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## **THERAPEUTIC POTENTIAL OF ANTI-INFLAMMATORY MEDICINAL PLANTS: EVIDENCE FROM PHYTOCHEMISTRY AND EXPERIMENTAL STUDIES**

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### **ABSTRACT**

Inflammation is a protective biological response that maintains tissue integrity following injury, infection or exposure to harmful stimuli. However, dysregulated or persistent inflammation contributes to the development of chronic diseases such as arthritis, diabetes, cardiovascular disorders and neurodegenerative conditions. Synthetic drugs including NSAIDs, COX-2 inhibitors, corticosteroids, DMARDs and biologics remain the primary therapeutic options, but their long-term use is often limited by gastrointestinal injury, renal toxicity, immunosuppression and cardiovascular complications. This has encouraged a growing interest in medicinal plants as safer, multi-targeted alternatives for managing inflammatory disorders. This review systematically highlights a wide range of medicinal plants traditionally used in inflammation, along with their extraction types, phytochemical profiles and mechanisms of action. Many of these plants contain flavonoids, alkaloids, terpenoids, saponins, phenols and essential oils that exert anti-inflammatory effects by inhibiting COX/LOX enzymes, suppressing NF- $\kappa$ B activation, modulating cytokines such as TNF- $\alpha$ , IL-1 $\beta$  and IL-6, and reducing oxidative stress. Experimental evidence from various in-vivo models—including carrageenan-induced paw edema, FCA-induced arthritis, LPS-stimulated macrophages and granuloma assays—supports their pharmacological potential. Overall, medicinal plants represent promising therapeutic resources with significant anti-

inflammatory activities. Further research on standardization, toxicity profiling and clinical validation is required to facilitate their safe integration into modern medicine.

## INTRODUCTION

Inflammation is a complex biological response of the immune system and serves as an essential defence mechanism that evolved to protect living organisms and maintain tissue homeostasis<sup>1</sup>. This protective reaction is activated when normal physiological balance is disrupted by injury, infection, toxins or other harmful stimuli<sup>2</sup>. The classic clinical signs of acute inflammation redness, heat, swelling and pain have been recognized since the first century A.D. and reflect the vascular and cellular changes that occur at the site of tissue damage<sup>3</sup>.

Inflammation is broadly classified into acute and chronic types. Acute inflammation represents the immediate and innate response aimed at the removal of harmful agents and restoration of normal function<sup>4</sup>. However, when this response becomes uncontrolled or persists over time, it transitions into chronic inflammation, a prolonged pathological state characterized by continuous tissue destruction, dysregulated immune activity and impaired healing<sup>5</sup>. Chronic inflammation is now well established as a major contributing factor to a wide range of non-communicable diseases, including cancer, diabetes, cardiovascular disorders, arthritis and neurodegenerative diseases<sup>6</sup>. Globally, chronic inflammatory disorders account for nearly 60% of total deaths, highlighting their immense public health burden<sup>7</sup>.

At the molecular level, inflammation is orchestrated by a complex network of mediators, including prostaglandins, leukotrienes, cytokines, chemokines and transcription factors<sup>8</sup>. The arachidonic acid cascade plays a central role, regulated by enzymes such as cyclooxygenase (COX) and lipoxygenases (LOX), which generate key inflammatory mediators<sup>9</sup>. The inducible isoform COX-2 is particularly associated with pain, fever and inflammatory tissue damage. Similarly, activation of transcription factors such as nuclear factor- $\kappa$ B (NF- $\kappa$ B) and pattern-recognition receptors like Toll-like receptors (TLRs) further amplifies inflammatory gene expression and propagates the inflammatory response<sup>10</sup>.

To manage inflammatory conditions, nonsteroidal anti-inflammatory drugs (NSAIDs) remain among the most widely used therapeutic agents<sup>11</sup>. However, prolonged NSAID use is associated with significant adverse effects, including gastrointestinal injury, renal impairment and cardiovascular risks, creating a need for safer and more effective alternatives<sup>12</sup>. In this

context, medicinal plants have gained considerable attention as potential therapeutic candidates. Many plant species contain bioactive phytochemicals with proven anti-inflammatory, antioxidant and immunomodulatory properties, often with fewer side effects compared to synthetic drugs<sup>13</sup>.

