

# Diversity of metabolites produced by White Rot Fungi possess different biological properties: A review

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## ABSTRACT

White rot fungi is a well known wood decaying macro-fungi being studied for many years for the potential applications in the bio-degradation of environmental pollutants because they produces a varieties of enzyme primarily responsible for the degradation of complex organic polymers. Along with these white rot fungi produces a vast number of cellular and extracellular metabolites having significant biological properties like antimicrobial, antioxidant, immunomodulatory, and nutritional properties. Because of these properties white rot fungi are being used in many field of science like Environmental Biotechnology, Bio-remediation technology, dye decolourization technology, Pharmaceutical industries, Nutritional researches etc. Aim of the study is to find out the different types of metabolites they produce, in which bio-synthetic pathways and their significant biological properties.

*Key words:* White rot fungi, Wood decay, Enzymes, Metabolites, Pathways, Biotechnology.

## Introduction

White rot fungi belongs to a large group of phylum Basidiomycota that contains nearly 30-32% of entire fungal diversity and comprising about 30,000 different species having cosmopolitan distribution in nature and involved in a varieties of relationships like, pathogenic, mutualistic, parasitism and symbiosis etc (Riley *et al.*, 2014; Webster and Weber, 2007). They are heterogenous group of saprophytic Fungi can grow everywhere on different organic substrates like lignocellulosic polymers of wood, waste organic materials, dead organisms and other biological substrates (Tuomela and Hatakka, 2019, Goyal *et al.*, 2016; Magan, 2008).

In a forest ecosystem, the saprotrophic wood decomposing fungi plays an important role in Earth's carbon cycle (Hatakka and Hammel, 2011). Certain basidiomycetes, specially the white-rot fungal

group, have the ability of degrading complex cell wall polymers of plants such as cellulose, hemicellulose and lignin and other organic materials including carbohydrates, polysaccharides and other biopolymeric substances thus release carbon dioxide and water into the environment that plays an significant role in Earth's ecosystem (Goodell *et al.*, 2020; Fukushima, 2020; Baldrian, 2008; Hatakka and Hammel, 2011). This ability due to production of a varieties of different cellular and extracellular enzymes like hydrolytic enzymes, cellulases, hemicellulose, laccases, peroxidases and others (Wang *et al.*, 2018; Glazunova *et al.*, 2018). As a result of degradation wood became pale and white spongy fibres and cause rot on wood (Abdel *et al.*, 2013; Gao *et al.*, 2010). Around 90% of wood rotting fungi are reported to white rot types and they are the only organism can efficiently degrade and mineralize the complex polymer lignin (Tuomela and Hatakka,

2019; Fernando *et al.*, 2016; Hatakka and Hammel, 2011). Based on the nature of action of wood degradation a group of basidiomycetes have been classified into brown rot and white rot and sometimes cause soft rot where, brown rot and soft rot fungi are responsible for degrading cellulosic and hemicellulosic materials (Yilikal, 2015; Schu and Anke, 2009) and white rot fungi is responsible for the degradation of lignin by producing extracellular enzyme lignin peroxidase (Sevindik, 2018).

Beside these white rot fungi possess a wide range of biological properties like they can be poisonous, edible with degrading, nutritional and therapeutic properties (Ukwuru *et al.*, 2018; Remya *et al.*, 2019; Swann and Hibbett, 2007). Some white rot basidiomycetes are known for producing fruiting bodies called as mushrooms (Swann and Hibbett, 2007). Since past centuries mushroom is being considered as functional food as they are contains nutrition rich compounds such as lipids, proteins, fat, carbohydrates, fibres, vitamins and minerals (Sevindik 2018). The mycelium and fruiting body consists of a wide range of metabolites with biologically active compounds that helps them to survive in nature by competing with other organisms for colonization, growth and nutritions (Wang *et al.*, 2018 ). A large no of enzymes along with others secondary metabolites are produced by white rot fungi have high therapeutic, nutritional, medicinal and degradation capabilities (Gebreyohannes *et al.*, 2019; Chowdhury *et al.*, 2015; Rodriguez, 2017). Those bioactive compounds such as cellular and extracellular components like enzymes and other metabolites are different in molecular weight such as phenolic compounds, high molecular weight polysaccharides, lipids, proteins, glycoproteins and tocopherols have shown bioactive properties in some isolated higher macro fungi. While secondary metabolites such as terpenoids, polyphenols, sterols, flavonoids, alkaloids, benzoic acid derivatives, quinolones, anthraquinones, lactones having bioactive properties mentioned by authors (Jaszek *et al.*, 2013; Srikrum and Supapvanich, 2016; Bains and Tripathi, 2016). White rot fungal metabolites having potential industrial, agricultural, environmental and biotechnological applications, in pharmaceutical industries, pulp industries such as bioleaching of pulp, in paper industries, waste water treatment, degradation and transformation of environmental pollutants, improvement of cellulose and lignin digestibility in animal, production of renewable raw materials from

