



# Pharmacognostic characterization of leaves of ajuru plant species chrysobalanus icaco (l.) And its phytochemical activity

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#### **1 INTRODUCTION**

The use of medicinal plants has been described in history since the dawn of humanity. Over time, to obtain a cure for diseases, our ancestors observed everything that nature offered, especially plants, to discover which species would serve to cure diseases, which would be suitable for food preparations, and which were poisonous. and hallucinogenic (GASPAR, 2009). The World Health Organization (WHO) estimates that 80% of the world's population uses some kind of plant to improve or treat signs and/or symptoms of pathological processes (ANVISA, 2002).

Among the possible species to be found in the Brazilian flora, several studies have shown the high potential for use/therapeutic purpose of a species linked to the Chrysobalanaceae family, the species Chysobalanus icaco L., popularly known in Brazil as guajuru, abageru, ajiru, among other names, and can be found in the states of Pará, Ceará, Maranhão, Pernambuco, Rio de Janeiro, and Rio Grande do Norte (PRESTA; PEREIRA, 1987; SILVA; PEIXOTO, 2009; AGUIAR, 2010), it is

classified as an angiosperm, belonging to the genus Chysobalanus. Initially, the Chysobalanaceae family, in which it is found, has a wide distribution in subtropical and tropical regions, and geographically it is present in European countries, Africa and Latin America, since its morphology, commonly of woody and shrubby species, is known in countries English-speaking ones like coco-plum, icaque-ponne, pork-fat-apple, and zicate; while in French-speaking countries it is known as Prunier de cacao (KRUEL; PEIXOTO, 2004).

Ajuru is a species that occurs in places with water deficit or flooding, such as swamps, varying



according to the seasons, developing in leached, sandy and shallow soils, it can be found in Restinga areas in the zones coastal and coastal areas and preferentially inhabits dunes (VARGAS, 1998; KRUEL; PEIXOTO, 2004; FREITAS; LOCATELLI, 2009). For being a kind

medium-sized wild, C. icaco has a height that varies between 1.5 and 3 meters (AGUIAR, 2010), with a trunk diameter that varies from 15 cm to 30 cm and hard, oval leaves, of a shiny dark green color on the upper part, which are up to 4 cm long and 3 to 6 cm wide (PRESTA; PEREIRA, 1987; MATOS, 1999; AGUIAR, 2010). Its fruit is rounded, with a width between 2 and 5 cm, with a diversified color that varies between creamy white and pink, with small variations for black, presenting the pulp in the color white with a spongy, sweet, or insipid characteristic that, when maturing, is quite astringent (AGUIAR, 2010; PERES, 2012).

As for medicinal use/application, the Chrysobalanaceae family comprises species with great therapeutic value for popular use, with a relative emphasis on the species considered in the present study, which is traditionally used in the northern region of Brazil for the treatment of diabetes (AGUIAR, 2010; PERES, 2012), in the treatment of leukorrhea, hemorrhages and chronic diarrhea (PRESTA; PEREIRA, 1987; GOMES, 2006), and is also known for its diuretic and hypoglycemic effect (FRANCIS, 2003). In Brazil, in coastal regions, it has its prominence in food, as for riverside people, but in American countries, Mexico and Cuba, it has economic importance, for the manufacture of sweets and preserves (AGUIAR, 2010). For medicinal purposes, seed oil is popularly used to prepare an antidiarrheal emulsion and for ointments. The roots, bark, and leaves are used against dysentery, leukorrhea, and kidney stones, due to their astringent characteristic. Its potential use as an antitumor agent and in the fight against diabetes mellitus is widely explored (TORRES et al., 2009; AGRA et al., 2008; KRUEL et al., 2006).

As for the biological activity, the authors Presta and Pereira (1987), Castilho et al. (2000), and Aguiar (2010) exposes that the leaves of C. icaco have analgesic, and anti-inflammatory characteristics, and properties with genotoxic effects. The extracts and fractions of the species have antibacterial activity against Staphylococcus aureus and Streptococcus pyogenes and hypoglycemic activity (CASTILHO et al., 2000). In their research, Gustafson et al. (1991) and Peres (2012), respectively, concluded that the diterpenes of Chrysobalanus icaco also have anti-HIV activity and that the hydroalcoholic extract of the leaves has been used in the treatment of mycoses and fungal infections of candidiasis. Concerning, popular medicine, the leaves of C. icaco have analgesic and anti-inflammatory, anesthetic, and hypoglycemic characteristics (BARBOSA et al., 2005: TORRES et al., 2009; CASTILHO et al., 2000; FRANCIS, 2003; AGUIAR, 2010).

