

International Journal of Dentistry and Oral Science (IJDOS) ISSN: 2377-8075

Health Benefits Of Annona Muricata - A Review

Review Article

Edala Venkata Gana Karthik¹, Vishnu Priya .V², Gayathri .R³, Dhanraj Ganapathy^{4*}

¹ Graduate Student, Department of Prosthodontics, Saveetha Dental college and Hospitals, Saveetha Institute of medical and Technical Sciences, Saveetha University, Chennai, India.

² Professor and Head of Department of Biochemistry, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Saveetha University, 162, Poonamallee High Road, Chennai - 600077, Tamil Nadu, India.

³ Associate Professor ,Department of Biochemistry, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Saveetha University, 162, Poonamallee High Road, Chennai - 600077, Tamil Nadu, India.

⁴ Professor and Head of Department of Prosthodontics, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Saveetha University, 162, Poonamallee High Road, Chennai - 600077, Tamil Nadu, India.

Abstract

Natural derivatives from plants have proven to be useful in maintaining men's health. In recent past years, phytochemicals derived from plants have been a backbone for pharmaceutical discoveries. *Annonamuricata L.*, commonly known as soursop, graviola, guanabana, paw-paw and sirsak, is a member of the Annonaceae family. *Annonamuricata* is a coveted tropical tree, and a wealth of phytochemical investigations have been conducted for this fruit plant. In addition to being an important source for the food industry and an indigenous medicinal plant, *Annonamuricata* is proven to possess numerous biological activities such as anti-arthritic activity, anti-cancer activity, antimicrobial activity, anticonvulsant activity, Anti-diabetic and hypolipidemic activity, anti-inflammatory and anti-nociceptive activity, antioxidant activity, anti-hypertensive activity, anti-parasitic activity, molluscicidal activity, wound-healing activity. This review enlightens various biological activities of *Annona Muricata*.

Keywords: AnnonaMuricata; Graviola; Biological Activities; Anti-Cancer; Anti-Oxidant.

Introduction

Natural derivatives from plants have proven to be useful in maintaining human health. In recent years, phytochemicals derived from plants have been a backbone for pharmaceutical discoveries.[1] Hence the biologically active ingredients of plants have to be inspected thoroughly concerning their potential role in nature. Among the lot, AnnonaMuricata is one of the most extensively used traditional plants.[2, 3]

Annonamuricata L., commonly known as soursop, graviola, guanabana, paw-paw and sirsak, is a member of the Annonaceae family comprising approximately 130 genera and 2300 species.[4, 5]. *Annonamuricata* being native to the warmest tropical areas in South and North America is now widely distributed throughout tropical and subtropical parts of the world, including Malaysia and India. Annonamuricata is an evergreen, terrestrial, erect tree reaching 5–8 m in height and features an open, roundish canopy with large, glossy, dark green leaves. The edible fruits of the tree are large, heart-shaped and green in colour, and the diameter varies between 15 and 20 cm [4, 6, 7].

Fruits of Annonamuricata are taken internally to cure worms, fever, to increase mother's milk after child birth, and as an astringent for diarrhoea and dysentery; unripe fruit mixed with olive oil was used for neuralgia, rheumatism, and arthritic pain [8]. The leaves are used in traditional medicine to treat headaches, hypertension, cough, and asthma and used as antispasmodic, sedative, and nervine for heart condition [9]. Annonaceous acetogenins, from *Annonamuricata L.*, were found to be a promising new antitumor and anticancer agent in numerous in vitro studies [4]. These acetogenins were demonstrated to be selectively toxic against vari-

Dhanraj Ganapathy,

Professor and Head of Department of Prosthodontics, Saveetha Dental College and Hospitals, Saveetha Institute of Medical and Technical Sciences, Saveetha University, 162, Poonamallee High Road, Chennai - 600077, Tamil Nadu, India. Tel: 9841504523

E-mail: dhanrajmganapathy@yahoo.co.in

Received: May 28, 2021 Accepted: June 16, 2021 Published: July 01, 2021

Citation: Edala Venkata Gana Karthik, Vishnu Priya .V, Gayathri .R, Dhanraj Ganapathy. Health Benefits Of Annona Muricata - A Review. Int J Dentistry Oral Sci. 2021;8(7):2965-2967. doi: http://dx.doi.org/10.19070/2377-8075-21000602

Copyright: Dhanraj Ganapathy[©]2021. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution and reproduction in any medium, provided the original author and source are credited.

