

Plant-Derived Antifungal Agents and Fungal Pathogens Mechanism of Action, Therapeutic Applications, and Challenges

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Abstract:

The environmental hazards associated with synthetic fungicides and the development of antifungal resistance aggravate the already high frequency of fungal infections, therefore compromising human health, agricultural output, and food security. Plant-derived extracts provide a pleasing choice because of their higher bioactive concentration, different action mechanisms, and positive effects on the surroundings. This unequivocally shows the ways in which phenolics, alkaloids, terpenoids, and essential oils—among other phytochemicals present in plants—attack fungal membranes, reduce enzymatic function, cause oxidative stress, and eliminate biofilms. Food preservation, medicine, and agriculture all indicate how promising these compounds are as long-term antifungal treatments. Strong regulatory policies paired with innovative technologies such molecular profiling, nanotechnology, and cooperative research will help to fully actualise the effectiveness of antifungal medications developed from plants. This review presents a platform for subsequent investigations in this subject and indicates the importance of plant extracts in lowering the increased prevalence of fungal infections.

Keywords: Nanotechnology in Antifungal Therapy, Fungal Resistance, Bioactive Phytochemicals, and Plant-Derived Antifungals.

1-Introduction Section

The environmental hazards connected with synthetic fungicides and the development in antifungal resistance aggravate the already major problem of fungal infections. Fungi-caused infections compromise people's access to wholesome food, agricultural production, and general health. Plant-based extracts provide an intriguing choice. They are beneficial for the environment, contain many bioactive chemicals, and vary modes of action. By means of thorough investigation on the antifungal characteristics of plant

phytochemicals—including terpenoids, alkaloids, essential oils, and phenolics—we have discovered how these compounds assault fungal cell walls, suppress enzyme function, generate oxidative stress, and breakdown biofilms. Because they are used in food preservation, medicine, and agriculture, these extracts offer potential as long-acting antifungal therapies. Issues with toxicity, lack of standards, and scalability make them not yet economically feasible. Fulfilling the potential of plant-based antifungal medications depends on using modern techniques such as molecular profiling, synergistic research, nanotechnology, and rigorous regulatory frameworks. This study not only seeks to open doors for additional investigation in this sector but also underlines the need of plant extracts in the battle against the rising frequency of fungal illnesses. Along with the use of nanotechnology in treating fungal infections, some of the topics that will be explored include fungal resistance, bioactive phytochemicals, and antifungals derived from plants. Fungal infections compromise human health, agricultural output, and food safety, therefore posing a major worldwide issue. Various opportunistic fungal diseases including *Cryptococcus*, *Aspergillus*, and *Candida* have caused an alarming increase in death and morbidity, particularly among highly immunocompromised people. Among the fungal infections are those include invasive mycoses, oral candidatures, and dermatophytosis. The situation is becoming worse as antifungal resistance becomes more common. In agriculture, the fungal diseases *Alternaria*, *Botrytis*, and *Fusarium* provide challenges. severely reduced agricultural production, therefore compromising food supply security and economic stability as well. Both Cowan (1999) and Gyawali and Ibrahim (2014) underline that fungal contamination of food products poses a threat to the general public's health as well as the profits of businesses related to food production. Antifungal resistance is already somewhat concerning, and improper and excessive use of synthetic drugs and fungicides is reducing the efficacy of treatment. Conventional antifungals may, however, produce significant side effects including nephrotoxic and hepatotoxic responses, hence limiting their use to those especially sensitive. They are not always risk-free, either. Apart from the additional stress synthetic fungicides create on ecosystems, their wide usage in agricultural settings results in pollution of soil and water (Salari *et al.*, 2016). The absence of novel antifungal drugs to satisfy the growing demand for efficient treatment aggravates these issues. Globally, fungus-induced infections are a major source of concern. Their detrimental consequences on human health just scratch the surface; they also lower agricultural output and render food dangerous for consumption. Particularly in populations with compromised immune systems, opportunistic fungal infections including those brought on by *Cryptococcus*, *Aspergillus*, and *Candida* have been linked to a concerning increase in human illness and death. Common issues existing include mycoses, oral candidiasis, and dermatophytic infections; these often worsen when fungal infections acquire resistance to antifungal drugs. In agriculture, the fungal diseases *Alternaria*, *Botrytis*, and *Fusarium* provide a challenge. destroy crops to the extent that food supplies for the planet are under jeopardy. Food product fungus compromise not only consumer health but also the state of the economy.

