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# Comprehensive review on Passiflora incarnata Linn

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#### Abstract

Passilora Incarnata commonly known as passiflower, belongs to family Passiflora. The fruit of Passiflora has several seeds which contain drinkable juice. Its medicinal use has been reported in the traditional system of medicine such as Ayurveda, Siddha and Unani. Passiflora incarnate contains phytochemicals such as alkaloids, flavonoids which are effective sedative that is not addictive. Phenols and flavonoids present in Passiflora have high antioxidant properties that show significant scavenger activity of free radicals. The genus Passiflora incarnate has long been used in traditional herbal medicine for treatment of insomnia and anxiety. It has been used as sedative tea and also been used to cure subjects affected by opiate dependence in India.

This review aims to provide up to date information about habitat, ecology, pharmacognostic evaluation, phytochemical studies, pharmacology, toxicology and herbal preparatioin which used to reduce anxiety particularly before surgery, also for variety of problems including pain, ADHD, stress, sleeplessness.

Keywords: Passiflora Incarnata, stress, ADHD, sedative, anxiety, scavenger

# Introduction

Passiflora comes from Latin word "passio" that was first time discovered by Spanish discoverers in 1529 and was described as symbol of "Christ" <sup>[1]</sup>. It also Known as maypop, purple passionflower, wild apricot, and wild passion Vine. The maypop, a species of passionflower belonging to the family Passiflora, features enormous, elaborate blooms with pronounced styles and stamens. The climbing vine known as the passion flower has white and purple Blossoms. The constituents of passion flower are releasing. South and Central America, as well as the Southeast United States, are the original homes of the passion flower. This is a fine climber suitable for covering arbours, verandahs and arches. It can be propagated by seed or layering. The fruit is edible when ripe <sup>[2]</sup>. Its medicinal usage has been reported in the traditional systems of medicine such as Ayurveda, Siddha and Unani. It has long been used to promote sleep <sup>[3]</sup>. From a phytochemical point of view, the plant contains several compounds including alkaloids, phenols, glycosyl flavonoids and cyanogenic compounds. Which are an effective sedative that is not addictive and does not cause drowsiness.

The plant contains flavonoids and alkaloids, with the leaves containing the greatest concentration of flavonoids. Other flavonoids present in P. incarnata include chrysin, apigenin, luteolin, quercetin, kaempferol, and isovitexin. Polyphenols belonging mainly to the C-glucoside class of flavones are present in P. incarnata and these phenols and flavonoids have high potential antioxidant properties that show a significant scavenger activity of free radicals. Passion flower is used by people to treat anxiety, particularly anxiety before surgery. For a variety of problems, including pain, ADHD, sleeplessness, and stress, some people also use passion flower. However, there isn't any reliable scientific data to back up these uses. Passion flower is used as a flavoring in several dishes and drinks<sup>[4]</sup>.

# Literature Review

M. Miroddi, G. Calapai et al. (2013). Passiflora incarnata, with 520 species having different pharmacological action like insomnia and anxiety in Europe and analgesic, anti-spasmotic in Brazil in different parts of world was found. This was due to the inadequate methodologies at different places [10].

Evans WC. Etal. (1996). The study prioritized to determine the phytochemical analysis of total phenolic, flavonoids, alkaloids, and tannins contents as well as the antioxidant

properties through 1,1-diphenyl-2-picrylhydrazyl (DPPH) quenching assay, superoxide and hydrogen peroxide radical scavenging assays of ethanol extract of *P. incarnata* leaves <sup>[11]</sup>.

Rubashiny veeramohanand noorma wati haron (2015). Taxonomic studies of four *Passiflora* species- *Passiflora* edulis (Passion fruit), *Passiflora coccinea*, *Passiflora* foetida, and *Passiflora incarnata* (Fragrant passion flower) focus to distinguish their macromorphological features like leaves, flowers, and fruits which revealed variations in color, texture, and shape by using a binocular microscope <sup>[12]</sup>.

Rehwald A., Meier B. *et al.* (1994). A qualitative and quantitative reversed- phase high-performance liquid chromatographic (HPLC) method has been developed for *Passiflora incarnata* flavonoid analysis like whichseprate out the major C-glycosylflavones like isovitexin, isoorientin, schaftoside <sup>[18]</sup>.

