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Abstract

Myristica fragrans is an aromatic evergreen tree which belongs to Myristicaceae family. Nutmeg (seed) and Mace (arillus) which are two separate spices derived from the fruit of tree Myristica fragrans used as spices in culinary and in traditional systems of medicine. This study was carried out to give an overview on Myristica fragrans according to traditional systems of medicine and to review the recent scientific evidences of phytochemical and pharmacological studies systematically. While reviewing the literature, it reveals Myristica fragrans shows therapeutic actions such as appetizer, carminative, digestive, stomachic and aphrodisiac activities and Myristica fragrans shows therapeutic uses such as indigestion, loss of appetite, diarrhoea, cough, asthma and sexual debility. Numerous studies have indicated that M. fragrans contains diverse phytochemicals such as Myristicin, Myristic Acid, Trimyristin, Elemicin, Safrole, Lignans, Neolignans, Maceneolignans etc. which exhibit many of pharmacological activities such as Anti-allergic activity, Antibacterial activity, Anti-cancer activity, Anticonvulsant activity, Anti-diabetic activity, Anti-diarrhoel activity, Anti-depressant activity, Antifungal activity, Anti-inflammatory activity, Anti-microbial activity, Anti-oxidant activity, Analgesic activity, Aphrodisiac activity, Gastro-protective activity, Hepatoprotective activity, and Immunomodulatory effect. Myristicin, Myristic Acid, Trimyristin are the most active compounds among them. The aim of this review is to comprehensively summarize the phytochemical and pharmacological properties of Myristica fragrans that have reported to date.

Keywords: Jaiphal, Javitri, Jatiphala, Jayapatri, mace, nutmeg

Introduction

A spice is a dried seed, fruit, root, bark or flower of a plant or a herb used in small quantities for flavor, color or as a preservative. The spices and Herbs used for flavor, aroma and medicinal properties derive a special value from the said factors ^[1]. Spices and herbs have been in use for centuries both for culinary and medicinal purposes. Spices not only enhance the flavor, aroma, and color of food and beverages, but they can also protect from acute and chronic diseases. Long before modern medicine, spices were valued for their ability to help individuals in disease prevention and health promotion ^[2].

Myristica fragrans is an aromatic evergreen tree which belongs to Myristicaceae family. Nutmeg (seed) and Mace (arillus) which are two separate spices derived from the fruit of tree *Myristica fragrans*. Nutmeg is the seed kernel inside the fruit and mace is the red lacy covering (aril) on the kernel. These are used as spices in culinary and in traditional systems of medicine ^[3].

Methodology

A systematic literature search was carried out to review articles and to gather the information available in the literature regarding *Myristica fragrans* in the view of description of the plant, chemical constituents, part used, therapeutic action and therapeutic uses, and recent scientific evidences of phytochemical and pharmacological activities. All the available information on *Myristica fragrans* was compiled from Unani, Ayurveda and Siddha textbooks & Pharmacopoeias and electronic databases such as Google scholar and PubMed.

Results

Scientific Classification of *Myristica fragrans*^[4] Kingdom: Plantae Division: Tracheophytes Class: Magnoliopsida Order: Magnoliales

Corresponding Author: MMM Nifras Demonstrator, Institute of Indigenous Medicine, University of Colombo, Sri Lanka Family: Myristicaceae Genus: Myristica Species: *M. fragrans* Houtt Botanical name: *Myristica fragrans Houttuyn*^[5]. Synonyms: M. *moschata* Thunb., *M. officinalis* Linn., *M. armocatica* Lamk^[5].

Vernacular names [5]

English: Nutmeg, Mace tree Tamil: Sadikkay Sinhala: Sadikka (Nutmeg), Wasawasi (Arillus/Mace) Unani Tibbi name: Jaiphal, Javitri/ Bisbasa ^[6] (Arillus/Mace) Sanksrit name: Jatiphala, Jayapatri ^[7] (Arillus/Mace)



Fig 1: Seeds of Myristica fragrans



Fig 2: Arills/ Mace of Myristica fragrans

Description of the plant^[5]

A moderate sized evergreen tree, 8-13 m high with numerous spreading branches covered with greyish-brown rather smooth bark, young branches green;

Leaves: simple, alternate, shortly petiolate, estipulate, somewhat convex above, 10-15cm long, ovate or oblong-

ovate, acute at both ends, entire, smooth, strongly veined, dark green and paler beneath;

