



## A Comprehensive Review Of Ethnopharmacology, Phytochemistry And Pharmacology Of Some Medicinal And Edible Mushrooms

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Article History	Abstract
Received: 25-11-2020 Revised: 20-12-2020 Accepted: 05-01-2021	<p>Since mushrooms have long been utilized throughout traditional medical practices curiosity about exploring mushrooms scientifically and their possible health advantages is rising. Polysaccharides, sterols, and phenolic compounds are just a few of the bioactive substances that can be found in mushrooms and have a positive impact on human health. The study of how different cultures have used mushrooms in traditional medicine is what makes mushrooms relevant from an ethnopharmacological perspective. Mushrooms have a wide variety of ethnopharmacological uses, and these uses change according to culture and mushroom species. Although the scientific study of mushrooms is still in its infancy, the preliminary findings are encouraging. There is proof that mushrooms can boost the immune system, lower cholesterol, reduce inflammation, stop the spread of cancer cells, and protect cells from free radical damage. We will probably discover much more information on the health advantages of mushrooms as further research is conducted. A useful source of knowledge about the possible health advantages of mushrooms is their ethnopharmacological relevance. From the present review, knowledge about ethnobotany, phytochemistry, and pharmacology can be utilised to direct scientific study and create new health-improving products made from mushrooms.</p>
CC License CC-BY-NC-SA 4.0	<b>Keywords:</b> Bioactive chemicals, Health benefits, Ethnopharmacology, Mushrooms.

### 1. INTRODUCTION

The discovery of novel bioactive chemicals in natural sources may facilitate drug development (Debbab et al., 2011, 2012; Newman and Cragg, 2007). With 3-5 million species, fungi may rank as the second most varied group of living things (Blackwell 2011). Fungi from Basidiomycota and certain Ascomycota that produce fleshy fruit bodies, which generate bioactive substances, may treat a number of ailments (Aly et al.,

2011; de Silva et al., 2012a). For many years, especially in Asia (Rapior et al., 2000; Ferreira et al., 2010; Xu et al., 2011), mushrooms have been used to treat or prevent a variety of illnesses. Unstudied species of macrofungi may aid in the discovery of compounds with biological effects. The fascinating substances produced by basidiomycota have a variety of biogenetic origins. Numerous research (Bao et al. 2001; Keller et al. 2002; Poucheret et al. 2006; Jeong 2011) have indicated that these substances have pharmacological effects on bacteria, fungi, viruses, and other species.

Medicinal mushrooms, which have historically dominated mushroom production in the Far East, are increasingly making their way into the Western market because of their excellent organoleptic properties and acceptance as real functional foods. The employment of complementary medical techniques, such as traditional medicine, in which medicinal mushrooms play a significant role, as well as rising consumer desire for wholesome foods, have all helped medicinal mushrooms gain traction on the world market. According to studies, there are 14,000–22,000 different varieties of mushrooms. Only 10% of the 140,000 species of mushrooms have been identified. There are still 7,000 kinds of mushrooms that haven't been found that may be helpful to human civilization if we assume that 5% of unknown and untested mushrooms will be (Hawksworth, 2001). Only a tiny portion of mushroom species have undergone in-depth research. Bioactive substances may be produced by microorganisms such as *Aspergillus*, *Tolyptocladium inflatum* W. Gams, and *Claviceps purpurea* (Fr.) Tul. (Lindequist et al. 2005).

In this review, we have discussed the incidents in the ethnomedicinal use of mushrooms in terms of ethnopharmacology, phytochemistry, pharmacology, and other increasing parameters, as well as providing support for the utilization of selected mushrooms as a common drug to produce bioactive secondary metabolites. We may presume that mushrooms have tremendous potential for successful bioprospecting given their enhanced capabilities for biological, pharmacological, and chemical investigation. This is something that we can say with confidence. In the present review, we have compiled the mushrooms related to their edible, non-edible, and medicinal importance.

## 2. BOTANY AND ETHNOBOTANY

There is no taxonomic category known as "mushroom." The term "mushroom" should be used here in the sense given by Chang and Miles, who define a mushroom as "a macrofungus with a distinctive fruiting body, that can be both hypogeous or epigeous, sufficiently big to be considered with the naked eye and to be picked by hand." Taxonomically speaking, mushrooms are primarily classified as basidiomycetes, but there are a few ascomycete species that fall under this category as well (Chang and Miles 1992).

### 2.1 *Inonotus obliquus* (Ach. ex Pers.) (Wild Chaga)

The fruit bodies look like tumors (a sterile lump without a stalk in the dimension of 25 to 40 cm and has a dark coloration of grey and a deep cleavage); The parental section is five mm thick, cruciform, and dark-brown in colour, while surrounding area is light yellow in colour. The tube length is 3 to 10 mm, and its front end is open (Huang, 2002). The pore is circular, smooth, and shiny, 9–10, and the cleavage is a parasitic black fungus called *Inonotus obliquus* grows on mature birch tree trunks. birch's brown *Ganoderma lucidum*, also known as chaga and birch antler, is a medicinal white-rot fungus. The white-rot fungus *Inonotus obliquus* grows on older birch trees' living trunks. *Inonotus obliquus* is the cause of the host tree's white heart rot. Chaga spores enter the tree through wounds, especially branch stubs. Due to white rot, the host's heartwood will degrade. The infection's sapwood can only be punctured near the mycelium mass on the sterile surface. For 10–80 years, the chaga fungus will degrade the living tree. While the tree is alive, the black exterior conk will only grow sterile mycelium and go to the sexual stage after damaging the tree. Under the bark, *Inonotus obliquus* produces fertile fruiting bodies. These organisms start white and turn brown. The fruiting body is rarely visible since most reproductive activity happens under the bark. These fruiting structures produce basidiospores, which transmit the infection to more sensitive plants (Lee et al. 2008).

