Inhibitory Effects of Edible and Medicinal Plant Extracts on the Enzymatic Activity of Pancreatic Lipase

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Received December 21st, 2022; Accepted March 17th, 2023.

DOI: http://dx.doi.org/10.29356/jmcs.v67i3.2004

Abstract. Plants with a strong activity to reduce the digestion of lipids from the diet are a possible way to prevent and combat obesity. This study evaluated 37 ethanol extracts of plants, some edible, medicinal, or belonging to a family that has the inhibitory activity of pancreatic lipase, aimed at looking for a new anti-obesity agent. Inhibition of pancreatic lipase (PL) was measured in vitro and in vivo assay. The plasma triacylglycerol levels after 1, 2, and 3 h in fasted male Wistar rats fed, by oral administration, with a lipid emulsion were measured. The antioxidant activities were evaluated using DPPH (1,1-diphenyl-2-picrylhydrazyl radical scavenging activity), FRAP (ferric reducing antioxidant power) and ABTS (2,2'-azino-bis(3-ethylbenzthiazoline-6-sulphonic acid), and the total phenol and flavonoid content were determined. Nine plants exhibited low inhibition (˂41 %), nine showed medium inhibition (41-50 %), eight demonstrated high inhibition (51-60%) and 11 plants had the highest inhibition (≥ 61 %). Hibiscus rosa-sinensis, dried leaves extract displayed the highest inhibitory activity of pancreatic lipase 71.90 % at 400 µg/mL, with a dose-dependent inhibition against PL similar to orlistat. The kinetic study showed uncompetitive inhibition. Moreover, H. rosa sinensis extract also reduced the elevation of plasma triacylglycerol levels after lipid emulsion administration similar as orlistat did. H. rosa-sinensis dried leaves had higher antioxidant activity and total phenolic compounds than fresh leaves. H. rosa-sinensis presented the strongest anti-lipase activity and could be used as an anti-obesogenic agent or as a food additive to reduce the absorption of fats from the diet.

Keywords: Obesity; lipase; triacylglycerol; inhibition; antioxidant capacity.

Resumen. Las plantas con una fuerte actividad para reducir la digestión de los lípidos de la dieta son una posible forma de prevenir y combatir la obesidad. Este estudio evaluó 37 extractos etanólicos de plantas, algunas comestibles, medicinales o pertenecientes a una familia que tienen una actividad inhibidora de la lipasa pancreática, con el objetivo de buscar un nuevo agente anti-obesogénico. Se midieron la inhibición de la lipasa pancreática (PL) in vitro e in vivo, se midieron los niveles de triacilglicerol en plasma 1,2 y 3 h después de la administración oral de una emulsión lipídica a ratas Wistar machos en ayunas. Las actividades antioxidantes se evaluaron utilizando DPPH (1,1-diphenyl-2-picrylhydrazyl actividad atrapadora de radical), FRAP (poder antioxidante reductor del fierro) y ABTS acido (2,2'-azino-bis(3-ethylbenzthiazoline-6-sulphonico), también se midió el contenido total de fenoles y flavonoides. Nueve plantas exhibieron una inhibición baja (<41 %), otras nueve con una inhibición media (41-50 %), ocho demostraron una inhibición alta (51-60 %) y once plantas mostraron la inhibición más alta (≥ 61 %). El extracto de hojas secas de Hibiscus rosa-sinensis, mostró la mayor actividad inhibidora de la PL con un 71.90 % a 400 µg/mL, con una inhibición dependiente de la dosis contra PL similar a orlstat. El estudio cinético mostró una inhibición acompetitiva. Además, el extracto de H. rosa-sinensis también redujo la elevación de los niveles de triacilglicerol en plasma después de la administración de una emulsión de lipidos similar a como lo hace orlistat. Las hojas secas de H. rosea-sinensis tuvieron mayor actividad antioxidante y cantidad de compuestos.
fenólicos totales que las hojas frescas. *H. rosa-sinensis* presentó la mayor actividad anti-lipasa y podría usarse como un agente anti-obesogénico o como un aditivo alimentario para reducir la absorción de grasas de la dieta. **Palabras clave:** Obesidad; lipasa; triacilglicerol; inhibición; capacidad antioxidante.

