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Fragaria & Frangula Combination (FIBROMELT)

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ABSTRACT

The article will present the indications, analyze of the component actions, cautions, and contraindications of the herbal formula Fragaria & Frangula Combination / Fibromelt /. The indications and analyze of the component actions will be done from the points of view of the Western and Traditional Chinese Medicines. The medical information of the Fragaria & Frangula Combination / Fibromelt / effectivity is done on the base of the 35 years practice of the author.

Keywords

Fibromyalgia relief, Natural pain relief, Herbal remedy for pain, Fragaria supplement, Frangula benefits.

Introduction

Fragaria & Frangula Combination / Fibromelt / is the herbal formula created on the base of the Traditional Bulgarian Medicine. Bulgaria has the thousand-years old history of the selection, cultivation, and export of the medical herbs, same as the China.

Indications and Analyze of the Component Actions Indications

Western medicine - Fibromyoma of the uterus with the chronic infection, menstrual over bleeding, and bleeding between the menstruation.

Traditional Chinese medicine - Heat and blood stasis in the uterus; phlegm congelation in the uterus.

Effects: Antibacterial, anti-inflammation, hemostatic, formation dissolving, anti-anemia, alleviating premenstrual and menopausal symptoms, regulate menstruation cycle, antioxidant, antianxiety, and antidepression effects.

Analyze of the Component Actions

In Fragaria & Frangula Combination / Fibrolyse / are included the next herbs:

Folium Fragaria vesca / Strawberry alpine / Herba Hypericum perforatum / St John's wort (stalks) /

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Folium Calendula arvensis / Calendula (flower) / Cortex Frangula alnus / Glossy buckthorn (bark) / Herba Polygonum hydropiper / Bachelor`s buttons (stalks) / Flower Matricaria chamomilla / Chamomile (flower) / Folium Urtica dioica / Common nettle (leaves) / Herba Achillea millefolium / Yarrow stalks/

Folium Fragaria Vesca / Strawberry alpine / [1,2]

Compounds: Caffeic acid derivatives, flavonoids: rutin, quercetin, tannins: ellagic acid tannins, oligomeric proanthocyanins, organic acids, and Vit C.

Effects: astringent and diuretic

Indications: Metrorrhagia, anemia, gastritis, enterocolitis, diarrhea, jaundice, catarrh of the respiratory and urinary tracts, kidney stones, gout, rheumatoid arthritis, nervous tension, hysteria, insomnia, hypertonia, and eczemas.

Cautions and adverse effects: No adverse effects with the administration of therapeutic doses. Not to be used during pregnancy.

Daily dosage: 1 to 2 g per day

Herba Hypericum Perforatum / St John`s wort (stalks)/ [1,2]

Compounds: Anthracene derivatives: naphthodianthrones, hypericin, pseudo hypericin; flavonoids: hyperoxide, quercitrin, rutin, isoquercitrin, amentoflavone; xanthones; acylphloroglucinols; volatile oil; oligomers; procyanidines and other catechin tannins; imanin; carotin, choline, caffeic acid derivatives, and Vit C.

Effects: Anti-anxiety, antidepressant, alleviating premenstrual and menopausal symptoms, hemostatic, anti-inflammatory, antibacterial, antifungal, antineoplastic, antioxidant, neuroendocrine, weight loss, and wound-healing effects.

Indications: Anxiety, depression, white vaginal discharge, wounds and burns, bronchitis, asthma, enuresis, gout, rheumatism, and dermatosis

Cautions and adverse effects: Contraindicated in the pregnancy as well as in a history of the photosensitivity or hypersensitivity to Hypericum perforatum. Decreasing activity of the irinotecan, protease inhibitors, amiodarone, anticoagulants, cyclophosphamide, cyclosporine, darunavir, etoposide, phenytoin, tamoxifen, benzodiazepines, beta-adrenergic blockers, calcium channel blockers, digoxin, statin medications, estrogen medications, methadone, nortriptyline, omeprazole, theophylline, tramadol, verapamil, and iron. Increasing activity of the rasagiline, selegiline, anesthetics, erlotinib, imatinib, monoamine oxidase inhibitors, SSRI, antidiabetic agents, buspirone, nefazodone, opioid analgesics, and venlafaxine.

