



## Phytochemistry and Pharmacology of Santalum Album L.

\*Mr . Ajit T. Gorde

Research scholar:- b pharmacy

Ashvin college of pharmacy,manchi hill, ashvi bk

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### ABSTRACT:

Santalum album L. (Santalaceae) , commonly known as Indian sandalwood, is one of the oldest and most valuable sources of natural fragrances of great medicinal and commercial importance. P. Album has been cultivated in India for 25 centuries and is highly valued around the world as a sweet, long-lasting, fragrant oil with medicinal value. Essential oils extracted from sandalwood and sandalwood heartwood are used in various traditional medical systems such as Ayurveda, Siddha medicine, and Unani medicine for the treatment and prevention of various diseases. Sandalwood's versatile therapeutic and health-promoting effects are thought to be due to its rich source of phytochemicals, especially sesquiterpenes. By analyzing globally recognized scientific databases (Pub Med, SciFinder, Scopus, ACS, and Web of Science), recognized books, indexed and non-indexed journals, A thorough bibliographic search was conducted. Modern pharmacological studies have shown a wide range of pharmacological activities, from antibacterial to anticancer effects. Sandalwood oil and its individual ingredients have no significant toxicity. However, to understand the complete phytochemical profile of and the complex pharmacological effects of this plant, further studies on the chemical components of and their mechanisms exhibiting specific biological activities are required. The increasing commercial use of sandalwood and the low productivity of this endangered plant have raised concerns about its conservation and increased productivity with modern tools and techniques. The review discusses the traditional uses, ethnopharmacology, phytochemistry, and biological activities of sandalwood and highlights its medicinal and industrial value as well as gaps requiring future research. I'm doing it.

**KEY WORDS:** Santalum album,, Phytochemistry, Biological activity, Ethnopharmacology

### I. INTRODUCTION :

Santalum album L. Belongs to the Santalum family and is one of the most valuable trees in the world. [1] Commonly known as White Sandalwood (English), Safed Chandan (Hindi) and Sriganda (Sanskrit), this tree is considered a revered gift of the plant world that is woven into Indian culture and tradition. It has been. Sandalwood is one of the oldest fragrances with a history of more than 2,000 years, and has maintained its importance as a popular fragrance from ancient times to modern times. If we look back at history, we find that sandalwood is mentioned in Indian mythology, folklore, and ancient scriptures. Certain cultures value its aroma and medicinal properties. Sandalwood species are generally thought to be native to the Indian peninsula, as their occurrence dates back at least 2,500 years. Sandalwood trees grow under different eco-climatic conditions and are very well adapted in terms of growth, heartwood and oil content [2]. The best wood grows in the driest regions, especially red and stony soils, while on rocky soils the trees often remain small but produce the highest oil yields. The heartwood is moderately hard, heavy and durable, yellow or brown in appearance and oily in texture, making it the perfect material for carving intricate designs. Sculptures of gods and mythical figures are in high demand in the market. A variety of items are made from sandalwood, including boxes, cabinet panels, jewelry boxes, combs, picture frames, fans, pen holders, card cases, letter openers, and bookmarks. Heartwood represents the central part of the tree and is valued for its aroma. However, parts of the tree such as the bark and outer wood (sapwood) have no scent. This plant is primarily used to extract sandalwood oil, which is obtained by steam distillation of the heartwood. The roots also contain essential oils [3]. Oil yield and quality vary depending on location, age of the tree, and distillation method. Sandalwood is commercially known as East India Sandalwood and its oil is known as East India Sandalwood Oil.

#### PLANT DESCRIPTION :

*Santalum album* is an evergreen tree that grows up to 20 m attaining girth of up to 2.4 m with slender drooping branchlets. There are two major commercial species of Sandalwood named Indian Sandalwood (*S. Album*) and Australian Sandalwood (*S. Spicatum*). Australian Sandal Trees are shorter in height. Bark of the tree is tight, dark brown, reddish, dark grey or nearly black in young trees it is black and smooth, but in older trees it is rough with deep vertical cracks and has a red interior. The sapwood is white and odorless, while the heartwood is yellowish to dark brown and strongly scented. Leaves are thin, almost opposite, ovate or oval-ovate, 3-8 x 3-5 cm, glabrous and light green above, glossy and slightly paler below. The tip is rounded or pointed. Stem grooved, 5-15 cm long. The veins are clearly reticular. Flowers are purplish-brown, small straw yellow, reddish-green or purple, about 4-6 mm long, up to 6 in small terminal or axillary clusters, odorless in conical secondary or terminal racemes. The fruit is a spherical, fleshy drupe. It is red, purple to black when ripe, about 1 cm in diameter, has a hard, ribbed endocarp, the crown of which has a stigma, almost stalkless, smooth, and single-seeded. [4] In India, flower spikes appear from March to April, and fruits ripen during the cold season. In Australia, flowers bloom from December to January and June to August, and fruit ripens from June to September. This species spreads rapidly by dispersal of seeds. Trees produce viable seeds when they reach 5 years of age. Trees over 30 years old will have a circumference of 18 to 38 inches. Sandalwood tolerates a wide range of site conditions and grows naturally in a variety of locations throughout the tropics, although growth is more active in some conditions than others. *S. Album* grows in well-drained loamy soils, preferably on sunny hillsides. It requires at least 20 to 25 inches of rainfall per year. It does not tolerate frost or waterlogging, but is drought tolerant and requires a lot of light during the seedling stage and later stages. The ongoing drought and fires have killed 4,444 trees. This plant is primarily used for the production of fragrant sandalwood oil.