Such studies denote, therefore, the need for more in-depth exploratory studies about the still unknown potential of the species, as well as pointing to the enormous possibility of discovering new herbal medicines, drugs, and/or pharmaceutical raw materials that legitimize its traditional use. Therefore, based





on the presented potential, it was proposed through the study to carry out the pharmacognostic characterization of the leaves of Ajurú collected in the urban area of the city of Macapá/AP, the objective of this research was to determine and identify, through the phytochemical study, the main classes of secondary metabolites found in the leaves of the plant species C. icaco.

## **2 METHODOLOGY**

Based on the methodology of Barbosa et al. (2001), collections of plant material were carried out, the leaves of C. icaco, located in the city of Macapá – Amapá (0°5'57.772"; 51°2'46.701"), and specimens were produced for cataloging. At the Laboratory of Pharmacognosy and Phytochemistry of the Federal University of Amapá, the collected material was separated and dried in an oven at 45°C, shortly after grinding, the powder obtained was weighed, resulting in 400g of material. For the process of obtaining the crude extract, according to the basic methodology, the material was placed in alcoholic extraction, with Hydrated alcohol at 92.8° INPM for 72 hours at room temperature, it undergoes a rota evaporation process, obtaining, after 3 extractions and drying of the extract, 3.31g of crude extract was obtained.

The phytochemical analyses carried out were: cardiac glycosides, saponins, phenols and tannins, alkaloids, catechins, reducing sugars, steroids and triterpenoids, coumarin derivatives, organic acids, proteins and amino acids, purines, anthraquinones, flavonoids, carotenoids, sesquiterpene lactones, depsides, and depsidones, according to the methodology of Barbosa (2001).

#### **3 CONCLUSION**

According to the phytochemical analysis of the classes of secondary metabolites carried out in the crude extract of the leaves of C. icaco, it was possible to positively identify: foaming saponins, catechins, carotenoids, phenols, tannins, depsides and depsidones, steroids and triterpenes (**Table 1**).

Secondary metabolite class	Results
Organic Acids	-
Reducing sugars	-
alkaloids	-
anthraquinones	-
Carotenoids	+
Catechin	+
Depsides and Depsidones	+
coumarin derivatives	-

Table 1- Phytochemical tests performed to identify classes of secondary metabolites





Steroids and triterpenoids	+
Phenols and Tannins	+
flavonoids	-
cardiac glycosides	-
Proteins and Amino Acids	-
Purines	-
saponins	+
Sesquiterpecolactones and other lactones	-
Parameters: Negative (-); Positive (+)	

From the results of the phytochemical prospection, it was possible to observe the presence of foaming saponins, historically in the literature as anti-inflammatory, varying the mechanism, being able to act by inhibiting the degradation of corticoids, causing a direct interference in the mediators, resulting in this biological action. Hemolytic activity can be due to the possible ability to complex with steroids, proteins, and phospholipids, changing the permeability of cell membranes, thus being able to have cytotoxic, antitumor, and antiviral action (BESSA et al., 2013; CASTEJON, 2011; MACEDO et al., 2007). Saponins are steroid glycosides or polycyclic terpenes, have a high molecular mass, and generally have complex structures (BARBOSA et al., 2020).

Saponins can act as a cholesterol reducer due to their irritating activity, possibly due to the mechanism of intestinal cholesterol interaction, causing exfoliation with loss of function and reduction of the area of absorption, which may cause a consequent reduction in cardiac congestive failure by inhibiting cellular efflux of Na+, acts as an expectorant, entropic, molluscicidal and healing agent (CASTEJON, 2011; MACEDO et al., 2007). Concomitantly, the class of catechins also known as flavonoids subclass of flavonoids found in Ajuru, belong to the class of polyphenols and have characteristics of colorless, water-soluble compounds, which favor the bitterness and astringency of the vegetable (STRACK; SOUZA, 2012; LAMARÃO; FIALHO, 2009).

