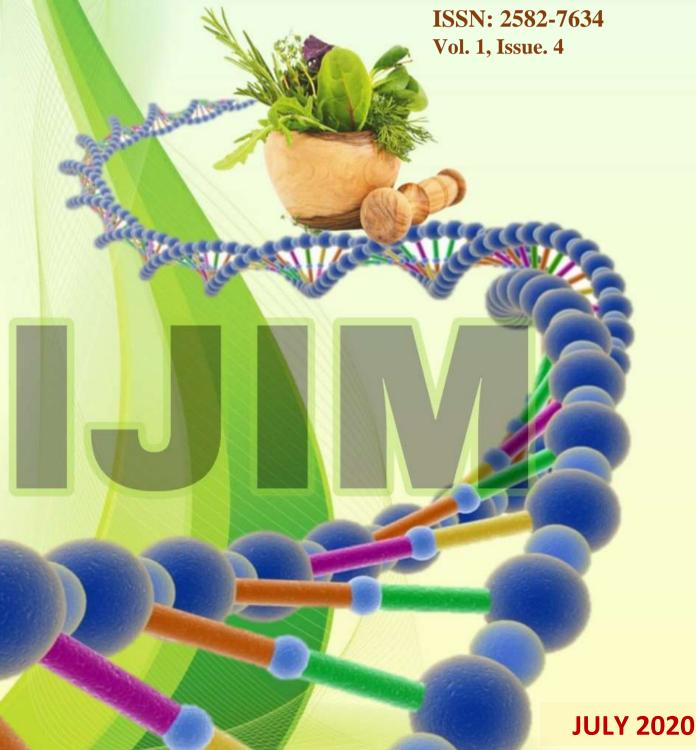


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# "SYSTEMIC REVIEW OF AMRUTMANJIRI, A POTENT AYURVED HERBOMINERAL COMBINATION AGAINST COVID-19 PANDEMIC."

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#### **Abstract:**

A Novel corona virus was identified as the causative agent and was subsequently termed COVID-19 by the World Health Organization and there is no scientific treatment for the disease till date. *Ayurveda* is a science of life main focus of *ayurveda* to cure the disease root causes and maintain the health of healthy individuals. *Amrutmanjiri* ras is a compound drug used in *sannipataj jwar*, *Kasa Shwas*, *Aamavat*, The content present in *Amrutmanjiri* are having properties such as antiviral ,expectorant, reducing infection of upper respiratory tract. An *Amrutmanjiri* content has been included in Indian pharmacopeia as well as in literature of many other ancient civilization. In *Ayurveda Amrutmanjiri* is one of the important medicine which is been referred in many texts with many therapeutic uses. Thus the content of *Amrutmanjiri* is found to have prominent antiviral activity .so we can use *Amrutmanjiri* as Antiviral Drug.

Keywords: Amrutmanjiri, Sannipatik jwar, Covid -19 treatment, Fever, Saam jwar, Aamvat.

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#### INTRODUCTION:

COVID-19 is caused by Beta corona virus named SARS-CoV-2 that affect the lower respiratory tract and manifest as pneumonia in Humans<sup>[1]</sup>, structurally the SARS corona virus (SARS-CoV) has a well - defined composition comprising 14 binding residues that directly interact with human angiotensin-converting enzyme 2.of these amino acids, 8 have been conserved in SARS CoV-2<sup>[2]</sup>. Most SARS-CoV-2 infected patients have developed mild symptoms such as dry cough, sore throat, and fever. The majority of cases have spontaneously resolved. However, some have developed various fatal complications including organ failure, septic shock, pulmonary oedema, severe pneumonia, and Acute Respiratory Distress Syndrome (ARDS)[3]At present, no effective antiviral treatment or vaccine is available for COVID-19. However, a randomized multicentre controlled clinical trial is currently underway to assess the efficacy and safety of patients with COVID-19 (ChiCTR2000029573). First-line treatment for fevers includes antipyretic therapy such as paracetamol, while ex- pectorants such as guaifenesin may be used for a non-productive cough. [4] There are currently few studies that define the patho physiological characteristics of COVID-19, and there is great uncertainty regarding its mechanism of spread. Current knowledge is largely derived from similar corona viruses, which are transmitted from human-to-human through respiratory fumets [5] Same in ayurveda sankramik disease.

Ancient wisdom has been the basis of modern medicine and therapeutics. An impressive number

of drugs have been isolated from natural sources. Many of this isolation are based on the use of these agents in traditional medicine. Amrutmanjiri ras are mentioned in our classics, among which the familiar one which is described in Bhaishajya ratnavali has been taken for review in this article. Man has been using Natural products, including plants, animals and minerals have been the basis of treatment of disease. In the History of human The disease which persist at the time of death is jwara (HYPER PYREXIA), In Ayurveda jwara is given top most importance because it is believed that jwara is the first disease to trouble mankind and it becomes the basic for other disease to exist. Many type of *jwaras* are explained acharyas/scholars and one of them is sannipataj  $jwara^{[6]}$ .

