# Resistance training and its effect on the development of mammary tumors in MNU-induced Wistar rats

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## Introduction

Mammary cancer is the most common cancer in women . Wistar rats have been generally recommended in experimental protocols to study mammary carcinogenesis. This outbred strain is very sensitive to chemical carcinogen agents, such as N-methyl-N-nitrosourea (MNU). According to many publications, physical activity improve body condition and may help in the breast cancer prevention. In order to evaluate the effects of resistance training in the mammary tumors' development, we evaluated several parameters such as, the latency period, total number of tumors *per* group, mean number of tumors *per* animal, mean volume and weight of tumor *per* group.

### Materials and Methods

Twenty-five female Wistar rats were divided into four groups (n=7): Sedentary (SED); SED+MNU; Exercised (EX); and EX+MNU. SED+MNU and EX+MNU animals received an intraperitoneal injection of the carcinogen MNU (50mg/Kg), at seven weeks of age. Exercised animals were trained 3 days/week for 18 weeks, by climbing a 1-meterhigh homemade ladder, 8-12 dynamic movements/climb and 4-8 climbs/session. Three animals (two from SED+MNU and one from EX+MNU groups) were humanely sacrificed before the end of the experiment and excluded from the study. All experiments were approved by the Ethics Committee and National Competent Authority. Data were analyzed using Microsoft Excel<sup>®</sup>.

#### SED n=7 Exercise: 8-12 dynamic movements/climb 4-8 climbs/session SED+MNU 3 days/week n=7 ΕX $\bigcirc$ Quarantine n=7 Acclimatization MNU administration (50mg/Kg, i.p.) EX+MNU SED - Sedentary; n=7 MNU - N-methyl-N-nitrosourea; Sacrifice EX - Exercised; Weeks of age

Fig.1 – Schematic representation of the experimental protocol.

#### Results

Animals from control groups (SED and EX) did not develop any mammary tumor. In order to ensure animals' welfare, two animals from SED+MNU group were sacrificed at

twelfth and seventeenth weeks and one animal from EX+MNU was sacrificed at seventeenth week of the experiment. The first mammary tumor was detected at the tenth week after MNU administration in SED+MNU group. Two weeks later, EX+MNU developed the first mammary tumor. At the end of the experiment, we detected a total of four tumors in SED+MNU group ( $0.8 \pm 1.17$  tumors *per* animal) and 10 tumors in EX+MNU group ( $1.67 \pm 2.87$  tumors *per* animal). The average tumor volume was slightly higher in SED+MNU group ( $6.71 \pm 6.28$ ) when compared with EX+MNU group ( $3.05 \pm 0.98$ ) when compared with SED+MNU group ( $2.31 \pm 1.95$ ).

SED+MNU	EX+MNU
10 weeks	12 weeks
$0.8 \pm 1.17$	$1.67 \pm 2.87$
4	10
$6.71\pm 6.28$	$6.43 \pm 7.88$
$3.05\pm0.98$	$2.31 \pm 1.95$
	SED+MNU         10 weeks $0.8 \pm 1.17$ 4 $6.71 \pm 6.28$ $3.05 \pm 0.98$

f exercise on the development of mammary tumors.





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The difference of two weeks between the appearance of the first mammary tumor in the SED+MNU group and the EX+MNU group suggests that exercise effectively delays the appearance of mammary tumors by increasing the latency period. The higher number of tumors *per* animal and tumors' weight in EX+MNU group can be explained by the enhancement of blood perfusion caused by exercise activity. The disparity in the results obtained regarding the volume and weight of the tumor may be related to its histology, which is currently being analyzed.

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