Its bioactivity may contribute to the reduction and incidence of some cancers, an effect induced by apoptosis, in the reduction of body fat, this is suggested as a modulator of norepinephrine in the sympathetic system, acting on thermogenesis and fat oxidation, as well as its possible role as a potent antioxidant, as a scavenger of free radicals, and in aiding the reduction of total cholesterol by a probable mechanism of action in the inhibition of oxidized LDL (low-density lipoprotein); this may come to act as a protective agent in the release of chemical mediators from the immune system, portraying also its effects in reducing platelet reactivity, anti-inflammatories, and insulin sensitivity, benefits correlated to the decrease of dyslipidemia and cardiovascular diseases with increased intake of food sources (PEREIRA; CARDOSO,





# 2012; STRACK; SOUZA, 2012; LAMARÃO; FIALHO, 2009).

The class of carotenoids is characterized by the presence of the polyene chain, with a structure of a long system of conjugated double bonds, this can be associated with the biological and therapeutic activity as its probable antioxidant action, it is associated that the chain for presenting groups cyclic terminals with substituents, in which, with oxygen, it provides a conjugated system in polyene electrons, helping in the absorption of singlet oxygen and the free radicals present, through the interruption of the chain reactions that they involve, in addition, these can be correlated the in vitro peroxidation inhibition of lipids at low oxygen pressures (McNULTY, et al., 2007; QUIRÓS; COSTA, 2006; SIKORA et al., 2008).

The class of phenols identified in the analysis may have biological activity associated with their composition and form of use, it is suggested that internally it may have anti-diarrheal activity, through the inhibition of some enzymes in the digestive process, and external antiseptic action, which may be associated with its astringency, waterproofing the most exposed layers of the skin and mucous membranes, a fact that also leads to toxic effects (high mortality of treated insects) related to condensed tannins by associating greater weight of the molecule and by the easy mechanism of complexing with metallic ions (ANDREO; JORGE, 2006).

However, some studies suggest the physiological properties with antiallergenic, anti-arteriogenic, anti-inflammatory, antithrombotic, antimicrobial, cardioprotective (reduction of LDL), and vasodilator effects, but the main effect is its action as a powerful antioxidant, through the capture of concentrated intercellular free radicals intercepting oxygen that can cause the formation of stable radicals, which can act in the prevention of degenerative diseases such as cancer and the delay of aging, in low molecular weight and adequate amounts for easy absorption (MONTEIRO et al., 2005; CARVALHO, 2007).

The tannins identified in the aqueous extract of abageru leaves, this class being commonly associated with medicinal activities against infections and dysentery when roots, fruit peels, flowers, and leaves are used (PRESTA; PEREIRA, 1987; KRUEL et al., 2006; AGRA, FREITAS, BARBOSA, 2007;

LORENZI; MATOS, 2002). In this way, it can be correlated with the vasoconstrictor action, preventing external aggressions and fluid decrease; it heals, being able to regenerate tissues in cases of wounds or burns; has biological action against certain microorganisms, such as carcinogenic agents and antitumor agents, and may supposedly be correlated with the fight against diabetes mellitus, but they are still widely explored (TORRES et al., 2009; AGRA et al., 2008; KRUEL et al., 2006).

The pharmacological actions of tannins derive from their ability to form complexes with proteins and polysaccharides, thus contributing to the healing of wounds and burns. Similarly, this affinity of polyphenols for proteins is of essential importance in the activity of inactivating enzymes, thus preventing the growth of some microorganisms (SIMÕES, 2010).

The use of C. icaco leaves associated with cytotoxic activity may be associated with the presence





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of diterpenes and triterpenes (FERNANDES et al., 2003; GUSTAFSON; MUNRO, 1991), terpenoids, from C. icaco leaves, these may be associated with the in vitro inhibitory action on the Acquired Immunodeficiency Virus (HIV) (GUSTAFSON et al., 1991), supposedly composes the species, compounds such as pomolic acid and betulinic acid, homologous series of hydrocarbons of C27 and C33 (BARBOSA et al., 2006; FERNANDES et al., 2003; MENDEZ et al., 1995; CASTILHO et al., 2000), and trienoic and tetraenoic acids and their derivatives from seed oils (GUNSTONE; SUBBARAO, 1967; VERMA; RAYCHAUDHURI, 1972).

