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Medical Response to Radiation: Navigating Emergencies, Enhancing Survival

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Lt Col Justin Peacock serves as Associate Dean for Research at the Uniformed Services University School of Medicine. A nuclear radiologist, he focuses on molecular imaging and therapy research and the responsible integration of artificial intelligence (AI) in medical education—particularly how to teach AI-enabled tools and how AI can improve educational practice.

Lt Col Peacock earned a BS in Biochemistry (Honors) from Brigham Young University, a PhD in Molecular Biophysics and Biochemistry from Yale University (2010), and an MD from the Mayo Clinic Alix School of Medicine (2014). He completed a transitional year internship, diagnostic radiology residency, and nuclear medicine fellowship with the San Antonio Uniformed Services Health Education Consortium (SAUSHEC).

Across the National Capital Consortium and SAUSHEC, Lt Col Peacock has held education and operational leadership roles, including Nuclear Medicine Fellowship Program Director, Diagnostic Radiology Residency Assistant Program Director (Quality Improvement/Patient Safety), Associate Air Force Program Director, and Armed Forces Radiobiology Research Institute (AFRRI) Military Medical Operations Department Head and Educational Director. He has authored 45 peer-reviewed manuscripts, launched the AFRRI Virtual MEIR 3- and 5-day courses, served as a Dean's Fellow, and developed the USU Faculty Development AI curriculum. He also serves on national and international radiology and nuclear medicine committees.



Disclosures/Disclaimers



- Dr. Justin Peacock has no relevant financial or non-financial relationships to disclose relating to the content of this activity.
- The opinions and assertions expressed herein are those of the author and do not necessarily reflect the official policy or position of the Uniformed Services University or the Department of War.
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- The author will discuss off-label uses of medications and medications in current development.



Learning Objectives



At the conclusion of this activity, participants will be able to:

1. Identify the key signs and symptoms of acute radiation syndrome aiding in prompt patient triage and initial management.
2. Describe the protocols for medical response to radiation exposure, including immediate and supportive care measures.
3. Evaluate the effectiveness of current radiation prophylaxis and post-exposure treatments in mitigating the health impacts of radiation.



Polling Question



How comfortable do you feel responding to a potential radiation emergency?

- A. Not at all comfortable
- B. Slightly comfortable
- C. Moderately comfortable
- D. Comfortable
- E. Very Comfortable



Chernobyl, Ukraine (1986)



- Chernobyl nuclear plant explosion
- 50-185 million Ci released
- Many with long-term effects



(Taylor, A. 2019)





Fukushima, Japan (2011)



- Tsunami-induced power loss
- Hydrogen gas explosions due to lack of cooling
- Deaths from disaster, evacuation and one from radiation



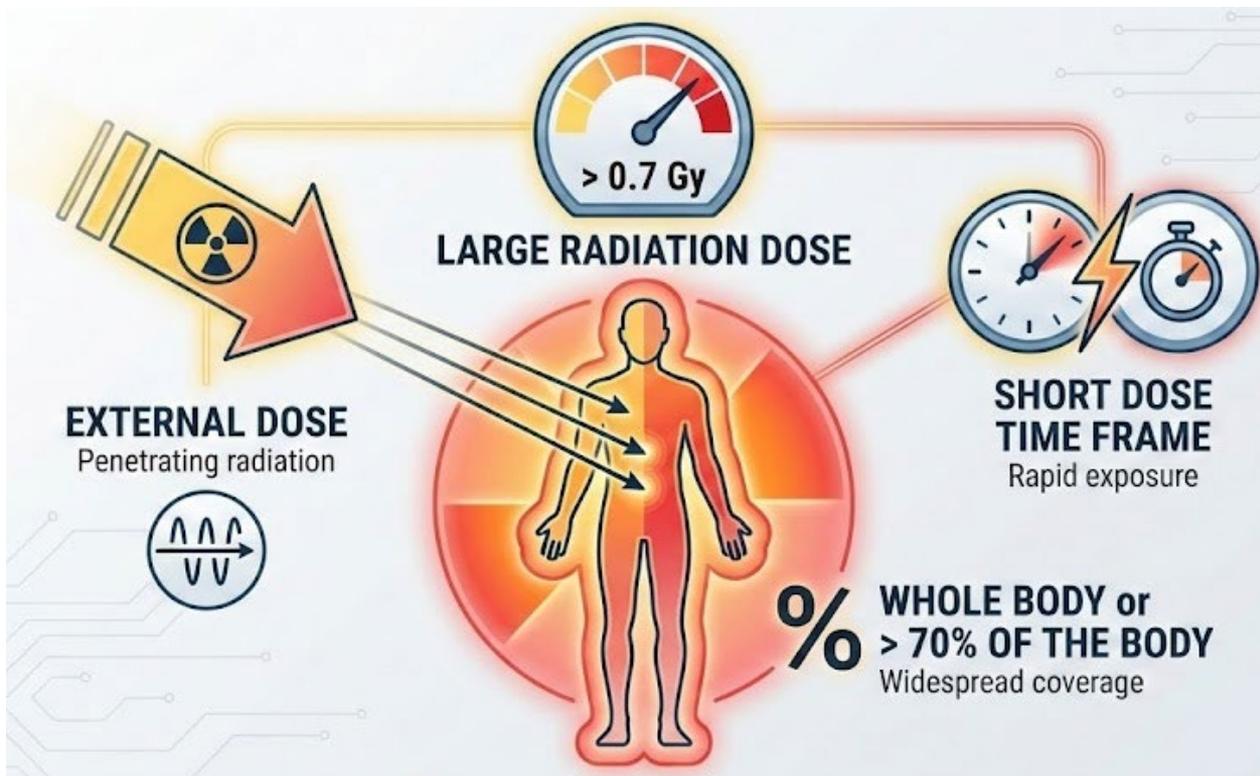
(Wikipedia contributors, n.d.)



Acute Radiation Syndrome (ARS)

