

EURADOS Annual Meeting  
Karlsruhe, Germany  
February 27 – March 2, 2017

## Plutonium Biokinetics in the Human Body Following Decorporation Treatment

Sara de Souza Zanotta Dumit, PhD candidate  
[sara.dumit@wsu.edu](mailto:sara.dumit@wsu.edu)

United States Transuranium and Uranium Registries  
Pharmaceutical Sciences Graduate Program  
College of Pharmacy, Washington State University

*“Learning from Plutonium and Uranium  
Workers”*



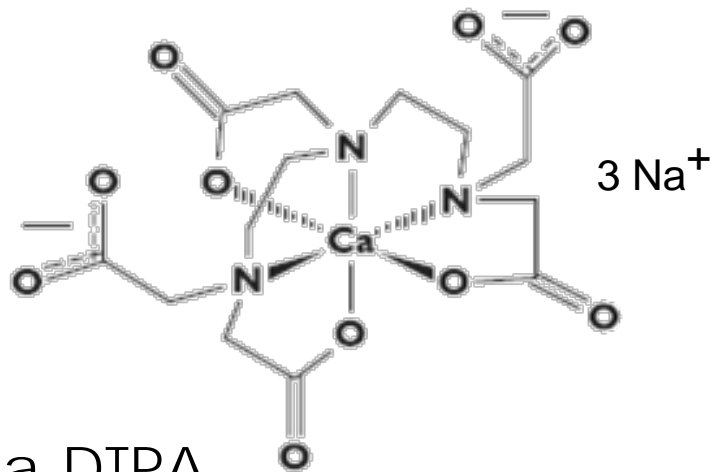


# Plutonium Facts

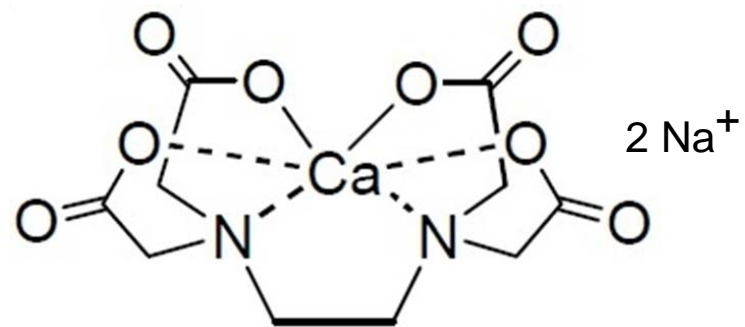
- $^{239}\text{Pu}$  is a major anthropogenic long-lived ( $T_{1/2} = 24,110 \text{ y}$ ) radionuclide of plutonium (Pu)
- Used in nuclear weapons and nuclear power generation
- Enters human body as the result of global fallout, industrial accidents, and occupational incidents
- Has a long-term retention in a human body with skeleton and liver as major depository sites
- Highly exposed individuals treated with chelating agents (decorporation therapy)

# Decorporation Therapy

- Removal of radioactive elements from the body using specific drugs called chelating agents
- Enhances metal excretion by forming stable complex
- Agents for Pu and Am decorporation:



Ca-DTPA  
diethylenetriaminepentaacetic  
acid

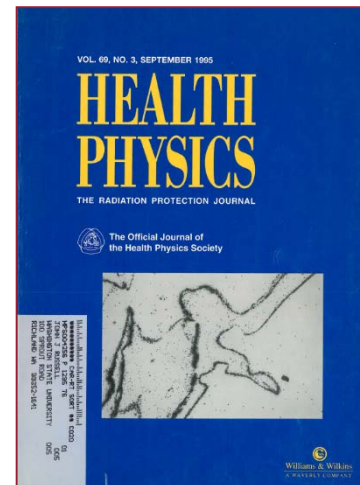
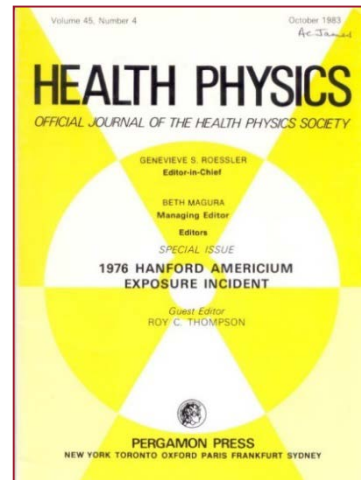