#### AMRUTMANJIRI IN DETAIL[7]

Amrutmanjiri rasa an Ayurvedic herb mineral formulation is indicated in the management of *jwara* (Hyper pyrexia) and Sannipataj jwara. It contains two mineral drugs and five herbal drugs.

Matra-125 to 250 mg

**Anupan**-Aadrak swaras, Madhu,Tambul patraras, Ushndak.

Swad- Aamla.

Varna- Raktabha

**Indication of Amrutmanjiri** – Sannipat jwara, Aagnimandya, Aajirna, Aamavat, Kasa five types, Shwasa, Sarvang graham, Jirnajwara, Kshya.

Table 1 showing Amrutmanjiri Ingredients In detail-

Content	Quantit	Rasa	Viry	Vipak	Guna	Doshaghnata	Indication
	у	(taste)	а				
Shudha Hingula [8]	1 part	Tiktha	Ushn	-		Tridoshaghna	Jwara, Aamavat, Rasayan,
(cinnabar)		Kasha	а				Deepan, Vrushya, Sarva

		ya , Katu .					Rogaghna, Kamala,Pliha, Kushta and Vishanashak.
Vatsanabh <sup>[9]</sup> (Aconi tum ferox)	1 part	Madhu r	Ushn a	Katu	Laghu, Ruksh, Teekshna, Vyavayi,	Jwarahara, Jangamvishaha ra, Madakari,	Jwara(Sannipataj), Madhumeha, Shwasa, Kasa, Pleehodara, Agnimandya, Vata rogas,
Shudha Tankan <sup>[10]</sup> (Borax)	1 part	Katu	-	-	Vikasi Ushna,Ruks ha, Teekshna, Sara.	Kushthaghna, Vatakaphaghna	Kaphavishleshaka, Hrudhya, Kasa Shwsa Hara, Sthavara vishanashaka,AgniDeepak
Marich <sup>[11]</sup> (Piper nigrum)	1 part	Katu	Ushn a	Aardr a- Madh ur	Ruksha, Tikshna	Pittakar, Kaphahara, Vatahara	Shwasa, Shula, Krumirog nashak
Pippali <sup>[12]</sup> (Piper longum)	1 part	Katu	Ushn a	Madh ur	Laghu, Sneegdha, Teekshna.	Vatashleshmah ara, Deepana, Vrushya, Rasayan.	Jwara,Aamavat, Udara, Pliha Roga,Kushta,Prameha,Gu Ima, Arsha, Shula.
Javitri [13] patra(Myristica fragrans)	1 part	Tikta,	Ushn a	Katu	Laghu, Tikshna,	Kaphaghna, Vataghna, Deepan, Grahi, Ruchikar	Kasa,Krumi,Shwasa, Shosha,Rhudrog, Pinas.Mukhadurgandha hara
Jambiri nimbu swaras <sup>[14]</sup> (Citrus lemon)	For trituratio n.	Aamla	Ushn a	Aamla	Guru,	Vata, Kaphahara	Kasa, Kaphotkleshaghna,Krumi ghna, Aasyavairasyahara. Rhutpida, Shulaghna,

#### 1)Shuddha Hingul (Cinnabar)

Therapeutic Utility of *Shuddha Hingula* [15,16,17] - *Shuddha Hingula* (*cinnabar*) when judiciously used in the form of suitable compound formulation and in proper dosage is beneficial in all types of eye diseases (*Netra Roga Hara*), it mitigates all the three vitiated *Doshas*. It is good appetizer, rejuvenator and aphrodisiac. It is the best remedy for the disorders of liver, pancreas, rheumatoid (arthritis *Aamavata*), diabetes mellitus (*Prameha*) and fever, is nutritive, It enhances body strength,

memory, facial glow and power of digestion. *Cinnabar* is used as sedative for its neuroactvity <sup>[18]</sup>. HgS adjust the imbalance of Pro-oxidant and Antioxidant, provides a protective effect in oxidative stress which is induced by Hypoxia<sup>[19]</sup>.

#### 2)Vatsanabh

Vatsanabh is having properties like Rasayan(rejuvenating) Balavardhanam( improve strength and immunity), jwarahara<sup>[20]</sup> (useful in fever), Shirashool<sup>[21]</sup> Vatsanabh acts as Yogavahi –a catalyst for other medicine Vatsanabh used with equal quantity of tankan (purified Borax) and double quantity of black paper (Marich)<sup>[22]</sup> will not

have any toxic effects. It is also *Rasayan*, *Sukhaprada*, *Balya*, *Shwas Kasa Bhagandar Gulma*, *Pandu*, *Aarsha*.<sup>[23]</sup>

#### 3) Tankana (Borax)

*Tankan* contain Na2B4O7 10H2O which is composed of boric acid and soda<sup>[24]</sup>.

