

## TROPICAL FRUIT PEELS AS SOURCES OF BIOACTIVE COMPOUNDS: A REVIEW

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### Abstract

Tropical ecosystems are rich in plant species of special interest to the pharmacological industry. Tropical fruit peels contain several bioactive compounds and show biological activities, unfortunately they don't get enough attention. This is an updated review of the bioactive compounds present in tropical fruits peels: *Actinidia deliciosa* (Kiwifruit), *Mangifera indica* L. (Mango), *Annona squamosa* L. (Sugar-apple), *Annona reticulata* L. (Custard apple), *Ananas comosus* (Pineapple), *Canarium odontophyllum* Miq. (Dabai), *Carica papaya* L. (Papaya), *Cucumis sativus* L. (Cucumber), *Persea americana* Mill. (Avocado), *Punica granatum* L. (Pomegranate), *Musa acuminata*, *Musa sapientum*, *Musa paradisiaca* (Banana), *Psidium guajava* L. (Guava), *Genipa americana* L. (Jagua), *Citrus maxima* - *Citrus grandis* L. (Pomelo), *Citrus paradisi* Macfad (Grapefruit), *Citrus sinensis* (Orange), *Citrus macroptera* (Wild Orange), *Citrus reticulata* (Mandarine), *Citrus limon* (Lemon), *Citrus medica* L. (Citron), *Melicoccus bijugatus* Jacq. (Spanish Lime), *Euphoria longana* Lam. (Longan), *Nephelium lappaceum* L. (Rambutan) and *Chrysophyllum cainito* L. (Star Apple) belonging to 15 different families. The peels of tropical fruits are a potential source of various bioactive compounds, such as polyphenols, flavonoids, carotenoids, terpenoids, alkaloids, among others. They also possess properties and antioxidant, antiviral, anticancer and antidiabetic activities that could be used to create new therapeutic agents. This review shows the pharmacological potential of tropical fruit peels and also pretends to be a basis to encourage cultivation and conservation of these rare (non-commercial) fruit species and promote their integral and sustainable use.

**Key words:** Fruit peels, Neotropical fruit shells, Biological activities, Pharmacological potential.

### Introduction

Natural products for the development of new pharmacological agents have had a long-standing history in the annals of medicine (for a thorough review on the matter see Newman & Cragg, 2020). Fortuitous observation and chance have undoubtedly played a vital role in this process, though. Other important and more controlled approaches include ethnobotanical bioprospecting and chemotaxonomics. Both strategies have relied heavily on the analysis of leaves, bark and the inflorescences of plants for their success. However, new studies have recently shed light on the use of fruits and their peels for the same purposes (Vílchez *et al.*, 2011; Chaouch & Benvenuti, 2020). Considering the above, the present review has the aim to provide a general overview of the pharmacological potential of the peels from eighteen tropical fruits: *Actinidia deliciosa* (Kiwifruit/ Actinidiaceae), *Mangifera indica* L. (Mango/ Anacardiaceae), *Annona squamosa* L. (Sugar-apple/ Annonaceae), *Annona reticulata* L. (Custard apple/ Annonaceae), *Ananas comosus* (Pineapple/ Bromeliaceae), *Canarium odontophyllum* Miq. (Dabai/ Burceraceae), *Carica papaya* L. (Papaya/ Caricaceae), *Cucumis sativus* L. (Cucumber/ Cucurbitaceae), *Persea americana* Mill. (Avocado/Lauraceae), *Punica granatum*

L. (Pomegranate/ Lythraceae), *Musa acuminata*, *Musa sapientum*, *Musa paradisiaca* (Banana/Musaceae), *Psidium guajava* L. (Guava/Myrtaceae), *Genipa americana* L. (Jagua/ Rubiaceae), *Citrus maxima* - *Citrus grandis* L., *Citrus paradisi* Macfad, *Citrus sinensis*, *Citrus macroptera*, *Citrus reticulata*, *Citrus limon*, *Citrus medica* L. (Pomelo, Grapefruit, Orange, Wild Orange, Mandarine, Lemon and Citron/Rutaceae), *Melicoccus bijugatus* Jacq. (Spanish Lime/ Sapindaceae), *Euphoria longana* Lam. (Longan/ Sapindaceae), *Nephelium lappaceum* L. (Rambutan/ Sapindaceae) and *Chrysophyllum cainito* L. (Star Apple/ Sapotaceae). Hopefully with the information presented here, future studies can delve deeper into the pharmacological potential of tropical fruits and their peels. We also wish to promote the sustainable use of the listed species.