lignocellulosic materials, in bioremediation and bioaugmentation technology, agricultural wastes revalorization etc (De Jong, 1993; Wang *et al.*, 2017; Rodriguez, 2017; Jurado *et al.*, 2011).

The purpose of the study was to find out different types of bioactive metabolites produced by white rot fungi during their growth phase and developments, in which biosynthetic pathways the metabolites are produced and their biological properties including their degradation capabilities, antimicrobial properties, nutraceutical and health benefits. Various literatures data showed that the metabolites produced by white rot fungi having a lot of contributions in researches, and in biotechnological, Industrial and environmental applications.

### **Diversity of metabolites produced by white rot fungi**

Metabolism is a biochemical process by which all the life processes are accomplished and it's a fundamental process for all cellular life. White Rot Fungi are one of the most beneficial organism on Earth due to the abilities of synthesizing numerous number of bioactive metabolites for survival in environment (Jaszek *et al.*, 2013; Rodriguez, 2017). Mycelial structure and the fruiting bodies produces a vast number of metabolites having different biological properties (Schu and Anke, 2009, Xiao and Zhong, 2016, Wang *et al.*, 2017). Their metabolites can be classified into two categories are primary and secondary metabolites.

#### **Primary metabolites**

The primary metabolites of white rot fungi are produced by different cellular metabolisms during the exponential growth phase that helps in their growth and development (Sanchez and Demain, 2008). The end products of metabolism are used by the cells as building blocks for the production of potential biomolecules are like amino acids and peptides such as lysine, methionine, phenyl alanine, tyrosine and repeated amino acid chain of leucine, valine (Lapadatescu *et al.*, 2000; Beck, 1997; Stergiopoulos *et al.*, 2013; Rahi and Malik, 2016), Some monomers or the subunits of nucleosides or nucleic acid derivatives like Eritadenine, Lentinacin and Clitocine were isolated from the white rot fungi species of *Lentinus edodes* and *Clitocybe inversa* (Fukushima, 2020, Rahi and Malik, 2016).

Organic acids such as oxalic acid, acetic acid, citric acid and malonic acids, fumaric acid, shikimic

acid etc (Dashtban *et al.*, 2010), carbohydrates and polysaccharides such as glucose, xylose, galactose, mannose and arabinose etc (Hatakka and Hammel, 2011). Vitamins a primary metabolite is very much needed for fungal growth and developments such as vitamin B12 or riboflavin, vitamin B1 or thiamine, niacin, biotin, ascorbic acid, pyridoxine, tocopherols etc have been reported to present in several white rot fungal mushroom (Ukwuru *et al.*, 2018). The secretion of extracellular and hydrolytic enzymes such as pectinase, chitinase, cellulase, laccase, xylanase, manganese peroxidase (Tuomela and Hatakka, 2019; Abdel *et al.*, 2013) lignin peroxidases enzymes during their secondary metabolism while degradation of components polymer (Alves *et al.*, 2012 a, Fonseca *et al.*, 2015; Shu, 2007; Alves *et al.*, 2013b, Rodriguez, 2017), some alcohols such as ethanol, benzyl alcohol and vanillyl alcohol (Lapadatescu *et al.*, 2000; Dashtban *et al.*, 2010; Mattila *et al.*, 2020). It has been reported that various enzymes are involved in the degradations of complex polymers that results in the formation of biproducts for cellular metabolism (Goyal *et al.*, 2016; Bezalel *et al.*, 1996; Agrawal *et al.*, 2016).