Ca-EDTA  
ethylenediaminetetraacetic  
acid



# Atomic Man: Chelation Therapy Saved His Life

- Explosion of ion-exchange column with ~ 150 g of  $^{241}\text{Am}$  (19 TBq)
- Estimate of uptake > 40 MBq
- Ca/Zn-DTPA chelation therapy administered
- Systemic deposition after treatment – 0.5 MBq
- Treatment efficacy factor: 80





# Motivation and Significance

Motivation:

- No recommended model for plutonium decorporation

Significance:

- Interpretation of bioassay measurements during the therapy
- Optimization of plutonium decorporation treatment



# Cases Available at the USTUR

USTUR Case	Route of intake	Treatment			
		Agent	Dosage, g	# of i.v.	Years
0212	Wound	Ca-DTPA	0.5 – 1.0	47	0.5
0269	Inhalation	Ca-DTPA/EDTA	0.2 – 8.0	161	4.0
0785	Complex	Ca-DTPA/EDTA	1.0 – 2.0	77	7.4

USTUR Case	Route of intake	Treatment			
		Agent	Dosage, g	# of i.v.	Days
0031	Wound	Ca-DTPA	1.0	9	9
0202	Inhalation	Ca-DTPA	1.0	5	5
0303	Wound	Ca-DTPA	0.5 – 1.0	16	60
0407	Inhalation	Ca-DTPA	1.0	5	5
0706	Inhalation	Ca-DTPA	1.0	6	8
0821	Inhalation	Ca-DTPA	1.0	4	4



# Materials and Methods

## Human Data

- In vivo (lung counts)
- In vitro (urine measurements)
- Post mortem radiochemical analyses of tissues

## Software

- Integrated Modules for Bioassay Analysis (IMBA) Professional Plus<sup>®</sup>
- ModelMaker<sup>®</sup> version 4 (MM4)

## Models

- Human Respiratory Tract Model (ICRP 130, 2015)
- Wound Model (NCRP 156, 2006)
- Systemic Model (Leggett *et al.* 2005)
- CONRAD Model for DTPA therapy (2009)



