Organization

Medicinal Properties of Illicium Verum Hook. F.

Asra Khan Pahore^{1*}, Shagufta Khan², Nasim Karim³

ABSTRACT

Medicinal plants have active elements that can be used for cures or making drug precursors. According to the World Health Organization, 80% of the world's population depends on these plants as traditional treatments to meet their health care needs. Illicium verum Hook. F. (star anise) is an extensively used spice in traditional Chinese cuisine as a flavoring agent or a historical medicine for vomiting, stomachache, flatulence, and rheumatism and is included in the Chinese Pharmacopoeia. Some active components of star anise are essential oil, anethole, trans-anethole, sesquiterpenoids, shikimic acid, lignans, etc., which possess anxiolytic, neuro-protective, antibacterial, antifungal, anti-inflammatory, antiemetic, anti-flu, antidiarrheal, antispasmodic, anti-diabetic, antioxidant, anti-thrombotic, and antitumour effects that are discussed in this review article.

KEYWORDS: Medicinal, Flavoring agents, Flatulence, Vomiting, Pharmacopoeia, Antiemetic.

INTRODUCTION

Illicium verum (star anise) is a classic scented tree that grows in China and Vietnam^{1,2}. The star anise fruit is a regularly used spice in the Chinese Pharmacopoeia. It has been used as an old Chinese medicine for the treatment of emesis, toothache. depressed liver, flatulence, stomach pain, insomnia, inflammation of the skin, and rheumatic pain. Traditionally, its essential oil is topically applied as an antiseptic3-5. Different biological benefits of star anise are due to its chemical constituents, mainly flavonoids, polysaccharides, trans-anethole, sesquiterpenes, lactones, and shikimic acid. Many favorable functions of shikimic acid have been reported, including anticancer, antibacterial, antidepressant, antioxidant, anti-inflammatory, coagulant, and anti-thrombotic effects. Shikimic acid is also a vital chemical originator of the oseltamivir drug used to treat avian and swine flu⁶. Trans-anethole is another significant component of star anise, which also has many reported therapeutic effects like antianti-hypertensive, thrombotic, anti-inflammatory, antifungal, anti-cataract, and anti-obesity effects^{7,8}.

Medicinal plants

According to primitive records, it has been almost 60,000 years since humans used plants medicines9. Medicinal plants have components that can be used for curative purposes or making drug

estimates that more than 80% of the world's population uses plants as a traditional medicine to fulfil their primary healthcare needs¹¹. In traditional cultures, medicinal plants have played a significant role in reducing human suffering, and preference is usually given to them due to their efficacy in some diseases, easy accessibility, cost-effectiveness, and cultural acceptance¹². The different parts of medicinal plants that can be used are seeds, leaves, roots, fruits, flowers, skin, or a complete plant. Active compounds are present in the bodies of these plants, which are used as medicinal drugs and produce direct or indirect biological effects on living bodies¹³.

World

Health

The

Illicium Verum

precursors¹⁰.

Illicium verum (i.verum) is an average-sized, timeless tree native to Asian tropical and sub-tropical areas¹⁴. Due to the similarity of the anise flavor from Illicium verum star-shaped pericarp fruit, it is also known as star anise, Chinese star anise, and badian. It is extensively cultivated commercially in China, India, and many other Asian countries. It is a commonly used spice in Chinese, Indian, Malay, and Indonesian cuisines. In the Indian subcontinent, star anise is a known spice to make biryani and masala chai. It is one of the significant constituents of traditional fivespice powder in China. In Vietnam, it is also an essential ingredient in noodle soup¹⁵.

