

Available online on 15.4.2022 at <http://ajprd.com>

# Asian Journal of Pharmaceutical Research and Development

Open Access to Pharmaceutical and Medical Research

© 2013-21, publisher and licensee AJPRD, This is an Open Access article which permits unrestricted non-commercial use, provided the original work is properly cited



Open Access

Review Article

## A Review on Phytochemical Pharmacological and Biological Activities of Thuja Occidentalis

**Keshav Bharti, Manmohan Sharma, Girish Kumar Vyas, Dr. Shailesh Sharma**

School of Pharmaceutical Studies, Faculty of Health Sciences, Dr. K. N. Modi University, Newai, Rajasthan

### ABSTRACT

Thuja occidentalis plant of Cupressaceae family originated in Eastern North America and is now planted as an ornamental tree in Europe and Brazil, where it is known as the "tree of life" or "white cedar." It's often used to treat liver problems, enuresis, amenorrhea, bullous bronchitis, psoriasis, cystitis, uterine carcinomas, diarrhoea, and rheumatism in traditional medicine. Due to the presence of essential oil, coumarins, flavonoids, tannins, and proanthocyanins, the chemical ingredients of T. occidentalis have piqued researchers' curiosity for decades. Antioxidant, anti-inflammatory, antibacterial, antifungal, anticancer, antiviral, gastrointestinal tract protective action, radioprotection, antipyretic, and lipid metabolism regulating activity are all examples of pharmacology. As a result, the current study is a compilation of all essential material for T. occidentalis, including phytochemical, biological and a detailed examination of their pharmacological properties. The current review represents a synthesis of all relevant information for Thuja occidentalis, including phytochemical, biological and a thorough analysis of their pharmacological activities, in order to promote all of the biological activities demonstrated thus far, rather than focusing on a single biological activity.

**Keywords:** Thuja occidentalis, thujone, pharmacological, biological activities**ARTICLE INFO:** Received; 23 Jan 2022   Review Complete; 22 Feb. 2022   Accepted; 19 March 2022   Available online; 15 April. 2022**Cite this article as:**

Bharti K, Sharma M, Vyas GK, Sharma S, A Review on Phytochemical Pharmacological and Biological Activities of Thuja Occidentalis, Asian Journal of Pharmaceutical Research and Development. 2022; 10(2):111-114.

DOI: <http://dx.doi.org/10.22270/ajprd.v10i2.11075>**\*Address for Correspondence:**

School of Pharmaceutical Studies, Faculty of Health Sciences, Dr. K. N. Modi University, Newai, Rajasthan

### INTRODUCTION

The application of diverse complementary treatments is becoming a more popular therapeutic option. The use of plants to cure various diseases plays an important role in this area<sup>[1-3]</sup>. Even though it is not legally recognised in many countries, folk medicine is extensively utilised across the world<sup>[4]</sup>. Plants and plant extracts are used by over 80% of the people in Africa and Asia to cure various diseases<sup>[5]</sup>. The basic herbs offer a solution with no side effects and effective treatments, and the greatest part is that anybody of any age may utilise herbal medicine<sup>[6]</sup>. Secondary metabolites such as steroids, phenolic compounds, flavonoids, alkaloids, and other chemicals can be produced by herbal plants. These secondary metabolites are used to treat a variety of diseases<sup>[7]</sup>. Natural goods and traditional therapies are quite valuable. Modern medicine will not be the exclusive remedy to today's diseases. As a result, humans favourably discern "return to nature" and converse with it in the same way as plant products are used

in medications<sup>[8]</sup>. Herbal medicines are becoming increasingly popular all over the world for a variety of reasons: They have long-term therapeutic benefits and are known for their efficacy, safety, and low side effects<sup>[9,10]</sup>. Ethnopharmacological research are now promoted across the world as a means of identifying species that contain compounds or beneficial products that may be exploited in the pharmaceutical, nutritional, and cosmetic industries<sup>[11,12]</sup>.