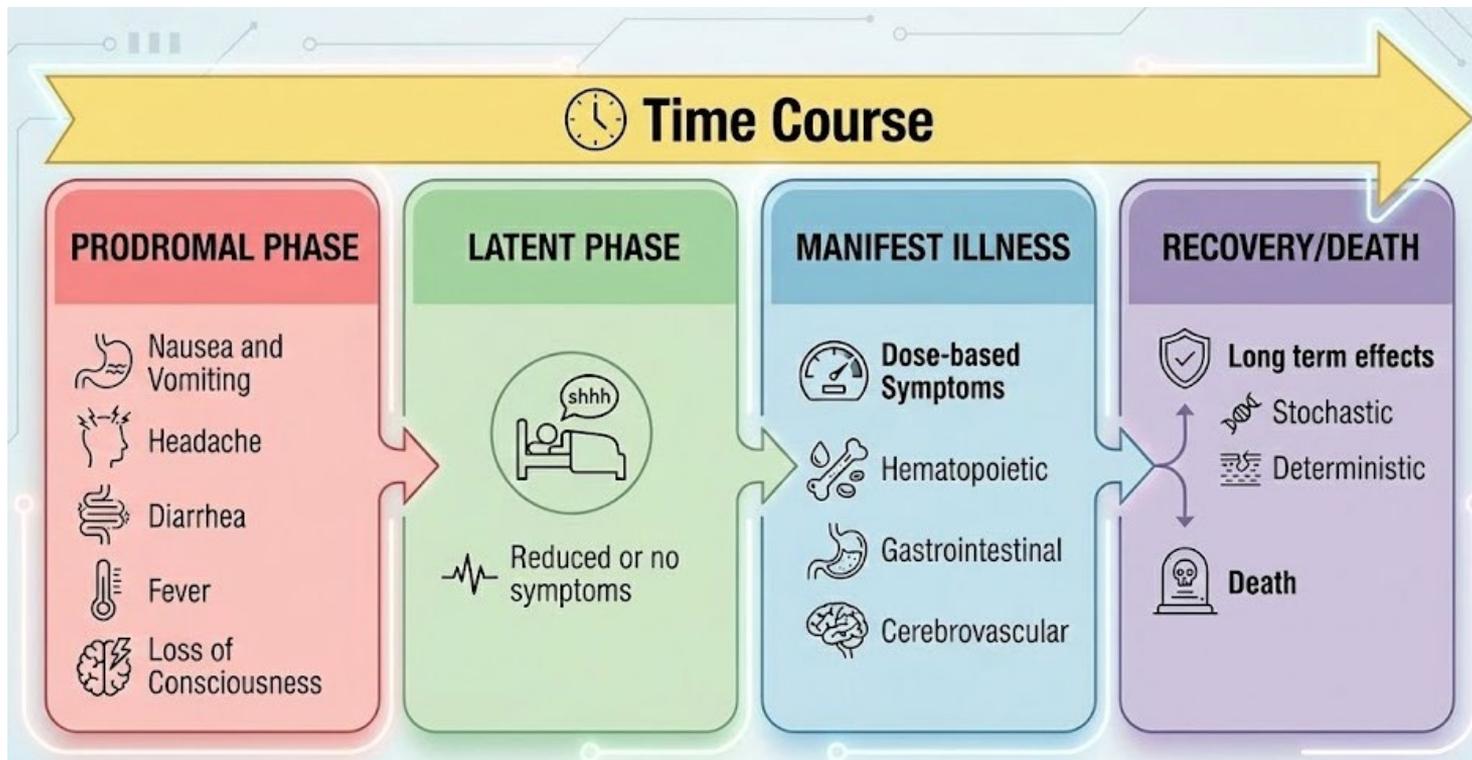


Acute Radiation Syndrome (ARS)



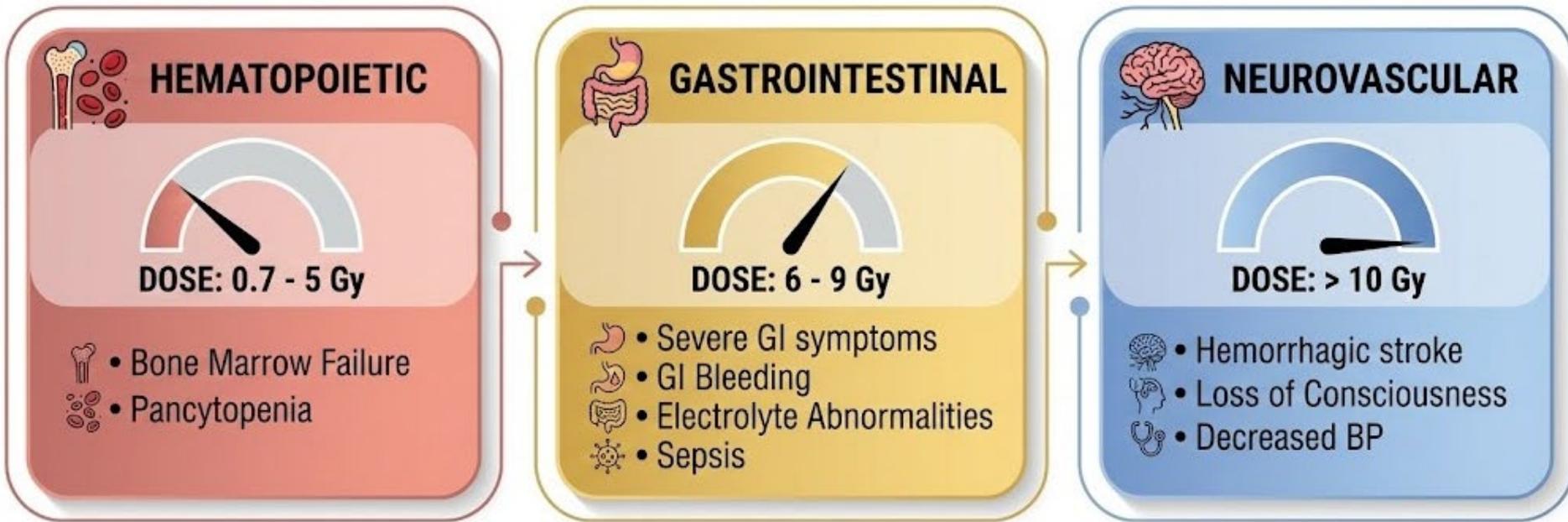


ARS Phases





ARS Subsyndromes



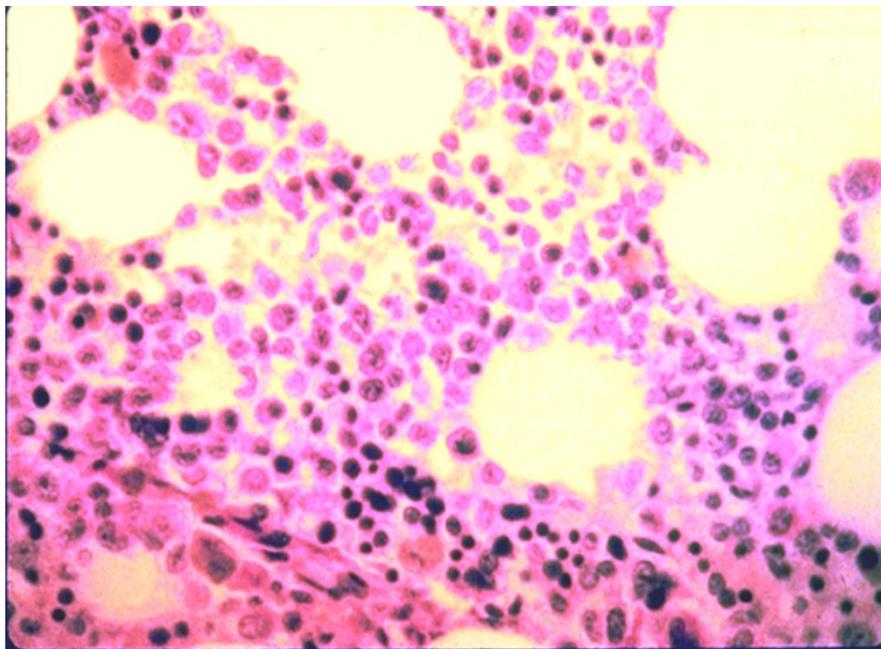
GI – gastrointestinal
BP – blood pressure



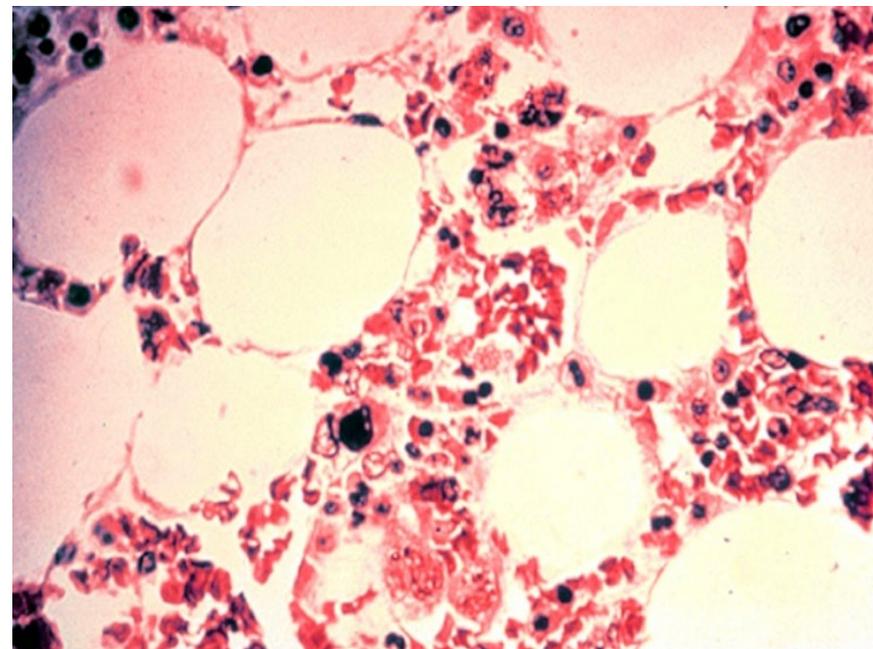
Hematopoietic Subsyndrome



Normal Marrow



Irradiated Marrow



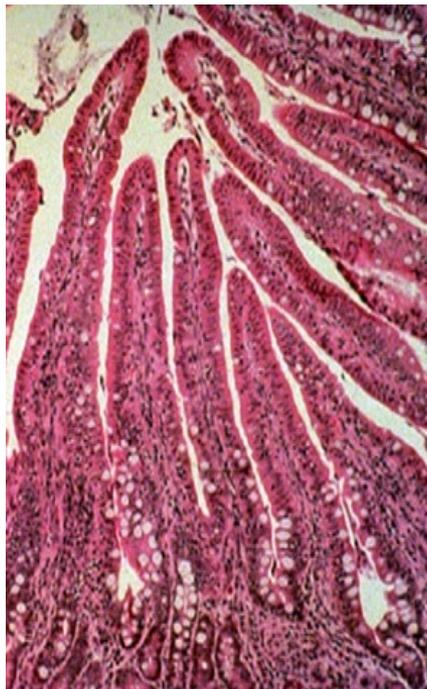
(Armed Forces Radiobiology Research Institute, n.d.)



