

65th Annual Meeting of the Radiation Research Society
San Diego, Florida, November 2 – 7, 2019

From Autopsies to Synchrotrons to Mars - Why the Brain Matters -

Sergei Y. Tolmachev¹, Tatjana Paunesku², Gayle E. Woloschak²,
John D. Boice Jr^{3,4}

¹ United States Transuranium and Uranium Registries, College of Pharmacy and
Pharmaceutical Sciences, Washington State University

² Department of Radiation Oncology, Feinberg School of Medicine, Northwestern
University

³ National Council on Radiation Protection and Measurements

⁴ Vanderbilt University Medical Center



Northwestern
University



National Council on Radiation
Protection and Measurements



College of
Pharmacy and
Pharmaceutical Sciences
WASHINGTON STATE UNIVERSITY



Growing Interest in Brain Dosimetry for Internal Emitters

- **U.S. Million Worker Study:** estimating brain doses and evaluating dementia, Alzheimer's, and other motor neuron diseases as possible adverse effects of radionuclide depositions in the brain
- **National Aeronautics and Space Administration:** interested in adverse effects of alpha dose on brain as a limited but perhaps informative analogy of behavioral and cognitive effects of galactic cosmic ray (high Z and high energy ions) exposure on astronauts
- **NCRP Scientific Committee 6-12:** Development of Models for Brain Dosimetry for Internally Deposited Radionuclides (2018 – 2020)

