

International BioScience Conference Novi Sad, Serbia November

2021

INHIBITION OF TUMOR GROWTH IN DISULFIRAM TREATMENT OF FIBROSARCOMA INOCULATED TO HAMSTERS

<u>Kosta J. Popović</u>¹, Dušica J. Popović², Dejan Miljković², Dušan Lalošević², Ivan Čapo² and Jovan K. Popović³

¹Department of Pharmacy, Faculty of Medicine, University of Novi Sad, Novi Sad, Republic of Serbia,

²Department of Histology and Embryology, Faculty of Medicine, University of Novi Sad, Novi Sad, Republic of Serbia,

³Department of Pharmacology, Toxicology and Clinical Pharmacology, Faculty of Medicine, University of Novi Sad, Novi Sad, Republic of Serbia

KEYWORDS: disulfiram; hamster fibrosarcoma; anticancer effects

INTRODUCTION

We investigated the anticancer effect of an clinically used anti-alcoholism drug disulfiram on *an in vivo* solid tumor model of fibrosarcoma in hamsters. Disulfiram inhibits growth of various cultured cancer cell lines.¹⁻³.

OBJECTIVES

Objective of the research was to prove that disulfiram inhibits growth of fibrosarcoma inoculated to hamsters.

METHOD / DESIGN:

20 Syrian golden hamsters of both sexes (10 males and 10 females), weighing approximately 70 g, were randomly allocated experimental and control group hamsters/group). 2 x 106 BHK-21/C13 cells in 1 ml were injected subcutaneously into the animals' back in both groups. The experimental group started peroral treatment with disulfiram 200 mg/kg daily via a gastric probe 3 days before tumor inoculation. After 19 days, when the tumors were approximately 2-3 cm in the control group, all animals were sacrificed. The blood was collected for glucose and other analyses. The tumors were excised and weighed and their volume (by water displacement method) and diameters were measured (Figure). The tumor samples were histologically and immunohistologically assessed and the main organs toxicologically analyzed. Tumor volume was also determined using the formula LxS²/2, where L was the longest and S the shortest diameter. Ki-67-positive cells in the tumor samples were quantified; images were taken and processed by software UTHSCSA Image Tools for Windows Version 3.00. Statistical significances of differences in tumor weight, volume, number of Ki-67-positive cells and other parameters were determined by the one way ANOVA.

RESULTS

Disulfiram inhibited fibrosarcoma growth in hamsters without toxicity and without influence on blood analyses.

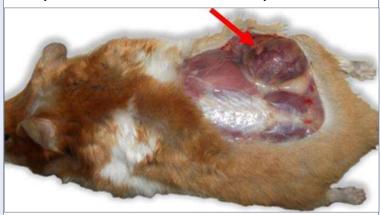


Figure. BHK fibrosarcoma: subcutaneous localization in a hamster

CONCLUSIONS

Inhibition of proteasome activity by disulfiram as an antitumor strategy might be an effective and safe therapeutic approach in novel nontoxic therapies and relapse prevention for human cancers.

ACKNOWLEDGEMENTS

This study was supported by the Republic of Serbia, Autonomous Province of Vojvodina, Provincial Secretariat for High Education and Scientific Research [Project title: Discovery of effective non-toxic anticancer drug combinations on experimental fibrosarcomas, grant no. 142-451-2498/2021-03 (Project leader Dušica Popović)] and Republic of Serbia, Ministry of Science [grant no. 172013].

REFERENCES

- Yip, N.C., Fombon, I.S., Liu, P., Brown, S., Kannappan, V., Armesilla, A.L., Xu, B., Cassidy, J., Darling, J.L., Wang, W., 2011. Disulfiram modulated ROS-MAPK and NFkB pathways and targeted breast cancer cells with cancer stem cell-like properties. Br. J. Cancer., 104(10), 1564–1574.
- Wang, W., Darling, J. L., 2013. How could a drug used to treat alcoholism also be effective against glioblastoma?. Expert. Rev. Anticancer. Ther., 13(3), 239–241.
- Popović KJ, Popović DJ, Miljković D, Popović JK, Lalošević D, Čapo I. Co-treatment with nitroglycerin and metformin exhibits physicochemically and pathohistologically detectable anticancer effects on fibrosarcoma in hamsters. Biomed. Pharmacother. 2020; 130: 110510

CONTACT

Corresponding author:

jovan.popovic@mf.uns.ac.rs; jovapopmf@gmail.com