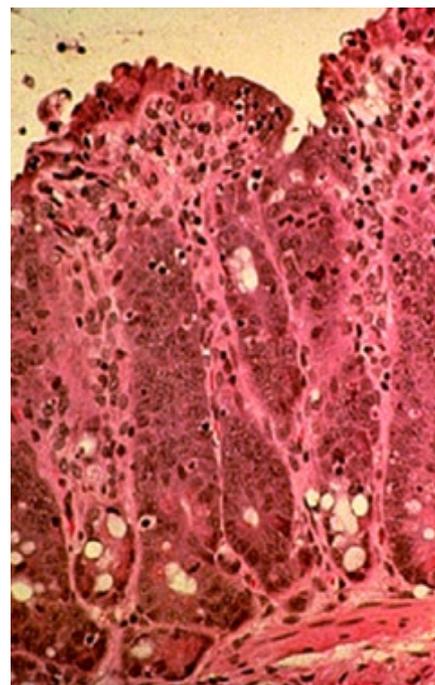
Gastrointestinal Subsyndrome



Normal GI Mucosa



Irradiated GI Mucosa



(Armed Forces Radiobiology Research Institute, n.d.)



Polling Question



Knowing about ARS, how would you go about triaging radiological-injured patients?



ARS Triage



Provider Protection



Class C (left)

- Consider for initial decontamination
 - Protects from airborne radioactive material
 - Protects from skin contamination
- Air-purifying respirator
- Face shield

Class D (right)

- Standard isolation personal protective equipment (PPE)
- Utilize after initial decontamination
- Exposure-only concerns





Triage Considerations



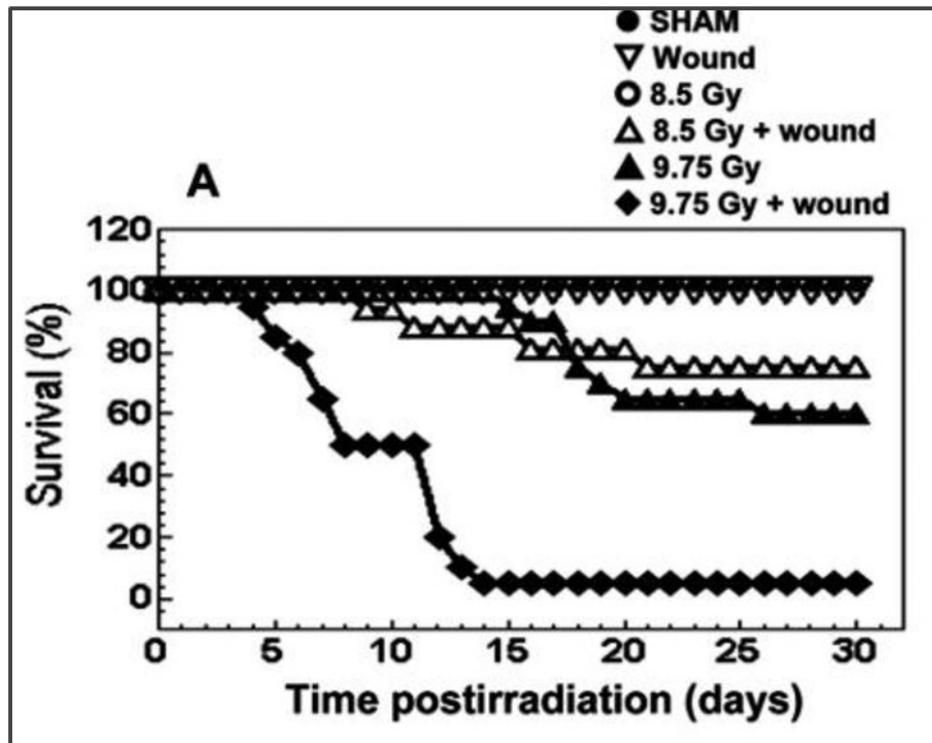


Don't delay life-saving care for decontamination!



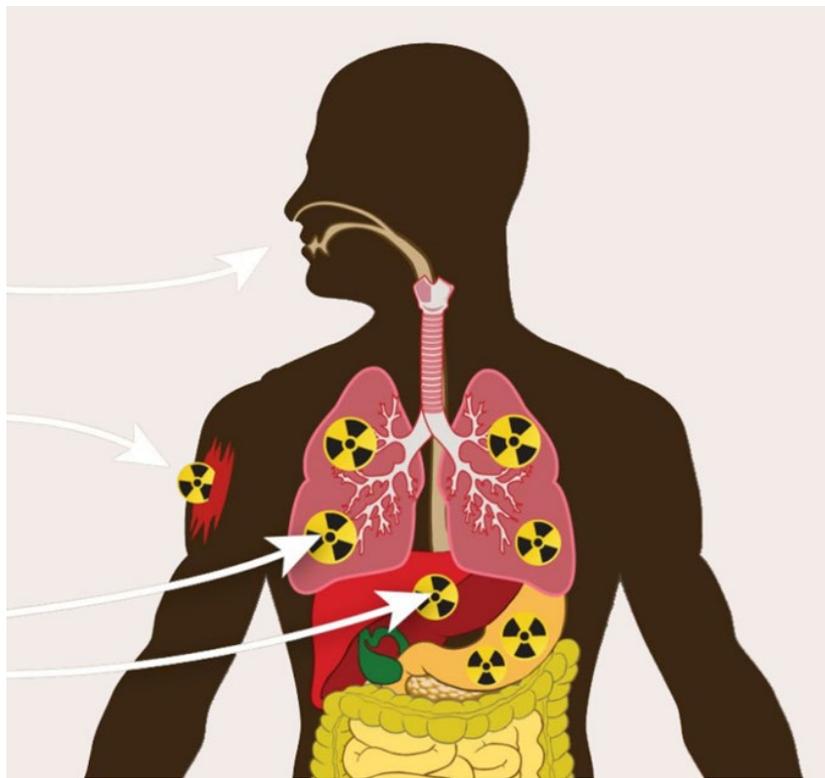


Combined Injury





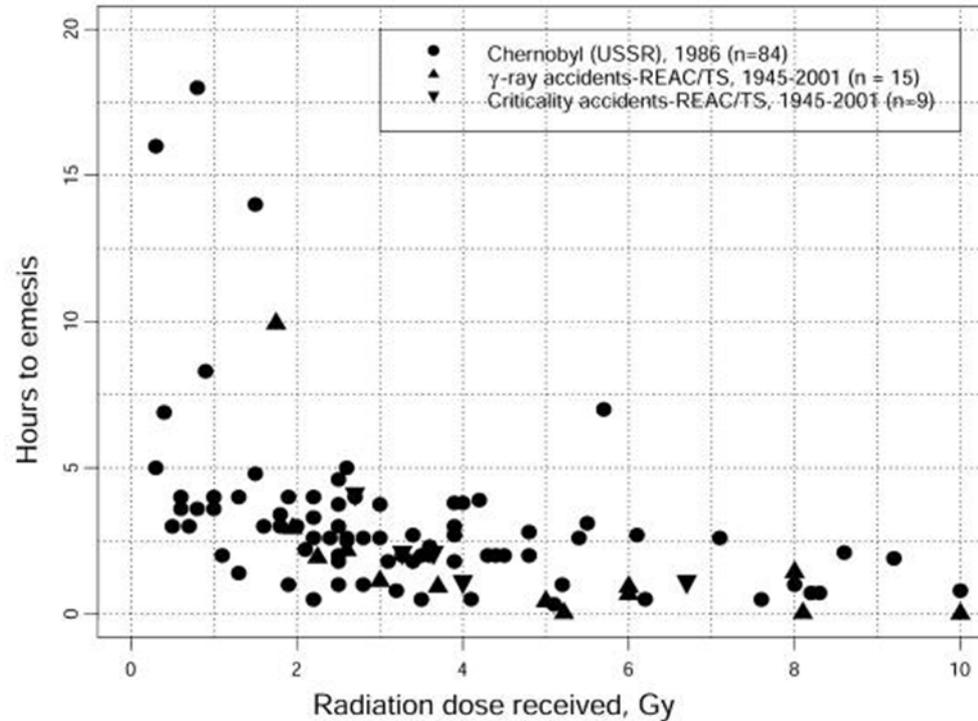
Internal Contamination



(Centers for Disease Control and Prevention, 2024)



