

Full Length Research Paper

Biotechnological potential of the *Carapa guianensis*, *Bertholletia excelsa* and *Copaifera* spp. oils

Naila Fernanda Sbsczk Pereira Meneguetti^{1*}, Dionatas Ulises de Oliveira Meneguetti^{1,2,3,4}
and Amauri Siviero^{1,5}

¹Programa de Pós-Graduação Stricto Sensu em Biodiversidade e Biotecnologia da Amazônia Legal,
Universidade Federal do Acre, Rio Branco, AC, Brasil.

²Programa de Pós-Graduação Stricto Sensu em Ciência da Saúde na Amazônia Ocidental,
Universidade Federal do Acre, Rio Branco, AC, Brasil.

³Programa de Pós-Graduação Stricto Sensu em Ciência Inovação e Tecnologia para a Amazônia,
Universidade Federal do Acre, Rio Branco, AC, Brasil.

⁴Colégio de Aplicação, Universidade Federal do Acre, Rio Branco, AC, Brasil.

⁵Empresa Brasileira de Pesquisa Agropecuária, Rio Branco, AC, Brasil.

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The use of vegetable species for treatment or prevention of diseases is one of the most ancient medicine techniques in humanity. In Amazon rainforest, there are numerous of plant species that possess biotechnological potential, and due to this, this study aimed at carrying out a bibliographic review describing the biotechnological potential in *Carapa guianensis*, *Bertholletia excelsa*, and *Copaifera* spp. oils. This study is a systematic review of literature in the databases: SCIELO, VHL and PUBMED, by using the descriptors: *C. guianensis* oil, *B. excelsa* oil and *Copaifera* spp. After the selection, 87 articles were selected; 48 on *C. guianensis*, 9 on *B. excelsa*, and 30 on *Copaifera* spp. Oil seeds from Amazon rainforest possess a good biotechnological potential to be explored. This review has shown that *C. guianensis* and *Copaifera* spp. are strong candidates for the search of new insecticide, antiparasitic, anti-inflammatory and healing products. *Copaifera* spp. has also shown to be promising for the production of antibiotic and antifungic medicines, and *C. guianensis* for acaricide drugs. Future studies are indicated to deepen the knowledge already described and to investigate new biotechnological potentialities of Amazonian oil seeds. However, for this to happen, greater financial support is required from the Brazilian government and agencies, thus increasing the level of research in the region, which will consequently maximize the likelihood of new product discovery.

Key word: Biotechnology, oilseeds, Amazon rainforest.

INTRODUCTION

Ethnopharmacology is defined as “the interdisciplinary scientific exploration of biologically active agents

*Corresponding author. E-mail: naila_sbsczk@hotmail.com.

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Table 1. Amount of articles found in literature search.

Descriptor	SCIELO	VHL	PUBMED	Total
<i>Carapa guianensis</i> oil	26	52	30	108
<i>Bertholletia excelsa</i> oil	12	24	12	48
<i>Copaifera</i> spp oil [#]	23	64	12	99
Total	61	140	54	255

[#]Species searched: *Copaifera duckei* oil, *Copaifera langsdorffii* oil, *Copaifera martii* oil, *Copaifera paupera* oil, *Copaifera pubiflora* oil, and *Copaifera reticulata* oil.

traditionally employed or observed by man" (Bruhn and Holmstedt, 1981; Elisabetsky, 2003). This area combines information gained from medicinal flora users (communities and traditional specialists) with chemical and pharmacological studies (Elisabetsky, 2003).

The use of vegetable species for treatment and prevention of diseases is one of the most ancient medicine techniques in humanity (Dutra, 2009). Even though plants have been used for centuries for medicinal purposes, the vast majority have unknown biotechnological features by science (Meneguetti and Facundo, 2015).

Data show that about 95% of Brazilian native species still do not possess scientific studies on their possible applicabilities (Santos, 2009), representing a huge economic and biotechnological potential to be explored, especially in vegetable species from Amazon rainforest, as many have still not been catalogued by science (Cechinel and Rosendo, 1998; Abreu et al., 2001; Meneguetti and Facundo, 2015). It can present new chemical compounds and consequently new specific and efficient drugs (Lang et al., 2008).

There are plenty of plants species in Amazonian rainforest that possess biotechnological potential (Osakada, 2009), among them are presented *Carapa guianensis* (Andiroba), *Bertholletia excelsa* (Amazônia's nut) and *Copaifera* spp. (Copaíba). Furthermore, this study aimed at carrying out a bibliographic review describing the biotechnological potential of these species' oil.