### Secondary Metabolites

The synthesis of secondary metabolites basically involves with the polymerization of certain enzymes those referred to as the backbone of primary metabolisms and then secondary metabolites are produced by using the enzymes or intermediate by-products of primary metabolisms (Keller *et al.*, 2005). Though secondary metabolites are not very much essential for growth and development but they possess different biological properties such as antimicrobial, anti-parasites, insecticides, microbial interactions, such as competition and symbiosis, production of important enzymes for biodegradations (Schu and Anke, 2009; Thirumurugan *et al.*, 2018; Verduzco and Gutierrez, 2020). Some important types of secondary metabolites are Terpenoids such as Taxol, lanosterol, pleuromutilin and illudin, ganoderic acid, hirsutane and ergosterols etc (Schu and Anke, 2009; Kuhnert *et al.*, 2018; Rahi and Malik, 2016). Phenols and phenolic compounds such as flavonoids, quinones, tannins, cinnamic acids, coumarins, gallic acid, p-hydroxybenzoic, ferulic acid, syringic acid, Tyrosol and hydroquinone (Gan *et al.*, 2019; Jong, 2010, Osinka *et al.*, 2015). Polyketides, Alkaloids and fatty acid such as sterenins, orsellinic acid, Pestalochloride,

Terphenyl, and Strobilurin (Jong, 2010; Tian *et al.*, 2020; Nofiani *et al.*, 2018), non-ribosomal peptides etc (Keller *et al.*, 2005). Carbohydrates and Polysaccharides such as 5-d-glucans, glucose, mannose, galactose (Hatakka and Hammel, 2011; Keller *et al.*, 2005; Wang *et al.*, 2017). Proteins and Glycoproteins such as lectins, pectin, heme peroxidase like manganese, lignin and versatile peroxidase, anthraquinones etc (Schu and Anke, 2009; Jaszek *et al.*, 2013; Thirumurugan *et al.*, 2018; Dashtban *et al.*, 2010; Jong, 2010). Veratryl alcohol is also a secondary metabolite that produce lignin peroxidase important enzymes for lignin polymer degradations (Jensen *et al.*, 1994). Several colorless fungal secondary metabolites are considered to be the pigments The genus *Thelephora* and *Hypoxyylon* has shown to be rich sources of pigment p-terphenyl (Chen and Liu, 2017) on the other hand an extracellular metabolite melanin pigment is produced from white rot fungus *Schizophyllum commune* (Arun and Gunasekaran, 2015).

### Metabolic biosynthesis

Fungi are the most ubiquitous organism in nature as they are capable to synthesize broad range of compounds naturally from simple molecules which referred to as primary metabolites to the complex molecules are called secondary metabolites (Rahi and Malik, 2016). Although the commercial production of fungal metabolites depends on the amount of nutrients, its growth environments and culture conditions that promotes metabolite biosynthesis (Bills *et al.*, 2008, Moreira *et al.*, 2003; Barberel and Walker, 2000). Different chemicals or substrates are generally used to transform the actions of cellular enzymes by the organisms for the production of bioactive metabolites that might be used as building blocks for the production of secondary metabolites (Nielsen, 2018, Bezalel *et al.*, 1996; Agrawal *et al.*, 2016). Naturally so many cellular and extracellular enzymes are involved in the degradations of various compounds or polymers in the environment by cellular metabolisms results in the formations of cellular biproducts (Goyal *et al.*, 2016; Bezalel *et al.*, 1996; Agrawal *et al.*, 2016). Secondary metabolites are produced by using those biproducts of primary metabolism as building blocks (Nord, 2014) and are well known bioactive compounds with structural diversities are synthesized in various metabolic pathways with the actions of various enzymes or intermediates (Goyal *et al.*, 2016; Lin *et al.*, 2019).