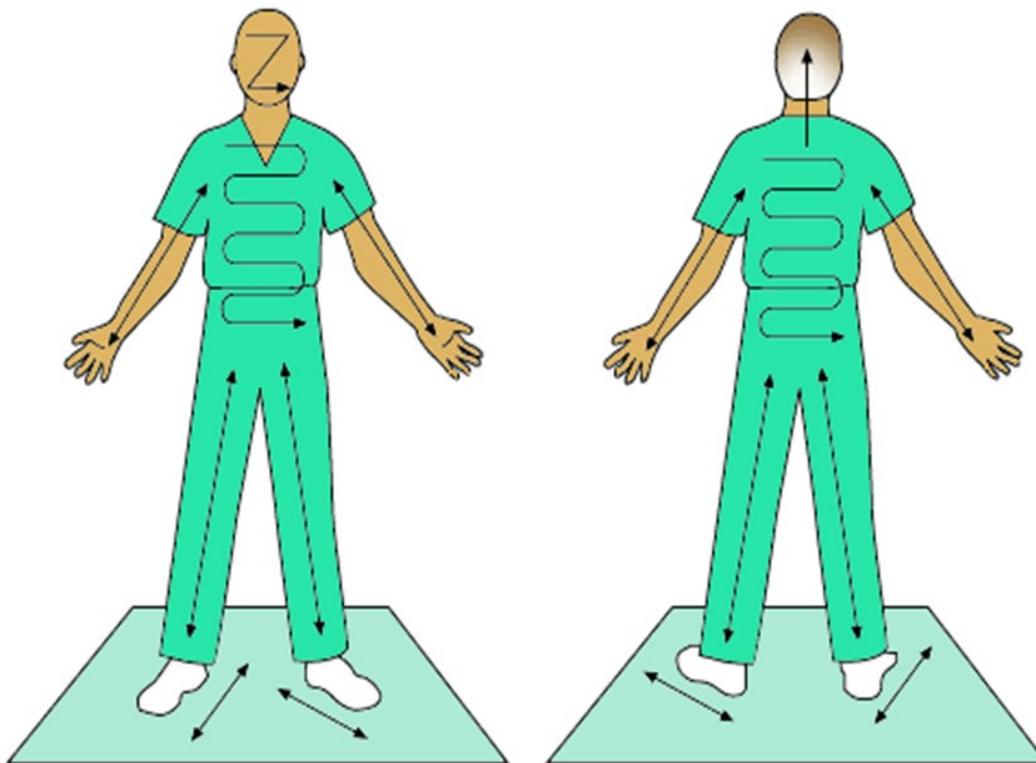
Time to Emesis



(Demidenko et al, 2009)



External Contamination Survey



(Radiation Emergency Medical Management, n.d.)



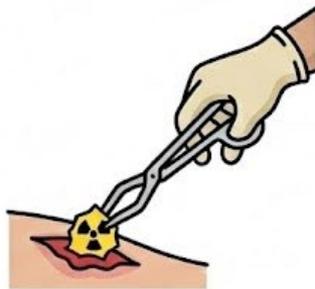
PROVIDER RADIATION SAFETY: HANDLING RADIOACTIVE SHRAPNEL

IDENTIFY THE RISK



Most significant radiation risk for providers

SAFE REMOVAL



Remove shrapnel with metal tongs/forceps

PROPER DISPOSAL

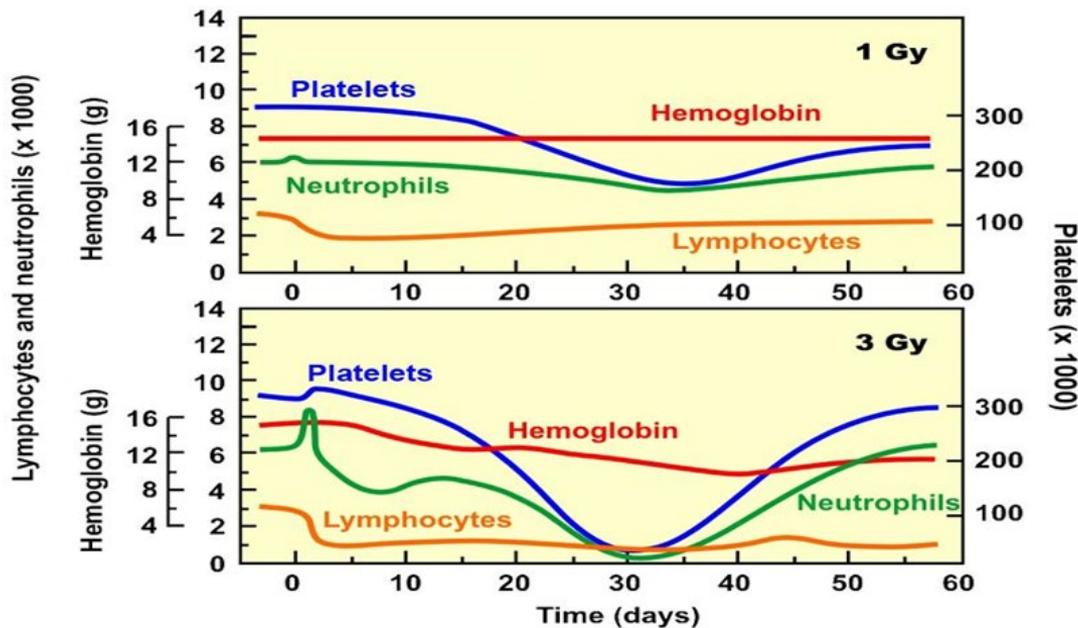


Dispose in containers per Radiation Safety Officer

Following these steps minimizes exposure and ensures safe handling of radioactive materials.

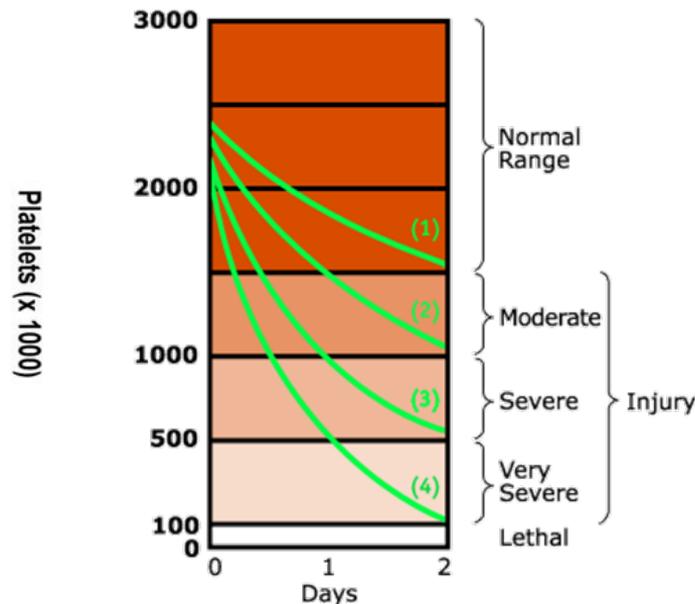


Lymphocyte Depletion Kinetics



(Armed Forces Radiobiology Research Institute, n.d.)

