



# Natural Products Isolated from Various Parts of Mangosteen (*Garcinia mangostana* L.) as Therapeutic Agent: A Review

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## ABSTRACT

<sup>12</sup>his review provides a comprehensive analysis of the therapeutic potential of natural products derived from mangosteen (<sup>13</sup>*Garcinia mangostana* L.). Mangosteen, a tropical fruit native to Southeast Asia, has long been valued for its medicinal properties. The review focuses on the isolation and characterization of bioactive compounds from different parts of the plant, including the fruit, rind, and leaves. The bioactive compounds identified in mangosteen exhibit a diverse range of therapeutic activities, making them promising candidates for the development of various therapeutic agents. These compounds include xanthenes, flavonoids, tannins, and phenolic acids, among others. Xanthenes, in particular, have shown potent antioxidant, anti-inflammatory, anticancer, antimicrobial, and neuroprotective properties. Additionally, they have been reported to have beneficial effects on various chronic diseases such as cardiovascular disorders, diabetes, and obesity. This review highlights the mechanisms of action underlying the therapeutic effects of mangosteen-derived compounds.

**Keywords:** *drug discovery, herbs, secondary metabolite, therapy*

## 1. INTRODUCTION

Mangosteen (*Garcinia mangostana* L.) is a tropical fruit native to Southeast Asia that has gained considerable attention in recent years due to its potential therapeutic properties. It has been traditionally used in folk medicine to treat various ailments such as diarrhea, dysentery, wounds, and skin infections [1]. This fruit is known for its unique flavor and nutritional value, but it is the bioactive compounds found in mangosteen that have gained significant scientific interest [2].

Extensive research has been conducted to isolate and characterize the bioactive compounds present in different parts of the mangosteen plant, including the fruit, rind, and leaves. These studies have revealed a wide array of phytochemicals, such as xanthenes, flavonoids, tannins, and phenolic acids, which possess various pharmacological activities [3]. Among these, xanthenes have been identified as the major class of bioactive compounds in mangosteen, with over 60 different types identified to date [4].

The therapeutic potential of mangosteen-derived natural products is extensive and covers a broad spectrum of health conditions. These bioactive compounds have been found to exhibit antioxidant, anti-inflammatory, anticancer, antimicrobial, and neuroprotective activities [5,6]. Additionally, they have demonstrated beneficial effects on chronic diseases, including cardiovascular disorders, diabetes, and obesity [7,8]. Understanding the mechanisms of action and exploring the potential applications of these natural products could lead to the development of novel therapeutic agents for a wide range of diseases.

## 2. AN OVERVIEW OF MANGOSTEEN (*GARCINIA MANGOSTANA* L.)

Mangosteen is a tropical fruit tree belonging to the family Clusiaceae. It is native to Southeast Asia and is highly regarded for its delicious flavor and potential health benefits. The fruit is small, round, and purple in color, with a thick rind and juicy white pulp. Mangosteen is often referred to as the "queen of fruits" due to its unique taste and nutritional value [9].

One of the most notable features of mangosteen is its rich content of bioactive compounds. Xanthenes, a class of polyphenolic compounds, are the primary bioactive constituents found in mangosteen. These compounds have been extensively studied for their potential health-promoting effects, including antioxidant, anti-inflammatory, antimicrobial, and anticancer properties [10]. They have been shown to scavenge free radicals, reduce inflammation, inhibit the growth of bacteria and fungi, and exhibit toxicity effects on cancer cells.

In addition to xanthenes, mangosteen contains other important phytochemicals such as flavonoids, tannins, and phenolic acids. These compounds contribute to the fruit's antioxidant activity and may play a role in its potential therapeutic benefits. Mangosteen has been traditionally used in Southeast Asian folk medicine to treat various ailments, including skin infections, diarrhea, and dysentery [11].