### Materials and Methods

Between October 2017 and August 2021, an exploration of scientific literature was conducted through the search engines: Pubmed, Science Direct, Springer Link, Science.Science.gov, SciELO, Dialnet and Google Scholar to collect publicly available information on the bioactive compounds and biological activities of tropical fruit shells.

Search terms included scientific and common names of each fruit species, along with terms such as tissues, peel, shell, skin, rind, husk, epicarp, exocarp, peels review, traditional medicine, chemical composition, phytochemical study, pharmacological study, *In vivo* study, *In vitro* study, bioactivities and pharmacological properties, as well as the types of chemical compounds and kinds of bioactivities of each one. The reference lists of scientific articles were cross-checked and only relevant works were included. The search for data was limited to reports of fruits from tropical places and included articles and books published between 1987 and 2021. The summaries presented in the tables were taken from the data of different authors. Chemical composition and bioactivities tables were also included.

## Results

**Fruits of the neotropics:** The fruit species present in the tropical regions of the Americas (i.e. the Neotropics) has been estimated at more than 2,000. From these, 259 are commercially grown, with 10% of such lands being exclusively destined for the cultivation of just 64 species. These same species are also responsible for generating 14% of the total revenue derived from local agriculture (Segura-Ledesma *et al.*, 2009; Blancke, 2016). Introduced species are commercially important as well, and most of them have their origins either in the old world or in the Far East. Today, such plants exist in a naturalized state; however, phenotype selection and persistent cultivation have given rise to new varieties vastly different from those present in their ancestral habitats. The Neotropics, therefore, represent an exceptional opportunity for the innovative use of novel biological resources in diverse sectors of the economy, including those that pertain to the pharmacological industry alone. In this sense, tropical ecosystems are particularly rich in plant species of special interest to such sectors (Valli *et al.*, 2012; Blancke, 2016). This comes in addition to their prior role as reservoirs of staple food crops. Already, many tropical food products (including certain fruits) are known to possess a relatively high nutritional content (Kehlenbeck *et al.*, 2013; Kumoro *et al.*, 2020), a fact that has helped to increase their demand in many international markets.

**Botanical description:** The eighteen species presented here are tropical fruits with different botanical characteristics and are described in Table 1.

Use in Traditional medicine of tropical fruit peels  
Diverses tissues like leaves, flowers and bark of Kiwifruit, Mango, Sugar-apple, Custard apple, Pineapple, Papaya, Cucumber, Avocado, Pomegranate, Banana, Guava, Jagua, Pomelo, Grapefruit, Orange, Wild Orange, Mandarine, Lemon, Citron, Spanish Lime, Longan, Rambutan and Star Apple are used in traditional medicine to treat various diseases such as asthma, cancer, diabetes, rheumatoid arthritis, syphilis, gonorrhea, dengue, fever, hypertension, among others, (Morton, 1987; Rangkadilok *et al.*, 2005; Ojewole & Amabeoku, 2006; Khare, 2007;

Yasir *et al.*, 2010; Da Conceicao *et al.*, 2011; Gajalakshmi *et al.*, 2011; Sanda *et al.*, 2011; Bystrom, 2012; Rajendra *et al.*, 2012; Anjum *et al.*, 2013; Mukherjee *et al.*, 2013; Chavan *et al.*, 2014; Jiménez *et al.*, 2014; Shailajan & Gurjar, 2014; Yogiraj *et al.*, 2014. Haque *et al.*, 2015; Sangeetha & Jayaprakash, 2015; Singh-Pal *et al.*, 2015; Chaudhari *et al.*, 2016; Chawla *et al.*, 2016; Mishra *et al.*, 2017; Sukmandari *et al.*, 2017; Faustino *et al.*, 2019). On the other hand, only pomegranates, banannas and oranges have reports on the use of their peels as a remedies to protect against skin infections (Manodeep *et al.*, 2012), dysentery, diarrhea and ulcers (Lavanya *et al.*, 2016) and anthelmintic (Mabberley, 2004), respectively. Concerning the buah dabai (*Canarium odontophyllum* Miq.), to the best of our knowledge, there are currently no reports that mention its use in traditional medicine.