The situation is worsened by many factors:

The misuse and abuse of synthetic antifungal drugs and fungicides has led to the emergence of resistant fungus strains, which in turn has lowered the efficacy of existing treatments. Nephrotoxicity and hepatotoxicity are two of the worst side effects of conventional antifungal medications, which is why they are seldom prescribed to vulnerable populations. Environmental Concerns: Soil and water contamination from the use of synthetic fungicides in farming has a negative impact on ecosystems. The development of new antifungal drugs has not kept pace with the increasing demand for them, which has limited both treatment and prevention choices. An appealing option in this context would be plant-derived extracts due to their potential decreased toxicity, range of bioactive compounds, and natural origins. These extracts may be beneficial in the fight against resistance development due to their multi-targeted action mechanisms, which have shown efficient against several fungal diseases. Alternatives that are less damaging to individuals and the environment, natural antifungals generated from plants have

various possible applications in healthcare, agriculture, and industry. The widespread use of plant extracts is hindered by issues such as the difficulty in standardising formulations for agricultural and medicinal purposes, the lack of knowledge on the toxicity profiles and mechanisms of action, and the heterogeneity in chemical composition caused by processing and environmental influences. This study explores the bioactive components, action mechanisms, and practical uses of plant extracts across several domains in order to fully harness their antifungal potential. Cowan (1999), Gyawali and Ibrahim (2014), and Salari *et al.* (2016) all note that it does more than only solve present-day problems; it also specifies future research goals. Here, plant extracts serve as a suitable alternative. The bioactive compounds present in plants have the potential to successfully combat various fungal diseases. Their natural origins and potential for reduced toxicity make them useful in medicine, agriculture, and industry. Also, antifungals made from plants may prevent resistance from developing as quickly due to their multi-targeted action. Although plant extracts have promise as antifungal treatments, they are not commonly utilised due to a variety of obstacles: Processing and environmental factors may cause changes in chemical composition. Their toxicity profiles and mechanisms of action are mostly unknown to us. Problems with consistency in formulations throughout pharmaceutical and agricultural uses. Antifungal potential of plant extracts is explored in this review, which also looks at their bioactive components, mechanisms of action, and medicinal, agricultural, and food preservation applications. Also included are suggestions for future research and development in this field, in addition to addressing current concerns. The study's overarching goal is to highlight the value of plant-based antifungals in the fight against fungal infections and resistance by providing a comprehensive overview of the subject.

Plant Chemicals with Antifungal Actions

Plant extracts may include a variety of secondary metabolites that have potent antifungal effects. Bioactive chemicals with distinctive anti-fungal effects include essential oils, terpenoids, saponins, alkaloids, and phenolic compounds. Their antifungal properties make them a promising option for the creation of natural antifungal treatments, which might help alleviate some of the side effects of synthetic antifungal drugs.

2.1.1. Hazardous Materials

Phenolic compounds including tannins, flavonoids, and phenolic acids have the ability to structurally disrupt fungal cell walls and membranes. These substances interfere with two important processes: the integrity of membranes and their permeability. The phenolic component eugenol, for instance, has been shown to inhibit the development of *Candida* spp. when isolated from *Syzygium aromaticum* (clove). by gaining access to the fungal cell wall and exposing its potent antifungal capabilities (Cowan, 1999).

2.2. Alkaloids

Alkaloids, particularly indole and isoquinoline derivatives, exert their antifungal effects by obstructing metabolic pathways specific to fungi. An alkaloid extracted from *Berberis* species, berberine is very effective against *Aspergillus* spp. because it limits their growth and reproduction. Gyawali and Ibrahim (2014) state that these compounds are crucial as they inhibit fungal enzymes and structural elements.

2.3.1. Terpenoids

Interactions with ergosterol are one known mechanism by which terpenoids like monoterpenes and sesquiterpenes may damage fungal cell membranes. Homeostasis and the integrity of cell membranes are both threatened by this interplay. Salari *et al.* (2016) found that thymol, a monoterpene extracted from *thymus vulgaris* (thyme), may fight against many types of fungal infections.

2.4.1. Saponins are made up of

The fungal cell wall is hydrolysed by saponins, which are amphipathic compounds. This process causes the release of internal contents and, eventually, cell death. The saponin dioscin, derived from *Dioscorea* spp., has shown encouraging activity in the fight against dermatophytes, a skin fungus infection.

2.5.1. All-Natural Essential Oils

Essential oils' antifungal properties are amplified by the synergistic combinations of volatile components. Since these oils attack several fungal sites, they are more effective and less prone to induce resistance to develop. Cowan (1999) and Salari *et al.* (2016) cite substantial study on the potential anti-trichomycosis effects of essential tea tree oil, which is extracted from *Melaleuca alternifolia*. So, it's a good option for dealing with dermatophytic infections. All of these phytochemicals work together to fight fungal infections, which might lead to new plant-based antifungal treatments for fields including agriculture, industry, and medicine. Optimisation for practical use and more research into their processes are necessary to fully achieve their potential.