Research on mangosteen and its bioactive compounds has gained momentum in recent years, leading to a better understanding of their potential health applications. Studies have suggested that mangosteen may have a positive impact on cardiovascular health by reducing oxidative stress, improving lipid profiles, and exerting anti-inflammatory effects [8,9]. It also shows promise in managing diabetes by regulating blood glucose levels and enhancing insulin sensitivity [3-5]. Additionally, mangosteen extracts have exhibited neuroprotective effects, which could have implications for the prevention and treatment of neurodegenerative diseases [10].

While the therapeutic potential of mangosteen and its bioactive compounds is intriguing, further research is needed to fully understand their mechanisms of action and optimize their application in various health conditions. Nevertheless, mangosteen continues to attract attention as a promising natural ingredient with potential health benefits, and its inclusion in functional foods, dietary supplements, and pharmaceutical products is being explored.

## 3. PHARMACOLOGICAL ACTIVITIES OF *GARCINIA MANGOSTANA* L.

The bioactive compounds of mangosteen fruit demonstrated various pharmacological properties, including antioxidant, anti-inflammatory, anticancer, antimicrobial, and neuroprotective activities. The main component in mangosteen fruit is xanthenes with 68 various xanthenes. Whilst mangosteen peel contained phenol and xanthone. Moreover, the extract of hot water derived from mangosteen peel demonstrated epicatechin as the main component. Furthermore, based on Suhartati et al (2018), mangosteen leaves demonstrated four active compounds, such as tannins, flavonoids, alkaloids, and saponins [12-14].

In general, the xanthone compound is abundant in mangosteen. This xanthone demonstrated protection on cells and tissues from oxidative damage, reducing the risk of chronic diseases associated with oxidative stress, such as cardiovascular diseases and neurodegenerative disorders. Furthermore, mangosteen exhibits significant anti-inflammatory effects. The bioactive compounds present in mangosteen can inhibit the production of pro-inflammatory mediators, thereby suppressing inflammation. This anti-inflammatory activity has implications for the management of inflammatory

diseases, such as arthritis and inflammatory bowel disease [15].

Recent studies have highlighted the anticancer potential of mangosteen. The xanthenes and other bioactive compounds in mangosteen have demonstrated cytotoxic effects on various cancer cell lines. They can induce cell cycle arrest, promote apoptosis (programmed cell death), and inhibit the growth and spread of cancer cells [6-8]. Mangosteen has been found to possess antimicrobial properties as well. Its bioactive compounds have shown inhibitory effects against bacteria, fungi, and even some drug-resistant strains. This antimicrobial activity suggests the potential use of mangosteen-derived compounds as natural alternatives for the treatment of infections [7]. Additionally, mangosteen exhibits neuroprotective effects, which are attributed to its ability to modulate neuronal signaling pathways, reduce neuroinflammation, and enhance antioxidant defenses. These properties make mangosteen a promising candidate for the prevention and treatment of neurodegenerative diseases, such as Alzheimer's and Parkinson's disease [15].

In brief, the pharmacological activities of mangosteen and its bioactive compounds demonstrate their potential in various health conditions. From their antioxidant and anti-inflammatory effects to their anticancer, antimicrobial, and neuroprotective properties, mangosteen exhibits a wide range of pharmacological activities that warrant further exploration and clinical research for their therapeutic applications [17-27].

#### 4. CONCLUSION

In conclusion, the review highlights the significant therapeutic potential of natural products derived from mangosteen. The diverse array of bioactive compounds, particularly xanthenes, exhibit various pharmacological activities, including antioxidant, anti-inflammatory, anticancer, antimicrobial, and neuroprotective effects. These compounds have demonstrated promising results in the treatment of chronic diseases such as cardiovascular disorders, diabetes, obesity, and neurodegenerative disorders. The synergistic interactions among the different compounds in mangosteen extract further enhance their therapeutic efficacy. Despite the promising findings, further research and clinical studies are needed to fully understand the mechanisms of action, optimize their application, and explore their potential as therapeutic agents in various diseases. Overall, the review emphasizes the need for continued investigation and utilization of these natural products from mangosteen to harness their full therapeutic potential.