Thuja occidentalis, sometimes known as Arbor vitae or white cedar, is a native of eastern North America and a popular decorative tree in Europe<sup>[13]</sup>. Native Indians in Canada discovered the herb as a medicine during a 16th century voyage, and it was proven to be useful in the treatment of scurvy weakness<sup>[14]</sup>. Thuja occidentalis has been used in traditional medicine to treat bronchial catarrh, enuresis, cystitis, psoriasis, psoriasis, uterine carcinomas, amenorrhea, and rheumatism<sup>[15-18]</sup>. It is now mostly used as a mother tincture or dilution in homoeopathy<sup>[19,20]</sup>. This

medicinal plant is also used as evidence-based phytotherapy for acute and chronic infections of the upper respiratory tract <sup>[21,22]</sup>, and as an adjuvant to antibiotics in severe bacterial infections such as bronchitis, angina, pharyngitis, otitis media, and sinusitis <sup>[23,24]</sup> when combined with other immunomodulating plants like *Echinacea purpurea*, *Echinacea pallida*, and *Baptisia tinctoria* <sup>[23,24]</sup>.

*Thuja occidentalis* is a shrub that was originally grown in Asia and North America. It is a natural European tree that

grows up to 15–20 metres tall. It exhibits pyramidal coniferous characteristics, with flattened branches and twigs in one plane and tiny scale-like leaves <sup>[25]</sup>. The leaves are green throughout the year, with a brighter green on the bottom side where resin glands are also found. The seeds are contained in little, 1–2 cm long green to brown coniferous pins <sup>[26,27]</sup>.



**Figure 1:** Thuja leaves and plant Image <sup>[28]</sup>

*T. plicata* essential oil has generally been used for wood preservation and insect repellent due to its antibacterial and insecticidal qualities <sup>[29–31]</sup>. Aborigines in the Pacific Northwest have long utilised *T. plicata* leaf oil to treat a variety of upper respiratory tract disorders <sup>[29]</sup>. Its essential oil also contains antibacterial and antifungal properties <sup>[30]</sup>. The high concentration of - and -thujone, the major physiologically active molecules in other essential oils with comparable antibacterial capabilities <sup>[32,33]</sup>, is assumed to be responsible for its antimicrobial action.

Table 1 summarises the bioactive potential of *Thuja* species based on their uses. The current study provides an overview of the state-of-the-art in ethnobotany, phytochemistry, and a detailed analysis of the pharmacological properties of *T. occidentalis*, including antioxidant, anti-inflammatory, antibacterial, antifungal, anticancer, antiviral, gastrointestinal tract protection, radioprotection, antipyretic, and lipid metabolism regulatory activity, in order to promote all of the biological activities shown thus far, rather than the antitumoral activity.

**Table 1:** *Thuja occidentalis*' bioactive potential in comparison to those of other *Thuja* genus members.

Thuja occidentalis' bioactive potential in comparison to those of other Thuja genus members.			
Sr. No.	Species of Thuja	Biological Activities-Related Benefits	
		Medicinal Use	Industrial Use
1	<i>T. koraiensis</i>	antimicrobial <sup>[34,35]</sup> ; antioxidant <sup>[36]</sup>	
2	<i>T. occidentalis</i>	antimicrobial <sup>[30,37–39]</sup> ; antioxidant and anti-inflammatory <sup>[37, 40–44]</sup> ; antitumor <sup>[45–48]</sup> ; hepatoprotective and gastroprotective <sup>[49,50]</sup> ; antidiabetic <sup>[51]</sup> ; antiatherosclerosis <sup>[52]</sup> ; radioprotective <sup>[53]</sup> ; antipyretic <sup>[54]</sup>	antifungal agent for biocontamination control in libraries and archives storage areas <sup>[55]</sup> ; insecticidal activity <sup>[56]</sup>
3	<i>T. plicata</i>	Antimicrobial <sup>[29,30,57]</sup> ; Anti-inflammatory, immunomodulatory, and tissue remodelling <sup>[58]</sup>	Antimicrobial agent for decontamination of buildings <sup>[29]</sup>
4	<i>T. standishii</i>	Antimicrobial <sup>[61–63]</sup> ; antitumor <sup>[61–63]</sup>	
5	<i>T. suchuenensis</i>	antimicrobial <sup>[59,60]</sup>	