MATERIALS AND METHODS

This study is a systematic review of literature based on Galvão and Pereira (2014), following the steps below:

- (a) Creation of research question: What is the biotechnological potential described in literature of Andiroba, Amazônia's Nut, and Copaíba oils?
- (b) Literature search: Search for scientific articles in the following databases: Scientific Electronic Library Online (SCIELO), Virtual Health Library (VHL) and National Institute of Health (PUBMED) by using the descriptors: *C. guianensis* oil, *B. excelsa* oil and *Copaifera* spp. oil of species (*Copaifera duckei* oil, *Copaifera langsdorffii* oil, *Copaifera martii* oil, *Copaifera paupera* oil, *Copaifera pubiflora* oil and *Copaifera reticulata* oil) because they were the

main copaíba species in the Brazilian Amazon rainforest. Articles in Portuguese, English and Spanish published by 2018 were included. The amount of articles found in the survey is presented in Table 1.

(c) Article selection: This selection was performed by three researchers, and the articles rejected by two or three of them were removed from this research. The criteria for exclusion were repeated articles (found in more than one database), out of the research's aim, low methodological quality and with negative results for biotechnological potential.

d) Data gathering: After the selection, 87 articles were selected: 48 on *C. guianensis*, 9 on *B. excelsa*, and 30 on *Copaifera* spp. Those were used in the results of this study. Besides the selected articles, others were used for creation of introduction and improvement of this article's discussion.

e) Data synthesis: The data were organized into tables, and described in the text according to their biotechnological potential.

f) Results discussion writing: Description and discussion of the data are at "Results and Discussion" present in the sequence.

RESULTS AND DISCUSSION

Insecticide activity

The data showing that the oils searched have insecticide activity are shown in Table 2.

The *C. guianensis* and *Copaifera* spp. oils have shown a good potential in the medical entomology for vector control of tropical diseases, such as *A. aegypti*: Dengue, Chikungunya, and Zika virus vector (Aragão et al., 2018); *A. albopictus*: Yellow fever vector (IEC, 2018); and *C. quinquefasciatus*: that has been also found infected by Zika virus (Smartt et al., 2018).

C. guianensis develop products against ectoparasitosis due to their activity against *F. subrostratus* (popularly known as lice) that is a public health problem in Brazil. It is estimated that up to two thirds of deprived population and communities were affected in the beginning of the year 2000 (Heukelbach et al., 2003).

In livestock, *C. guianensis* has proven to be promising against two flies species: *M. domestica* and *H. irritans*, with this last one known popularly as "Horn Fly", which causes several problems in the Brazilian cattle (Klauck et al., 2014).

C. guianensis and *Copaifera* spp. may also be beneficial to agriculture, acting as anti-*S. frugiperda* popularly known as "Armyworm" (Santos et al., 2016), *A.*

Table 2. Insecticide oilseeds performance from Amazon rainforest.

Insect	<i>C. guianensis</i>	<i>B. excelsa</i>	<i>Copaifera</i> spp	Reference
<i>Aedes aegypti</i> (larvicide)	X	-	X	Mendonça et al. (2005)* Silva et al. (2006)* Abed et al. (2007)*** Silva et al. (2007)*** Geris et al. (2008)*** Rodrigues et al. (2014)***
<i>Aedes aegypti</i> (repellent)	X	-	-	Miot et al. (2004)* Jesus et al. (2017)*
<i>Aedes albopictus</i> (larvicide)	X	-	-	Silva et al. (2004)*
<i>Atta sexdens rubropilosa</i>	X	-	-	Ambrozin et al. (2006)*
<i>Culex quinquefasciatus</i> (larvicide)	-	-	X	Silva et al. (2003)***
<i>Diabrotica speciosa</i>	-	-	X	Barbosa et al. (2013)***
<i>Felicola subrostratus</i>	X	-	-	Barros et al. (2012)*
<i>Haematobia irritans</i>	X	-	-	Klauck et al. (2014)*
<i>Musca domestica</i>	X	-	-	Klauck et al. (2014)*
<i>Spodoptera frugiperda</i>	X	-	X	Santos et al. (2016)* Santos et al. (2016)***
<i>Tribolium castaneum</i>	-	-	X	Melo et al. (2015)***
<i>Zabrotes subfasciatus</i>	-	-	X	França et al. (2012)***

*Citations reference to *C. guianensis*; ** Citations referent to *B. excelsa*; *** Citations referent to *Copaifera* spp.

sexdens rubropilosa or “Sauba Ant” (Ambrozin et al., 2006), *D. speciosa* known as “vaquinha verde” (Barbosa et al., 2013), *Z. subfasciatus* known as “caruncho do feijão” (França et al., 2012) and *T. castaneum* as “May Beetle” (Melo et al., 2015), and they may cause jeopardy to different types of crops.