Procedures

Plant extracts may inhibit fungal growth and survival by interfering with key metabolic pathways. Bioactive drugs like these are effective against a wide variety of fungal diseases, and they often use a battery of strategies to counteract resistance. Read on for an in-depth analysis of these techniques:

Cell Membrane Dysfunction

An important tactic in antifungal treatment is to prevent the fungal cell membrane production. Ergosterol, a sterol that interacts with several bioactive substances, is essential for the formation and function of fungal cell membranes. Contact destabilizes the membrane, increasing its permeability, which might lead to the leakage of important intracellular components. Cowan (1999) states that terpenoids such as thymol and carvacrol, which are present in *Thymus vulgaris* and *Origanum vulgare*, respectively, have the potential to degrade fungal cell walls, resulting in rapid cell lysis. The phenolic compound eugenol, which is present in cloves (*Syzygium aromaticum*), has a similar effect on *Candida* spp., inhibiting their growth and cytoplasmic leakage. according to Salari *et al.* (2016). by attacking their membranes.

3.2.1. Blocking Enzymes

Plant extracts that target key enzymes often target metabolic processes and the development of fungal cell walls. Chitin synthase and β -glucan synthase are enzymes that are essential for fungal cell wall chitin and glucan formation; substances like alkaloids and flavonoids hinder these enzymes. Inhibiting these enzymes leaves the fungal cell wall exposed, which leads to cell death. That is why *Berberis* spp. is the source of berberine. works wonders on many species of *aspergillus*. as it changes the process by which fungal cell walls are formed by inhibiting glucan synthase. Gyawali and Ibrahim authored the 2014

publication. Moreover, plant-derived compounds inhibit fungal adaptation and survival by targeting enzymes like cytochrome P450 subunits that are involved in energy metabolism and detoxification.

3.3. Producing Dangers of Oxidative Stress

Some plant extracts induce oxidative stress in fungal cells by, among other things, producing reactive oxygen species (ROS). When these highly reactive compounds damage the DNA, proteins, and lipids of fungal cells, the cells die. Gallic acid and caffeic acid are phenolic compounds that may increase the production of reactive oxygen species (ROS) by fungal cells, which may disrupt their normal cellular activity (Salari *et al.*, 2016). By simultaneously impacting several cellular components, oxidative stress makes adaptation harder for fungus and may be used to overcome fungal resistance.

Interrupting Biofilms

Because fungal biofilms are structured communities of fungal cells encased in an extracellular matrix, they are inherently resistant to the majority of antifungal therapies. Extracts from plants have showed promise in a number of areas, including preventing the formation of biofilms and removing existing biofilms. Cinnamomum zeylanicum oil and Melaleuca alternifolia oil are two examples of essential oils that have the potential to disrupt biofilm architecture and impede fungal cell adhesion to surfaces (Cowan, 1999; Gyawali & Ibrahim, 2014). Flavonoids and saponins not only attack fungal cells, but also the extracellular matrix, weakening it and exposing the cells to antifungal medications.

Mycoplasmal Efflux Pump Interference

Proteins called fungal efflux pumps are responsible for flushing out antifungal medications and other potentially toxic chemicals from fungal cells. These pumps are hindered by some plant-generated compounds. By binding to these pumps, plant extracts enhance the intracellular concentration of antifungal medicines, making them more potent. Examples of substances that may block the efflux pumps of drug-resistant fungus strains include alkaloids such as berberine and flavonoids such as quercetin. (Salari *et al.*, 2016).

A Multi-Target Mechanism's Critical Role

Plant extracts are quite effective against a spectrum of fungal illnesses because to their multi-pronged approach. Reduction of biofilms, oxidative stress, enzyme inhibition, and membrane disturbance renders resistance formation difficult. Synthetic antifungals lower the possibility of resistance development by concentrating on one route. Plant extracts are appealing choices for antifungal therapy whether taken by itself or in combination with other medications because of their particular feature. Scientists should be able to develop more efficient and long-lasting antifungal medicines for pharmaceutical, agricultural, and industrial usage if they can better grasp these mechanisms and identify better methods to use molecules generated from plants.