**Introduction**

Some strategies to prevent obesity are control food intake or alter lipids metabolism by inhibiting fat absorption [1]. Pancreatic lipase (triacylglycerol acylhydrolase) is the most important enzyme in the digestion of 50-70 % of the fat diet and, it is responsible for the hydrolysis of triacylglycerols (triglycerides) to monoacylglycerols and free fatty acids [2].

Tetrahydrolipstatin (Orlistat TM), a derivative of lipstatin obtained from *Streptomyces toxytricini*, is the only pancreatic lipase inhibitor currently approved for the treatment of long-term obesity. Its use can result in up to 10 % weight loss when used in combination with diet and physical activity. However, this drug can cause adverse liver effects (cholelithiasis, colostatichepatitis, and sub-acute liver failure) and troublesome gastrointestinal symptoms [3]. The inhibition of pancreatic lipase in many plants has been studied for its anti-obesogenic potential. In addition, the use of natural products to control obesity increased the interest in finding inhibitors of pancreatic lipase without the orlistat side effects [4].

Many medicinal, herbal, and edible plant extracts have been reported with *in vitro* inhibitory activity against pancreatic lipase, mainly in plants of eastern countries e.g., Malaysia, China, Jordan, and Korea [5,6,7]. One of the studies with the highest number of plants (400 plants) searching to inhibit pancreatic lipase activity was reported by Roh and Jung [8]. Mexico is a country with a high biodiversity of plants. However, there are only two studies focused on searching for a plant with a high inhibitory activity of pancreatic lipase. Ramirez *et al.* [9] reported that among 23 medicinal plants used in traditional medicine for diabetes, only two plants had 31.4 % and 27.2 % of lipase inhibitory activity. The second study [10] reported four plants with pancreatic lipase inhibition ≥ 60 % by analyzing a suite of 30 medicinal plants from Oaxaca State. This study aims to analyze 37 ethanol extracts from different parts of plants, easy to obtain, to assess their anti-lipase activity.

**Experimental**

**Plant materials**

Table 1 shows the plants and the parts tested. Some plants are edible, medicinal, or belong to a family that has the inhibitory activity of pancreatic lipase. The parts of the plants used in this study were based on the lack of studies in these tissues. The compound’s composition changes with the use of fresh or dry material and for this reason, both were studied [11,12]. Some species were obtained from a local supermarket and others were acquired from a local greenhouse in Puruándiro, Michoacán, Mexico. The plant identity was confirmed by the taxonomist Patricia Silva of the Department of Botany of the University of Michoacán, and a voucher specimen was deposited in the herbarium of the University of Michoacán (EBUM). The voucher number of *Hibiscus rosa-sinensis* is EBUM240774.

**Plant extractions**

Leaves, fruits, seeds, roots, peel, *Opuntia* sp. cladode, strawberry sepal, and rhizomes were washed with water. Some leaves were dried in the convection oven at a temperature of 40 °C for two weeks. Dried and fresh samples were grounded into fine powder. Three grams of the sample were placed in 30 mL of 98 % ethanol for seven days at room temperature. The solvent was evaporated under 45 °C and low pressure. The dried extracts were dissolved in 1 % of dimethyl sulfoxide (DMSO).
Pancreatic lipase inhibition

The plant extracts obtained at 1 mg/mL were diluted in 1 % DMSO at 50, 100, 200, 300, and 400 μg/mL to perform an inhibition curve. 0.2 mL of the different concentrations were added to 0.1 mL of a pancreatic lipase solution (1 mg/mL); then, Tris-HCl buffer pH 7.4 was added to up to 1 mL and incubated for 15 min at 37 °C. After that, 0.1 mL of p-nitrophenyl-butyrate (100 mM) was added and incubated for 30 min at 37 °C; then, the samples were read at 405 nm. Orlistat was used as a reference control [13]. IC$_{50}$ is the concentration that gives 50 % lipase inhibition.

