



## ANTIMICROBIAL ACTIVITY OF FLAVONOIDS FROM *Manilkara zapota* (L.) LEAVES

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*Manilkara zapota* (L.) (Synonyms: *M. zapotilla*, *M. achras*, *Mimusopus manilkara*, *Achras zapota*, and *A. sapota*), commonly known as Sapodilla belongs to the Sapotaceae family [1]. It is an evergreen, glabrous tree, 8-15 m in height, and native to Mexico and Central America. In folk medicine, the leaves are used to treat coughs, colds and diarrhea [2]. For this reason, the antimicrobial properties of ethanolic extract, fractions and isolated compounds from *M. zapota* leaves were assessed by determining the minimal inhibitory concentrations (MICs) in relation to a Gram-positive bacterium, a yeast and a dermatophyte with the broth microdilution technique [3]. The MIC values were obtained after 24 hours, 18 hours and 5 days for their respective pathogens. Moreover, the separation of compounds was performed by means of chromatographic techniques (LC flash and preparative-HPLC, with H<sub>2</sub>O–CH<sub>3</sub>CN gradients) using the high polarity fraction obtained from the ethanol extract of the leaves. The structures of three compounds (4, 5 and 6 samples in the MIC) were elucidated through 1D and 2D NMR spectroscopic data (in CD<sub>3</sub>OD on a Bruker 500 MHz spectrometer) as well as MS data (High-resolution ESITOFMS techniques). The pseudomolecular ions were obtained at  $m/z$  481.0986 [M + H]<sup>+</sup>; (481.0982 calcd for C<sub>21</sub>H<sub>21</sub>O<sub>13</sub> – );  $m/z$  463.0877 [M–H]<sup>–</sup> (463.0877 calcd for C<sub>21</sub>H<sub>19</sub>O<sub>12</sub>.) and  $m/z$  = 477.1039 [M–H]<sup>–</sup> (477.1033 calcd for C<sub>22</sub>H<sub>21</sub>O<sub>12</sub>.). Thus, the structural elucidation showed that the isolated compounds from *M. zapota* leaves are flavonoids (Flavon-3-ol glycosides). The MIC values for the samples are reproduced in Table 1. In conclusion, the ethanol extract fractions and isolated compounds from *M. zapota* leaves demonstrated potential antimicrobial action. Furthermore, the compound 4 (myricetin-3-glucoside) proved to be potentially active against *S. aureus* (MIC of 8 µg/mL) and also showed high activity against *T. rubrum* (MIC of 16 µg/mL).

Table 1: Minimum Inhibitory Concentration (MIC) of the ethanolic extract (1), fractions (2, 3) and isolated compounds (4, 5 and 6) from *M. zapota* leaves.

Strain	MIC (µg/mL)						Pos. ctrl <sup>*</sup>
	1	2	3	4	5	6	
<i>C. albicans</i> (ATCC 10231)	16	16	16	>128	>128	>128	8 <sup>a</sup>
<i>T. rubrum</i> (SNB-TR1)	256	128	>512	16	64	128	3.125x10 <sup>-3a</sup> 2 <sup>b</sup>
<i>S. aureus</i> (ATCC29213)	128	32	128	8	32	32	2 <sup>c</sup>

\*Positive control: <sup>a</sup> fluconazole; <sup>b</sup> itraconazole; <sup>c</sup> oxacilin.

### References:

- [1] Quattrocchi F.L. 2000 CRC World Dictionary of Plant Names-Common Names, Scientific Names, Eponyms, Synonyms and Etymology. Boca Raton, London, New York, Washington, DC: CRC Press; p. 1609-10.
- [2] Mohiddin H.M.Y.B, Chin W., Holdsworth D. 1992. Traditional medicinal plants of Brunei, Darussalam Part III. Sengkurong. Int J Pharmacog; 30: 105-108.
- [3] Clinical and Laboratory Standards Institute, 'Reference Method for Broth Dilution Antifungal Susceptibility Testing of Yeasts; Approved Standard', 2008 3<sup>rd</sup> ed, Document M27-A3, CLSI, Wayne.

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