Flowers: regular, unisexual, yellowish, dioecious in small axillary racemes of 2-6 flowers, flowers sometimes solitary in the female tree, pedicels slender, drooping with a single, quickly deciduous, rounded bract just below the flower; perianth about 1cm long, fleshy, bell-shaped or urceolate, nearly smooth, pale yellow, cut into 3 (rarely 4) spreading or erect, triangular, acute teeth, aestivation valvate;

Fruit: pendulous, about 7.5cm by 5cm, grooved by a longitudinal furrow which passes through the somewhat lateral apiculus marking the positions of the stigmas, smooth, yellow; pericarp nearly I.2 cm thick, tough and fleshy, yellowish-white, dehiscing from above along the furrow into two equal halves and containing a single erect seed which completely fills the cavity, out of which it readily falls when ripe;

Seed: about 3 cm long, broadly ovoid, blunt, closely enveloped and almost completely covered by an irregularly cut, fleshy arillus (mace) which is cup-shaped round the basal hilum and much folded over the top of the seed, brilliant scarlet when fresh, but yellow and brittle when dry, testa very hard and thick, dark brown, smooth and shining, marked with impressions from the tightly ariilus, inner seed coat thin, membranous, pale brown, nucleus of the seed (nutmeg) wrinkled externally, mainly consisting of the abundant endosperm, which is rather soft but firm, whitish and marbled with numerous reddish-brown vein-like partitions into which the inner seed coat penetrates (ruminated), embryo at base of the hilum, cotyledons foliaceous, lacineated.

Parts used

Seed, Arillus^[6] (mace)

Chemical Constituents

The principal constituents of Nutmeg are a Fixed oil, Volatile oil 3% containing D-pinene, Myristin, Myristic acid and its Ester, Myristicin, Fatty acids etc^[7].

Properties of *Myristica fragrans* according to Unani, Ayurveda and Siddha systems of medicine

Table 01 shows the properties of the *Myristica fragrans* according to the traditional systems of medicine.

Table 1: Properties of Myristica fragrans according to Unani, Ayurveda and Siddha systems of medicine

| Unani ^[6] | Ayurveda ^[7, 8] | Siddha ^[9] |
|---------------------------------------|--|---|
| 1. Taste: | 1. Rasa (Taste): | 1. Cuvai (Taste): |
| Pungent and slightly bitter (Nutmeg), | Katu (Pungent), Tikta (Bitter) and Astringent | Kaarppu (Pungent), Thuvarppu (Astringent) |
| Slightly bitter (Arillus/Mace) | 2. Guna (Attribute): | 2. Gunam (Character): |
| 2. Mizaj (Temperament): | Laghu (Light), Snigdha (Oily) and Tiksna (Sharp) | Ilaku (Light), |
| Hot 2^0 Dry 3^0 | 3. Virya (Potency): | Koormai (Sharp) |
| | Usna (Hot) | 3. Virium (Potency): |
| | 4. Vipaka | Veppam (Hot) |
| | (Post digestive effect): | 4. Pirivu (Class): |
| | Katu (Pungent) | Kaarppu (Pungent) |

Therapeutic actions of *Myristica fragrans* according to **Unani, Ayurveda and Siddha systems of medicine** Table 02 shows the Therapeutic actions of *Myristica* fragrans according to the traditional systems of medicine.

Table 2: Therapeutic actions of Myristica fragrans according to Unani, Ayurveda and Siddha systems of medicine

| Unani ^[6] | Ayurveda ^[7, 8] | Siddha ^[9] |
|---|---------------------------------------|---|
| Jaiphal (Nutmeg) | 1. Deepana (Appetizer) | 1. Akadduvayvakatri (Carminative) |
| 1. Muqawwi-e-Meda (Stomachic) | 2. Digestant | 2. Kaamamperukki (Aphrodisiac) |
| 2. Muqawwi-e-Bah (Aphrodisiac) | 3. Grahi (Anti-diarrhoeal) | 3. Manamootti (Aromatic) |
| 3. Muqawwi (Tonic) | 4. Vrsya (Aphrodisiac) | 4. Moorchchaiyundaakki (Sedative/ Hypnotic) |
| 4. Muhallil-e-Waram (Anti-inflammatory) | 5. Mukhalednasaka | 5. Uramaakki (Nutrient) |
| Javitri/ Bisbasa (Arillus/Mace) | 6. Mukhadaurgandhyasnasaka | 6. Veppamundaakki (Stimulant) |
| 1. Muqawwi-e-Meda (Stomachic) | (Anti-hallitosis) | |
| 2. Hazim (Digestive) | 6. Kaphavatapana (Alleviates Vata and | |
| 3. Kasir-e-Riyah (Carminative) | Kapha dosas) | |
| 4. Muqawwi-e-Qalb (Cardiotonic) | 7. Stambhana | |

