leaf suspension (10% w/v) with 100% growth inhibition in tested S. aureus and B. subtilis<sup>51</sup>. The methanol and ethanol extracts of the E. foetidum have shown a quite similar and good antibacterial potential to the aqueous extracts<sup>42</sup>. Pretreatment with essential oil of E. foetidum as an antimicrobial agent at 15 mg/L concentrations reduced

the pasteurization temperatures of pineapple juice to 60°C, where customarily conducted at 80 to 95°C for 15 to 30 seconds. Due to the antibacterial effect of *E. foetidum* essential oil, there is inactivation of *L. monocytogenes*<sup>28</sup>.

Ndip *et al.*<sup>22</sup> found that the methanol extract from *E. foetidum* leaves showed moderate antibacterial activity against six clinical strains of *Helicobacter pylori* out of 15 tested strains, using the disk diffusion method. Another study has shown that the methanol extract of *E. foetidum* has anti-helicobacter activity *in vitro* and *in vivo* compared to ciprofloxacin<sup>19</sup>. The ZnO nanoparticles prepared by incorporating leaf extracts of *E. foetidum* showed broadspectrum antibacterial activity<sup>52</sup>. The polyacetylenes of this plant also demonstrated antibacterial abilities<sup>1</sup>. Most of the studies evaluate the antibacterial activity of extracts only. Further studies need to evaluate the antibacterial activity of isolated phytoconstituents<sup>53</sup>.

#### Antifungal activity

Leaf and stem extracts of *E. foetidum* inhibited some fungi including *Candida albicans, Candida guilliermondi,* and *Cryptococcus neoformans* with minimum inhibitory concentrations of 256, 1024, and 32 µg/mL, respectively. Additionally, this extract showed a significant inhibitory effect towards strains of *C. neoformans*<sup>54</sup>. However, relatively few tests were carried out on fungi compared to bacteria.

### Anti-helminthic activity

The methanol extract of *E. foetidum* exhibited a dosedependent helminthicidal activity against *Paramphistomum sp*<sup>13</sup>. In another *in vitro* study using the infective larvae at the  $3^{rd}$  stage of *Strongyloides stercoralis, E. foetidum* showed the most effectiveness at a 50 mg/mL concentration among 25 Jamaican herbal extracts. Eryngial (trans-2-dodecenal) possessed anti-helminthic activity against infective larvae of *S. stercoralis* and was significantly more effective than ivermectin as a comparison drug<sup>50</sup>.

#### Anti-tumor activity

Extracts from E. foetidum can reduce the risk factors of cancer development associated with different inflammatory mediators. Inhibition of TNF-a expression by E. foetidum leaf extract would be helpful in cancer prevention involving reducing invasiveness of cancer cells<sup>37</sup>. This plant can also inhibit cell division, which is another significant controlling factor in tumor development<sup>39</sup>. The essential oil and methanol extract of leaves had selective inhibition towards the proliferation of PC-3 and A-549 cell lines<sup>45</sup>. Another study revealed no clastogenicity associated with freeze-dried leaves of E. foetidum and has anticlastogenic potential in mice using erythrocyte micronucleus assay. This study further suggested the potential health benefit of E. foetidum leaves<sup>55</sup>.

#### Anti-diabetic activity

Eryngium foetidum leaf extracts showed a amylase inhibitory activity in vitro56. Further, in vivo studies reported the reduced blood glucose levels (comparable to glibenclamide) in streptozotocin-induced diabetic rat models at doses of 250 and 500 mg/kg of the extract<sup>57</sup>. A study reported 52.2% inhibition of a-amylase activity using E. foetidum ethanol extract, where the slightest inhibition was observed with an aqueous extract<sup>42</sup>. Kusirisin et al.58 revealed that E. foetidum potentially prevents glycation associated with diabetes subjects. However, another study found that the aqueous extract E. foetidum was ineffective as an anti-hyperglycaemic agent on STZ induced rat models subjected to oral glucose tolerance test<sup>59</sup>. Another study mentioned that the E. foetidum plants of Manipur, India, could be used as herbal remedies for the treatment of diabetes mellitus<sup>42</sup>.