Botanicals

I.verum is categorized in the class Magnoliopsida and the family Illiciaceae. Its plant is an average-sized shrub, 8 to 15 meters high and 30 centimetres deep. The bay is either white or silver. Leaflets are simple, glossy, leathery, alternative, and 6 to 12 centimetres long. Flowers are oversized, 1 to 1.5 centimetres in diameter, bisexual, pinkish white to red or vellow with a hint of green, solitary, and axillary. Fruit is capsule type and forms clusters in a star shape (Figure I). Each arm is a seed pod. The picking of fruit occurs before it is fully grown and dries. Seeds are sparkly

Correspondence: asra.pahore@gmail.com

doi: 10.22442/jlumhs.2023.00962

Received: 29-04-2022 Revised: 16-05-2023 Accepted: 17-05-2023 Published Online: 31-07-2023



¹Department of Pharmacology, Altamash Institute of Dental Medicine, Karachi, Sindh-Pakistan.

²Department of Biological & Biomedical Sciences, Aga Khan University, Karachi, Sindh-Pakistan.

³Department of Pharmacology, Bahria University Medical & Dental College, Sailors Street, Adjacent PNS Shifa, DHA Phase II, Karachi, Sindh-Pakistan.

Pahore et al.

coffee-coloured or red with a significant amount of oil.

The flowers of star anise mature from March to May, while the fruits are full-flavoured from September to October 16,17.

Figure I: Morphology of the dried fruit of Illicium verum Hook. F.

Dried fruit of Illicium Verum

Illicium verum dried fruit contains alkaloids, essential oils, tannins (9-10%), cis-anethole, trans-anethole (85 -90%), Limone, α-pinene, α-terpinoil, β-phellandrene and farnesol^{18,19}. Out of these, anethole and fatty oil are the primary components of star anise. Anethole is an organic bioactive compound with many medicinal effects like anti-inflammatory, antioxidant, antibacterial, anticancer, anti-spasmotic, anti-thrombotic, anti-diabetic, hypotensive, neuroprotective, and immuno-modulatory. Star anise oil is topically used for rheumatic pain, earache, fever, scables, insomnia, and constipation and as an antiseptic^{20,21}. Star anise can be used internally and externally in many preparations. Its most common internal use is dyspepsia, while its external use is topical inhalation for respiratory tract congestion. It is also a significant component of anti-tussive drugs and is currently used as a medicine for the flu (**Table I)** ²².

Table I: Components of *Illicium verum* and their pharmacological actions²²

Components	Pharmacological actions
α- Pinene	Anti-inflammatory and anti- cholinestrase activity
α- Terpineol	Anti-tumor, anti-convulsant and antioxidant activity
β-Phellandrene	Flavoring agent
Anethole	Antibacterial and antifungal activity
Lignans	Anti-inflammatory and anti-tumor activity
Phenylpropanoids	Protective activity against pathogens
P-coumaric acid	Anti-tumor and antioxidant activity
Quercetin	Maintenance of cardiac and mental health
Quercetin-3-o-alpha- d-xyloside	Antioxidant activity
Sesquiterpenoids	Protective agents
Trans-anethole	Antibacterial and antifungal activity
Shikimic acid	Anti-coagulant and anti-thrombotic activity
Terpinen-4-ol gamma	Antibacterial and antifungal activity

Medicinal Uses of Illicium Verum

Antifungal effect

The crucial oil of *I.verum* comprises monoterpenes that result in antifungal activity by causing degradation of the

fungi's cell wall, injury to the cytoplasmic membrane, and ultimately, protein destruction²³. It has been reported that an absolute and 50% methanol extract of *I.verum* have shown significant zone inhibition of *Aspergillus fumigatus* with 14.3mm and 10 mm diameters, respectively²⁴.

Antioxidant effect

The star anise fruit essential oil has shown its antioxidant activity through DPPH (1,1-diphenyl-2-picrylhydrazyl) and ABTS (2, 2-azino-bis(3-ethylbenzothiazoline-6-sulfonic acid)) assays with an IC $_{50}$ value of 47µg/ml and 75.9µg/ml, respectively 25 . Another study has shown the antioxidant activity of a methanolic extract of star anise using DPPH with an IC $_{50}$ value of 61µg/ml 26 .

Anti-diabetic effect

The seed extract of star anise has anti-diabetes action through in vitro dose-dependent inhibition of α -amylase²⁷.