#### Historical Background :

There are references to sandalwood in Indian mythology, folklore, and scriptures. It is mentioned in Indian literature in AD and is as old

as Milindapana (200 BC), Patanjali Mahabhasya AD (100 BC), Dhammapada, Anguttara and Vinaya Pitaka (400-300 BC). Theophrastus described a type of sandalwood in his *Arthashastra* (200 BC). Sandalwood is also mentioned in the epics *Ramayana* and *Mahabharata*. It is unlikely that some of the references to sandalwood in ancient Indian texts refer to his *Pterocarpus santalinus*, called Red Saunders or Red Sandalwood. There is ample evidence that *S. Album* was cultivated in India over the past 25 centuries. Sandalwood oil is highly valued all over the world, with India being the main exporter. Sandalwood, known as "sandalwood" in China and "sandalwood" in Japan, is known in ancient Chinese texts as well as ancient Sanskrit texts and manuscripts. It is also used in various forms of initiation ceremonies to open the minds of students.

#### Origin and distribution:

*S. Album* is native to the tropical regions of the Indian peninsula, eastern Indonesia, and northern Australia. The main distribution is in the drier tropical regions of India and the Indonesian islands of Timor and Sumba. It is native to the highlands of southern India, with the main sandal regions being most of Karnataka and adjoining districts of Maharashtra, Tamil Nadu and Andhra Pradesh. In these regions, this species primarily inhabits dry deciduous forests and shrublands. The vegetation type is a typical monsoon vine thicket growing on pure sand. This record was recorded on a coastal dune just above standard high water mark and near mangroves. It also grows on low lateritic cliffs above beaches. This tree is actually an obligate hemiparasitic plant to various hosts, *Cassia siamea*, *Pongamia glabra*, and *Lantana acuminata*. It is currently grown in India, China, Sri Lanka, Indonesia, Malaysia, the Philippines, and northern Australia.

#### Traditional Uses :

*S. Album* is grown primarily for its timber and aromatic oil. 870 kg/cubic meter wood is durable and stable. The densely grained heartwood is used for ornaments and carvings. Wood was used as fuel, but is generally considered too valuable for that purpose. Sandalwood oil is distilled from the heartwood and is a pale to yellow viscous liquid with a sweet, aromatic, long-lasting, spicy, warm, woody, animalistic, milky, and nutty note. Widely used in perfume, cosmetics, aromatherapy and pharmaceutical industries. Being an excellent fixative, it is highly valued by the perfume and

toiletory industry, especially for certain delicate scents that are very rare and fragile. No heavy or oriental perfume composition of type is complete without a generous infusion of sandalwood oil. Most Indian attars use sandalwood oil as a base. This is due to its ability to absorb most of the essential notes of other whole herbs and flowers, which can enhance scent status and stability. This oil is used as a flavoring agent in foods such as frozen dairy desserts, sweets, bread masala, bakery products, gelatin, pudding, alcoholic and non-alcoholic beverages. The United States Food and Drug Administration, the European Council of Flavor and Extract Manufacturers Associations and the FAO/WHO Joint Expert Committee have approved the use of sandalwood oil as a food additive [8].

#### Ethnopharmacology :

Therapeutically, sandalwood has a calming and relaxing effect. It is said to reduce stress, depression, anxiety, nervous fatigue, restlessness, fatigue, insomnia, and promote meditation. It was believed to promote spiritual practices, peaceful relaxation, openness, and grounding.[6] Sandalwood is used in various traditional medicine systems such as Ayurveda, Siddha medicine, and Unani medicine to treat various ailments. In Ayurveda, India's traditional medical system, sandalwood is primarily used as a sedative, diuretic, and mild stimulant. In Ayurveda, sandalwood is considered an antiseptic, antipyretic, antiscabies, diuretic, expectorant, and stimulant, and its antibacterial and antifungal properties can treat bronchitis, urinary disorders, urinary tract infections, and gonorrhea. It is prescribed for the treatment of. [12] Other therapeutic uses mentioned in Ayurveda include bile, fever, papules, diarrhea with internal bleeding, eye infections, hemorrhage, hiccups, omphalitis, poisoning, early stages of smallpox, hives, etc. Including its use in the treatment of various diseases.[3,8]