Patterns of early lymphocyte response in relation to dose.



(Radiation Emergency Medical Management, n.d.)



Radiation Emergency Triage Categories

Immediate

-  Life-threatening injuries
-  Combined injuries
-  Internal Contamination

Minimal

-  Radiation exposure without injury
-  Worried well

Delayed

-  Evidence of radiation injury (ARS)
-  Time to emesis < 4 hours
-  Lymphocyte drop

Expectant

-  High dose (> 20 Gy)



Triage by Resource Availability

Triage Category and Myeloid Cytokine Recommendation Category^X with Combined Injury (Radiation Exposure ≥ 2 Gy and Trauma/Burn*)

Est. Radiation Dose (gray)	Resource Availability at Medical Venue: "Normal or Good"		Resource Availability at Medical Venue: "Fair or Poor"	
	Moderate Trauma	Severe Trauma	Moderate Trauma	Severe Trauma
>10 Gy	Expectant ³	Expectant ³	Expectant ³	Expectant ³
≥ 6 Gy - 10 Gy	Delayed ²	Expectant ³	Expectant ³	Expectant ³
≥ 2 Gy - < 6 Gy	Immediate ¹	Delayed ²	Delayed ²	Expectant ³
	Resource Availability at Medical Venue: "Normal or Good" Standard of Care: Conventional to Contingency		Resource Availability at Medical Venue: "Fair or Poor" Standard of Care: Crisis	

(Radiation Emergency Medical Management, n.d.)



Initial Radiation Injury Management



Radiological Response Resources



- Armed Forces Radiobiology Research Institute (AFRRI)
 - Medical Radiobiology Advisory Team (MRAT)
 - <https://afri.usuhs.edu/>
- Radiation Emergency Assistance Center/Training Site (REAC/TS)
 - <https://orise.orau.gov/reacts/index.html>
- Radiation Emergency Medical Management (REMM)
 - <https://remm.hhs.gov/>
- Radiation Injury Treatment Network (RITN)
 - <https://ritn.net/>



Contaminated or Exposed?



Polling Question

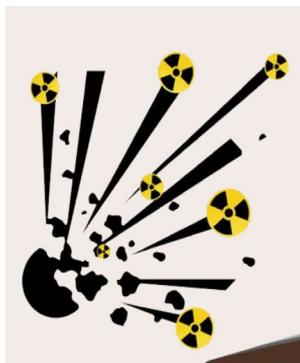
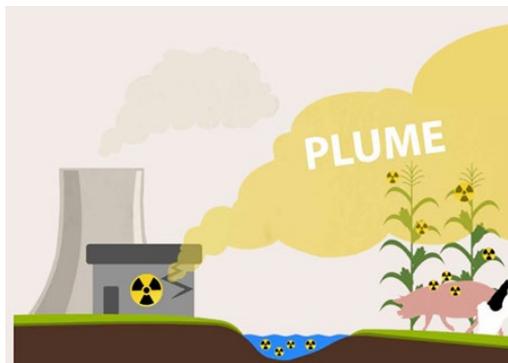


For each of the following scenarios, would patients primarily be contaminated or exposed?

- A. Contaminated
- B. Exposed



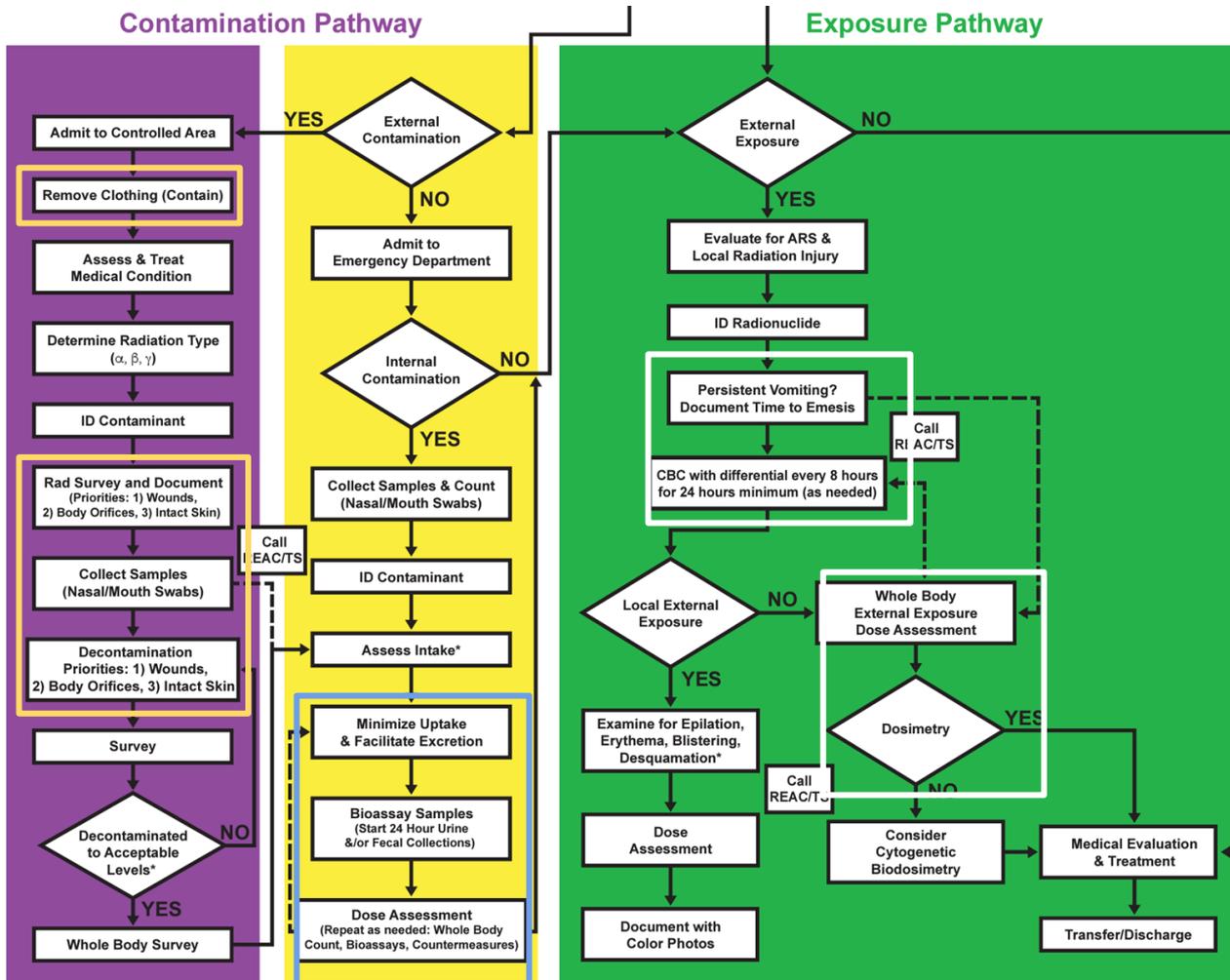
First, consider the source



(Centers for Disease Control and Prevention, 2024)

Management Algorithms

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(Oak Ridge Associated Universities, 2020)



Removing a patient's clothing and washing their skin and hair removes >90% of contamination.