Antibacterial effect

The antibacterial activity of star anise is due to its components like anisyl aldehyde, alcohol, and acetone, which inhibit the growth of *Pseudomonas aeruginosa*, *Staphylococcus aureus*, and *Acinetobacter baumannii*²⁸. The aqueous methanol extract of star anise also showed significant inhibition of bacterial growth with a zone diameter of 13mm in both *Acinetobacter baumannii* (AB5057) and *Methicillin-resistant Staphylococcus aureus* (MRSA USA 300).²⁹

Anti-inflammatory effect

The essential oil of *I.verum* produces an anti-inflammatory effect similar to an antioxidant effect through the production of free radicals, inhibition of lipid peroxidation, and protein breakdown³⁰. Furthermore, different concentrations of star anise extract prevent heat-induced albumin breakdown³¹. Star anise component trans-anethole also possesses anti-inflammatory activity by making fewer IL1b, IL6, MDC (macrophage-derived chemokine), and TARC (thymus activation regulated chemokine)³².

Antiemetic effect

Flavonoids, phenolic acid, and alkaloid components of star anise have been reported to produce effects against emesis³³.

Anti-flu effect

In plants, bacteria, and fungi, shikimic acid is produced as an intermediate product of amino acid synthesis, while in mammals, this mechanism is absent. Shikimic acid is present in the star anise fruit and is a primary production source. It is the precursor for forming oseltamivir against avian and swine influenza, both A and B (H1N1 and H5N1) ^{34,35}. It also relieves blockage of the respiratory tract, inflammation of the bronchi, and whooping cough ³⁶.

Neuroprotective effect

According to the cholinergic hypothesis, the abnormal cholinergic transmission of acetylcholine in the central nervous system is the cause of memory loss in Alzheimer's disease. Cholinesterase inhibitors are the first option for treating Alzheimer's disease. A significant proportion of trans-anethole in star anise gives it its anti-cholinesterase properties³⁷.

Sedative and anxiolytic effects

It has been reported that the sedative and anxiolytic activity of star anise is due to the presence of trans-anethole and water extract, which cause the initiation of the GABA-nergic inhibitory system³⁸.

Antispasmodic

Star anise is well known for its anti-flatulence and eupeptic effects³⁹. It is also a famous herbal medicine for treating infant colic. Its antispasmodic action is due to anethole and

Pahore et al.

terepene hydrocarbons (limonene, phellandrene, and dipentene) $^{\rm 40}.$

Antidiarrheal effect

The presence of anethole and dicycloether in star anise has the potential for antidiarrheal effects. Also, flavonoids and glycosides of star anise may have protective action against the damaging impact of ricinoleic acid on the mucosa, so severe diarrhea does not happen⁴⁰.

Anti-thrombotic effect

The rise in prothrombin time and activated partial thromboplastin time occurs due to the methanol extract of star anise. The foremost mechanism of warfarin is that it inhibits the vitamin K epoxide reductase enzyme, which is essential for clotting factors 2, 7, 9, and 10. It is probably true that star anise also has a similar mechanism of action as warfarin⁴¹.

Anticancer effect

It has been reported that essential oils from *I.verum* fruit possess anti-proliferative activity against a human colon cancer cell line with an IC $_{50}$ value of 50.34µg/ml by inducing apoptosis and controlling cell metastasis. Another study showed the trans-Anethole component of *I.verum* has an anti-proliferative effect in human osteosarcoma cell lines with an IG $_{50}$ value of 60.25µM. It also restricted the cell cycle in the G1 and G0 phases by producing reactive oxygen species and reducing mitochondrial membrane potential 25,42 .

CONCLUSION

Even though it is frequently utilized, the precise mechanisms underlying some of its therapeutic qualities and any possible adverse effects are still not completely understood. Therefore, further clinical research must be done to understand better the potential of this natural medicine to replace synthetic drugs. These studies may provide fresh insights into the efficacy and safety of *I.verum*, which may assist in advancing medical practices and better meeting patient demands.

Conflict of Interest: No conflicts of interest.

Financial Disclosure / Grant Approval: No funding agency was involved in this research.