Table 1: Natural Antifungal Agent Advances — Research on Bioactive Compounds Derived from Plants (2024–2025)

<i>Year</i>	<i>Study Title</i>	<i>Methodology</i>	<i>Findings</i>	<i>Implications</i>
2024	The Fungicidal Effects of Cinnamon and How It Works	Extracted essential oils; conducted bioassays; used microscopy to study fungal cell damage	Inhibited spore germination and disrupted fungal cell walls	Cinnamon is effective against <i>Aspergillus flavus</i> and <i>Penicillium citrinum</i>
2025	What Makes Matcha Green Tea Bioactive	In vitro antifungal testing; gas chromatography for compound separation	Reduced growth of fruit-borne fungi; enhanced antioxidant defenses	Post-harvest use of tea tree oil as an antibacterial
2024	Plant-Based Antifungal Nanoparticles	Used <i>Swertia</i> spp. isolates in nanoparticle synthesis; employed spectroscopy and electron microscopy	Blocked fungal growth with minimal cytotoxicity	Plant-derived nanoparticles may offer sustainable antifungal treatments
2024	Guava Leaf Extract: A Comprehensive Review for the Textile Industry	Phytochemical analysis and in vitro antifungal testing	Safe for textile use and effective against fungi	<i>Psidium guajava</i> extract is a dual-purpose eco-friendly antimicrobial
2024	An Unconventional Deduction of How β -Sitosterol Fights <i>Candida albicans</i>	Investigated biofilms; optical microscopy; virulence gene expression studies	Significantly reduced fungal adherence and biofilm formation	β -Sitosterol targets fungal pathogenicity and shows therapeutic promise
2024	Phenolic Compounds from <i>Zanthoxylum armatum</i>	Isolated phenolic acids; used electron microscopy to observe structural effects	Induced oxidative stress and damaged fungal membranes	<i>Zanthoxylum</i> -derived compounds show potential as antifungal drugs
2024	Yeast Supports the Fruiting Body of <i>Hibiscus sabdariffa</i>	Used microscopy and molecular assays to study antioxidant impact	Reduced oxidative stress in yeast cells	<i>Hibiscus</i> extracts have antifungal and stress-reducing properties
2024	Analysis of Stereotypic Activity and Phytotoxicity of <i>Stephania venosa</i> Root Extract	LC-MS phytochemical profiling; antifungal and toxicity testing	Fungicidal but safe for non-target organisms	<i>Stephania venosa</i> root is a safe antifungal candidate
2024	<i>Piper arboretum</i> Extract and the Function of Dermatophytes	Evaluated antifungal efficacy; fractionated alkaloids	Inhibited dermatophyte growth by disrupting fungal cell wall formation	Alkaloids from <i>Piper arboretum</i> are promising agents against dermatophytes

Table 2: Plant-Based Bioactive Compounds and Their Extraction Methods

Plant Type	Active Compounds	Extraction Method
Allium (Onion with cruciferous)	Allium, a herb	The use of steam in distillation
Centaurea longa	Cinnamaldehyde, Eugenol	The method called hydro-distillation
Hibiscus (An aromatic Hibiscus)	Beta-3 sitosterol	Maximising solvent efficiency
Phyllanthus armatum	Aromatic compounds	Extraction with methanol
Costa Rican pathogen	Colourants, Antioxidants	Using ethanol for extraction
Stephania venosa	Alkaloids	LIMS evaluation
Salvia officinalis	Proteins, Antioxidants	Making ethanol from scratch
Piper (The gardens of Piper)	Alkaloids	The process of dichloromethane extraction
Swertia chirayita (Wrath of Chirayita)	Carotenoids, Alkaloids	Nanoparticle synthesis
Cinnamomum sinensis L. (The cinnamon tree)	Catechins, polyphenols	Maximising solvent efficiency
Cinnamon clove	Cinnamaldehyde, coumarins	Supercritical CO ₂ extraction
Artemisia (An artemisia plant)	Flavours of lavender and aromatic oils	The use of steam in distillation
Centaurea longa	Piperine	The Soxhlet separation
Divine peak	Clary sage, eugenol	Removing effluent
Azadirachta indica (According to azadirachta)	Hydroxyanisole, Azadirachtin	Pressing the extract cold
Cymbopogon species	Clerolid, Citral	The method called hydro-distillation
Centaurea serpentine	Fresh Cucumber and Turmeric	The Soxhlet separation
Evergreen tropical	Tannins, one, eight-cineoles	The use of steam in distillation
Moringa oleifera (Oil of Moringa persica)	Thiocyanates, polyphenols	Methanol extraction
The black asparagus	K1 vitamin	Maximising solvent efficiency

Research has shown that these compounds may effectively combat many different types of fungal infections, such as those caused by dermatophytes, *Aspergillus* species, *Candida albicans*, and agricultural fungus.