Given the global disease burden of inflammation and the limitations of current pharmacotherapy, exploring medicinal plants as natural anti-inflammatory agents is both scientifically valuable and clinically relevant. This review provides a comprehensive overview of medicinal plants traditionally used for inflammatory conditions, their phytochemical constituents and the molecular mechanisms through which they exert anti-inflammatory effects.

## **SYNTHETIC DRUGS**

### **1. Nonsteroidal Anti-Inflammatory Drugs (NSAIDs).<sup>14,15</sup>**

Nonsteroidal anti-inflammatory drugs are the most commonly used synthetic agents for managing inflammation, pain and fever. They act primarily by inhibiting cyclooxygenase enzymes (COX-1 and COX-2), which are responsible for converting arachidonic acid into prostaglandins—key mediators of inflammation. Drugs such as ibuprofen, diclofenac, naproxen and aspirin reduce swelling, redness and pain by decreasing prostaglandin levels at the site of injury. NSAIDs are effective in acute conditions like sprains, headaches and minor injuries, as well as chronic inflammatory disorders such as arthritis. However, long-term use is limited by side effects including gastric irritation, peptic ulceration, kidney damage and increased cardiovascular risk. Despite their limitations, NSAIDs remain the cornerstone of anti-inflammatory therapy due to their strong efficacy and easily accessible dosage forms.

### **2. Selective COX-2 Inhibitors (Coxibs).<sup>16,17</sup>**

Selective COX-2 inhibitors were developed to address the gastric toxicity associated with traditional NSAIDs. Drugs such as celecoxib and etoricoxib selectively inhibit the inducible COX-2 enzyme which is elevated during inflammation while sparing COX-1 which protects the stomach lining. This selective action provides powerful anti-inflammatory and analgesic effects with fewer gastrointestinal side effects. Coxibs are particularly useful in chronic inflammatory conditions such as osteoarthritis and rheumatoid arthritis. However, their prolonged use may increase the risk of cardiovascular complications such as hypertension, stroke and myocardial infarction. Therefore, despite their therapeutic benefits, selective

COX-2 inhibitors must be prescribed with caution, especially in patients with underlying heart disease.

### 3. Corticosteroids.<sup>18,19</sup>

Corticosteroids are among the most potent synthetic anti-inflammatory agents and are used when rapid suppression of inflammation is required. Drugs such as prednisolone, dexamethasone, hydrocortisone and betamethasone mimic the natural glucocorticoids produced by the adrenal cortex. They inhibit phospholipase A2, thus blocking the entire arachidonic acid cascade and suppressing the formation of prostaglandins and leukotrienes. Additionally, corticosteroids downregulate multiple inflammatory cytokines such as TNF- $\alpha$ , IL-1 and IL-6, leading to strong immunosuppressive effects. While they are extremely effective for severe inflammation, autoimmune diseases and allergic reactions, long-term use can cause serious side effects including weight gain, immunosuppression, diabetes, osteoporosis, mood disturbances and adrenal suppression. Therefore, corticosteroids are generally used for short-term or controlled therapy.

### 4. Disease-Modifying Anti-Rheumatic Drugs (DMARDs).<sup>20,21</sup>

DMARDs are a class of synthetic drugs specifically designed to prevent disease progression in chronic inflammatory conditions such as rheumatoid arthritis, psoriatic arthritis and autoimmune disorders. Common DMARDs include methotrexate, leflunomide, sulfasalazine and hydroxychloroquine. Unlike NSAIDs, which only provide symptomatic relief, DMARDs act on the underlying immune processes by inhibiting lymphocyte proliferation, reducing cytokine production and modulating immune signaling pathways. Methotrexate, the most widely used DMARD, reduces joint damage by decreasing T-cell activation and pro-inflammatory cytokines. Although highly effective, DMARDs require regular monitoring because of potential adverse effects such as liver toxicity, bone marrow suppression and gastrointestinal disturbances. They are often used in combination with NSAIDs or corticosteroids to enhance therapeutic outcomes.