**Chemical constituents and pharmacological activity of tropical fruit peels:** For many years, botanical bioprospecting relied almost exclusively on the use of leaves, bark and the inflorescences of plants as sources for clinically relevant bioactive compounds, with other organs or structures being analyzed relatively recently (Vílchez *et al.*, 2011; Majeed *et al.*, 2019; Fierascu *et al.*, 2020). An important example of the latter is the study completed by Deng *et al.*, (2012) concerning the antioxidant potency and phenolic content present in the residues (i.e. peels and seeds) of 50 different types of fruits. Included in this analysis were 19 taxonomic families, chief among them the Annonaceae, Sapindaceae, Myrtaceae, Rutaceae, and Cactaceae. The authors concluded that the wastes of certain fruit species do contain relatively high levels of antioxidants and phenolics, the most abundant being the compounds catechin, cyanidin 3-glucoside, epicatechin, galangin, gallic acid, homogentisic acid, kaempferol and chlorogenic acid. The wastes of *A. squamosa* in particular (commonly known as sweetsop) were especially high in antioxidant activity, and it was determined that such residues—like those of other species—could be used as sources of inexpensive and readily available bioactive compounds of potential interest to the pharmaceutical industry.

Similar studies (namely phytochemical and pharmacological analyses) have also been conducted on the peels of other tropical fruits. What follows is the result of a bibliographical analysis of several fruit species with proven pharmacological properties. They appear organized according to their botanical family, with salient features summarized in Tables 2 and 3.

**Family actinidiaceae:** Phenols, tannins, and alkaloids have been detected in the peels of kiwifruit. Pharmacological studies also report the presence of cytotoxic, cytotoxicity-modulating, antibacterial, antioxidant, and anti-HIV activities (Motohashi *et al.*, 2001; Lucas *et al.*, 2003; Duda-Chodak & Tarko, 2007; Indhumathi *et al.*, 2011; Singh-Pal *et al.*, 2015; Alim *et al.*, 2019).

Table 1. Geographical origin and description of selected fruits.