Data Sharing Statement: The corresponding author can provide the data proving the findings of this study on request. Privacy or ethical restrictions bound us from sharing the data publically.

AUTHOR CONTRIBUTIONS

Pahore Ak: Designed and wrote the original draft of the manuscript

Khan S: Review of manuscript, editing, approval of final draft Karim N: Review of manuscript, editing, approval of final draft

REFERENCES

- Elmasry TA, Al-Shaalan NH, Tousson E, El-Morshedy K, Al-Ghadeer A. Star anise extracts modulation of reproductive parameters, fertility potential and DNA fragmentation induced by growth promoter Equigan in rat testes. Braz J Pharm Sci. 2018; 54(1): 17261-70. doi: 10.1590/s2175-97902018000117261.
- Destro BG, Jorge RM, Mathias AL. Optimization of high -concentration trans-anethole production through hydro distillation of star anise. Braz J Chem Eng. 2019; 36(2): 823-30. doi: 10.1590/0104-6632.20190362s20180203.
- Boota T, Rehman R, Mushtaq A, Kazerooni EG. Star anise: A review on benefits, biological activities and

- potential uses. Int J Chem Biochem. 2018; 14: 110-4.
- Peng W, Lin Z, Wang L, Chang J, Gu F, Zhu X. Molecular characteristics of Illicium verum extractives to activate acquired immune response. Saudi J Biol Sci. 2016; 23(3): 348-52. doi: 10.1016/j.sjbs.2015.10.027.
- Nguyen TT, Trinh NY, Le PT. Recovery Yield and Bioactivities Evaluation on Essential Oil and Ethanolic Extract of Star Anise (Illicium verum Hook. f.). Chem Eng Trans. 2021; 83: 205-10. doi: 10.3303/CET2183 035.
- Li SN, Sun JF, Wang JM, Jin L, Zong TQ, Zhou W et al. Two new phenolic glycosides from the fruits of Illicium verum. J Asian Nat Prod Res. 2022; 24(1): 31-8. doi: 10.1080/10286020.2021.1871606.
- Shi Y, Chen G, Chen K, Chen X, Hong Q, Kan J. Assessment of fresh star anise (Illicium verum Hook. f.) drying methods for influencing drying characteristics, color, flavor, volatile oil and shikimic acid. Food Chem. 2021; 342: 128359-70. doi: 10.1016/j.foodchem.2020. 128359.
- Sharafan M, Jafernik K, Ekiert H, Kubica P, Kocjan R, Blicharska E et al. Illicium verum (Star Anise) and Trans -Anethole as Valuable Raw Materials for Medicinal and Cosmetic Applications. Molecules. 2022; 27(3): 650-65. doi: 10.3390/molecules27030650.
- de Oliveira Silva MR, Santos MH. Medical plants: from Colonial Brazil to their indication by the National Single Health System–SHS. Int J Adv Eng Res Sci. 2019; 6(9): 165-173. doi: 10.22161/ijaers.69.19.
- Boy HI, Rutilla AJ, Santos KA, Ty AM, Alicia IY, Mahboob T et al. Recommended medicinal plants as source of natural products: a review. Digital Chin Med. 2018; 1(2):131-42. doi:10.1016/S2589-3777(19)30018-7.
- 11. Veiga M, Costa EM, Silva S, Pintado M. Impact of plant extracts upon human health: A review. Crit Rev Food Sci Nutr. 2020; 60(5): 873-86. doi: 10.1080/10408398. 2018.1540969.
- Nankaya J, Gichuki N, Lukhoba C, Balslev H. Medicinal plants of the Maasai of Kenya: a review. Plants. 2019; 9 (1): 44-60. doi: 10.3390/plants9010044.
- Jamshidi-Kia F, Lorigooini Z, Amini-Khoei H. Medicinal plants: Past history and future perspective. J HerbMed Pharmacol. 2017; 7(1): 1-7. doi: 10.15171/jhp.2018.01.
- Assiry AA, Karobari MI, Bhavikatti SK, Marya A. Crossover analysis of the astringent, antimicrobial, and anti-inflammatory effects of Illicium verum/star anise in the oral cavity. Biomed Res Int. 2021; 2021: 1-6. doi: 10.1155/2021/5510174.
- 15. Sachan AK, Kumar S, Kumari K, Singh D. Medicinal uses of spices used in our traditional culture: Worldwide. J Med Plants Stud. 2018; 6(3): 116-22.
- Vecchio MG, Gulati A, Minto C, Lorenzoni G. Pimpinella Anisum and Illicium Verum: The Multifaceted Role of Anise Plants. Open Agric. 2016; 10(1): 81-6. doi: 10.2174/1874331501610010084.
- 17. Shahrajabian MH, Sun W, Cheng Q. Survey on Chemical Constituent, Traditional and Modern Pharmaceutical and Health Benefits of Chinese Star Anise, a Treasure from the East. Phcog Commn. 2021; 11(1): 31-5. doi: 10.5530/pc.2021.1.7.
- 18. Patrá JK, Das G, Bose S, Banerjee S, Vishnuprasad CN, del Pilar Rodriguez-Torres M et al. Star anise (Illicium verum): Chemical compounds, antiviral properties, and clinical relevance. Phytother Res. 2020; 34(6): 1248-67. doi: 10.1002/ptr.6614.
- 19. Nguyen TT, Trinh NY, Le PT. Recovery Yield and