In the databases, there were no articles found showing that *B. excelsa* has insecticide activity.

Acaricide activity

The *C. guianensis* oil was the most promising in this study; shown to be anti-*Anocentor nitens* (Farias et al., 2009; Farias et al., 2012), *Rhipicephalus sanguineus* (Farias et al., 2009; Farias et al., 2012; Vendramini et al., 2012a; Vendramini et al., 2012b; Roma et al., 2013; Roma et al., 2015), and *Rhipicephalus (Boophilus) microplus* (Farias et al., 2012; Chagas et al., 2012). The *B. excelsa* oil has also shown to be anti-*R. (B.) microplus*, however it was considered low (Villarreal et al., 2017).

The *A. nitens* tick is one of the species that affects horses, and it is vector of *Babesia caballi* protozoan, which is an etiologic agent of equine babesiosis, disease that promotes low development and death in animals (Borges and Leite, 1993; Bello et al., 2008). It is known as “Spinose Ear Tick”, and it is responsible for injuries in the pinna, productivity drop, irritation, blood spoliation,

propensity to myiasis and secondary bacterial infections (Borges and Leite, 1993; Bello et al., 2008).

R. sanguineus has high prevalence in urban dogs, and is one of the species of parasites of these animals (Labruna and Pereira, 2001; Szabó et al., 2001; Soares et al., 2006; Paz et al., 2008). It is one responsible for pathogenic agent’s transmission, such as *Babesia canis* and *Ehrlichia canis* (Smith et al., 1976; Gothe et al., 1989). There are records of infestation of this species in humans (Guglielmone et al., 2006), as they may become a secondary vector of Ehrlichiosis, Babesiosis and Spotted Fever (Fernandes et al., 2001; Paz et al., 2008).

R. (B.) microplus is the main species of ticks that affects cattle in Brazil, which causes reduction in milk production; the main economic impact due to it (Rodrigues and Leite, 2013). It may also cause jeopardy in meat and leather production, besides of several other diseases that can be transmitted by them (Massard and Fonseca, 2004; Andreotti et al., 2011; Santos et al., 2018).

Antiparasitic activity

Among the oils studied, parasitic activity was found against eight parasites species (Table 3).

The *C. guianensis* oil has shown activity against three species of goats and sheep gastronintestinal nematodes:

Table 3. Antiparasitic activity of Amazonian oleaginous oils.

Parasite	<i>C. guianensis</i>	<i>B. excelsa</i>	<i>Copaifera</i> spp.	Reference
<i>Haemonchus</i> sp.	X	-	-	Farias et al. (2010)*
<i>Leishmania amazonenses</i>	-	-	X	Santos et al. (2008a)*** Meneguetti et al. (2015)***
<i>Leishmania chagasi</i>	-	-	X	Rondon et al. (2012)*** Meneguetti et al. (2015)***
<i>Oesophagostomum</i> sp.	X	-	-	Farias et al. (2010)*
<i>Plasmodium falciparum</i>	X	-	-	Miranda juúnior et al. (2012)* Pereira et al. (2014)* Nardi et al. (2016)*
<i>Plasmodium berghei</i>	X	-	-	Pereira et al. (2014)*
<i>Trichostrongylus</i> sp	X	-	-	Farias et al. (2010)*
<i>Trypanosoma evansi</i>	-	-	X	Dorneles et al. (2013)

*Citations reference to *C. guianensis*; ** Citations referent to *B. excelsa*; *** Citations referent to *Copaifera* spp.

Haemonchus sp., *Oesophagostomum* sp., and *Trichostrongylus* sp. (Farias et al., 2010). It has positive performance in veterinary and potential for future studies with helminths that affect humans.

C. guianensis was also antiparasitic against two species of *Plasmodium* gender, being the etiologic agent of malaria; disease among the four main epidemics in Latin America, with about 100 thousand new cases each year (Braz et al., 2006; Ferreira et al., 2012; Meneguetti et al., 2014). The anti-malaria potential is important because there is a need to search for new drugs against malaria, as it is resistant to nowadays drugs used, and being a threat to the disease control (Meneguetti et al., 2014).