### 5. Biologic Anti-Inflammatory Agents.<sup>22,23,24</sup>

Biologic agents represent the most advanced category of anti-inflammatory therapies. These drugs are genetically engineered to specifically target key molecules involved in inflammation. Examples include anti-TNF- $\alpha$  antibodies (infliximab, adalimumab), IL-6 receptor blockers (tocilizumab) and IL-1 receptor antagonists (anakinra). By neutralizing specific cytokines that drive inflammation, biologics offer a level of precision that traditional

drugs cannot achieve. They are particularly effective in severe, refractory autoimmune conditions where conventional therapies have failed. However, biologics are expensive and require injection or infusion. They may also increase the risk of serious infections, including tuberculosis, due to their powerful immunosuppressive actions. Despite these limitations, biologics have revolutionized the management of chronic inflammatory diseases.

Although synthetic anti-inflammatory drugs are highly effective, their long-term use is often limited by adverse effects. This has increased the global interest in plant-based anti-inflammatory agents, which are believed to act through multi-target mechanisms with potentially fewer side effects. Comparing synthetic drugs with natural alternatives helps highlight the therapeutic gap and supports the scientific need to investigate medicinal plants as safer and holistic options for managing inflammation.

#### List of Medicinal Plants With Reported Anti-Inflammatory Activity.

Sl no	Plant Name (Scientific Name)	Common Name	Extraction Used	Phytoconstituents Responsible for Activity	Method Used for Induction / Assay	Reference
1	<i>Achillea millefolium</i>	Yarrow	Oil extracts, Ethanolic extract	Sesquiterpenes, Flavonoids, Alkaloids, Salicylic acid	Tested on artificially irritated skin (human volunteers), Inhibition of polymorphonuclear leukocytes/oxidative stress	25
	<i>Adhatoda vasica</i>	Adosa or Malabar nut tree, Vasaka	Pyrroloquinazoline alkaloids (from roots), Methanolic leaf extract	Vasicine and Vasicinone (quinazoline alkaloids)	Carrageenan and Freund's Complete Adjuvant (FCA)-induced paw edema in rats, Collagen-induced arthritis (CIA) model	26
	<i>Aegle marmelos</i>	Bilwa	Aqueous extract (root bark)	- (Anti-inflammatory effect reported generally)	Carrageenan induced paw oedema model, Cotton pellet induced granuloma model	27

	<i>Agrimonia eupatoria</i>	Common Agrimony	Infusion, Polyphenol-enriched fraction	Polyphenols (isoquercetin, tiliroside, kaempferol O-acetyl-hexosyl-O-rhamnoside)	Carrageenan-induced paw oedema, Inhibition of inflammatory cytokines (IL-1, IL-4, IL-6)	28
	<i>Agropyrum repens</i>	Couch Grass, Pirevina	Ethanol extract (rhizomes), Dry couch grass extract (in cream)	Carbohydrates, mucilages, saponins, essential oils	Carrageenan foot oedema (rat hind-paw), Tested for allergic contact dermatitis in rats	29
	<i>Albizia lebbeck</i>	Shirisha or Siris	Bark extract (petroleum ether/ethyl acetate/methanol mixture)	- (Anti-inflammatory reported generally)	Carrageenan induced rat paw edema model	30
	<i>Allium cepa</i>	Onion	Aqueous extract/Poultices (traditional use)	Quercetin, Thiosulfinates, Cepaenes, Flavonoids	Inhibition of chemotaxis of human polymorphonuclear leukocytes, Inhibition of COX and LOX enzymes	31
	<i>Allium sativum</i>	Garlic	Garlic oil, Garlic extracts	Allicin, Thiocreatonone (sulfur compound)	Inhibition of cytoskeleton formation/disassembly,	32
	<i>Aloe ferox</i>	aloe	Extract (gel/resin)	Malic acid acylated polysaccharide	Anti-swelling (general mechanism)	33