Fruit data/ Number, scientific name, common name and family	Geographical origin	Size	Weight	Shape	Peel
1. <i>Actinidia deliçiosa</i> Kiwifruit Actinidiaceae <sup>a</sup>	China	5 to 6.3 cm	0.03-0.15 kg	Ovoid or oblong	Bright-green flesh and a dull-brown skin, covered by short epidermal hairs known as trichomes. Thick and leathery, generally smooth, waxy, and aromatic with light green, dark green, light yellow, yellow, yellow-orange, reddish-pink, bright red, dark red, or even purple skin when mature.
2. <i>Mangifera indica</i> L. Mangoes Anacardiaceae <sup>b</sup>	India	6.2 to 25 cm	0.18 to 2.3 kg	Round, oval, ovoid-oblong, or kidney-shaped	The exterior is marked by polygonal tubercles which correspond to the fused carpels from which the fruit is formed. The ripe fruit is light yellowish-green or purple and the exterior readily separates along the lines between the tubercles.
3. <i>Annona squamosa</i> L. Sugar-apple Annonaceae <sup>c</sup>	Probably from either the tropical regions of Central America or the Antilles	5 to 12 cm (diameter)	0.200 to 0.80 kg	Globose to oviform and almost heart-like in shape	The skin, thin but tough, may be yellow or brownish when ripe, with a pink, reddish or brownish-red bluish, and faintly, moderately, or distinctly reticulated.
4. <i>Annona reticulata</i> L. Custard apple Annonaceae <sup>d</sup>	Central America and the Caribbean	8 to 15 cm in diameter	Up to 1 kg	Heart-shaped or spherical	The hard and waxy shell is formed by interlocking hexagonal units and can be a dark green, yellow, orange-yellow, or reddish color when the fruit is ripe.
5. <i>Ananas comosus</i> Pineapple Bromeliaceae <sup>e</sup>	South America	30 cm or more	0.5 to 4.5 kg or more	Oval to cylindrical-shaped	Drupe with thin skin, mostly white when immature, and bluish-black when ripe.
6. <i>Canarium odontophyllum</i> Miq. Dabai Burceraceae <sup>f</sup>	Malaysia	3 to 4 cm	0.010 to 0.013 kg	Oval to ellipsoid	Very thin skin which is green when immature, and yellow, orange, or red-orange when ripe.
7. <i>Carica papaya</i> L. Papaya Caricaceae <sup>g</sup>	Perhaps in Southern Mexico	5 to 50 cm or more	0.2 to 9 kg	Large oval-shaped berry	Smooth, dark-green skin
8. <i>Cucumis sativus</i> L. Cucumber Cucurbitaceae <sup>h</sup>	Native to Southern Asia	10 to up to 50 cm	0.050 to 4 kg	Elongated, ellipsoidal, convex, and laterally striated	The exocarp (i.e. rind or skin) is variable in both thickness and texture, and can be of a green, black, purple, or even reddish color at maturity.
9. <i>Persea americana</i> Mill. Avocado Lauraceae <sup>i</sup>	Central America	7-20 cm long and 7-10 cm in diameter	0.100 and 1 kg	Usually spherical or piriform, the pointed calyx	The tough, leathery skin or rind is typically yellow overlaid with light or deep pink or rich red.
10. <i>Punica granatum</i> L. Pomegranate Lythraceae <sup>j</sup>	Northern Africa and the Caucasian	5 to 12.7 cm	0.250 to 0.300 kg	Nearly round and crowned by the pointed calyx	The skin is thin and tender to thick and leathery, yellow, green, or red in color.
11. <i>Musa acuminata</i> , <i>Musa sapientum</i> , <i>Musa paradisiaca</i> Banana Musaceae <sup>k</sup>	Southeast Asia	3-40 cm long, and 2-8 cm in diameter.	Average 0.125 kg	Elongated, curved, and more or less round in cross section and triangular ovary.	Smooth or rough epicarp and the skin turns a characteristic yellow (or reddish-yellow) color at ripe stage.
12. <i>Psidium guajava</i> L. Guava Myrtaceae <sup>l</sup>	Probably from southern Mexico to South America	5 to 10 cm long	Up to 0.500 kg	Round, pyriform or ovoid berry	Thin leathery skin of greenish-brown or reddish color.
13. <i>Genipa americana</i> L. Jagua Rubiaceae <sup>m</sup>	Native to the tropical forests of North and South America	8-10 cm long and 6-7 cm in diameter	0.200 to 0.400 kg	Elliptic or rounded-oval berry, tapering briefly at the stem end	Tough outer skin or epicarp, commonly known as flavedo that can be green, yellow, orange, or even reddish in color.
14. <i>Citrus maxima</i> - <i>Citrus grandis</i> L., <i>Citrus paradisi</i> Macfad., <i>Citrus sinensis</i> , <i>Citrus macroptera</i> , <i>Citrus reticulata</i> , <i>Citrus limon</i> , <i>Citrus medica</i> L. Genus <i>Citrus</i> (Pomelo, Grapefruit, Orange, Wild Orange, Mandarin, Lemon and citrón) Rutaceae <sup>n</sup>	Probably Southeast Asia.	Fruit of the genus <i>Citrus</i> vary greatly in size and range from about 3 cm (kumquats) to more than 30 cm in diameter (pummelos)	0.01 to 2 kg	Round to oblong	Thin, leathery, brittle skin that is green to yellowish when mature.
15. <i>Melicoccus bijugatus</i> Jacq. Spanish Lime Sapindaceae <sup>o</sup>	Northern regions of South America	2.5 to 4 cm in diameter	0.009 to 0.022 kg	Round or ovoid-shaped drupes	The peel is tan, thin, and leathery with tiny hairs
16. <i>Euphoria longana</i> Lam. Longan Sapindaceae <sup>p</sup>	Southern China	1.2-2.5 cm in diameter	0.006 to 0.019 kg	Conical, heart-shaped or spherical	Thick and leathery skin covered entirely by (flexible) spines. When ripe, the skin turns red in color.
17. <i>Nephelium lappaceum</i> L. Rambutan Sapindaceae <sup>q</sup>	Indonesia and Malaysia	Up to 7 x 5 cm	0.020 to 0.095 kg	drupe. Oval to spherical in shape	Smooth skin of a purple-red, dark purple, or even pale green color.
18. <i>Chrysophyllum cainito</i> L. Star Apple Sapotaceae <sup>r</sup>	Greater Antilles and the West Indies	5 to 10 cm in diameter	Average 0.129 kg	Subglobose berry	