Antioxidant activity determinations

Radical-scavenging activity (RSA) assay

Free radical scavenging capacity was analyzed by the 2, 2-diphenyl-1 picryl hydrazyl (DPPH) assay according to the method of Hatano et al. [14]. Trolox was used as standard in a range of 25-800 μM.

Ferric reducing antioxidant power (FRAP) assay

The FRAP assay was performed using the method described by Thaipong et al. [15].

Trolox equivalent antioxidant capacity (TEAC)

The assay was performed as Rufino et al. [16] with modifications that reduced the time to obtain the ABTS$^+$ radical cations to two hours. It was produced by mixing an equal volume of 2,2'-Azino-bis(3-ethylbenzothiazoline-6-sulfonic acid) liquid substrate system (ready-to-use) trade Sigma, and 2.4 mM potassium persulfate. The mixture was shaken for 2 h at room temperature in the dark. At this time the absorbance reached a value of 3.0; the ABTS$^+$ solution was diluted with ethanol to get an absorbance of 0.70 ± 0.02 at 734 nm. Fresh ABTS$^+$ solution was prepared for each assay. A sample (0.05 mL) was added to ABTS solution (0.95 mL) after mixing, and the reaction was monitored for 7 minutes. Trolox standard was in the range of 25 - 600 μM. TEAC values were expressed as Trolox equivalents (TE)/g fresh mass.

Total phenolic content

The amount of the total phenolic was determined with the Folin-Ciocalteu reagent using the method of Pripdeevech et al. [17]. Gallic acid was in the range (0.01-0.4 mM). The results were expressed as mg of gallic acid equivalent (GAE)/g fresh mass.

Total flavonoid content

The amount of the total flavonoid was determined using the method of Chang et al. [18]. Rutin acid was used to calculate the standard curve (0.025 to 0.5 mg/mL).

Animals and experimental design

Male Wistar rats from 8-12 weeks old (200–300 g), were housed at a room temperature of 24 °C, with a 12 h light/dark cycle, fed with chow standard diets and water ad libitum. The rats were from the animal house of the Universidad Michoacana de San Nicolás de Hidalgo (UMSNH). The protocol for these experiments was approved by the Institutional Committee for Use of Animals of the UMSNH and followed the recommendations of the regulatory standard for the use of animals issued by the Mexican Ministry of Agriculture in its Federal Regulations for the Use and Care of Animals (NOM062-ZOO-1999). Thirty male Wistar rats that had fasted overnight were divided into five groups: one control group and the rest four groups were administered with oral gavage of 3 mL of a lipid emulsion, (6 mL corn oil, cholic acid 80 mg, 5 mL egg yolk and 6 mL of saline solution), with slight modifications [19]. Groups 2, 3, and 4 were administered orally with dried leaves extract of Hibiscus rosa-sinensis at a concentration of 62.5, 125, and 250 mg/kg respectively, and the 5th group with orlistat (50 mg/kg p.o.), as a positive control. Blood samples were collected from the tail every hour for three hours. Triacylglycerol concentrations were determined using Spinreact Kit.
Statistical Analysis
Pancreatic lipase inhibition activity and antioxidant assay results were expressed as mean ± standard deviation (SD) (n =4). Group differences were analyzed by one-way ANOVA. Values were considered to be indicative of statistical significance *p≤0.05.