#### PHYTOCHEMISTRY :

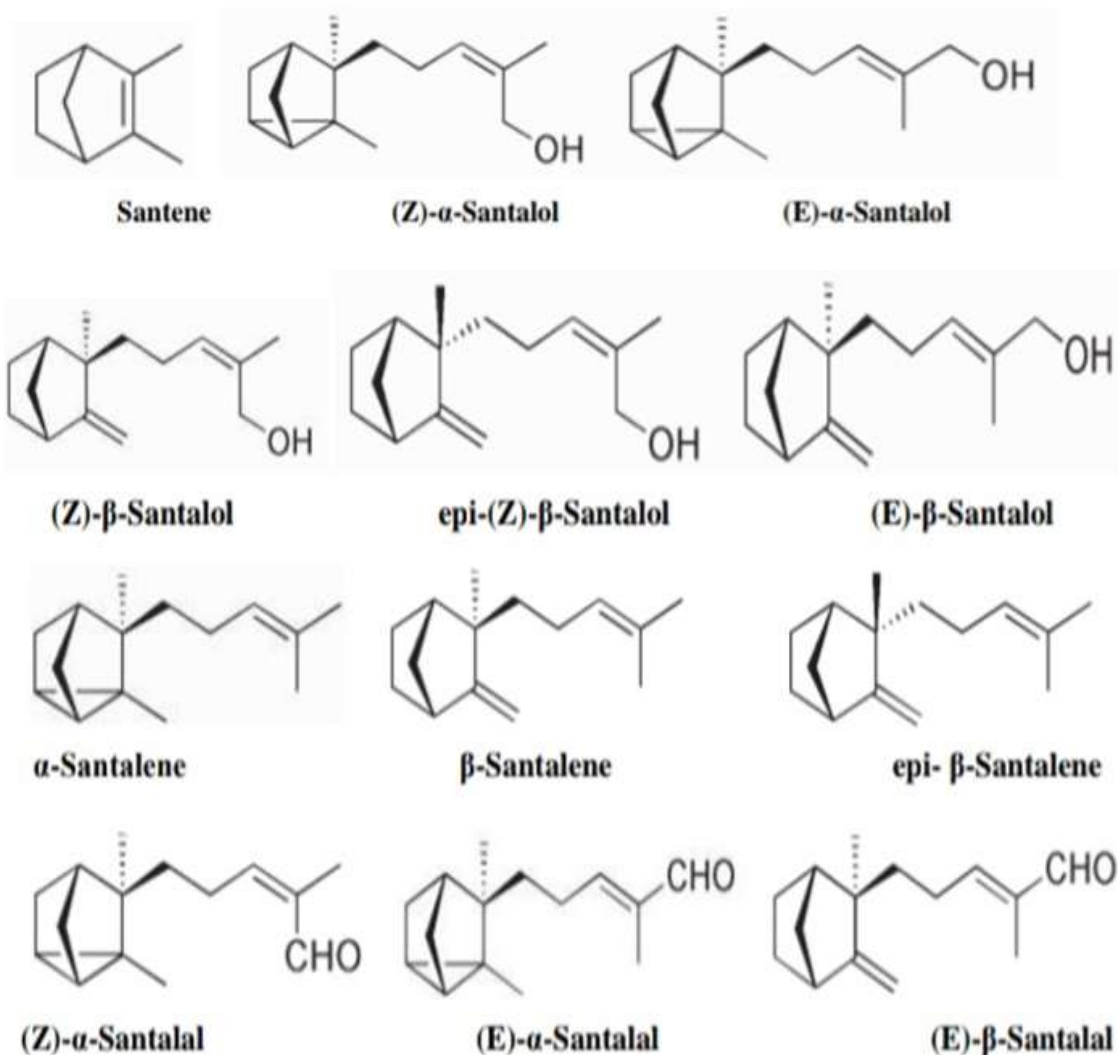
Sandalwood, the most valuable wood and the main source of essential oils, has been extensively studied for its chemical composition. Sandalwood oil accumulates in the heartwood only after it has grown for 30 years under natural conditions. [37] The yield and composition of all essential oils are strongly influenced by tree age, heartwood color, organ maturity, individual tree, location within the tree, and environmental and genetic factors of the plant. [38-40] Traditionally,