Daily dosage: 2 to 4 g taken 3 times daily for the dried herb.

Folium Calendula Arvensis / Calendula (flower) / [1,2]

Compounds: Triterpene saponins, triterpene alcohols, flavonoids, hydroxycoumarins, carotenoids, volatile oil, water-soluble polysaccharides, and plyynes.

Effects: Anti-inflammatory, antimicrobial (Staphylococcus aureus, Klebsiella pneumoniae, Sarcina lutea, Candida monosa), antiviral (HIV, vesicular stomatitis virus, Rhinovirus), spasmolytic, choleretic, antilipid, antihypotensive, dermatitis, and wound-healing effects.

Indications: Inflammatory conditions of the internal organs, dysmenorrhea, irregular menstruation, stomach and intestinal ulcer, constipation, cholecystitis, atherosclerosis, wounds, and inflammatory skin diseases.

Cautions and adverse effects: No adverse effects with the administration of therapeutic doses.

Daily dosage: 1 to 2 g

Cortex Frangula alnus / Glossy buckthorn (bark) / [1,2]

Compounds: Anthracene derivatives, naphthalene derivatives, and peptide alkaloids.

Effects: Anti-absorptive effects

Indications: Constipation, hemorrhoids

Cautions and adverse effects: Not to be used during pregnancy and nursing. Not to be used in the intestinal obstruction and acute intestinal inflammations. Can cause hypokalemia. *Daily dosage:* 20 to 180 mg

Polygonum hydropiper / Bachelor`s buttons stalks / [1,2]

Compounds: Flavonoid glucosides, polygopeparin, tannins, phytostirrings, organic acids, Vit C, Vit K, etheric oil, and others *Effects:* Hemostatic, analgetic, and antibacterial effects

Indications: Bleeding from the gastrointestinal tract, metrorrhagia, white discharge from uterus

Cautions and adverse effects: No adverse effects with the administration of therapeutic doses.

Daily dosage: 3 to 6 g daily

Flower Matricaria Chamomilla / Chamomile (flower) / [1,2]

Compounds: Volatile oil, sesquiterpene lactones, flavonoids, caffeic and ferulic acid ester, polyynes, coumarin, apigenin, matricin, herniarin, choline, Vit A, and Vit C.

Effects: Antibacterial (gram-positive bacteria, dermatomycoses), hemostatic, analgetic, spasmolytic, cholagogue, and anti-anxiety effects.

Indications: Menstrual complaints – dysmenorrhea, nervousness, metrorrhagia, white vaginal discharge, hematuria, bloating, fullness, spasmodic gastrointestinal disturbances, hysteria, inflammation of the mouth, throat, rhinitis, toothache, earache, headache, and influenza.

Cautions and adverse effects: No adverse effects with the administration of therapeutic doses. Not to be used during pregnancy.

Daily dosage: 3 g

Folium Urtica dioica / Common nettle (leaves) / [1,2]

Compounds: Histamine, serotonin, acetylcholine, formic acid, leukotrienes, flavonoids, silicic acid, volatile oil, and potassiumions.

Effects: Hemostatic, hemopoietic, anti-inflammatory, analgetic, and anti-arthritic effects

Indications: Metrorrhagia, acne, infections of the urinary tract, kidney, and bladder stones, pulmonal and gastric hemorrhages, hemorrhoid, decreased lactation, and rheumatism.

Cautions and adverse effects: No adverse effects with the administration of therapeutic doses. Not to be used during pregnancy. Gastric irritation. Must be separated 1-2 hours from the iron medications.

