

WORLD JOURNAL OF PHARMACEUTICAL AND MEDICAL RESEARCH

www.wjpmr.com

<u>Review Article</u> ISSN 2455-3301 WJPMR

TAXUS BACCATA (STHAUNEYA): POTENTIAL ANTICANCEROUS MEDICINAL PLANT – A REVIEW

Rajput S.*¹, Chaubey S.² and Singh R.³

¹M.D. Scholar, Dept. of *Dravyaguna*, Rishikul, Campus Haridwar, Uttarakhand Ayurved University, Uttarakhand, India.

²Professor, Dept. of *Dravyaguna*, Rishikul Campus, Haridwar, Uttarakhand Ayurved University, Uttarakhand, India. ³Associate Professor, Dept. of *Dravyaguna*, Motherhood Ayurvedic Medical College, Haridwar, Uttarakhand Ayurved University, Uttarakhand, India.



*Corresponding Author: Rajput S.

M.D. Scholar, Dept. of Dravyaguna, Rishikul, Campus Haridwar, Uttarakhand Ayurved University, Uttarakhand, India.

Article Received on 10/10/2024

Article Revised on 31/10/2024

Article Accepted on 21/11/2024

ABSTRACT

Cancer is a serious public health concern all over the world due to a lack of traditional treatments and the harsh side effects of chemotherapy. It is the second greatest cause of death in the world, following heart disorders. Medicinal plants have been used to treat a wide range of ailments since ancient times. Natural products are critical in the fight against cancer and provide a valuable resource for the creation and testing of new therapeutic treatments. This review explains an overview of potential therapeutic effect of *Taxus baccata (Sthauneya)* as anticancer plants. Peclitaxel, an active chemical identified in taxus sp., is important for its capacity to cure various tumors. PTX and its variants are effective against a variety of cancer cell lines.

KEYWORDS: Taxus baccata, Sthauneya, taxol, peclitaxol, anticancerous agent.

INTRODUCTION

Taxus \times media (Taxus \times media Rehder), a member of the genus Taxus in the family Taxaceae, is a plant that combines medicinal, timber, and ornamental values, thus possessing significant economic and research importance.^[1] As a species within the Taxus genus, it plays a crucial role in the ecosystem and is highly valued in the field of medicine due to its rich bioactive compounds.^[2] Many advancements have been made in developing anticancer medications due to studies on the molecular pathways involved in cancer growth. However, despite these efforts, using chemically made medications has not significantly improved overall survival rates. Treating cancer remains a challenging task with limited success. Available treatment choices comprise surgery, radiation therapy, and systemic chemotherapy. In the chemotherapy drug category, medications like methotrexate (antimetabolites), cisplatin and doxorubicin (DNA-interactive drugs), taxanes (antitubulin) are most widely used in addition to other hormones, and molecular targeting drugs.^[3]

ORIGIN AND GEOGRAPHICAL LOCATION

The genus *Taxus* (Taxaceae) contains seven to nine species^[4] among which, *Taxus baccata*, *T. brevifolia*, *T. canadensis*, *T. cuspidata* and *T. floridana* are the most well-known species. *T. baccata* Hook. F. syn. *T.*

wallichiana Zucc is found in Europe, North Africa, the Caucasus, Iran and temperate areas of the Indian subcontinent. Popularly known as Himalayan yew, it is distributed from Pakistan to southwest China, Nepal and Bhutan, mainly at elevations of 1800-3300 m a.s.l. In Indian Himalaya, it is locally called 'Rakhal' or 'Thuner', and is the only species of the genus in Jammu Himachal and Kashmir, Pradesh, Uttarakhand, Meghalaya, Nagaland and Manipur, mainly associated with oak (kharshu) and silver fir species, rarely with spruce, deodar cedar and oak (mohru) in the western Himalaya, and in Eastern Himalaya it is mostly associated with silver fir, hemlock-spruce and rhododendron. This evergreen tree flowers during March-April and seeds ripen from September to November. It can tolerate frost, drought and strong winds, and usually grows on limestone-derived moist soil.^[5]

Yew is a valuable medicinal plant, chiefly for its Taxine and DAB (10-deacetyl baccatin III) content. Initially, the tree was exploited for bark but later demand shifted to leaves, and it is a source of taxine, a precursor to the drug Taxol, used in preparation of Paclitaxel for treatment of ovarian cancer. Oncologists working with taxol regard it as one of the best anticancer agents. According to the Wealth of India.^[5] Taxol was first extracted from bark of *T. brevifolia*, and anti-cancer properties were first reported in 1964.^[7]