# USTUR Whole Body Case 0212

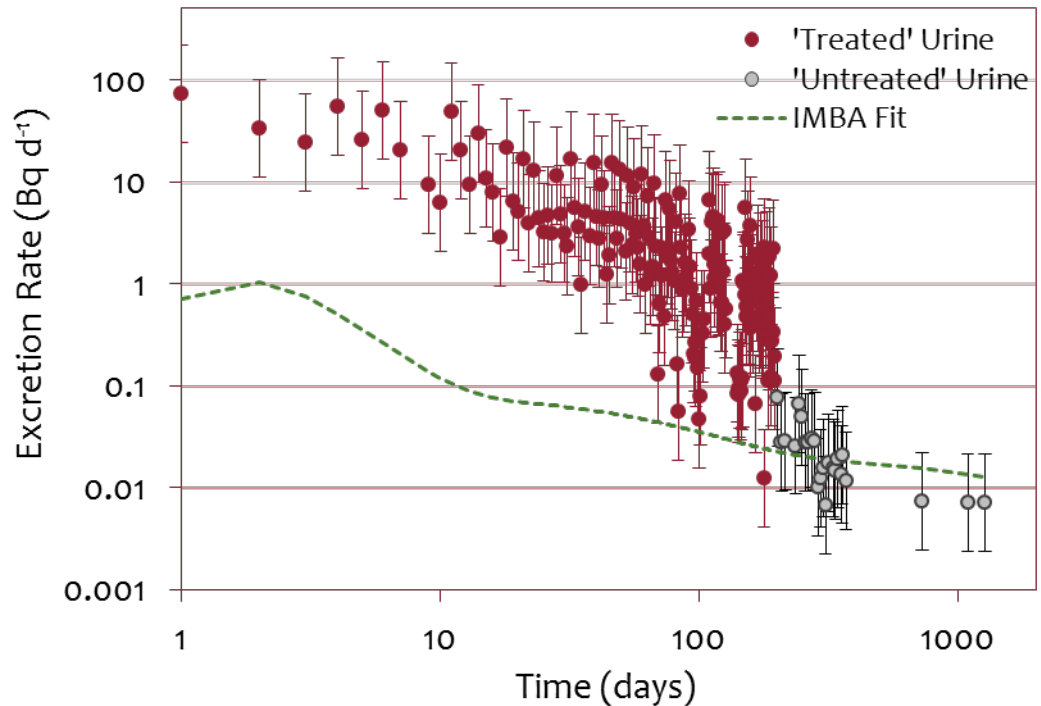
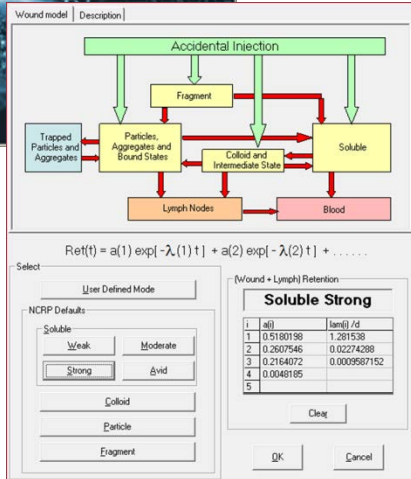
- Exposure: Wound
- Treatment: Tissue excision & Ca-DTPA
- Donation year: 1984
- Time post-intake: 17 y
- Age: 56 y
- Cause of death: Pulmonary emphysema



