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p-coumaric acid: A review of its chemistry, biological activities and therapeutic potential

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Abstract

p-Coumaric acid (p-CA) is a natural phenolic compound widely distributed in plants, fruits, vegetables, and cereals. It is recognized for its broad spectrum of biological activities, including antioxidant, antimicrobial, anti-inflammatory, and anticancer effects. As a hydroxycinnamic acid derivative, p-CA plays an important role in protecting cells against oxidative stress and modulating various biochemical pathways. This review provides a detailed overview of its chemistry, biosynthesis, dietary sources, mechanisms of biological activity, therapeutic potentials, and recent advances in formulation and biomedical applications. The article also discusses limitations, bioavailability challenges, and future perspectives for developing p-CA-based nutraceutical and pharmaceutical formulations.

Keywords: *p*-Coumaric acid; hydroxycinnamic acid; antioxidant; antimicrobial; anticancer; phenolic compounds

1. Introduction

Phenolic compounds are secondary metabolites synthesized by plants to protect against ultraviolet radiation, pathogens, and oxidative damage. These compounds have attracted considerable scientific interest due to their potential health-promoting effects in humans [1-6]. Among phenolic acids, hydroxycinnamic acids such as caffeic, ferulic, and *p*-coumaric acids are particularly significant because of their antioxidant and therapeutic properties [7]. *p*-Coumaric acid (4-hydroxycinnamic acid) (Figure 1) is one of the most abundant and biologically active phenolic acids in nature, widely found in edible plants, grains, fruits, and vegetables [2-4].

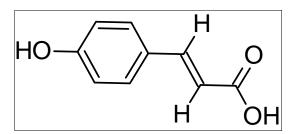


Fig 1: Chemical structure of p-Coumaric acid.

The compound's simple aromatic structure, consisting of a hydroxyl group attached to the para position of the phenyl ring, enables it to participate in redox reactions and neutralize free radicals. Over the past decade, numerous studies have demonstrated its antioxidant, antimicrobial, anti-inflammatory,

anticancer, and anti-aging effects [8-11]. This review provides an integrated understanding of the chemistry, biosynthesis, biological activities, and therapeutic applications of p-coumaric acid, with emphasis on its biomedical significance.

2. Chemical Nature, Biosynthesis and Dietary Sources

Chemically, p-coumaric acid (p-CA) belongs to the hydroxycinnamic acid family derived from the phenylpropanoid pathway in plants [12]. It originates from phenylalanine through enzymatic deamination and hydroxylation reactions that yield cinnamic acid and subsequently p-coumaric acid [10]. The molecule possesses a phenolic hydroxyl group and an α, βunsaturated carboxylic acid chain, both of which contribute to its antioxidant activity [13]. p-CA is commonly present in cereals such as wheat, corn, and barley, especially in the bran layers, as well as in fruits, peanuts, tomatoes, garlic, and grapes [4]. It exists in both free and bound forms, the latter often conjugated with sugars, esters, or polymers. The presence of p-CA in a wide variety of dietary sources makes it a major contributor to the total polyphenol intake in humans [8].

3. Antioxidant Activity

The antioxidant mechanism of p-coumaric acid involves hydrogen donation, free radical scavenging, and modulation of antioxidant enzymes [12]. Experimental studies have shown that p-CA

neutralizes reactive oxygen species (ROS) and protects cellular components from oxidative damage. In endothelial cells exposed to high glucose, p-CA reduces oxidative stress and lipid peroxidation [13-16]. Similarly, in keratinocytes subjected to ultraviolet (UV) radiation, p-CA attenuates oxidative damage and enhances cellular viability [11]. Table 1 depicts the antioxidant activities of p-Coumaric acid in different biological and food systems. These results suggest that p-CA could play a protective role in cardiovascular, metabolic, and skin-related disorders.

Table 1: Antioxidant activity of p-coumaric acid in various systems [2-4, 8, 11].