BOTANICAL DISCRIPTION

- A small or medium sized evergreen tree, stem fluted, branches horizontal, wide-spreading, not whorled.
- Leaves 2.5-3.8 cm. long, linear, flattened, distichous, acute, narrowed into a short petiole which is decurrent along the twig, dark green and shining above, pale yellowish brown or rusty red below.
- Flowers usually dioecious. Male flowers in catkins which are sub globose and solitary in the leaf axils; stamens about 10, pollen sacs 5-9, globose, arranged

around the filament beneath the peltate tip of the stamens. Female flowers solitary, axillary, resembling leaf- buds, each consisting of a few imbricate scales round a single erect ovule, which is surrounded at the base by a membranous cup-shaped disk.

• In fruit the disk enlarges, becomes succulent and bright red, about 7.5 mm. long, and surrounds the olive-green seed of which only the tip is exposed. Testa woody; embryo with 6-7 cotyledons.^[8]





Fig. 5. Tree



Fig. 3. Leaves



Fig. 4. Bark

Source of fig. 1., fig. 2., fig. 3., fig. 4., fig. 5. from <u>https://identify.plantnet.org/k-world-</u>flora/species/Taxus%20baccata%20L./data

DESCRIPTION AS PER AYURVEDIC TEXTS

Taxus baccata explained in the controversy of *Talish patra*, it is now explained as Sthounayak⁹.

स्थौणेयक के गुण

स्थौणेयं कफवातघ्नं सुगन्धि कटुतिक्तकम्। पित्तप्रकोपशमनं बलपुष्टिविवर्द्धनम् ।। (राज निघण्टु चंदनादि वर्ग/130)^[10]

स्थौणेयकं कटु स्वादु तिक्तं स्निग्धं त्रिदोषनुत। मेधा शुक्रकरं रूच्यं रक्षोघ्नं ज्वरजन्तुजीत्। हन्ति कृष्ठास्रतृडदाहदौर्गन्ध्य तिलकालकान्। (भावप्रकाश निघण्ट् कर्पूरादि वर्ग/110-111)[11]

In Charak Samhita

Agruadi Taila ^[12]
Mritsanjivan Agad ^[13]
Bala Taila ^[14]
Madanphal Utkarika Modak Yog ^[15]

In Sushruta Samhita

Elaadi gana of Sushruta Samhita Sutra 38/24.^[16]

According to Ayurvedic Pharmacopoeia of India^[17]

SYNONYMS

Sanskrit: Sukapuspa, Vikarna Beng.: Birmi, Bhirmie, Talish Patra, Bhada Getela Eng.: Himalayan Yew Guj.: Gethela Barmi Hindi: Thuner, Talispatra Bhed Kan.: Sthauneyak Mal.: Thuriangam, Tuniyankam Mar.: Sthauney Barmi Ori.: Talisabhed, Chalisa Patra Punj.: Birmi Tam.: Talispatri-Bhedam Tel.: Taleesa Patri Bhedamu Urdu.: Birmi, Zarnab

DESCRIPTION

a) Macroscopic: Drug occurs as whole or broken leaf pieces, entire leaf flattened, linear with recurved margins, 1.3-4.0 cm long and 0.1-0.3 cm wide, tip sharp pointed and prickly. entire, thick, brown above, but paler below, petiole, very short: odour, pleasant, taste. acrid, bitter and disagreeable.

b) Microscopic

Leaf

Lamina shows dorsiventral structure, margin slightly turned downward, upper epidermis single layered covered with thick, striated cuticle, lower epidermis single layered with papillate projection, sunken stomata present only on lower surface, overhung by subsidiary cells, palisade two layered, spongy parenchyma 3-5 layered, thin-walled, oval or irregular in shape, containing reddish-brown contents, vascular bundle single, present in the midrib within an endodermis.

Powder- Brown; Shows fragments of reddish-brown spongy parenchyma cells and very rarely xylem tracheid, polygonal epidermal cells with striated cuticle and a few sunken stomata in surface view.

IDENTITY, PURITY AND STRENGTH

Foreign matter-Not more than 2 per cent, Appendix 2.2.2.