Lyca R. da Fonseca *et al.* (2020). This article explores *Passiflora* species with anxiolytic properties, dealing with their secondary metabolites, pharmacological studies, and registered products. Clinical trials are limited. Yet numerous other *Passiflora* species remain underexplored for their potential in medicinal and nutraceutical applications <sup>[32]</sup>.

Ruggy GH, Smith CS. (1940). The dried aerial parts of *Passiflora incarnate* L., was carried out in this article which give detail study of pharmacological effects span analgesic, antipyretic, anti-inflammatory, antimicrobial, cardiovascular, CNS depressant, and uterine stimulant<sup>[41]</sup>.

Dhawan K, Kumar S, Sharma A. (2003). This article focuses on threatment of dysmenorrhea, neuralgia, and nervous tachycardia, while also serving as a mild sedative for anxiety and sleeplessness. In addition to this it have certain clinical pharmacology which includes its impact on nausea, menopause, dysmenorrhea, diabetes <sup>[51]</sup>.

# Habitat

The purple passionflower thrives in average, medium, welldrained soil with full sun to partial shade. The flower typically grows in southeastern U.S., particularly in Florida and Texas, but can grow in central and eastern U.S. as far north as Delaware. The plant is also known to be found in Bermuda as well. It thrives in soil with clay, loam, or sand texture with high organic matter. It can be found roadsides, prairies, plains, meadows, pastures, savannas, woodlands, and riverbanks <sup>[7]</sup>.

# Taxonomical Classification <sup>[8]</sup>

Kingdom: Plantae Division: Magnoliophyta Class: Magnoliopsida Family: Passifloraceae Genus: Passiflora L. Species: P. incarnata L.

# Common Names [9]

English: Passion flower, Apricot, Maypop. Hindi: Krishna kamal. Marathi: Krishna kamad

# Pharmacognstic Evaluation Morphology <sup>[10]</sup>

**Stems:** The stem is herbaceous or woody, generally climbing, very rarely tree-like Vine, glabrous to slightly pubescent, tendrils present.

**Leaves:** Leaves alternate, sometimes simple, entire, rarely tendril axillary, arising from sterile pedicels. 3-lobed, serrate, petiolate. Petioles contains two glands near the base of the leaf blade.

**Flowers:** Flowers can be bisexual or unisexual, regular. The corolla consisting of a corolla and stamens. The diameter of the flower is usually 6-7 cm. The flower has 5 petals and 5 sepals, which are purplish to whitish, similar and alternate. The flower has 3 types, typically 3 stamens, 5 green-white sepals with terminal appendages.

Fruit: fleshy, initially green, yellow-red when ripe.

 Table 1: (Morphology of Passiflora Incarnata.)
 [11]

Appearance	Coarse powder		
Colour	Green		
Odour	No Characteristic odour		
Taste	Slightly bitter		
Percent loss on drying	7.12		
Crude fibers	9.4		



Fig 1: (Passiflora Flower)





Fig 2: (Passion Flower with fruit)



Fig 3: (Floral parts of the Purple Passionflower (*Passiflora Incarnata*)

#### Stigma

The stigma is located at the center of the flower and serves as the landing site for pollen.

#### Anther

Anthers are the male reproductive parts of the flower. Anthers contain pollen, which contains the male genetic material necessary for fertilization.

# Ovary

The ovary is the base of the female reproductive system within the flower. It contains ovules, which are the future seeds. After successful fertilization, the ovary develops into the fruit of the plant.

# Petal

Petals are the colorful, surrounded the reproductive organs. The petals are typically purple, lavender, or pink, and they serve to attract pollinators with their vibrant colors and unique patterns.

# Sepal

Sepals are the protective, leaf-like structures that encase the flower bud before it opens. They are typically green and serve to shield the developing flower.

# Microscopy [12]



Fig 4: (Abaxial surface of P. incarnata leaf)



Fig 5: (Trichomes of P. incarnata)



Fig 6: (Amphipericytic stoma of P. incarnate)



Fig 7: (Adaxial surface of P. incarnate leaf)

*Passifloraincarnata* L. leaves are alternate and palmately tri-lobed. Leaf apex is cuspidate while the leaf base is cordate with serrate margin. The adaxial surface of the leaves is dark green whereas the abaxial surface is dull green. The scanning electron microscopy view of a *P*. *incarnata* leaf shows papillose epidermal surface with

amphipericytic stomata distributed on the abaxial surface (Fig. 1). Trichomes are simple, unicellular and sparsely distributed throughout the abaxial and adaxial surfaces of the leaf (Figs. 2 and 4). Both stomata and trichomes distribution are more dense on the abaxial surface compared to the adaxial surface.



