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# Crotalus molossus Venom-Loaded PLGA Nanoparticles as Potential Drug Delivery Systems for Breast Carcinoma Cells

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#### INTRODUCTION & AIM

Cancer is one of the leading causes of death in developed and in under-development countries. Specifically in Mexico, breast cancer is amongst the most common types in both incidence and mortality.

**Encapsulating** bioactive agents such as **peptides** and **proteins** in biocompatible **nanoparticles**, such as PLGA, may enhance their therapeutic properties.

Snake venoms contain a high diversity of bioactive agents, some of which have been reported with:

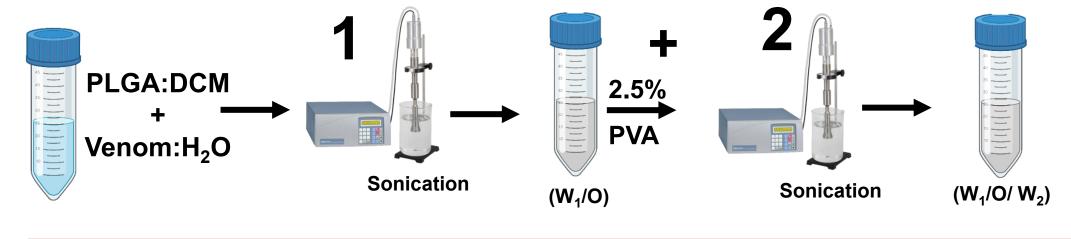


Antibacterial activity
Antiparasistic activity
Antitumoral activity

Our aim was to entrapp the venom of the northern blacktailed rattlesnake (*Crotalus molossus*) into PLGA NPs, and evaluate their drug delivery system potential against a breast carcinoma cell line (T-47D).

#### **METHOD**

Nanoparticles (NPs) were obtained by a double emulsion-solvent evaporation process:



#### CONCLUSION

- The **encapsulation** of **bioactive components** from *C. molossus* venom **was effective** through a double emulsion-solvent evaporation method.
- **Venom** from the **PLGA NPs** was **released** in a **sustained** manner following a first-order model kinetic, driven mainly by diffusion and matrix relaxation process.
- The PLGA-Venom NPs delivered the bioactive components into the breast carcinoma cells, supporting their role as potential drug delivery systems.

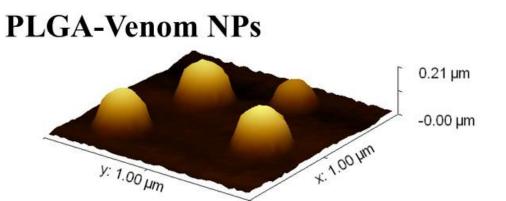
#### FUTURE WORK / REFERENCES

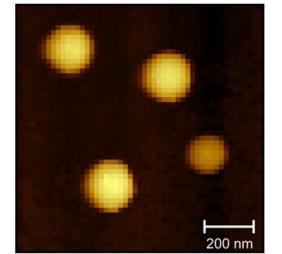


#### **RESULTS & DISCUSSION**

### PLGA-Venom NPs have suitable sizes and high EE%

Nanoparticles	D <sub>H</sub> (nm)	PDI	Z-Potential (mV)	EE%	LC%
PLGA NPs	$286.9 \pm 3.49$	0.11 ± 0.02	$\textbf{-32.5} \pm 0.65$	N/A	N/A
<b>PLGA-Venom NPs</b>	$310.2 \pm 5.36$	$0.15 \pm 0.02$	$\textbf{-32.8} \pm 0.46$	$74.54 \pm 4.22$	$9.19 \pm 0.92$

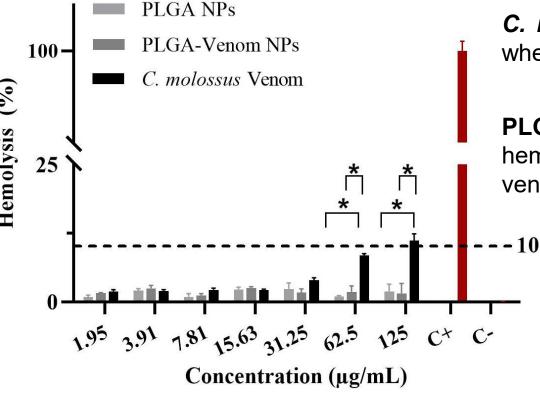




**PLGA-Venom NPs** have a smooth surface and are spherically shaped.

**AFM** images confirm **DLS** size data.

### **Snake Venom Encapsulation Inhibits Hemolytic Activity**

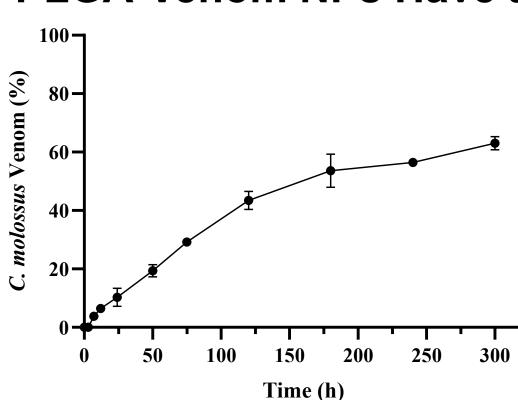


**C. molossus** venom can be considered as hemolytic when evaluating concentrations >60 μg/mL.

**PLGA-Venom NPs** did not exhibit a significant hemolytic activity when evaluated at the same venom concentrations.

Data suggests that by using concentrations <60 µg/mL of *C. molossus* venom **may** be **safe** against red blood cells.

#### PLGA-Venom NPs Have a Sustained Protein Release

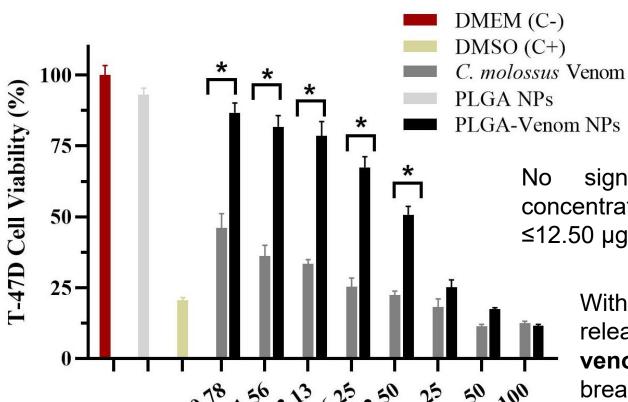


*C. molossus* venom from the PLGA NPs was released in a sustained manner, similarly to what's been reported in other works.

Approximately 12.5% of *C. molossus* venom was released during the first 24 h and, around 60% cumulatively released at 300 h.

**First-order** model **best fits** our data (R<sup>2</sup>: 0.997), while **zero-order** and **Higuchi** models had **decent**, **but inferior** values (R<sup>2</sup>: 0.931 and 0.981, respectively).

## PLGA-Venom NPs are Cytotoxic Against Breast Carcinoma Cells



IC<sub>50</sub> of *C. molossus* venom was determined at 1.55 μg/mL at 24 h.

No significant differences were found at concentrations  $\geq 25 \, \mu \text{g/mL}$ , in contrast to those  $\leq 12.50 \, \mu \text{g/mL}$  ( $p \leq 0.001$ ).

With 12.50 μg/mL of loaded venom, the released amount at 24 h is around **1.56 μg of venom**, reaching approximately **50%** of the breast carcinoma **cells viability**.

0.78 1.56 3.13 6.25 1.50 25 50 100

C. molossus Venom Concentration (μg/mL)