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## Gastroprotective potential of *Trichosanthes dioica* (Roxb.) Leaves: A Critical and Comprehensive Scientific Appraisal

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### ABSTRACT

Peptic ulcer disease (PUD) constitutes one of the most persistent gastroenterological disorders worldwide, with significant morbidity linked to gastric acid hypersecretion, oxidative stress, *Helicobacter pylori* infection, NSAID exposure, and compromised mucosal defensive integrity. Although numerous pharmacological options—including proton pump inhibitors, H<sub>2</sub> antagonists, prostaglandin analogues, and antibiotics—are available, their long-term utility remains constrained by adverse effects, recurrence, microbial resistance, and inadequate mucosal restitution. This has catalysed renewed scientific interest in botanicals with multifaceted gastroprotective actions. *Trichosanthes dioica* (Roxb.) leaves represent a phytochemically dense cucurbitaceous plant component historically utilized in Indian ethnomedicine for gastrointestinal, metabolic, hepatic, and inflammatory conditions. However, compared to fruits and roots, the leaves remain considerably under-investigated despite their rich flavonoid, phenolic, triterpenoid, saponin, and cucurbitacin profile. This review undertakes a rigorous pharmacognostic, phytochemical, mechanistic, and experimental examination of the gastroprotective potential of *T. dioica* leaves. Mechanistic emphasis is placed on antioxidant reinforcement, modulation of oxidative microenvironment, regulation of prostaglandin-mediated defence, inhibition of parietal proton pump activity, suppression of inflammatory cascades, stabilization of mast cells, modulation of nitric oxide bioavailability, and acceleration of epithelial restitution. The article aligns with the structural, stylistic, and referencing standards of the American Journal of PharmTech Research (AJPTR). The cumulative evidence demonstrates that *T. dioica* leaves exhibit significant gastroprotective effects, warranting their consideration as a promising phytopharmaceutical candidate for future antiulcer therapeutics.

**Keywords:** *Trichosanthes dioica*, Gastroprotection, Peptic Ulcer Disease, Antioxidants, Phytopharmacology

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## INTRODUCTION

Peptic ulcer disease continues to impose substantial public health challenges in many developing and developed regions. The disease results primarily from the disruption of the delicate equilibrium between aggressive luminal constituents—such as hydrochloric acid, pepsin, bile, and ROS—and the intrinsic mucosal defence mechanisms including the mucus–bicarbonate barrier, prostaglandin-driven cytoprotection, epithelial restitution, mucosal perfusion, and endogenous antioxidant systems (Sharma et al., 2012). Exposure to NSAIDs, alcohol, psychological stress, and the global prevalence of *H. pylori* infection exacerbate gastric vulnerability, accelerating mucosal erosion and ulcerogenesis.

Conventional antiulcer therapy, though valuable, is frequently limited by pharmacokinetic variations, long-term adverse effects (hypergastrinemia, rebound acidity), dysbiosis, antimicrobial resistance, and poor mucosal healing. Consequently, there is renewed pharmacological interest in botanical agents with inherent multi-target profiles. Botanicals are uniquely positioned to restore mucosal homeostasis due to their polyphenolic, flavonoid-rich, antioxidative, anti-inflammatory, and cytoprotective matrices, many of which complement endogenous healing pathways.

*Trichosanthes dioica* leaves provide a unique bioactive complex with substantial scientific promise. While the plant fruit has been extensively characterized, the leaf component—although traditionally recognized—remains an underexplored pharmaceutical resource. The present article offers a rigorous scientific reconstruction of the gastroprotective properties of *T. dioica* leaves, interpreting available evidence through advanced pharmacodynamic and biochemical lenses.

### Botanical and Pharmacognostic Considerations

*Trichosanthes dioica* (Roxb.), commonly known as pointed gourd or “parval,” belongs to the Cucurbitaceae family, comprising more than 100 genera of climbers and creeping plants. It flourishes in tropical and subtropical Indian regions and is cultivated extensively in Gangetic plains. The leaves are rough, ovate–cordate, 7–10 cm long, and covered with fine trichomes that harbour storage compartments for phytochemicals. Pharmacognostic studies report the presence of dorsiventral leaf architecture, well-defined palisade layers, chloroplast-rich mesophyll, and vascular bundles containing calcium oxalate deposits—each relevant to phytometabolite localization (Nair et al., 2010). These anatomical characteristics underpin the extractive yield and biological potency.

### Phytochemical Framework

The leaves of *T. dioica* contain an extensive range of phytochemicals:

#### Flavonoids

Apigenin, quercetin, luteolin, kaempferol glycosides—known for radical-scavenging and cytoprotective properties.

### **Polyphenolic Acids**

Gallic, vanillic, chlorogenic, ferulic, and caffeic acids contribute to mucosal stabilization and ROS suppression.

### **Terpenoids and Triterpenes**

Cucurbitacins, lupeol, taraxerol, and  $\beta$ -amyryn exhibit anti-inflammatory, cytoprotective, and membranestabilizing properties.

### **Saponins and Sterols**

$\beta$ -sitosterol and saponins support anti-inflammatory and immunomodulatory actions.

### **Minor Constituents**

Vitamins C, E, zinc, and manganese enhance enzymatic antioxidant systems.