### 2.2 *Pleurotus ostreatus* (Jacq. Ex Fr.) P. Kumm.

Both the scientific name and the colloquial name of the fruiting body of *P. ostreatus* take into account the morphology of the fruiting body (Sher et al., 2011). The way in which the stem grows with respect to the cap is referred to as "pleurotus," which is a Latin term that means "sideways." The word "oyster" comes from the Latin word "ostreatus," which refers to the shape of the cap, which resembles a bivalve with two shells. The cap of *P. ostreatus* ranges in length from 5 to 25 centimeters and is broad, fan-shaped, or oyster-shaped. Natural specimens might be white, gray, tan, or dark brown in color, and their edges are smooth when they

are young but frequently have lobes and waves on them as they mature. Because of the way the stipes are organized, the thickness of the white, firm flesh can vary. The gills of the mushroom have a color range from white to cream, and they dangle downward from the stalk, if it has one. The stipe is not located in the middle of the piece of wood and is instead linked to one of the sides. On a dark surface, the mushroom's spore print, which ranges in color from white to lilac-gray, is most visible (Rajarithnam et al., 1987; Menolli et al., 2010).

### 2.3 *Trametes versicolor* (L.) Lloyd

*Trametes versicolor*, also referred to as Turkey Tail, is a widespread lignicolous fungal species with white rot that grows on various trees such as oak and Prunus and on various conifers such as fir or pine trees (Gautam, 2013; Janjušević et al., 2017; WA et al., 2020). Despite the fact that this genus is inedible, it has been traditionally used in the medicinal practices of a number of different nations. Numerous studies have offered scientific confirmation of their vast range of pharmacological behavior including anti-oxidant, anti-cancer, and anti-inflammatory characteristics, amongst others. (Kamiyama et al., 2013; Janjušević et al., 2017). The upper surface of the cap displays the characteristic concentric zones of various hues, and the border is always the palest part of the surface. A dark layer sits just below a layer of tomentum, on top of which lies a layer of white flesh (PUIA et al., 2018). The thickness of the flesh itself ranges from one to three millimeters, and it has a leathery consistency. Older specimens, like the one in the image, may have areas where green algae is forming, giving them a green appearance. On the fallen branches and fallen trees of deciduous trees, it will often develop in tiled layers that are arranged in groups or rows. The mushroom does not have a stem, and the cap might be a rusty brown or a deeper brown color. It occasionally has black zones. The area of the cap may range from 0.5 to 1 centimeter and be up to 8 cm by 5 cm. It is often triangular or circular in shape, and it has zones of fine hairs. White to light brown in color, the surface of the pores is circular and, with increasing age, becomes twisted and labyrinthine. 3–8 pores per millimeter (Gautam, 2013).

### 2.4 *Ganoderma applanatum* (Pers.) Pat.

The mycelium of this parasitic and saprophytic fungus develops on both living and dead trees. One by one or in clusters, *Ganoderma applanatum* grows. Its fruiting bodies are inedible, leathery, and woody, measuring 3 to 30 cm in width by 5 to 50 cm in length. They begin white before turning a deep red-brown color. Reddish-brown conidia are found on top of the fruiting body. Brown spores come from the underside of pores. Due to their high concentration, a piece of conk measuring 10–10 cm can produce 4.65 billion spores in a day. 4–12 mm tubes have 4-6 round pores per millimeter at the end. Perennial fruiting structures are constantly growing and developing new pores. These strata can be seen in cross-sections or in concentric rings on fruiting bodies. Thus, the age of the fruiting body can be determined by tree rings. In many trees, *Ganoderma applanatum* causes the heartwood to decay. It can develop as a pathogen in the living sapwood of elder trees that are moist, old, or diseased, while it mostly grows on dead trees (Jogaiah et al., 2019).

### 2.5 *Agaricus campestris*

The white, 3 to 12 centimeter-diameter cap may have fine scales. It starts out hemispherical before maturing into a flat shape. Both the spores and the valves have a pinkish colour at first, then a reddish-brown colour, and finally a dark brown colour. (Davis et al., 2012). According to Davis et al. (2012) and Nilsson & Persson (1977), the stipe is 3 to 10 cm in height, 1 to 2 cm wide, and primarily white in colour. In addition to that, it has a single delicate ring. It has a mild flavor. White tissues bruise a rusty crimson brown, unlike *Agaricus xanthodermus* and comparable species, which have a fleshy yellow bruise. The spores are 5.5 to 8 mm by 4 to 5 mm in size and elliptical in shape, while the sporewall is thick. Dark brown in colour. Cheilocystidia are not present (Miller & Miller, 2006).

## 3. TRADITIONAL USES OF MUSHROOMS

Since ancient times, macrofungi have been the norm in human culture. Around the world, oral tradition has been passed down from one generation to the next, describing the health benefits of mushrooms. Ancient scriptures like the Vedas reference and value the conventional understanding of mushroom use throughout the world. According to Roman and Chinese mythology, mushrooms have capacity and a life energy that gives them power. The inhabitants of prehistoric China, India, and Iran employed mushrooms in their religious rituals and thought of them as gifts from the deity Osiris (Sharma, 2003; Lowy, 1971).