#### Antimalarial activity

A survey on the use of herbal remedies for malaria and leishmaniasis in Loreto of Peru reported that boiled plant material from *E. foetidum* had been used to treat malaria/leishmaniasis. Nevertheless, *in vitro* study was unable to confirm such activity<sup>60</sup>. However, another study identified that *E. foetidum* had an *in vitro* antiplasmodial activity with IC<sub>50</sub> more than 25  $\mu$ g/mL when traditionally using antimalarial herbal medications among Quechua and Mestizo populations native to Loreto in Peru-were tested against parasite cultures<sup>61</sup>. Ruiz *et al.*<sup>62</sup> found *in vitro* study that this plant has an antimalarial potential with IC<sub>50</sub> of more than 10  $\mu$ g/ml, where 59 locally using plants to treat malaria in Nanay river banks of Peru.

Aqueous extracts of the whole plant of *E. foetidum* have shown an anti-plasmodial activity against *Plasmodium gallinaceum in vivo*, in chicken models, where 476 plants from the American continent were reviewed activity against different strains of plasmodium<sup>63</sup>. Both *n*-hexane and ethyl acetate fractions from extracts of aerial parts of *E. foetidum* showed anti-leishmanial activity against *Leishmania tarentolae* and *Leishmania donovani*<sup>64</sup>. An extract rich in eryngial was patented for the treatment of parasites in humans and other mammals<sup>65,66</sup>.

#### Larvicidal activity

As much as 3% of brine shrimp larvae mortality was identified with aqueous and methanol extracts of *E. foetidum*, showed no mortality; 226 amazonian plants were tested to determine their lethal effect on brine shrimp larvae (nauplii) in an *in vitro* assay<sup>67</sup>. An *in vitro* study reported that the crude extract of this plant achieved 100% mortality against mosquito larva after 24 hours. Further, this study suggested that terpenes present in these plants could be the reason for this mosquito larvicidal activity<sup>68</sup>. Sumitha *et al.*<sup>69</sup> reported that the effectively used against fourth-instar *Aedes albopictus* larvae, in which 90% of larvicidal activity was exhibited. Further, this study emphasizes using this plant's essential oil as a natural insecticide with minimum side effects on

humans. Crude extract of *E. foetidum* showed the most toxic activity against the mosquito larvae. This plant can be used as a substitute for mosquito repellent coils<sup>70</sup>.

#### Anti-convulsant activity

This plant has been extensively used in traditional medicine to treat fits in Jamaica<sup>71,72</sup>. In an *in vivo* study, intraperitoneal administration of the extract of *E. foetidum* with the concentration of 110 g/250 mL showed antiepileptic activity against picrotoxin-induced convulsions in rats<sup>73,74</sup>. The aqueous extract of chopped *E. foetidum* leaves, when given orally at a dose of 100 mg/kg, was effective against picrotoxin-induced convulsions in albino rats<sup>71</sup>. Furthermore, Nsour *et al.* <sup>73</sup> reported that intraperitoneal injection of aqueous extracts of leaves and stem of *E. foetidum*, in rat models, exhibited anticonvulsant effects equal to phenobarbital.

foetidum		
Parts	Preparation / extract	Activities
Leaves	<i>n</i> -hexane	Anti-inflammatory <sup>40</sup>
	Ethanol	Anti-inflammatory <sup>37</sup>
	Aqueous	Anti-inflammatory <sup>38</sup>
	bioaccessible fraction	
	Methanol	Antioxidant <sup>26,27,44,45,52</sup>
	Methanol, water &	Antibacterial <sup>44</sup>
	chloroform	
	Methanol	Antibacterial <sup>19,22,49</sup>
	Methanol, ethanol &	Antibacterial <sup>42</sup>
	aqueous	
	Methanol & ethanol	Antibacterial <sup>42</sup>
	Essential oil	Antibacterial <sup>28</sup>
	Aqueous extract in	Antibacterial <sup>52</sup>
	ZnO nanoparticles	
	Methanol	Antifungal <sup>54</sup>
	Methanol	Anti-helminthic <sup>13,50</sup>
	Essential oil &	Anti-tumor <sup>45</sup>
	methanol	
	Aqueous	Antidiabetic <sup>57</sup>
	Ethanol	Antidiabetic <sup>42</sup>
	Aqueous	Antidiabetic <sup>58</sup>
	Aqueous	Anti-convulsant71,73,74
	Freeze dried leaves	Anticlastogenic <sup>55</sup>
	Aqueous	Anti-inflammatory <sup>38</sup>
	bioaccessible fraction	
Leaves	Essential oil	Antioxidant <sup>5,25,45</sup>
and stem		
Whole	Methanol	Antifungal &
plant		antioxidant <sup>54</sup>
	Aqueous	Antiplasmodial <sup>63</sup>
	Extract	Larvicidal <sup>67,70</sup>
	Essential oil	Larvicidal <sup>69</sup>
Aerial	<i>n</i> -hexane and ethyl	Antiplasmodial <sup>63</sup>