### Phytochemical constituents of *T. occidentalis*:

Fresh plant includes essential oil, reducing sugar, water soluble polysaccharides, water soluble minerals, free acid, tannic agents<sup>[64]</sup>, flavonoids, saponins, glycosides, and alkaloids<sup>[65]</sup>, according to biochemical investigations. The primary monoterpenes in the essential oil of fresh leaves (related to the monoterpene fraction) are 65 percent thujone, 8 percent isothujone, 8 percent fenchone, 5 percent sabinene, and 2 percent  $\alpha$ -pinene<sup>1,3</sup>. Carvotanacetone, origanone, origanone, myrcene, and camphor are some of the other monoterpenes that have been described<sup>[64, 67-69]</sup>. The heartwood of *T. orientalis* includes aroma-dendrin, taxifolin, widdrol, cedrol, thujopsadiene, dehydro-

curcumene, -isobiotol, and curcumenol, while the leaves contain rhodoxanthin. For 100 percent of the whole oil, *T. orientalis* essential oil revealed the presence of 38 compounds.  $\alpha$ -pinene (22.25 percent), 3-carene (20.65 percent), cedrol (18.71 percent),  $\alpha$ -Caryophyllene (6.13 percent),  $\alpha$ -humulene (5.68 percent), terpinolene (4.53 percent), and limonene (3.35 percent) were the primary components<sup>[70]</sup>. An article discovered twenty-one molecules, with quantitative variations mostly between cone and needle oils. Both oils were high in monoterpene hydrocarbons, with the primary components being  $\alpha$ -pinene (64.2 and 49.3 percent in cones and needles, respectively), phellandrene (6.7–9.6 percent), and  $\alpha$ -cedrol (6.7–9.6 percent) (3.9 and 8.2 percent)<sup>[71]</sup>.

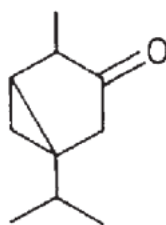


Figure 2: Thujone API of *Thuja occidentalis* plant

It is the main element in the essential oil extracted from the dried herbal content of *Thuja occidentalis* plants. It contains 85%  $\alpha$ -thujone and 15%  $\beta$ -thujone<sup>[72]</sup>.

### Biological Activities

Antioxidant<sup>[40,42]</sup>, anti-inflammatory<sup>[73]</sup>, antibacterial, antifungal<sup>[30,37]</sup>, antitumoral<sup>[46-48]</sup>, antidiabetic<sup>[51]</sup>, hypolipidemic, antiatherosclerosis<sup>[52]</sup>, gastroprotective<sup>[49]</sup>, antiviral, immunostimulant<sup>[74,75]</sup>, radioprotector<sup>[53]</sup>, and sedative<sup>[76]</sup> are just a few of the pharmacological properties of *Thuja occidentalis*. The impacts of thujone biological activities have begun to be used in pharmacological and therapeutic research.

#### a. Antipyretic Activity

In rabbits, a methanolic extract of *Thuja occidentalis* has antipyretic action. At dosages of 100 mg and 200 mg/kg body, it lowers fever and normalises body temperature, similar to paracetamol<sup>[54]</sup>.

#### b. Antiviral Activity

Antiviral and immunostimulant properties of polysaccharides isolated from *T. occidentalis* have been demonstrated, with the capacity to suppress HIV-1 and influenza A<sup>[38,78]</sup>.

#### c. Antioxidant Activity

The capacity of phenolic compounds to neutralise free radicals generated by cell metabolism is best defined as antioxidant capability<sup>[77]</sup>. The antioxidant capacity of the *Thuja occidentalis* methanolic extract was estimated to have high DPPH radical scavenging activities, ABTS, NO, and lipid peroxidation tests in a 2016 published research by Nazir et al.<sup>[40]</sup>.

#### d. Anticancer Activity

On the malignant melanoma cell line A375, thujone produced from *T. occidentalis* ethanolic extract was demonstrated to exhibit anticancer effects. Thujone was

found to have an antiproliferative impact and the potential to trigger apoptosis in the same research<sup>[45]</sup>.

#### e. Lipid Metabolism Regulation

At a dosage of 200 mg/kg, the ethanolic fraction of *T. occidentalis* was demonstrated to exhibit hypoglycaemic characteristics in rats with alloxan-induced diabetes, with no effect on body weight. It has also been found to enhance lipid profiles and to protect against oxidative stress by boosting glutathione levels in the blood<sup>[51]</sup>.