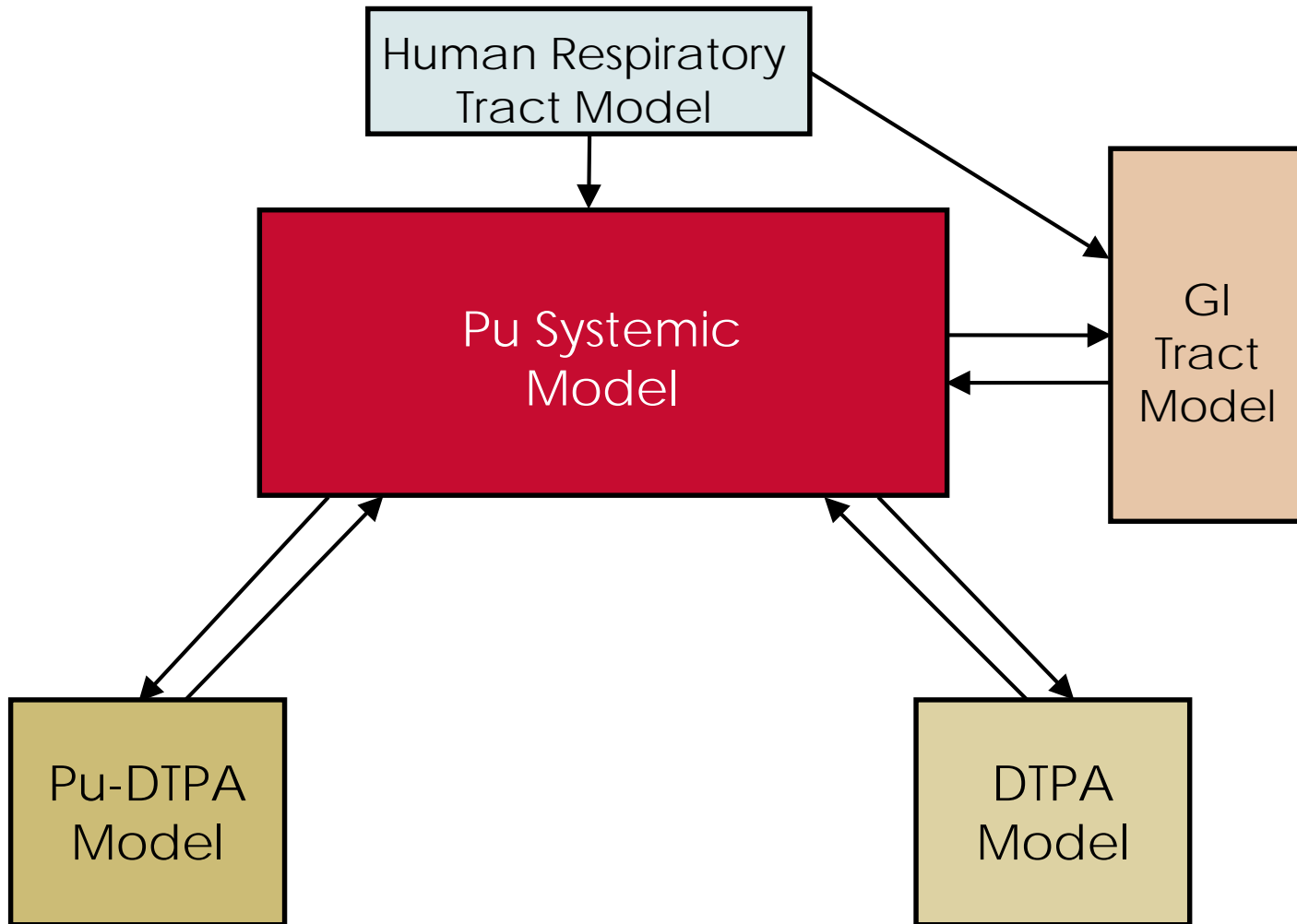
# USTUR Whole Body Case 0212

## Intake

- IMBA estimate: 294 Bq
  - +
  - Pu excreted with DTPA: 916 Bq
- } 1,210 Bq

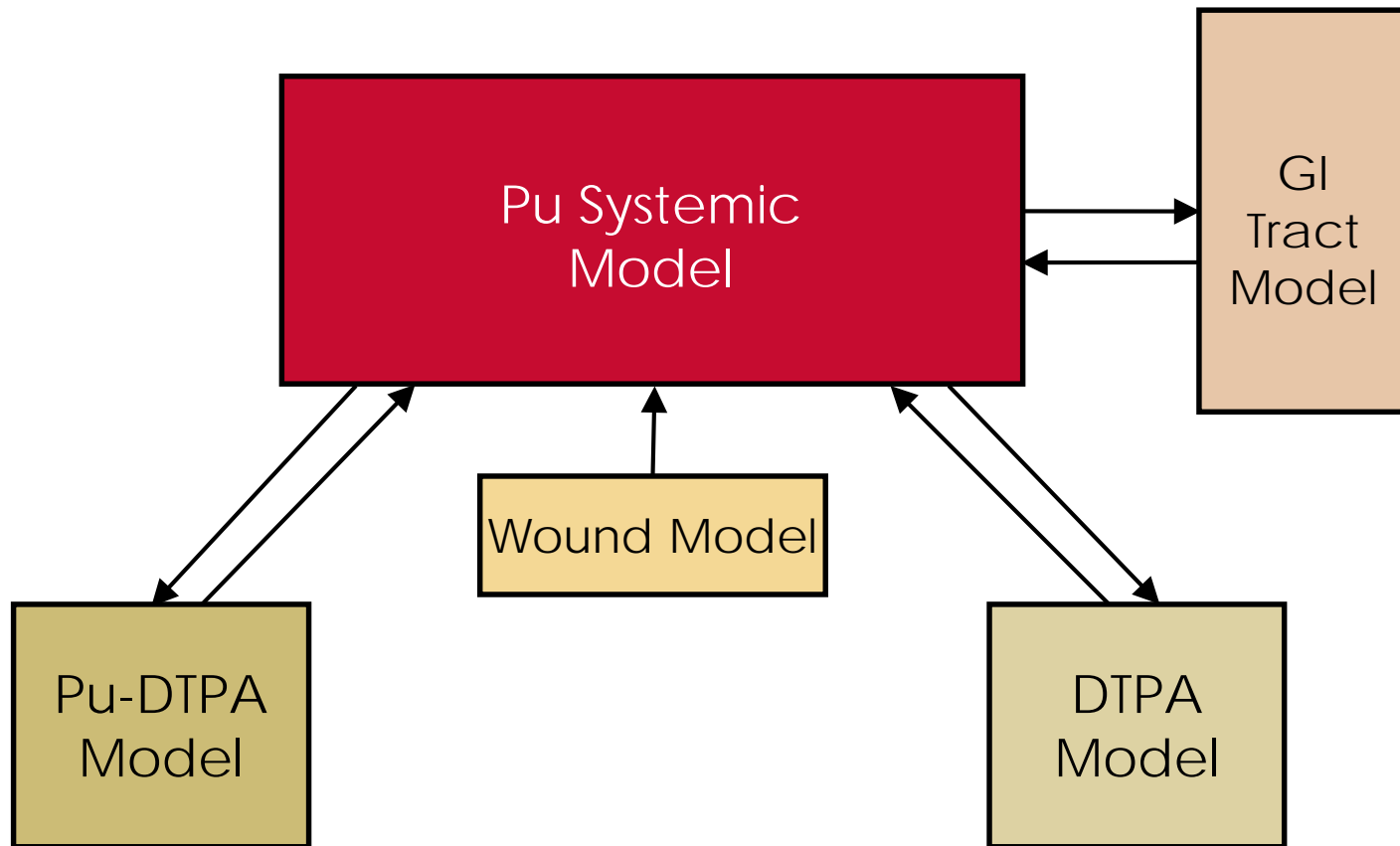


# CONRAD Model



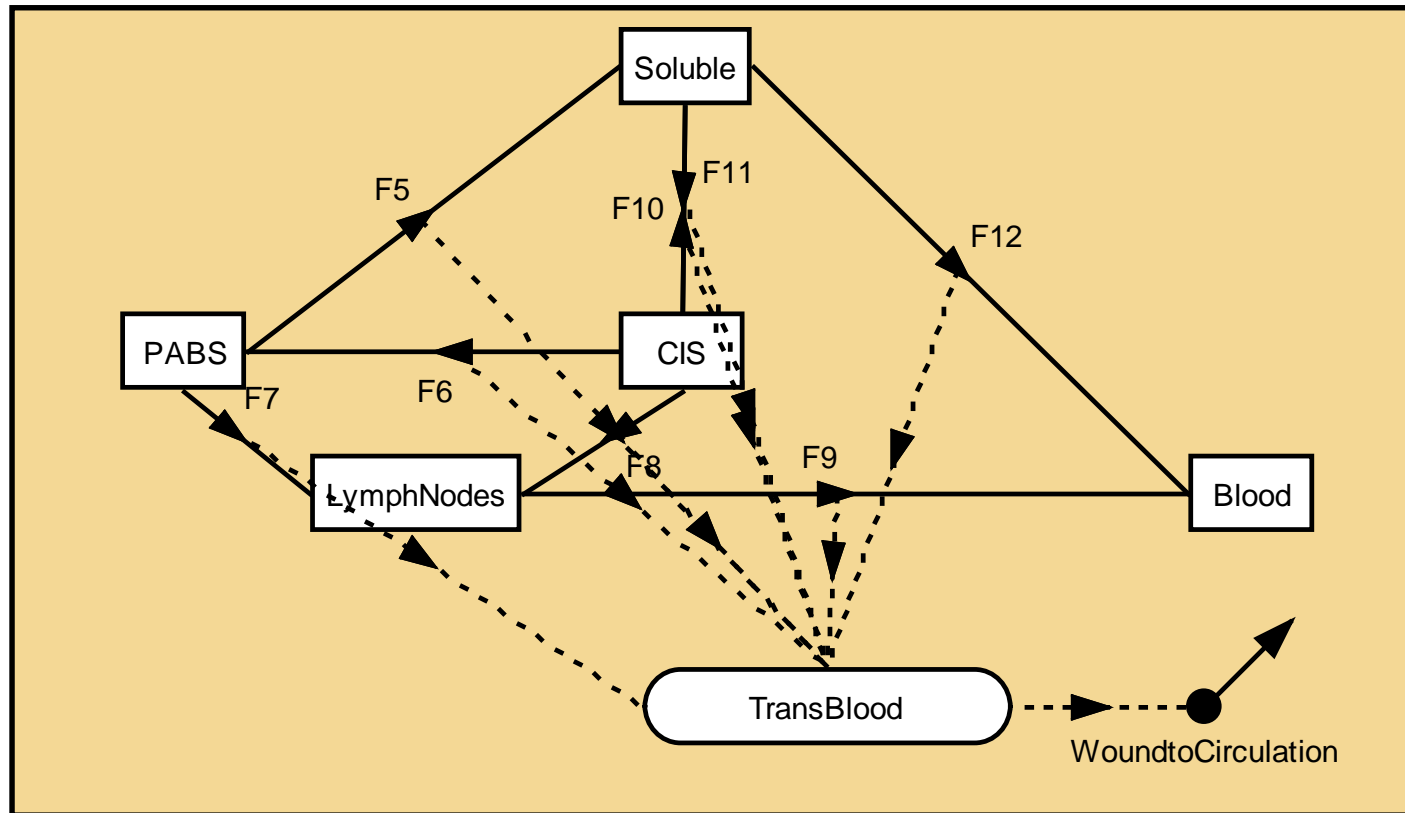


# CONRAD Model + Wound Model



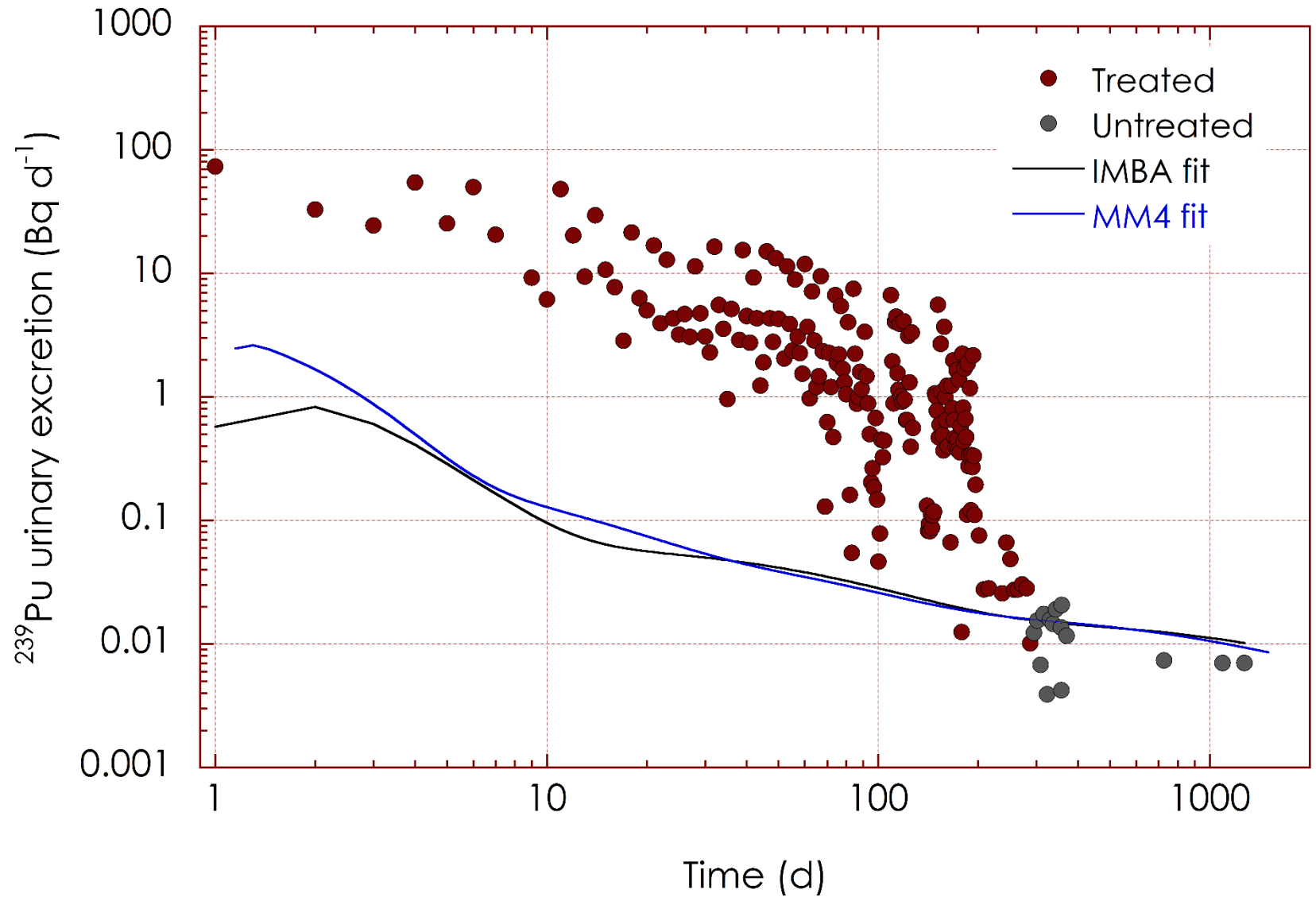