### TOXICITY STUDIES

The acute toxicity study of hydro-alcoholic extracts of E. foetidum did not show any morbidity, mortality, or post toxicity signs in albino mice, even at a dose of 2000 mg/kg74. Chandira et al.57 reported that the oral dose at 2500 mg/kg of aqueous extract of *E. foetidum* was lethal for healthy, adult, and young rodents. Acute oral toxicity of methanolic extract of leaves of E. foetidum was studied using rats to which extract was fed. This study revealed that  $LD_{50}$  of the extract showed 2000 mg/kg, and toxicity of the plant was extremely low upon oral administration<sup>48</sup>. Further genotoxicity of the methanolic extract of the plant revealed that plants protect damages to DNA induced by some substances. Also, the even higher concentration of extract did not produce DNA damages. It could be related to the higher antioxidant potential of the plant<sup>75</sup>.

Consumption of *E. foetidum* above 0.8%, which is more than 35 times human consumption, resulted in weight loss, kidney and spleen damage, as demonstrated through chronic toxicity animal studies. Further elevation of blood urea nitrogen was observed due to kidney damage by renal tubular-nephrosis, interstitial nephritis, and spleen hemosiderosis due to toxic metabolites of *E. foetidum*<sup>76</sup>. *Eryngium foetidum* was influential in the amelioration of carbon tetrachlorideinduced hepatotoxicity in mice<sup>75</sup>. This study demonstrated the hepatoprotective potential of this plant. Furthermore, the compound 'eryngial' as the main constituent of essential oil of *E. foetidum* showed significant inhibition on enzyme cytochrome P450 2E1<sup>6</sup>.

### CONCLUSION

*Eryngium foetidum* L is a potential aromatic crop and is widely used for ethnomedical and culinary purposes. Furthermore, it exhibited many pharmacological applications such as anti-inflammatory, antioxidant,

parts

acetate fractions

antimicrobial, anticonvulsant, antimalarial, anthelminthic, larvicidal, anticancer, anti-diabetic, and hepatoprotective activities with no cytotoxic effects. Even though several research studies were done to screen pharmacological activities, limited studies have been conducted to isolate and screen phytoconstituents of the plant. Studies should be focused on identifying the bioactives with potential activities. This review promised that potential new chemical entities could be elicited from phytoconstituents of *E. foetidum*, and also this plant can also be used as a substitute for mosquito repellent coils. Therefore, further studies should be done to prove this effect in the future.

# ACKNOWLEDGMENT

None.

## **AUTHORS' CONTRIBUTION**

Galamula Hewage Tharuka Kanchana Hemachandra: conceived and design the analysis, writing of the paper. Sinnadurai Thuvaragan: conceived and design the analysis. Vinotha Sanmugarajah: collected data, contributed data or analysis tools.

# DATA AVAILABILITY

None.

# CONFLICT OF INTEREST

The authors declare no conflict of interest.

### REFERENCES

1. Parham S, Kharazi AZ, Bakhsheshi-Rad HR, Nur H, Ismail AF, Sharif S, et al. Antioxidant, Antimicrobial and Antiviral Properties of Herbal Materials. Antioxidants. 2020;9(12):1309. doi:10.3390/antiox9121309