Therapeutic uses of *Myristica fragrans* according to Unani, Ayurveda and Siddha systems of medicine

according to the traditional systems of medicine.

Table 03 shows the Therapeutic uses of Myristica fragrans

Table 3: Therapeutic uses of Myristica fragrans according to Unani, Ayurveda and Siddha systems of medicine

| Unani ^[6] | Ayurveda ^[7, 8] | Siddha ^[9] |
|------------------------------------|-------------------------------|--|
| | 1. Atisara (Diarrhoea) | 1. Pasiththeekkuraivu (Loss of Appetite) |
| | 2. Svasa (Asthma) | 2. Iraippu (Tuberculosis) |
| Jaiphal (Nutmeg) | 3. Chardi (Vomiting) | 3. Irumal (Cough) |
| 1. Ishal (Diarrhoea) | 4. Kasa (Cough) | 4. Naalpatta Kazhichchal (Chronic Dysentery) |
| 2. <i>Falij</i> (Paralysis) | 5. Pinasa (Catarrh) | 5. Perunkazhichchal (Diarrhoea) |
| 3. Wajaul Mafasil (Arthritis) | 6. Grahani | 6. Vinthuk Kuraivu (Oligospermia) |
| | 7. Mukharoga (Oral diseases) | |
| Javitri/ Bisbasa (Arillus/Mace) | 8. Sukrameha (Spermatorrhoea) | |
| 1. Amraz-e-Qalb (Cardiac diseases) | 9. Loss of appetite | |
| 2. Su-e-Hazm (Indigestion) | 10. Indigestion | |
| 3. Zof-e-Bah (Sexual debility) | 11. Worm infestation | |
| | 12. Premature ejaculation | |
| | 13. Dysmenorrhea. | |

Compound formulations of *Myristica fragrans* according to Unani, Ayurveda and Siddha systems of medicine

fragrans according to the traditional systems of medicine.

Table 04 shows the Compound formulations of Myristica

Table 4: Compound formulations of Myristica fragrans according to Unani, Ayurveda and Siddha systems of medicine

| Unani ^[6] | Ayurveda ^[7, 8] | Siddha ^[9] |
|---------------------------------------|------------------------------------|---|
| Jaiphal (Nutmeg) | 1. Jatiphaladi Curna | 1. Ashtapayiravak Kulikai |
| 1. Laboob-e-Kabir | Jatiphaladi Vati | 2. Saampiranippoo Pathangam |
| 2. Laboob-e-Asrar | | Surapunga Vilvathi Ilakam |
| Jawarish-Ood-Shirin | | 4. Ilaku Santhanaathi Thailam |
| 4. Habb-e-Mumsik. | | 5. Kapaada Maathirai |
| Javitri/ Bisbasa (Arillus/Mace) | | |
| 1. Jawarish Bisbasa | | |
| 2. Jawarish Zarooni | | |
| Jawarish Zanjabeel | | |
| 4. Habb-e-Azaraqi | | |
| 5. Habb-e-Jadwar | | |
| 6. Habb-e-Mumsik | | |
| 7. Laboob-e-Kabir | | |
| 8. Laboob-e-Hazik | | |

Dosage of Fenugreek according to Unani, Ayurveda and Siddha systems of medicine

to the traditional systems of medicine.

 Table 04 shows the Dosage of Myristica fragrans according

Table 4: Dosage of Myristica fragrans according to Unani, Ayurveda and Siddha systems of medicine

| Unani ^[6] | Ayurveda ^[7, 8] | Siddha ^[9] |
|--|----------------------------|-----------------------|
| 4-5g (Nutmeg) 1.5-10.5g (Arillus/ Mace) | 0.5-1g (Powder) | 500mg – 1g (Powder) |

Scientific evidences on phytochemical studies of *Myristica fragrans*

phytochemical studies of Myristica fragrans.