				es, Aloe resin, Enzymes (carboxypeptidase, bradykinase)		
	<i>Alstonia scholaris</i>	Devils Tree or Dita Bark Tree	Alkaloids fraction of ethanolic leaf extract, Ethanolic extract (EEAS) leaves	Alkaloids, Flavonoids	Xylene-induced ear edema, Carrageenan-induced air pouch formation, FCA-induced arthritis in rats	34
	<i>Althaea officinalis</i>	Marshmallow, Beli Slez	Root extracts	Mucilage, Flavonoids, Caffeic acid, p-coumaric acid	induced cytotoxicity/ROS in macrophages, LPS-induced release of TNF-alpha and IL6	35
	<i>Anacardium occidentale</i>	Cashew or Kaju	Leaf extract	Oleamide	(Anti-inflammatory property determined but specific induction method is not detailed in the excerpt)	36
	<i>Asparagus racemosus</i>	Shatavari or Satavar	Ethanolic leaf extract, Hydroalcoholic roots extract	Steroidal saponins (Shatavarin I-X), Flavonoids (quercetin, rutin)	Carrageenan-induced paw edema in rats, FCA-induced arthritis in rats	37
	<i>Betula pendula</i>	Common Birch, Bela Breza	Aqueous extract, Dense	Flavonoids, Tannins, Resins,	In vitro study on human peripheral blood mononuclear cells (PBMC), Carrageenin-	38

			extract (tablets)	Essential oils	induced inflammation in rats	
	<i>Bryophyllum pinnatum</i>	Parnabeeja	Aqueous extract (plant leaf)	Flavonoids, Polyphenols	Fresh egg albumin-induced pedal (paw) oedema model	39
	<i>Calendula officinalis</i>	Calendula, Neven	Calendula oil (flower extract), Methanolic extract of flowers	Triterpenoids, Flavonoids	LPS-stimulated macrophages (NO inhibition), Carrageenan-induced paw oedema (implied in anti-oedematous effect)	40
	<i>Camelia sinensis</i>	Tea (Green/Black tea)	Aqueous green tea extract (GTE), Aqueous black tea extract (BTE)	Polyphenols (Catechins, Flavonoids), EGCG	Rat adjuvant-induced arthritis (AIA) model, FCA-induced arthritis rat model	41
	<i>Cassia fistula</i>	Golden shower tree	Bark extracts	Flavonoids and Bioflavonoids	Acute and chronic anti inflammatory models of inflammation in rats	42
	<i>Cassia occidentalis</i>	Coffee senna	Ethanolic extract (whole plant)	(Anti-inflammatory activity reported generally)	Carrageenan-induced paw oedema model	43
	<i>Cedrus deodara</i>	Himalayan Cedar or Deodar	Wood oil, Aqueous stem bark extract	Essential oils, Flavonoids, Sesquiterpenes (himachalenes)	Carrageenan-induced paw edema, FCA-induced arthritis, Croton oil-induced granuloma pouch	44



	<i>Cissampelos sympodialis</i>	-	Alkaloids total fraction and ethanolic extract (aerial parts)	Alkaloids (implied)	Reduced tumor necrosis factor and interleukin-1 (IL-1) levels	45
	<i>Citrus limetta</i>	Mosambi / Mousambi	Essential oils (EOs)	Limonene (monoterpene hydrocarbon)	Lipopolysaccharide-induced Inflammation (macrophages), 12-O-tetradecanoylphorbol- 13acetate-induced ear inflammation (in vivo)	46
	<i>Citrus limon</i>	Nimbu and Lemon	Essential oils (EOs)	(Anti- inflammatory activity reported generally)	Reduced the sum of writhes and paw licking	47
	<i>Cuminum cuminum</i>	Cumin	Volatile oil, Essential oils (EOs)	(Anti- inflammatory activity reported generally)	Carrageenan-induced Rat-paw oedema, LPS-stimulated RAW 264.7 cells (inhibited COX-2, IL-1, IL-6)	48
	<i>Curcuma longa</i>	Turmeric or Haldi	Oil-free aqueous extract (COFAE) of rhizome, Polar extract (PCL) of rhizome	Curcuminoids (curcumin, demethoxycur cumin), Volatile oils	Xylene induced ear edema/Cotton pellet granuloma, Monosodium Iodoacetate induced osteoarthritis in rats	49
	<i>Cynodon</i>	Bermuda grass	Aqueous	(Anti-	Carrageenan, serotonin,	50