Table 3

Main Results and Antifungal Mechanisms of Bioactive Compounds

Major Outcomes	Antifungal Mechanism
Exhibits anti-fungal action over a wide range	Fungal cell wall degradation
Gets rid of the fungus that causes food to spoil.	The breaking of membranes and the suppression of spore germination
Reduces the potential damage caused by fungus	Keeping biofilms from forming and fungi from adhering
Great potential for application in food and medicine	Activation of oxidative stress and change in the

Protection against textile-related fungal diseases	cellular composition of fungi
Broad antifungal effectiveness	Blocking the fungus's ability to produce cell walls
	Harmful effects on mitochondria lead to harmful effects on fungus.
Can withstand fungal strains that have evolved resistance mechanisms	Limiting the pathways that fungi may use to spread
Perfect for eliminating skin fungus when used topically	Eliminating dermatophytes and the cell walls they produce
Efficacious green nanoparticles that combat fungal infections	Destroying fungal cells and amplifying oxidative stress
Increases the storage life of perishable foods.	Inhibition of fungal enzymes; disruption of energy metabolism
Aspergillus and Candida are significantly stunted in their development.	Formation of rhodopsin; disintegration of membrane potential
Exciting new antifungal	Fungus respiration pathway blockage and development of oxidative stress
Combines well with other treatments for maximum effect.	Inhibition of efflux pumps; disturbance of fungal ion homeostasis
Infectious fungi that have developed resistance to standard antifungal treatments	Inhibition of DNA synthesis by fungus
Strong effectiveness in combating agricultural illnesses	Interfering with fungal enzyme activity
A first-rate antifungal for usage in cosmetics	When spores' membranes are disrupted, they are unable to germinate.
Great for making medications	Biofilm and cell membrane disruption in fungi
An effective antifungal medication for pulmonary infections	The release of reactive oxygen species; membrane disruption
Effective against food poisoning mushrooms	Fungi that limit protein synthesis
Highly effective in eradicating fungal infections	Mitochondrial interference in fungus

Using Plant Extracts for Antifungal Therapeutic Purposes

The antifungal properties of plant extracts have led to their use in several sectors. Among these sectors are the food processing, medical, and agricultural industries. These applications demonstrate their versatility as natural alternatives to synthetic antifungal medications.

Fourth Section. Treatment for Individuals

Researchers have long sought to improve efficiency and decrease the side effects of conventional antifungals by using plant extracts as monotherapy or as an adjuvant for human fungal infections. Aloe vera extract shows promise as a therapy for skin-related fungal infections due to its antibacterial and anti-inflammatory properties. When applied to the skin, it soothes inflammation and stops the growth of fungus. For both oral and systemic candidiasis, research has shown that garlic extract (*Allium sativum*) has antifungal characteristics that effectively combat many species of *Candida*. It may be used topically or taken orally since its primary element, allicin, destroys fungal cell walls and halts their growth. Studies have shown that when used with existing antifungals, garlic may increase treatment efficacy and minimise drug resistance (Cowan, 1999; Gyawali and Ibrahim, 2014).

Agriculture

Plant-based substitutes for industrial fungicides might help to manage fungal diseases in agricultural environments in a more ecologically friendly manner. Derived from the *Azadirachta indica* plant, neem oil is a strong natural fungicide that targets several species causing crop illnesses. Among these things are species of *Alternaria* and *Fusarium*. Neem oil is healthy for the environment and efficient against fungal growth and reproduction as it is biodegradable and non-toxic. Basil (*Ocimum basilicum*) extracts assist postharvest fruit and crop fungal diseases free-from. Salari *et al.* (2016) claim that as basil extract includes essential oils with antifungal capabilities, it may help to lower postharvest losses resulting from species of *Aspergillus* and *Penicillium*.

4.3.1. Protecting Consumption

Plant extracts may be able to limit the rate of spoilage and mycotoxin production in perishable foods, therefore increasing their shelf life, due to their antifungal characteristics. Essential oils, like cinnamon's, have anti-fungal properties that could be useful for extending the shelf life of almost spoilt baked goods and other foods. The active component of cinnamon oil, cinnamonaldehyde, has antifungal characteristics that prolong the freshness of food without altering its taste or consistency. As a means of ensuring the safety and quality of stored grains, antimicrobials derived from plants may reduce the number of mycotoxins present (Cowan, 1999; Gyawali and Ibrahim, 2014). The significance and potential outcomes. Plant extracts have shown great promise as a potential alternative to synthetic antifungal medications due to their versatility and broad range of applications. When used to medical treatments, agricultural operations, and food preservation measures, they fight fungal illnesses and are in harmony with global efforts to promote eco-friendly solutions. Additional research and standards are necessary to maximise their use, guarantee consistency, and enable widespread implementation.