As antidote-

Tankan is might be to minimize toxic effects of *Vatsanabh*. This concept highlights the antidote property of Tankan in *Vatsanabh* toxicity.<sup>[25]</sup>

#### Wound healing -

Tankan is used in the process of repair that follows injury to the skin and other soft tissues. The objective in wound management is to heal the wound in the shortest time possible, with minimal pain, discomfort, and scarring to the patient. [26]

#### **Another Uses-**

- -It is used in the treatment of cough, bronchitis.
  - -It is also used in treating food poisoning.
  - -It improves digestion power, relieves bloating.
- -It induces menstruation in women suffering with amenorrhea or oligomenorrhea (Scanty menstrual flow).
- -It is useful in dandruff Tankan is applied on hair and keeps for 5 to 10 min and washes out.
- -It is useful ulceration of mouth and cracks and fissures of tongue.
- -Dose varying from 10-30 grains are given in prolonged labour.
- -Useful in other uterine infection.
- -Useful in chronic tonsillitis.

The gargling with *Tankan bhasma* dissolved water gives significant result in relieving the tonsillitis [27].

#### 4)Marich

Pharmacological Activities of *P. Nigrum* - Antidiarrheal<sup>[28]</sup>, Antimicrobial<sup>[29]</sup>, Antioxidant<sup>[30]</sup>,

Anticancer<sup>[31]</sup> and tumour reduction activity<sup>[32]</sup>, Antihypertensive<sup>[33]</sup>, Anti-asthmatic, Anti-inflammatory<sup>[34]</sup>, Anti-obesity<sup>[35]</sup>, Antidiabetic<sup>[36]</sup>, Hepatoprotective activity <sup>[37]</sup>, Digestive activity<sup>[38]</sup>, Antidepressant<sup>[39]</sup>, Analgesic and Antipyretic activity <sup>[40]</sup>, Anti-adipogenesis activity<sup>[41]</sup>, Effects in metabolism<sup>[42]</sup>, Effects in neurodegeneration and cognitive impairment <sup>[43]</sup>, Effect in Bioavailability<sup>[44]</sup>, Toxicity activity<sup>[45]</sup>, Effects in stomach<sup>[46]</sup>, Effect in bile secretion<sup>[47]</sup>, Effect in pharmacokinetic of drugs<sup>[48]</sup>.

#### 5)Pimpali

Pharmacological Activity of *Piper longum*:-Insecticidal and acaricidal activity<sup>[49]</sup>:, Antifungal activity<sup>[50]</sup>, Antiamoebic activity<sup>[51]</sup>, Antimicrobial activity<sup>[52]</sup>, Effect on respiratory system<sup>[53]</sup>, Antiasthmatic activity<sup>[54]</sup>, Effect on cardiovascular system<sup>[55]</sup>, Antidiabetic activity<sup>[56]</sup>, Hypochoesterolaemic activity<sup>[57]</sup>, Antioxidant activity<sup>[58]</sup>, Analgesic activity <sup>[59]</sup>, Anti-inflammatory activity <sup>[60]</sup>, Immunomodulatory activity<sup>[62]</sup>, Anti-cancer activity<sup>[62]</sup>, Antidepressant activity<sup>[63]</sup>, Antiulcer activity<sup>[64]</sup>, Effect on Reproductive system<sup>[65]</sup>, Bioavailability enhancement <sup>[66]</sup>, Hepatoprotective activity<sup>[67]</sup>.

#### 6) Javitri.

Pharmacological Activity- Anti-bacterial, Antimicrobial and Anti-fungal property [68] – Effective against Gram-positive and Gram-negative microbes . Hypoglycemic and antidiabetic activities [69] - It is potential anti-diabetic agent for the treatment of type 2 diabetes study done in rats. Hypolipidaemic and platelets antiaggregatory activity [70]- Reducescholesterol in heart and liverLDL and VLDL levels were also reduced in albino significantly study. Hepatoprotective activity [71] - due to the inhibition of Tubour Necrosis Factor(TNF)-alpha release from macrophages in mice ,Antiinflammatory activity [72] - The anti-inflamatory property of myristicin might be due to inhibition of chemokines, cytokines, nitrous oxide and

growth factors in double stranded RNA stimulated macrophages via the calcium pathway, Anticancer activity  $^{[73]}$ , Memory enhancing activity  $^{[74]}$ , Aphrodisiac activity  $^{[75]}$ , Anti-diarrhoeal activity  $^{[76]}$ , Antidepressant activity  $^{[77]}$ , Anti-oxidant activity  $^{[78]}$ , Pesticide activity: insecticidal  $^{[79]}$ 

#### 7) Jambir

Anticancer activity [80] – Inhibition of the proliferation of cancer cells . Activation of "TRAIL" apoptotic cell death .Inhibition of tumour growth in chronic erogenous leukaemia (CML)

Antioxident activity<sup>[81]</sup> – Augmentation of antioxidant cellular defences via ERK/Nrf2 signalling pathway.

Anti-inflammatory activity [82] Antibacterial activity [83] – Inhibiting activity against Gram –positive bacteria ,Enterococcus faecalis, Bacillus subtitles, Inhibiting activity against Gram – negative bacteria Salmonela typhimurium, Shigella sonnei , Helicobacter pylori.

Antifungal activity [84] – Inhibition activity against Candida glabrata strains.

Antiviral activity [85] – Inhibition of replication of herpase simplex

Anti-allergic activity [86] – Inhibition of histamine secretion in peretonial cell of rats.

Hepatogenerative activity<sup>[87]</sup> – Normalization of alanine aminotraneferase, alkaline phospatase and bilirubine.

