



Available Online at EScience Press

Journal of Plant and Environment

ISSN: 2710-1665 (Online), 2710-1657 (Print)

<https://esciencepress.net/journals/JPE>

Uses of Ethnomedicinal Plants for the Treatment of Ulcer in Pakistan

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ARTICLE INFO

Article History

Received: January 05, 2022

Revised: March 20 2022

Accepted: April 03, 2022

Keywords

Ulcer treatment

Ethnomedicinal plants

Pharmacological research

Anthropogenic activities

ABSTRACT

The main objective of the study is to summarize the uses of plants by local communities in Pakistan for the treatment of ulcer. In the present review, the data regarding antiulcerogenic plants were collected by searching different online data banks various research publications, PubMed, different books, thesis copies from libraries, and other national organizations. In the current review article, a total of 109 species belonging to 55 families were found to be used against ulcer treatment. Among these Asteraceae was the predominant family containing (11species) followed by Fabaceae (6 species), Asclepiadaceae (5 species), Chenopodiaceae (5 species), Pinaceae (5 species), Brassicaceae (4 species), Polygonaceae (4 species), and, so on. The highest number species used were herbaceous (67.88 %) followed by trees (17.43 %) and shrubs (14.67 %). The majority of the plants reported have significant and useful properties, but certain are cited in several papers that have sparked interest in further pharmacological research. It was occluded that whole plant and anthropogenic activities can make special interest to be exploited sustainably.

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INTRODUCTION

Ulcer syndrome has been a leading cause of gastrointestinal surgical treatments, with high morbidity and fatality rates (Chung *et al.*, 2017). Ulcer is a disorder that causes severe bleeding in patients, open lesions on the epidermis of skin or a mucosa depicted by collapse of tissue (Chan *et al.*, 2004). Generally, ulcer is a well-known disorder that can build up in the gastrointestinal system for example in the (mouth, esophagus, stomach, duodenum, jejunum, Zollinger-Ellison syndrome, and in connection with a Meckel's diverticulum having ectopic gastric mucosa). The ulcer also causes gentle abdominal

suffering catastrophic harm and bleeding to patients, and even mental disorders (Pradip *et al.*, 2012). Ulcer are classified into various forms peptic, mouth, esophagus and peptic ulcer of stomach (Debjit *et al.*, 2010). Gastric and duodenal ulcers are the 2 most common types of peptic ulcers. In the stomach patients feel severe pain and such ulcer is most common in the older age group. Other manifestations are sickness; weight loss and vomiting for the reason that too much eating increases the stomach pain (Vyawaharen *et al.*, 2009).

Gastric ulcers are present in the small intestine because of severe pain in the upper abdomen. Ulcers often appear

on the posterior and anterior walls of duodenum (Wong *et al.*, 2004). In several cases, peptic ulcer cause death due to the symptoms of abdominal pain, bloody stool, and contractions along with blood vomiting (Chung and Shelat, 2017). The peptic ulcer physiology infection involves an unevenness between offensive and defensive factors (Hoogerwerf *et al.*, 2001, Tarnawski 2005, Kaunitz, 2004, Bandyopadhyay *et al.*, 2001). Peptic ulcer has been found by researchers due to bacterial infections (*Helicobacter pylori*) or result in a variety of medications (Marshall *et al.*, 1984). *Helicobacter pylori*, stress, nonsteroidal anti-inflammatory drugs, alcohol abuse, and smoking are the major causative factors associated with gastric ulcer (Malfertheiner *et al.*, 2009). A gastric ulcer is a worldwide known gastrointestinal disease that affects 10 % of the world's population (Zapata-Colindres *et al.*, 2006).

Pakistan is situated in South Asia and is at the seam of Central Asia having a land boundary 6,774 km length Pakistan geographical position is very important and it has India in the east while Afghanistan and Iran in the west surrounding China in the northeast. Pakistan has diverse flora represented by just about 6000 wild plant species are habitually common in the Himalaya and Karakorum regions (Abbasi *et al.*, 2010; Tardio *et al.*, 2008). In Pakistan, little interest has been paid to the ethnomedicinal standards of medicinal plants (Qureshi *et al.*, 2006; Husain *et al.*, 2008; Qureshi *et al.*, 2009; Mahmood *et al.*, 2011). About 75-80 % of the world population still utilize herb medicine mostly in growing countries for basic health needs (Kumar *et al.*, 2011). The presence of major secondary metabolites including flavonoids and tannins for antiulcer action was discovered by a phytochemical screening of medicinal plants (Patel *et al.*, 2010). The present reserach was aim to review the medicinal herbs for ulcers and put together potency for different biological processes in the latest investigation. The current review was focused to document the ethnomedicinal plant's knowledge and usually medicinal plants against ulcer patients in Pakistan.