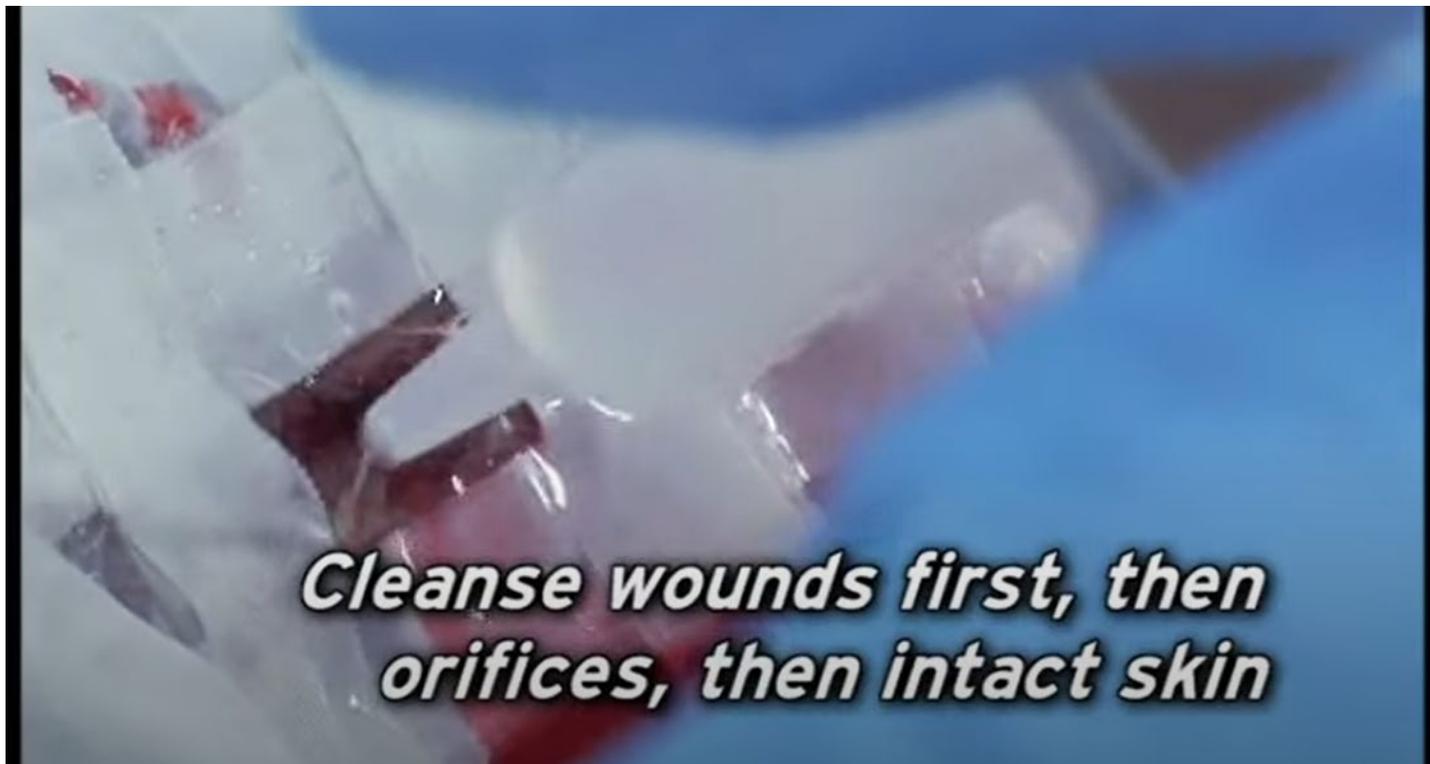


Nasal/Oral Sample Collection





Prioritized Decontamination



(Centers for Disease Control and Prevention, 2024)



Biodosimetry



**LOW RESOURCES,
RAPID ASSESSMENT**
(Hours)



- Time to emesis (< 4 hours, likely ARS)



**HIGH RESOURCES,
MEDIUM-TERM
ASSESSMENT** (Days)



- NM imaging and dosimetry techniques for internal contamination



**LOW RESOURCES,
24-48 HOUR
ASSESSMENT**
(Days)



- CBC with differential (48 hour lymphocyte depletion by 50%, > 4Gy dose)



**HIGH RESOURCES,
MEDIUM TO LONG-TERM
ASSESSMENT**
(Weeks/Months)



- Dicentric Chromosome Assay (DCA)
- FISH analysis of chromosome abnormalities
- Electron Paramagnetic Resonance (EPR)



Medical Radiation Injury Management



Supportive Measures





Transfusion Guidance



Criteria for Substitution Therapy for Hematopoietic Type Acute Radiation Syndrome

Patient's individual condition	Threshold value	Substitution therapy
Close monitoring possible, no other complication, no bleeding	Platelets: 10,000/ μ L	Irradiated and leukoreduced platelet concentrates
Close monitoring not possible, increased risk of manifest bleeding	Platelets: 20,000/ μ L	Irradiated and leukoreduced platelet concentrates
Additional trauma, surgery, mass transfusion, cerebral edema	Platelets: 50,000/ μ L	Irradiated and leukoreduced platelet concentrates
Anemia	Hemoglobin: 10 g/dL	Irradiated and leukoreduced packed red cells

(Radiation Emergency Medical Management, n.d.)



Internal Contamination Countermeasures

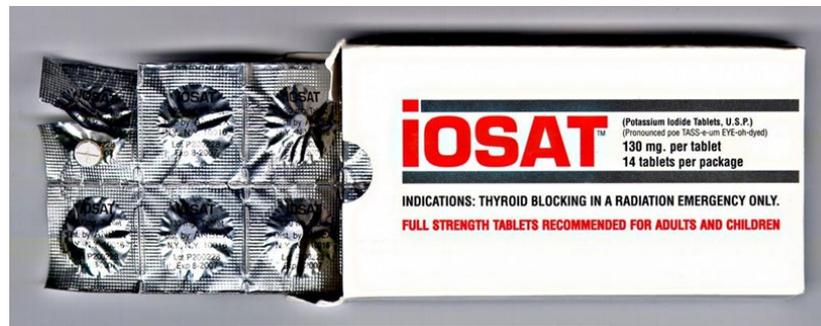
Radioisotope	Primary Target	Medical Treatment	How it Works
Radioiodine (^{131}I)	 Thyroid	Potassium Iodide (KI)	Blocks uptake: Saturates the thyroid with "good" iodine so it can't absorb the radioactive version.
Radiostrontium (^{90}Sr)	 Bone	Calcium / Strontium Salts	Competitive Inhibition: Floods the body with stable minerals to reduce the "bone-seeking" isotope's absorption.
Tritium (^3H)	 Whole Body Water	Fluids & Diuretics	Dilution & Flush: Increases the turnover of body water to accelerate the natural excretion of radioactive water.
Cesium (^{137}Cs)	 Muscle / Soft Tissue	Prussian Blue	Ion Exchange: Traps cesium in the intestines, preventing it from being reabsorbed and moving it out through waste.
Transuranics (Pu, Am, Cm)	 Lungs / Liver / Bone	Chelation (DTPA/EDTA)	Chemical Binding: Acts like a "claw" that grabs heavy metal atoms in the blood so the kidneys can filter them out.



Potassium Iodide (KI)



- Indication: Protection against radioiodine exposures.
- Administration: Oral tablet (130 mg or 65 mg) & solution (65 mg/mL)
- Timing:
 - Ideally 24 hours before exposure, but up to 4 hours post exposure. Then every 24 hours.
- Dosing:
 - Adult: 130 mg
 - Children: Varies based on age
- Prioritization:
 - Pregnant women, neonates and young children

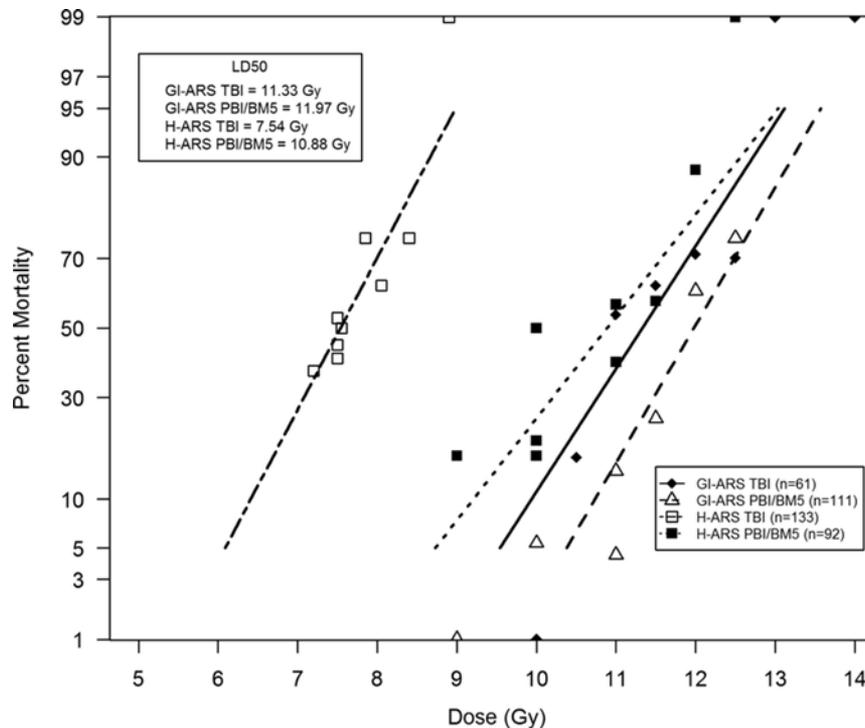




Cytokine Countermeasures



- FDA-approved cytokines
 - Filgrastim (Neupogen®)
 - PEGylated filgrastim (Neulasta®)
 - Sargramostim (Leukine®)
- Predominantly promote neutrophils
 - Leukine promotes many cell lineages