# MM4: NCRP 156 Wound Model

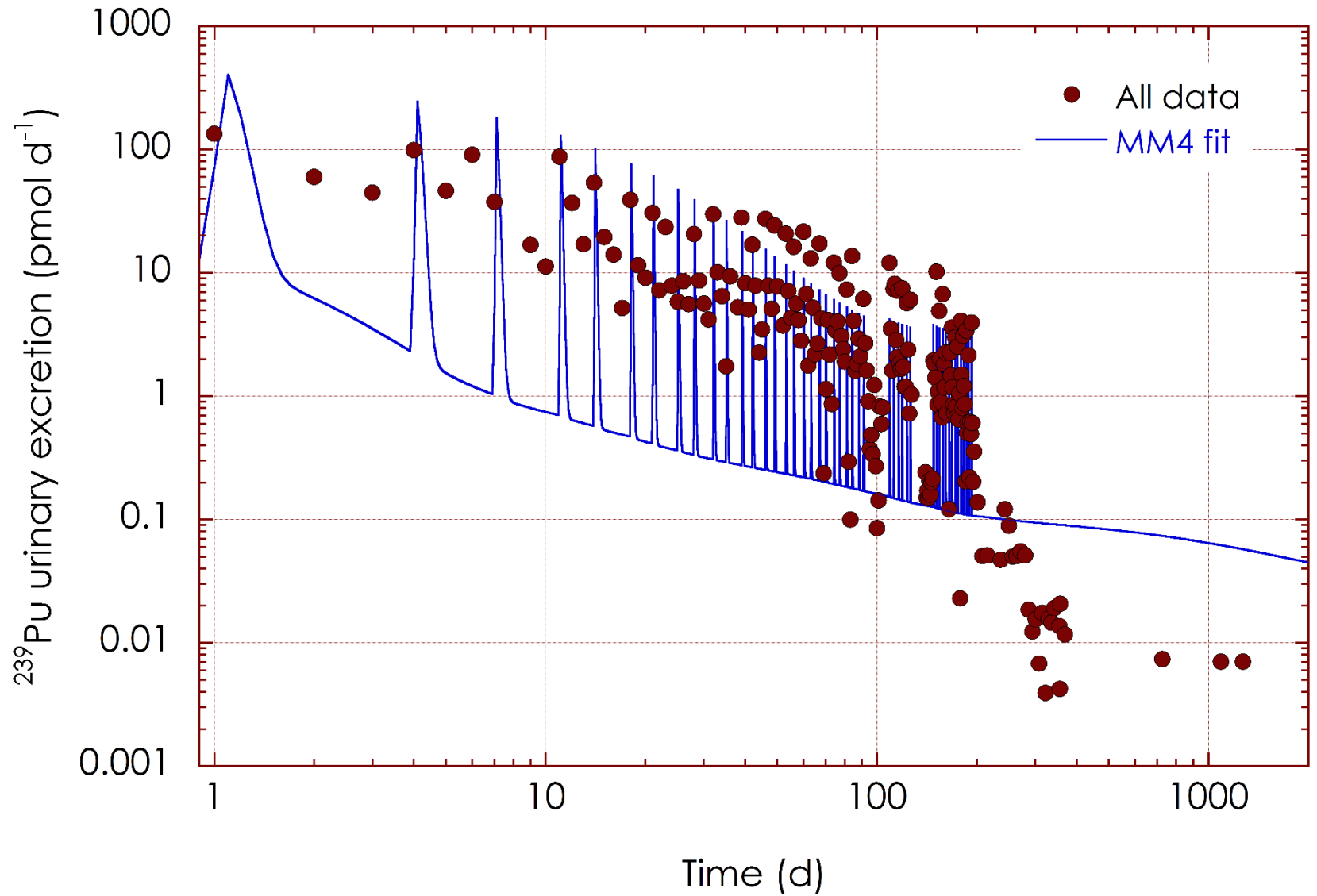




# Baseline Fit: IMBA vs MM4



# Case 0212 Urinary Excretion: MM4 Fit





# Summary

- Interpretation of bioassay measurements during the therapy is still challenging
- Preliminary results showed that CONRAD Pu-DTPA model implemented with NCRP Wound Model is able to fit Pu urinary excretion during decorporation treatment
- However, it has limitation in fitting the data not affected by the therapy (baseline)
- Modifications are needed to better fit both: data affected by treatment and baseline data



# Future Directions

- Consideration:
  - Initial systemic distribution between Blood1 (70%) and ST0 (30%) compartments
  - Intracellular chelation
  - Evaluation of enhancement factor



# Acknowledgments

## USTUR:

- Maia Avtandilashvili
- Margo D. Bedell
- Florencio Martinez
- Stacey L. McComish
- George Tabatadze
- Elizabeth M. Thomas
- Sergei Y. Tolmachev

## Graduate Committee:

- Dr. Sayed S. Daoud, *Chair*
- Dr. Kathryn E. Meier
- Dr. Jeannie Padowski
- Dr. Daniel J. Strom
- Dr. Sergei Y. Tolmachev, *Advisor*

## Collaborator:

- Dr. Bastian Breustedt



**Sponsors:** LASPAU, CAPES, WSU/COP and US DOE/USTUR