- Erdem SA, Nabavi SF, Orhan IE, Daglia M, Izadi M, nabavi SM. Blessings in disguise: a review of phytochemical composition and antimicrobial activity of plants belonging to the genus Eryngium. Daru. 2015;23:53. doi:10.1186/s40199-015-0136-3
- Hossain MA, Jashimuddin M, Nath TK, O'Reilly P. Spiny coriander (Eryngium foetidum L.) cultivation in the Chittagong Hill Tracts of Bangladesh: Sustainable agriculture innovation by indigenous communities. Indian J Tradit Knowl. 2017;16(1):59-67. handle:123456789/37014
- 4. Singh BK, Ramakrishna Y, Ngachan SV. Spiny coriander (Eryngium foetidum L.): a commonly used, neglected spicing-culinary herb of Mizoram, India. Genet Resour Crop Evol. 2014;61:1085-90. doi:10.1007/s10722-014-0130-5
- Thomas PS, Essien EE, Ntuk SJ, Choudhary MI. Eryngium foetidum L. Essential Oils: Chemical Composition and Antioxidant Capacity. Medicines. 2017;4(2):24. doi:10.3390/medicines4020024
- Paul JHA, Seaforth CE, Tikasingh T. Eryngium foetidum L.: a review. Fitoterapia. 2011;82(3):302-8. doi:10.1016/j.fitote.2010.11.010
- Geck MS, Cristians S, Berger-González M, Casu L, Heinrich M, Leonti M. Traditional Herbal Medicine in Mesoamerica: Toward Its Evidence Base for Improving Universal Health Coverage. Front Pharmacol. 2020;11:1160. doi:10.3389/fphar.2020.01160
- Wunderlin RPBF, Hansen ARF, Essig FB. Atlas of Florida Plants [Internet]. (Landry SM, Campbell KN, USF Water Institute). Institute for Systematic Botany, University of South Florida, Tampa; 2021 [cited 2021 Jun 28]. Available from https://florida.plantatlas.usf.edu/
- 9. Duke JA, Bogenschutz-Godwin MJ, duCellier J, Duke PAK. Handbook of Medicinal Herbs. 2nd ed. Florida (US): CRC Press; 2002.
- Joshi RK, Satyal P, Setzer WN. Himalayan Aromatic Medicinal Plants: A Review of their Ethnopharmacology, Volatile Phytochemistry, and Biological Activities. Medicines. 2016;3(1):6. doi:10.3390/medicines3010006
- Banout J, Havlik J, Kulik M, Kloucek P, Lojka B, Valterova I. Effect of solar drying on the composition of essential oil of sacha culantro (Eryngium foetidum L.) Grown in the peruvian amazon. J Food Process

Eng. 2010;33(1):83-103. doi:10.1111/j.1745-4530.2008.00261.x

- 12. Martins AP, Salgueiro LR, da Cunha AP, Vila R, Cañigueral S, Tomi F, et al. Essential Oil Composition of Eryngium foetidum from S. Tomé e Príncipe. J Essent Oil Res. 2003;15(2):93-5. doi:10.1080/10412905.2003.9712077
- 13. Swargiary A, Daimari A, Daimari M, Basumatary N, Narzary E. Phytochemicals, antioxidant, and anthelmintic activity of selected traditional wild edible plants of lower Assam. Indian J Pharmacol. 2016;48(4):418-23. doi:10.4103/0253-7613.186212
- 14. Wong KC, Feng MC, Sam TW, Tan GL. Composition of the Leaf and Root Oils of Eryngium foetidum L. J Essent Oil Res. 1994;6(4):369-74. doi:10.1080/10412905.1994.9698401
- 15. Xu Y, Liang D, Wang GT, Wen J, Wang RJ. Nutritional and Functional Properties of Wild Food-Medicine Plants from the Coastal Region of South China. J Evid Based Integr Med. 2020;25:2515690X20913267. doi:10.1177/2515690X20913267
- Mabeku LBK, Flaurant TT, Jacques K. Screening of some Cameroonian Medicinal Plants against Bacterial and Yeasts involved in Gastrointestinal Disorders. Int J Biol. 2017;9(3):56-64. doi:10.5539/ijb.v9n3p56
- 17. Usharani L, Singh WRC, Surodhani S, Singh WN. Pharmacognostical evaluation and antibacterial activity of medicinally important spices occurred in local area of Manipur. Asian J Plant Sci Res. 2016;6(2):42-5.
- Shavandi MA, Haddadian Z, Ismail MHS. Eryngium foetidum L. Coriandrum sativum and Persicaria odorata L.: A Review. J Asian Sci Res. 2012;2(8):410-26.
- Mabeku LBK, Bille BE, Ngeupi E. In Vitro and In Vivo Anti-Helicobacter Activities of Eryngium foetidum (Apiaceae), Bidens pilosa (Asteraceae), and Galinsoga ciliata (Asteraceae) against Helicobacter pylori. Biomed Res Int. 2016;2016:2171032. doi:10.1155/2016/2171032
- 20. Nataraj K, Girisha ST, Raghavendra VB. Miracle plant Eryngium foetidum Linn- A review on Ethnobotanical, Phytochemical composition and Pharmacology. Int J Adv Sci Technol. 2020;29(2):1099-115.

