

## Anti-parasitic activity of terpenoids from *Casearia sylvestris*

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*Casearia sylvestris* Swartz (Salicaceae) is a plant known as "guaçatonga" that has valuable pharmacological arsenal because of the presence of clerodane diterpenes known as casearins A-X and casearvestrins A-C that have been described as compounds with anticancer activity<sup>[1][2]</sup>. Besides that, the anti-parasitic effects of casearins A, B, G and J was established<sup>[2]</sup>. Leishmaniasis and Chagas disease are neglected diseases caused by the *Leishmania* protozoa and *Trypanosoma cruzi* respectively. The treatments for these diseases are based in chemotherapy drugs, that has many undesirable side effects and high toxicity<sup>[3][4]</sup>. This work describes the antileishmanial and antitrypanosomal activities of  $\delta$ -cadinol and casearin L, metabolites isolated from the hexane phase of MeOH extract from the leaves of *Casearia sylvestris* by the bioguided chromatographic fractionation. The compounds were identified by RMN <sup>1</sup>H, <sup>13</sup>C and mass spectra (Figure 1).

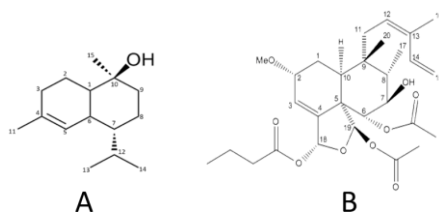


Figure 1. Structure of  $\delta$ -cadinol (A) and Casearin L (B)

$\delta$ -cadinol showed IC<sub>50</sub> of 9,47  $\pm$  0,88  $\mu$ g/mL against amastigotes of *L. infantum* and CC<sub>50</sub> of 35,39  $\pm$  0,04  $\mu$ g/mL against mammalian fibroblasts, while casearin L showed IC<sub>50</sub> of 2,73  $\pm$  0,34  $\mu$ g/mL and CC<sub>50</sub> of 75,06  $\pm$  0,58  $\mu$ g/mL. Against tripomastigotes of *T. cruzi*,  $\delta$ -cadinol showed IC<sub>50</sub> of 5,3  $\pm$  0,4  $\mu$ M and CC<sub>50</sub>>200  $\mu$ M and casearin L, IC<sub>50</sub> > 100  $\mu$ M and CC<sub>50</sub> of 22,54  $\pm$  1,88  $\mu$ M. These results demonstrated that  $\delta$ -cadinol has a better activity against *T. cruzi* while casearin L against *L. infantum*, because of a higher selectivity index of >37,73 and 27,49 respectively, meaning that it needs a small concentration to kill the parasite, but a higher one to kill the healthy mammalian cells. Therefore, the compounds have a selective toxicity against these protozoans. Considering the search of new alternative and selective drugs with less adverse effects, these results suggest that *C. sylvestris* is a promising tool for a possible development of new anti-parasitic drugs.

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