Prevention of diabetes and treatment of its symptoms [88]\_ Inhibition of Gluconeogenesis, Reducing wound healing time,Increasing tissue growth rate, collagen synthesis and protein and hydroxyproline concentration.

Anti obesity activity [89] – Lowering blood lipids, Reducing the level of insulin, leptin and adiponectin in the blood

**Effect on cardiovascular system**<sup>[90]</sup> – Limiting myocardial damage, Decreasing blood fibrinogen, Lowering blood pressure in people with hypertension.

**Effects on nervous system** [91] – Strengthen short term memory.

**Effects on the respiratory system** [92] – Treatment in chronic pneumonia

Effects on skeletal system [93] – Increases bone density, Decreases osteoclast activity, Decreasing bone resorption activity (nomilin)

Treatment on menstrual disorder [94] – Period induction in case of irregular menstrual cycles.

#### **DISCUSSION-**

COVID-19 Which is caused by droplet infection of corona virus and enters in to body through Nasal and Buccal mucosa and increases in number. it primarily acts on respiratory system and patient may develop Acute Respiratory Distress Syndrome.

Amrutmanjiri is usefl in Aamashaya udbhavjanya vyadhi such as,Sannipataj jwarw Shwas, Kasa Jwara Aamavata. Main cause of sannipataj jwara is aama and Aam is nothing but the Endogenous toxins. Formed due to digestive error , metabolic error ,immunological reaction and hypersensitivity reaction. Amrutmanjiri is having Krumighna property due to which it acts as Antiviral, Antibacterial, Antifungal, Anticancer and Antimicrobial activity.

Amrutmanjiri is having immunomodulatory activity which is useful to avoil the disease also having the *swedakar* property which is responsible to reduce the *jwarw* (hyper pyrexia) and act on all type of *vyadhi*. Due to these properties of *Amrutmanjiri* we can use it in COVID-19 disease.

#### **CONCLUSION:**

As the *Amrutmanjiri ras* is indicated in *shwas*, *kaas*, *jwar*, *sarvanggrah*, *kshay*, etc. And same symptoms were found in COVID -19, so we canbe used *amrutmanjiri ras* in COVID-19 symptomatically. This is my small efforts, how this medicine will act by their ingredients *doshaghnata* as well as by their qualities. But there is a need of further preclinical study.

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#### **Conflicts of interest**

Nil

#### **References:**

- 1. World Health Organization, WHO Director-General's Remarks at the Media Briefingon 2019-nCoV on 11 February 2020, (2020) https://www.who.int/dg/speeches/detail/who-director-general-s-remarks-at-the-media-briefing-on-2019-ncov-on-11- february-2020.
- 2. (A.R. Fehr, S. Perlman, Coronaviruses: an Overview of Their Replication and Pathogenesis, (2015), pp. 1–23.
- 3. N. Chen, M. Zhou, X. Dong, et al., Epidemiological and clinical characteristics of 99 cases of 2019 novel coronavirus pneumonia in Wuhan, China: a descriptive study, Lancet (2020), https://doi.org/10.1016/S0140-6736(20)30211-7.
- 4. D. Wang, B. Hu, C. Hu, et al., Clinical characteristics of 138 hospitalized patients with 2019 novel coronavirus–infected pneumonia in wuhan, China, J. Am. Med. Assoc. (2020),
- 5. Centers for Disease Control and Prevention, 2019 Novel Coronavirus, (2020) https://www.cdc.gov/coronavirus/2019-ncov/about/transmission.html.
- 6. Kaviraj Govind Das sen, Bhaishajyaratnavali, edited with sidhiprada Hindi commentary by prof.Sidhinandan Mishra, Chaukhamba surbharti prakashan, Varanasi,Edition in 2013.chapter 5/552,553 page no.138.
- 7. Kaviraj Govind Das sen, Bhaishajyaratnavali, edited with sidhiprada Hindi commentary by

- prof.Sidhinandan Mishra, Chaukhamba surbharti prakashan, Varanasi,Edition in 2013.chapter 5/552,553 page no.138.
- 8. shri vagbhatacharya rasa ratnasamuchay edited by pandit shri Dharmanandan Sharma second edition Varanasi Motilal Banarasi Das 1996. Chapter version 105, 60 pp
- 9. Shri Sadanand Sharma Rasa tarangini 11 edition Varanasi Motilal Das Banarasi Publication 1989 14<sup>th</sup> Taranga<sup>-</sup>26-31 shlok , 653pp
- 10. Shri Sadanand Sharma Rasa tarangini 11edition Varanasi Motilal Das BanarasiPublication 1979 . Taranga 13 Verses 79 81 ,667pp
- 11. Shatri J.L.N. Dravyaguna Vignana 2<sup>nd</sup> edition Varanasi: Chaukamba orientalia, 2005, second part, 4449 pp
- 12. Prof . Levekar G.S. et,al Data Base on Medicinal Plants used in Ayurveda and siddha . new Delhi : published by CCRAS 2007 Vol 3, 472 pp .
- 13.Dr. Gangasahay pandey, Dr Krushnachandra Chunekar Bhavprakash Nighantu nighantu Indian Materia medica reprint year 2006 Publication Chaukhamba bharti Academy Varanasi. page no 216
- 14.Dr. Gangasahay pandey , Dr Krushnachandra Chunekar Bhavprakash Nighantu nighantu Indian Materia medica reprint year 2006 Publication Chaukhamba bharti Academy Varanasi. page no 594
- 15. Bhudev Mukharjee, Rasa Jala Nidhi, Vol-2,