Polycyclic steroids or terpenes, which in their structural form have a lipophilic part (triterpene or steroid) and a hydrophilic part (sugars), which may be correlated to antifungal activity (BORELLA et al., 2006), among the main classes are the polyenes, azoles, allylamines, 5-fluorocytosine, griseofulvin, among others. However, polyenes and azoles are related in the treatment of candidiasis, in the hexanic extract of C. icaco leaves, antifungal activity was revealed against the

Candida albicans (CPRobin) Berkhout (CASTILHO et al., 2000; FERNANDES et al., 2003).

The action of depsides and depsidones is still vaguely elucidated, however, they are associated with activities against gram-positive bacteria, mycobacteria, nematodes, and insects, in addition to having antiinflammatory, anticancer, analgesic, antiviral, and antipyretic properties (ZHAO et al. ., 1997). It is known that depsides are an example of polyketides and are metabolites produced by lichens and endophytic microorganisms, which have as a chemical characteristic the presence of two or more aromatic rings that are linked by a carboxylic unit (SILVA et al. 2020).

Therefore, it was possible to associate these classes of metabolites identified in the crude extract of Ajuru leaves with antibacterial and anti-inflammatory activity correlated to C. icaco, which may have therapeutic potential, such as tannins, which are presented, according to the literature, effects against dysentery, thus associating the popular use. However, a microbiological study of the obtained extract is necessary to investigate the pharmacological potential of C. icaco leaves.





## REFERENCES

Agra, m.f. *Et al.* Survey of medicinal plants used in the region northeast of brazil. Revista brasileirade farmacognosia, n. 18, p. 472-508, 2008.

Agência nacional de vigilância sanitária, re nº 1185, de 09 de julho de 2002. Disponível em: <a href="http://www.anvisa.gov.br/legis/resol/2002/1185\_02re.htm">http://www.anvisa.gov.br/legis/resol/2002/1185\_02re.htm</a>>. Acesso em: 19 nov. 2022.

Aguiar, t. M. Caracterização química e física de folhas, frutos e sementes do bajuru (chrysobanalusicaco l.) E avaliação do chá dessas folhas em camundongos (suiss) normais e diabéticos. 2010. 171f. Dissertação (mestrado em ciência e tecnologia de alimentos) - universidade federal rural do rio dejaneiro.

Andreo, d.; jorge, n. Antioxidantes naturais: técnicas de extração. Boletim do centro de pesquisade processamento de alimentos. 2006, v.24, p.319-336.

Barbosa, w. L. R., *et al.* Manual para análise fitoquímica e cromatográfica de extratos vegetais.Revista científica da ufpa. Pará, v. 4, n. 5, 2001. (http://www.ufpa.br/rcientífica)

Barbosa, j.m; vasconcelos, t.h.c; alencar, a.a; batista, l.m; oliveira, r.a.g;guedes, d.n; falcão, h.s; moura, m.d; diniz, m.f.f.m; modesto, j. Plantas e seus

Constituintes ativos da américa do sul, central e do norte com hipoglicemia atividade. Revista brasileira de farmacognosia, ed.15 p. 392-413. 2005.

Barbosa, w.l.r; peres, a.; gallori, s.; vincieri, f.f. Determinação de derivados de miricetina em *chrysobalanus icaco* l. (chrysobalanaceae) revista brasileira de farmacognosia, joãopessoa, v. 16, n. 3 july/sept. 2006.

Barbosa, i.i.a. *Et al.* Estudo fitoquímico do extrato hidroalcoólico bruto de folhas de *ficus benjamina* l. (moraceae). In: ii jornada de botânica e ecologia e ii jornada amapaense de botânica,1., 2020, macapá. Anais da ii jornada de botânica e ecologia e ii jornada amapaense de botânica. Amapá: universidade do estado do amapá, 2020. P. 98.