Fig 8: (T.S. of Lamila)

**Fig 9:** (T.S. of Stem)



Fig 10: (T.S. of Petiole)

# **Powder microscopy**

**Stomata:** A baxial epidermal peelings showed the presence of epidermal cells and stomata. The epidermal cells are amoeboid in outline with wavy anticlinal walls. Stomata were densely distributed and they are anamocytic type. The guard cells are elliptical and thin walled.

Fig 11: (T.S. Tendril)

**Parenchyma cells:** Parenchyma cells are abundant and scattered in the powder. They vary in shape and size. They may be narrow and rectangular or wide and elongated. Some of the parenchyma cells are squarish and thin walled. The parenchyma cells are up to 140 mm long and 20 mm thick.



# Fig 12: (T.S. of Podwer)

# **Botanical Voucher Specimen**



# Analytical Parameter<sup>[9]</sup>

Table 2: (Analytical Parameter)

Parameters	Value in w/w
Total ash	9.23
Acid- insoluble ash	10.3
Water- insoluble ash	6.50
Sulfated ash	18.20

# **Phytochemical Studies**

Table 3: (Phytochemical Parameter)

Sr. no.	Part of plant	phytoconstituents	Examples
1.	Leaves	Flavonoid	C- glycosylflavones, Isovitexin, vitexin, orientin, isoorientin, saponarin.
2.	Leaves	Alkaloid	Harman, harmol, harmine.
3.	Leaves and roots	carbohydrates	raffinose, sucrose, D- glucose and Dfructose

In phytochemical studies, orientin and vitexin are found in small amounts in the leaves of *P. incarnata*, *P. alata*, and *P. edulis* <sup>[10, 11]</sup>. Furthermore, iso-orientin is one of the major flavonoids in *P. incarnata* and *P. edulis* var. flavicarpa, but it is only detected in small concentrations in *P. alata*. <sup>[10, 12]</sup> A comparative metabolite profiling and fingerprinting of the genus *Passiflora* leaves were reported using a multiplex

approach of UPLC-MS and NMR analyzed by chemometric tools <sup>[14]</sup>. Thin layer chromatography can also be used in order to differentiate species <sup>[24]</sup>.

The aqueous extract is phytochemically characterized by a set of C-glycoside flavonoids such as vitexin (1), isovitexin (2), schaftoside (3), isoschaftoside (4), orientin (5), isoorientin (6), and swertisin (7). <sup>[26, 27]</sup> Moreover, the free International Journal of Pharmacognosy and Pharmaceutical Sciences

flavonoids apigenin (8), luteolin (9), quercetin (10), kaempferol (11), and chrysin (12) are also found <sup>[15]</sup>.

# Flavonoids

Flavonoids are reported to be the major phyto-constituents of *P. incarnata*. C- glycosylated flavonoids found in *P. incarnata* consist mainly of a glucose group directly linked to the aromatic nucleus only in positions 6 and 8 of the flavonoid core <sup>[18]</sup>. The aqueous extract is phytochemically characterized by a set of C-glycoside flavonoids <sup>[8]</sup>.



 Table 4: (Flavonoid present in Passiflora Incarnata)
 [32]

Flavonoids	R1	R2	R3	R4	R5	R6
(1) Vitexin	OH	Н	Н	Н	OH	Glucose
(2) Isovitexin	OH	Н	Н	Glucose	OH	Н
(3) Schaftoside	OH	Н	Н	Glucose	OH	Arabinose
(4) Orientin	OH	Н	Н	Arabinose	OH	Glucose
(5) Isoorientin	OH	OH	OH	Н	OH	Glucose
(6) Isoorientin	OH	OH	OH	Glucose	OH	Н
(7) Swertisin	OH	Н	Н	Glucose	OCH3	Н
(8) Apigenin	OH	Н	Н	Н	OH	Н
(9) Luteolin	OH	OH	OH	Н	OH	Н
(10) Quercetin	OH	OH	OH	Н	OH	Н
(11) Kaempferol	Н	OH	OH	Н	OH	Н
(12) Chrysin	Н	Н	H	Н	OH	Н