Plant Extracts: A Comparison Based on Conte

Table 4: Comparative Analysis of Plant Extracts by Context

Plant Species	Active Compound(s)	Methodology	Food Safety	Clinical Use	Agriculture
<i>Allium sativum</i>	Allicin	Steam distillation	Prevents fungal spoilage in stored grains and fruits	Antifungal creams for skin infections	Effective for against <i>Fusarium</i> and <i>Botrytis cinerea</i>
<i>Cinnamomum verum</i>	Cinnamaldehyde, Eugenol	Hydro-distillation	Used as a preservative in bakery products	Potential antifungal in oral candidiasis	Prevents fungal infestation in horticultural crops
<i>Hibiscus rosa-sinensis</i>	β -Sitosterol	Solvent extraction	Limited direct applications in food safety	Reduces virulence in <i>Candida albicans</i>	Enhances soil fungal balance by targeting

<i>Plant Species</i>	<i>Active Compound(s)</i>	<i>Methodology</i>	<i>Food Safety</i>	<i>Clinical Use</i>	<i>Agriculture</i>
					pathogens
<i>Zanthoxylum armatum</i>	Phenolic acids	Methanol extraction	Prevents fungal contamination in spices	Supports antifungal in treatment of respiratory infections	Broad-spectrum fungicide for crop protection
<i>Psidium guajava</i>	Flavonoids, Tannins	Ethanol extraction	Reduces fungal spoilage of juices and fruit-based products	May have adjunct use in antifungal medications	Protective coating for crops
<i>Stephania venosa</i>	Alkaloids	LC-MS profiling	Not reported	Investigational applications for systemic fungal infections	Promising antifungal for seed treatments
<i>Caralluma indica</i>	Glycosides, Flavonoids	Ethanol extraction	Low application in food safety	Adjunct to systemic antifungal treatments	Resistance management in fungal pathogens
<i>Piper arboretum</i>	Alkaloids	Dichloromethane extraction	Prevents fungal growth in fermented products	Treats dermatophyte-related skin conditions	No reported agricultural applications
<i>Swertia chirayita</i>	Terpenoids, Flavonoids	Nanoparticle synthesis	Enhances shelf life of stored grains	Green adjunct for nanotechnology antifungal therapies	Sustainable fungicide alternative
<i>Camellia sinensis</i>	Catechins, Polyphenols	Solvent extraction	Extends shelf life of dairy and bakery products	Antifungal activity against opportunistic infections	Controls fungal growth in tea crops
<i>Cinnamomum cassia</i>	Cinnamaldehyde, Coumarins	Supercritical CO ₂ extraction	Widely used in food preservatives	Treats resistant <i>Candida</i> strains	Effective against seed-borne fungal pathogens
<i>Artemisia eriantha</i>	Artemisinin, Essential oils	Steam distillation	Not reported	Promising antifungal for hepatocarcinoma-associated infections	Supports biopesticides for fungal pest control
<i>Piper nigrum</i>	Piperine	Soxhlet extraction	Prevents fungal contamination in	Enhances antifungal drug	Limited

Plant Species	Active Compound(s)	Methodology	Food Safety	Clinical Use	Agriculture
			dried spices	activity	reported use
<i>Ocimum sanctum</i>	Eugenol, Rosmarinic acid	Water extraction	Protects food oils from fungal contamination	Effective against drug-resistant <i>Candida</i> spp.	Potential for fungal disease suppression in crops
<i>Azadirachta indica</i>	Azadirachtin, Limonoids	Cold press extraction	Effective against fungal contamination in stored cereals	Antifungal adjunct in skin infections	Broad-spectrum bio-pesticide
<i>Cymbopogon citratus</i>	Citral, Geraniol	Hydro-distillation	Used in beverages and essential oils to prevent fungal spoilage	Suitable for cosmetic antifungal formulations	Controls fungal pests in leafy vegetables
<i>Curcuma longa</i>	Curcumin, Turmerones	Soxhlet extraction	Prevents fungal contamination in processed food products	Used in topical formulations for fungal infections	Effective against soil-borne pathogens
<i>Eucalyptus globulus</i>	1,8-Cineole, Tannins	Steam distillation	Limited direct application	Respiratory therapy	Controls fungal pathogens in timber production
<i>Moringa oleifera</i>	Isothiocyanates, Phenolic acids	Methanol extraction	Enhances microbial stability in dairy products	Antifungal supplements in immunocompromised patients	Soil application to suppress fungal disease cycles
<i>Nigella sativa</i>	Thymoquinone	Solvent extraction	Prevents fungal spoilage in oils	Highly effective systemic antifungal in infections	Potential antifungal in crop disease management

There is a large market for natural preservatives made from plants, such as citral (*Cymbopogon citratus*) and cinnamon (*Cinnamomum cassia*). Their potential to inhibit fungal growth might extend the shelf life of foods including spices, fruits, oils, and baked goods without the need of artificial preservatives.

Practical Uses in Healthcare: *Ziziphus rosa-sinensis* and *Zantholum armatum* have promising anti-Candida albicans and anti-dermatophyte characteristics. Active ingredients like curcumin (*Curcuma longa*) and thymoquinone (*Nigella sativa*) are being developed by the antifungal supplement business.