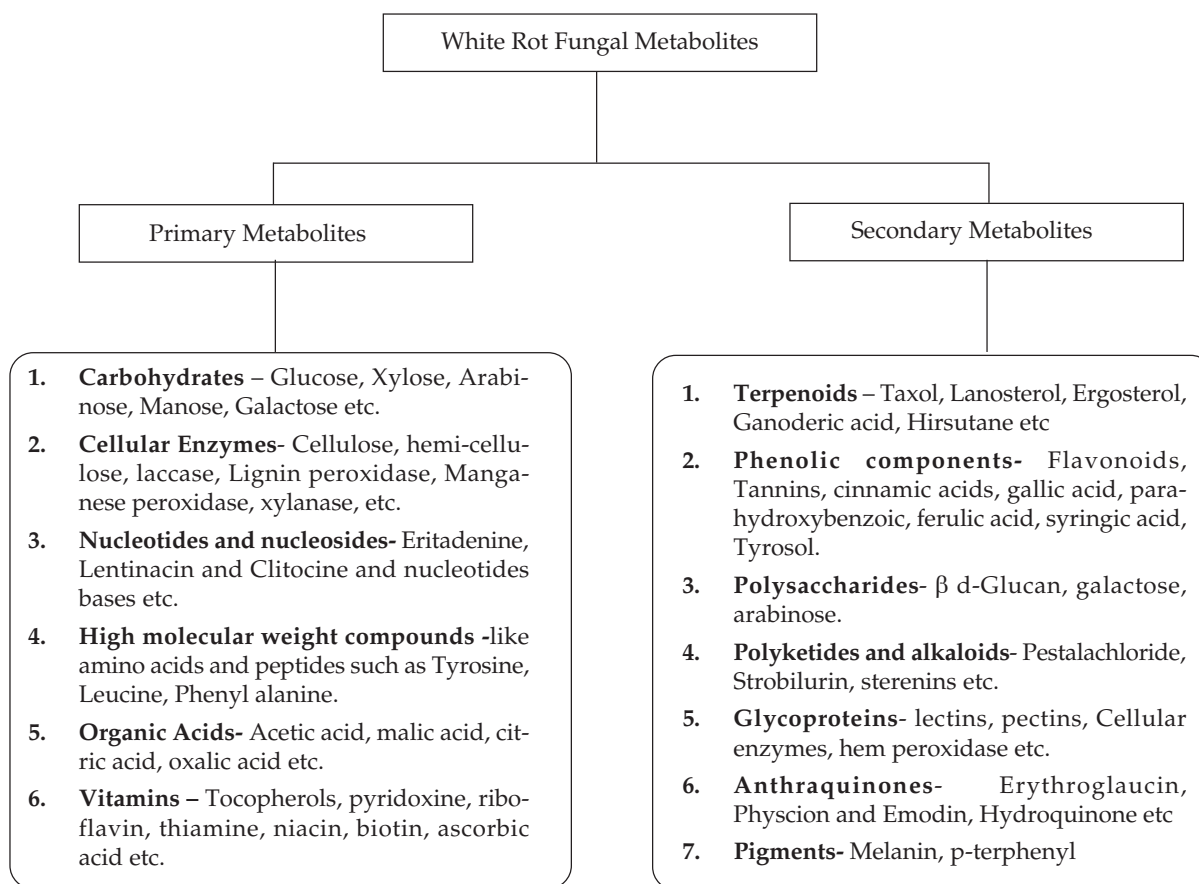


Fig. 1. Types of metabolic components produced by White Rot fungi

## Biosynthetic pathways are involved like

### i. Mevalonic acid pathway

It is a common pathway of many plants and including fungi (Hansson, 2013). Several class of terpenoids like sesquiterpenoids, diterpenoids and triterpenoids, steroids secondary metabolites of white rot fungi are synthesized by following the mevalonic acid pathway by integration of primary and secondary metabolic products (Goyal *et al.*, 2016; Kaller *et al.*, 2005; Thirumurugan *et al.*, 2018). White rot fungi mostly follows this mevalonic acid pathway for the synthesis of several classes of terpenoids and its derivatives (Hansson, 2013).

### ii. Shikimic acid pathway

Shikimic acid pathway is a biosynthesis pathway for the production of various aromatic amino acids and organic compounds like L-phenylalanine, L-tryptophan and L-tyrosine, glycine, various proteins

and sterols etc (Goyal *et al.*, 2016; Moore *et al.*, 2020). The shikimic acid pathway involved with the carbohydrate metabolisms and pentose phosphate pathway to produce shikimic acids and other by products (Hansson, 2013). Shikimic acid phosphorylated with a Phosphoenol pyruvate molecules an important compounds of Krebs cycle and attached to another group of shikimic acid and helps in the formation of many other important compounds such as aromatic amino acids such as Benzoic acid derivatives and phenolic compounds are originated from this pathway and tannic acid is formed by the reaction of glucose and shikimic acids following the shikimic acid pathway (Goyal *et al.*, 2016; Kojima *et al.*, 2015; Thirumurugan *et al.*, 2018; Hansson 2013; Nowicka and Kruk, 2010).