### 3.1 *Inonotus obliquus* (Ach. ex Pers.) (Wild Chaga)

Rationality and mysticism coexist in traditional medicine. Finding substances that can cure diseases by observing nature makes sense. In many countries, this is the only way to improve health because of poverty or a lack of modern treatment. *I. obliquus* was used in ancient traditional medicine. One of the first texts referencing its medicinal use is Hippocrates' "Corpus Hippocraticum" (Szczepkowski et al. 2013). The father of medicine used mushroom infusions to treat wounds. Since the 12th century, chaga mushrooms have been used throughout Eastern Europe. In the past, Kiev Kniaz treated a lip malignancy. *Inonotus obliquus* is used in Siberian traditional medicine for a number of things (Perevedentseva, 2013). The fungus was helpful because of its gastrointestinal, anti-inflammatory, anti-parasitic, and anti-tuberculosis properties (Harpe, 1801). It was also suggested for liver and heart conditions. The most often used techniques were infusions, inhalations, and aqueous macerates (Harpe, 1801). Antibacterial soaps for external *I. obliquus* were created. In Siberia, infusions prepared from chaga mushrooms started to supplant tea in the latter half of the 20th century.

Before modern oncology, *Inonotus obliquus* was studied for its cancer-fighting abilities. Popular literature like Aleksander Solzhenitsyn's Cancer Ward (Saar, 1991) has such depictions. Chaga mushroom extracts were used to enhance cardiac function, the plasma lipid system, and their anti-bacterial, anti-inflammatory, and anti-cancer actions in Asian and Baltic countries as well as in Russia (Shashkina et al., 2006). *I. obliquus* may also aid in the prevention of diseases including atherosclerosis, cancer, diabetes, accelerated aging, and central nervous system degeneration that are linked to free radicals in modern society (Park et al., 2005). Chaga extracts slow down the multiplication of HCV and HIV (Shibnev et al., 2011).

### 3.2 *Pleurotus ostreatus* (Jacq. Ex Fr.) P. Kumm.

It is common knowledge across a number of continents, including Asia, Europe, South America, and Africa, that mushrooms belonging to the genus *Pleurotus* contain medicinal characteristics (Deepalakshmi and Sankaran, 2014). The antibacterial properties of *Pleurotus ostreatus* are most effective. The extract that was taken from this species of mushroom, which is usually referred to as the shimeji mushroom, was shown to be effective as an antioxidant agent in vitro (Stefanello et al., 2012). A positive correlation between the extract's total phenolic content and its antioxidant capacity served as evidence of this.

### 3.3 *Trametes versicolor* (L.) Lloyd

Traditional East Asian botanicals (Yun-zhi), such as *Trametes* formulations, are still widely used today and are said to have a number of health advantages when taken orally. According to Scarpari et al. (2017), *Trametes versicolor* has been utilized to restore qi and essence. According to the Compendium of Chinese Materia Medica (Ch, 1995), there are more than 120 different strains of *Trametes versicolor*. Traditional Chinese Medicine (TCM) practitioners claim that *Trametes versicolor* has a mildly sweet taste and a chilly quality and that it acts on the liver and spleen to produce its effects (Yang et al., 1993; Ng, 1998). *Trametes versicolor* is regarded as a medicinal herb that has been used to treat illnesses in various TCM classics. It is thought to be effective at clearing heat, removing toxins, stimulating the immune system, and strengthening the body. *Trametes versicolor* is frequently recommended for treating different cancers, chronic liver disease, and infections of the upper respiratory, urinary, and gastrointestinal systems in TCM therapeutic practice (Yang et al., 1993; Ng, 1998; Chu et al., 2002). While most therapeutic mushrooms promote a wide range of immune activity, different species frequently have distinctive immunological characteristics. According to Ramberg et al. (2010), *Trametes versicolor*, also called "turkey tail" and formerly *Coriolus versicolor*, has been shown to improve the immune system's innate as well as adaptive responses.

### 3.4 *Ganoderma applanatum* (Pers.) Pat.

*Ganoderma* is a therapeutic fungus that dates back more than two thousand years and has been mentioned in Shennong's herbal classic. The Shennong herbal classic included the recordings "Chizhi," "Qingzhi," "Heizhi," "Baizhi," and "Huangzhi," in addition to "Zizhi." These six different varieties of *Ganoderma* are said to improve eyesight, build muscles and bones, strengthen kidneys, and ease nerves. *Ganoderma* has been associated to a number of health benefits, including a relaxed body, an eased aging process, and a longer life span. On the other hand, recent research has shown that the *Ganoderma* genus has a history that extends further back in time than 6,000 years. Five prehistoric samples of the *G.* species were discovered at three Neolithic sites in the province of Zhejiang in China (Jong and Birmingham, 1992; Sukowska-Ziaja et al., 2019). *G. applanatum* is a non-edible fungus that may be identified in this scene by its unpolished, wrinkled, and white underside. In West Cameroon's indigenous tribes, it is common knowledge that the plant

has specific medical benefits, such as the treatment of headaches, hypertension, fatigue, diabetes, and hepatitis (Wang et al. 2020), despite the fact that these claims have not been confirmed.

### 3.5 *Agaricus campestris*

One of the most commonly ingested wild mushrooms is *Agaricus campestris*. A wide variety of biomolecules with both nutritional and therapeutic characteristics can be found in both wild and domesticated mushrooms. They are good agents for food and medicine because of these qualities. Such qualities have been acknowledged as functional food attributes as well as a source for the creation of pharmaceuticals and nutraceuticals. Active metabolites that have built up in the fruiting bodies, spores, and mycelia have anticancer, antibacterial, cardiac-protecting, anti-inflammatory, immunomodulatory, anti-diabetic, and antiviral characteristics; as a result, they are mostly employed for their medicinal and nutritional benefits (Wani et al., 2010). They have historically been applied to pharmacology and therapeutics. Another illustration is the *Agaricus campestris* Linnaeus ex Fries (field mushroom), a fungus that can be found in the Caribbean and the northern and southern temperate zones. According to reports, *A. campestris* was cooked in milk to treat throat cancer (Dias et al. 2012).