- Bioactivities Evaluation on Essential Oil and Ethanolic Extract of Star Anise (Illicium verum Hook. f.). Chem Eng Trans. 2021; 83: 205-10. doi: 10.3303/CET218 3035.
- Shahrajabian MH, Sun W, Cheng Q. Chinese star anise (Illicium verum) and pyrethrum ('Chrysanthemum cinerariifolium') as natural alternatives for organic farming and health care-a review. Aust J Crop Sci. 2020; 14(3): 517-23. doi: 10.21475/ajcs.20.14.03. p2209.
- 21. Zhang G, Yuan C, Sun Y. Effect of selective encapsulation of hydroxypropyl-β-cyclodextrin on components and antibacterial properties of star anise essential oil. Molecules. 2018; 23(5): 1126-40. doi: 10.3390/molecules23051126.
- Sayyar HT. Exotic Spice Illicium Verum Hook. F. J Bahria Univ Med Dent Coll. 2018; 8(3): 189-93. doi: 10.51985/JBUMDC2018044.
- 23. Yu C, Zhang J, Wang T. Star anise essential oil: Chemical compounds, antifungal and antioxidant activities: a review. J Essent Oil Res. 2021; 33(1): 1-22. doi: 10.1080/10412905.2020.1813213.
- Alhajj MS, Qasem MA, Nabi AR, Al-Mufarrej SI. In-vitro antibacterial and antifungal effects of high levels of Chinese star anise. Braz J Poultry Sci. 2019; 21(1): 001-008. doi: 10.1590/1806-9061-2016-0427.
- 25. Asif M, Yehya AH, Al-Mansoub MA, Revadigar V, Ezzat MO, Ahamed MB et al. Anticancer attributes of Illicium verum essential oils against colon cancer. S Afr J Bot. 2016; 103: 156-61. doi: 10.1016/j.sajb.2015.08.017.
- Bhatti HA, Khan S, Faizi S, Abbas G, Ali I, Jawaid S et al. Protocatecheuic acid underlies the antioxidant activity exhibited by Illicium verum fruit. J Anal Pharm Res. 2017; 6(3): 00177-81. doi: 10.15406/japlr.2017.06. 00177.
- Anise PE. Antioxidant and Anti-Diabetic activities of Polyphenol-enriched Star Anise (Illicium verum) seeds extract. Int J Biotechnol Biochem. 2018; 14(2): 77-84.
- Yang EC, Hsieh YY, Li-Yeh C. Comparison of the Phytochemical Composition and Antibacterial Activities of the Various Extracts from Leaves and Twigs of Illicium verum. Molecules. 2021; 26(13): 3909-24. doi: 10.3390/molecules26133909.
- Salem MA, El-Shiekh RA, Hashem RA, Hassan M. In vivo Antibacterial Activity of Star Anise (Illicium verum Hook.) Extract Using Murine MRSA Skin Infection Model in Relation to Its Metabolite Profile. Infect Drug Resist. 2021; 14: 33-48. doi: 10.2147/IDR.S285940.
- Luís Â, Sousa S, Wackerlig J, Dobusch D, Duarte AP, Pereira L et al. Star anise (Illicium verum Hook. f.) essential oil: Antioxidant properties and antibacterial activity against Acinetobacter baumannii. Flavour Fragr J. 2019; 34(4): 260-70. doi: 10.1002/ffj.3498.