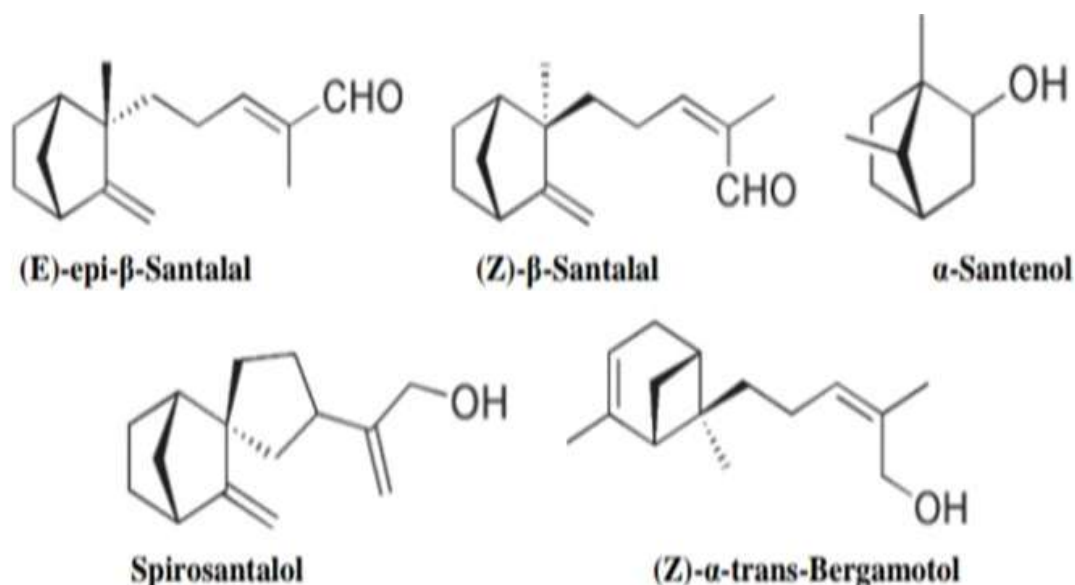
according to ancient customs, steam distillates of heartwood are sold as marketable sandalwood essential oil. [41] Additionally, the composition of the oil extracted from young and mature sandalwood trees is different, [43] while the content and composition of the oil from the heartwood, which is extracted in varying amounts from the tree, is different. [44]. Sandalwood oil has been extensively studied for its chemical composition and its isolation, synthesis, and quantitative estimation.[45-57] Shankaranarayana and his colleagues have carried out extensive research on the phytochemical and other related aspects of sandalwood. Chromatographic separation of  $\alpha$ - and  $\beta$ -santalol [58]  $\alpha$ - and  $\beta$ -santalole by column chromatography [59] Simple method for high-yield extraction of sandalwood oil by rectification of benzene extract, [60] Sodium santarubate dimethyl sulfate inclusion complex, [61] Estimation of oil content in sandalwood depot-based sapwood, [62] Possibility of developing fragrant products from low-odor sandalwood oil [41] Estimation of oil content and composition from sandalwood Central and transitional regions of sandalwood slices [63], Use of sandalwood extract [64], Hydrolyzed Depleted Sandalwood Powder (HESP) extraction of essential oil from [65-66], isolation of santalol from sandalwood oil [67]. The main components of commercially available sandalwood oil are sesquiterpene alcohols such as  $\alpha$ - and  $\beta$ -santalol (C<sub>15</sub>H<sub>24</sub>O), bergamotol, and several stereoisomers thereof, while minor components include linsol, Includes sesquiterpene hydrocarbons such as pherol, bisabolol, and  $\alpha$ -  $\Phi$ -santarene (C<sub>15</sub>H<sub>24</sub>), bergamotene,  $\alpha$ -,  $\phi$ -, and  $\omega$ -curcumene,  $\phi$ -bisabolene [68-72], and phenylpropanoids [73]. A-santalol is usually more common than  $\tilde{O}$ -santalol.

[74] With Verghese Collaborators reported the sesquiterpene alcohols cis- $\square$ -santalol and cis- $\square$ -santalol,  $\square$ -trans—bergamotol and epi-cis- $\square$ -santalol as the main components of essential oils [75]. Minor components include trans- $\square$ -santalol and cis-lantheol, [72] hydrocarbon santarene (C<sub>9</sub>H<sub>14</sub>),  $\square$ -santarene,  $\square$ - santarene,  $\square$ -bergamotene, epi- $\square$ -santarene,  $\square$ -curcumene,  $\square$ -Curcumene,  $\square$ -Curcumene,  $\square$ - Bisabolene,  $\square$ - Bisabolol [72,76] and heterocycles. Other ingredients mentioned in sandalwood oil include alcohol, sunthenol (C<sub>9</sub>H<sub>16</sub>O), and telecentalol (C<sub>10</sub>H<sub>16</sub>O). Aldehydes, nor-tricycloecasantral (C<sub>11</sub>H<sub>16</sub>O) and isovaleraldehyde. The ketones L-santenone (C<sub>9</sub>H<sub>14</sub>O) and Santalone (C<sub>11</sub>H<sub>16</sub>O), the acids telesanthalic acid (C<sub>10</sub>H<sub>14</sub>O) (partially

free and partially esterified form of  $\alpha$ - and  $\beta$ -santalnic acid (C<sub>15</sub>H<sub>22</sub>O<sub>2</sub>). The genetic diversity among sandal populations from different localities in India [78] and the relationship between abundance and heartwood/oil yield in sandal localities [2] were investigated. Indian Sandalwood Oil and Australian Sandalwood Oil contain similar ingredients, but the concentrations of these ingredients are different, resulting in two similar but completely different oils. The main components of Australian sandalwood oil are sesquiterpenes, with the two major sesquiterpene alcohols  $\alpha$ -santalol and  $\beta$ -santalol, and E, E-farnesol and  $\beta$ -bisabolol being the predominant occupies. It is worth mentioning here that the content of secondary metabolites depends on the intrinsic

properties of the plant material, environmental and genetic aspects, or external aspects such as extraction methods. [80] also noted that the amount of santalyl acetate and santarene in 10-year-old trees was slightly higher than in 30-year-old trees [81]. Roots of Indian sandalwood tree yeast were also tested for yield and essential oil composition. By solvent extraction of sandalwood, oil recovery from the roots of was determined to be 10.3%. GC and GC/MS analysis of the oil revealed that it accounts for 99.9% of the total oil, including 30 sesquiterpenols (78.5%), 9 sesquiterpenes (7.8%), and 1 terpene acid (0.4%). Compounds of 53 types were detected. Five sesquiterpenoid isomers (4.4%);