^{*}Corresponding Author:

ous types of the cancerous cells without harming healthy cells. Various other plants from this family have also been reported for their cytotoxic potential [10].

The leaves, root and stem barks of Annonamuricata afforded seven isoquinoline alkaloids: reticuline, coclaurine, coreximine, atherosperminine, stepharine, anomurine and anomuricine [11] The essential oil of the fresh fruit pulp of A. muricata yielded 2-hexenoic acid methyl ester (23.9%), 2-hexenoic acid ethyl ester (8.6%), 2-octenoic acid methyl ester (5.4%), 2-butenoic acid methyl ester (8.6%), 2-octenoic acid methyl ester (5.4%), 2-butenoic acid methyl ester (2.4%), β -caryophyllene (12.7%), 1,8-cineole (9.9%), linalool (7.8%), α -terpineol (2.8%), linalyl propionate (2.2%), and calarene (2.2%). [12, 13] The seeds of Annonamuricatanfforded annomuricatin A.[14] Our research experience has prompted us in pursuing this review. [15-24] This article would help in understanding all the biological health benefits of Annonamuricata.

Biological Activities

There are several known biological activities likeanti-depressive [25, 26], toxicity against prostate PC-3 cancer cells [27], toxicity against pancreatic MIA PaCa-2 and colon HT-29 cancer cells [27, 28], toxicity against brine shrimp and different cancer cells [29], toxicity against lung A549, toxicity against human hepatoma cells, neurotoxic, breast MCF7, toxicity against oral KB cancer cells and brine shrimp larva, antileishmanial, molluscicidal and various others.[30] Several significant activities among these are:

Antioxidant Activity

The precursor for oxidative stress and which subsequently catalyses metabolic deficiency and cellular death through physiological and biochemical lesions are the immoderate generation of intracellular reactive oxygen species (ROS) [31] Identification of these antioxidants from natural products is very important as they play a crucial role in nullifying the destructive effects of ROS [31, 32]. Certain tests such as DRSA, FRAP and HRSA methanolic and aqueous leaf extracts of AnnonaMuricata leaves revealed the antioxidant properties. These extracts also help in protecting DNA against H2O2 (hydrogen peroxide) induced toxicity. Among the other species of AnnoaceaeFamily, Annonamuricata was found to have a stronger antioxidant activity through different models such as ABTS, nitric oxide and hydroxyl radicals [33, 34]. The seeds and leaves of the plant are reported to possess enzymatic antioxidants, including catalase and superoxide dismutase, and non-enzymatic antioxidants, including vitamin C and E [1]. The stem bark also showed antioxidant activity through DPPH test [35]. Thus, these above findings strongly suggest the potential use of Annonamuricataas a natural source of antioxidants.

Antihypertensive Activity

To evaluate the anti-hypertensive properties of Annonamuricata, Sprague-Dawley rats were administered with aqueous leaf extract(9.17-48.5mg/kg) [36]. The results have shown that the leaf extracts have quite significantly decreased the blood pressure in these rats without affecting the heart rate. This effect was suggested to be induced through peripheral mechanisms involving the antagonism of Ca2+ [37].

Gastroprotective Activity

To evaluate the gastro-protective of AnnonaMuricataleaves were examined against ethanol-induced gastric injury [37, 38] Oral administration of the ethyl acetate extract (200 and 400 mg/kg) have showed a significant activity in anti-ulcer activity, which had protective effects against gastric wall mucosal damage [37-39]. These findings strongly suggest the Gastroprotective activity of Annonamuricata.

Anti-cancer Activity

Cancer mortality rates have increased in the developed countries throughout this century and already as the cause death in some Western countries. Extracts of AnnonaMuricata have proven to show significant anti-cancer activity.[40] Ethyl acetate extract of leaves have proven to supress lung A549 cancer cells, colon HT-29 and HCT-116 [41] cancer cells by mitochondrial-mediated apoptosis, cell cycle arrest at G1 phase. Water extract of the leaves had an effect on the rats prostate cancer by reduction in the prostate size. Ethanolic extract of leaves hada effect on breast tissues of mice prevention of DMBA by induced DNA damage [42], DMBA/croton oil induced mice skin papillomagenesis by suppression of tumor initiation and promotion, DMH induced colon cancer by reduction of ACF formation, K562 chronic myeloid lukemia cells by induction of apoptosis [42, 43] Leaves boiled in water had effect on metastatic breast cancer by stabilization of disease. Ethyl acetate of the leaves has also proven to have effects on azoxymethane induced colon cancer by reduction in ACF formation These findings strongly suggest the Anti-cancer activity of Annonamuricata.^[44]