Bessa, n.g. F.; borges, j. C. M.; beserra, f. P.; carvalho, r. H. A.; pereira, m. A. B.; fagundes, r.; campos, s. L.; ribeiro, l. U.; quirino, m. S.; chagas junior, a. F.;

Alves, a. Prospecção fitoquímica preliminar de plantas nativas do cerrado de uso popular medicinalpela comunidade rural do assentamento vale verde – tocantins. Rev. Bras. Plantas med. Campinas, v.15, n. 4, supl. I, p. 692-707, 2013.

Borella, j. C. *Et al.* Variabilidade sazonal do teor de saponinas de *baccharis trim*era (less.) Dc (carqueja) e isolamento de flavona. Revista brasileira de farmacognosia, joão pessoa, v. 16, n. 4, oct./dec. 2006.

Carvalho, e.b. Estudos da interação entre proteínas e taninos: influência da presença de polissacarídeos. 2007. 193f. Tese (doutorado em química) - universidade do porto, portugal, 2007.

Castejon, f.v. Taninos e saponinas. 2011. 29f. Tese (mestrado) - seminário apresentado junto à disciplina seminários aplicados do programa de pós-graduação em ciência animal da escola de medicina veterinária e zootecnia – universidade federal de góias, goiânia.

Castilho, r.o. Et al. A survey of chemical and biological activities of chrysobalanaceae. Anais daacademia



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brasileira de ciências, 72, p. 292-293. 2000.

Fernandes, j.; castilho, r. O.; costa, m.r.; souza, k.w.; kaplan, m.a.c.; gattass, C.r. Triterpenos pentacíclicos de espécies de chrysobalanaceae: citotoxicidade em linhas celulares de leucemia sensível e resistente a múltiplas drogas. Cancer lett. V. 190, p. 165-169. 2003.

Francis, j. Chrysobanalus icaco, l. Coco-plum. Forest service of united. Estates department of agriculture (usda). 2003. (http://www.fs.fed.us/global/iitf/pdf/chrysobanalus%20icaco20l.pdf)

Freitas, t.; locatelli, e. Anais do ix congresso de ecologia do brasil. São lourenço – mg,2009. P.1-3. Gaspar, lúcia. Plantas medicinais. Pesquisa escolar online, fundação joaquim nabuco, recife. Disponível em: <a href="http://basilio.fundaj.gov.br/pesquisaescolar/">http://basilio.fundaj.gov.br/pesquisaescolar/</a>. Acesso em: 19 nov. 2022.

Gomes, m.l. *Et al.* Usos medicinais e composição química das folhas de licania macrophylla benth. (chrysobalanaceae). Revista brasileira de farmácia. V. 87, p. 26-29, 2006.

Gunstone, f.d.; subbarao, r. Novos óleos de sementes tropicais. Parte i. Conjugou os ácidostrienoico e tetraenóico e seus oxidantes nos óleos de sementes de *chrysobalanus icaco* e *parinarium laurinum*. Chemistry and physics of lipids, p. 349-359. 1967.

Gustafson, k. R. *Et al.* Hiv inhibitory natural products 3-diterpenes from homalonthus acumonathus and chrysobalanus icaco. Tetrahedron, v.47, n.26, p. 4547-4554. 1991.

Kruel, v.s. F.; peixoto, a.l. Etnobotânica na reserva extrativista marinha de arraial do cabo, rj,brasil. Acta botânica brasileira, porto alegre, v. 18, n. 1, p. 177-190. 2004.

Kruel, v.s.f.; peixoto, a.l.; sá, c.f.c.; araújo, d.s.d.; silva, w.l.; ferreira, a.j. Plantas úteis da restinga: o saber dos pescadores artesanais de arraial do cabo. Rio de janeiro: institutode pesquisas jardim botânico do rio de janeiro. 2006.

Lamarão, r.c.; fialho, e. Aspectos funcionais das catequinas do chá verde no metabolismo celular e sua relação com a redução da gordura corporal. Rev. Nutrição. Campinas, v.22, n. 2, p. 257-269, 2009.

Lorenzi, h.; matos, f. J. A. Plantas medicinais no brasil. Nova odessa: instituto plantarum. 2002.

Macedo, f. M.; martins, g. T.; rodrigues, c. G.; oliveira, d. A. Triagem fitoquímica do Barbatimão [stryphnodendron adstringens (mart) coville]. Revista brasileira de biociências. Porto alegre, v. 5, supl. 2, p. 1166-1168, 2007.