Plants used for treatment of Ulcer and their diversity in Pakistan

Ethnomedicinal investigations in various parts of Pakistan have revealed the indigenous use of medicinal herbs in the treatment of ulcers. In previous articles, a large number of local plants were used for the ulcer treatment. ((Hussain *et al.*, 2018; Parvaiz 2014; Akhtar *et al.*, 2013).

In the recent review, it was discovered that the majority of these research lacked adequate information on the oral usage of the herbs. Approximately twenty percent of the plant species of the world are probably used in health management systems (Baillie *et al.*, 2004). In the current review research, 109 species from 55 families were discovered to be helpful for the treatment of ulcers.

Among these Asteraceae was leading family having (11species) followed by 6 Fabaceae species, 5 Asclepiadaceae species, 5 Chenopodiaceae species, 5 Pinaceae species, 4 Brassicaceae species, 4 Polygonaceae species, 3 Apiaceae species, 3 Boraganaceae species, 3 Convolvulaceae species, 3 Cucurbitacea species, 3 Lamiaceae species, 3 Mimosaceae species, 2 Aizoaceae species, 2 Capparidaceae species, 2 Cyperaceae species, 2 Malvaceae species, 2 Plantaginaceae species, 2 Portulacaceae species, 2 Rosaceae species, 2 Saxifragaceae species, 2 Solanaceae species, 2 Tamaracaceae species, 1 Alliaceae species, 1 Amaranthaceae species, 1 Apocynaceae species, 1 Araliaceae species, 1 Aristolochiaceae species, 1 Asphodelaceae species, 1 Berberidaceae species, 1 Cactaceae species, 1 Caryophyllaceae species, 1 Cuscutaceae species, 1 Dioscoreaceae species, 1 Dryopteridaceae species, 1 Euphorbiaceae species, 1 Geraniaceae species, 1 Juglandiaceae species, 1 Linaceae species, 1 Meliaceae species, 1 Molluginaceae species, Nyctaginaceae 1 species, Oleaceae 1 species, Orobanchaceae 1 species, Poaceae 1 species, 1 Punicaceae species, 1 Rhamanaeae species, 1 Rubiaceae species, 1 Salicaceae species, 1 Scrophulariaceae species, 1 Ulmaceae species, 1 Verbenaceae species, 1 Violaceae species and 1 Salvadoraceae species. Compare to previous literature indicates that the Asteraceae was frequently represented in the study areas (Wazir *et al.*, 2004; Gorsl and Mairaj, 2002; Abbas *et al.*, 2013, Abbasi *et al.*, 2013).

Different plants have been reported due to their wide distribution pattern and ethnomedicinal potential in 8, 7, 6, 4, and 3 research papers, these plants are well-known as antiulcerogenic plants. Among different studies, *Withania somnifera* is reported in 8 publications have the highest value as an antiulcerogenic plant. *Withania somnifera* is followed by *Calotropis Procera* having (7 citations), *Carthamus oxycantha* (6 citations), *Fragaria nubicola* (6 citations), *Bergenia ciliate* (4 citations), *Cleome viscosa* (4 citations), *Pinus roxburghii* (4 citations)

and *Tamarix dioica* (4 citations). Eleven plants *Ajuga bracteosa*, *Asphodelus tenuifolius*, *Berberis lyceum*, *Bergenia ciliata*, *Bistorta amplexicaulis*, *Capparis deciduas*, *Chenopodium album*, *Dryopteris ramosa*, *Geranium wallichianum*, *Olea ferruginea* and *Solanum nigrum* are reported in 3 publications each is ethnomedicinally used against ulcer.

Plants sections used

In Pakistan, practically all components of the plant, including the whole plant, leaves, seeds, roots, bark, and fruit, flower, rhizome, latex, stem, aerial parts, oil, resin, bulb, shoot, and wood are used in the ulcer treatments (Arif *et al.*, 2021). Data analysis shows that Whole plant (28.48 %) are mostly used against ulcer followed by leaves (24.24 %), seeds (10.30 %), roots (9.09 %), bark (7.87 %), fruit (7.27 %), flower (6.66 %), rhizome (2.42 %), latex (2.42 %), stem (1.81 %), aerial parts (1.21 %), oil (1.21 %), resin (1.21 %), bulb (0.60 %), shoot (0.60 %) and wood (0.60 %). Leaves were reported as frequently used part in the ethnobotanical study of tehsil Kabal, district Swat, Pakistan (Khan *et al.*, 2015). However, if mistakenly exploited, the use of root and whole plant in remedy compositions may exert pressure on the flora.

