Cytotoxicity evaluation of *nor*-neolignans from the fruits of two Styrax species.

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Introduction

Styrax camporum and S. ferrugineus belong to the Styracaceae family. Phytochemical investigations of *Styrax* species have reported lignans, neolignans, nor-neolignans, phenolic acids, saponins and triterpenes.¹ We now describe the isolation and structural identification of chemical constituents from the methanol extract of *S. camporum* and *S. ferrugineus* fruits, as well as their cytotoxicity activities.

Results and Discussion

The powdered fruits of *S. ferrugineus* (95 g) and *S. camporum* (63 g) were extracted with methanol. Subsequently filtration, the solvents were removed under reduced pressure, to yield 8.6 g e 13.3 g of the extract. The extracts (3.5 g, *S. ferrugineus*) e (5 g, *S. camporum*) were chromatographed on silica using a gradient of *n*-hexane/EtOAc as eluent. Compounds **1**, **2**, **3** and **4** were isolated from *S. camporum*. Preparative RP-HPLC and prep-TLC were employed to purify **1** and **2**, respectively. The *S. ferrugineus* extract yield compounds **3**, **4**, and **5**. Compound **5** was isolated by prep-TLC. Compounds **3** and **4** were acetylated (**3a**) and (**4a**).

The spectral data of all the isolated compounds (Figure 1) are in agreement with previously published data and allowed for identification of demethoxyhomoegonol (1), demethoxyegonol (2), homoegonol (3), egonol (4), and egonol-2-methylbutanoate (5).

The screening for cytotoxic activity against cell line GM07492A (normal human lung fibroblasts) was assessed using the Colorimetric Assay *In Vitro* Toxicology - XTT Kit (Roche Diagnostics).

The extracts of *S. ferrugineus* and *S. camporum* shown IC₅₀ of 270.8 \pm 14.2 µg/mL and 164.7 \pm 10.9 µg/mL, respectively. Treatment of the cells with compounds **1**, **2**, **3a**, and **4a**, did not reveal statistically significant reduction in cell viability at assayed concentrations. Although, compound **5**

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shown IC₅₀ of 50.26 \pm 1.97 μM . In this case, IC₅₀ was considered promising, when compared with the other compounds and with previously publish data for compounds **3** and **4**.²

The most promising compound (5) exhibited similar chemical structure, differing only in the presence of a 2-methylbutanoate moiety, which increased the hydrophobicity and the activity.



Figure 1. Chemical structures of the isolated compounds.

Conclusions

The compound **5** was more toxic to cell line, and the hydrophobicity could be explored to improve cytotoxic activity.

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