	<i>dactylon</i>		extract	inflammatory activity reported generally)	histamine, dextran induced rat paw edema, cotton pellet method	
	<i>Cyperus rotundus</i>	Nutgrass	Essential oils (EOs)	(Anti-inflammatory activity reported generally)	Carrageenan injection (reduced paw edema)	51
	<i>Dendropanax morbifera</i>	Korean Dendropanax Gold-Leaf Tree	Methanolic extracts	Phenolic compounds (quercetin, rutin, resveratrol, chlorogenic acid)	Inhibited LPS induced pro inflammatory cytokines/mediators	52
	<i>Emblica officinalis</i>	Amla / Indian gooseberry	Water fraction of methanol extract (leaves)	(Anti-inflammatory reported generally)	Tested on the synthesis of leukotriene $B_4$ , PAF, and thromboxane	53
	<i>Glycyrrhiza glabra</i>	Liquorice, Radix dulcis	Hydroalcoholic extract, Methanol extract	Glycyrrhizin, Glycyrrhizic acid.	Carrageenan-induced paw oedema, COX-2 inhibitory screening assay, Reduced antigen induced arthritis symptoms in mice	54
	<i>Hibiscus rosa-sinensis L. / H. rosa-sinensis var alba</i>	Red Hibiscus / White Hibiscus	Ethanol extracts (flower and leaf)	Flavonoids, Saponins, Steroids	Carrageenan-induced Paw Edema, PNL Infiltration study, Formalin-induced Paw Licking Test	55
	<i>Hypericum</i>	St. John's Wort,	Lipophilic	Hyperforin,	Antiedematogenic/antinocicept	56

	<i>perforatum</i>	Kantarion	extract, Hypericum oil extracts	Hypericin, Flavonoids, Tannins	ive properties in rats	
	<i>Ipomoea pescaprae</i>	Goat Foot Vine and Beach Morning Glory	Leaf extracts	- (Compounds inhibiting prostaglandin synthesis, implied)	Oedema caused by ethyl phenyl propiolate in animals, Treating dermatitis caused by jellyfish stings	57
	<i>Juglansregi a</i>	Walnut, Orah	Ethanolic extracts (leaves), Walnut septum extract	Tannins, Naphthoquino ne derivatives, Flavonoids	Carrageenan-induced hind paw edema model in mice, Citric acid-induced cough model in rats	58
	<i>Leonotis ocymifolia</i>	Granthiparni	80 percent methanolic leaf extract	- (Anti- inflammatory activity reported generally)	Reduced paw oedema by carrageenan induction, Slowed granuloma synthesis	59
	<i>Matricaria chamomilla</i>	Chamomile, Kamilica	Volatile essential oil, Aqueous extract	Essential oils, Sesquiterpene lactones, Coumarins	Carrageenan-induced pedal swelling in rats, Xylol-induced ear swelling in mice	60
	<i>Mentha piperita</i>	Peppermint, Nana	Essential oil and extract	Essential oils, Phenolics, Flavonoids	Croton oil-induced mouse ear oedema model, Inhibitory effect on NO and $\text{PGE}_2$ production	61
	<i>Moringa olifera</i>	shigru	Aqueous and ethanolic extract (stem bark)	- (Anti- inflammatory activity reported generally)	Carrageenan induced rat paw edema	62