## 4. PHYTOCHEMISTRY OF MUSHROOMS

### 4.1 *Inonotus obliquus* (Ach. ex Pers.) (Wild Chaga)

#### Polysaccharides

These polysaccharides are the most prevalent in chaga mushrooms and have been linked to a number of biological functions. In this regard, Huang et al. (2012) used column chromatography to identify five polysaccharides from *Inonotus obliquus*. Monosaccharide compositions and average molecular weights were examined using GC and HPGPC. IOP1b only contained the 1.5–105 Da molecule glucose. Rha, Man, and Glu made up the components of IOP2a, which had a molecular weight of 9.3104 Da and a molecular proportion of 1.0:2.3:1.7. IOP2c included Man, Glu, and Gal in the molecular proportions of 1.0:2.9:0.8 and had a mass of 2.3 105 Da. IOP3a was discovered to be a glycoprotein with an O-link between the protein and the carbohydrate by alkali.

#### Triterpenoids

There have been 34 different triterpenoids extracted from the sclerotia of *Inonotus obliquus*. These are examples of compounds that were isolated. Inotodiol, 3 $\beta$ -hydroxy-lanosta-8,24-dien-21-al isolated by Kahlos et al. (1984), Lanosterol, Trametenolic acid, 3 $\beta$ -Hydroxy-8,24-dien-lanosta-21,23-lactone, 21,24-Cyclopenta-3 $\beta$ ,21,25-triol-8-ene, 3 $\beta$ ,22,25-Trihydroxy-Lanosta-8-ene, and inonotsuoxide isolated by Yusoo et al., (2001). Taji et al. (2007) isolated inonotsulides A, B, and C, whereas Taji et al. (2008) isolated lanosta-8,23E-diene-3, 22R, 25-triol, lanosta-7:9, 23E-triene-3, 22 R, 25-triol, lanosta-8,24-dien-3, 21-diol. Nakamura et al. (2009) were able to obtain inoterpenes A through F under these circumstances. Handa and his colleagues discovered seven different triterpenoids in this landscape in 2010, including spiroinonotsuoxodiol and inonotsudiol A, B, and C. Two more triterpenoids, epoxyinonotsudiol and methoxyinonotsutrial, were discovered in 2012 (Handa et al., 2010; Handa et al., 2012).

#### Phenolics

Lee et al. (2007) extracted six polyphenols from the methanolic extract of *Inonotus obliquus*: inonoblins A, B, and C and phelligridins D, E, and G. Seven low-molecular-weight phenolics, 4-hydroxy-3,5-dimethoxy benzoic acid, 2-hydroxy-1-hydroxymethyl ethyl ester, protocatechic acid, caffeic acid, 3,4-dihydroxybenzaldehyde, 2,5-dihydroxyterephthalic acid, syringic acid, and 3,4-dihydroxybenzalacetone, were isolated from *Inonotus obliquus* and tested for antioxidant and anticancer activity (Nakajima et al., 2009). 2,5-dihydroxybenzalactone and gallic acid were extracted and tested on cancer cells (Kim et al., 2011; Kuriyama, 2013; Babitskaya et al., 2000).

#### Proteins

Leucine, glycine, aspartic acid, tyrosine, leucine, methionine, lysine, threonine, serine, and histidine were the 15 amino acids generated by the protein hydrolysis of *Inonotus obliquus*, accounting for 40% of the total (Shashkina et al., 2006). Hyun et al. found an *Inonotus obliquus* platelet aggregation-inhibiting peptide in 2006. In this panorama, ethanol-derived peptides are isolated by reverse-phase HPLC. By using LC/MS analysis, it was discovered that the peptide from *Inonotus obliquus* had a molecular size of 365 Da, an amino acid composition of Glycine (50%), Cystine (25%), and Tryptophen (25%), and a sequence of Trp-Gly-Cys.

**Lignin**

The extracts of *Inonotus obliquus* boiling water were filtered many times using ultrafiltration and Superdex 200 columns. IR, UV, and <sup>1</sup>H NMR spectroscopy all indicated that fraction 23 contained the majority of the hydrophilic lignin derivatives. Coniferyl alcohol and sinapyl alcohol, both of which are components of lignin, did not significantly inhibit HIV-1 protease activity; however, fraction 23 did. According to Ichimura et al. (1998), lignin that is water-soluble has the ability to suppress HIV-1 because a derivative of it inhibits HIV reverse transcriptase.

**4.2 *Pleurotus ostreatus* (Jacq. Ex Fr.) P. Kumm.****Polysaccharides**

Numerous polysaccharides, including  $\alpha$ -glucans and  $\beta$ -glucans, are found in the cell wall of the mushroom. These polysaccharides have garnered a lot of scientific attention. These substances have been found to have immunomodulatory and antioxidant characteristics, which make them interesting candidates for therapeutic uses (Zhang et al., 2012).

**Proteoproteins and amino acids**

The protein-rich fish *Pleurotus ostreatus* contains both needed and optional amino acids. For their bioactive qualities, such as antioxidant and antihypertensive effects, several peptides produced from the mushroom have been examined (Chirinang & Intarapichet., 2009).

