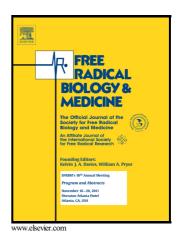
# Author's Accepted Manuscript

9-Norbornyl-6-Chloropurine (NCP) induces Cell death through GSH Depletion-Associated ER stress and Mitochondrial Dysfunction

Pavla Plačková, Michal Šála, Markéta Šmídková, Milan Dejmek, Hubert Hřebabecký, Radim Nencka, Hendrik-Jan Thibaut, Johan Neyts, Helena Mertlíková-Kaiserová



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# **ACCEPTED MANUSCRIPT**

**Title** 

9-NORBORNYL-6-CHLOROPURINE (NCP) INDUCES CELL DEATH THROUGH GSH DEPLETION-ASSOCIATED ER STRESS AND MITOCHONDRIAL DYSFUNCTION

### Author names and affiliations

Pavla Plačková<sup>a</sup>, Michal Šála<sup>a</sup>, Markéta Šmídková<sup>a</sup>, Milan Dejmek<sup>a</sup>, Hubert Hřebabecký<sup>a</sup>, Radim Nencka<sup>a</sup>, Hendrik-Jan Thibaut<sup>b</sup>, Johan Neyts<sup>b</sup>, Helena Mertlíková-Kaiserová<sup>a</sup>\*

<sup>a</sup>Institute of Organic Chemistry and Biochemistry, Academy of Sciences of the Czech Republic, v.v.i., 166 10 Prague 6, Czech Republic

<sup>b</sup>Rega Institute for Medical Research, KU Leuven, Minderbroedersstraat 10, BE-3000 Leuven, Belgium

\*Corresponding author: mertlikova@uochb.cas.cz

Abbreviations: NCP, 9-norbornyl-6-chloropurine; GST, glutathione-S-transferase; GSH, L-glutathione reduced, MMP, mitochondrial membrane potential; GR, glutathione reductase; PI, propidium iodide; 4-PBA, sodium phenylbutyrate; QUER, quercetin; ALLO, allopurinol; ABT, 1-aminobenzotriazole; CCCP, carbonyl cyanide 3-chlorophenylhydrazone; CAPE, caffeic acid phenethyl ester; MDA, 1,1,3,3-tetraethoxypropane; DPI, diphenyleneiodonium chloride; TBA, 2-thiobarbituric acid; EDTA, ethylenediaminetetraacetic acid; TFA, trifluoroacetic acid; GSSG, L-glutathione oxidized; SA, sulfosalicylic acid; CDNB, 1-chloro, 2,4-dinitrobenzene; LO, lipoxygenase

#### **Abstract**

9-Norbornyl-6-chloropurine (NCP) is a representative of a series of antienteroviral bicycle derivatives with selective cytotoxicity towards leukemia cell lines. In this work we explored the mechanism of the antileukemic activity of NCP in T-cell lymphoblast cells (CCRF-CEM). Specifically, we searched for a potential link between its ability to induce cell death on the one hand and to modulate intracellular glutathione (GSH) that is necessary to its metabolic transformation via glutathione-S-transferase on the other hand. We have observed that GSH

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