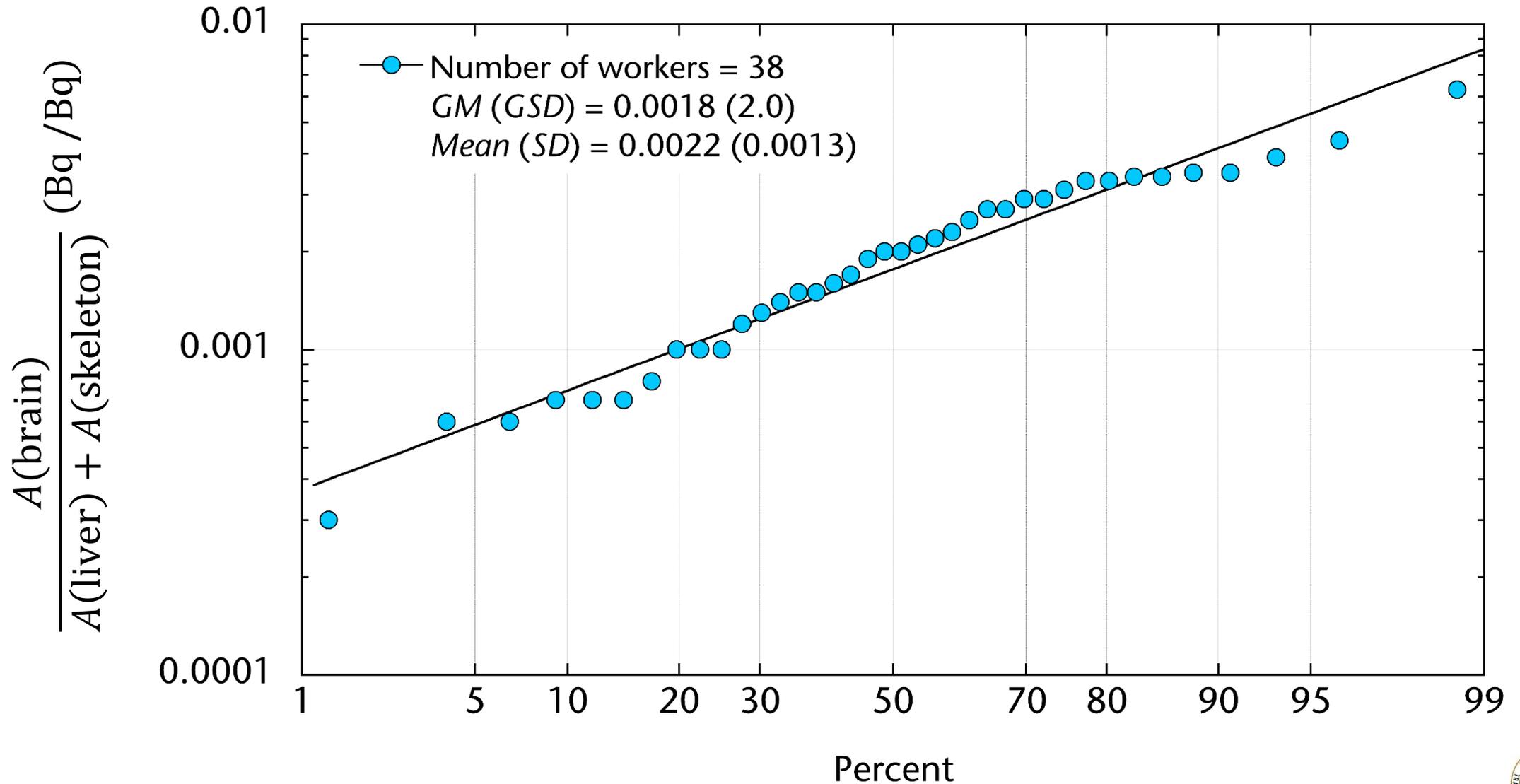


US Transuranium and Uranium Registries: Plutonium in Brain of Occupationally Exposed Individuals





Plutonium: How Much is in the Brain?





Distribution of Plutonium and Radium in Brain

The Bionanoprobe: Synchrotron-based Hard X-ray Fluorescence Microscopy for 2D/3D Trace Element Mapping

Si Chen^{1,*}, Tatjana Paunesku², Ye Yuan², Qiaoling Jin³, Benjamin Hornberger⁴, Claus Flachenecker⁴, Barry Lai¹, Keith Brister⁵, Chris Jacobsen^{1,3,4,6}, Gayle Woloschak², and Stefan Vogt^{1,3} *Micros Today*. 2015 May ; 23(3): 26–29.



Northwestern
University



College of
Pharmacy and
Pharmaceutical Sciences
WASHINGTON STATE UNIVERSITY



USTUR Case 0785

- Dumit *et al.* Health Physics 117: 156-167; 2019
- ^{239}Pu in brain = $1.9 \pm 0.1 \text{ Bq kg}^{-1}$



Brain stem
Hippocampus (R)

Cortex (L)
Hippocampus (L)

