

ABSTRACT

Charles University, Faculty of Pharmacy in Hradec Králové, Department of Pharmaceutical Botany

Author: Hana Šimková

Supervisor: prof. Ing. Lucie Cahlíková, Ph.D.

Title of diploma thesis: Alkaloids of genus *Narcissus*: isolation, structural identification, biological activity

Key words: *Narcissus*, alkaloids, biological activity, Alzheimer's disease, cytotoxic activity

The aim of the diploma thesis was an isolation of alkaloids with a focus on minor fractions. These fractions were obtained from the summary alkaloid extract of *Narcissus pseudonarcissus* cv. Carlton. The method of preparative TLC was used for the isolation of alkaloids. Three substances of alkaloid origin marked as Fj 3-4/kr, F 7/2-1, F 7/2-3 were isolated from the assigned fractions. These substances were identified as alkaloids of homolycorine type lycorenine, homolycorine and hippeastrine by using GC-MS, NMR and optical rotation. The results were also compared with data in the literature.

These three alkaloids were tested for their inhibitory activity against AChE, BuChE, POP and GSK-3 β . The inhibitory activity against AChE and BuChE was compared with the reference substances galanthamine (IC_{50} AChE = $1,71 \pm 0,07 \mu\text{M}$, IC_{50} BuChE = $42,3 \pm 1,3 \mu\text{M}$) and huperzine A (IC_{50} AChE = $0,033 \pm 0,001 \mu\text{M}$, IC_{50} BuChE > $1000 \mu\text{M}$). The inhibitory activity against POP was compared to Z-Pro-prolinal (IC_{50} POP = $3,27 \pm 0,02 \text{ mM}$) and berberine (IC_{50} POP = $0,14 \pm 0,02 \text{ mM}$). The most active one of the isolated alkaloids was homolycorine. Inhibitory activity of homolycorine against AChE (IC_{50} = $63,7 \pm 4,3 \mu\text{M}$), BuChE (IC_{50} = $151 \pm 19 \mu\text{M}$) and POP ($40,6 \pm 1,3 \text{ mM}$). Moderate inhibitory activity against GSK-3 β was observed in alkaloids homolycorine (% inhibition = 54 ± 1) and lycorenine (% inhibition = 48 ± 3).

Isolated alkaloids were tested for their cytotoxic activity using nine cancer cell lines including Jurkat, MOLT-4, A549, HT-29, PANC-1, A2780, HeLa, MCF-7, SAOS-2. The results did not indicate any significant cytotoxic activity.