Daily dosage: 8 to 12 g

Achillea millefolium / Yarrow stalks/ [1]

Compounds: Volatile oil, sesquiterpene lactones, polyynes, alkamids, flavonoids, betaine

Effects: Cholagogue, spasmolytic, cardiovascular (lowered blood pressure, decreased LDL and triglycerides, and increased HDL), anti-edema, anti-inflammatory, and antimicrobial effects (Escherichia coli, Salmonella typhosa, Shigella sonnei, Staphylococcus aureus, Candida albicans)

Indications: Loss of appetite, spastic gastrointestinal discomfort, liver, and gall bladder disorders (cirrhosis, dyslipidemia, bile stasis), hypertonia, bleeding hemorrhoids, and menstrual problems. *Cautions and adverse effects:* Not to be used during pregnancy and breastfeeding

Daily dosage: 4.5 g

Cautions and Contraindications

Not recorded in the proper administration. Not to be used during the pregnancy.

Daily Dosage: 1 to 2 capsules

Chemical Analysis

The Fragaria & Frangula Combination / Fibromelt / had a chemical and antibacterial analysis in the Technology Center of the Plovdiv University "Paisii Hilendarski" EOOD, Bulgaria, by prof. Ilia Iliev

Methods of Analysis

Determination of the total polyphenolic content by the method of Singleton and Rossi

Determination of the total flavonoid content by spectrophotometric method with AlCl3 (Aluminium chloride)

Quantification of condensed tannins using the methyl cellulose precipitation method

Quantification of polyphenolic substances, representatives of the group of phenolic acids, flavonoids and tannins using the HPLC method (High-performance liquid chromatography)

Quantification of arbutin according to the European Pharmacopeia 10.0

Determination of antioxidant activity using the DPPH method LC-Mass Spectrometry analysis

The chemical analysis of the Fragaria & Frangula Combination / Fibromelt / shows the existing of the many useful chemical substances. The chemical substances with the highest concentration are grouped according to their physiological effects:

Effects of the Isolated Chemical Substances

Antibacterial - Gallic acid (Campylobacter, Escherichia coli, Listeria monocytogenes, and Staphylococcus aureus, Streptococcus mutans, Pseudomonas) [3-5]; Syringic acid [6]; Quercetin (Pseudomonas aeruginosa, Salmonella enteritidis, Staphylococcus aureus, Escherichia coli, Proteus, and Aspergillus flavus, Candida albicans and antiparasitic properties Leishmania, Trypanosoma, and Plasmodium) [7-9]; Kaempferol [10-12]; Isorhamnetin (Influenza virus, Staphylococcus aureus, Escherichia coli, Klebsiella pneumoniae, Proteus, Staphylococcus aureus and Bacillus, Mycobacterium tuberculosis) [13-15]; Rutin (Escherichia coli, Proteus vulgaris, Shigella sonnei and Klebsiella sp, Pseudomonas auruginosssa and Bacillus subtilis; Mycobacterium smegmatis; Antifungal activities - Candida gattii; Antimalarial activity - Plasmodium (Bennettinia) juxtanucleare, Plasmodium falciparum) [16-19]; Aucubin (Staphylococcus epidermidis, S. aureus, Enterococcus faecalis, and Bacillus subtilis, Proteus vulgaris, Enterobacter aerogenes, Klebsiella pneumoniae, Proteus mirabilis, and Citrobacter diverse, Candida albicans) [20-22].

Anti-inflammation – Gallic acid [3-5]; Chlorogenic acid; Caffeic acid; P-coumaric acid; Salicylic acid; Syringic acid [6]; Quercetin [7,9]; Myricetin; Kaempferol [10-12]; Isorhamnetin [13-15]; Rutin [16-19]; Aucubin [20,22-24]; Catalpol [25-28].