The main components of the essential oil were  $\alpha$ -santalol and  $\beta$ -santalol, and their percentages were 19.6% and 16.0%, respectively.  $\beta$ -Santalol content was below the recommended range of 41–55%. However, in , the  $\beta$ -santalol content was close to the specification of 18%. The total content of 4,444 bisabolene A, B, C, D and their isomers was also high, being 25.0% of the oil [82]. In another study, GC-MS studies identified a total of 32 active phytochemicals in stem extracts of Santalum album. In another study, his GC-MS analysis detected 4,444 volatile metabolites in the heartwood of 15-year-old trees. [84] Bioassay-controlled fractions of Santalum album heartwood were performed and seven  $\beta$ -Santas including (9R,10E)-9-hydroxy- $\beta$ -Santalol, (10R,11R)-10,11 Role derivative was isolated. Dihydroxy- $\beta$ -santalol, (9E)-11,13-dihydroxy- $\beta$ -santalol, and (10E)-12-hydroxy- $\beta$ -santalol acid . Their structures were determined by spectroscopic analysis.[85] HPTLC-based evaluation of sesquiterpenoids from sandalwood oil was developed to profile metabolites such as n-alkanes, sesquiterpenes, sesquiterpenoids, fatty acids, alcohols, and hydrocarbons in sandalwood oil. Ta. Srivastava et al. Studied the functional characterization of a novel -sesquiterpene synthesis from Indian sandalwood. The possibility of reliably identifying wood based on its anatomical structure, color of the hot water extract, chemical composition of the oil (mainly santalol content), and DNA fingerprints was investigated. [87] The structure-activity relationship of sandalwood-derived odorant compounds was studied. [88- 89]

Furthermore, the relationship between the structure and odor of (Z)- $\alpha$ -santalol, the main component of sandalwood essential oil, which has a unique wood odor derivative, was also studied. , focused on the relationship between the side chain structure and odor of types of compounds. This study found that the odor of the Z isomer is similar to that of the corresponding saturated compound of , but significantly different from that of the corresponding E isomer of (odorless, fresh or oily). Did. These results indicated that the relative configuration of the side chain to the santalol backbone plays an important role in the odor of  $\alpha$ -santalol. The E configuration of the side chain eliminates the woody aroma character of  $\alpha$ -santalol and its studied derivatives, but the Z configuration or the saturation of the carbon side chains do not [90]. Studies on sandalwood tree seeds and seed oil [91], partial hydrogenation of sandalwood seed oil [92], removal of unsaponifiable components from sandalwood seed oil, fatty acid composition of seed coat [93], and seed Overstorage of petroleum has also been reported [94]. Isolation of betulinic acid from sandalwood seed coat and its reduction by Baggins have been reported. The use of bark and seeds of depleted sandalwood was investigated. [91,96] Flavonoid components vicenin-2, vitexin, isovitexin, orientin, isoorientin, chrysin-8-C- $\beta$ -D-glucopyranoside, chrysin-configuration 6- C - $\beta$ -D- glucopyranoside and isorhamnetin was isolated from the leaves of S. And characterized. Albums. [97-98] Sandalwood oil research, synthetic substitutes, industrial and therapeutic uses were summarized by Ranade.

Furthermore, Makoto published a research report on natural and synthetic fragrances of sandalwood [100]. The synthesis of sandalwood odor derivatives from camporenic aldehyde has been reported [101]. Process of synthesizing fragrances with similar properties Flavor patented oxidation of santalol with MnO<sub>2</sub> in hexane to give Z- $\alpha$ -santalal, which isomerized with 0.5-2% glacial acetic acid to produce the product E- $\alpha$ -santalal . [102] Safety evaluation of oils in food has been studied [103]. A comparative study of the chemical composition of fragrant sandalwood species including *S. Album*, *S. Spicatum*, and *S. Austrocaledonicum* was conducted.[104] The use of many essential oils , including sandalwood oil, in aromatherapy was described by Setzer. Gleason discussed the possibilities and prospects for using Indian sandalwood oil as an ingredient in luxury perfumes. Methods have been developed to identify and detect adulterants.[107-108]

#### Pharmacology :

Sandalwood and its oil have demonstrated a wide range of pharmacological activities, in addition to its relevance in perfumery and cosmetics. *S. The album* has been extensively studied to confirm traditional therapeutic claims and reveal additional biological activities. Her series of 4,444 pharmacological studies on sandalwood and its oil reported biological effects ranging from antibacterial to anticancer. The reported pharmacological activities of his sandalwood and its oil are summarized below.