**Terpenoids and sterols**

Terpenoids and sterols, particularly ergosterol and lanosterol, can be found in abundance in mushrooms. Their anti-inflammatory and antioxidant capabilities, which highlight their potential therapeutic benefits, are a result of these substances (Corrêa et al., 2016).

**Phenolic Compounds**

Numerous phenolic compounds, including flavonoids and phenolic acids, have been found in *Pleurotus ostreatus*, and they have been shown to have anti-inflammatory and antioxidant properties. This section investigates their possible contribution to the prevention of diseases caused by oxidative stress and inflammation (Gąsecka et al., 2016).

**Fatty acids and lipids**

Unsaturated fatty acids, which have been connected to cardiovascular health and anti-inflammatory properties, are among the interesting lipids present in the mushroom (Pedneault et al., 2007).

**Volatile oils**

*Pleurotus ostreatus*' distinctive flavor and aroma are a result of its volatile constituents. Further research is necessary because some of these chemicals may also have health effects (Omarini et al., 2010).

**4.3 *Trametes versicolor* (L.) Lloyd****Polysaccharides**

The cell wall of the mushroom contains a variety of polysaccharides, including  $\beta$ -glucans and heteropolysaccharides, which have immunomodulatory, anticancer, and antioxidant activities. Polypeptides are also present in the mushroom cell wall. *Trametes versicolor* polypeptides have been identified, and they have antiviral and anticancer properties, making them potential candidates for medicinal research (Chen et al., 2015).

**Triterpenoids**

*Trametes versicolor* has a significant amount of lanostane- and ergostane-type triterpenoids. As natural medicinal agents, these triterpenoids exhibit anti-inflammatory, antioxidant, and anticancer effects (Jin et al., 2019).

**Phenolic Compounds**

The mushroom contains phenolic compounds, including flavonoids and phenolic acids, which have anti-inflammatory and antioxidant properties. This section explains how they can prevent diseases caused by oxidative stress and inflammation (Rašeta et al., 2020).

### Terpenoids and Sterols

Ergosterol, among other terpenoids and sterols, is found in abundance in *Trametes versicolor*. Researchers have looked at the possible immunomodulatory and anticancer properties of these substances (Bains et al., 2021).

### Fatty acids and Lipids

The mushroom has a lipid profile that is high in essential fatty acids, which are crucial for the immune system and cardiovascular health (Bains et al., 2021).

#### 4.4 *Ganoderma applanatum* (Pers.) Pat.

##### Triterpenoids

These substances have anti-inflammatory, anti-cancer, and antioxidant effects. Triterpenoids such ganoderic acids, lucidenic acids, and ganodermediols have been identified in *G. applanatum* (Elkhateeb et al., 2018).

##### Flavonoids

These substances, which are also antioxidants, have been demonstrated to provide a number of health advantages, including as lowering the risk of cancer, heart disease, and stroke. *G. applanatum* contains flavonoids as quercetin, rutin, and myricetin among others (Li et al., 2016).

##### Saponins

The biological effects of these substances range from anti-inflammatory to anti-cancer to immunomodulatory. Ganodarins, ganodiols, and ganosporins are a few of the saponins discovered in *G. applanatum* (Luo et al., 2015).

##### Other compounds

Sterols, polysaccharides, and proteins are some of the additional substances included in *G. applanatum*. Furthermore, numerous health advantages of these substances have been demonstrated (Wang et al., 2020).

#### 4.5 *Agaricus campestris*

##### Phenolic compounds

A class of organic substances known as phenolic compounds has anti-inflammatory, anti-cancer, and antioxidant effects. Gallic acid, caffeic acid, ferulic acid, quercetin, and rutin are some of the phenolic chemicals found in *A. campestris*. These substances have been demonstrated to neutralise free radicals, reduce inflammation, and slow the growth of cancer cells .

##### Indoles

Indoles are a class of organic substances that are present in a wide variety of plants, including mushrooms. *A. campestris* contains indol-3-carbinol (I3C) and diindolylmethane (DIM), among other indoles. The anti-cancer property of I3C and DIM have been demonstrated, and they might possibly be advantageous for cardiovascular health (Bains et al., 2021).

##### Sterols

Sterols are a class of organic substances that are present in both plants and mammals. Ergosterol, campesterol, and stigmasterol are among the sterols found in *A. campestris*. It has been demonstrated that ergosterol, a precursor of vitamin D, has anti-cancer properties. It has been demonstrated that stigmasterol and campesterol reduce cholesterol levels (Goulston et al., 1975).

##### Other Compounds

*Campestris* includes lentinan, beta-glucans, and chitin in addition to phenolic compounds, indoles, and sterols. Immune-boosting beta-glucan lentinan. Beta-glucans, a dietary fibre, decrease cholesterol and inflammation. Mushroom cell walls include chitin. It aids ins weight loss and digestive health .

## 5. PHARMACOLOGICAL ACTIVITY

### 5.1 *Inonotus obliquus* (Ach. ex Pers.) (Wild Chaga)

#### Anti-diabetic activity

A study conducted by Wang et al. (2017) In STZ-induced diabetic mice, *I. obliquus* raised insulin levels and corrected glucose enzyme metabolism. *I. obliquus* polysaccharide extract was able to block  $\beta$ -glucosidase, which resulted in a delay in the digestive organs' ability to absorb glucose shortly after eating. The extract of *I. obliquus* prepared in ethyl acetate decreased the activity of  $\alpha$ -amylase and increased HDL cholesterol levels in mice with alloxan-induced diabetes. In this order, an increased level of insulin was reported in diabetic nephropathic mice (C57BL/6) treated by the polysaccharide extracts of *I. obliquus*. Many researchers also reported a significant reduction in blood glucose level, plasma insulin level, and body weight in STZ-induced diabetic mice when treated with an IOP-chromium (III) complex (Wang et al. 2017; Cha et al. 2006; Chen et al. 2010).