Conclusion

Annonamuricatais a coveted tropical tree, and a wealth of phytochemical investigations have been conducted for this fruit plant. In addition to being an important source for the food industry and an indigenous medicinal plant, Annonamuricatais proven to possess numerous biological activities such as anti-arthritic activity, anti-cancer activity, anti-microbial activity, anti-convulsant activity, Anti-diabetic and hypolipidemic activity, anti-inflammatory and anti-nociceptive activity, anti-oxidant activity, anti-hypertensive activity, anti-parasitic activity, anti-plasmodial activity, hepatoprotective and bilirubin-lowwering activity, insecticidal activity, gastroprotective activity, molluscicidal activity, wound-healing activity[30]. Previously, this plant trails were neglected. This article is thus hoped to provide enlightment and motivation to other researcher' in conducting clinical trails and developing new pharmaceutical and agricultural agents.

References

- Kumaran T. A Review of Recent Studies on The Phytochemical and Pharmacological Activity of Annona Muricata. Forestry & Agriculture Review. 2021 Jan 10;2(1):1-5.
- [2]. Prasad SK, Varsha V, Devananda D. Anti-cancer properties of Annona muricata (L.): A Review. Med. Plants - Int. J. Phytomed. Relat. Ind. 2019;11(2):123-34.
- [3]. Banerjee A, Das D, Maji BK, Mukherjee S. Anticancer effects of Annona muricata with its acetogenins as bioactive compound. CHIJ. 2018;2(1):1-6.
- [4]. Viegas A, dos Santos T, Costa L. Phytochemistry and anticancer potential of graviola (Annona muricata) [Internet]. Proceedings of MOL2NET 2018,

International Conference on Multidisciplinary Sciences, 4th edition. 2018. Available from: http://dx.doi.org/10.3390/mol2net-04-05527