Results

Pancreatic lipase (PL) inhibition activity
37 plant extracts were tested at the concentrations of 50, 100, 200, 300, and 400 μg/mL against the pancreatic lipase activity. Nine plants had a low percentage of inhibition of lipase activity < 41 %; nine extracts displayed a medium percentage of inhibition (41-50 %); eight extracts presented high inhibition in the range of range 51-60 %, and 11 extracts produced the highest percentage of lipase inhibition, ≥ 61 %. Table 1 shows that some extract plants had IC₅₀ values up to 400 μg/mL. Although this study worked with a few specimens (37 plant extracts), 19 plants showed high inhibitory pancreatic lipase activity which represents 51.3 % of the plants. Moreover, all plant extracts presented inhibition against pancreatic lipase (Fig. 1).

<table>
<thead>
<tr>
<th>Scientific Name</th>
<th>Family</th>
<th>Part Used</th>
<th>IC₅₀ (µg/mL)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cucumis sativus</td>
<td>Curcubitaceae</td>
<td>P</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Heliopsis longipes</td>
<td>Asteraceae</td>
<td>Rt</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Nelumbo nucifera</td>
<td>Nelumbonaceae</td>
<td>FL</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Rubus sp.</td>
<td>Rosaceae</td>
<td>FR</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Salvia hispánica</td>
<td>Lamiaceae</td>
<td>FL</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Solanum melongena</td>
<td>Solanaceae</td>
<td>FR</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Tamarindus indica</td>
<td>Fabaceae</td>
<td>FR</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Vaccinium corymbosum</td>
<td>Ericaceae</td>
<td>FR</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Vitis sp.</td>
<td>Vitaceae</td>
<td>FL</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Amorphophallus konjac</td>
<td>Araceae</td>
<td>FRh</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Fragaria vesca.</td>
<td>Rosaceae</td>
<td>S</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Heidichium coronarium</td>
<td>Zingiberaceae</td>
<td>FRh</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Hibiscus rosa-sinensis</td>
<td>Malvaceae</td>
<td>FL</td>
<td>&gt;400</td>
</tr>
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<td>Lamiaceae</td>
<td>DL</td>
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<td>Cactaceae</td>
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<td>Araceae</td>
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<td>&gt;400</td>
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<td>Tigridia pavonia</td>
<td>Iridaceae</td>
<td>B</td>
<td>&gt;400</td>
</tr>
<tr>
<td>Vitis sp.</td>
<td>Vitaceae</td>
<td>DL</td>
<td>&gt;400</td>
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<tr>
<td>Annona muricata</td>
<td>Annonaceae</td>
<td>FL</td>
<td>324.37</td>
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<tr>
<td>Callistemon citrinus</td>
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<td>383.46</td>
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<td>249.06</td>
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<tr>
<td>Morus sp.</td>
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<td>DL</td>
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<tr>
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<td>Spinacia oleracea</td>
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<tr>
<td>Amorphophallus konjac</td>
<td>Araceae</td>
<td>DRh</td>
<td>218.21</td>
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<tr>
<td>Annona comosus</td>
<td>Annonaceae</td>
<td>FR</td>
<td>39.53</td>
</tr>
<tr>
<td>Byrsonima crassifolia</td>
<td>Malpighiaceae</td>
<td>FR</td>
<td>163.93</td>
</tr>
<tr>
<td>Capsicum sp.</td>
<td>Solanaceae</td>
<td>FL</td>
<td>278.03</td>
</tr>
</tbody>
</table>
Carica papaya  |  Caricaceae  |  Se  |  44.63
Fragaria sp.  |  Rosaceae  |  FR  |  45.35
Hibiscus rosa-sinensis  |  Malvaceae  |  DL  |  73.82
Hibiscus sabdariffa var. rubra  |  Malvaceae  |  FL  |  44.73
Lavandula  |  Lamiaceae  |  FL  |  3
Passiflora sp.  |  Passifloraceae  |  FL  |  121.93
Syzygium jambos  |  Myrtaceae  |  DL  |  207.27

Plant parts: FL: Fresh Leaves, Se: Seeds, DL: Dry Leaves, FR: Fruits, Rt: Roots, P: Peel, FRh: Fresh Rhizome, DRh: Dry Rhizome, S: Sepal of a flower, C: Cladode, B: Bulb. The results are the mean ± SD (n = 4).