- 21. Anam EM. A novel triterpenoid saponin from Eryngium foetidum. Indian J Chem Sect B. 2002;41B:1500-3. handle:123456789/22011
- 22. Ndip RN, Tarkang AEM, Mbullah SM, Luma HN, Malongue A, Ndip LM, et al. In vitro anti-Helicobacter pylori activity of extracts of selected medicinal plants from North West Cameroon. J Ethnopharmacol. 2007;114(3):452-7. doi:10.1016/j.jep.2007.08.037
- 23. Wang P, Su Z, Yuan W, Deng G, Li S. Phytochemical Constituents and Pharmacological Activities of Eryngium L. (Apiaceae). Pharm Crop. 2012;3:99-120. doi:10.2174/2210290601203010099
- 24. Chowdhury JU, Nandi NC, Yusuf M. Chemical Constituents of Essential Oil of the Leaves of *Eryngium foetidum* from Bangladesh. Bangladesh J Sci Ind Res. 2007;42(3):347-52. doi:10.3329/bjsir.v42i3.674
- 25. Jaramillo BE, Duarte E, Martelo I. Volatile chemical composition of the essential oil from colombian Eryngium foetidum L. and determination of its antioxidant activity. Rev Cubana Plant Med. 2011;16(2):140-50.
- 26. Leitãoa DdSTC, Siqueira FC, de Sousa SHB, Mercadante A, Chisté RC, Lopes AS. Amazonian Eryngium foetidum leaves exhibited very high contents of bioactive compounds and high singlet oxygen quenching capacity. Int J Food Prop. 2020;23(1):1452-64. doi:10.1080/10942912.2020.1811311
- 27. Ashwaty PM, Saj OP. Carminative, Phytochemical and Antioxidant Potentialities of the Leaf Extracts of Eryngium foetidum L. (Apiaceae). World J Pharm Pharm Sci. 2014;3(6):2269-80.
- 28. Ngang JJE, Nyegue MA, Ndoye FC, Kamgain ADT, Kamdem SLS, Lanciotti R, et al. Characterization of Mexican coriander (Eryngium foetidum) essential oil and its inactivation of Listeria monocytogenes in vitro and during mild thermal pasteurization of pineapple juice. J Food Prot. 2014;77(3):435-43. doi:10.4315/0362-028x.jfp-13-323
- Thi NDT, Anh TH, Thach LN. The Essential Oil Composition of Eryngium foetidum L. in South Vietnam Extracted by Hydrodistillation under Conventional Heating and Microwave Irradiation. J Essent Oil-Bear Plant. 2008;11(2):154-61. doi:10.1080/0972060X.2008.10643612