#### f. Gastrointestinal Tract protective effect

The ethanol fractions of *T. occidentalis* demonstrated a hepato-protective effect in acute and chronic liver-induced HCV, according to an article. The ethanolic fraction of *T. occidentalis*, according to the same researchers, has a significant impact against stomach lesions<sup>[50]</sup>.

#### g. Radioprotective Activity

*T. occidentalis* was shown to protect Swiss albino mice from gamma-induced toxicity in this investigation. Alkaline phosphatase, pyruvate transferase, and lipid peroxidation were all decreased by the *Thuja occidentalis* alcoholic extract<sup>[53]</sup>.

#### h. Anti-Inflammatory Activity

Inflammation is a basic defensive response that occurs in response to damaging stimuli and is used to restore damaged tissue. Gastric toxicity was not seen at high dosages of aqueous extract and polysaccharide fraction derived from *T. occidentalis* of 300 mg/kg<sup>[73]</sup>.

### REFERENCES

- Adetutu A, Morgan WA, Corcoran O. Ethnopharmacological survey and in vitro evaluation of wound-healing plants used in South-western Nigeria. *Journal of ethnopharmacology*. 2011; 137(1):50-6.
- Shenoy RR, Sudheendra AT, Nayak PG, Paul P, Kutty NG, Rao CM. Normal and delayed wound healing is improved by sesamol, an active constituent of *Sesamum indicum* (L.) in albino rats. *Journal of Ethnopharmacology*. 2011;133(2):608-12.