Preparation, forms, dose and using time of herbal remedies

Out of total 109 plant species, different people in Pakistan against ulcer use different methods of preparation of remedies. However, in the majority of the reports preparation methods and the dosage time is not declared. Various people employ different portions of the plant in their cures, such as fresh parts, dry parts, or both (fresh and dry) at a time. Water is used mostly as a medium for preparing recipes while occasionally milk, oil, and butter are used for application. Decoction (22.44 %) is the most commonly utilized formulation method, followed by infusion (19.72 %), powder (18.36 %), extract (12.92 %), vegetable (7.48 %), directly used (5.44 %), juice (4.08 %), paste (4.08 %), poultice (2.72 %), ointment (2.04 %) and demulcent (0.68 %). According to the present review article, the people of Pakistan have kept medicinal plant remedies on hand in case of emergency. The highest number species used for ulcer treatment were an herbaceous habit (67.88%) followed by trees (17.43 %) and shrubs (14.67 %) (Ullah *et al.*, 2013).

Ethnomedicinal and pharmacological validation of important plants

Medicinal plants are an important element in pharmaceutical industry to find out active ingredients

and develop a new drug. An appropriate way for selecting plants for complete pharmacological screenings is to employ ethnomedicinal assessment (Redzi, 2007; Sari - Kundali *et al.*, 2010). The reported plants in the current review investigate various parts of Pakistan and the rest of the World with most of the plants having strong curative and beneficial effects as antiulcerogenic plants.

Calotropis Procer is a perennial shrub, component of Asclepiadaceae. The whole plant decoction, extract, powder and infusion is used to treat ulcer (Ahmad *et al.*, 2014). The plant extracts were tested for antiulcer activity in pylorus ligated rats and prominent protection was observed in histamine-induced duodenal ulcers in guinea pigs (Yadav *et al.*, 2011).

Carthamus oxycantha is a small herb and belongs to Asteraceae. The seeds and leaves paste and extract are used to treat ulcer (Adeel *et al.*, 2011, Mahmood *et al.*, 2013, Taj *et al.*, 2009, Hussain *et al.*, 2010, Hayat *et al.*, 2008, Choudhary *et al.*, 2014). Methanolic extract of seeds of *Carthamus oxycantha* was evaluated in different acute gastric ulceration in albino mice prevented the increase of gastric mucosal lesion and decreased the gastric toxicity produced by ulcerogenic agents (Hussain *et al.*, 2015).

Chenopodium album is an annual herb and belongs to Chenopodiaceae. The whole plant is used as a vegetable and the infusion is used in the treatment of ulcer (Khan, S. W. and S. Khatoon 2008, Ahmad *et al.*, 2014, Khan *et al.*, 2013b). The ethanolic extract was analyzed in rats to check the antiulcer activity through three models and ranitidine was used as reference standard. The alcoholic withdraw significantly lessens the capacity of gastric acid secretion and ulcer ratio with respect to control group (Nigam *et al.*, 2011).

Cleome viscosa is herbaceous and belongs to Brassicaceae and the leaves, seeds and roots decoction and infusion are used in ulcer management (Arshad *et al.*, 2011, Hussain *et al.*, 2010, Nisar *et al.*, 2011, Weiss *et al.*, 2014). The 70 % ethanolic extract of the aerial parts was investigated in various stomach ulcer models in rats. The *Cleome viscosa* extract showed significant inhibition ($p < 0.01$) lesion ratio in ethanol (15.93–42.30 percent), PL (26.34–59.28 percent) and (CRS 22.58–54.03 percent), correspondingly and inhibit the oxidative injuries of gastric mucosus membrane by preventing peroxidation of lipids and by a remarkably decrease ($p < 0.001$) in SOD, and an increment in catalase activity. A prominent decrease ($p < 0.01$) happened in the amount

of H⁺K⁺ATPase, the degree of stomach juice and overall increase in pH (Gupta *et al.*, 2013).

Heliotropium indicum is herbaceous and belongs to Boraginaceae. The leaves extract is used against ulcer (Taj *et al.*, 2009). The ethanolic withdraw of *Heliotropium indicum* exhibited antiulcer activity against pylorus ligated induced gastric ulcer in rats, HCl- Ethanol induced ulcer in mice and water immersion stress induced ulcer in rats that indicates that *Heliotropium Indicum* leaves extract to have possible anti-ulcer action in the 3 models tested (Ashoka *et al.*, 2011).

Hippophae rhamnoides is herb and belongs to Asteraceae. The whole plant decoction is used to cure ulcer (Shedayi, *et al.*, 2012). It significantly lessens ulcer development in H₂O-immersion ($P < 0.05$) and reserpine-induced ($P < 0.01$) models in rats and it is recommended that the CO₂-extracted sea buckthorn seed and pulp oils have both preventive and curative effects against gastric ulcers in the study in rats (Xing *et al.*, 2002).