# Alkaloids

The alkaloids present in *Passiflora* are of the indole type ( $\beta$ -carbolines), which are the second major known group of alkaloids (54). Some of them have value in medicine as tranquilizers and for the treatment of hypertension (55). *P. incarnata* is the most widely studied species containing

alkaloids (24). In studies carried out in the 1960s, harmine (13), harmol (14), harmaline (15), harmalol (16), and Harman (17) were detected (Figure 2). Although the presence of traces of these alkaloids was confirmed. they were undetectable in most commercial materials <sup>[6]</sup>.

Table 5: (Alkaloid present in P. incarnata)<sup>[6]</sup>

Sr. No.	Alkaloid	Structure
1.	Harman	L Z Z Z Z
2.	Harmaline	H <sub>3</sub> C
3.	Harmalol	HO NH CH3



**Miscellaneous Phyto-constituents:** Various other constituents which have been reported from *P. incarnata* include  $\gamma$ -benzo-pyrone derivative maltol (27), carbohydrates such as raffinose, sucrose, D-glucose and Dfructose (28); essential oil containing hexanol (1.4%),

benzyl alcohol (4.1%), linalool (3.2%), 2-phenylethyl alcohol (1.2%), 2-hydroxy benzoic acid methyl ester (1.3%), carvone (8.1%), trans- anethol (2.6%), eugenol (1.8%), isoeugenol (1.6%),  $\beta$ ionone (2.6%),  $\alpha$ -bergamotol (1.7%) and phytol (1.9%) <sup>[6]</sup>.

# **Pharmacological Activities**

<b>Fable 6:</b> (Major pharmaco)	ological a	activity of	Passiflora	species)
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Part of plant System		Effects		Ref.
	Chemical activities	Anxiolytic, spasmolytic, hypnotic, sedative, narcotic and anodyne		33
Flower	Biological and Pharmaceutical	Antimicrobial Allergic reactions, asthma, irritated sinuses, skin rashes, and skin	Products	34
		Blood vessel inflammation (Vasculitis). Congestive heart failure restlessness, anxiety, and agitation	Extract	
	Traditional uses	Smoked	Dried	35
	Biological and	Antibacterial Antioxidant	Ethanol	36
Laguag	Pharmaceutical	Anti-inflammatory	extracts	37
Leaves		Hemolytic	Extracts	
			Aqueous	
			extract	
Fruit	Biological and Pharmaceutical	Antibacterial Anti-tumor	Ethanol extracts	38
Stem	Biological and Pharmaceutical	Antioxidant	Extracts	39

#### Anti-Deprasant

(Ruggy G. H et al., 1940, Medina, et al., 1990 Speroni, E et al., 1996) initiated investigations by 25 mg/kg intraperitoneal injection of the water extract resulted in decreased organization and locomotion, while doses of 60-250 mg/kg of 30% or 40% ethanol extracts caused a cessation of movement. The 40% ethanol extract at 60 mg/kg increased sleep duration, and at 50 mg/kg, it delayed the onset of seizures. In vivo studies involving rats receiving a daily dose of 5 g/kg of 30% ethanol extract for three weeks showed no significant changes in body weight, temperature, or motor organization, yet a decrease in motor activity was observed. Furthermore, chrysin, identified in Herb Passiflora, demonstrated a high affinity for benzodiazepine receptors, affecting animal locomotion and enhancing pentobarbital-induced hypnosis at 30 mg/kg<sup>[36, 37,</sup> 38, 39]

# Anti-inflammatory activity

(Borrelli, F et al., et al. 1994, Guerin, J. C et al., 1985) gives tale of Herb Passiflora unfolded with animals taking center

stage. Intriguingly, Animals receiving an intragastric injection of 75 to 500 mg/kg bw of Herb Passiflora ethanol extract experience a reduction in inflammation one hour later. Herb Passiflora has anti- inflammatory and antioxidant properties that can be used to cure and prevent numerous diseases, including intricate inflammatory processes. After a dose of 500 mg/kg bow of Herb Passiflora ethanol extract, it was discovered that total leukocyte passage to the animal pleural cavity had decreased by 40%. This outcome was brought about by a decline in the movement of polymorphonuclear and mononuclear These outcomes were leukocytes. equivalent to acetylsalicylic acid at 250 mg/kg [40, 41].