Chaukambha publishers, Varanasi, 3<sup>rd</sup> Edition:1998, Chapter 3, Pp-225

16.Acharya Yashodhara Bhatta, Rasa Prakash Sudhakara, Siddhiprada Hindi Commentry By Dr Siddhinandan Mishra, Chaukambha Orientalia, Varanasi, 3<sup>rd</sup> Edition, 2004, Verse No-6/87-88, Pp-130.

17.Acharya Somadeva, Rasendra Chudamani, Siddhiprada Hindi Commentry By Dr Siddhinandan Mishra, Chaukambha Orientalia, Varanasi, 2<sup>nd</sup> Edition, 1999, 7/108, Pp-107.

18.Chuu J.J.Liu.,Lin S.H.,Lin-Shiau S.Y.Differential neurotoxic effect of methylmercury and mercuric sulfide in rats.Toxicol.Lette.2007;169:10120.doi:10.1016/j.toxlet.2006.12.006.

[PubMed][CrossRef][Google Scholar]

19. Qiangqiang He, Ji Ma. HgS Inhibits Oxidative stress caused by Hypoxia through Regulation of 5-HT Metabolism Pathway, A Research Article Int. J. Mol. Sci. 2019. Mar; 20(6):1364 doi:10.3390/ijms 20061364 [PubMed] [Cross Ref] [Google Scholar]

20. Dr. Gangasahay pandey, Dr Krushnachandra Chunekar Bhavprakash Nighantu nighantu Indian Materia medica reprint year 2006 Publication Chaukhamba bharti Academy Varanasi. Bhavprakash purvasthan 7/253-254 21.Shri Sadanand Sharma Rasa tarangini 11 edition Varanasi Motilal Das Banarasi Publication 1979. Rasatarangini 24/26-31 22.Vd.Laxmipati shastri Yogratnakar with vidyotini Hindi comentry, Edition 2012. Chaukhamba prakashan Varanasi, page no 166. 23.Vd.Laxmipati shastri Yogratnakar with vidyotini Hindi comentry,Edition 2012. Chaukhamba prakashan Varanasi, page no 166. 24. Mustafa K, Histological Evaluation of the Effects of Borax Obtained from Various Sources in Different Rat Organs, Int. J. Morphol, 2015;33(1), 255-261

25. Nilesh K, Evaluation of Prativisha Properties (Antidote Properties) of Tankana (Borax) In Vatsanabha Vishaktata (Aconite Poisoning), International Ayurvedic Medical Journal, 2015;3(7),1991-1998.

26. Swati R, Development and Study of Wound Healing Activity of an Ayurvedic Formulation, Asian J. Res. Pharm. Sci., 2011; 1(1), 26-28.

27. <a href="http://ayurmedinfo.com/tankan-bhasma-benefits-dosage-side-effects ingredients,2012">http://ayurmedinfo.com/tankan-bhasma-benefits-dosage-side-effects ingredients,2012</a>.

28. Shamkuwar PB, Shahi SR and Jadhav ST: Evaluation of anti-diarrheal effect of Black pepper (*P. nigrum* L). Asian Journal of Plant

29. Zang J, Ye KP, Zang X, Pan DD, Sun YY and Cao JX: Antibacterial activity and mechanism of action of black pepper essential oil on meatborne *Escherichia coli*. Front. Microbiol. 2017; 7: Article-

https://doi.org/10.3389/fmicb.2016.02094.

Science and Research. 2012; 2: 48-53.

30. JS Swathy, Mishra P, Thomas J, Mukherjee A and Chandrasekaran N: Antimicrobial potency of high-energy emulsified black pepper oil nanoemulsion against aquaculture pathogen. Aquaculture. 2018; 491: 210-220. https://doi.org/10.1016/j.aquaculture.2018.03.045.

- 31. Ramji MT, Deepthi K, Lakshmi KA and Uma P: *In silico* docking analysis of piperine amino acid analogues against carcinogenic activating enzymes. Biotechnology. 2011; doi:10.4172/jpb.1000240.
- 32. Pradeep CR and Kuttan G: Effect of piperine on the inhibition of lung metastasis induced B16F-10 melanoma cells in mice. Clin Exp Metastas. 2002; 19: 703-708.
- 33.JS Swathy, Mishra P, Thomas J, Mukherjee A and Chandrasekaran N: Antimicrobial potency of high-energy emulsified black pepper oil nanoemulsion against aquaculture pathogen. Aquaculture. 2018; 491: 210-220.

https://doi.org/10.1016/j.aquaculture.2018.03. 045.