3. Sanwal R, Chaudhary AK. Wound healing and antimicrobial potential of *Carissa spinarum* Linn. in albino mice. *Journal of Ethnopharmacology*. 2011 Jun 1; 135(3):792-6.
4. Tlili N, Sarikurku C. Bioactive compounds profile, enzyme inhibitory and antioxidant activities of water extracts from five selected medicinal plants. *Industrial Crops and Products*. 2020; 151:112448.
5. Caruntu S, Ciceu A, Olah NK, Don I, Hermenean A, Cotoraci C. *Thuja occidentalis* L.(Cupressaceae): Ethnobotany, phytochemistry and biological activity. *Molecules*. 25(22):5416.
6. Keshav Bharti, Manmohan Sharma, Girish Kumar Vyas, Dr. Shailesh Sharma, *Asian Journal of Pharmaceutical Research and Development*. 2022; 10(2):17-22.
7. Raj M, Vyas GK, Dr. Sharma S, Sharma A, Phyto Analysis, Formulation, And Evaluation of Herbal Lotion Produced from *Allium Sativum* and *Phyllanthus Emblica* Alcoholic Extracts, *Asian Journal of Pharmaceutical Research and Development*. 2022; 10(2):37- 43.
8. Raj M, Vyas GK, Sharma S, Bishnoi H, A Comparative Review on *Allium Sativum* and *Phyllanthus Emblica*, *Asian Journal of Pharmaceutical Research and Development*. 2022; 10(2):77-82.
9. Mirghafourvand M, Mohammad-Alizadeh-Charandabi S, Ahmadvpour P, Javadzadeh Y. Effects of *Vitex agnus* and Flaxseed on cyclic mastalgia: A randomized controlled trial. *Complementary therapies in medicine*. 2016; 24:90-5.
10. Rai MK. Herbal medicines in India: retrospect and prospect. *Fitoterapia (Milano)*. 1994; 65(6):483-91.
11. Harvey AL, Edrada-Ebel R, Quinn RJ. The re-emergence of natural products for drug discovery in the genomics era. *Nature reviews drug discovery*. 2015; 14(2):111-29.
12. Caruntu S, Ciceu A, Olah NK, Don I, Hermenean A, Cotoraci C. *Thuja occidentalis* L. (Cupressaceae): Ethnobotany, phytochemistry and biological activity. *Molecules*. 2020; 25(22):5416.
13. Chang LC, Song LL, Park EJ, Luyengi L, Lee KJ, Farnsworth NR, Pezzuto JM, Kinghorn AD. Bioactive Constituents of *Thuja occidentalis*. *Journal of natural products*. 2000; 63(9):1235-8.
14. Millsbaugh CF. American medicinal plants: an illustrated and descriptive guide to plants indigenous to and naturalized in the United States which are used in medicine. Courier Corporation; 1974.
15. Naser B, Bodinet C, Tegmeier M, Lindequist U. *Thuja occidentalis* (Arbor vitae): a review of its pharmaceutical, pharmacological and clinical properties. Evidence-based complementary and alternative medicine. 2005; 2(1):69-78.
16. Shimada K. Contribution to Anatomy of the Central Nervous System of the Japanese XI. Upon the Vernal Arbor Vitae. *Okajimas Folia Anatomica Japonica*. 1956; 28(1-6):207-27.
17. Baran D. Arbor vitae, a guarantee of health. *Revista medico-chirurgicala a Societatii de Medici si Naturalisti din Iasi*. 1991; 95(3-4):347-9.
18. Offergeld R, Reinecker C, Gumz E, Schrum S, Treiber R, Neth RD, Gohla SH. Mitogenic activity of high molecular polysaccharide fractions isolated from the cupressaceae *Thuja occidentalis* L. enhanced cytokine-production by thyapolsaccharide, g-fraction (TPSg). *Leukemia*. 1992 Jan 1; 6:189S-91S.
19. Naser B, Bodinet C, Tegmeier M, Lindequist U. *Thuja occidentalis* (Arbor vitae): a review of its pharmaceutical, pharmacological and clinical properties. Evidence-based complementary and alternative medicine. 2005 Feb 9;2(1):69-78.
20. Naser B, Bodinet C, Tegmeier M, Lindequist U. *Thuja occidentalis* (Arbor vitae): a review of its pharmaceutical, pharmacological and clinical properties. Evidence-based complementary and alternative medicine. 2005 Feb 9;2(1):69-78.
21. Reitz HD. Immunomodulatoren mit pflanzlichen Wirkstoffen: ein wissenschaftliche Studie am Beispiel Esberitox N. *Notabenmedici*. 1990; 20:362-6.
22. Vorberg G. Bei Erkältungsspezifische Immunabwehrstimulieren. *ÄrztPrax*. 1984; 36:97-8.
23. Verma SK, Singh SK, Singh S, Mathur A. Evaluation of immunomodulatory and microbicidal potential of *Thuja occidentalis*. *Environment Conservation Journal*. 2010; 11(3):85-8.
24. Zimmer M. Gezielte konservative Therapie der akuten Sinusitis in der HNO-Praxis. *Therapiewoche*. 1985;35(36):4024-8.
25. Nahed GA, Balbaa LK. Influence of tyrosine and zinc on growth, flowering and chemical constituents of *Salvia farinacea* plants.
26. Vanherweghem JL, Tielemans C, Abramowicz D, Depierreux M, Vanhaelen-Fastre R, Vanhaelen M, Dratwa M, Richard C, Vandervelde D, Verbeelen D, Jadoul M. Rapidly progressive interstitial renal fibrosis in young women: association with slimming regimen including Chinese herbs. *The lancet*. 1993 Feb 13; 341(8842):387-91.