<sup>a</sup>Lucas et al., 2003; Motohashi et al., 2001; Chawla et al., 2016. <sup>b</sup>Jahurul et al., 2015; Morton, 1987; Chel-Guerrero et al., 2018a. <sup>c</sup>Chavan et al., 2014. <sup>d</sup>Rashad et al., 2015; Morton, 1987. <sup>e</sup>Basri et al., 2014a; 2014b; Yograj et al., 2014. <sup>f</sup>Mukherjee et al., 2013; Yasir et al., 2010. <sup>g</sup>Manodeep et al., 2012; Haque et al., 2015. <sup>h</sup>Ross, 2001; Dahham et al., 2015. <sup>i</sup>Morton, 1987; Mishra et al., 2017. <sup>j</sup>Bazilio-Omena et al., 2012. <sup>k</sup>Mabberley, 2004. <sup>l</sup>Morton, 1987; Crane & Balardi, 2016. <sup>m</sup>Rangkadilok et al., 2005. <sup>n</sup>Sukmandari et al., 2017. <sup>o</sup>Morton, 1987; Chel-Guerrero et al., 2018a.

**Table 2. Chemical components in the peels of several tropical fruits.**

Compound	Fruit																	
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18
Polyphenols	x	x	x	x		x	x	x	x	x	x	x		x	X	x	x	x
Flavonoids		x	x	x	x	x	x	x	x	x	x	x	x	x	X	x		x
Anthocyanins		x				x				x	x							
Anthocyanidins					x							x						
Carotenoids		x				x	x		x		x			x				
Coumarins				x						x			x	x				
Terpenoids				x	x		x	x	x	x			x	x				x
Saponins				x	x		x	x		x	x	x		x				x
Tannins	x		x		x		x		x	x	x	x		x		x	x	
Alcaloids	x			x	x		x	x		x				x		x	x	
Esteroids				x	x		x	x	x	x	x			x				
Esterols				x	x		x			x	x		x					
Anthraquinones											x		x					
Anthrones													x					
Glycosides										x	x							
Carbohydrates											x			x				

The number in the columns corresponds to the fruit with the same number in Table 1. Source: Documented chemical studies; specific references are listed in the main text, in the sections corresponding to each family

**Table 3. Pharmacological activities reported in the peels of several tropical fruits.**

Activity	Fruit																	
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18
Antioxidant	x	x	x	x	x	x	x	x	x	x	x	x	x	x	x	x	x	x
Antiviral		x				x												x
Anti-immunomodulatory										x								x
Anticancer								x		x	x	x		x				x
Cytotoxic or antiproliferative		x	x	x					x	x	x							
Antitumor				x														
Modulated of the cytotoxicity		x																
Antidiabetic						x	x			x		x		x				x
Hypoglycemic										x						x		
Anti-inflammatory			x			x				x	x							
Analgesic										x								
Anti-arthritis									x									x
Antiulceric												x						
Antibacterial		x				x		x	x	x	x			x				x
Antimicrobial				x	x			x		x	x	x	x	x				x
Anti-allergic														x				
Antifertility				x						x								
Antiparasitic						x												
Antifungal						x								x				
Diuretic												x						
$\alpha$ -amilase and $\alpha$ -glucosidase inhibiting																		x
Anti-diarrheal													x					
Hipolipidemic												x		x				
Antihyperlipidemic																		x
Antihypercolesterolemic														x				x
Anti-alzheimer														x				
Anticholinesterase							x							x				
Anti-aging																		x
Anti-thyphoid														x				
Antihelminthic														x				
Estrogenic														x				
Antidepressant														x				
Antinociceptive																x		
Hepatoprotective										x								
Gastroprotective							x			x								
Cardioprotective							x											
Radioprotective										x								
Anti osteoporotic										x								
Protective effect against prostatitis										x								