#### AUTHORS' CONTRIBUTIONS

ANMA and YA participated in the writing and reviewing draft of manuscript. AAAM, VDK, BM, and

MKJK contributed in the design of the study. DDRT and IR carried out the data collection. VJ, MR, TP, and RZ support the data gathering. All authors read and approved the final manuscript.

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#### REFERENCES

- [1] A.F. Ahmed, G.M. Sulaiman, A.K.M. Omar, A review on phytochemical constituents and pharmacological activities of mangosteen (*Garcinia mangostana* L.), *Research Journal of Pharmacy and Technology*, 2019, pp. 5073-5079
- [2] A.N.M. Ansori, A. Fadholly, S. Hayaza, R.J.K. Susilo, B. Inayatillah, D. Winarni, S.A. Husen, A review on medicinal properties of mangosteen (*Garcinia mangostana* L.), *Research Journal of Pharmacy and Technology*, 2020, pp. 974-982. DOI: 10.5958/0974-360X.2020.00182.1
- [3] Y.W. Chin, M.J. Balunas, H.B. Chai, A.D. Kinghorn, Drug discovery from natural sources, *The AAPS Journal*, 2008, pp. E239-E253. DOI: <https://doi.org/10.1007/BF02854894>
- [4] R.S.M. Labban, H.A. Alfawaz, M. Amina, R. S. Bhat, W. M. Hassan, A. El-Ansary, Synergism between Extracts of *Garcinia mangostana* Pericarp and *Curcuma* in Ameliorating Altered Brain Neurotransmitters, Systemic Inflammation, and Leptin Levels in High-Fat Diet-Induced Obesity in Male Wistar Albino Rats, *Nutrients*, 2022, pp. 4630. DOI: <https://doi.org/10.3390/nu14214630>
- [5] D. Obolskiy, I. Pischel, B. Feistel, N. Glotov, M. Heinrich, *Garcinia mangostana* L.: A phytochemical and pharmacological review, *Phytotherapy Research*, 2009, pp. 1047-1065. DOI: <https://doi.org/10.1002/ptr.2730>
- [6] J. Pedraza-Chaverri, N. Cardenas-Rodríguez, M. Orozco-Ibarra, J.M. Pérez-Rojas, O. N. Medina-Campos, Review: medicinal properties of mangosteen (*Garcinia mangostana*), *Food and Chemical Toxicology*, 2008, pp. 3227-3239. DOI: <https://doi.org/10.1016/j.fct.2008.07.024>
- [7] M.M. Ashton, O.M. Dean, A.J. Walker, C.C. Bortolasci, C.H. Ng, M. Hopwood, B. H. Harvey, M. Möller, J.J. McGrath, W. Marx, A. Turner, S. Dodd, J.G. Scott, J.P. Khoo, K. Walder, J. Sarris, M. Berk, The Therapeutic Potential of Mangosteen Pericarp as an Adjunctive Therapy for Bipolar Disorder and Schizophrenia, *Frontiers in psychiatry*,