# Anti-bacterial

Borrelli, F *et al.*, in the year (1996) Research painted a vivid picture of resilience the growth of the following fungi was not inhibited by the 50% ethanol extract of Herb *Passiflora* over 500 mg/ml: Botrytis cinerea, *Aspergillus fumigatus*, *Rhizopus nigricans*, *Fusarium oxysporum*, *Penicillium digitatum*, and Candida albicans. Helicobacter pylori's

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growth was inhibited by a methanol extract of Herb *Passiflora* with a minimum inhibitory concentration of 50 g/ml. Passion fruit seeds contain volatile substances that are poisonous to 1- octanol and Meloidogyne incognita. The nematicidal impact of these substances is active against Meloidogyne incognita <sup>[40]</sup>.

# Anti-nauseating effect

(Fisher, A. A *et al.* 2000, Freson G, *et al.* 2008) unveiled a cautionary chapter in the story of *Passiflora incarnata*. In a clinical trial, a 34-year-old woman experiencing severe nausea after receiving *Passiflora incarnata* as directed. Thus, this patient experienced negative effects from *Passiflora* 14 ncarnate. Due to the carboline alkaloids (Harmaline, harmane, and tetrahydroharmine), which excite the central nervous system by slowing down the metabolism of amine neurotransmitters, *Passiflora incarnate* causes optical and auditory illusions, locomotor ataxia, nausea, vomiting, misperception, and anxiety <sup>[42]</sup>. According to a clinical investigation, passionflower is effective in strengthening resilience in individuals with nervous restlessness <sup>[43, 44, 45]</sup>.

# Anti-anxiety activity

Dhawan K, Kumar S, Sharma Ain the year (2001) take plant *Passiflora incarnate Linn*. Which Has been used to cure anxiety and insomnia since time immemorial. A fraction derived from the methanol extract of *P. incarnate* has been observed to exhibit significant anxiolytic activity at a dose of 10 mg/kg in mice using elevated plus-maze model of anxiety. This fraction comprises mainly two components which are visible as blue and turquoise colored fluorescent

spots at 366 nm of the UV light. The possibility of a phytoconstituent having benzoflavone nucleus as the basic moiety being responsible for the bioactivity of P. incarnate is highly anticipated <sup>[46]</sup>.

# Antitussive activity

Dhawan K, Kumar S, Sharma A, in the year (2003-2017). Studied on mice and found that when mice were exposed to sulphur dioxide, the leaves of P. 15ncarnate showed considerable antitussive action, with cough induction comparable to that of codeine phosphate <sup>[47, 48]</sup>.

# Hypertension

Benson VL *et al* in the year (2008). Delved into the enduring challenge of Cardiovascular disease continues to be the leading cause of morbidity and mortality worldwide despite advances in pharmacotherapies and mechanical treatments and there is a good possibility that this burden will rise. An associated species of *P. nepalensis* called P. 15 ncarnate has been known to have antihypertensive properties.

*P. incarnate* has an antihypertensive effect because it contains flavonoids and water- soluble component that has been identified as a mercury salt (C10H22O8 NHgCl2). An associated species of *P. nepalensis* called P. 15ncarnate has been known to have antihypertensive properties <sup>[53]</sup>.

# Herbal Preparation <sup>[62]</sup>

- Hard capsule (300 mg/capsule
- Tablet (400 mg/tablet)
- Oral liquid
- Herbal tea (2g per tea sachet)

No	<b>Preparation</b>	Dosage form	Posology	Indication
1.	Passion flower herb	Hard capsules (300mg/capsule)	Adults and the elderly: For relief of symptoms of mild anxiety 1- 2 capsules 3 times a day. To aid sleep, take 2 capsules 30 minutes before bedtime with an earlier dose of 2 capsules earlier in the evening if necessary. Maximum daily dose: 8 single doses	Temporary relief of mild anxiety and to aid sleep.
2.	Passiflora 16 nc arnate L.,herba	Tablet (400 mg/tablet)	Adults and adolescents over 12 years of age: 400 mg - 2 g (1-4 tablets per day)	Relief of mild symptoms of mental stress and to aid sleep. Also relief of mild nervous tension.
3.	Passifloraeherb a, cut	Herbal tea (2g per tea sachet)	Adults and adolescents over 12 years: 1 cup of tea from 1 tea sachet 2-4 times daily	To improve general condition in mental stress and to aid sleep.
4.	Powder	Hard capsule (1 capsule contains 300 mg of powdered drug)	Adults: 2 hard capsules 2 times daily Adolescents over 12 years of age: 1 hard capsule 2 times daily	Symptomatic treatment of neurotonic conditions of adults and children, notably in cases of mild disorders of sleep.