### iii. Acetic acid pathway

The Fungal polyketides are generally synthesized by polyketide synthases an enzyme is composed of

a multi domain proteins and non-ribosomal peptide synthase following by the polyketide and non-ribosomal peptide synthesis pathways (Kaller *et al.*, 2005; Nielsen, 2018; Valiante, 2017). These two enzymes contain short chain of carboxylic acids such as acetyl CoA and malonyl Co-A are two important by-products of Krebs cycle or Tri carboxylic Acid cycle are involved in polyketide and fatty acid biosynthesis those are produced by the acetic acid pathway and  $\beta$  oxidation of fatty acids (Moore *et al.*, 2020; Kaller *et al.*, 2005; Collemare and Seidl, 2019). Polyketide synthase simulates other intermolecular reactions where anthraquinones are synthesised in this pathway (Hansson, 2013; Nowicka and Kruk, 2010).

#### iv. Biosynthesis of carbohydrates and polysaccharides

The biosynthetic pathway of carbohydrates are common in all fungi in compared with the other organisms (Deveau *et al.*, 2008). Polysaccharides are composed of long chain of carbohydrate molecules joined together by galactosidase bonds are essential components for all living organisms produced by both primary and secondary metabolism (Kaller *et al.*, 2005; Hansson, 2013). Examples of carbohydrates and polysaccharides are glucose, mannose, galactose and glucan are produced via several reverse pathways are like glycolytic, gluconeogenesis, pentose phosphate pathway and krebs cycle from non-carbohydrate sources (Deveau *et al.*, 2008; Deacon, 2013).

#### Biological properties of white rot fungal metabolites

White rot fungi is one of the most beneficial organism on earth easily available in nature as they grows naturally in the environment produces unlimited no of metabolites and compounds possess different varieties of bioactive properties, some wild white rot basidiomycetes are not edible because of the presence of toxins might cause harm to human health and on the other hand wild edible white rot fungi has an importance in every aspects of life, from their nutritional properties to their potential applications in various field of science has increased based on their biological properties (Table 1).

#### Degradation properties

White rot fungi is a type of fungi having the capability to degrade and mineralize the most abundant

complex polymer lignin of wood or plants along with other lignocellulosic biomass and various organic pollutants that could be a possible way to degrade a wide varieties of environmental pollutants and chemicals that are insoluble and very recalcitrant in nature are easily mineralized by that fungi (Aust, 1995; Shah *et al.*, 1992; Abdel *et al.*, 2013). Although the capability of Lignin degradation by white rot fungi is being studied from last 30 years and it widely applied for its biotechnical applications in pulp mills and paper industries, bleaching processes, dyes decolorization in textile industries for removal of environmental chemicals and other xenobiotic pollutants (Tuomela and Hatakka, 2019; Aust 1995; Boer, 2018; Jaszek *et al.*, 2006; Zuleta *et al.*, 2016). The xenobiotic compounds such as some organochlorinated and other pesticides with diverse chemical classes, some of important chemicals are diuron, Linuron, metalaxyl, lindane, metribuzin, chlorpyrifos, atrazine or terbuthylazine, DDT or 1,1,1-trichloro-2,2-bis (4-chlorophenyl)ethane, TCDD or 2,3,7,8- tetrachlorodibenzo-p-dioxin, benzo (a) pyrene, polyaromatic hydrocarbons or PAHs, polychlorinated biphenyls, pentachlorophenol and various groups of pesticides (Magan 2008; Aust, 1995, Shah *et al.*, 1992; Bending *et al.*, 2002; Mohapatra *et al.*, 2018; Gouma *et al.*, 2019).

Their ability of degradation might be for the action of producing extracellular non-specific oxidative and hydrolytic enzymes during metabolisms that composed of cellulase, hemicellulose, lignin peroxidase, manganese peroxidase and laccase that allows the organisms to mineralize the cellulosic polymers, lignin and xenobiotics pollutants persistent in soil and water in the environment (Tuomela and Hatakka, 2019; Jaszek *et al.*, 2013; Chaturvedi *et al.*, 2018; Dewick, 2009). The ligninolytic system of white rot fungi have a great complexity involved with the varieties of non-specific enzymes those cleaves the carbon-carbon and carbon-oxygen bonds in the lignocellulosic recalcitrant polymers results in the depolymerisation of lignin and thus degrades varieties of aromatic and aliphatic groups containing xenobiotic compounds (Tuomela and Hatakka, 2019; Magan, 2008).