Experimental System	Observed Effect	Mechanism / Biomarker
Cereal extracts	Prevented lipid	Free radical
(in vitro)	oxidation in grains	scavenging
Lens epithelial cells	Reduced H ₂ O ₂ - induced oxidative stress	Upregulated catalase, SOD
Tomato peel	Enhanced antioxidant	Increased total
extract	activity	phenolic content
Human	Protected cells from	Decreased lipid
keratinocytes	UV damage	peroxidation
Food emulsions	Inhibited peroxidation in stored oils	Hydrogen atom donation

4. Antimicrobial and Antifungal Activity

p-Coumaric acid demonstrates strong antimicrobial and antifungal properties against a wide range of microorganisms. The mechanism involves disruption of bacterial membranes, leakage of cellular contents, and interference with DNA replication ^[4]. Studies have confirmed that p-CA is active against Grampositive and Gram-negative bacteria, including Staphylococcus aureus and Escherichia coli ^[5]. In addition, polymeric films containing p-CA exhibit improved wound healing by preventing microbial infection and promoting tissue regeneration ^[17-20]. Table 2 explains antimicrobial and antifungal activities of p-Coumaric acid and its derivatives.

Table 2: Antimicrobial and antifungal activities of *p*-Coumaric acid and derivatives [12, 15-18, 21-24].

Target Microorganism	Observed Effect	Mode of Action
E. coli and S.	Strong inhibition	Membrane
aureus	at 0.1 mg/mL	disruption
Candida albicans	Reduced fungal viability	ROS generation
Foodborne pathogens	Preserved meat quality	Cell wall permeability alteration
Mixed bacterial strains	Dose-dependent inhibition	Dual damage mechanism

5. Anticancer and Chemopreventive Effects

Several studies have demonstrated the anticancer activity of p-coumaric acid through multiple mechanisms such as inhibition of proliferation, induction of apoptosis, and cell cycle arrest $^{[4, 19-22]}$. In melanoma and breast cancer cell lines, p-CA suppresses tumor cell growth and migration $^{[6-10]}$.

Animal models of colon cancer have shown that *p*-CA decreases inflammatory responses and enhances antioxidant enzyme activity ^[25-27]. Furthermore, copper-assisted complexes of *p*-CA and its derivatives have been developed to improve anticancer efficiency ^[24-30]. Beyond its antioxidant and anticancer effects, *p*-coumaric acid exhibits anti-inflammatory, hepatoprotective, neuroprotective, and skin-protective actions ^[19-21]. It reduces pro-inflammatory cytokines, prevents oxidative damage in liver tissue, and improves neuronal antioxidant defenses ^[31-32]. In cosmetic formulations, *p*-CA serves as an antimelanogenic and UV-protective agent, reducing hyperpigmentation and photoaging ^[22, 32]

8. Conclusion and Future Prospects

Despite its promising biological activities, pcoumaric acid suffers from poor solubility and limited bioavailability. Encapsulation of p-CA in nanoparticles, liposomes, or biodegradable polymers enhances stability, absorption, and controlled release. Such delivery systems have shown improved antioxidant and anticancer performance both in vitro and in vivo. Although numerous in vitro and in vivo studies have established the therapeutic potential of p-coumaric acid, several challenges remain. The limited bioavailability, metabolic instability, and lack of human clinical data restrict its translation into pharmaceuticals. Future research should focus on enhancing delivery systems, conducting welldesigned clinical trials, and understanding long-term safety. Further studies on structure-activity relationships could lead to the design of more potent derivatives with improved therapeutic efficacy. p-Coumaric acid is a versatile natural compound with broad biological activities including antioxidant, antimicrobial, and anticancer effects. Its mechanisms of action involve modulation of oxidative stress, apoptosis, inflammation, and microbial membrane integrity. While its dietary abundance supports its safety, further pharmacological and clinical investigations are required to fully realize its potential as a nutraceutical and therapeutic agent.

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