Antioxidant - verified with antioxidant analyze – (TPC –total polyphenol content, mg GAE/g – equivalent of gallic acid: 21.1; IC50 DPPH mg GAE/g 15 min: 10.6: EC50 CUPAC mg GAC/g: 6.3; EC50 FRAP mg GAC/g: 21.1); Gallic acid [3-5]; Chlorogenic acid; Caffeic acid; P-coumaric acid; Syringic acid [6]; Quercetin [7,9]; Kaempferol [10-12]; Isorhamnetin [13-15]; Rutin [16-19]; Aucubin [20,23]; Catalpol [25-28].

Hemostatic – tannins: Gallic acid [3,29]; Caffeic acid; flavonoid: Myricetin; Isorhamnetin [14]; Rutin [16,18].

Formation dissolving – Gallic acid [3,4,30-32]; Chlorogenic acid; Caffeic acid; Syringic acid [6]; Quercetin [7,33,34]; Myricetin; Kaempferol [10,11]; Isorhamnetin [14,35,36]; Rutin [16,18,19,37]; Aucubin [20,23,24]; Catalpol [25-27].

Antianxiety – Gallic acid [3,4,38,39]; Chlorogenic acid; Caffeic acid; Quercetin [7,40]; Kaempferol [41,42]; Isorhamnetin [14]; Rutin [16,18,43]; Aucubin [20,44]; Catalpol [25,26,28]

Antidepression - Gallic acid [3,4,38,39]; Chlorogenic acid; Caffeic acid; Quercetin [7,40]; Kaempferol [41,42]; Isorhamnetin [14]; Rutin [16,18,43]; Aucubin [20,44]; Catalpol [25,26,28]

Methods and Methodology

For the article were analyzed 63 women, ages ranging from 27 to 68 years of age. The period of administration of Fragaria & Frangula Combination / Fibromelt / was from 2 months to 1 year and 11 months, in the dose 1 capsule daily. All women were diagnosed with fibromyoma (CA 125 was negative for all women). The size of the fibromyomas were from 0.9 to 7.3 cm. The quality of the menstruation regularity, dysmenorrhea, metrorrhagia, spotting between or around menstruation, premenstrual syndrome (headache, nervousness, insomnia), and the relapse of the fibromyoma were analyzed for every one woman. The follow up period of the patients were 10 years. Nine of the women with the fibromyoma were in the climax. The size of the cysts was measured by the Ultrasound diagnostic apparatus – LOGIQ P6.

Results

The 39 patients have fully dissolved of the fibromyoma nodule, 24 patients have 22.7 - 85.7% decreasing at the size, and 0 patients had no response to the therapy (p < 0.1). The relapse of the fibromyomas was found at 6 women, with much less size. Five of them were dissolved again by Fragaria & Frangula Combination / Fibromelt / and 1 which was not responded had surgical removal. The other clinical symptoms that were analyzed in the treatment with the Fragaria & Frangula Combination / Fibromelt / were the irregularity of the menstruation, metrorrhagia, dysmenorrhea, and pre-menstrual symptoms. The 6 women had the irregular menstruation (5 with earlier menstruation, before 21 days and 1 woman with late menstruation, after 30 days). All women recovered the regularity of the menstruation after 2 to 4 months therapy (p < 0.4). The 20 women had dysmenorrhea (slight to very strong menstrual crumps and pain). The 16 women were without of the menstrual pain and cramps and 4 women were with the decreased menstrual pain and cramps after the therapy (p < 0.4). The 35 women were with metrorrhagia (over bleeding at the menstruation, with blood clots, 5 to 15 days bleeding; 6 women were with perimenstrual spotting; 3 women with ovulation spotting). The 32 women recovered the normal menstrual bleeding (3 to 7 days without blood clots), 3 women had decreased menstrual bleeding, with less blood clots (with normal menstrual duration), and no one had the spotting before or at the middle of the menstruation cycle (p < 0.3). The 6 women had the premenstrual syndrome (6 with headache and nervousness and 1 with breast pain). After the therapy, all women were without the premenstrual symptoms.