Hibiscus rosa-sinensis (Malvaceae), commonly known as China rose. The dried leaf extract displayed the highest inhibitory activity of pancreatic lipase, 71.90 % at 400 µg/mL, very similar to orlistat 70% at 50 µg/mL. The inhibitory action of H. rosa-sinensis dried leaf extract was dose-dependent in a range of inhibitor concentrations (50-400 µg/mL). The enzyme kinetics of H. rosa-sinensis, using the graphical representation of the Lineweaver-Burk, showed an uncompetitive inhibition with \( K_m \) of 11.11 µM and \( V_{max} \) value of 10 µmol/min, whereas \( K_m \) value of orlistat for pancreatic lipase was of 28.5 µM and the \( V_{max} \) was 16.6 µmol/min, also showed an uncompetitive inhibition (Fig 2).
Fig. 2. Lineweaver-Burk plot of orlistat and *H. rosa-sinensis* against porcine lipase pancreatic at different concentrations of p-nitrophenyl butyrate as substrate.

**Antioxidant activity**

*Hibiscus rosa-sinensis* showed the highest PL inhibitory activity; as a result, this plant was selected to evaluate the antioxidant capacity using the DPPH, FRAP, and ABTS assays. In all assays, *H. rosa-sinensis* dried leaves had higher antioxidant activity and total phenolic compounds than fresh leaves. Conversely, fresh leaves contained a higher number of flavonoids than dried leaves (Table 2).

**Table 2.** Antioxidant activity and the total amount of phenolic and flavonoids of ethanol extracts of *Hibiscus rosa-sinensis* leaves

<table>
<thead>
<tr>
<th></th>
<th>Dry</th>
<th>Fresh</th>
</tr>
</thead>
<tbody>
<tr>
<td>DPPH (μM Trolox/mL)</td>
<td>500.03 ± 1.50</td>
<td>356.76 ± 8.08</td>
</tr>
<tr>
<td>EC₅₀ DPPH</td>
<td>293.23 ± 10.57</td>
<td>380.40 ± 0.38</td>
</tr>
<tr>
<td>FRAP (μM Trolox/mL)</td>
<td>646.62 ± 1.84</td>
<td>432.55 ± 1.55</td>
</tr>
<tr>
<td>EC₅₀ FRAP</td>
<td>175 ± 0.98</td>
<td>360 ± 0.87</td>
</tr>
<tr>
<td>ABTS (μM Trolox/mL)</td>
<td>443.86 ± 2.64</td>
<td>241.66 ± 3.24</td>
</tr>
<tr>
<td>Phenolics GAE (mg/g f.w)</td>
<td>26.91 ± 0.82</td>
<td>13.15 ± 0.80</td>
</tr>
<tr>
<td>Flavonoids Rutin (mg/g f.w)</td>
<td>30.33 ± 0.65</td>
<td>44.57 ± 1.65</td>
</tr>
</tbody>
</table>

DPPH: EC₅₀ (μg/mL) effective concentration at which 50 % of DPPH radicals are scavenged, Trolox 150 μg/mL.
FRAP: EC₅₀ (μg/mL) effective concentration at which the absorbance is 0.5, Trolox 75 μg/mL. (Values are given as mean ± SD (n = 4).

**Oral administration of lipid emulsion in rats**

Fig. 3 shows the reduced absorption of lipids effect of *H. rosa-sinensis* dried leaves in vivo. Plasma triacylglycerol levels were measured after oral administration of lipid emulsion. 125 mg/kg of *H. rosa-sinensis* significantly reduced the postprandial levels of triacylglycerol, after 2 h of oil emulsion administration, similar to orlistat. Meanwhile, triacylglycerol levels increased in the other groups.
Discussion

This study tested 37 plant extracts, with different uses in Mexico, to evaluate their anti-lipase activity. Pancreatic lipase is the main enzyme responsible for the digestion of dietary triacylglycerols [2]. Therefore, the inhibition of enzymes involved in the assimilation of lipids is an important issue to prevent obesity.