#### Anticancer Activity

According to the research done by Youn et al. (2008), obliquus extract diminished the survival rate of HepG2 cells in a manner that was dose-dependent. It also arrested cancer cells in the G<sub>0</sub> and G<sub>1</sub> phase and triggered apoptotic cell death by reducing the expression of p53, cyclins D1, Cdk2, Cdk4, and Cdk6. A 10-day intraperitoneal infusion of 20 mg/kg/day aqueous extracts from *I. obliquus* reduced tumour expansion in Balb/c mice inserted with B16-F10 melanoma cells (Youn et al., 2009). This was demonstrated by a decrease in the expression of pRb, p53, and p27, as well as an arrest in tumour cell proliferation at the G<sub>0</sub> and G<sub>1</sub> phase. According to research carried out by Lee et al. in 2014, polysaccharides derived from *I. obliquus* were able to inhibit the phosphorylation of MAPKs, PI3K, and AKT as well as lower the levels of expression of MMP-2, MMP-9, COX-2, and NF-B inside human A549 nonsmall lung cancer cells.

#### Immunomodulatory activity

Oral therapy using a hot-water extract from *I. obliquus* at a dose of 100 mg/kg led to a decrease in serum levels of IgE and IgG2a, as well as a suppression of IL-4 and an increase in IFN- in concanavalin-stimulated lymphocytes (Ko et al. 2011). In a study conducted by Harikrishna et al. (2012), the nutritional intervention of the olive flounder fish, *Paralichthys olivaceus*, with *I. obliquus* extract resulted in improved lysozyme, complement, and myeloperoxidase activity as well as higher resistance to *Uronemamarinum* pathogens. Oral administration to hens with *I. obliquus* products of fermentation during vaccination resulted in elevated mononuclear cells in the peripheral blood and Th1/Th2 ratios, as well as a reduction in the pathogenicity of the Newcastle disease virus (Zhang et al., 2018). *I. obliquus* lignin complexes elevated NO production, promoted phagocytic activity, and scavenged 1,1-diphenyl-2-picrylhydrazyl (DPPH) radicals in RAW 264.7 macrophages, according to research published by Niu et al. in 2016.

#### Antioxidant Activity

In mice with scopolamine-induced memory loss, a methanolic extract of *I. obliquus* increased levels of antioxidants (SOD and GSH) while decreasing levels of nitrite (the proportion of GSSG to GSH), MDA, and GSSG/GSH. This research was published in 2011 by Giridharan and colleagues. According to research by Nakajima et al. (2009), the *I. obliquus* compound 3,4-dihydroxybenzalacetone prevented damage from oxidation caused by H<sub>2</sub>O<sub>2</sub> in PC12 cells. In the hepatocyte tissue of alloxan-induced diabetic male Kunming mice, *I. obliquus* ethyl acetate extract scavenged DPPH-free radicals, lowered MDA, and raised GSH, according to research published by Lu et al. in 2010. According to Wang et al. (2017), IOP-chromium (III) complex treatment in T2DM male Kunming mice lowered MDA levels and enhanced hepatic enzymes that protect cells (SOD, catalase, and GSH-Px).

#### Antiinflammatory activity

*Inonotus obliquus* extracts suppressed proinflammatory mediators (NO, PGE<sub>2</sub>, iNOS, COX-2, TNF- $\alpha$ , IL-1 $\beta$ , and IL-6) in cells (RAW 264.7) triggered by lipopolysaccharide (Debnath et al. 2013). In this order, Kim et al. reported significant inhibition of Akt, I $\kappa$ B $\alpha$ , and MAPK in lipopolysaccharide-activated macrophages by 70% ethanolic extract of *I. obliquus* (Kim et al. 2007). NF-B and transforming growth factor expression were reduced, and renal tubular cells (LLC-PK1) were protected from streptozotocin (STZ) and advanced glycation end-products-induced glucotoxicity after three days of treatment with a low molecular mass polysaccharide extract of *I. obliquus* in mice with nephropathy brought on by HFD/STZ (Chou et al. 2016).



## 5.2 *Pleurotus ostreatus* (Jacq. Ex Fr.) P. Kumm.

### Antioxidant activity

Researchers have shown that the fruiting bodies of the *Pleurotus ostreatus* mushroom have the ability to chelate iron, fight free radicals, reduce power, and reduce power, respectively. According to Dubost and Beelman (2007), the antioxidant activity of extracts from *Pleurotus ostreatus* is quite low when compared to that of other mushrooms that are edible. According to Adebayo et al. (2014b) and Okafor (2017), oxidative stress is the root cause of several disorders, including cancer and hepatotoxicity. Both flavonoids and phenols are effective in preventing oxidative stress. An increase in the expression of the catalase gene and a reduction in the amount of protein oxidation caused by free radicals were discovered by Jayakumar et al. (2011) in their research on the effects of an extract of *Pleurotus ostreatus* on elderly rats.