The most unique feature of the fungi are, they occurs naturally having the ability of degradation with the degradative peroxidase enzymes are produced in response to differences in carbon nitrogen ratio or the presence of organic polymers, xenobiotics and heavy metals through various

metabolic pathways in natural or artificial environment (Tuomela and Hatakka, 2019; Shah *et al.*, 1992).

### Antioxidant properties

At present scenario unhealthy food habits, hypertension, climate conditions, ultraviolet radiations and other environmental factors responsible for oxidative damage or stress, free radicals, and numerous health diseases (Roman *et al.*, 2020) Numerous no of bioactive properties of secondary metabolites produced by Fungi have been described extensively in many studies with their antioxidant properties also reported is an important features could be used as natural source antioxidants where now a days doctors are suggesting natural food sources those posses antioxidants to prevent oxidative damages in the body (Fernando *et al.*, 2016). Oxidative stress is a condition of damage that is a vital problem for human and all living organisms including is basically caused by the uncontrolled formations of reactive oxygen species or ROS including free radicles having one or more unpaired electrons are generated during normal cellular metabolism that consequently exposed to stress factors, such as changes in temperature, environmental pollution, change in climates (Fernando *et al.*, 2016; Jaszek *et al.*, 2013; Rezaeian *et al.*, 2016).

An uncontrolled increase of level of free radicals can cause oxidative damage to cellular components including DNA, proteins, membrane lipids that leads to serious degenerative human diseases such as coronary heart disease, impaired immune functions, aging, cancer, cardiovascular disease, brain disfunctions and atherosclerosis (Fernando *et al.*, 2016; Jaszek *et al.*, 2013; Jain and Choudhary, 2012). Various authors have done antioxidant activity of different white rot fungi and other higher fungi and reported that the secondary metabolites produced from white rot basidiomycetes posses strong antioxidative mechanism that can protect mammalian cells against free radicals mediated toxicity generated in human body (Fernando *et al.*, 2016; Jaszek *et al.*, 2013, Rezaeian *et al.*, 2016).

### Antimicrobial properties

Several antimicrobial screening has been done to know about antimicrobial properties of white rot fungal metabolites and a large number of bioactive components both cellular, extracellular components and secondary metabolites have shown to be effective

against pathogenic microorganisms that could be used to treat variety of disease (Wang *et al.*, 2018; Jaszek *et al.*, 2013). From literature studies some bioactive compounds have been identified, including cellular and extracellular low and high molecular weight compounds are mainly secondary metabolites like some lipids, glycoproteins, peptides, terpenes, carbohydrates, pigment such as melanin (Arun and Gunasekaran, 2015), polysaccharides, alkaloids, sterols, polyphenols (Jaszek *et al.*, 2013) and flavonoid derivatives, anthraquinones, benzoic acid derivatives, and quinolines responsible for showing antimicrobial activity from different extracts of white rot basidiomycetes against both gram-positive and gram negative pathogenic bacteria (Wang *et al.*, 2018; Sharma *et al.*, 2017; Sevindik, 2018; Akyuz *et al.*, 2010; Jaszek *et al.*, 2013).

### Nutraceutical benefits

Some edible white rot fungi are able to produces fruiting bodies also referred as mushroom that is being consumed as prominent food from several centuries in many countries traditionally called as "Food of the Gods" to ancient Greeks because the fruiting bodies of edible white rot fungi or mushrooms were very good in taste with aroma and flavours (Rahi and Malik, 2016; Jennison *et al.*, 1957; Kandeler, 2007). As many as researches are being conducted with mushrooms and macro fungi several informations are being reported in literatures. Scientists have reported that all edible mushrooms including white rot fungi contains a large number of bioactive components are very much beneficial to human health (Rezaeian *et al.*, 2016; Dewick, 2009). Some white rot fungal species are *Agaricus campestris*, *Lentinula edodes*, *Cyclocybe aegerita*, *Pleurotus* sp (Yogita *et al.*, 2010) are composed of high content of carbon and nitrogenous components, dietary fibers, water composition, low fat and low-calorie protein content including the essential amino acids along with numerous no of minerals and vitamins B12 or riboflavin, B3, B5 and vitamin C and D, Copper, Selenium, Potassium, carbohydrates and nitrogen rich compounds, lipids etc (Ukwuru *et al.*, 2018, Remya *et al.*, 2019; Rahi and Malik, 2016; Xiao and Zhong, 2016, Yogita *et al.*, 2010). The bioactive compounds polyphenols and flavonoid components produced by some white rot fungal species are used as a functional food source having nutraceutical benefits with dietary components (Fernando *et al.*, 2016).