Thalamus

Internal
left capsule

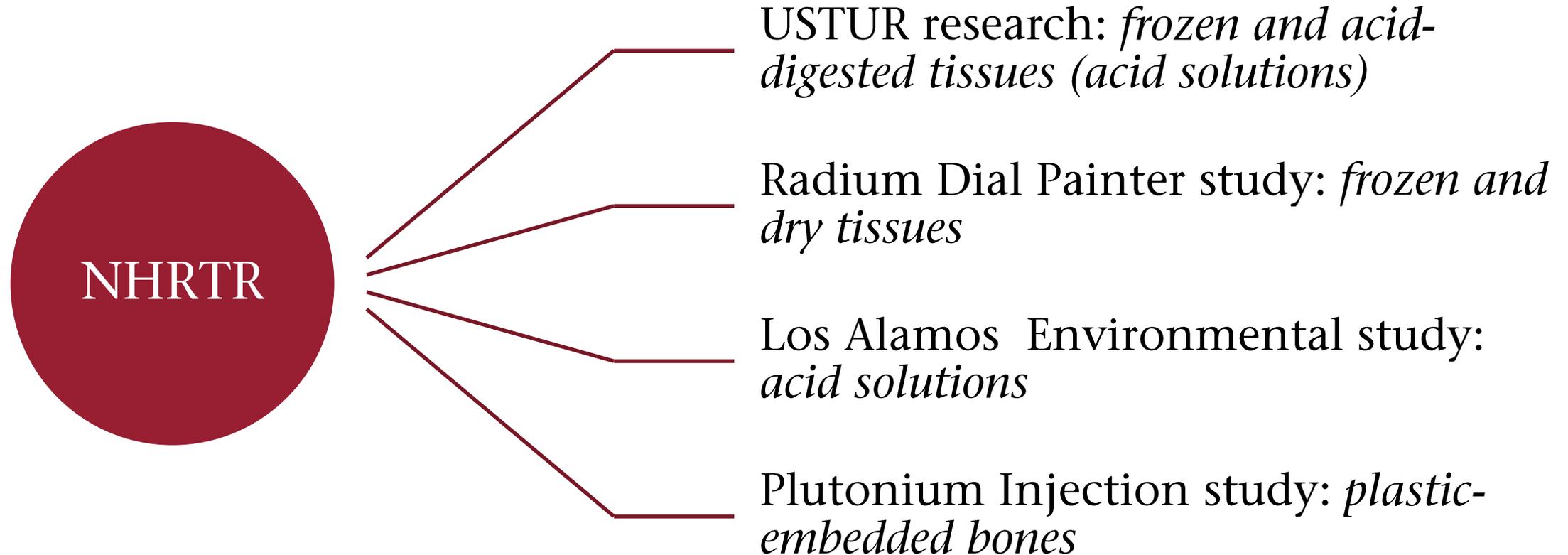
Occipital

Parietal
Cortex (R)

Cerebellum



National Human Radiobiology Tissue Repository (NHRTR) at the US Transuranium and Uranium Registries (USTUR)





Radium Dial Painter: Case 03-551

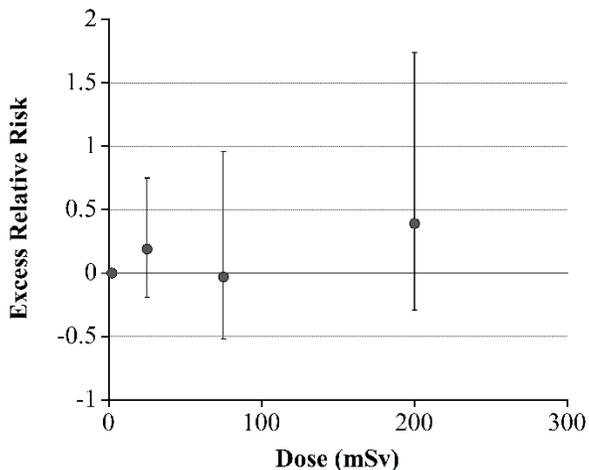
Rowland (1993) Radium in Humans – A review of U.S. Studies

- ^{226}Ra intake = 261.7 μCi (μg)
- ^{226}Ra total body activity = 1.077 μCi (μg)



Current Relevance to the Million Person Study (I)

- Mound Facility (Polonium intake):** Brain dose from polonium deposition and alpha particle decay estimated for the first time and related to the occurrence of dementia, Alzheimer's, Parkinson's and other Motor Neuron Disease (e.g. ALS)



Dose (mGy)	N Workers	N Deaths	RR	95% CI
<5	2,662	69	1.0	Ref
5 - <50	1,262	47	1.19	0.81 - 1.75
50 - <100	328	9	0.97	0.48 - 1.96
100+	324	10	1.39	0.71 - 2.74

p for the trend (two sided) 0.06
 RR at 100 mGy (95% CI) 1.23 (0.99 - 1.54)

- Radium Dial Painter Study (Radium intakes):** A new follow-up of 2,470 dial painters. Radium dose to brain, red bone marrow and lung will be assessed and incorporated into the new biokinetic models for the epidemiologic study



Current Relevance to the Million Person Study (II)

- **Rocky Flats Facility (Plutonium intakes):** Autopsy material is being used by Oak Ridge National Laboratory to assist in understanding the bioassay data (urine samples, chest counts) of over 700 workers with relatively high intakes of ^{239}Pu for accurate organ dose assessment
- The distribution of **beryllium**, a potential confounder, is to be examined for several autopsy cases and contrasted with radiation dose
- **Brain doses also have been estimated for workers at:** Los Alamos National Laboratory (^{238}Pu , ^{239}Pu), Malinckrodt Chemical Works (radium, uranium), Rocketdyne (uranium, plutonium, other) and will be for the Savannah River (plutonium) and Hanford (plutonium) sites



To Advance Knowledge the Future is the Integration of Biology (and Chemistry and Physics) with Epidemiology



Northwestern
University



National Council on Radiation
Protection and Measurements



VANDERBILT
UNIVERSITY



College of
Pharmacy and
Pharmaceutical Sciences
WASHINGTON STATE UNIVERSITY