#### Hepatoprotective activity :

Hydroalcoholic extract from leaves of *S. Album* shows significant hepatoprotective activity against CCl<sub>4</sub> and paracetamol and induces hepatotoxicity by reducing the activities of serum marker enzymes, bilirubin and lipid peroxidation. And showed a significant increase in glutathione levels. Superoxide dismutase, catalase and protein decreased in a dose-dependent manner, which was further confirmed by the decrease in total liver weight and histopathological studies. [109]

#### CNS Effect :

*Santarum Album L* It is known to be effective in improving memory. [110-111] Studies on the sedative effects of have shown that inhalation of East Indian sandalwood oil reduced motility in mice by 40-78% compared to 0% controls. [112] The soothing effects of sandalwood oil and aqueous extracts have already

been demonstrated. [113-114] Sandalwood oil is said to have a relaxing effect on the nerves and is used for hot or excited emotional states that cause headaches, insomnia, and nervous tension. . [32] Santalol, as a bioactive ingredient , has been reported to affect the central nervous system (CNS). Accordingly, the effects of depression have been shown to have a 4,444 impact on patients with sleep disorders [115]. The first study of its kind explores the sense of smell.Receptor neurons specifically stimulated by four synthetic sandalwood compounds and oil were identified. Additionally, solvent extracts of heartwood have been shown to have neuroleptic effects in mice.  $\alpha$ -santalol and  $\beta$ -santalol significantly increase the concentrations of homovanillic acid, 3,4-dihydroxyphenylacetic acid, and/or 5-hydroxyindoleacetic acid in the brain of mice when administered intragastrically and intracerebroventricularly. I let it happen. [113]  $\alpha$ -Santalol is a potent dopamine D2 antagonist – and serotonin 5-HT<sub>2A</sub>- receptor binding. Furthermore, the effects of  $\alpha$ -santalol were similar to those of chlorpromazine as an antipsychotic. [117] Alpha-santalol caused significant physiological changes such as relaxing and sedative effects, whereas sandalwood oil caused physiological inactivation after transdermal absorption but no behavioral activation. Ta.Recently, TLC254 bioautograph studies have shown that alpha-santalol, the main component of this essential oil, is a potent inhibitor of both tyrosinase and cholinesterase in vitro, and therefore this essential oil may be associated with Alzheimer's disease. It has been shown that it has great potential for use in the treatment of The same is true for skin care [119].

#### Anti-ulcer effect :

Oral treatment of hydroalcoholic extract of *S. Album* strain through effective inhibition of physical (stress) and chemical (both local irritants and pharmaceutical NSAIDs) induce gastric exposure. , was reported to induce good levels of gastroprotection in rats. Ulcer. [120]

#### Antifungal Activity :

Sandalwood oil is reported to have antifungal activity against *Microsporiumcanis*, *Trichophytonmentagrophytes*, and *T. Rubrum*. Sandalwood oil was found to be effective against human pathogenic fungal strains *Microsporiumcanis*, *Trichophytonmentagrophytes* , and *T. Rubrum*, but not against *Candida albicans*,

*Aspergillus niger*, and fumigated *A. niger*. It was ineffective against [132]

#### Antioxidant Activity :

Phytochemical and pharmacological studies have demonstrated the existence of principles that justify the traditional medicinal properties of antioxidant. *S. Album* and other Indian medicinal plants were tested in vitro for possible regulatory effects on nitric oxide (NO) levels using sodium nitroprusside as the NO donor. Most plant extracts showed significant direct, dose-dependent scavenging activity towards NO. [140] Reported to have nitrous oxide scavenging activity and DPPH antioxidant activity. [140] [141] *Santalum album* can protect cardiac tissues from cell damage and lipid peroxidation caused by oxidative stress, and also prevents DOX-induced inflammation and induction of apoptosis. [142] Recently, the anthocyanin pigment cyanidin-3-glucoside from *S. Album* was shown to have antioxidant properties and be nutritionally important. Furthermore, in a comparative study, nine in vitro antioxidant assays were performed and in vitro cultured callus cells were shown to exhibit antioxidant activity comparable to sandalwood oil. Sandalwood oil increased glutathione S-transferase (GST) activity and acid-soluble sulfhydryl (SH) levels in the livers of adult male Swiss albino mice. [145] Increased GST activity and acid-soluble SH levels suggested a possible chemopreventive effect of sandalwood oil on carcinogenesis through a blocking mechanism.

Similarly, methanol extract of sandalwood demonstrated inhibition of acetylcholinesterase, DPPH and superoxide radical scavenging activity in albino mice, indicating the potential of to combat dementia and memory loss associated with Alzheimer's disease. Was shown. Recently, an in vivo study investigated the antiglycemic and antioxidant effects of sandalwood oil and its main component  $\alpha$ -santalol in an alloxan- and D-galactose-mediated model using oxidative stress-induced diabetic male Swiss albino mice. Possibilities have been evaluated.

#### Antibacterial activity :

Some studies have focused on the antibacterial properties of East Indian sandalwood oil [121], while many other studies have focused on Australian sandalwood oil [122]. A comparative study conducted with 26 essential oils that evaluated their antibacterial activity against bacteria in the axilla showed the strongest activity