#### Table 7: (Herbal Preparation)

# Conclusion

*P. incarnata*, a plant widely employed in the Indian system of medicine. According to these investigations, this natural remedy is a cutting-edge option for drug development and bioprospecting for the treatment of conditions like anxiety, sleeplessness, convulsions, sexual dysfunction, cough, cancer, and postmenopausal syndrome. These plants' phytochemicals and minerals will make it possible to utilize them for therapeutic purposes. Flavonoid make up the majority of Herb *Passiflorae*'s components. For treatment of dysmenorrhea, neuralgia, and nervous tachycardia, Herb *Passiflora* is used. Herb *Passiflorae*'s fruit is used to make jams, jellies, and sweets. The juice is a preferred beverage flavoring. Some *Passiflora* species have not yet been thoroughly investigated for their medicinal potential. In the future, they can be examined for their medicinal potential.

# Reference

- Dhawan K, Dhawan S, Sharma A. *Passiflora*: A review update. J Ethnopharmacol. 2004;94:1-23. DOI: 10.1016/j.jep.2004.02.023.
- Fisher AA, Purcell P, Le Coureur DG. Toxicity of *Passiflora incarnata* L. Journal of Toxicology: Clinical Toxicology. 2000 Jan 1;38(1):63-6.
- 3. Patel SS. Morphology and pharmacology of *Passiflora edulis*: A review. Journal of Herbal Medicine and Toxicology. 2009;3(1):1-6.
- 4. Kaikade AR, Gurunani SG, Pandel TW, Sherekar SA, Kaikade PR, Mehare SR, *et al.* Phyto-Pharmacognostic review on *Passiflora* species. Journal of Medicinal Plants Studies. 2023;11(3):35-50.

- 5. Washington B. Purple Passionflower Pops in the Summer. Virginia Native Plant Society - Northern Neck Chapter. Vnps.org; c2020 Jul 30.
- 6. Patel SS, Verma NK, Gauthaman K. *Passiflora Incarnata* Linn: A Review on Morphology, Phytochemistry and Pharmacological Aspects. Phcog Rev. 2009;3(5):175-181.
- Kaikade AR, Gurunani SG, Pandel TW, Sherekar SA, Kaikade PR, Mehare SR, *et al.* Phyto-Pharmacognostic review on *Passiflora* species. Journal of Medicinal Plants Studies. 2023;11(3):35-50.
- Miroddi M, Calapai G, Navarra M, Minciullo PL, Gangemi S. *Passiflora incarnata* L.: Ethnopharmacology, clinical application, safety and evaluation of clinical trials. Journal of Ethnopharmacology. 2013;150:791-804.
- 9. Evans WC. Trease and Evans Pharmacognosy. W.B. Saunders Publication, Edinburgh, USA; c1996.
- Rehwald A, Meier B, Sticher O. Qualitative and quantitative reversed-phase high-performance liquid chromatography of flavonoids in *Passiflora incarnata* L. Pharmaceutica Acta Helvetiae. 1994;69:153-158. DOI: 10.1016/0031-6865(94)90017-5.
- Ulubelen A, Oksuz S, Mabry TJ, Dellamonica G, Chopin J. C-glycosylflavonoids from *Passiflora pittieri*, *P. alata*, *P. ambigua* and *Adenia mannii*. Journal of Natural Products. 1982;45:783. DOI: 10.1021/np50024a030.
- 12. Zucolotto SM, Fagundes C, Reginatto FH, *et al.* Analysis of C-glycosyl flavonoids from South American *Passiflora* species by HPLC-DAD and HPLC-MS. Phytochemical Analysis. 2012;23:232-239. DOI: 10.1002/pca.1348.
- Farag MA, Otify A, Porzel A, Michel CG, Elsayed A, Wessjohann L. Comparative metabolite profiling and fingerprinting of genus *Passiflora* leaves using a multiplex approach of UPLC-MS and NMR analysed by chemometric tools. Analytical and Bioanalytical Chemistry. 2016;408:3125-3143. DOI: 10.1007/s00216-016-9376-4.
- Gomes SVF, Portugal LA, Anjos JP, *et al.* Accelerated solvent extraction of phenolic compounds exploiting a Box-Behnken design and quantification of five flavonoids by HPLC-DAD in *Passiflora* species. Microchemical Journal. 2017;132:28-35. DOI: 10.1016/j.microc.2016.12.021.
- 15. Veeramohan R, Haron N. Macromorphological and micromorphological studies of four selected *Passiflora* species in peninsular Malaysia. Pak J Bot. 2015;47(2):485-492.
- Borrelli F, Pinto L, Izzo AA, Mascolo N, Capasso F, *et al*. Anti-inflammatory activity of *Passiflora incarnata* L. in rats. Phytotherapy Research; c1996.
- 17. Dhawan K, Dhawan S, Sharma A. *Passiflora*: a review update. Journal of Ethnopharmacology. 2004;94:1-23. DOI: 10.1016/j.jep.2004.02.023.
- Oliveira MS, Pinheiro IO, Silva FSBV. Vermicompost and arbuscularmycorrhizal fungi: an alternative to increase foliar orientin and vitexin-2-O-rhamnoside synthesis in *Passiflora alata* Curtis seedlings. Industrial Crops and Products. 2015;77:754-757. DOI: 10.1016/j.indcrop.2015.09.061.