Application in Farming:

The bio-pesticidal effectiveness of *Swertia chirayita* and *Azadirachta indica* extracts have been shown. These are safer alternatives for synthetic fungicides that may help farmers practise sustainable agriculture.

Critical Issues:

Cinnamomum cassia, for instance, has use in food safety, medicine, and agriculture, making it an example of an extract with versatile usage. It is common practice in clinical practice to use adjunct medications or enhanced formulations. One of the ways in which extracts with biopesticide potential might benefit the agriculture industry is less dependent on chemical fungicides.

Plant Extracts: A Comparison Based on Context

Using antifungal medications to treat fungal infections connected to hepatocarcinoma has demonstrated positive outcomes. Two much sought-after plant-based natural preservatives are *Cinnamomum cassia* and *cymbopogon citratus*. Their ability to stop fungal development promises them as a natural substitute for chemical preservatives for increasing the shelf life of foods like fruits, spices, oils, and baked products.

Practical Uses in Healthcare:

Positive developments include the anti-Candida albicans and anti-dermatophyte properties of *Zantholum armatum* and *Ziziphus rosa-sinensis*. The antifungal supplement manufacturer is now working on new active chemicals, such as curcumin (derived from *Curcuma longa*) and thymoquinone (derived from *Nigella sativa*).

Application in Farming:

Extracts from *Swertia chirayita* and *Azadirachta indica* have been shown to physically slow down pest development. Much research has validated this. Farmers aiming for a sustainable agricultural life should find these less damaging substitutes for industrial fungicides helpful. Issues Still To Be Solved: One such extract with several uses is *Cinnamomum cassia*. In food safety, medicine, and agriculture this plant has numerous uses. Furthermore, much benefit would be the domain of food safety by its use. In clinical practice, it is usual practice to use enhanced formulations or extra medications. Extracts with biopesticide potential might help the agriculture industry greatly. They might reduce the demand for chemical fungicides in agriculture, among other things, therefore helping in many ways.

Prospects and Challenges in Plant Extractive Research Although plant extracts might be antifungal agents, their extensive use is limited in agriculture, industry, and medicine owing various challenges. To allay the concerns about resistance development, toxicity, scalability, and standards, will need coordinated effort. We really need to cooperate to fix these problems.

5. Respect for accepted norms

One of the main issues the business deals with is the quite different chemical composition of plant extracts. The many factors that could influence the precise chemical content of a plant extract are just a few of the several possible ones: Many elements in the environmental surroundings might influence the bioactive chemical concentrations in plants. These factors include soil type, climate, and geographical location as well as others. Task Completion and Data Gathering: The moment of harvesting, the technique of removal, and the storage conditions all affect the content and efficacy of the finished product. Given the variety of plant extracts, standardising them to guarantee a uniform antifungal activity becomes much more challenging. Reaching regulatory clearance and economic success depends on the synthesis of extracts with repeatable bioactivity.

On the other hand, this becomes troublesome in the lack of reliable quality control mechanisms (Cowan, 1999). People's concerns and fears about their safety and security are really significant. Although plant extracts are sometimes thought to provide less of a danger than synthetic antifungal drugs, careful assessment of plant extract safety profiles is still required. Key points include Determining suitable dosages for use in humans and other animals remains a challenge due to the toxicity of some bioactive compounds at high concentrations. Long-Term Effects: In medical settings in particular, significant investigation into the potential side effects and drug interactions of using plant extracts over an extended period of time is required. According to Gyawali and Ibrahim (2014), comprehensive clinical and preclinical research is necessary to ensure that plant-based antifungals are just as safe as their synthetic counterparts. Only then can these concerns be addressed.

5.3.1. Maximising efficacy via expansion

Producing plant extracts on a large scale is not without its economic and logistical challenges:

An important concern about the availability of raw materials is the preservation of natural populations and the prevention of environmental damage in the process of obtaining sufficient plant materials. High Production Costs: It could be costly to isolate, purify, and stabilise bioactive compounds, particularly from plants that don't yield enough of the desired antifungal compounds. It is essential to verify that the processes used to extract plant materials adhere to ecologically acceptable standards in order to guarantee that the process does not harm the environment. Traditional farming methods may find a solution to these issues by using biotechnology techniques. Cultivating plant tissues and engineering plant metabolites by microbes are two examples of such techniques. "Al-Salih, along with others, 2021" part in part 5.4.1 of the Construction of a Defence Mechanism Although plant extracts have many uses and are often praised for their multi-targeting capabilities, fungal populations may nevertheless develop resistance to them if they are used too often or in the wrong manner. When it comes to agricultural uses, this becomes an even bigger worry since fungi might be subjected to selection pressure due to the substantial use of plant-based fungicides. This is due to the fact that natural selection may favour fungus. In order to forestall a repeat of the same: To prevent the development of resistance, it is recommended to utilise plant extracts in conjunction with standard antifungal drugs. Traditional antifungals and plant extracts with antifungal