27. Elsharkawy ER, Aljohar H, Donia AM. Comparative study of antioxidant and anticancer activity of *Thuja orientalis* growing in Egypt and Saudi Arabia. *Br. J. Pharmaceut. Res*. 2017 Jan 1; 15:1-9.
28. Hudson J, Kuo M, Vimalanathan S. The antimicrobial properties of cedar leaf (*Thuja plicata*) oil; a safe and efficient decontamination agent for buildings. *International journal of environmental research and public health*. 2011 Dec;8(12):4477-87.
29. Tsiri D, Graikou K, Poblacka-Olech L, Krauze-Baranowska M, Spyropoulos C, Chinou I. Chemosystematic value of the essential oil composition of *Thuja* species cultivated in Poland—antimicrobial activity. *Molecules*. 2009 Nov;14(11):4707-15.
30. Guleria S, Kumar A, Tiku AK. Chemical composition and fungitoxic activity of essential oil of *Thuja orientalis* L. grown in the north-western Himalaya. *Zeitschrift für Naturforschung C*. 2008 Apr 1;63(3-4):211-4.
31. Başer KH, Demirci B, Demirci F, Koçak S, Akıncı Ç, Malyer H, Gülyeriz G. Composition and antimicrobial activity of the essential oil of *Achillea multifida*. *Planta medica*. 2002 Oct;68(10):941-3.
32. Sivropoulou A, Nikolaou C, Papanikolaou E, Kokkini S, Lanaras T, Arsenakis M. Antimicrobial, cytotoxic, and antiviral activities of *Salvia fruticosa* essential oil. *Journal of Agricultural and Food Chemistry*. 1997 Aug 18;45(8):3197-201.
33. Zhang XW, Choe YH, Park YJ, Kim BS. Effect of Korean arbor vitae (*Thuja koraiensis*) extract on antimicrobial and antiviral activity. *African Journal of Pharmacy and Pharmacology*. 2014 Mar 15;8(10):274-7.
34. Aljos F. A monograph of Cupressaceae and Sciadopitys. Surrey: Royal Botanic Gardens. "Flora of China" editorial board (1978). *Flora of China*. Science. 2005; 7:318.
35. Mu H, Gao Y, Cheng F, Lin L, Wang G, Xia F. Transcriptomic analysis of different tissues in Korean arborvitae. *Dendrobiology*. 2019;81.
36. Bellili S, Aouadhi C, Dhifi W, Ghazghazi H, Jlassi C, Sadaka C, Beyrouthy ME, Maaroufi A, Cherif A, Mnif W. The influence of organs on biochemical properties of Tunisian *Thuja occidentalis* essential oils. *Symmetry*. 2018 Nov;10(11):649.
37. Gohla SH, Zeman RA, Bögel M, Jurkiewicz E, Schrum S, Haubeck HD, Schmitz H, Hunsmann G, Neth RD. Modification of the in vitro replication of the human immunodeficiency virus HIV-1 by TPSg, a polysaccharide fraction isolated from the Cupressaceae *Thuja occidentalis* L.(Arborvitae). In *Modern Trends in Human Leukemia IX 1992* (pp. 140-149). Springer, Berlin, Heidelberg.
38. Asha R, Nisha P, Suneer K, Mythili A, Shafeeq HA, Panneer SK, Manikandan P, Shobana CS. In vitro activity of various potencies of homeopathic drug *Thuja* against molds involved in mycotic keratitis. *Internat. J. Pharm. Pharm. Sci*. 2014; 6:555-9.
39. Nazir MZ, Chandel S, Sehgal A. In vitro screening of antioxidant potential of *Thuja occidentalis*. *J Chem Pharm Res*. 2016; 8:283-6.
40. Stan MS, Voicu SN, Caruntu S, Nica IC, Olah NK, Burtescu R, Balta C, Rosu M, Herman H, Hermenean A, Dinischiotu A. Antioxidant and anti-inflammatory properties of a *Thuja occidentalis* mother tincture for the treatment of ulcerative colitis. *Antioxidants*. 2019 Sep;8(9):416.
41. Dubey SK, Batra A. Antioxidant activities of *Thuja occidentalis* Linn. *Asian J. Pharm. Clin. Res*. 2009;2(1):73-6.
42. Yogesh K, Ali J. Antioxidant potential of *Thuja* (*Thuja occidentalis*) cones and peach (*Prunus persia*) seeds in raw chicken ground meat during refrigerated (4±1 °C) storage. *Journal of food science and technology*. 2014 Aug;51(8):1547-53.
43. Căruntu S, Olah NK, Balta C, ROȘU M, Mlădin B, Hermenean A. *Thuja Occidentalis* Mother Tincture Alleviate The Cox-2 Expression In Intestinal Mucosa Of The Tnbs-Induced Ulcerative Colitis Model In Mice. *Studia Universitatis VasileGoldisSeriastinteleVietii* (Life Sciences Series). 2020 Jul 1; 30(3).
44. Biswas R, Mandal SK, Dutta S, Bhattacharyya SS, Boujedaini N, Khuda-Bukhsh AR. Thujone-rich fraction of *Thuja occidentalis* demonstrates major anti-cancer potentials: evidences from in vitro studies on A375 cells. Evidence-Based Complementary and Alternative Medicine. 2011 Jan 1;2011.
45. Torres A, Vargas Y, Uribe D, Carrasco C, Torres C, Rocha R, Oyarzún C, San Martín R, Quezada C. Pro-apoptotic and anti-angiogenic properties of the α/β-thujone fraction from *Thuja occidentalis* on glioblastoma cells. *Journal of neuro-oncology*. 2016 May;128(1):9-19.
46. Sunila ES, Hamsa TP, Kuttan G. Effect of *Thuja occidentalis* and its polysaccharide on cell-mediated immune responses and cytokine levels of metastatic tumor-bearing animals. *Pharmaceutical biology*. 2011 Oct 1;49(10):1065-73.