11 plants presented inhibition ≥ 61%; in this group, dried rhizome and leaves of Amorphophallus konjac, Hibiscus rosa-sinensis, and Syzygium jambos presented higher inhibitory activity of pancreatic lipase, than the leaves in a fresh state. However, the opposite occurred in Lavandula spp., where the fresh leaves had a higher inhibitory activity of pancreatic lipase than dried leaves. These results could be explained by considering that in some species the drying process can alter the phytoconstituents (phenolics and flavonoids) [20,21]. Moreover, as reported by Dewi et al. [22] in Rubus fraxinifolius Hayata, fresh leaves had a higher content of phenolics and flavonoids and antioxidant capacity than dry leaves, but the α-glucosidase inhibitory activity was lower. However, the opposite occurred in dry leaves. Morus sp. did not show a difference in pancreatic lipase inhibition between fresh and dried leaves; both presented a high anti-lipase activity. Some plant components that have been reported for pancreatic lipase inhibitory activity are flavonoids, lignans, polyphenols, terpenes, and saponins [23].

Hibiscus rosa-sinensis dried leaves extract showed the highest pancreatic lipases inhibitor activity, presenting a dose-response curve. It is a glabrous shrub widely cultivated in the tropics and has several forms with varying colors of flowers. One of the first properties of flowers and leaves was hair growth-promoting. Other uses are to treat arterial hypertension, hypoglycemic, laxative, and oral contraceptives [24]. In some Asia countries and even in Mexico, leaves have been used for dysentery, diarrhea, and as analgesic [25]. In Nigeria, the young leaves of H. rosa-sinensis is eaten [26]. Rios-Chavez et al. [27], reported as the major constituents of H. rosa-sinensis leave extract: fructose (18.60 %), linoleic acid (9,12-octadecadienoic acid 18:2 n-6) (10.36 %), α-linolenic acid (9,12,15-octadecatrienoic acid 18:3 n-3, ALA) (7.67 %) and glycerol (6.l8 %). Seeds of H. rosa-sinensis are rich in unsaturated fatty acids and high content of ALA; however, H. rosa-sinensis leaves have an abundant composition of linoleic and α-linolenic acids. These compounds are important as a dietary component.

Linoleic acid and ALA are unsaturated fatty acids that humans cannot synthesize. These fatty acids are beneficial to health, besides being a precursor of arachidonic acid and having antioxidant properties, and anticarcinogenic and antiartherogenic effects [28].
This is the first study reporting the pancreatic lipase inhibitory activity of *H. rosa-sinensis*. In the genus *Hibiscus*, Buchholz and Melzig [29] showed that the methanol and aqueous extract of *H. sabdariffa* flowers has an inhibitory activity of pancreatic lipase of 100% at a concentration of 2500 μg/mL. However, our study showed 70% of inhibition of lipase activity at a concentration of 400 μg/mL. Moreover, Kumar *et al.* [30] found a hypolipidemic activity of *H. rosa-sinensis* roots ethanol extract, when used in a high-fat diet in rats.

Fig. 2 shows the inhibition mode of PL by *H. rosa-sinensis* at 400 μg/mL using a Lineweaver-Burk plot. The kinetic study revealed that *H. rosa-sinensis* extract decreases the $K_m$ and $V_{max}$ values, implying that it has uncompetitive inhibition of pancreatic lipase activity. In this type of inhibition, the inhibitor binds to the enzyme-substrate complex, and not to the enzyme, preventing the product formation, and making this mode of action useful for enzyme-target drug design [31].