- 30. Cardozo E, Rubio M, Rojas LB, Usubillaga A. Composition of the Essential Oil from the Leaves of Eryngium foetidum L. from the Venezuelan Andes. J Essent Oil Res. 2004;16(1):33-4. doi:10.1080/10412905.2004.9698645
- Leclercq PA, Duñg NX, Lô VN, Toanh NV. Composition of the Essential Oil of Eryngium foetidum L. from Vietnam. J Essent Oil Res. 1992;4(4):423-4. doi:10.1080/10412905.1992.9698097
- 32. Yeh PH. Essential Oils XVI Oil of Eryngium Foetidum, Linn. J Chin Chem Soc. 1974;21(3):139-47. doi:10.1002/jccs.197400019
- 33. Quynh CTT, Kubota K. Aroma Constituents and Enzyme Activities of Japanese Long Coriander Leaves (Culantro, Eryngium foetidum L.). Food Sci Technol Res. 2012;18(2):287-94. doi:10.3136/fstr.18.287
- 34. Pino JA, Rosado A, Fuentes V. Chemical Composition of the Seed Oil of Eryngium foetidum L. from Cuba. J Essent Oil Res. 1997;9(1):123-4. doi:10.1080/10412905.1997.9700731
- Acharya GC, Ponnam N, Kumari M, Roy TK, Shivashankara KS, Sahoo MR. Phytochemical profiling of spiny coriander (Eryngium foetidum L.)
  A potential perennial spicing-culinary herb of eastern India. Acta Chromatogr. 2021;34(2):197-202. doi:10.1556/1326.2021.00909
- Azab A, Nassar A, Azab AN. Anti-Inflammatory Activity of Natural Products. Molecules. 2016;21(10):1321. doi:10.3390/molecules21101321
- Mekhora C, Muangnoi C, Chingsuwanrote P, Dawilai S, Svasti S, Chasri K, et al. Eryngium foetidum suppresses inflammatory mediators produced by macrophages. Asian Pac J Cancer Prev. 2012;13(2):653-64. doi:10.7314/apjcp.2012.13.2.653
- Dawilai S, Muangnoi C, Praengamthanachoti P, Tuntipopipat S. Anti-Inflammatory Activity of Bioaccessible Fraction from Eryngium foetidum Leaves. Biomed Res Int. 2013;2013:958567. doi:10.1155/2013/958567
- 39. Suttisansanee U, Thiyajai P, Chalermchaiwat P, Wongwathanarat K, Pruesapan K, Charoenkiatkul S, et al. Phytochemicals and In Vitro Bioactivities of Aqueous Ethanolic Extracts from Common Vegetables in Thai Food. Plants. 2021;10(8):1563. doi:10.3390/plants10081563

- 40. García MD, Sáenz MT, Gómez MA, Fernández MA. Topical antiinflammatory activity of phytosterols isolated from Eryngium foetidum on chronic and acute inflammation models. Phytother Res. 1999;13(1):78-80. doi:10.1002/(sici)1099-1573(199902)13:1<78::AID-PTR384>3.0.CO;2-F
- 41. Sáenz MT, Fernández MA, García MD. Antiinflammatory and analgesic properties from leaves of Eryngium foetidum L. (Apiaceae). Phytother Res. 1998;11(5):380-3. doi:10.1002/(SICI)1099-1573(199708)11:5<380::AID-PTR116>3.0.CO;2-%23
- Malik T, Pandey DK, Roy P, Okram A. Evaluation of Phytochemicals, Antioxidant, Antibacterial and Antidiabetic Potential of Alpinia galanga and Eryngium foetidum Plants of Manipur (India). Pharmacogn J. 2016;8(5):459-64. doi:10.5530/pj.2016.5.8
- 43. Gyesi JN, Opoku R, Borquaye LS. Chemical Composition, Total Phenolic Content, and Antioxidant Activities of the Essential Oils of the Leaves and Fruit Pulp of Annona muricata L. (Soursop) from Ghana. Biochem Res Int. 2019;2019:4164576. doi:10.1155/2019/4164576
- 44. Dalukdeniya DACK, Rathnayaka RMUSK. Comparative Study on Antibacterial and Selected Antioxidant Activities of Different Eryngium foetidum Extracts. J Appl Life Sci Int. 2017;12(4):1-7. doi:10.9734/JALSI/2017/34378
- 45. Chandrika R, Jagath V, Thara-Saraswathi KJ. In Vitro Antioxidant and Anti-Proliferative Activities in Eryngium Foetidum L. Int J Pharm Res Health Sci. 2016;4(2):1110-6.
- 46. Mateos-Maces L, Chávez-Servia JL, Vera-Guzmán AM, Aquino-Bolaños EN, Alba-Jiménez JE, Villagómez-González BB. Edible Leafy Plants from Mexico as Sources of Antioxidant Compounds, and Their Nutritional, Nutraceutical and Antimicrobial Potential: A Review. Antioxidants. 2020;9(6):541. doi:10.3390/antiox9060541
- 47. Attah AF, Fagbemi AA, Olubiyi O, Dada-Adegbola H, Oluwatudon A, Elujoba A, et al. Therapeutic Potentials of Antiviral Plants Used in Traditional African Medicine With COVID-19 in Focus: A Nigerian Perspective. Front Pharmacol. 2021;12:596855. doi:10.3389/fphar.2021.596855
- 48. Raunelli P, Liviac D, Alvis R, Puente S, Best I, Reategui O. Cytoprotective Effect of the Eryngium

foetidum "Sacha Culantro" Methanolic Leaf Extract Versus Sodium Fluoride Exposed Mice using the Micronucleus Test and the Comet Assay. Pharmacogn J. 2019;11(3):461-5. doi:10.5530/pj.2019.11.72