(MacVittie et al, 2015)



Filgramostim (Neupogen®)



- Indication: Myelosuppressive acute radiation doses (≥ 2 Gy)
- Administration: Subcutaneous injection (300 mcg and 480 mcg vials)
- Timing:
 - Begin as soon as suspected or confirmed exposure to ≥ 2 Gy
 - Daily injections until absolute neutrophil count (ANC) greater than 1000/mm³ for three consecutive daily complete blood counts (CBCs)
- Dosing:
 - Adults: 10 mcg/kg (can start with 5 mcg/kg depending on supply)
 - Children: 5 mcg/kg



PEGylated filgrastim (Neulasta®)

- Indication: Myelosuppressive acute radiation doses ($\geq 2\text{Gy}$)
- Administration: Subcutaneous injection (0.6 mL syringe)
- Timing:
 - Begin as soon as suspected or confirmed exposure to $\geq 2\text{ Gy}$
 - Two doses, one week apart
- Dosing:
 - Adults and children weighing $\geq 45\text{ kg}$: 6 mg per dose
 - Children: Variable based on weight



Sargramostim (Leukine®)



- Indication: Myelosuppressive acute radiation doses (≥ 2 Gy)
- Administration: Subcutaneous injection (250 mcg vials)
- Timing:
 - Begin as soon as suspected or confirmed exposure to ≥ 2 Gy
 - Daily injections until ANC greater than 1000/mm³ for three consecutive daily CBCs
- Dosing:
 - Adults and children weighing ≥ 40 kg: 7mcg/kg
 - Children weighing 15-40 kg: 10 mcg/kg
 - Children weighing <15 kg: 12 mcg/kg



Platelet Radiation Countermeasures



- FDA-approved thrombopoietin (TPO) receptor activator
 - Romiplostim (Nplate[®])
 - Promotes megakaryocytes and platelets
- Additional platelet promoters are in development, including recombinant TPO, TPO activators, and IL-11.

Drug	Irradiation dose	Drug dose (µg/kg)	Injection time post-irradiation (days)	Group size (n=)	Percent mortality	Platelet Nadir (×10 ⁹ /L)	Duration of thrombocytopenia (days)
Vehicle	~LD70/60	Vehicle	1	40*	67.5	12.5	3.8
Nplate	6.8 Gy	5	1	40*	27.5	36.3	1.3
Nplate + PEGfilgrastim		5 and 0.3	1 (both drugs), 8 (only PEGfilgrastim)	40*	12.5	31.5	1.1

(MacVittie et al, 2015)



Romiplostim (Nplate[®])



- Indication: Myelosuppressive acute radiation doses (≥ 2 Gy)
- Administration: Subcutaneous injection (125, 250, or 500 mcg vials)
- Timing:
 - Begin as soon as suspected or confirmed exposure to ≥ 2 Gy
 - Single dose
- Dosing:
 - Adults and children (including neonates): 10 mcg/kg



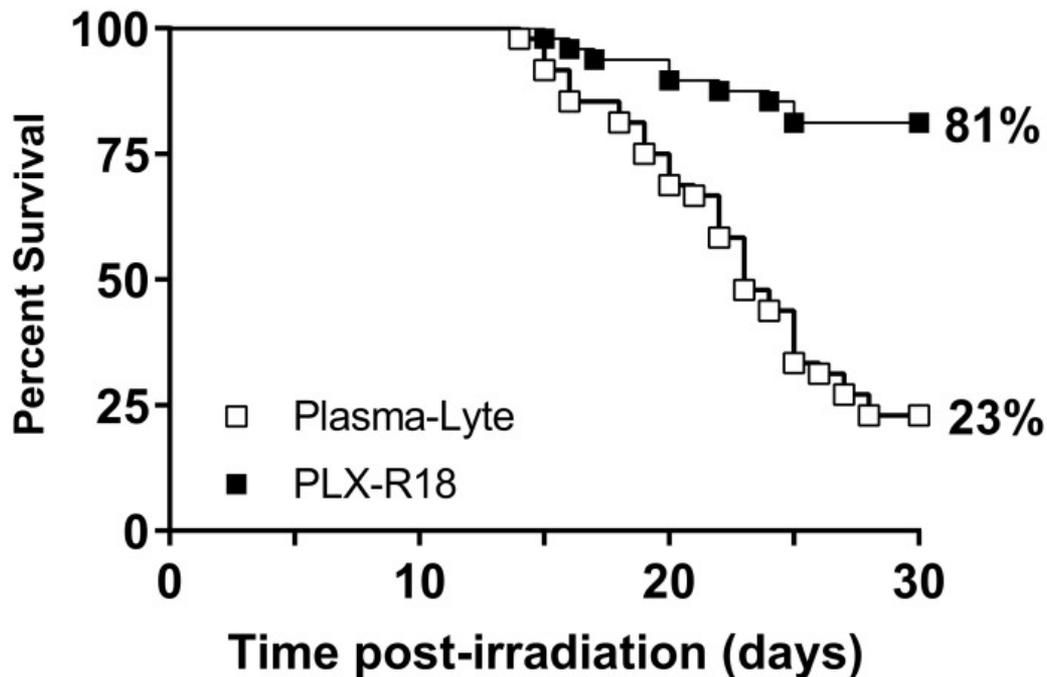
Stem Cell Transplant



- Allogeneic and syngeneic stem cell transplants have been attempted in several cases of high radiation exposure.
- In a review of 29 historical bone marrow transplant cases
 - Three patients survived longer than one year
 - Many died from GI subsyndrome conditions, traumatic injuries, or graft vs host disease
 - Engraftment may occur around the same time as the patient's own bone marrow reconstitutes



Stromal Cell Therapy



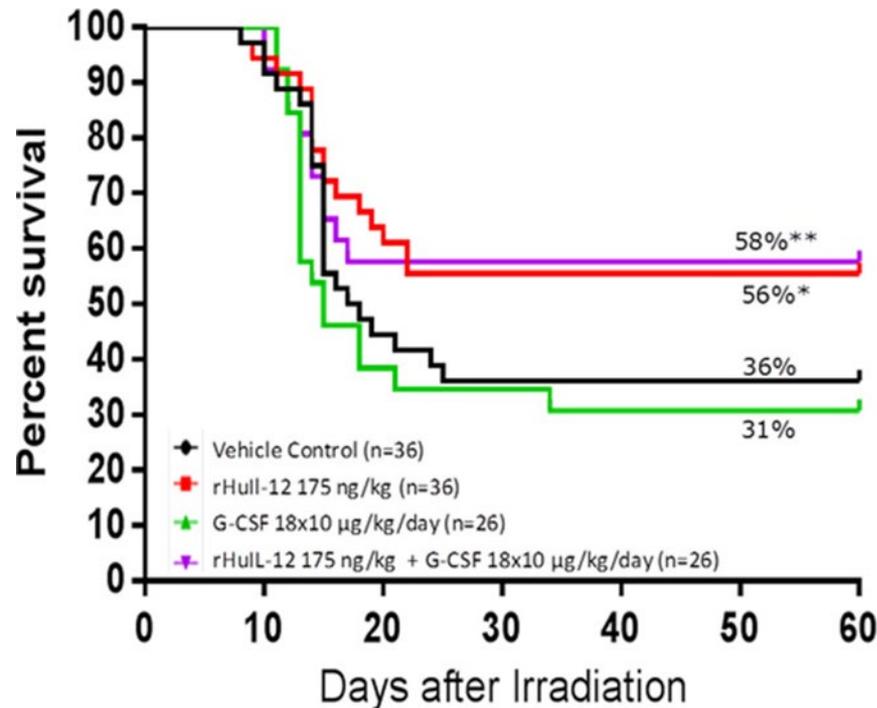
(Kumar et al, 2022)