- Wosch L, Santos KC, Imig DC. Comparative study of *Passiflora* taxa leaves: II. A chromatographic profile. Brazilian Journal Pharmacognosy. 2017;27:40-49. DOI: 10.1016/j.bjp.2016.06.007.
- Petry RD, De Souza KCB, Bassani VL, Petrovick PR, Ortega GG. Doseamento do teor de flavonoides totais em extratos hidroalcóolicos de *Passiflora alata* Dryander (Maracujá). Revista Brasileira de Farmácia. 1998;79:7-10.
- 21. Reginatto F, De-Paris HF, Petry RD, *et al.* Evaluation of *anxiolytic* activity of spray dried powders of two South Brazilian *Passiflora* species. Phytother Res. 2006;20:348-351. DOI: 10.1002/ptr.1853.
- Quercia V, Turchetto L, Pierini N, Cuozzo V, Percaccio G. Identification and determination of *vitexin* and *isovitexin* in *Passiflora incarnata* extracts. J Chromatogr A. 1978;161:396-402. DOI: 10.1016/s0021-9673(01)85261-4.
- 23. Jay M. *C*-glycosylflavonoids. In: Harbone JB, ed. The Flavonoids. London, UK: Chapman & Hall; c1996.
- Wohlmuth H, Penman KG, Pearson T, Lehmann RP. Pharmacognosy and chemotypes of passionflower (*Passiflora incarnata* L.). Biol Pharm Bull. 2010;33:1015-1018. DOI: 10.1248/bpb.33.1015.
- Zeraik ML, Pereira CAM, Zuin VG, Yariwake JH. Maracujá Um alimentofuncional? Rev Bras Farmacogn. 2010;20:459-471. DOI: 10.1590/s0102-695x2010000300026.
- 26. Da Fonseca LR, Rodrigues RA, Ramos AS, Da Cruz JD, Ferreira JLP, Silva JR de A, Amaral ACF. Herbal Medicinal Products from World Journal Article ID 6598434; c2020. p. 18.
- Pinheiro E, Silva I, Gonzaga LV, Amante E, Reinaldo F, Ferreira M, *et al.* Optimization of extraction of high ester pectin from passion fruit peel (*Passiflora edulis* flavicarpa) with citric acid by using response surface methodology. Bioresour Technol. 2008;99-113:5561-5566.
- 28. Puricelli L, Aicab I, Sartorb L, Garbisa S, Caniato R. Preliminary evaluation of inhibition of matrix metalloprotease MMP-2 and MMP-9 by *Passiflora edulis* and *P. foetida* aqueous extracts. Fitoterapia. 2003;74:302-304.
- 29. Akhondzadeh S, Naghavi HR, Vazirian M, Shayeganpour A, Rashidi H, Khani M. Passionflower in the treatment of generalized anxiety: a pilot doubleblind randomized controlled trial with oxazepam. J Clin Pharm Ther. 2001;26(5):363-367.
- 30. Ichimura T, Yamanaka A, Ichiba T, Toyokawa T, Kamada Y, Tamamura T, *et al.* Antihypersensitive effect of an extract of *Passiflora edulis* rind in spontaneously hypersensitive rat. Biosci Biotechnol Biochem. 2006;70(3):718-721.
- 31. Ikegami F, Fujii Y, Ishihara K, Satoh T. Toxicological aspects of Kampo medicines in Clinical Use. Chem Biol Interact. 2003;145:235-250.
- 32. Barbosa PR, Valvassori SS, Bordignon CL, Kappel VD, Martins MR, Gavioli EC, *et al.* The Aqueous Extracts of *Passiflora edulis* and *P. foetida* Reduce Anxiety Related Behaviors Without Affecting Memory Process in Rats. J Med Food. 2008;11:282-288.
- 33. Medina JH, Paladini AC, Wolfman C, De Stein ML, Calvo D, Diaz LE, *et al.* Chrysin (5,7-di-OH-flavone), a naturally occurring ligand for benzodiazepine receptors,