Table 1 represents summarized result of the therapy with the Fragaria & Frangula Combination / Fibromelt /

Clinic symptoms	Full	Fully	Partial
	number	compensated	compensated
1. Fibromyoma	63	39	24
1. Irregular menstruation	6	6	
* Earlier menstruation	5	5	
* Late menstruation	1	1	
2. Metrorrhagia	35	32	
* Spotting around and			
between the menstruations	9	9	
3. Dysmenorrhea	20	16	4
4. Premenstrual syndrome			
(headache, nervousness, and	6	6	
breast pain)			

Analyzes

The Fragaria & Frangula Combination / Fibromelt / is effective to dissolve the uterine fibromyomas (61.9 %), to stop the metrorrhagia, spotting around and between the menstruations (91.4 %), to prevent the dysmenorrhea (80.0 %), as well as to regulate the menstruation and premenstrual syndrome.

References

- 1. David Heber. PDF for Herbal Medicines. Fourth edition. Thomson Healthcare Inc. Montvale NJ. 2007.
- 2. Landgev Ilia. Trud Sofia. Encyclopedia of the healing herbs in Bulgaria. Bulgaria. 2005.
- 3. Milad Hadidi, Rafael Liñán-Atero, Mohammad Tarahi, et al. The Potential Health Benefits of Gallic Acid Therapeutic and Food Applications. Antioxidants Basel. 2024; 13: 1001.
- 4. Jinrong Bai, Yunsen Zhang, Tang Ce, et al. Gallic acid Pharmacological activities and molecular mechanisms involved in inflammation-related diseases. Biomed Pharmacother. 2021; 133: 110985.
- Kang Yang, Limeng Zhang, Pinfeng Liao, et al. Impact of Gallic Acid on Gut Health Focus on the Gut Microbiome Immune Response and Mechanisms of Action. Baichuan D Front Immunol. 2020; 11.
- Srinivasulu Cheemanapalli, Ramgopal Mopuri, Ramanjaneyulu Golla, et al. Syringic acid SAA Review of Its Occurrence Biosynthesis Pharmacological and Industrial Importance. Biomedicine & Pharmacotherapy. 2018; 108: 547-557.
- Dengyu Yang, Tiancheng Wang, Miao Long, et al. Quercetin Its Main Pharmacological Activity and Potential Application in Clinical Medicine. Oxid Med Cell Longev. 2020; 2020: 8825387.
- Thi Lan Anh Nguyen, Debanjana Bhattacharya. Antimicrobial Activity of Quercetin An Approach to Its Mechanistic Principle. National Center for Biotechnology Information. <u>https://www.ncbi.nlm.nih.gov/pmc/articles/PMC9029217</u>
- 9. Weidong Qi, Wanxiang Qi, Dongwei Xiong, et al. Quercetin Its Antioxidant Mechanism Antibacterial Properties

and Potential Application in Prevention and Control of Toxipathy. Academic Editor and Sara Vitalini Academic Editor. Molecules. 2022; 27: 6545. <u>https://www.ncbi.nlm.nih.</u> gov/pmc/articles/PMC9571766