against sandalwood oil and its synthetic analogs [123]. Sandalwood oil is an effective antibacterial agent against methicillin-resistant *Staphylococcus aureus* and antifungal-resistant *Candida* species. [124] Crude extract of sandalwood oil and  $\alpha$ - and  $\beta$ -santalol compounds exhibit antibacterial activity against *Helicobacter pylori*, a Gram-negative bacterium closely associated with the development of duodenal, gastric, and gastric ulcers. . Sandalwood oil also shows activity against herpes simplex virus type 1 [125], and  $\alpha$ -santalol shows anti-influenza activity against H3N2 virus. Another study found the greatest inhibitory effect of sandalwood oil on *Bacillus mycoides* and *Escherichia coli*. [126] Methanol extract from *S. Album* is said to be effective against *Bacillus subtilis*, *Salmonella typhi*, *Staphylococcus aureus*, and *Staphylococcus aureus*. Very effective against *Pseudomonas aeruginosa* and *Candida albicans*. [127] Sandalwood oil showed antidermatophyte activity against *Microsporium canis*, *Trichophyton robrum*, and *Trichophyton mentagrophytes*. [128] Furthermore, the sandalwood oil components  $\alpha$ - and  $\beta$ -santalol were found to be effective against *Salmonella typhimurium* and *Staphylococcus aureus*, and  $\alpha$ -santalole was found to be effective against *Salmonella typhimurium*. Did. [128] Santalbic acid (trans-11-octa-decene-9-ynoic acid), the main component of seeds, was found to inhibit Gram-positive bacteria and some pathogenic fungi. Santalol was found to be effective against yeast and Gram-positive and negative bacteria at high and/or medium concentrations, and showed better antibacterial activity even at low concentrations. Furthermore, immature tree buds have also been shown to have antibacterial effects against 4,444 13 different bacterial strains [130].

#### Antiviral Activity :

Sandalwood's antiviral activity has also been proven by biological studies. Rephrase Sandalwood oil has been proven to be used in the prevention and treatment of warts, age spots, and other skin tumors caused by viruses. [133-134] Traditional medical systems, including Ayurveda and Chinese medicine, also mention the antiviral effects of sandalwood oil. [135] An in vitro study showed that sandalwood oil has dose-dependent antiviral activity against herpes simplex virus (HSV)-1 and 2 by inhibiting the replication of the virus. Ta. Sandalwood oil was also thought to help protect cells by regulating the levels of glutathione, S-transferase, and acid-soluble sulfhydryls in the

liver. [136] Sandalwood oil showed an inhibitory effect on herpes simplex virus type 2 (HSV-2) on RC-37 cells in vitro. Interestingly, before being adsorbed to cells, sandalwood oil affected the virus only through non-specific inhibition of the interaction between the virus and host cells [25]. The components of sandalwood oil, alpha- and beta-santalol, their mixtures and derivatives, are relevant for the treatment of human warts, especially HPV and DNA poxviruses that cause warts. *Molluscum contagiosum* is said to be a treatment for HIV and other RNA viruses, as well as for seborrheic dermatitis, psoriasis, and the dryness, scaling, and desiccation associated with allergic and eczematous rashes. *Santalum album* is said to be a treatment for acne lesions on the face and body and removal of pustular acne – lesions caused by staphylococcal acne and streptococcal infections. Additionally, sandalwood oil and santalol derivative are said to be used in the treatment of cold sores and cold sores [137]. Recently, single cell and somatic embryo suspension cultures of Indian sandalwood have been shown to be an alternative and renewable source of shikimic acid, a precursor for the industrial scale synthesis of Tamiflu, the only commercially available neuraminidase inhibitor against influenza A. It has been shown. *Virus*. [138]

#### Antioxidant Activity :

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#### Hemolytic Activity :

The hemolytic activity of any compound is an indicator of general cytotoxicity to normal healthy cells. One study found that a leaf extract of this plant caused the lysis of 4,444 red blood cells. However, this hemolytic activity occurs only when administered parenterally. Studies have shown that the leaves of this plant contain saponins, which are responsible for hemolytic activity on blood. [148]

#### Anticancer Activity :

studies have demonstrated the chemopreventive effects and molecular mechanisms of  $\alpha$ -santalol on skin cancer development in both animal models and skin cancer cell lines. Rephrase Anticancer effects of the oil were demonstrated in chemically induced skin carcinogenesis in CD-1 and SENCAR mice, UV-B-induced skin carcinogenesis in SKH-1 mice, and in vitro models of melanoma, non-melanoma, breast and melanoma. It has been reported. Prostate cancer and its ability to induce cell cycle arrest and apoptosis in cancer cells has also been demonstrated. [150]  $\alpha$ -Santalol, a component of sandalwood oil, delayed the development of papillomas in both strains of mice. [151]  $\alpha$ -santalol at concentrations of 25–75  $\mu$ M was found to induce apoptotic death of human epidermal carcinoma A431 cells through caspase activation, along with loss of mitochondrial potential and cytochrome release. [152] In a similar study in female hairless mice of the SKH-1 strain, topical application of  $\alpha$ -santalol showed chemopreventive effects due to reductions in ornithine decarboxylase activity, tumor incidence, and proliferation. Has been proven. Furthermore,  $\alpha$ -santalol slows skin tumor development, reduces