34.Wang B, Zhang Y, Huang J, Dong L, Li T and Fu X: Anti- inflammatory activity and chemical composition of dichloromethane extract from *Piper nigrum* and *P. longum* on permanent focal cerebral ischemia injury in rats. Revista Brasileira de Farmacognosia. 2017; 27: 369-374. 35.Shah SS, Shah GB, Singh SD, Gohil PV, Chauhan K, Shah KA and Chorawala M: Effect of piperine in the regulation of obesity-induced dyslipidemia in high-fat diet rats. Indian J. of Pharmacology. 2011; 43(3): 296-299.

36. Vazhacharickal PJ, Mathew JJ and Babu A: Phytochemical and anti-diabetic activities of different plant part extracts among black pepper (*Piper nigrum*) varieties in comparison with *Piper longum* and Piper betel. Scientific Study. 2017.

37. Sarfaz M, Khaliq T, Khan JA and Aslam B: Effect of aqueous extract of black pepper and ajwa seed on liver enzymes in alloxan-induced diabetic Wister albino rats. Saudi Pharmaceutical Journal. 2017; 25(4): 449-452. https://doi.org/10.1016/j.jsps.2017.04.004.

38. Myers BM, Smith JL and Graham DY: Effect of red pepper and black pepper on the stomach. Am J Gastroenterol. 1987; 82: 211-214.

39. Lee SA, Hong SS, Han XH, Hwang JS, Oh GJ, Lee KS, Lee MK, Hwang BY and Ro JS: Piperine from the fruits of *Piper longum* with inhibitory effect on monoamine oxidase and antidepressant like activity. Chem. Pharma. Bull. 2005; 53: 832-835.

40. Sabina EP, Nasreen A, Vedi M and Rasool M: Analgesic, antipyretic and ulcerogenic effects of piperine: An active ingredient of pepper. J. pharm. Sci. and Res. 2013; 5(10): 203-206.

41. Park UH, Jeong HS, Jo EY, Park T, Yoon SK, Kim EJ, Jeong JC and Um SJ: Piperine, a component of Black Pepper, inhibits adipogenesis by antagonizing PPARy activity in

3T3-L1 Cells. J. Agrc. Food Chem. 2012; 60: 3853-3860.

42. Lee CS, Han ES, and Kim YK: Piperine inhibition of 1- methyl- 4- phenylpyridinium - induced mitochondrial dysfunction and cell death in PC12 cells. Europ. J. Pharma. 2006; 537: 37-44.

43. Chonpathompikunlert P, Wattanathorn J and Muchimapura S: Piperine, the main alkaloid of Thai black pepper, protects against neurodegeneration and cognitive impairment in animal model of cognitive deficit like condition of Alzheimer's disease. Food. Chem. Toxicol. 2010; 48: 798-802.

44. Mohapatra SS, Sarma J, Roy RK, Panigrahi S and Ganguly S: Ethnomedicinal plants used in Balasore District of Odisha: A Comprehensive Report. Int. J. Curr. Microbiol. App. Sci. 2018; 7(1): 1959-1963.

45. Piyachaturawat P, Glinsukon T and Toskulkao C: Acute and subacute toxicity of piperine in mice, rats and hamsters. Toxicology Letters. 1983; 16: 351-359.

46.Myers BM, Smith JL and Graham DY: Effect of red pepper and black pepper on the stomach. Am J Gastroenterol. 1987; 82: 211-214.

47.Bhat GB and Chandrasekhara N: Effect of black pepper and piperine on bile secretion and composition in rats. Nahrung. 1987; 31: 913-916. 48.Pattanaik S, Hota D and Prabhakar S: Effect of piperine on the steady-state pharmacokinetics of phenytoin in patients with epilepsy. Phytother Res. 2006; 20: 683-686.

49. Kokate CK, Tipnis HP, Gonsalvis LX, and D'cruz JL, Anti-insect and juvenile hormone mimicking activities of essential oils of Adhatoda vasica, *P. longum* and Cyperus rotundus in *4<sup>th</sup> Asian Symposium of medicinal plants*, Spices, Bangkok, Thailand, 1980,154-158.

50. Bhargava AK, Chauhan CS, Antibacterial activity of essentialoils, *Indian J Pharm*, 1968, 30, 150-152. 51. Sawangjaroen N, Sawangjaroen K and