- Dutta S, Bhattacharjee A, Yadav M, Shougrakpam P, Monin RG, Das A, Devi MN. Antibacterial activity of spiny coriander (eryngium foetidum linn.) on gram positive and gram negative bacteria. Int J Recent Sci Res. 2017;8(9):19959-62. doi:10.24327/ijrsr.2017.0809.0795
- 50. Forbes WM, Gallimore WA, Mansingh A, Reese PB, Robinson RD. Eryngial (trans-2-dodecenal), a bioactive compound from Eryngium foetidum: its identification, chemical isolation, characterization and comparison with ivermectin in vitro. Parasitology. 2014;141(2):269-78. doi:10.1017/s003118201300156x
- 51. Homer S, Baccus-Taylor GSH, Akingbala JA. Antibacterial Efficacy of Eryngium foetidum (Culantro) against Select Food-borne Pathogens. In: Hutchinson SD, editor. Proceedings of the 27th West Indies Agricultural Economics Conference. 2007 Jul; Belize City, Belize. St. Augustine (TT): Caribbean Agro-Economic Society; 2007. p. 179-92.
- 52. Begum S, Ahmaruzzaman M, Adhikari PP. Ecofriendly bio-synthetic route to synthesize ZnO nanoparticles using Eryngium foetidum L. and their activity against pathogenic bacteria. Mater Lett. 2018;228:37-41. doi:10.1016/j.matlet.2018.05.091
- 53. Altemimi A, Lakhssassi N, Baharlouei A, Watson DG, Lightfoot DA. Phytochemicals: Extraction, Isolation, and Identification of Bioactive Compounds from Plant Extracts. Plants. 2017;6(4):42. doi:10.3390/plants6040042
- 54. Tchuenguem RT, Kechia FA, Kuiate JR, Dzoyem JP. Ethnopharmacological survey, antioxidant and antifungal activity of medicinal plants traditionally used in Baham locality (Cameroon) to treat fungal infections. Arch Med Biomed Res. 2017;3(2):91-103. doi:10.4314/ambr.v3i2.5
- 55. Promkum C, Butryee C, Tuntipopipat S, Kupradinun P. Anticlastogenic effect of Eryngium foetidum L. assessed by erythrocyte micronucleus assay. Asian Pac J Cancer Prev. 2012;13(7):3343-7. doi:10.7314/apjcp.2012.13.7.3343
- 56. Singh S, Singh DR, Banu S, Salim KM. Determination of bioactives and antioxidant activity in Eryngium

foetidum L.: A traditional culinary and medicinal herb. Proc Natl Acad Sci India Sect B Biol Sci. 2013;83:453–60. doi:10.1007/s40011-012-0141-y

- 57. Chandira RM, Jaykar B. Extraction, Pharmacological Evaluation and Formulation of Selected Medicinal Herbs for Antidiabetic Activity. Int J Pharm Teach Pract. 2013;4(1):458-82.
- 58. Kusirisin W, Srichairatanakool S, Lerttrakarnnon P, Lailerd N, Suttajit M, Jaikang C, et al. Antioxidative activity, polyphenolic content and anti-glycation effect of some Thai medicinal plants traditionally used in diabetic patients. Med Chem. 2009;5(2):139-47. doi:10.2174/157340609787582918
- 59. Offiah NV, Seaforth CE, Sampath S, Rajack R, Mohammed A. An evaluation of the blood glucose lowering effects of Eryngium foetidum. In: Clement YN, Seaforth CE, editors. Proceedings of the Advancing Caribbean herbs in the 21st century. 2005 Jun 23-26; Rodney Bay, Saint Lucia. St. Augustine (TT): University of the West Indies; 2006. p. 30-5.
- 60. Kvist LP, Christensen SB, Rasmussen HB, Mejia K, Gonzalez A. Identification and evaluation of Peruvian plants used to treat malaria and leishmaniasis. J Ethnopharmacol. 2006;106(3):390-402. doi:10.1016/j.jep.2006.01.020
- 61. Roumy V, Garcia-Pizango G, Gutierrez-Choquevilca A-L, Ruiz L, Jullian V, Winterton P, et al. Amazonian plants from Peru used by Quechua and Mestizo to treat malaria with evaluation of their activity. J Ethnopharmacol. 2007;112(3):482-9. doi:10.1016/j.jep.2007.04.009
- Ruiz L, Ruiz L, Maco M, Cobos M, Gutierrez-Choquevilca A-L, Roumy V. Plants used by native Amazonian groups from the Nanay River (Peru) for the treatment of malaria. J Ethnopharmacol. 2011;133(2):917-21. doi:10.1016/j.jep.2010.10.039
- 63. Mariath IR, Falcão HdS, Barbosa-Filho JM, de Sousa LCF, Tomaz ACdA, Batista LM, et al. Plants of the American continent with antimalarial activity. Rev Bras Farmacogn. 2009;19(1a):185-92. doi:10.1590/S0102-695X2009000100026
- Rojas-Silva P, Graziose R, Vesely B, Poulev A, Mbeunkui F, Grace MH, et al. Pharm Biol. 2014;52(3):398-401. doi:10.3109/13880209.2013.837077
- 65. Forbes WM, Steglich C, inventors. Slippery Rock University of Pennsylvania, Slippery Rock,