- 2019, pp. 115. DOI: <https://doi.org/10.3389/fpsyt.2019.00115>
- [8] J.S. Park, E.Y. Ahn, Y. Park, Asymmetric dumbbell-shaped silver nanoparticles and spherical gold nanoparticles green-synthesized by mangosteen (*Garcinia mangostana*) pericarp waste extracts, *International Journal of Nanomedicine*, 2017, pp. 6895–6908. DOI: <https://doi.org/10.2147/IJN.S14190>
- [9] G.A. Mohamed, A.M. Al-Abd, A.M. El-Halawany, H.M. Abdallah, S.R.M. Ibrahim, New xanthenes and cytotoxic constituents from *Garcinia mangostana* fruit hulls against human hepatocellular, breast, and colorectal cancer cell lines, *Journal of Ethnopharmacology*, 2017, pp.302–312. DOI: <https://doi.org/10.1016/j.jep.2017.01.030>
- [10] V. Yuvanatemiya, P. Srean, W.K. Klangbud, K. Venkatachalam, J. Wongs, T. Parametthanuwat, N. Charoenphun, A Review of the Influence of Various Extraction Techniques and the Biological Effects of the Xanthenes from Mangosteen (*Garcinia mangostana* L.) Pericarps, *Molecules*, 2022, pp. 8775. DOI: <https://doi.org/10.3390/molecules27248775>
- [11] C. Palakawong, P. Sophanodora, P. Toivonen, P. Delaquis, Optimized extraction and characterization of antimicrobial phenolic compounds from mangosteen (*Garcinia mangostana* L.) cultivation and processing waste, *Journal of the Science of Food and Agriculture*, 2013, pp. 3792–3800. DOI: <https://doi.org/10.1002/jsfa.6277>
- [12] A. Ghasemzadeh, H.Z.E. Jaafar, A. Baghdadi, A. Tayebi-Meigooni, Alpha-Mangostin-Rich Extracts from Mangosteen Pericarp: Optimization of Green Extraction Protocol and Evaluation of Biological Activity, *Molecules*, 2018, pp. 1852. DOI: <https://doi.org/10.3390/molecules23081852>
- [13] R. Defri, R. Hartati, T. Nadhifa, I. Fidrianny, Chemical compounds and pharmacological activities of mangosteen (*Garcinia mangostana* L.) - updated review, *Biointerface Research in Applied Chemistry*, 2022, pp. 2503-2516. DOI: <https://doi.org/10.33263/BRIAC122.25032516>
- [14] R. Suhartati, F. Apriyani, Khusnul, D.P. Virgiannti, M. Fathurohman, Antimicrobial activity test of mangosteen leaves ethanol extract (*Garcinia mangostana* Linn) against *Pseudomonas aeruginosa* bacteria, *ICCOMSET*, 2018, pp. 1-6, DOI: 10.1088/1742-6596/1179/1/012167
- [15] B.R. Albuquerque, M.I. Dias, J. Pinela, R. C. Calhelha, T. C. S. Pires, M. J. Alves, R. C. G. Corrêa, I. C. F. R. Ferreira, M. B. P. P. Oliveira, L. Barros, Insights into the Chemical Composition and In Vitro Bioactive Properties of Mangosteen (*Garcinia mangostana* L.) Pericarp, *Foods*, 2023, pp. 994. DOI: <https://doi.org/10.3390/foods12050994>
- [16] T.Y. Pramana, B. Wasita, V. Widyaningsih, R. Cilmiaty, S. Suroto, A. Mudigdo, B. Purwanto, The ethanol extract of *Garcinia mangostana* L peel reduces the isoniazid-induced liver damage in rats, *Bali Medical Journal*, 2021, pp. 156–159. DOI: <https://doi.org/10.15562/bmj.v10i1.2108>
- [17] R.S. Indharty, I. Japardi, A.M. Siahaan, S. Tandeau, Mangosteen extract reduce apoptosis via inhibition of oxidative process in rat model of traumatic brain injury, *Bali Medical Journal*, 2019, pp. 227–232. DOI: <https://doi.org/10.15562/bmj.v8i1.1153>
- [18] V. D. Kharisma, A.N.M. Ansori, Y. Antonius, I. Rosadi, A.A.A. Muradlo, V. Jakhmola, M. Rebezov, N. Maksimiuk, E. Kolesnik, P. Burkov, M. Derkho, P. Scherbakov, M.E. Ullah, T.H. Sucipto, H. Purnobasuki, Garcinoxanthenes from *Garcinia mangostana* L. tackle SARS-CoV-2 infection and cytokine storm pathway inhibition: A viroinformatics study, *J Pharm Pharmacogn Res*, 2023, pp. 743–756. DOI: [https://doi.org/10.56499/jppres23.1650\\_11.5.743](https://doi.org/10.56499/jppres23.1650_11.5.743)
- [19] D. Winarni, F. N. Husna, M. F. Syadzha, R. J. K. Susilo, S. Hayaza, A. N. M. Ansori, M. A. Alamsjah, M. N. G. Amin, P. A. C. Wulandari, P. Pudjiastuti, K. Awang, Topical Administration Effect of *Sargassum duplicatum* and *Garcinia mangostana* Extracts Combination on Open Wound Healing Process in Diabetic Mice, *Scientifica*, 2022, pp. 9700794. DOI: <https://doi.org/10.1155/2022/9700794>
- [20] A. N. M. Ansori, R. J. K. Susilo, S. Hayaza, D. Winarni, S. A. Husen, Renoprotection by *Garcinia mangostana* L. pericarp extract in streptozotocin-induced diabetic mice, *Iraqi Journal of Veterinary Sciences*, 2019, pp. 13-19. DOI: <https://doi.org/10.3389/ijvs.2019.125513.1035>
- [21] S.A. Husen, Salamun, A. N. M. Ansori, S. Hayaza, R. J. K. Susilo, D. Winarni, W. Darmanto, Renal protective effects of gamma-mangostin in streptozotocin-induced diabetic mice, *Indian Journal of Forensic Medicine and Toxicology*, 2020, pp. 1221-1226.
- [22] L. Vania, M. H. Widyananda, V. D. Kharisma, A. N. M. Ansori, S. W. Naw, N. Maksimiuk, M. Derkho, A. Denisenko, N. I. Sumantri, A. P. Nugraha, Anticancer activity prediction of *Garcinia mangostana* L. against HER2-positive breast cancer