Total ash-Not more than 6 per cent, Appendix 2.2.3.

Acid-insoluble ash-Not more than 1.5 per cent, Appendix 2.2.4

Alcohol-soluble extractive-Not less than 10 per cent, Appendix 2.2.6.

Water-soluble extractive-Not less than 16 per cent, Appendix 2.27

CONSTITUENTS- Alkaloids -Taxine, Ephedrine, Glycoside, Tannins, Resins, Reducing Sugars and Formic Acid.

PROPERTIES AND ACTION

Rasa: Katu, Tikta, Madhura Guna:Snigdha, Guru Virya: Sita Vipaka:Madhura Karma: Medhya, Sukravardhaka, Kaphahara, Vatahara, Pittasämaka, Jantughna, Varna Prasadaria, Lomasanjanana

IMPORTANT FORMULATIONS: Mahānārāyana Taila, Balá Taila

THERAPEUTIC USES-Rakta-Vikāra, Trsná, Tila Kalaka, Daha, Kustha, Krmi Roga, Pidika, *Arbuda* (*Karkata*)

DOSE-1-3 g. of the drug in powder form.

ANTICANCER ACTIVITY

Paclitaxel, the principal antitumor component of *Taxus* × *media*, is a diterpenoid compound. Due to its concentration in the bark and leaves, it has been extensively harvested¹⁸ and used in various traditional medical systems to treat many diseases.^[19,20] Paclitaxel inhibits tumor cell mitosis and proliferation by promoting microtubule stabilization. This mechanism of action has led to its widespread use in the treatment of various types of cancer, including ovarian, breast, Kaposi's sarcoma, and lung cancer.^[21]



Fig. 6: From: Paclitaxel and its semi-synthetic derivatives: comprehensive insights into chemical structure, mechanisms of action, and anticancer properties.

Chemical structure of paclitaxel and its derivatives. A Paclitaxel, B 2-debenzoyl-2-trigloyl

paclitaxel, C baccatin and F Abraxane III, D docetaxel, E cabazitaxel,





Fig. 7: From: Paclitaxel and its semi-synthetic derivatives: comprehensive insights into chemical structure, mechanisms of action, and anticancer properties.

Mechanism of anticancer action of Paclitaxel (PTX): PTX intervention leads to (1) stabilization of microtubule, cell arrest, and subsequent apoptosis, (2) inhibition of the TLR4 signaling pathway, (3) increase in the immunomodulatory effects of the drug and (4) activates ER stress-mediated cell death in different cancers. BAK: Bcl-2 homologous antagonist/killer; BAX: Bcl-2 associated X protein; Bcl-2: B cell lymphoma 2; Cyt C: cytochrome C; eIF2 α : eukaryoic translation initiation factor 2 alpha; ER: endoplasmic reticulum; IRAK: interleukin 1 receptor-associated kinase; MAPK: mitogen-activated protein kinase; MEK: MAPK/extracellular signal-regulated (ERK) kinase; MyD88: myeloid differentiation primary response protein 88; NF κ B: nuclear factor light chain enhancer of kappa; PERK: PRKR-like endoplasmic reticulum kinase; PTX: paclitaxel; TLR4: toll-like receptor 4; TRAF: tumor necrosis factor (TNF) receptor-associated factor; TRIF: TIR domain-containing adaptor protein. created Biorender.com



Fig. 8: From: Paclitaxel and its semi-synthetic derivatives: comprehensive insights into chemical structure, mechanisms of action, and anticancer properties.

Paclitaxel vital role in several cancer types.

CONCLUSION

Taxus baccata is commonly utilized as an anticancer plant because it contains *taxol*, which acts as an anticancer agent. It is described in ayurvedic literature as *Sthauneya*, which is disputed by *Talish patra*, but subsequently *Vaidya Bapalal* interpreted it as *Sthauneya*, which is similar to *Taxus baccata*. It is a high-potential plant for usage as an anticancer agent. Researchers have now moved their focus to organic molecules that work as anticancer drugs. Further research is needed to investigate the potential effect of *Taxus baccata* as an anticancer medication.