Philadelphia, US, assignee. *Methods of treating infectious diseases*. International Patent (US) 20090047342. 2007-08-15.

- 66. Segarra-Newnham M. Manifestations, diagnosis, and treatment of Strongyloides stercoralis infection. Ann Pharmacother. 2007;41(12):1992-2001. doi:10.1345/aph.1k302
- 67. Quignard ELJ, Pohlit AM, Nunomura SM, Pinto ACdS, dos Santos EVM, de Morais SKR, et al. Screening of plants found in Amazonas State for lethality towards brine shrimp. Acta Amaz. 2003;33(1):93-104. doi:10.1590/1809-4392200331104
- 68. Liu M, Panda SK, Luyten W. Plant-Based Natural Products for the Discovery and Development of Novel Anthelmintics against Nematodes. Biomolecules. 2020;10(3):426. doi:10.3390/biom10030426
- Sumitha KV, Prajitha V, Sandhya VN, Anjana S, Thoppil JE. Potential Larvicidal Principles in Eryngium foetidum L. (Apiaceae), An Omnipresent Weed, Effective Against Aedes albopictus Skuse. J Essent Oil-Bear Plant. 2014;17(6):1279-86. doi:10.1080/0972060X.2014.958544
- 70. Mangalat S, Narayanan V, Janardhanan M. Herbal larvicides to control mosquito larvae: A preliminary study. Nat Prod Radiance. 2004;3(1):24-6.
- 71. Kumar S, Madaan R, Bansal G, Jamwal A, Sharma A. Plants and Plant Products with Potential Anticonvulsant Activity – A Review. Pharmacogn Commun. 2012;2(1s):3-99. doi:10.5530/pc.2012.suppl1.2
- 72. Júnior LJQ, Almeida JRGS, Lima JT, Nunes XP, Siqueira JS, de Oliveira LEG, et al. Plants with anticonvulsant properties: a review. Rev Bras Farmacogn. 2008;18(Suppl):789-819. doi:10.1590/S0102-695X2008000500026
- 73. Nsour WM, Lau CB, Wong IC. Review on phytotherapy in epilepsy. Seizure. 2000;9(2):96-107. doi:10.1053/seiz.1999.0378
- 74. Chakraverty R, Nath N, Debnath T, Ghosh A, Datta S. Pharmacological Assessment of the Hydroalcoholic Extracts of Leaves of Eryngium foetidum Linn. (EFHA). Int J Curr Trend Pharmacobiol Med Sci. 2016;1(2):75-80.
- 75. Rodrigues E. Plants of restricted use indicated by three cultures in Brazil (Caboclo-river dweller, Indian

and Quilombola). J Ethnopharmacol. 2007;111(2):295-302. doi:10.1016/j.jep.2006.11.017

76. Janwitthayanuchit K, Kupradinun P, Rungsipipat A, Kettawan A, Butryee C. A 24-Weeks Toxicity Study of Eryngium foetidum Linn. Leaves in Mice. Toxicol Res. 2016;32(3):231-7. doi:10.5487/tr.2016.32.3.231