has anticonvulsant properties. Biochem Pharmacol. 1990;40(10):2227-2231. https://DOI.org/10.1016/0006-2952(90)90716-X.of Chamomile and *Passiflora* Extracts in Sleep-Disturbed Rats, Biol. Pharm. Bull. 2005, 28, 808-810.

- Ruggy GH, Smith CS. A pharmacological study of the active principle of *Passiflora incarnata*. J Am Pharm Assoc. 1940;29(6):245-249. DOI: 10.1002/jps.3030290602.
- 36. Speroni E, Billi R, Mercati V, Boncompagni E, Toja E. Sedative effects of crude extract of *Passiflora incarnata* after oral administration. Phytother; c1996.
- Speroni E, Billi R, Perellino NC, Minghetti A. Role of chrysin in the sedative effects of *Passiflora incarnata* L. Phytother Res; c1996.
- Borrelli F, Pinto L, Izzo AA, Mascolo N, Capasso F, Mercati V, *et al.* Anti-inflammatory activity of *Passiflora incarnata* L. in rats. Phytother Res. 1996.
- 39. Caruso S, Cianci S, Cariola M, Fava V, Rapisarda AMC, Cianci A. Effects of nutraceuticals on quality of life and sexual function of perimenopausal women. J Endocrinol Invest. 2017;40:27-32. DOI: 10.1007/s40618-016-0500-2.
- Fisher AA, Purcell P, Le Coureur DG. Toxicity of *Passiflora incarnata* L. J Toxicol Clin Toxicol. 2000;38(1):63-6.
- Fisher AA, Purcell P, Le Couteur DG. Toxicity of *Passiflora incarnata* L. J Toxicol Clin Toxicol. 2000;38:63-66. DOI: 10.1081/CLT100100919. Available from: https://biointerfaceresearch.com/12899.
- 42. Frison G, Favretto D, Zancanaro F, Fazzin G, Ferrara SD. A case of β-carboline alkaloid intoxication following ingestion of *Peganum harmala* seed extract. Forensic Sci. Int. 2008;179(2-3):e37-43. DOI: 10.1016/j.forsciint.
- 43. Dhawan K, Kumar S, Sharma A. Anti-anxiety studies on extracts of *Passiflora incarnata* Linneaus. J Ethnopharmacol. 2001;78(2-3):165-70. DOI: 10.1016/s0378-8741(01)00339-7. PMID: 11694362.
- 44. Dhawan K, Kumar S, Sharma A. Aphrodisiac activity of methanol extract of leaves of *Passiflora incarnata* Linn. In mice. Phytother Res. 2003;17(4):401-3.
- 45. Dhawan K, Kumar S, Sharma A. Antiasthmatic activity of the methanol extract of leaves of *Passiflora incarnata*. Phytother Res. 2003;17(7):821-22.
- 46. Hameed IH. A new polymorphic positions discovered in mitochondrial DNA hypervariable region HVIII from central and north-central of Iraq. Mitochondrial DNA. 2016;27(5):3250-4.
- 47. Assessment report on *Passiflora incarnata* L., herba. EMA/HMPC/669738; c2013.