IL-12 Countermeasures



- rhIL-12 (HemaMax®)
- Pro-inflammatory cytokine
 - Improves hematopoiesis
 - Improves GI function
 - Improves survival



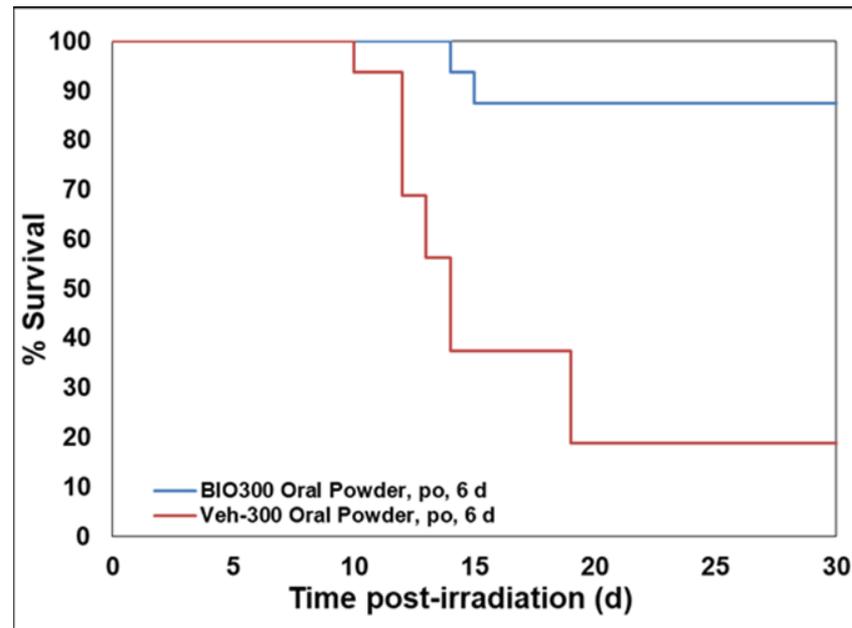
(Gluzman- Poltorak et al, 2014)



Genistein (BIO300®)



- Phytoestrogen that binds ERs
- Protects progenitor cells from radiation damage
 - Increases cell cycle checkpoints and DNA repair enzymes (inhibits NF-kB)
 - Reduces inflammatory cytokines (anti-oxidant)
 - Captopril combination increases radioprotection
- Mitigates radiation-induced pneumonitis/fibrosis

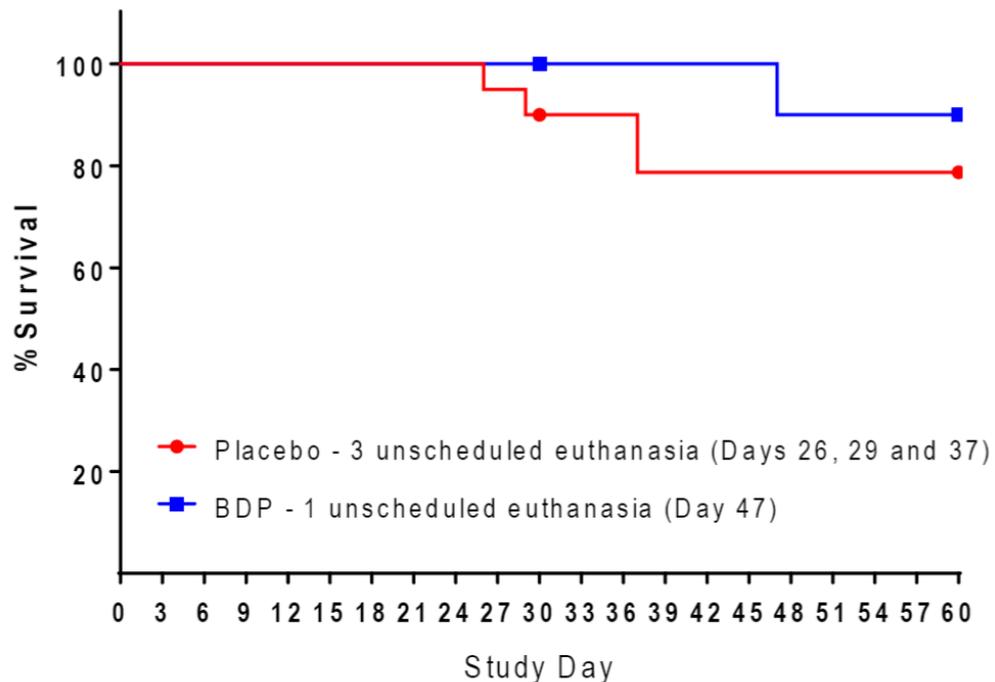


(Singh & Seed, 2020)

(Image courtesy of Dr. Singh)



OrbeShield® for GI-ARS



(Measey et al, 2016)



Prophylactic Medications in Development



- Piperazine derivatives
- Chlorobenzylsulfones (example Ex-RAD)
- Aminothiols (example amifostine)
- Enalapril

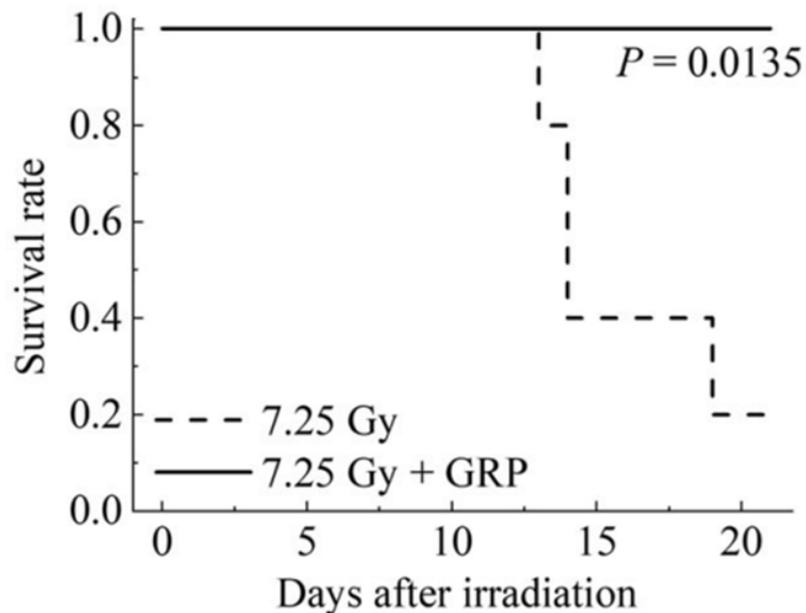




Combination Therapies



- ARS is complicated, impacting many systems.
- Combination therapies likely address the multi-system effects.
- rhG-CSF (Neutrogin®) and human TPOR agonist RP (Romiplat®) demonstrate improved survival in lethal dose mice models (right).



(Hirouchi et al, 2015)



Long-Term Effects of Radiation Exposure (Post-ARS/Injury)



STOCHASTIC EFFECTS

(More radiation, greater statistical likelihood)

-  Cancer



DETERMINISTIC EFFECTS

(More radiation, greater severity)

-  Cataract (Most Common)
-  Infertility
-  Atherosclerotic Disease
-  Pulmonary Pneumonitis/Fibrosis



CONCLUSION: Patients surviving ARS/radiation injury require ongoing patient care for these long-term risks.



Polling Question



Now, how comfortable do you feel responding to a potential radiation emergency?

- A. Not at all comfortable
- B. Slightly comfortable
- C. Moderately comfortable
- D. Comfortable
- E. Very comfortable



Key Takeaways



- Healthcare provider exposures are usually low after major nuclear events.
- Don't delay ABCs for decontamination—treat the life-threatening injuries first.
- ARS is predictable in its phases and dose-dependent in its timing and severity.
- Triage and treat according to resources—consider simple over more advanced measures when resources are limited.
- Patients exposed to radiation need supportive care (e.g., fluids, antimicrobial agents) and possible bone marrow support (e.g., G-CSF/GM-CSF), similar to oncology patients.



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