tumorproliferativeness, and inhibits lipid peroxidation in skin and liver microsomes in vitro, possibly by acting as an antioxidant, and thus has been shown to prevent UVB—induced skin tumor development. [154]  $\alpha$ -Santalol significantly upregulated apoptosis-related proteins, caspases 3 and 8, and tumor suppressor protein p53 through the extrinsic pathway in a UV-B-induced skin tumor development model in SKH-1 mice. It has been reported that it increases. -Santalol induced apoptosis in human prostate cancer cells through activation of caspase-3. [156] Approximately six novel sesquiterpenes were assayed for in vitro activation of Epstein-Barr virus early antigen (EBV-EA) in Raji cells to assess their antitumor-promoting activity. Terpenoids, two aromatic glycosides and several neolignans were identified. Furthermore, in vivo two-step carcinogenesis assays of demonstrated a strong inhibitory effect on EBV-EA activation and a strong inhibitory effect of on two-step carcinogenesis in mouse skin. [157] Furthermore, a derivative of  $\beta$ -santalol showed tumor-selective cytotoxicity in human HL-60 promyelocytes, Leukemia cells and normal human diploid TIG-3 fibroblasts. [158] Two lignans obtained from heartwood samples showed tumor cell apoptosis-inducing cytotoxicity against HL-60 human promyelocytic leukemia cells and A549 human lung adenocarcinoma cells. Alpha-santalol, an active ingredient in sandalwood essential oil, has been studied for its protective effects against skin cancer in mouse models of skin carcinogenesis. [159]

#### Antipyretic effect:

Santalwood oil at a dose of 200 mg/kg showed a highly significant antipyretic effect on yeast-induced fever in albino rats.

#### Anti-inflammatory effects :

Santalol has been reported to have significant anti-inflammatory effects in several experimental models. [33] Santalum album had anti-inflammatory and anti-ulcer effects as evidenced by significant inhibition in carrageenan-induced paw edema, cotton pellet-induced granulomas, and pyloric ligation-induced ulcers. These findings may support the inclusion of this plant in the effective treatment of inflammatory diseases such as ulcers in traditional medicine. Methanol extracts from heartwood have been shown to have in vitro antioxidant effects (and in vivo analgesic and anti-inflammatory effects) in mice. [160]

#### antihyperglycemic and antihyperlipidemic effects :

Effect of long-term oral administration of the petroleum ether fraction of sandalwood in streptozotocin-induced A study on the antihyperglycemic and antihyperlipidemic effects of long-term oral administration of the petroleum ether fraction of sandalwood in streptozotocin-induced diabetic rats showed a reduction in blood glucose levels. A decrease in blood glucose levels was also observed in the metformin treatment group, whereas an increase in blood glucose levels was observed in the diabetic control group. Additionally, total cholesterol (TC), low-density lipoprotein (LDL), and triglyceride (TG) levels were decreased in treated diabetic rats, whereas cardioprotective high-density lipoprotein (HDL) was increased. A significant improvement in the atherogenic index was observed around, leading to the conclusion that *S. album* has potential antihyperglycemic and antihyperlipidemic effects. [161]

#### Cardioprotective activity :

Aqueous extract of sandalwood has been reported to significantly inhibit cardiac tissue damage by reducing lipid peroxidation in doxorubicin-induced cardiotoxicity in a rat model [162] and ISO in albino Wistar rats. It has been reported that it has a significant protective effect against induced myocardial infarction. Dependent way.

#### Physiological Effects:

Santalwood oil and its main component  $\beta$ -santalol have been shown to influence several physiological functions and sensory stimulation. The oil reportedly increases pulse rate, skin conductance and systolic blood pressure, while  $\beta$ -santalol produces higher levels of alertness and mood than the oil. [163] Inhaling sandalwood oil has been reported to improve hearing. [164] Recently, sandalwood tea significantly increased myocardial contractility and heart rate in isolated and dysfunctional frog hearts, while having excellent anti-fatigue effects on smooth muscle contraction in isolated rabbit aortic strips. It has been shown to be effective. [165] Sandalwood oil did not show phototoxicity, but irritation and sensitization reactions in humans have been occasionally reported. [166]

#### Metabolic Effects :

Sandalwood oil has been reported to exhibit changes in neonatal hepatic xenobiotic-metabolizing enzymes in lactating mouse pups upon transmammary exposure. It was also observed that sandalwood oil and its constituents passed through milk and altered xenobiotic metabolizing enzymes in the liver, including a decrease in soluble sulfhydryl content and hepatic cytochrome P 450 content.

## II. CONCLUSION:

*Santalum album* is one of the most famous and widely used plants in perfumery and cosmetics. Apart from its perfume and cosmetic uses, sandalwood has a wide range of pharmacological activities and this plant is considered one of the important medicinal plants. Although this plant has been extensively studied since the past 20 years, there is still much scope to realize the full potential of using sandalwood for humanity. For more than a century, researchers around the world have focused on the study of interesting chemical constituents, especially the sesquiterpenoids of sandalwood, in terms of structure, synthesis, and pharmacological activity. There has been a recent surge in research activity validating the traditional medical uses of essential oils and their components through modern experimental approaches, driven by thorough pharmacological and mechanistic studies. Various studies have demonstrated the pharmacological activities of sandalwood and its oil, ranging from antibacterial to anticancer properties. It also shows prominent activity in various skin diseases. There are few toxicological studies on Sandalwood. It is necessary to summarize all activities reported about this plant. This review consolidates different reported activities of sandalwood plant as well as its oil.

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