- 31. Paul R, Geetha RV. Evaluation of anti-inflammatory action of Illicium verum-An in vitro study. Drug Discov Today. 2018; 10(12): 2441-2444.
- Sung YY, Kim YS, Kim HK. Illicium verum extract inhibits TNF-α-and IFN-γ-induced expression of chemokines and cytokines in human keratinocytes. J Ethnopharmacol. 2012; 144(1): 182-9. doi: 10.1016/j. jep.2012.08.049.
- 33. Maimoona A. Phytochemical analysis and antiemetic activity of Illicium verum Hook. f. fruit by chick emesis model. J Pharmacogn Phytochem. 2016; 5(6): 185-8.
- 34. Kshirsagar PR, Pai SR, Vyavahare GD, Bapat VA, Desai NS. Prospecting alternative source of shikimic acid from clusiaceae of Western Ghats, India and evaluating discrepancies with reference to seasonal variation, developmental stage and sex of Mammea suriga. Ind Crops Prod. 2020; 149: 112354-61. doi: 10.1016/j.indcrop.2020.112354.
- Díaz-Quiroz DC, Cardona-Félix CS, Viveros-Ceballos JL, Reyes-González MA, Bolívar F, Ordoñez M et al. Synthesis, biological activity and molecular modelling studies of shikimic acid derivatives as inhibitors of the shikimate dehydrogenase enzyme of Escherichia coli. J Enzyme Inhib Med Chem. 2018; 33(1): 397-404. doi: 10.1080/14756366.2017.1422125.
- 36. Palai S, Dehuri M, Patra R. Spices boosting immunity in COVID-19. Ann Phytomed. 2020; 9(2): 80-96.
- Sayyar HT, Raza ML, Baqir SR. Evaluation of Antiamnesic and Cholinesterase inhibitory effects of Illicium verum hook. f (Star anise) against Scopolamine induced memory impairment in Mice. J Bahria Univ Med Dent Coll. 2023; 13(01): 08-12.
- Chouksey D, Upmanyu N, Pawar RS. Central nervous system activity of Illicium verum fruit extracts. Asian Pac J Trop Dis. 2013; 6(11): 869-75. doi: 10.1016/j.fitote. 2010.11.003.
- McCallum RW, González-Stuart AE. Medicinal Plants Used for Digestive Disorders - What the Gastroenterologist Needs to Know. Pract Gastroenterol. 2015; 13: 12-25.
- Díaz A, Vargas-Perez I, Aguilar-Cruz L, Calva-Rodríguez R, Treviño S, Venegas B et al. A mixture of chamomile and star anise has anti-motility and antidiarrheal activities in mice. Rev Bras Farmacogn. 2014; 24: 419-24. doi: 10.1016/j.bjp.2014.07.016.
- Shaikh TA. Effect Effects of Methanol Extract of Illicium verum hook. f on Coagulation Parameters: Effects of Methanol Extract of Illicium verum hook. f. Pak J Med Dent. 2020; 9(1): 22-8. doi: 10.36283/PJMD9-1/006.
- Pandit K, Kaur S, Kumar A, Bhardwaj R, Kaur S. Trans-Anethole abrogates cell proliferation and induces apoptosis through the mitochondrial-mediated pathway in human osteosarcoma cells. Nutr Cancer. 2021; 73 (9): 1727-45. doi: 10.1080/01635581.2020.1803927.