- Argyrios Periferakis, Konstantinos Periferakis, Ioana Anca Badarau, et al. Kaempferol Antimicrobial Properties Sources Clinical and Traditional Applications. Int J Mol Sci. 2022; 23: 15054.
- 11. Jun Chen, Haopeng Zhong, Zhouyin Huang, et al. A Critical Review of Kaempferol in Intestinal Health and Diseases. Antioxidants Basel. 2023; 12: 1642.
- Bangar S, Vandana Chaudhary, Lorenzo J, et al. Kaempferol A flavonoid with wider biological activities and its applications. Crit Rev Food Sci Nutr. 2022; 63: 9580-9604.
- 13. Gomez-Zorita S, Trepiana J, Milton-Laskibar I, et al. Isorhamnetin Current knowledge and potential benefits for disease management. Handbook of Dietary Flavonoids. 2023; 1-61.
- 14. Gang Gong, Ying-Yun Guan, Zhong-Lin Zhang, et al. Isorhamnetin A review of pharmacological effects. Biomedicine & Pharmacotherapy. 2020; 128: 110301.
- 15. Christiana Eleojo Aruwa, Stephen O. Amoo, Neil Koorbanally, et al. Laccase-mediated modification of isorhamnetin improves antioxidant and antibacterial activities. Process Biochemistry. 2022; 112: 53-61.
- 16. Aditya Ganeshpurkar, Ajay K. Saluja. The Pharmacological Potential of Rutin. Saudi Pharm J. 2017; 25: 149-164.
- Mahesh Dilip Mane, Nilesh Shashikant Patole, Sanket Arun Metkari, et al. AN OVERVIEW OF ANTIMICROBEAL PROPERTIES OF RUTIN. IJNRD. 2024; 9.
- 18. Naif Abdullah Al-Dhabi, Mariadhas Valan Arasu, Chang Ha Park, et al. An up-to-date review of rutin and its biological and pharmacological activities. EXCLI J. 2015; 14: 59-63.
- 19. Koval'skii IV, Krasnyuk II, Krasnyuk II, et al. Mechanisms of Rutin Pharmacological Action Review. Pharmaceutical Chemistry Journal. 2014; 48: 73-76.
- 20. Kartini Kartini, Michelle Abigail Irawan, Finna Setiawan, et al. Characteristics Isolation Methods and Biological Properties of Aucubin. Molecules. 2023; 28: 4154.
- 21. Yunhui Liao, Feng Chen, Lujie Xu, et al. Study on extraction and antibacterial activity of aucubin from Eucommia ulmoides seed-draff waste biomass. Heliyon. 2022; 8: e10765.
- 22. Kairat Zhakipbekov, Aknur Turgumbayeva, Raushan Issayeva, et al. Antimicrobial and Other Biomedical Properties of Extracts from Plantago major Plantaginaceae. Pharmaceuticals. 2023; 16: 1092.
- 23. Xiangchang Zeng, Fei Guo, Dongsheng Ouyang. A review of the pharmacology and toxicology of Aucubin. Fitoterapia. 2020; 140.
- Min Shao, Ziyun Kuang, Wenlin Wang, et al. Aucubin Exerts Anticancer Activity in Breast Cancer and Regulates Intestinal Microbiota. Evid Based Complement Alternat Med. 2022; 2022: 4534411.

- 25. Subrat Kumar Bhattamisra, Kah Heng Yap, Vikram Rao, et al. Multiple Biological Effects of an Iridoid Glucoside Catalpol and Its Underlying Molecular Mechanisms. Biomolecules. 2020; 10: 32.
- Shuanglin Liu, Yuanfang Kong, Juntao Cai, et al. Advances in Structural Modification and Pharmacological Activity of Catalpol and its Derivatives. Chemistry Select. 2021; 6: 13520-13535.
- 27. Na Gao, Jian-Xin Tian, Yu-Hong Shang, et al. Catalpol Suppresses Proliferation and Facilitates Apoptosis of OVCAR-3 Ovarian Cancer Cells through Upregulating MicroRNA-200 and Downregulating MMP-2 Expression. Int J Mol Sci. 2014; 15: 19394-19405.
- 28. Xiaohui Wu, Chen Liu, Junming Wang, et al. Catalpol Exerts Antidepressant-Like Effects by Enhancing Anti-oxidation and Neurotrophy and Inhibiting Neuroinflammation via Activation of HO-1. Neurochem Res. 2022; 47: 2975-2991.
- 29. Yonghui He, Juan Wang, Wei Yan, et al. Gallic acid and gallic acid-loaded coating involved in selective regulation of platelet endothelial and smooth muscle cell fate. RSC Advances. 2014; 1.
- Subramanian AP, John AA, Vellayappan MV, et al. Gallic acid prospects and molecular mechanisms of its anticancer activity. RSC Advances. 2015; 45.
- Milad Ashrafizadeh, Ali Zarrabi, Sepideh Mirzaei, et al. Gallic acid for cancer therapy Molecular mechanisms and boosting efficacy by nanoscopical delivery. Food Chem Toxicol. 2021; 157: 112576.
- 32. Victoria Sanchez-Martin, María del Carmen Plaza. Gallic Acid A Natural Phenolic Compound Exerting Antitumoral Activities in Colorectal Cancer via Interaction with G-Quadruplexes. Cancers Basel. 2022; 14: 2648. <u>https://</u> www.ncbi.nlm.nih.gov/pmc/articles/PMC9179882
- 33. Mingxin Guo, Jiaqi Zeng, Zhangzhang Sun, et al. Research Progress on Quercetin's Biological Activity and Structural Modification Based on Its Antitumor Effects. Chemistry Select. 2023; 8: e202303167.
- 34. Si-Min Tang, Xue-Ting Deng, Jian Zhou, et al. Pharmacological basis and new insights of quercetin action in respect to its anticancer effects. Author links open overlay panel. Biomedicine & Pharmacotherapy. 2020; 121: 109604.