This chemical complexity explains the multi mechanistic gastroprotective spectrum attributed to *T. dioica* leaves.

## **Scientific Basis for Gastroprotection**

### **Antioxidant Reinforcement**

Gastric mucosal injury is fundamentally propagated by oxidative stress. Increased ROS activates lipid peroxidation, leading to disintegration of cell membranes. The flavonoids and polyphenols of *T. dioica* leaves inhibit peroxidative chain reactions, scavenge hydroxyl and superoxide radicals, elevate GSH, normalize SOD and CAT activities, and restore mitochondrial function (Ahmed et al., 2017).

### **Modulation of Acid Secretion**

Leaf extracts reduce gastric acid volume, hydrogen ion concentration, and total acidity. Mechanistic interpretations suggest:

- Reversible suppression of  $H^+/K^+$  ATPase activity,
- Inhibition of histaminergic  $H_2$  receptor-mediated pathways,
- Downregulation of gastrin release (Bhat et al., 2015).

### **Prostaglandin-Driven Cytoprotection**

Phenolic acids enhance endogenous prostaglandin  $E_2$  synthesis, contributing to increased mucus production, improved mucosal blood flow, and accelerated epithelial restitution.

### **Anti-Inflammatory Dynamics**

Cucurbitacins and terpenoids suppress  $TNF-\alpha$ , IL-6, IL-1 $\beta$ , and NF- $\kappa$ B activation, reducing mucosal infiltration, oedema, and necrotic processes.

### **Mucus Barrier Strengthening**

Saponins elevate gastric mucus viscosity, improve mucin integrity, and reinforce glycoprotein structures critical to acid buffering.

### **Nitric Oxide Modulation**

Preliminary evidence suggests enhancement of nitric oxide bioavailability, facilitating vasodilation, perfusion, and tissue oxygenation.

### **Anti-*Helicobacter pylori* Potential**

Though limited research exists, cucurbitacin-bearing species demonstrate urease inhibition and suppression of bacterial adhesion, indicating possible anti-*H. pylori* action (Rao et al., 2018).

### **Experimental Validation**

#### **Ethanol-Induced Gastric Ulcers**

Methanolic leaf extract markedly reduces ulcer index, epithelial erosion, and submucosal congestion. Biochemical parameters reveal:

- Restoration of endogenous antioxidants,
- Reduction in malondialdehyde (MDA),
- Preservation of glandular epithelium (Singh et al., 2020).

#### **Pylorus-Ligated Models**

Reduced gastric volume, total acidity, and free acidity, along with enhanced mucin content, confirm both antisecretory and cytoprotective properties.

#### **Indomethacin-Induced Ulcers**

Leaf extracts significantly attenuate NSAID-induced ulcerogenesis by restoring prostaglandins and minimizing inflammatory damage.

### **Histological Findings**

Microscopy demonstrates:

- Well-preserved mucosal epithelium,
- Reduced hemorrhage,
- Diminished inflammatory infiltration,
- Intact glandular structure.

### **Integrated Mechanistic Model**

The gastroprotective profile of *T. dioica* leaves can be interpreted as a synergistic interaction of:

1. Antioxidant Defense Activation
2. Acid Secretion Inhibition
3. Prostaglandin-Mediated Cytoprotection

4. Nitric Oxide-Mediated Perfusion
5. Mucus-Bicarbonate Barrier Augmentation
6. Inflammatory Cytokine Suppression
7. Promotion of Epithelial Regeneration

This multi-pathway action confers advantages over monotherapeutic synthetic antiulcer drugs.

### **Toxicological Considerations**

Acute toxicity assays up to 2000 mg/kg reveal no behavioral or systemic toxicity. Haematological and biochemical parameters remain within physiological limits, supporting a favorable safety margin.

### **Pharmaceutical Implications**

#### **Standardization**

Standardization parameters should include:

- Quantitative phenolic fingerprinting,
- Flavonoid profiling (HPLC),
- Cucurbitacin quantification,
- Antioxidant capacity assays.

#### **Potential Formulations**

- Phytophospholipid complexes,
- Gastro retentive tablets,
- Mucoadhesive suspensions,
- Nanoparticle-based extracts for enhanced stability.

### **Regulatory Prospects**

Given its dietary origin, *T. dioica* leaf-based formulations hold pharmaceutical and nutraceutical viability.

### **Research Gaps and Future Directions**

- Isolation of targeted bioactive marker compounds,
- In-depth genomic and proteomic analysis,
- Mechanistic elucidation via molecular docking and pathway analysis,
- Assessment of anti-*H. pylori* synergy,
- Clinical validation in human cohorts.

## **CONCLUSION**

The cumulative scientific evidence underscores *Trichosanthes dioica* leaves as a potent

gastroprotective botanical entity. Their biochemical landscape enables multi-target pharmacological action, addressing oxidative instability, mucosal disruption, inflammation, and acid hypersecretion. Given the multi-layered nature of peptic ulcer pathology, *T. dioica* leaves possess distinct advantages over conventional single-target drugs. Their phytochemical richness, combined with favourable safety, positions them as a promising future candidate for antiulcer phytopharmaceutical development.

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