Linum usitatissimum is an herb and belongs to Linaceae. The seed, bark, leaves, flower and oil decoction are used to treat ulcer (Iqbal *et al.*, 2011). In a rat model of ethanol-induced gastric ulcer, the oil and mucus were evaluated, and they dramatically reduced the number and length of ethanol-induced gastric ulcers. An oral dose of flax seed oil (5 ml/kg) reduced ulcer severity more than ranitidine (50 mg/kg), showing that both flax (seed oil) and flax (seed mucus) may protect rats from ethanol-induced stomach ulcers. (Dugani *et al.*, 2008).

Momordica charantia is an herb and belongs to Cucurbitaceae. The fruits are used as a vegetable and are used to cure ulcer (Khan *et al.*, 2013b). In pylorus-ligated rats were given a methanol extract of *Momordica charantia* fruits to explore duodenal and gastric duodenal ulcers which showed control in the ulcer index and an elevation in gastric mucosa content. In ethanol-induced, stress-induced, indomethacin-induced stomach ulcers, and cysteamine-induced duodenal ulcers, the extract reduces the ulcer ratio. (Alamed *et al.*, 2009).

Plantago lanceolata is an herb and belongs to Plantaginaceae. The whole plant decoction is used to treat ulcer (Awan *et al.*, 2013, Sultana *et al.*, 2006). The aqueous extract of *Plantago lanceolata* demonstrated that higher doses offer better safety against gastroduodenal ulcers than the normal drugs implemented through antisecretory and cytoprotective mechanisms (Endale *et al.*, 2011).

Portulaca oleracea is herbaceous and belongs to Portulacaceae. The whole plant is used as a vegetable and is useful in ulcer (Wariss *et al.*, 2014). The ethanol and aqueous extracts were investigated in mice and both extracts exhibited a dose-related depletion in severity of ulcers. The retroperitoneal and oral management of extracts lessens the gastric increase in pH in pylorus-ligated mice and this suggests that *Portulaca oleracea* has gastroprotective action and validates its utilization in traditional medicine for stomach and intestine diseases (Gholamreza *et al.*, 2004).

Solanum nigrum is an annual herb and belongs to Solanaceae. The aerial parts and whole sap are used to treat ulcer (Sultana *et al.*, 2006, Wariss *et al.*, 2014, Gulshan *et al.*, 2012). The anti-ulcer effect of *Solanum nigrum* fruits extract on cryopreservation stress, indomethacin, pyloric ligation, and ethanol-induced gastric ulcer models, as well as ulcer healing activity on acetic acid-induced ulcer model in rats, significantly inhibited the CRU induced gastric lesions (76.6 percent), IND (73.8 percent), PL (80.1 percent), and EtOH (70.6 percent), with greater potential than omeprazole. These results reveal that *Solanum nigrum* extract has antiulcerogenic and ulcer-healing capabilities, which could be attributed to its antisecretory action (Mallika *et al.*, 2006).

Tephrosia purpurea is a shrub and belongs to Fabaceae. The whole plant powder and decoction is used locally for ulcer treatment (Ahmad *et al.*, 2014a). When compared to control vehicle animals, the ulcer ratio in *Tephrosia purpurea* treated animals was much lower in all models, and this antiulcer action was even more obvious in animals with ulcers produced by HCl, indomethacin, or pyloric ligation. Comparative to the control group, omeprazole (8 mg/kg¹) provided considerable stomach and duodenal ulcer prevention. (Deshpande *et al.*, 2003).

Withania somnifera is shrubby and belongs to Solanaceae. The juice, powder, poultice, and paste of the whole plant is used against ulcer (Muslim *et al.*, 2010, Arshad *et al.*, 2011, Kamal *et al.*, 2009, Hussain *et al.*, 2010, Nisar *et al.*, 2011, M. Hamayun, 2007, Zereen *et al.*, 2013). Preventer ulcer activities of uproot of *Withania somnifera* and its measure against stress plus posterior ligated induced peptic ulcer in rats has minimize ulcer index as compared to control group.

Moreover, *Withania somnifera* have the property to control the release of hydrochloric acid in the stomach, however, it boosts body's defense mechanisms, such as

the antioxidant defense system to avoid gastric mucosal damage. (Bhatnagar *et al.*, 2005).

CONCLUSION

Herbal plants are best choice for the treatment of ulcer available in large quantity as compared to other synthetic drugs. In Pakistan, treatment of ulcer is done with various local plant remedies. However, in most cases, the ethnomedicinal research reports gave important information about the recipes preparation method, amount of dose, and time of application of remedies. Future pharmacological investigation shows the majority of the plants with various positive and therapeutic properties. In conclusion, the viable utilization and alteration of primitive knowledge will be useful and enhance the lifestyle of deprived communities.

CONFLICT OF INTEREST

The authors declares that they have no conflict of interest.

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