Most of the studies of the antioxidant capacity in *Hibiscus rosa-sinensis* do not mention the flower color; this characteristic, as reported by Patel *et al.* [32] is important. These authors studied the antioxidant properties of five cultivars (red, yellow, orange, pink, and white) of *H. rosa-sinensis*, finding differences in their DPPH radical scavenging activities. The leaf extract of the red cultivar showed $EC_{50}$ 389.2 μg/mL. This finding was consistent with our results that showed an $EC_{50}$ 380.40 and 293.20 μg/mL in fresh and dried leaf extracts of the red cultivar of *H. rosa-sinensis*.

There are few reports about the reducing power of *H. rosa-sinensis*. Patel *et al.* [32] found that the yellow cultivar presented the highest reducing power at 1000 μg/mL and the lowest reducing power was found in the white cultivar. Divya *et al.* [33] reported a high ferric reducing power in methanol extract of leaves using 95 μg/mL. Our study showed $EC_{50}$ values of 360 and 175 μg/mL in fresh and dry leaves of *H. rosa-sinensis*, respectively. Most of the studies use dry tissues because some conditions as storage, time, and temperature could affect antioxidant activity [14,34]. Conversely, there are reports showing that drying alters the phytoconstituents as phenolic, flavonoids, total antioxidant capacity, and some enzymatic activities [18,19,20].

In this study, the total phenolic and flavonoid contents were similar as reported Ghaffar and El-Elaimy [35]. Moreover, Wong *et al.* [36] screening six *Hibiscus* species in Malaysia and found that antioxidant properties change as a function of the environmental conditions. The total phenolic content in the leaves and flowers of *H. rosa-sinensis* was lower than our results. In addition, a study on the leaves and flowers of *Hibiscus roseus* presented lower phenolic and flavonoid content. However, phenolic compounds have potential applications in the skin [37].

Our study showed, for the first time, that dried leaves of *H. rosa-sinensis* presented higher antioxidant capacity, phenolic content, and inhibitory pancreatic lipase activity than fresh leaves and revealed a positive correlation between the dried state and biological activities.

Dietary fat promotes more body fat storage than dietary carbohydrate causing obesity, hyperglycemia, hypertension, and atherosclerosis, pathologies included in the metabolic syndrome [38]. Therefore, plants that possess the capacity to inhibit digestion and absorption of dietary fat are a good alternative to treating obesity. Tetrahydrolipstatin (Orlistat TM) is a pancreatic lipase inhibitor that does not affect the nervous system or enter the bloodstream [39]. However, it has many gastrointestinal side effects [40]. Few works measured plasma triacylglycerol levels after oral administration of a lipid emulsion in rats. As shown in Fig. 3, 125 mg/kg of *H. rosea-sinensis* leaf extract presented a reduction in triacylglycerol levels, similar to orlistat. Li *et al.* [41] reported that sesame meal presented pancreatic lipase inhibitory activity and the major compounds were linoleic acid and oleic acid. They also showed that these two fatty acids reduced fat digestion. As mentioned above, linoleic acid is one of the major compounds in *H. rosa-sinensis*, so its anti-lipase activity may be produced by this compound. The finding that *H. rosa-sinensis* reduced the elevation of plasma triacylglycerol in a similar way to orlistat did, suggests a relationship with the inhibitory pancreatic lipase activity. *H. rosa-sinensis* dried leaf extract may be useful for body weight control.

**Conclusions**

Based on the analysis of 37 plants, *H. rosa-sinensis* showed the highest inhibition of pancreatic lipase activity that displayed a positive correlation with the antioxidant capacity. Consequently, *H. rosa-sinensis* can
be used to reduce body weight. Further studies are required to identify the active compound that shows inhibitory pancreatic lipase activity.

Acknowledgments

The authors are grateful and wish to highlight that this line of research that is cultivated in the group had its origin in the contributions and leadership of Professor Victor M Loyola-Vargas in the field of chemistry and natural products.

References