- Poonpanang P, Antiamoebic effects of *Piper longum* fruit, *Piper sarmentosum* root and *Quercus infectoria* nut gall on caecal amoebiasis in mice, *J Ethnopharmacol*, 91(2-3), 2004,357-360.
- 52. Khan M and Siddiqui M, Antimicrobial activity of fruits of *Piperlongum*, *Nat prod Rad*, 6, 2007, 111-113.
- 53. Kulshresta VK, Singh N, Shrivastava RK, Kohli RP, Rastogi SK, A study of central stimulant activity of *Piper longum*, *J Res Indian Med*, 6(1), 1971,17-19.
- 54. Kulshresta VK, Singh N, Shrivastava RK, Kohli RP, A study of central stimulant effect of *Piper longum*, *Indian J Pharmacol*, 1(2), 1969,8-10
- 55.Lee SW, Rho MC, Park HR, Choi JH, Kang JY, Lee JW, Kim K, Lee HS, Kim YK, Inhibition of diacylglycerol acyltransferase by alkamides isolated from the fruits of *Piper longum*, *J Agri Food Chem*, 54(26), 2006, 9759-9763.
- 56. Dhar ML, Dhar MM, Dhavan BN, Malhotra BN, Ray C, Screening of Indian Plants for biological activity-1, *Indian J Exp Bio*, 6, 1968, 232-235.
- 57.Wu E, Bao Z, Effects of unsaponificable matter of *Piper longum* oil on cholesterol biosynthesis in experimental hypocholestrolaemic mice, *Honggacayano*, 23(4), 1992, 197-200.
- 58.Natarajan KS, Narasimhan M, Shanmugasundaram KR, and Shanmugasundaram ER, Antioxidant activity of a salt-spice-herbal mixture against free radical induction , *J Ethnopharmacol*, 105(1-2),2006,76-83.
- 59. Vedhanayaki G, Shastri GV, Kuruvilla A, Analgesic activity of Piper longum Linn. Root, *Indian J Exp Biol*, 41(6), 2003, 649-651
- 60. Sharma AK, and Singh RH, Screening of antiinflammatry of certain indigenous drugs on carrageeninducedhind pawedemainrats, *Bull Med Ethanobot Res*, 2, 1980, 262-264.
- 61. Devan P, Bani S, Suri KA, Satti NK, and Qazi GN, Immunomodulation exhibited by piperinic acid of *Piper longum L.*, through suppression of

- proinflammatory cytokines, *Int Immunopharmacol*, 7(7), 2007,889-899.
- 62.Arion, 1967. Report of the Composite Drug ReseachScheme,ICMR,NewDelhi,pp.243-245.
- 63. Song L, Che W, Minwei W, Wei L, Kinzo M and Yiyuan T, Antidepressant like effects of piperine in chronic mild stress treated mice and its possible mechanisms, *Life Sci*, 80(15), 2007,1373-1381.
- 64. Bajad S, Bedi KL, Singla AK, Johri RK, Piperine inhibits gastric emptying and gastrointestinal transitin rats and mice, *Planta Med*, 67(2), 2001, 176-179.
- 65.Kholkute SD, Kekere MB, Munshi SR, Antifertility effects of the fruits of *P.longum* in female rats, *Indian J Exp Biol*, 17, 1979, 289-290.
- 66. Atal CK, Zutshi U and Rao PG, Scientific evidence on the role of Ayurvedic herbals on bioavailability of drugs. *J Ethnopharmacol*, 4(2), 1981, 229-232.
- 67.Indu BK, Aruna K, Evaluation of the liver protective potential of piperine, an active principle of long pepper, *Planta Med* 59(5), 1993, 413-417.
- 68. Dorman HJ, Deans SG. Antimicrobial agents from plants: antibacterial activity of volatile plant oils. J Appl Microbiol 2000;88:308-16.
- 69. Somani RS, Singhai AK. Hypoglycemic and antidiabetic activities of seeds of *Myristica fragrans* in normoglycemic and alloxan-induced diabetic rats. Asian J Exp Sci 2008;22:95-102.
- 70. Sharma A, Mathur R, Dixit VP. Prevention of hypercholesterolemia and atherosclerosis in rabbits after supplementation of *Myristica fragrans* seed extract. Indian J Physiol Pharmacol 1995;39:407-10.
- 71. Singh A, Rao AR. Modulatory effect of areca nut on theaction of mace (*Myristica fragrans*, Houtt) on the hepatic detoxification system in mice. Food Chem Toxicol 1993;31:517-21.
- 72. Dewi K, Widyarto B, Erawijantari PP, Widowati W. *In vitro* study of *Myristica fragrans* seed (Nutmeg)

ethanolic extract and quercetin compound as an antiinflammatory agent. Int J Res Med Sci 2015;3:2303-10.

73. Chirathaworn C, Kongcharoensuntorn W, Dechdoungchan T, Lowanitchapat A, Sa-nguanmoo P, Poovorawan Y. *Myristica fragrans* houtt. methanolic extract induces apoptosis in a human leukemia cell line through SIRT1 mRNA downregulation. J Med Assoc Thai 2007;90:2422-8.

74. Parle M, Dhingra D, Kulkarni SK. Improvement of mouse memory by *Myristica fragrans* seeds. J Med Food 2004;7:157-61.

75. Tajuddin, Ahmad S, Latif A, Qasmi IA. Aphrodisiac activity of 50% ethanolic extracts of *Myristica fragrans* Houtt. (nutmeg) and *Syzygium aromaticum* (L) Merr. & Perry. (clove) in male mice: a comparative study. BMC Complementary Altern Med 2003;20:3-6.

76. Gupta S, Yadava JNS, Mehrotra R, Tandon JS. Antidiarrhoea1 profile of an extract and some fractions from *Myristica fragrans* (Nutmeg) on escherichia coli enterotoxin-inducedsecretory response. Pharm Biol 1992;30:179-83.

77. Dhingra D, Sharma A. Antidepressant-like activity of n-hexane extract of nutmeg (*Myristica fragrans*) seeds in mice. J Med Food 2006:9:84-9.