**Table 1.** Bioactive metabolites produced by white rot fungi and their biological properties:

Diversity of Metabolites	White Rot Fungal Species	Significant Features	References
1. Nucleosides and nucleotide derivatives	<i>Lentinus edodes</i> <i>Clitocybe inversa</i>	Nucleosides and nucleotide derivatives serves the energy within the cell in the form of the nucleoside triphosphates like ATP, GTP, CTP, and UTP and also play central roles in cellular metabolisms.	Fukushima, 2020; Rahi and Malik, 2016
2. Laccase and laccase type of enzyme named phenol oxidases	<i>Crassostrea gigas</i> <i>Cerrena unicolor</i>	Antimicrobial activities and immune defence mechanisms Degradation of lignin polymers, Antioxidative properties	Tuomela and Hatakka, 2019; Luna <i>et al.</i> , 2011 Rodríguez, 2017
3. Lignin peroxidase, Manganese peroxidase and other enzymes	<i>Trametes versicolor</i> <i>Phanerochaete chrysosporium</i> <i>Pleurotus ostreatus</i> <i>Pleurotus sajor-caju</i> <i>Ganoderma lucidum</i> <i>Trametes sp</i>	Degradation of lignin, cellulose and hemicellulose Degradation of xenobiotics including pesticides, aromatic hydrocarbons, Industrial dyes decolourization, Acts as antioxidants.	Tuomela and Hatakka, 2019; Jaszek <i>et al.</i> , 2013; Koroleva <i>et al.</i> , 2015, Rodríguez, 2017
4. Phenolic compounds and Flavonoids	<i>Pestalotiopsis sp.</i> <i>Stereum sp.</i> <i>Ganoderma lucidum</i> <i>Phellinus linteus</i> <i>Agaricus campestris</i> <i>Lentinus edodes</i> <i>Agaricus blazei</i> <i>Pleurotus ostreatus</i> <i>Flavodon flavus</i> <i>Xylaria feejeensis</i> <i>Schizophyllum sp.</i>	Antioxidant activities, Oxidative stress reduction Nutraceutical properties Antimicrobial activities against MRSA Cytotoxic and Scavenging activity in human body.	Fernando <i>et al.</i> , 2016; Ukwuru <i>et al.</i> , 2018; Sharma <i>et al.</i> , 2017; Rahi and Malik, 2016; Verduzco and Gutierrez, 2020; Tian <i>et al.</i> , 2020; Rezaeian <i>et al.</i> , 2016 Jain and Choudhary, 2012
5. Veratryl alcohol	<i>Phanerochaete chrysosporium</i>	This compound stabilizes the lignin peroxidase enzymes to promote oxidation of a variety of lignocellulosic and recalcitrant substrates, Acts as antioxidant	Jennison <i>et al.</i> , 1957; Morales <i>et al.</i> , 2010
6. Terpenoids and Several Lipids	<i>Cystoderma sp.</i> <i>Conocybe sp</i> <i>Pestalotiopsis spp</i> <i>Ganoderma lucidum</i> <i>Stereum hirsutum</i> <i>Clitocybe illudens</i> <i>Coprinus sp.</i> <i>Limacella illinita</i> <i>Fomitella fraxinea</i> <i>Clavariadelphus truncates</i> <i>Grifola frondosa</i> <i>Ganoderma lucidum</i>	Antitumor properties, Anticancer, Neurotrophic Anti-infectious and anti-inflammatory activities Cytotoxic activity Fungicidal and Antibacterial and Nematocidal activities Acts against gram positive methicillin resistant <i>staphylococcus aureus</i> or MRSA	Schu and Anke 2009, Zhong and Xiao 2009, Rahi and Malik 2016, Jong 2010; Farhat <i>et al.</i> , 2018