### Antimicrobial activity

*Pleurotus ostreatus* is antibacterial and antifungal. p-anisaldehyde in *P. ostreatus* hexane-dichloromethane extract inhibits *Bacillus subtilis*, *Pseudomonas aeruginosa*, *Aspergillus niger*, and *Fusarium oxysporum*. Pleuran is in *P. ostreatus* fruiting bodies. Bacterial-susceptible mice survived -D glucan (Karacsonyi & Kuniak, 1994). According to research that was conducted by Cowan in 1999, the phenolic and tannin components of *P. ostreatus* were able to prevent cell membrane destruction, synthesis of proteins, the activity of enzymes that break down proteins, and microbial attachments.

Cells are unharmed by antiviral chemotherapy. Thus, innovative, side-effect-free antivirals that diminish viral resistance are necessary. EI-Fakharany et al. (2010) found that *P. ostreatus* laccase stopped the hepatitis C virus from becoming active again and getting into HepG2 cells in the liver and blood vessels in the peripheral zone. In this context, An HIV-1 reverse transcriptase-inhibiting ubiquitin-like protein was discovered by Wang and Ng (2000b) from *P. ostreatus* extract.

Karaman et al. (2010) said that the methanolic and chloroformic extracts of *P. ostreatus* may be antibacterial and effective against gram-positive bacteria. Mirunalini et al. assessed *P. ostreatus* inhibitory zone widths and biosynthesized AgNPs against different gram-positive bacteria in 2012. *P. ostreatus*-biosynthesized AgNPs inhibited all bacteria.

### Anti-diabetic activity

In this context, evaluated the significant anti-diabetic impact of synergistic solution of *P. ostreatus* with *Murraya koenigii* on the level of blood glucose in insulin-dependent and insulin-independent diabetics. In case of, hyperglycemic rats, *P. ostreatus* extract lowers blood glucose less than amaryl (Krishna and Usha, 2009).

### Anti-hypercholesterolic activity

*P. ostreatus* includes ergothioneine, lovastatin, and chrysin, which prevent and treat atherosclerosis, according to Mohamad et al. (2017). In normal Wistar male rats, *P. ostreatus*' ethanolic extract reduced hyperlipidemia. Alam et al. (2009) fed hypercholesterolic rats 5% *P. ostreatus* powder, which lowered TC by 37% and TG by 46%. Lovastatin in this mushroom caused this.

### Antitumor activity

Mycelial extract of *P. ostreatus* reduced the tumor growth in the both individually as well as with the combination of cyclophosphamide (Facchini et al., 2014) while this combination also reduced leucopenia (Meerovich et al., 2005). Similarly, aqueous mycelial extract caused human cancer cell apoptosis greater than other mushroom extracts (Sarangi et al., 2006; Saat et al., 2019). Wang et al. (2000a) observed that *P. ostreatus* lectin suppressed sarcoma and hepatoma cell proliferation and increased mice lifetime. In an another study proteoglycans identified in aqueous mycelial extract prevented sarcoma in mice (Sarangi et al., 2006).

### Immunomodulatory activity

Oyster mushrooms' immuno-modulatory qualities and low cytotoxicity could aid radiation and chemotherapy patients by enhancing immune resistance and minimizing toxicity (El-Enshasy & Hatti-Kaul, 2013). Wang et al. (2000a) identified many immune-modulating components (lectins, polysaccharides, polysaccharide-peptides, and polysaccharide-protein complexes) from *P. ostreatus*. The aqueous extract obtained from *P. ostreatus* fruit flesh and mycelia increases neutrophil ROS production and modulates all immunologically competent cells.

### 5.3 *Trametes versicolor* (L.) Lloyd

#### Antitumor and immune-modulating action

Polysaccharopeptide inhibits tumour formation and metastatic activity in experimental models in preclinical investigations (Xu et al. 2008). Hepatocellular malignancy, nasal cavity cancer, sarcoma, breast atypical swelling, Ehrlich ascites tumour, cancer of the breast, ascetic hepatoma, and malignancy are among the conditions that polysaccharopeptide cures (Lin, 2014). Polysaccharopeptide usually inhibits more than other treatment groups. The standard cancer therapy medicine inhibits more than polysaccharopeptide; however, the combined drug treatment has fewer side effects. Polysaccharopeptide boosts immunity (Zheng, 2004) and prevents mammary gland epithelial cell hyperplasia (Wu et al., 2001). Polysaccharopeptides with cyclophosphamide improve immunity, anticancer effects, and side effects (Hu et al., 1988).

#### Anti-hyperlipidemia activity

Hyperlipidemia is characterized by high serum lipids and lipoproteins, generally cholesterol, triglycerides, and low-density lipoproteins. Statins and fibrates are clinically utilized anti-hyperlipidemia medicines (Liu and Li, 2004). Statins and fibrates increase the risk of myopathy and rhabdomyolysis. Ezetimibe, sequestrants, and niacin of bile acids are often found in statins. PSP decreased cholesterol levels in 240 patients (Rao et al., 2007). In a rodent high cholesterol model, PSP consistently lowers the level of complete cholesterol, triglycerides, and LDL while increases the HDL. Published data show that PSP lowers serum lipids without harming liver or kidney function, suggesting It could be a secure and successful lipid lowering drug.

#### Effects against chronic bronchitis

Aging reduces respiratory defenses. Thus, the elderly can develop chronic bronchitis (Zheng, 2004). Chronic bronchitis is characterized by coughing, phlegm, and dyspnea (Hong, 1986). Immunological laboratory tests and a cough scavenging assay demonstrated that PSP improves innate immunity and lowers cough, phlegm, and dyspnea in treated patients by 89.3%. Polysaccharopeptide is also safe and non-toxic.