The number in the columns corresponds to the fruit with the same number in Table 1. Source: Documented chemical studies; specific references are listed in the main text, in the sections corresponding to each family

**Family Anacardiaceae:** Phenols, flavonoids, anthocyanins and carotenoids have often been detected in the peels of *M. indica*. Numerous pharmacological studies also report the presence of antioxidant, anti-inflammatory and antiproliferative activities (Knödler *et al.*, 2008; Masibo & He, 2009; Kim *et al.*, 2010; Ali *et al.*, 2012; Solidum, 2012; Ahmad *et al.*, 2015; Jahurul *et al.*, 2015; Masud-Parvez, 2016; Fierascu *et al.*, 2020).

**The family annonaceae:** Several bioactive compounds including polyphenolics and flavonoids have been detected in the peels of *A. squamosa*. Pharmacological studies also report the presence of antioxidant, antimicrobial, cytotoxic, and antiviral activities, as well as modulation of acetylcholinesterase activity (Sharma *et al.*, 2013., Kaladhar *et al.*, 2015; Chel-Guerrero *et al.*, 2018a, 2018b; Alcántar-Ramírez *et al.*, 2018; Shehata *et al.*, 2021). Phenols, flavonoids, alkaloids, cholesterol, coumarins, tannins, saponins, steroids, and terpenoids have likewise been detected in the peels of *A. reticulata*, as have antioxidant, cytotoxic, antiviral, anti-inflammatory and antimicrobial activities (Kaladhar *et al.*, 2015; Chel-Guerrero *et al.*, 2018a, 2018b).

**The family bromeliaceae:** Alkaloids, flavonoids, saponins, tannins, steroids, triterpenes and phytosterols have often been detected in the peels of *A. comosus*. Pharmacological studies also report the presence of antioxidant, antiviral, anti-inflammatory, antibacterial, antiparasitic and antifungal activities (Azizan *et al.*, 2020; Rashad *et al.*, 2015; Lawal, 2013).

**The family burceraceae:** Phenols, flavonoids, anthocyanins, anthocyanidins and carotenoids have all been detected in the peels of *C. odontophyllum*. Pharmacological studies also report the presence of antioxidant, cardioprotective, gastroprotective, antidiabetic and anticholinesterase activity (Xuan & Azlana, 2016, Salahuddin-Mirfat *et al.*, 2018).

**The family caricaceae:** Phenols, flavonoids, terpenoids, tannins, alkaloids, saponins, carotenoids, steroids and sterols have been detected in the peels of *C. papaya* (Mendes dos Santos *et al.*, 2014; Sihombing *et al.*, 2015; Dada *et al.*, 2016; Siddique *et al.*, 2017). Pharmacological studies also report antioxidant, anticancer, antimicrobial and antibacterial activities (Dada *et al.*, 2016; Siddique *et al.*, 2017; Undugodage-Dulanjali & Kanchana-Perera, 2018; Villacís-Chiriboga, 2020).

**The family cucurbitaceae:** Phenols, flavonoids, alkaloids, saponins, terpenoids and steroids have all been detected in the peels of *C. sativus* (Mukherjee *et al.*, 2013; Foong *et al.*, 2015; Sonia *et al.*, 2016; John *et al.*, 2018). Pharmacological studies also report antioxidant, cytotoxic, antidiabetic, antimicrobial and antibacterial activities (Dixit & Kar, 2010; Mukherjee *et al.*, 2013; Foong *et al.*, 2015; John *et al.*, 2018).

**The family lauraceae:** Phenols, flavonoids, carotenoids, terpenoids, tannins and steroids have all been detected in the peels of *P. americana* (Vinha *et al.*, 2013; Putta and

Kilari, 2015; Rodríguez-López *et al.*, 2015; Smitha-Grace *et al.*, 2015; Roser *et al.*, 2019; Salazar-López, *et al.*, 2020). Pharmacological studies also report antioxidant, cytotoxic, anti-inflammatory, antimicrobial, antiarthritic and antibacterial activities (Rodríguez-Carpena *et al.*, 2011, Vinha *et al.*, 2013, Enwa, 2016, Smitha *et al.*, 2017; Tremocoldi *et al.*, 2018; Roser *et al.*, 2019; Figueroa *et al.*, 2021).