78. Assa JR, Widjanarko SB, Kusnadi J, Berhimpon S. Antioxidant potential of flesh seed and mace of Nutmeg (*Myristica fragrans* Houtt). Int J ChemTech Res 2014;6:2460-8.

79. Jung WC, Jang YS, Hieu TT, Lee CK, Ahn YJ. Toxicity of *Myristica fagrans* seed compounds against *Blatella germanica* (Dictyoptera: Blattellidae). JMed Entomol 2007;44:524-9.

80. Kim, J.; Jayaprakasha, G.K.; Uckoo, R.M.; Patil, B.S. Evaluation of chemopreventive and cytotoxic effect of lemon seed extracts on human breast cancer (MCF-7) cells. *Food Chem. Toxicol.* **2012**, *50*, 423–430. [CrossRef] [PubMed] 81. Xavier, S.M.; Barbosa, C.O.; Barros, D.O.; Silva, R.F.; Oliveira, A.A.; Freitas, R.M. Vitamin C antioxidant effects in hippocampus of adult

Wistar rats after seizures and status epilepticus induced by pilocarpine. *Neurosci. Lett.* **2007**, *420*, 76–79. [CrossRef]

82. Parhiz, H.; Roohbakhsh, A.; Soltani, F.; Rezaee, R.; Iranshahi, M. Antioxidant and anti-inflammatory properties of the citrus flavonoids hesperidin and hesperetin: An updated review of their molecular mechanisms and experimental models. *Phyther. Res.* **2015**, *29*, 323–331. [CrossRef] [PubMed]

83.Otang, W.M.; Afolayan, A.J. Antimicrobial and antioxidant efficacy of *Citrus limon* L. peel extracts used for skin diseases by Xhosa tribe of Amathole District, Eastern Cape, South Africa. *S. Afr. J. Bot.* **2016**, *102*, 46–49. [CrossRef]

84.Otang, W.M.; Afolayan, A.J. Antimicrobial and antioxidant efficacy of *Citrus limon* L. peel extracts used for skin diseases by Xhosa tribe of Amathole District, Eastern Cape, South Africa. *S. Afr. J. Bot.* **2016**, *102*, 46–49. [CrossRef]

85.Bhatia, H.; Pal Sharma, Y.; Manhas, R.K.; Kumar, K. Traditional phytoremedies for the treatment of menstrual disorders in district Udhampur, J&K, India. *J. Ethnopharmacol.* **2015**, *160*, 202–210.

86. Tsujiyama, I.; Mubassara, S.; Aoshima, H.; Hossain, S.J. Anti-histamine release and anti-inflammatory activities of aqueous extracts of citrus fruits peels. *Orient. Pharm. Exp. Med.* **2013**, *13*, 175–180. [CrossRef

87. Bhavsar, S.K.; Joshi, P.; Shah, M.B.; Santani, D.D. Investigation into hepatoprotective activity of *Citrus limon. Pharm. Biol.* **2007**, *45*, 303–311. [CrossRef]

88. Mohanapriya, M.; Ramaswamy, L.; Rajendran, R. Health and medicinal properties of lemon (*Citrus limonum*). *Int. J. Ayurvedic Herb. Med.* **2013**, *1*, 1095–1100.

89.Kim, M.J.; Hwang, J.H.; Ko, H.J.; Na, H.B.; Kim, J.H. Lemon detox diet reduced body fat, insulin resistance, and serum hs-CRP level without hematological changes in overweight

Korean women. *Nutr. Res.* **2015**, *35*, 409–420. [CrossRef][PubMed]

90.Kato, Y.; Domoto, T.; Hiramitsu, M.; Katagiri, T.; Sato, K.; Miyake, Y.; Aoi, S.; Ishihara, K.; Ikeda, H.; Umei, N.; et al. Effect on blood pressure of daily lemon ingestion and walking. *J. Nutr. Metab.* **2014**, *2014*. [CrossRef] [PubMed]

91.Riaz, A.; Khan, R.A.; Algahtani, H.A. Memory boosting effect of *Citrus limon*, pomegranate and their combinations. *Pak. J. Pharm. Sci.* **2014**, *27*, 1837–1840.

92.Nie, Y.C.; Wu, H.; Li, P.B.; Luo, Y.L.; Long, K.; Xie, L.M.; Shen, J.G.; Su, W.W. Anti-inflammatory effects of naringin in chronic pulmonary

neutrophilic inflammation in cigarette smoke-exposed rats. *J. Med. Food* **2012**, *15*, 894–900. [CrossRef]

93. Shen, C.L.; Smith, B.J.; Lo, D.F.; Chyu, M.C.; Dunn, D.M.; Chen, C.H.; Kwun, I.S. Dietary polyphenols and mechanisms of osteoarthritis. *J. Nutr. Biochem.* **2012**, *23*, 1367–1377. [CrossRef]

94.Bhatia, H.; Pal Sharma, Y.; Manhas, .K.; Kumar, K. Traditional phytoremedies for the treatment of menstrual disorders in district Udhampur, J&K, India. *J. Ethnopharmacol.* **2015**, *160*, 202–210.

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