### 5.4 *Ganoderma applanatum* (Pers.) Pat.

#### Anticancer Activity

The whole metabolite extract showed  $160 \pm 4.08$   $\mu\text{g/ml}$  IC<sub>50</sub> against Caco-2 human colon cancer cells. *G. applanatum* methanolic extract altered cell morphology and elevated glutathione. At 80 $\mu\text{g/ml}$  and 160  $\mu\text{g/ml}$ , *Ganoderma applanatum* significantly increased Bax and Bcl-2 percentages on the Caco-2 through an irreversible p53-independent route while reducing p53 expression of proteins and increasing Cas-3 transcript. *G. applanatum* (200  $\mu\text{g/ml}$ ) therapy for five days reduced solid Ehrlich tumor (SEC) volume in vivo. mRNA Bax/Bcl-2 and p53/Cas-3 upregulation verified the apoptotic p53-dependent mechanism. In conclusion, *G. applanatum* could induce apoptosis in Caco-2 and SEC via p53-independent and p53-dependent pathways. *G. applanatum* showed promise as an alternate or co-anticancer drug (Elkhateeb et al. 2018).

#### Hepatoprotective activity

This study looked at the possible mechanisms behind the protective effects of a triterpanoid made from *G. applanatum* alongside inflammatory and oxidative stress caused by benzo(a)pyrene (BaP) in the livers of mice. This triterpanoid prevented BaP-induced increases in mouse hepatic antioxidant levels (Cu/Zn-SOD, CAT, GPx, and GST) as well as drastically decreased liver ROS, MDA, and GSH/GSSG ratios. Triterpanoid successfully decreased hepatic inflammation by decreasing proinflammatory cytokines (IL-1, COX-2, and NF-B translocation) in BaP-treated rats, as shown by Western blot analysis (Ma et al. 2011).

#### Spleen cell proliferation activity

In vitro, *G. applanatum* polysaccharides increased the growth of mouse spleen cells. High concentrations of these polysaccharides (125–500 mg/mL) boosted the IL-2 and IFN-g production of ConA-stimulated mouse spleen cells. When triggered by 5 mg/mL ConA, normal mice administered *G. applanatum* polysaccharides (20 mg/kg/d for 10 days, s.c.) produced more IL-2 and IFN-g and enhanced spleen cell proliferation (Guo and Yen, 1991). Initial immune responses of spleen cells to sheep RBCs were enhanced. NK activity, IL-2, and IFN-g production in the spleen cells were restored after *G. applanatum* polysaccharides (20 mg/kg/d for 15 days, s.c.) administration to mice with sarcoma transplants for 15 days straight (Guo and Yen, 1991).

**Immunomodulatory effect**

According to Zhang (1989), the *G. applanatum* extract had a 60% inhibiting effect on the formation of HbsAg in PLC/PRF/5 cells.

**5.5 *Agaricus campestris*****Antioxidant activity**

This study examines bioassay-guided fractionation's antioxidant activity and active compounds and components. Fractions yielded three chemicals including methoxydiazine, afrormosin, and candidol (AC-1–3). They had high antioxidant activity at 100, 200, 300, 400, and 500 µg/mL (Reddy et al. 2023).

**Antiinflammatory activity**

The findings demonstrated that both alcoholic extracts of *Agaricus campestris* significantly ( $p < 0.005$ ) inhibit hyaluronidase. The alcoholic extract of *Agaricus campestris* has significantly ( $p < 0.005$ ) less hyaluronidase inhibition activity than standard diclofenac. During inflammation, hyaluronidase activity in the blood increases, and when inflammation subsides, hyaluronidase activity decreases. According to the above findings, *Agaricus campestris* has anti-inflammatory activity via hyaluronidase inhibition.

**Hepatoprotective activity**

AEAC ensured the auxiliary uprightness of the hepatocellular film in a subordinate measuring manner as a part of the investigation. This was in accordance with the pledge supplied by Silymarin (100 mg kg<sup>-1</sup> b.w.t.; p.o.), a well-known hepatoprotective specialist. This metabolite causes glutathione (GSH) to be depleted, which makes it possible for cells to pass through. It is evident that the AEAC concentrate brought all significant levels of AST, ALT, ALP, and total blood bilirubin back down to normal levels. This suggests that plasma layer adjustment and hepatotoxin-induced hepatic tissue healing have been impacted.

**6. CONCLUSIONS AND FUTURE OUTLOOK**

Fungi have been highly prized for their many health benefits since ancient times, and not only in the context of culinary use. The scientific study of mushrooms has gained popularity in recent years, and the results so far have been promising. Polysaccharides, sterols, and phenolic compounds are only a few of the many bioactive substances found in mushrooms. Immunomodulators, anti-inflammatories, cancer fighters, antioxidant powerhouses, and cardiovascular health boosters are just some of the health benefits linked to these compounds.

Mushrooms have promising future uses in both the medical and culinary fields. More investigation into the beneficial impacts of these versatile fungi on human health is likely to reveal even more. Mushroom cultivation is also a growing economic sector and a sustainable kind of agriculture. An expanding field of study is the creation of novel products based on mushrooms. These include powders, extracts, and beverages. The healthy nutrients present in mushrooms are concentrated in these products, making them an easy-to-use and useful option. There is a growing interest in spreading mushroom farming to new places, and there are many diverse settings that are favourable for the production of mushrooms. This could help people all throughout the world gain access to high-quality mushrooms at more affordable prices.

The outlook for the development of the above-documented edible and medicinal mushrooms is generally optimistic. The phytochemistry and pharmacological sections of this review provided key information to conduct more research to develop new products for human health.

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