REFERENCES

- Hao, D.C., Xiao, P.G., Peng, Y., Liu, M. and Huo, L., Research progress and trend analysis of biology and chemistry of Taxus medicinal resources. *Yao xue xue bao= Acta Pharmaceutica Sinica*, 2012; *47*(7): 827-835.
- Li, M., Geng, W., Wang, Z., Wang, Q., Pang, L., Wang, B., Wang, P., Qu, F. and Zhang, X., Analysis of the utilization value of different tissues of Taxus× Media based on metabolomics and antioxidant activity. *BMC Plant Biology*, 2023; 23(1): 285.
- 3. Choudhari, A.S., Mandave, P.C., Deshpande, M., Ranjekar, P. and Prakash, O., Phytochemicals in cancer treatment: From preclinical studies to clinical practice. *Frontiers in pharmacology*, 2020; *10*: 1614.
- 4. Chatterjee, S. and Dey, S., A preliminary survey of the status of Taxus baccata in Tawang District of Arunachal Pradesh. *Indian Forester*, 1997; *123*; 746-754.
- 5. Clapham, A.R., Tutin, T.G. and WARBURG, E.F., 1962. Flora of the British Isles. By AR Clapham... TG Tutin... and EF Warburg... University Press.
- 6. Council of Scientific & Industrial Research (India), 1972. *The Wealth of India: a dictionary of Indian raw materials and industrial products* (Vol. 9). Council of Scientific and Industrial Research.
- 7. Appendino, G., 1993. Taxol (Paclitaxel): historical and ecological aspects.
- K.R. Kirtikar and B.D. Basu, 2005, Indian Medicinal Plants, text, Taxus Linn., 2006; 3: 2383-2385.
- Dr. Bapalal Vaidya, 2014, Some Controversial Drugs in Indian Medicine, chapter 4, *Talisa Patra*, 109-110.
- 10. Dr. Indradev Tripathi, Rajnighantu, edition 5, Chaukhamba Krishandas Academy, Varanasi, Chandanadi varga, Sthounayak, 2006; 422.
- 11. Padamshree Prof. Krishanchandra Chunekar, Bhavprakash Nighantu, Chaukhamba Bharti Academy, Varanasi, 2010, karpuradi Varga, 242-247.
- 12. Sastri K. Agnivesha, Carak Samhita, Revised by Carak and Drdhabala, Vidyotini Hindi Commentery, Chaukhambha Bharati Academy, Varanasi, India, Chikitsasthan, 2020; 2: 143.

- 13. Sastri K. Agnivesha, Carak Samhita, Revised by Carak and Drdhabala, Vidyotini Hindi Commentery, Chaukhambha Bharati Academy, Varanasi, India, Chikitsasthan, 2020; 2: 574.
- 14. Sastri K. Agnivesha, Carak Samhita, Revised by Carak and Drdhabala, Vidyotini Hindi Commentery, Chaukhambha Bharati Academy, Varanasi, India, Chikitsasthan, 2020; 2: 731.
- 15. Sastri K. Agnivesha, Carak Samhita, Revised by Carak and Drdhabala, Vidyotini Hindi Commentery, Chaukhambha Bharati Academy, Varanasi, India, Kalpasthan, 2020; 2: 823-824.
- 16. Kaviraj Dr. Ambikadutt Shastri, Susruta Samhita, Ayurveda Tattva Sandipika, Hindi Commentary, Chaukhamba Sanskrit Sansthan, Varanasi, India, Part 1, Sutrasthan, 2020; 185.
- 17. THE AYURVEDIC PHARMACOPOEIA OF INDIA, PART-I, VOLUME-III, First edition, Government of India, Sthauneya(leaf), 2000; 203-204.
- Alqahtani, F.Y., Aleanizy, F.S., El Tahir, E., Alkahtani, H.M. and AlQuadeib, B.T., Paclitaxel. Profiles of drug substances, excipients, and related methodology, 2019; 44: 205-238.
- 19. Zhu, L. and Chen, L., Progress in research on paclitaxel and tumor immunotherapy. *Cellular & molecular biology letters*, 2019; 24(1): 40.
- 20. Weaver, B.A., How Taxol/paclitaxel kills cancer cells. *Molecular biology of the cell*, 2014; 25(18): 2677-2681.
- Mu, L. and Feng, S.S., A novel controlled release formulation for the anticancer drug paclitaxel (Taxol®): PLGA nanoparticles containing vitamin E TPGS. *Journal of controlled release*, 2003; 86(1): 33-48.