- through inhibiting EGFR, HER2 and IGF1R protein: a bioinformatics study, *Biochemical and Cellular Archives*, 2021, pp. 3313-3321.
- [23] S. A. Husen, M. F. Syadzha, M. F. Setyawan, P. Pudjiastuti, A. N. M. Ansori, R. J. K. Susilo, S. Hayaza, D. Winarni, W. Darmanto, Evaluation of the combination of *Sargassum duplicatum*, *Sargassum ilicifolium*, *Abelmoschus esculentus*, and *Garcinia mangostana* extracts for open wound healing in diabetic mice, *Systematic Reviews in Pharmacy*, 2020, pp. 888-892. DOI: <https://doi.org/10.31838/srp.2020.9.129>
- [24] A. N. M. Ansori, V. D. Kharisma, A. A. Parikesit, F. A. Dian, R. T. Probojati, M. Rebezov, P. Scherbakov, P. Burkov, G. Zhdanova, A. Mikhalev, Y. Antonius, M. R. F. Pratama, N. I. Sumantri, T. H. Sucipto, R. Zainul, Bioactive Compounds from Mangosteen (*Garcinia mangostana* L.) as an Antiviral Agent via Dual Inhibitor Mechanism against SARS-CoV-2: An In Silico Approach, *Pharmacognosy Journal*, 2022, pp. 85-90. DOI: <https://doi.org/10.5530/pj.2022.14.12>
- [25] A. N. M. Ansori, R. J. K. Susilo, A. Fadholly, S. Hayaza, A. P. Nugraha, S. A. Husen, Antidiabetes type 2 phytomedicine: Mangosteen (*Garcinia mangostana* L.)-A review, *Biochemical and Cellular Archives*, 2020, pp. 3173-3177. DOI: <https://doi.org/10.35124/bca.2020.20.S1.3173>
- [26] S. A. Husen, D. Winarni, Salamun, A. N. M. Ansori, R. J. K. Susilo, S. Hayaza, Hepatoprotective Effect of Gamma-mangostin for Amelioration of Impaired Liver Structure and Function in Streptozotocin-induced Diabetic Mice, *IOP Conference Series: Earth and Environmental Science*, 2019, pp. 012031. DOI: <https://doi.org/10.1088/1755-1315/217/1/012031>
- [27] S. A. Husen, D. Winarni, F. Khaleyla, S. H. Kalqutny, A. N. M. Ansori, Activity assay of mangosteen (*Garcinia mangostana* L.) pericarp extract for decreasing fasting blood cholesterol level and lipid peroxidation in type-2 diabetic mice, *AIP Conference Proceedings*, 2017, pp. DOI: <https://doi.org/10.1063/1.5004303>

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