	<i>Nicotiana tobacum</i>	Dhumrapatra or Tamraparna	Leaf extract	Alkaloids (nicotine, anabasine), Quercitrin, Scopoletin	Used as an anti-inflammatory (general mention)	63
	<i>Olea europaea</i>	Olive tree	Extra virgin olive oil	- (Anti-inflammatory activity reported generally)	Carrageenan-induced paw edoema in rats	64
	<i>Oroxylum indicum</i>	Sonapatha or Shyonaka	Ethanol extract (stem bark), Petroleum ether root bark extract	Flavonoids (baicalein, chrysin), Phenolic compounds	Xylene-induced ear edema, Formalin-induced paw edema in mice, Adjuvant-induced arthritis in rat model	65
	<i>Persicaria chinensis</i>	Chinese knotweed	Methanolic extract	- (Anti-inflammatory activity reported generally)	LPS-induced nitric oxide and $\text{PGE}_2$ in RAW264.7 macrophages	66
	<i>Phyllanthus acidus</i>	StarGooseberry and Harpharauri	Methanolic extract (aerial parts)	Flavonoids (kaempferol and quercetin)	Inhibited $\text{PGE}_2$ and NO production in LPS-treated RAW 264.7 cells	67
	<i>Piper nigrum</i>	Piper or Black Pepper	Hexane/ethanolic fruit extracts	Piperine (alkaloid)	Carrageenan-induced paw inflammation in rats, Carrageenan-induced acute paw pain and arthritis in rats	68
	<i>Salix alba</i>	White Willow, Bela Vrba	Methanolic and	Salicylic acid, Phenolic	Xylene-induced ear oedema, Carrageenan-induced paw	69

			aqueous extracts (barks)	glycosides, Flavonoids	oedema	
	<i>Salvia officinalis</i>	Sage, Žalfija	Chloroform extracts	Flavonoids, Triterpenes (ursolic acid, carnosol)	Croton oil-induced ear oedema in mice, Formalin-induced inflammatory phase	70
	<i>Sida cordifolia</i>	Malva-branca	Aqueous extract	- (Anti-inflammatory activity reported generally)	Anti-inflammatory and analgesic activity evaluation (general test methods)	71
	<i>Solanum melongena</i>	Eggplant	Aqueous extract (leaves)	Ascorbic acid, Alanine, Arginine, Caffeic acid	Anti inflammatory activity (general assessment)	72
	<i>Swertia chirayita</i>	Chiretta or Chiratitka	Ethanollic extract (root), Aqueous stem extract	Xanthones, Terpenoids, Flavonoids, Glycosides	Carrageenan-induced paw edema in the rat model, Adjuvant-induced arthritis in mice	73
	<i>Syzygium caryophyllatum</i>	South Indian Plum and Dwarf Black Plum	Aqueous root extract	- (Anti-inflammatory activity reported generally)	Heat-induced egg albumin denaturation bio assay technique (in-vitro)	74
	<i>Tephrosia purpurea</i>	Wild Indigo	50 percent alcoholic extract (root)	- (Anti-inflammatory activity reported generally)	Carrageenan and produced paw oedema method	75
	<i>Vitex</i>	Chaste Tree or	Aqueous	Lignans,	Carrageenan-induced paw	76

	<i>negundo</i>	Nirgundi	leaves extract, Ethanolic seeds extract (total lignans)	Flavonoids, Essential oils	edema in rats, Collagen-induced arthritis in rats	
	<i>Withania somnifera</i>	Ashwagandha or Indian Ginseng	Methanolic leaves extract, Aqueous roots extract (WSAq)	Steroidal withanolides, Alkaloids, Polyphenols	Stainless steel-induced inflammation in adult zebra fish, Collagen-induced arthritis in rats	77
	<i>Zingiber officinale</i>	Ginger, Red ginger	40% ethanolic extract (dried red ginger), Aqueous rhizome extract	Essential oil, Polyphenolic compounds, Gingerol	Acute and chronic inflammation models, Carrageenan-induced paw edema in rats, Collagen-induced arthritis in mice	78

## CONCLUSION

This review demonstrates that medicinal plants possess significant anti-inflammatory potential supported by strong phytochemical composition and mechanistic evidence. Their bioactive constituents—such as flavonoids, terpenoids, alkaloids, and phenolic compounds—act through multiple pathways, including inhibition of COX/LOX enzymes, suppression of NF-κB signaling, modulation of pro-inflammatory cytokines and enhancement of antioxidant defenses. While synthetic anti-inflammatory drugs remain effective, their long-term adverse effects highlight the need for safer, natural alternatives. The findings of this review clearly indicate that medicinal plants offer multi-targeted, accessible and biologically safe therapeutic options for managing both acute and chronic inflammation. However, despite the promising preclinical evidence, further research involving standardization, bioavailability

studies, toxicity assessments and well-designed clinical trials is essential to ensure their effective translation into clinical practice.

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