47. Sunila ES, Kuttan G. A preliminary study on antimetastatic activity of *Thuja occidentalis* L. in mice model. *Immunopharmacology and Immunotoxicology*. 2006 Jan 1; 28(2):269-80.
48. Das S, Rani R. Antioxidant and gastroprotective properties of the fruits of *Thuja occidentalis* Linn. *Asian J. Biochem. Pharm. Res.* 2013; 3(3):80-7.
49. Dubey SK, Batra A. Hepatoprotective Activity from Ethanol Fraction of *Thuja occidentalis* Linn. *Asian Journal of Research in Chemistry*. 2008; 1(1):32-5.
50. Dubey SK, Batra A. Anti-diabetic activity of *Thuja occidentalis* Linn. *Research Journal of Pharmacy and Technology*. 2008; 1(4):362-5.
51. Dubey SK, Batra A. Role of Phenolics in Anti-Atherosclerotic Property of *Thuja occidentalis* Linn. *Ethnobotanical Leaflets*. 2009; 2009(6):12.
52. Sunila ES, Kuttan G. Protective effect of *Thuja occidentalis* against radiation-induced toxicity in mice. *Integrative Cancer Therapies*. 2005 Dec;4(4):322-8.
53. Aziz A, Khan IA, Ahmed MB, Hussain S. Evaluation of antipyretic activity of *Thuja occidentalis* Linn. in PGE1 and TAB-Vaccine induced pyrexia models in rabbits.
54. Rakotonirainy MS, Lavédrine B. Screening for antifungal activity of essential oils and related compounds to control the biocontamination in libraries and archives storage areas. *International biodeterioration & biodegradation*. 2005 Mar 1;55(2):141-7.
55. Pavela R. Insecticidal activity of some essential oils against larvae of *Spodoptera littoralis*. *Fitoterapia*. 2005 Dec 1;76(7-8):691-6.
56. Selvarani V, James H. The activity of cedar leaf oil vapor against respiratory viruses: Practical applications. *Journal of Applied Pharmaceutical Science*. 2013 Nov 1;3(11):11.
57. Han X, Parker TL. *Arborvitae* (*Thuja plicata*) essential oil significantly inhibited critical inflammation-and tissue remodeling-related proteins and genes in human dermal fibroblasts. *Biochimie open*. 2017 Jun 1; 4:56-60.
58. Lei H, Wang Y, Su C, Liang F, Su W, Hui M, Shaw P, Luo Y. Chemical composition and antifungal activity of essential oils of *Thuja sutchuenensis*, a critically endangered species endemic to China. *Natural Product Communications*. 2010 Oct;5(10):1934578X1000501032.
59. Wang M, Zhao L, Chen K, Shang Y, Wu J, Guo X, Chen Y, Liu H, Tan H, Qiu SX. Antibacterial sesquiterpenes from the stems and roots of *Thuja sutchuenensis*. *Bioorganic Chemistry*. 2020 Mar 1; 96:103645.
60. Tanaka R, Ohtsu H, Iwamoto M, Minami T, Tokuda H, Nishino H, Matsunaga S, Yoshitake A. Cancer chemo preventive agents, labdane diterpenoids from the stem bark of *Thuja standishii* (Gord.) Carr. *Cancer letters*. 2000 Dec 20;161(2):165-70.
61. Kinouchi Y, Ohtsu H, Tokuda H, Nishino H, Matsunaga S, Tanaka R. Potential antitumor-promoting diterpenoids from the stem bark of *Picea glehnii*. *Journal of natural products*. 2000 Jun 23; 63(6):817-20.
62. Iwamoto M, Ohtsu H, Tokuda H, Nishino H, Matsunaga S, Tanaka R. Anti-tumor promoting diterpenes from the stem bark of *Thuja standishii* (Cupressaceae). *Bioorganic & medicinal chemistry*. 2001 Jul 1; 9(7):1911-21.