Copaifera spp. has shown activity against trypanosomatids genders: *Leishmania* and *Trypanosoma*, Leishmaniasis and Chagas' disease etiologic agents, diseases included in the group of Neglected Diseases, in which only 10% of the world's expenses with research in health are destined to diseases that account for 90% of the global ill people (Bezerra et al., 2012).

Leishmaniasis has shown the need for urgente new candidates for treatment drugs (Santos et al., 2013a), because today the first choice for leishmaniasis treatment have been pentavalent antimonials (Sb^{5+}), amidines, polyene, aminoglycosides, and hexadecylphosphocholine (miltefosine). However, some present toxicity for the patient (Bezerra et al., 2004; Figueiredo et al., 2014; Meneguetti et al., 2015).

Currently the treatment of Chagas' disease presents only one drug available in Brazil, benzonidazole (2-nitroimidazole) introduced in the therapy in 1967 and to date a drug with satisfactory cure potential has not been developed and the drug used is inefficient and presents various side effects (Bezerra et al., 2012).

In the searched databases no articles were found that demonstrate the antiparasitic action of *B. excelsa*.

Antimicrobial activity

The oils studied showed antimicrobial activity against bacteria and fungi, as can be observed in Table 4. *Copaifera* spp. presented antimicrobial action; presented activity against 14 species, followed by *C. guianensis* that had action against 3 species. In the searched databases, no articles were found that demonstrate the antimicrobial action of *B. excelsa*.

The action against the microorganisms observed in Table 4, demonstrates the potential of the *Copaifera* spp. and *C. guianensis* oils for the development of drugs for the treatment of periodontal diseases, since it has had action against *A. actinomycetemcomitans*, *E. faecalis* and *P. gingivalis*. When they are present, several periodontal diseases are associated with cases of failure of endodontic treatment (Gasparetto et al., 2000; Carvalho and Cabral, 2007; Nacif and Alves, 2010).

Microbicidal activity against *E. coli*, *S. aureus*, *S. epidermidis* and *L. monocytogenes* is also highlighted, as they may cause several problems to human health, especially infections (Farbe and Peterkin, 1991; Lowy, 1998; Otto, 2009; Matos et al., 2015), which are often resistant to antibiotics (Santos, 2014), which demonstrates the importance of research for the development of new drugs.

The *Copaifera* spp. oil has an antifungal potential, with action against the dermatophyte *T. rubrum* (Dias et al., 2015) and species of the genus *Candida*, which causes the disease candidiasis (Calderone and Fonzi, 2001), which according to the *Associação de Obstetrícia e Ginecologia do Estado de São Paulo* (SOGESP - Association of Obstetrics and Gynecology of São Paulo State), affects 75% of women and of these, almost half will have a second episode and about 5% will present the condition more than once a year (Galileu, 2018). There is a large pharmacological market to be exploited for the

Table 4. Antimicrobial activity of Amazonian oleaginous oils.

Microrganism	<i>C. guianensis</i>	<i>B. excelsa</i>	<i>Copaifera</i> spp.	Reference
<i>Actinobacillus actinomycetemcomitans</i>	-	-	X	Dias et al. (2015)***
<i>Bacillus cereus</i>	-	-	X	Santos et al. (2013a)***
<i>Bacillus subtilis</i>	-	-	X	Santos et al. (2008b)***
<i>Candida glabrata</i>	-	-	X	Alencar et al. (2015)***
<i>Candida krusei</i>	-	-	X	Alencar et al. (2015)***
<i>Enterococcus faecalis</i>	X	-	X	Meccia et al. (2013)* Santos et al. (2008b)***
<i>Escherichia coli</i>	X	-	-	Brito et al. (2001)*
<i>Listeria monocytogenes</i>	-	-	X	Santos et al. (2013b)***
<i>Microsporum canis</i>	-	-	X	Dias et al. (2015)***
<i>Porphyromonas gingivalis</i>	-	-	X	Dias et al. (2015)*** Brito et al. (2001)* Meccia et al. (2013)*
<i>Staphylococcus aureus</i>	X	-	X	Santos et al. (2008b)*** Santos et al. (2013a)*** Alencar et al. (2015)***
<i>Staphylococcus epidermidis</i>	-	-	X	Alencar et al. (2015)*** Santos et al. (2008b)***
<i>Streptococcus mitis</i>	-	-	X	Dias et al. (2015)***
<i>Streptococcus</i> sp	-	-	X	Ziech et al. (2013)***
<i>Trichophyton rubrum</i>	-	-	X	Dias et al. (2015)***

*Citations reference to *C. guianensis*; ** Citations referent to *B. excelsa*; *** Citations referent to *Copaifera* spp.

production of antifungal agents.