- [5]. Velázquez-Aponte RA, Cassé C. ANTIPROLIFERATIVE PROPERTIES OF ETHANOLIC AND AQUEOUS GRAVIOLA LEAF EXTRACTS ON TONGUE SQUAMOUS CELL CARCINOMA CELL LINE-25. J Med Rev Case Rep. 2020 Aug 21;3(08).
- [6]. Adewole SO, Caxton-Martins EA. Morphological changes and hypoglycemic effects of Annona muricata linn.(annonaceae) leaf aqueous extract on pancreatic β-cells of streptozotocin-treated diabetic rats. Afr J Biomed Res. 2009;9(3).
- [7]. Chavez ML. Medicinal Plants of the World. Chemical Constituents, Traditional and Modern Medicinal Uses. Am. J. Pharm. Educ. 1999 Jul 1;63(2):4.
- [8]. George VC, Kumar DR, Rajkumar V, Suresh PK, Kumar RA. Quantitative assessment of the relative antineoplastic potential of the n-butanolic leaf extract of Annona muricata Linn. in normal and immortalized human cell lines. Asian Pac J Cancer Prev. 2012;13(2):699-704.Pubmed PMID: 22524847.
- [9]. Pharmacognostic and Safety Assessment of Hydro-Methanol Leaf Extract of Annona muricata Linn (Annonacea) in Male Wistar Rats. Trop. J. Nat. Prod. Res. 2021;4:1182–9. Available from: http://dx.doi.org/10.26538/ tjnpr/v4i12.26
- [10]. Hasmila I, Natsir H, Soekamto NH. Phytochemical analysis and antioxidant activity of soursop leaf extract (Annona muricata Linn). J. Phys. Conf. Ser. 2019 Oct 1; 1341(3): 032027.
- [11]. R S, SoumyaS R, Bishop Moore College, 110. M 690, Ramachandran A, DineshRaj R, et al. MOSQUITO LARVICIDAL AND CYTOTOXIC STUDIES ON ANNONA RETICULATA L. Int. j. adv. res. 2017;5:414–9. Available from: http://dx.doi.org/10.21474/ijar01/5545
- [12]. Makabe H. ChemInform Abstract: Synthesis of Annonaceous Acetogenins from Muricatacin. ChemInform. 2008;39. Available from: http://dx.doi. org/10.1002/chin.200848264
- [13]. Paige MA. Modular synthesis of annonaceous acetogenins and their activity against H-116 human solid colon tumor cells. University of Virginia; 2003.
- [14]. Makabe H. Synthesis of annonaceous acetogenins from muricatacin. BiosciBiotechnolBiochem. 2007 Oct 23;71(10):2367-74.
- [15]. Hemalatha R, Dhanraj S. Disinfection of Dental Impression- A Current Overview. Cuddalore. 2016 Jul;8(7):661–4.
- [16]. Ramya G, Pandurangan K, Ganapathy D. Correlation between anterior crowding and bruxism-related parafunctional habits. Drug invent. today. 2019 Oct 15;12(10).
- [17]. Anjum AS, Ganapathy D, Kumar K. Knowledge of the awareness of dentists on the management of burn injuries on the face. Drug invent. today. 2019 Sep 1;11(9).
- [18]. Inchara R, Ganapathy D, Kumar PK. Preference of antibiotics in pediatric dentistry. Drug invent. today. 2019 Jun 15;11:1495-8.
- [19]. Philip JM, Ganapathy DM, Ariga P. Comparative evaluation of tensile bond strength of a polyvinyl acetate-based resilient liner following various denture base surface pre-treatment methods and immersion in artificial salivary medium: An in vitro study. Contemp Clin Dent. 2012 Jul;3(3):298-301. Pubmed PMID: 23293485.
- [20]. Gupta A, Dhanraj M, Sivagami G. Implant surface modification: review of literature. The Internet J Dent Sci. 2009;7(1):10.
- [21]. Indhulekha V, Ganapathy D, Jain AR. Knowledge and awareness on biomedical waste management among students of four dental colleges in Chennai, India. Drug invent. today. 2018 Dec 1;10(12):32-41.
- [22]. Mohamed Usman JA, Ayappan A, Ganapathy D, Nasir NN. Oromaxillary prosthetic rehabilitation of a maxillectomy patient using a magnet retained two-piece hollow bulb definitive obturator; a clinical report. Case Rep Dent. 2013;2013:190180.Pubmed PMID: 23533823.
- [23]. Ganapathy DM, Joseph S, Ariga P, Selvaraj A. Evaluation of the influence of blood glucose level on oral candidal colonization in complete denture wearers with Type-II Diabetes Mellitus: An in vivo Study. Dent Res J (Isfahan). 2013 Jan;10(1):87-92.Pubmed PMID: 23878569.
- [24]. Menon A, Ganapathy DM, Mallikarjuna AV. Factors that influence the colour stability of composite resins. Drug invent. today. 2019 Mar 1;11(3).
- [25]. Hasrat JA, De Bruyne T, De Backer JP, Vauquelin G, Vlietinck AJ. Isoquinoline derivatives isolated from the fruit of Annona muricata as 5-HTergic 5-HT1A receptor agonists in rats: unexploited antidepressive (lead) products. J Pharm Pharmacol. 1997 Nov;49(11):1145-9.Pubmed PMID: 9401954.
- [26]. Hasrat JA, Pieters L, De Backer JP, Vauquelin G, Vlietinck AJ. Screening

of medicinal plants from Suriname for 5-HT(1A) ligands: Bioactive isoquinoline alkaloids from the fruit of Annona muricata. Phytomedicine. 1997 Jun;4(2):133-40.Pubmed PMID: 23195401.