Following table shows the recent evidences on

| Name of the Chemical constituents | Part Used | References | |
|--|--|----------------------------------|--|
| Aryltetralin and Diarylbutane lignans ^[10] 1-5 | Methanol extracts of Seeds | Kwon HS, et al. (2008) | |
| Dihydrobenzofuran Neolignans and Myticaganal A-C ^[11] (1-3) | Seed | Chumkaew P, et al (2019) | |
| Eight New Neolignans and Maceneolignans A-H ^[12] (1-8) | Methanol extract of the Arils | Morikawa T, et al. (2016) | |
| Four Lignans, Meso-Dihydroguaiaretic acid, Macelignan, Fragransin A2 & Nectandrin B ^[13] | Seeds | Thuong PT, et al. (2014) | |
| Lignan Compound ^[14] | N-Hexane Root Extract | Ginting B, et al. (2020) | |
| Macelignan ^[15] | Nutmeg Mace | Paul S, et al. (2013) | |
| Myristicin ^[16] | Aril (Mace) | Naikodi MA, et al. (2011) | |
| Myristicin ^[17] | Essential oil | Wang Y, et al. (2004) | |
| Myristicin, Myristic Acid, Trimyristin, Elemicin & Safrole ^[18] | Essential Oil | Setty JV, et al. (2020) | |
| Myristic Acid, Myristicin, Terpinen-4-ol, Alpha-Pinene & Safrole ^[19] | Essential oil | Qiu Q, et al. (2004) | |
| Myristicin, Methyleugenol, Safrole, Dehydrodiisoeugenol, Guaiacin & Myrisisolignan ^[20] | Seed | Yang XW, et al. (2008) | |
| Myristigranol (a new Diarylpropane derivative), one Diarylpropanoid & three Stilbenoids ^[21] | Methanol extract of Wood | Hiranrat A, et al. (2019) | |
| Myrifralignan ^[22] | Seed | Cao GY, et al. (2015) | |
| Phenolic compounds ^[23] | Fruit | Duan L, et al. (2009) | |
| Phenylpropanoid Compound ^[24] | Extract of Seed | Maeda A, et al. (2008) | |
| Trimyristin ^[25] | Chloroform extract of seeds | Narasimhan B, et al. (2006) | |
| 3',4',7-trihydroxyflavone ^[26] | Methanolic extract of crude Seed Kernel | Dzotam JK, et al. (2018) | |
| α-phellandrene, 3-Carene, p-cymene, Limonene, α-Thujene, α-pinene, Camphene, Sabinene, β-pinene, and Myrcene, α-terpinene, γ-Terpinene, Terpinolene & Myristicin ^[27] | Essential Oil | Ibrahim MA, <i>et al.</i> (2020) | |

Table 5: Phytochemical studies of Myristica fragrans

Scientific evidences on pharmacological activities of *Myristica fragrans*

Following table shows the recent evidences on

pharmacological activities of Myristica fragrans.

| Table 6: Pharmaco | logical activities | of Myristica fragrans |
|-------------------|--------------------|-----------------------|
|-------------------|--------------------|-----------------------|