63. Patel S, Sharma V, S Chauhan N, Thakur M, Dixit VK. Hair growth: focus on herbal therapeutic agent. *Current drug discovery technologies*. 2015 Mar 1;12(1):21-42.
64. Jasuja ND, Sharma SK, Saxena R, Choudhary J, Sharma R, Joshi SC. Antibacterial, antioxidant and phytochemical investigation of *Thuja orientalis* leaves. *Journal of Medicinal Plants Research*. 2013 Jul 3; 7(25):1886-93.
65. Witte L, Berlin J, Wray V, Schubert W, Kohl W, Höfle G, Hammer J. Mono- and diterpenes from cell cultures of *Thuja occidentalis*. *Planta medica*. 1983 Dec; 49(12):216-21.
66. Berlin J, Witte L, Schubert W, Wray V. Determination and quantification of monoterpenoids secreted into the medium of cell cultures of *Thuja occidentalis*. *Phytochemistry*. 1984 May 14;23(6):1277-9.
67. Kawai S, Hasegawa T, Gotoh M, Ohashi H. 4-O-Demethylatein from the branch wood of *Thuja occidentalis*. *Phytochemistry*. 1994 Dec 1;37(6):1699-702.
68. Khubeiz MJ, Mansour G, Zahraa B. Antibacterial and phytochemical investigation of *Thuja orientalis* (L.) leaves essential oil from Syria. *Int. J. Curr. Pharmaceut. Rev. Res.* 2016; 7:243-7.
69. Ismail A, Mohsen H, Bassem J, Lamia H. Chemical composition of *Thuja orientalis* L. essential oils and study of their allelopathic potential on germination and seedling growth of weeds. *Archives of Phytopathology and Plant Protection*. 2015 Jan 2;48(1):18-27.
70. Naser B, Bodinet C, Tegtmeier M, Lindequist U. *Thuja occidentalis* (Arbor vitae): a review of its pharmaceutical, pharmacological and clinical properties. Evidence-based complementary and alternative medicine. 2005 Feb 9;2(1):69-78.
71. Silva IS, Nicolau LA, Sousa FB, de Araújo S, Oliveira AP, Araújo TS, Souza LK, Martins CS, Aquino PE, Carvalho LL, Silva RO. Evaluation of anti-inflammatory potential of aqueous extract and polysaccharide fraction of *Thuja occidentalis* Linn. in mice. *International journal of biological macromolecules*. 2017 Dec 1; 105:1105-16.
72. Caruntu S, Ciceu A, Olah NK, Don I, Hermenean A, Cotoraci C. *Thuja occidentalis* L.(Cupressaceae): Ethnobotany, phytochemistry and biological activity. *Molecules*. 2020 Jan; 25(22):5416.
73. Tegtmeier M, Harnischfeger G. Die Abhängigkeit des Thujongehaltes vom Extraktionsverfahren bei Zubereitungen aus Thujaherba. *Pharmazie*. 1994; 49(1):56-8.
74. Aziz A, AHMAD KI, Hussain M, Raza MA. Pharmacological Evaluation of Sedative activity of methanolic extract of *Thuja occidentalis* in mice.
75. Bodinet C, Freudenstein J. Effects of an orally applied aqueous-ethanolic extract of a mixture of *Thuja occidentalis* herba, *Baptisia tinctoria* radix, *Echinacea purpurea* radix and *Echinacea pallida* radix on antibody response against sheep red blood cells in mice. *Planta medica*. 1999 Dec; 65(08):695-9.
76. Gohlsh, zemanra, gartner s, popovic m, jurkiewicz e, haubeckhd, schrum s, gallorc, nethrd, hunsmann g. Inhibition of the replication of hiv-1 by tpsg, a polysaccharide-fraction isolated from the cupressaceae *Thuja-occidentalis* l. In: *Inaids research and human retroviruses* 1990 jan 1 (vol. 6, no. 1, pp. 131-131). 2 madison avenue, larchmont, ny 10538: maryannliebertainc publ.

77.