**The family lythraceae:** In the peels of *P. granatum*, tannins, saponins, quinones, terpenoids, steroids, flavonoids, phenols, alkaloids, glycosides, cardiac glycosides, coumarins and anthocyanins have been determined (García *et al.*, 2021; Chen *et al.*, 2020; Foujdar *et al.*, 2020; Magangana *et al.*, 2020; Machado-Brêtas *et al.*, 2020; Sangeetha & Jayaprakash, 2015). Pharmacological studies also report antioxidant activity, anticancer, antidiabetic, cytotoxic, analgesic, immunomodulatory, antifertility, anti-inflammatory, gastrointestinal protection activities, with uterine stimulant effect, protective effect against prostatitis, hypoglycemic activity, radioprotective, gastroprotective, antibacterial, antityrosinase, antimicrobial, antihyperlipidemic, hepatoprotective, antiproliferative and antiosteoporotic (Fawole *et al.*, 2012; Ismail *et al.*, 2012; Manodeep *et al.*, 2012; Howel, 2013; Sreekumar *et al.*, 2014; Haque *et al.*, 2015; Mphahlele *et al.*, 2016; John *et al.*, 2017; Benguiar *et al.*, 2020; Chen *et al.*, 2020; Foujdar *et al.*, 2020; Karim *et al.*, 2020; Magangana *et al.*, 2020; Peršurić *et al.*, 2020; More & Aria, 2021).

**The family musaceae:** Different bioactive compounds including phenols, flavonoids, anthocyanins, carotenoids, carbohydrates, tannins, saponins, anthraquinones, steroids, glycosides, sterols and terpenoids have been detected in the peels of *Musa* spp. Pharmacological studies also report antioxidant, diuretic, anticancer, antiulceric, antiviral, antibacterial and antimicrobial activities as well as mutagenic effect and inhibition of the cholesterol crystallization (Swanson *et al.*, 2010; Imam *et al.*, 2011; Ehiowemwenguan *et al.*, 2014; Dahham *et al.*, 2015; Kibria *et al.*, 2019; Oyeyinka & Jide-Afolayan, 2020a, 2020b).

**The family myrtaceae:** Phenols, tannins, flavonoids, anthocyanidins and saponins have all been detected in the peels of *P. guajava*. Several pharmacological studies also report the presence of antioxidant, antimicrobial, anticancer, hypoglycemic, antidiabetic, antidiarrheal and hypolipidemic activities (Thaiponga *et al.*, 2006, Rai *et al.*, 2009; Rai *et al.*, 2010; Barbalho *et al.*, 2012; Wang *et al.*, 2014; Abdelmalek *et al.*, 2016; Issa *et al.*, 2016; Gupta *et al.*, 2018; Liu *et al.*, 2018; Angulo-López *et al.*, 2021; Ganoza-Yupanqui *et al.*, 2021).

**The family rubiaceae:** Flavonoids, anthraquinones, anthrones, coumarins, terpenoids and sterols have all been detected in the peels of *G. americana*. Antioxidant activity has also been reported (Bazílio-Omena *et al.*, 2012; Náthia-Neves *et al.*, 2017).

**The family rutaceae (Syn. aurantiaceae):** Phenols, flavonoids, tannins, carotenoids, terpenoids, steroids, carbohydrates, coumarins, saponins and alkaloids have often been detected in the peels of pomelos, oranges, grapefruits, mandarins, lemons and acid limes. Pharmacological studies also report the presence of antioxidant, anti-cancer, anti-Alzheimer, antimicrobial, antibacterial, antifungal, antithyroid, antidiabetic, hypoglycemic, hypolipidemic, antihypercholesterolemic, anti-allergic, lipolytic, anthelmintic, estrogenic, antidepressant and anticholinesterase activities (Panara *et al.*, 2012; Lagha-Benamrouche and Madani, 2013; Oikeh *et al.*, 2013, Kumar *et al.*, 2014; Madhuri *et al.*, 2014; Natesan *et al.*, 2014; Vijayalakshmi and Radha, 2014; Muhtadi *et al.*, 2015; Aktar and Foyzum, 2017; Vijayalakshmi and Radha, 2016; Amutha *et al.*, 2017; Nongalleima *et al.*, 2017; Safdar *et al.*, 2017; Rafiq *et al.*, 2018; Montero-Calderon *et al.*, 2019; Klimek-Szczykutowicz *et al.*, 2020; Santoshkumar-Jayagoudar *et al.*, 2020; Singh *et al.*, 2020; Czech *et al.*, 2021; Nieto *et al.*, 2021; Taktak *et al.*, 2021).