| Pharmacological activity | Part Used | References |
|---|--|-------------------------------|
| Anti-allergic activity ^[28] | Extract of wood, nutmeg and mace | Champasuri S, et al. (2016) |
| | Extract of Seed | Narasimhan B, et al. (2006) |
| | Extract of Seed | Paul S, et al. (2013) |
| Antihastarial astivity [25, 15, 29, 26, 30, 31] | Essential Oil | Wang J, et al. (2019) |
| Antibacterial activity (20, 20, 20, 20, 20, 20, 20, 20, 20, 20, | Methanolic extract of crude seed kernel | Dzotam JK, et al. (2018) |
| | Ethyl acetate & ethanol extracts from Flesh, Seed & Mace | Shafiei Z, et al. (2012) |
| | Hydrolats and Essential Oil | Matulyte I, et al. (2020) |
| | Nutmeg Mace | Paul S, et al. (2013) |
| | Extract of the Leaves | Akinboro A, et al. (2012) |
| | Essential Oil | Piaru SP, et al. (2012) |
| Anti concer activity [15, 32, 33, 34, 35, 36, 14, 13] | Water extract of Seed | Kim EY, et al. (2016) |
| Anti-cancel activity | Methanolic extract | Chung JY, et al. (2006) |
| | Ethanolic Mace Extract | Suthisamphat N, et al. (2020) |
| | N-Hexane Root Extract | Ginting B, et al. (2020) |
| | Seeds | Thuong PT, et al. (2014) |
| Anticonvulsant activity [37] | Volatile oil of nutmeg, the dried seed kernel | Wahab A, et al. (2009) |
| Anti diabetic activity [15, 38] | Nutmeg Mace | Paul S, et al. (2013) |
| Anti-diabetic activity | Extract of Nutmeg | Pashapoor A, et al. (2020) |
| Anti-diarrhoel activity [39] | Serial extracts of the Seeds | Pillai NR, et al. (1991) |
| Anti depressant activity [40, 41] | Herbal extract | Moinuddin G, et al. (2012) |
| Anti-depressant activity to 3 | N-hexane extract of Seeds | Dhingra D, et al. (2006) |
| Anti-fungal activity ^[42] | Methanolic extract of Seeds | Cho JY, et al. (2007) |
| Anti-inflammatory activity ^[43, 15, 44, 45, 46] | Extract of Wood, Nutmeg and Mace | Champasuri S, et al. (2016) |
| | Nutmeg Mace | Paul S, et al. (2013) |
| | Seed | Cuong TD, et al. (2011) |
| | Seed | Lee JY, et al. (2011) |
| | Pericarp of Nutmeg | Zhang CR, et al. (2015) |
| Anti-microbial activity [18] | Essential Oil | Setty JV, et al. (2020) |
| Anti oxidant activity [47, 33, 24, 32, 46] | Seed | Li CW, et al. (2020) |
| Anti-oxidant activity [17, 55, 21, 52, 16] | Essential oil | Piaru SP, et al. (2012) |

| | Extract of Seed | Maeda A, et al. (2008) |
|---|--------------------------------------|---------------------------|
| | Extract of the Leaves | Akinboro A, et al. (2011) |
| | Pericarp of Nutmeg | Zhang CR, et al. (2015) |
| Analgesic activity ^[48] | Nutmeg seed kernels | Hayfaa AA, et al. (2013) |
| Approducing activity [49, 50] | Ethanolic extracts of Nutmeg | Tajuddin, et al. (2003) |
| Aphilouisiae activity | Ethanolic extracts of Nutmeg | Tajuddin, et al. (2005) |
| Gastro-protective activity ^[51] | Extract of the Seeds | Sattar A, et al. (2019) |
| | Nutmeg Mace | Paul S, et al. (2013) |
| Hepatoprotective activity ^[15, 52, 53] | Essential Oil | Morita T, et al. (2003) |
| | Kernal extract of Nutmeg | Dkhil MA, et al. (2019) |
| Immunomodulatory effect ^[54] | Aqueous Extract of Fresh Nutmeg Mace | Checker R, et al. (2008) |

Discussion

While reviewing the literature, it reveals Myristica fragrans shows therapeutic actions such as appetizer, carminative, digestive, stomachic and aphrodisiac activities and Myristica fragrans shows therapeutic uses such as indigestion, loss of appetite, diarrhoea, cough, asthma and sexual debility. Numerous studies have indicated that M. fragrans contains diverse phytochemicals such as Myristicin, Myristic Acid, Trimyristin, Elemicin, Safrole, Lignans, Neolignans, and Maceneolignans etc. which exhibit many of pharmacological activities such as Antiallergic activity. Antibacterial activity. Anti-cancer activity. Anticonvulsant activity, Anti-diabetic activity, Antidiarrhoel activity, Anti-depressant activity, Anti-fungal activity, Anti-inflammatory activity, Anti-microbial activity, Anti-oxidant activity, Analgesic activity, Aphrodisiac Gastro-protective activity, Hepatoprotective activity, activity, and Immunomodulatory effect. Myristicin, Myristic Acid, Trimyristin are the most active compounds among them.

Conclusion

Myristica fragrans is an aromatic evergreen tree which belongs to Myristicaceae family. Nutmeg (seed) and Mace (arillus) which are two separate spices derived from the fruit of tree *Myristica fragrans*. Nutmeg is the seed kernel inside the fruit and mace is the red lacy covering (aril) on the kernel. These are used as spices in culinary and in traditional systems of medicine. This review is to comprehensively summarized the phytochemical and pharmacological properties of *Myristica fragrans* that have reported to date.

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