characteristics have been the subject of research as potential combination treatments. Not only are these therapies less likely to cause treatment resistance, but they also have a better chance of actually working. Increasing our understanding of how fungi adapt to antifungal chemicals derived from plants is crucial if we are to conquer fungal resistance. To discover the full potential of plant extracts, conduct research on them, and bring them to market, it is crucial for groups of experts from many professions to interact. Despite the many obstacles, this remains true. Incorporating these chemicals into antifungal therapies requires a standardised extract composition, safety assurances, the elimination of scaling impediments, and the reduction of resistance threats. It would be incomplete without these items. If we can solve these problems, we can increase the chances of using plant-based antifungals and discover new ways to treat fungal infections in agriculture, medicine, and food preservation. Number six on the list is a look at what's to come. Research on antifungal medications produced from plant extracts, as well as proactive improvements in research, technology, and regulation, are of the highest significance. These potential initiatives will be able to surpass the current constraints since antifungals made from plants are going to be thoroughly integrated into diverse applications.

6.1. 4. Studies on Collaboration

By integrating plant extracts with conventional antifungal drugs, we may be able to increase effectiveness while minimising resistance. Tasks that are complementary to one another might The antifungal medicine and plant extract are more effective when administered in smaller doses, which means less side effects. To Delay the Adaptation and Resistance Development of Fungi, It Is Recommended to Use Multiple Targets Simultaneously. Extending the Scope of Action: Synergistic chemicals, when administered together, have the potential to boost their effectiveness against resistant or otherwise difficult-to-treat fungal strains. Tea tree oil and fluconazole are two examples of essential oil combinations that have shown to be more efficient against *Candida* species, according to research. Gyawali and Ibrahim authored the 2014 publication. We need further research to find the best combinations and strategies.

6.2. Miniature molecule-based technology

When applied to plant-based antifungal compounds, nanotechnology has the potential to improve their transport and stability. Nanoparticles coated with plant extracts may be able to:

Nanoparticle encapsulation improves bioavailability by increasing the absorption and utilisation of bioactive substances, including those with low solubility. Encapsulation enhances the stability of fragile phytochemicals by preventing their degradation due to exposure to light, heat, or oxygen. To avoid systemic toxicity and increase local effectiveness, plant extracts may be administered directly to the infection site using tailored nanocarriers. An example of how nanotechnology has the potential to revolutionise plant-based antifungal medications is the improved antifungal effectiveness of thymol-loaded nanoparticles against *Aspergillus* spp. (Salari *et al.*, 2016; Al-Salih, 2021).

6.3. Proteomics and Genomics

Research on modern genomes and proteomics might provide fresh insights on the connection between fungal infections and plant extracts. Investigating fungal reactions at the molecular level might allow the following: By demonstrating which genes and pathways relate to the effects of plant extracts, genomic studies might assist in the identification of molecular targets by researchers. Find the proteins that

provide fungal resistance to molecules derived from plants. Proteomic research might identify which proteins enable certain medications to be resistant against fungus. Development of more customised and successful plant-based therapies depends on a deeper knowledge of the molecular basis of antifungal action. Maybe this knowledge might help new approaches of chemical synthesis.

Transcriptome studies have shown changes in gene expression linked with oxidative stress response and membrane formation in *Candida albicans* subjected to plant extractions. These findings would help to clarify the action mechanisms (Cowan, 1999; Al-Salih, 2021).

Significant Policies and Procedures Before they are commercialised, strict control systems are essential to ensure the effectiveness, safety, and moral usage of plant-based antifungals. Here are some key factors to give thought:

Standardising the content and potency of plant extracts helps to guarantee consistency in quality control management. A vital component of toxicology and safety is the evaluation of toxicity profiles, which have to include both the short- and long-term effects as well as pharmaceutical interactions.

Implementing sustainable practices in the acquisition and processing of plant materials may help reduce the likelihood of overharvesting and environmental harm. An ethical use would be to encourage fair trade practices and guarantee that indigenous communities that provide traditional knowledge get a fair compensation for their contributions. It is critical that researchers, lawmakers, and industry stakeholders collaborate to construct these frameworks and facilitate the global adoption of plant-based antifungal treatments. The creation of plant-based antifungals relies on interdisciplinary collaboration and innovative thinking. Strong regulatory legislation, nanotechnology, molecular biology, and collaborative research are essential for plant extracts to reach their full potential. The war against fungal infections in food production, farming, and human health may be won if we can overcome the current challenges and make room for plant-derived antifungals to replace toxic synthetic alternatives.

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The authors declare that there is no conflict of interest regarding the publication of this paper.

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