**The family sapindaceae:** Both phenols and flavonoids have been detected in the peels of *M. bijugatus* and pharmacological studies report antioxidant activity and modulation of acetylcholinesterase activity (Chel-Guerrero *et al.*, 2018a). In *E. longana*, phenols, flavonoids, tannins and alkaloids are present in its peel (Rangkadilok *et al.*, 2005, Ripaa *et al.*, 2015), and hypoglycemic activities have been described (Li *et al.*, 2015, Ripaa *et al.*, 2015). The peels of *N. lappaceum* also contain several bioactive compounds, including phenols, tannins, terpenoids, saponins, and alkaloids (Thitilertdech *et al.*, 2010; Fila *et al.*, 2012; Rakariyatham *et al.*, 2020). In this case, pharmacological studies reveal antioxidant, anti-aging, anticancer, antidiabetic, antihypercholesterolemic, anti-inflammatory, antihyperlipidemic, anti-immunomodulatory, antidepressant, antinociceptive, antimicrobial, antibacterial and antiviral activities. Protective effects against arthritis, as well as an inhibitory effect against  $\alpha$ -amylase and  $\alpha$ -glucosidase are likewise mentioned (Sukmandari *et al.*, 2017; Phuong *et al.*, 2020; Rakariyatham *et al.*, 2020).

**The family sapotaceae:** Phenolic compounds and flavonoids have been detected in the peels of *C. cainito*. Recent studies also report antioxidant, cytotoxicity and antiviral activities (Moo-Huchin *et al.*, 2015; Chel-Guerrero *et al.*, 2018a, 2018b; Alcántar-Ramírez *et al.*, 2018; Doan & Le, 2020).

Thus, it is clear that fruit peels will often harbor several pharmacologically-relevant bioactive compounds and should not simply be regarded as polluting waste. Doing so can result in several unnecessary expenditures as the cost of eliminating such wastes may be rather high, especially for producers. If, on the other hand, fruit peels are separated from polluting wastes and commercialized independently for use in other products or applications (e.g. pharmaceutical ones), the cultivars from which they originate could benefit from the increased demand. An interesting case in point (though one that does not involve peels) concerns the pharmaceutical product known commercially in Mexico as QG5. As incredibly as it may

seem, guava leaves are now priced at a similar value to that of actual fruits – at least in certain regions of Mexico (Rivera-Arce *et al.*, 2003).

Moreover, the confirmation of both medically-relevant and economically-important species would guide the conservation and sustainable use of the plants involved, as well as those of the ecosystems from which they originate.

**Conclusion and future prospects:** The work done by many researchers and reviewed here shows the tremendous potential of tropical fruit peels as sources of beneficial bioactive compounds, which can be used to develop new therapeutic agents. This in turn, confirm the importance of further research into the pharmacological potential of these tissues. Nowadays, few studies correlate chemical composition with pharmacological activity, which makes bioprospecting and chemical screening more relevant. Generally speaking, the issues that still need to be resolved in the near future are: 1) Obtaining and analyzing the metabolomic profile of tropical fruit peels; 2) Identifying the specific types of compounds present in the peels (i.e. not just the total content of their respective molecular classes); 3) Empirically correlating pharmacological activity with chemical composition; 4) Identifying the active compounds responsible for such activities; 5) Establishing environmentally-friendly methods of extraction and purification; 6) Standardizing the composition of extracts; and 7) Reaching Phase I of drug development. Such challenges must be adequately addressed by future studies aiming to characterize the chemical and pharmacological potential of tropical fruit peels. Ideally, this should also include molecular labeling data to permit the creation of biological parameters that are specific for the species being studied.

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