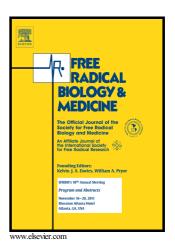
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ACCEPTED MANUSCRIPT

VAOS, a novel vanadyl complexes of alginate saccharides, inducing apoptosis via activation of AKT-dependent ROS production in NSCLC

Ling Zhou^{ab1}, Yuetao Yi^{c1}, Qing Yuan^a, Jing Zhang^a, Youjie Li^a, Pingyu Wang^a, Maolei Xu^{b*}, Shuyang Xie^{a*}

^aKey Laboratory of Tumor Molecular Biology in Binzhou Medical University, Department of Biochemistry and Molecular Biology, Binzhou Medical University, YanTai, ShanDong, 264003, P.R.China.

^bThe Key Laboratory of Traditional Chinese Medicine Prescription Effect and Clinical Evaluation of State Administration of Traditional Chinese Medicine, School of Pharmacy, Binzhou Medical University, YanTai, ShanDong, 264003, P.R.China.

^cYantai Institute of Coastal Zone Research Chinese Academy of Sciences, Yantai 264003, P.R.China.

shuyangxie@aliyun.com (Shu-Yang Xie)

xumaolei1234@163.com (Mao-Lei Xu)

* Corresponding authors. Tel: +86 535 6913070. Fax: +86 535 6913163.

Abstract

Previous studies have confirmed that protein tyrosine phosphatase 1B (PTP1B) can promote tumour progression in non-small cell lung cancer (NSCLC). Vanadyl alginate oligosaccharides (VAOS) is a new coordination compounds that possesses a good PTP1B inhibitory activity. However, the potent anticancer efficacy of VAOS in human NSCLC requires further study. In this study, VAOS exhibited effective inhibitory effects in NSCLC both in cultured cells and in a xenograft mouse model. VAOS was further identified to induce NSCLC cell apoptosis through activating protein kinase B (AKT) to elevate intracellular reactive oxygen species (ROS) levels by increasing in oxygen consumption and impairing the ROS-scavenging system. Neither silencing of *PTP1B* by siRNA nor transient overexpression of *PTP1B* had an effect on the AKT phosphorylation triggered by VAOS, indicating that PTP1B inhibition was not involved in VAOS-induced apoptosis. Through phosphorus

¹ These authors contributed equally to this work.

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