Matos, f. J. A. Plantas da medicina popular do nordeste. Ufc edições. 1999.

Mcnulty, h.p.; byun, j.; lockwood, s.f.; jacob, r.f.; mason, p. Differential effects of Carotenoids on lipid peroxidation due to membrane interactions: x-ray diffraction analysis. Biochimicaet biophysica acta, v. 1768, n. 1, p. 167-174, 2007.

Mendez, j.; bilia, a.r.; morelli, i. Investigações fitoquímicas do gênero licania. Flavonoides etriterpenoides de *licania pittieri*. Pharmaceutica acta helvetiae, p. 223-226. 1995. Monteiro, j.m.; albuquerque, u.p.; araújo, e.l. Taninos: uma abordagem da química àecologia. Rev. Química nova, v. 28, n. 5, p. 892-896, 2005.





Peres, a.r m.n. Caracterização farmacognóstica e avaliação antifúngica das folhas de chrysobalanus icaco (lin) em espécies de candida. 2012. 112 f. Dissertação (mestrado) - universidadefederal do pará, instituto de ciências da saúde, belém, 2012. Programa de pós-graduação em ciências farmacêuticas.

Presta, g. A.; pereira, n. A. Atividade do abagerú (crysobalanus icaco lin, crysobalanaceae) em modelos experimentais para o estudo de plantas hipoglicemiantes. Revista brasileira de farmácia, v.68, p. 91. 1987.

Quirós, a. R.; costa, h. S. Analysis of carotenoids in vegetable and plasma samples: a review. Journal of food composition and analysis, v. 19, p. 97-111, 2006.

Sikora, e.; cieslik, e.; leszczynska, t.; filipiak-florkiwuacz, a.; pisulewski, P.m. The antioxidant activity of selected cruciferous vegetables subjected to aqua thermal processing.Food chemistry, london, v. 107, p. 50-55, 2008.

Silva, i. M.; peixoto, a. L. O abajuru (*chrysobalanus icaco* l. E eugenia rotundifolia casar.) Comercializado na cidade do rio de janeiro, brasil. Revista brasileira de farmacognosia, v. 19, n.1b,p. 325-332. 2009.

Silva, m.l.c. *Et al.* Compostos fenólicos, carotenóides e atividade antioxidante em produtos vegetais. Semina: ciências agrárias, v. 31, n. 3, p. 669-681, 2010. (https://www.redalyc.org/pdf/4457/445744097017.pdf).

Silva, a.s. *Et al.* Análise fitoquímica das folhas de heliconia psittacorum l.f. (heliconiaceae). In: iijornada de botânica e ecologia e ii jornada amapaense de botânica, 1., 2020, macapá. Anais da ii jornada de botânica e ecologia e ii jornada amapaense de botânica. Amapá: universidade do estadodo amapá, 2020. P. 37.

Simões, c. M. O. *Et al.* Farmacognosia: da planta ao medicamento. 6 ed. Porto alegre: ufrgs, p.1102. 2010.

Strack, m.h.; souza, c.g. Antocianinas, catequinas e quercetina: evidências na prevenção e notratamento de doenças cardiovasculares. Rev bras nutr clin 2012; v.27, n.1, p: 43-50. Torres, d.f.; oliveira, e.s.; alves, r.r.n.; vasconcellos, a. Etnobotánica y Etnozoología em unidades de conservación: uso de la biodiversidade em la apa de genipabu, rio grande del norte, brasil. Inci, v. 34, n. 9, p. 623-629. 2009.

Vargas, s.g.f. Icaco (chrysobalanus icaco l.): análisis químico de flavonoides y propagación porestacas. 1998. 65 f. Dissertação (mestrado em ciências) – colégio de postgraduados en ciencias agrícolas, montecillo edo. De méxico.

Verma, v. S.; raychaudhuri, s. P. Efeito de taninos de catecol isolados de uma planta medicinal chrysobalanus icaco na infectividade do vírus da batata x. Zentralbl bakteriol parasitenkdinfektionskr hyg, v. 127, p. 178-179. 1972.

Zhao, h. *Et al.* Coumarin-based inhibitors of hiv integrase. Journal of medicinal chemistry, v. 40, p.242-249, 1997.



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