- [27]. Sun S, Liu J, Kadouh H, Sun X, Zhou K. Three new anti-proliferative Annonaceous acetogenins with mono-tetrahydrofuran ring from graviola fruit (Annona muricata). Bioorg Med ChemLett. 2014 Jun 15;24(12):2773-6.
- [28]. Wu FE, Gu ZM, Zeng L, Zhao GX, Zhang Y, McLaughlin JL, et al. Two new cytotoxic monotetrahydrofuran Annonaceous acetogenins, annomuricins A and B, from the leaves of Annona muricata. J Nat Prod. 1995 Jun;58(6):830-6.Pubmed PMID: 7673926.
- [29]. Zeng L, Wu FE, McLaughlin JL. Annohexocin, a novel mono-THF acetogenin with six hydroxyls, from Annona muricata (Annonaceae). Bioorganic Med. Chem. Lett. 1995 Aug 17;5(16):1865-8.
- [30]. Moghadamtousi SZ, Fadaeinasab M, Nikzad S, Mohan G, Ali HM, Kadir HA. Annona muricata (Annonaceae): a review of its traditional uses, isolated acetogenins and biological activities. Int J Mol Sci. 2015 Jul;16(7):15625-58.
- [31]. George VC, Kumar DR, Suresh PK, Kumar RA. Antioxidant, DNA protective efficacy and HPLC analysis of Annona muricata (soursop) extracts. J Food Sci Technol. 2015 Apr;52(4):2328-35.Pubmed PMID: 25829616.
- [32]. Chen W, Weng YM, Tseng CY. Antioxidative and antimutagenic activities of healthy herbal drinks from Chinese medicinal herbs. Am J Chin Med. 2003;31(4):523-32.Pubmed PMID: 14587875.
- [33]. P N, Nalini P. Phytochemical Screening and Antioxidant Potential in the Leaf Extract of Annona Muricata. IJRASET. 2018;6:2744–7. Available from: http://dx.doi.org/10.22214/ijraset.2018.4459
- [34]. Purnamasari F, Yulianty R, Latief S. Effectiveness of Soursop Leaf Extract (Annona muricata l.) on IL-6 Levels in Mammary Sprague dawley Female Rats Induced by Staphylococcus aureus. Unnes j. public health. 2020 Jan 31;9(1):56-63.
- [35]. Agunloye OO, Onifade AK. Annona muricata: Comparative Assessment of the Antibacterial Activities of the Leaf and Stem Extracts against Multiple Antibiotic Resistant Clinical Isolates. J Adv Microbiol . 2020 Jun 2:12-21.
- [36]. Nwokocha CR, Owu DU, Gordon A, Thaxter K, McCalla G, Ozolua RI, et al. Possible mechanisms of action of the hypotensive effect of Annona muricata (soursop) in normotensive Sprague-Dawley rats. Pharm Biol. 2012 Nov;50(11):1436-41.Pubmed PMID: 22950673.
- [37]. Kuswinarti K, Savira K, Rudiman R. The Analgesic Effect of Ethanol Extract Soursop (Annona muricata L.) Leaves in Wistar Rats. Althea Med. J. 2018 Dec 28;5(4):196-200.
- [38]. Hamid RA, Foong CP, Ahmad Z, Hussain MK. Antinociceptive and antiulcerogenic activities of the ethanolic extract of Annona muricata leaf. Revista Brasileira de Farmacognosia. 2012 Jun;22(3):630-41.
- [39]. de Sousa OV, Vieira GD, de Jesus R G de Pinho J, Yamamoto CH, Alves MS. Antinociceptive and anti-inflammatory activities of the ethanol extract of Annona muricata L. leaves in animal models. Int J Mol Sci. 2010 May 6;11(5):2067-78.Pubmed PMID: 20559502.
- [40]. Moghadamtousi SZ, Kadir HA, Paydar M, Rouhollahi E, Karimian H. Annona muricata leaves induced apoptosis in A549 cells through mitochondrial-mediated pathway and involvement of NF-κB. BMC Complement Altern Med. 2014 Aug 15;14:299.Pubmed PMID: 25127718.
- [41]. Zorofchian Moghadamtousi S, Karimian H, Rouhollahi E, Paydar M, Fadaeinasab M, Abdul Kadir H. Annona muricata leaves induce G₁ cell cycle arrest and apoptosis through mitochondria-mediated pathway in human HCT-116 and HT-29 colon cancer cells. J Ethnopharmacol. 2014 Oct 28;156:277-89.Pubmed PMID: 25195082.
- [42]. Nalini P, Durairaj BR. Antitumor Potential of hydroethanolic extract of Annona muricata leaves against dalton's lymphoma ascites-induced tumor in mice. Asian J Pharm Clin Res. 2018;11(3):364-7.
- [43]. Hamizah S, Roslida AH, Fezah O, Tan KL, Tor YS, Tan CI. Chemopreventive potential of Annona muricata L leaves on chemically-induced skin papillomagenesis in mice. Asian Pac J Cancer Prev. 2012;13(6):2533-9.
- [44]. Zorofchian Moghadamtousi S, Rouhollahi E, Karimian H, Fadaeinasab M, Firoozinia M, Ameen Abdulla M, et al. The chemopotential effect of Annona muricata leaves against azoxymethane-induced colonic aberrant crypt foci in rats and the apoptotic effect of Acetogenin Annomuricin E in HT-29 cells: a bioassay-guided approach. PLoS One. 2015 Apr 10;10(4):e0122288. Pubmed PMID: 25860620.