Other activities with biotechnological potential

Other activities of *C. guianensis*, *B. excelsa* and *Copaifera* spp. can be observed in Table 5.

The anti-inflammatory and healing effects of *C. guianensis* and *Copaifera* spp. are in agreement with their popular use, since both are used for these purposes, demonstrating the importance of the empirical knowledge of traditional populations for scientific research.

The antioxidant effect of *C. guianensis* and *B. excelsa* is very well seen in the pharmacological and food industry, since antioxidants prevent the formation of free radicals in the body, retarding cellular aging (Silva and Ferrari, 2011), which if added to the antigenotoxic effect of *C. guianensis* (Lemes et al., 2017) may be an alternative for future studies of cancer treatment. In the case of *B. excelsa*, which is used for the production of olive oils (Valdez et al., 2009) and other beverages (Felberg et al., 2009), the antioxidant characteristic, together with its nutritional power (Spini et al. 2006) further yields economic value to the product.

In addition to all the activities observed in Table 5, in a

study with rats, *C. guianensis* did not produce toxic effects (Costa-Silva et al., 2008), genotoxic (Milhomem-Paixão et al., 2016), did not provoke chromosome aberrations in bone marrow cells (Arrebola et al., 2013) and did not interfere in the fertility and development of offspring (Costa-Silva et al., 2006). These data demonstrate a safety for the production of drugs from this species.

Copaifera spp. has also demonstrated safety for the use of oil as a therapeutic agent, since it does not present acute toxicity and neurotoxic effects (Sachetti et al., 2009) and still has neuroprotective action (Santos et al., 2012). When used in combination with vaginal cream and applied to rats, it has been shown to be safe during pregnancy (Lima et al., 2011).

B. excelsa has also been shown to be a safe species for use, since its almond is already widely used in food. This species still presents biodiesel production potential, as was observed in *C. guianensis* (Stachiw et al., 2016), which has also been shown to be a good catalyst (Tiosso et al., 2014).

Conclusion

It was verified that the Amazonian oilseeds have a good

Table 5. Other activities of Amazon rainforest oleaginous oils.

Activity	<i>C. guianensis</i>	<i>B. excelsa</i>	<i>Copaifera spp</i>	Reference
Antiallergic	X	-	-	Penido et al. (2005)* Nardi et al. (2016)* Ferraris et al. (2011)* Ferraris et al. (2012)* Henriques and Penido, 2014)*
Anti-inflammatory	X	-	X	Penido et al. (2005)* Penido et al. (2006)* Henriques and Penido, 2014)* Nardi et al. (2016)* Higuchi et al. (2017)* Wanzeler et al. (2018)* Carvalho et al. (2005)*** Muniz et al. (2009)*** Teixeira et al. (2017)***
Antioxidant	X	X	-	Milhomem-Paixão et al. (2016)* Vieira and Regitano-D'arce, 1999)* Gomes et al. (2016)*
Antigenotoxic	X	-	-	Lemes et al. (2017)* Botelho-Brito et al. (2001)* Santos et al. (2013a)* Silva et al. (2015)*
Healing	X	-	X	Wanzeler et al. (2018)* Estevão et al. (2009)*** Estevão et al. (2013)*** Feitosa Junior et al. (2018)***
Neuroprotective	-	-	X	Santos et al. (2012)***

biotechnological potential to be explored. The present review showed that *C. guianensis* and *Copaifera spp.* are strong candidates in searching for new products with insecticidal, antiparasitic, anti-inflammatory and cicatrizing activity.

Copaifera spp. has also been shown to be promising for the production of antibiotics and antifungals and *C. guianensis* for acaricidal drugs.

Future studies are indicated to deepen the knowledge already described and to investigate new biotechnological potentialities of Amazonian oilseeds. However, for this to happen, greater financial support is required from the Brazilian government and agencies and the interaction between different researchers, laboratories and research groups to form multidisciplinary and interdisciplinary teams, thus increasing the level of research in the region, which will consequently maximize the likelihood of new product discovery.

CONFLICT OF INTERESTS

The authors have not declared any conflict of interests.

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