**Table 1.** Continued ...

Diversity of Metabolites	White Rot Fungal Species	Significant Features	References
7. Sterols	<i>Stereum hirsutum</i> <i>Lentinula edodes</i> <i>Cordyceps sinensis</i> <i>Tricholomopsis rutilans</i>	Acts against both Gram positive and negative bacteria, Antioxidant properties	Jaszek <i>et al.</i> , 2013; Rahi and Malik 2016; Kandeler 2007
8. Polyketides, Alkaloids and Fatty acid derivatives	<i>Stereum</i> sp. <i>Pestalotiopsis adusta</i> <i>Ceriporia subvermisporea</i> <i>Strobilurus tenacellus</i>	Antibacterial, Antifungal, Anticancer and Immunosuppressive properties	Schu and Anke, 2009; Jaszek <i>et al.</i> , 2013, Rahi and Malik, 2016; Jong 2010; Nofiani <i>et al.</i> , 2018
9. Glycoproteins	<i>Agaricus campestris</i> <i>Tricholoma</i> <i>Mongolicum</i> <i>Phallus</i> sp. <i>Lactarius</i> sp.	Cellular mechanisms like Cellular transport, Adhesions of cell to cell, Cellular Recognition, Cell Differentiation, Growth regulation in living organisms	Schu and Anke, 2009; Rahi and Malik, 2016; Elisashvili, 2012
10. Polysaccharides	<i>Stereum</i> sp. <i>Ganoderan</i> sp. <i>Lentinan</i> sp. <i>Grifolan</i> sp. <i>Schizophyllum commune</i> <i>Trametes villosa</i>	Anti- Cancer Anti-Tumor, Immuno-Modulatory Properties, Antiviral Activities, Antimicrobial Properties, Anti-Inflammatory, Hypoglycaemic Activities, Antioxidative, Hepatoprotective Effects	Wang <i>et al.</i> , 2018; Jaszek <i>et al.</i> , 2013, Rahi and Malik 2016, Elisashvili 2012, Rajasekar <i>et al.</i> , 2008, Osinka <i>et al.</i> , 2015
11. Anthraquinones	<i>Lentinus</i> sp. <i>Cortinarius</i> sp.	Antimicrobial and antioxidant activities.	Fernando <i>et al.</i> , 2016; Zahmatkesh <i>et al.</i> , 2018
12. Pigments	<i>Thelephora</i> sp. <i>Hypoxylon</i> sp. <i>Schizophyllum commune</i>	Nutraceuticals Antimicrobial properties	Chen and Liu 2017 Arun and Gunasekaran, 2015
13. Vitamins and minerals	<i>Pleurotus spp.</i> , <i>Lentinus edodes</i>	Vitamins regulate the growth of body. Selenium is a mineral antioxidant that prevents cell damage and reducing the risk of cancer and other diseases. Potassium is an extremely important mineral that regulates blood pressure.	Rahi and Malik, 2016

Literatures data shown that the bioactive compounds such as polysaccharides, proteins, glycoproteins, unsaturated fatty acids, phenolic compounds, tocopherols, ergosterols, lectins etc produced by white rot fungi having therapeutic properties like antidiabetic, anti-obesity, antioxidants, anticancer, antimicrobial and immunomodulatory properties, hypocholesteraemia and anti-aging along with they lowering the cholesterols, hypertension and stress factors in human body (Rahi and Malik, 2016; Xiao and Zhong, 2016; Sevindik, 2019).

Numerous no of investigations shown different bioactive molecules obtained from white rot fungi by different screening methods and the studies revealed that white rot fungi could be an alternative and natural source of bioactive compounds with therapeutic and nutraceutical benefits.

### Conclusion

From the study we have got to know about varieties of primary and secondary metabolites including



cellular enzymes white rot fungi produces during their growth and metabolisms. Moreover they are composed of varieties of nutritional components and due to the presence of those components some edible white rot fungi or mushrooms are being considered as functional food and dietary supplements. Beside nutritional properties they possess significant biological properties as they are well known macro fungi having great potential in the field of bioremediation technology and other biotechnological industrial applications, their antioxidant and antimicrobial properties. Various studies have been done with white rot fungi, some authors applied them into biotechnological applications while, other authors only focused on their antimicrobial and medicinal properties. In this review we have found different biologically active metabolites produced by white rot fungi, the metabolites produced in which biosynthetic pathways, their biodegradation capabilities towards complex organic polymers and environmental pollutants, then their nutritional properties to human health and diseases. So it's been concluded that white rot fungi not only will lead to the contribution in biotechnological applications and bioremediation technology but also could make some significant contributions in pharmaceutical industries, nutritional and biomedical research field.

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