- 35. Partha Biswas, Abu Kaium, Mohaimenul Islam Tareq, et al. The experimental significance of isorhamnetin as an effective therapeutic option for cancer A comprehensive analysis. Biomedicine & Pharmacotherapy. 2014; 176: 116860.
- Qiong Li, Fu-Qiang Ren, Chun-Lei Yang, et al. Antiproliferation effects of isorhamnetin on lung cancer cells in vitro and in vivo. Asian Pac J Cancer Prev. 2015; 16: 3035-3042.
- 37. Atefeh Satari, Sorayya Ghasemi, Solomon Habtemariam, et al. Rutin A Flavonoid as an Effective Sensitizer for Anticancer Therapy Insights into Multifaceted Mechanisms and Applicability for Combination Therapy. Evid Based Complement Alternat Med. 2021; 23: 9913179.
- Neurobiological effects of gallic acid: current perspectives. National Center for Biotechnology Information. 2023. <u>https://www.ncbi.nlm.nih.gov/pmc/articles/PMC10015939</u>
- 39. Maria Lúcia da Silva Cordeiro, Verônica Giuliani de Queiroz Aquino Martins, Ariana Pereira da Silva, et al. Phenolic Acids as Antidepressant Agents. Nutrients. 2022; 14: 4309.
- 40. Shen Chen, Yueheng Tang, Yang Gao, et al. Antidepressant Potential of Quercetin and its Glycoside Derivatives A Comprehensive Review and Update. Front Pharmacol. 2022; 13: 865376. <u>https://www.ncbi.nlm.nih.gov/pmc/articles/</u> <u>PMC9024056</u>
- Jéssica Silva dos Santos, João Pedro Gonçalves Cirino, Patrícia de Oliveira Carvalho, et al. The Pharmacological Action of Kaempferol in Central Nervous System Diseases A Review. Front Pharmacol. 2020; 11: 565700.
- 42. Wenqi Gao, Wei Wang, Yan Peng, et al. Antidepressive effects of kaempferol mediated by reduction of oxidative stress, proinflammatory cytokines and up-regulation of AKT/βcatenin cascade. Metab Brain Dis. 2019; 34: 485-494. <u>https://</u> <u>pubmed.ncbi.nlm.nih.gov/30762138</u>
- 43. Ahmed I Foudah, Mohammed H Alqarni, Aftab Alam, et al. Rutin Improves Anxiety and Reserpine-Induced Depression in Rats. Molecules. 2022; 27: 7313.
- 44. Ping Yang, Qiaoyue Zhang, Hengyan Shen, et al. Research progress on the protective effects of aucubin in neurological diseases. Pharm Biol. 2022; 60: 1088-1094.

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