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Cautions

- Authored by <u>REMM</u> and <u>RITN</u> physicians, this set of orders is a prototype only.
- Orders must be customized for each patient and incident.
- Specific drugs are suggested for function only. Patients may not need any/every category of drug listed.
- No HHS, CDC, FDA, or other US government entity endorsement of specific drugs or drug doses is intended or implied by inclusion in this order set.
- Consult the notes at the end of this document for additional, key information.

Internal contamination (decorporation treatments)

- This **Adult Orders Prototype** lists only FDA-approved medications as radioisotope countermeasures.
- Some, but not all of these drugs are currently in the Strategic National Stockpile.
- Prescribers should consult the FDA drug label for complete prescribing information.
- Decorporation drugs should be used in children with great caution.
- The online version of REMM has additional recommendations about <u>additional</u> <u>countermeasure drugs that may be considered</u>.
- This prototype does **not** address threshold levels of <u>internal contamination</u> that would trigger initiation, continuation, or discontinuation of decorporation treatment. See <u>REMM Countermeasures Caution and Comment</u>, which discusses this issue

Drug dosages

- All adult drug doses in this prototype are based on a 70 kg adult with normal renal and hepatic function.
- Appropriate dose adjustments should be made based on age, weight, drug-drug interactions, nutritional status, renal, and hepatic function.

• If this adult order set, **Version date 4/17/2017**, has been printed for use offline, consult the online version of REMM to see if updates are available. <u>https://www.remm.nlm.gov/adult-order.pdf</u>

[•] After a mass casualty incident, practitioners may encounter counterfeit drugs. This <u>FDA website</u> will provide information on avoiding and detecting counterfeit drugs and assist with reporting of suspected counterfeit medications.

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Unique Identifier	
Jnique Identifier:	
Address:	
Phone:	-
Spoken language:	
Unaccompanied minor:	
Date of Birth:	_
Age (years:	
Gender:	
Next of kin contact informati	ion (home phone, cell phone, email, or address):

2. Admit to:

Inpatient Service	Area
Team:	PICU
Hem/Onc:	Hematopoietic Stem Cell Transplantation:
Admitting Physician:	Pager:
Attending Physician:	Pager:
Other Physician:	Pager:

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3. Diagnoses

Acute/Chronic Non-radiation Related Admission Diagnoses:

a
b
C
d
e
f
Acute Radiation-related Admission Diagnoses:
a. Radiation contamination? Yes No
See REMM Body Chart (page 19) to record whole body radiation survey.
External contamination with Isotope (Specify or unknown)
Internal contamination with Isotope (Specify or unknown)
Contamination suspected, Isotope uncertain
b. Radiation Exposure / Acute Radiation Syndrome (ARS)?
Yes No
 Estimated whole body dose from exposure(units of gray/Gy)
 See also Item #24, page 11 for additional radiation details and work-up
Other potential complicating factors
Mass casualty incident
Other, Specify
Specific populations potentially requiring more customized management?
Yes No
Age > 65 y

___ Pregnant/Possibly pregnant Duration of Pregnancy (weeks): _____

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- ___ Immunosuppressed
- ___ Other, Specify _____
- See REMM page about <u>at-risk populations</u>

4. Precautions:

Infectious

- ___ Contact
- __ Droplet
- ___ Airborne
- ___ Reverse Isolation/Neutropenic

Radiation precautions

- For persons with known or suspected external or internal contamination.
- Persons with <u>exposure</u> but NO <u>contamination</u> are NOT radioactive.
- Patients with exposure only do not need Radiation Precautions.
- __ Precautions: Single room, gown, mask, cap, boots, and gloves
- Use medical facility procedures for discarding all biological/physical/radioactive waste, including linens/towels/trash/personal protective equipment.
- Contact Radiation Safety Officer for additional instructions. Phone: Pager:
- Place Radiation Safety Sign on door if patient has internal or external radioactive contamination
- ___ Notify pregnant staff that entry to room is prohibited if patient is/may be contaminated.
- Everyone entering room/area of contaminated patient must wear personal radiation dosimeter assigned by Radiation Safety.
- Use medical facility procedures for disposal of radiation waste, including linens/towels/trash/personal protective equipment.

• See guidance

 <u>2007 Guideline for Isolation Precautions: Preventing Transmission of</u> <u>Infectious Agents in Healthcare Settings</u> Healthcare Infection Control Practices Advisory Committee (HHS/CDC)

5. Urgent consultations: specify	
Intensive Care	Transfusion Medicine
Hematopoietic Stem Cell Transplantation	Radiation Oncology
Mental Health / Psychiatry	Endocrinology
Ophthalmology	Palliative Care and Pain Service
Dermatology / Plastic Surgery	Gastroenterology
Radiation Safety	Burn Therapy
Surgery:GeneralTraumaThor	acic Orthopedics
Hepatology	Infectious Disease
Pulmonary	Plastic Surgery
Cardiology	Nephrology
ENT	
Other	
6. Condition:	
Good Fair Stable Gu	arded Critical
7. Vital Signs:	
q 2 hours X 4 Ward routine q 4 hours X 4	e
	Other:

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8. Allergies:

- ___ No Known Drug Allergies (NKDA) ___ Allergies (drugs, foods) If yes, specify: ____ 9. Activity: ___ Bed rest ___ Bathroom privileges ___ Out of bed/up to chair every ____ hrs. ___ Ambulate as tolerated ___ Confine to room 10. Diet: ___ Regular Diet ___ Liquids (full, clear) ___ NPO ___ Advance as tolerated ___ Neutropenic diet ___ Special dietary needs/requests: 11. Height, weight: Height: ____ cm Weight: ____ kg Repeat body weight: q ____ days q _____ hours 12. Admission studies: Labs ___ CBC w/differential ____ w/ Platelet count ___ Comprehensive Metabolic Panel (CMP) / Chem 14 ___ PT or INR/PTT/fibrinogen/TT ___ Urinalysis - Collection method: _____ ___ Urine culture ___ Blood culture - Collection method: _____ Sets: _____ Type of culture: Bacteria, fungal, aerobic, anaerobic ___ Sputum culture ____ Urine HCG (for all girls \geq 10 years or post-menarche, whichever is earlier)
- ___ Serum HCG (for any girls ≥10 years or post-menarche, whichever is earlier)

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___ Thyroid Function Tests (Specify) _____

____ Wound cultures

Serologies:

- ____ Herpes Simplex Virus type 1 (HSV-1)
- ____ Herpes Simplex Virus type 2 (HSV-2)
- ___ Cytomegalovirus (CMV)
- ____ Varicella-zoster virus (VZV)
- ___ Epstein Barr Virus (EBV)

13. Standing labs / studies

- ___ CBC w/diff and platelets q ____ hours, x ____ days, Followed by q ____ until further orders
- Comprehensive Metabolic Panel (CMP) / Chem 14 Followed by q _____ hours, x _____ days Followed by q _____ until further orders

___ Other _____ (specify test and frequency)

14. Blood bank

(May set institutional transfusion parameters, e.g.: PRBC transfusion for Hgb < (7 g/dl) and platelet count < 20000/micL unless otherwise specified by medical staff.)

___ Type and cross match

___ Type and screen

For _____ units or _____ ml of packed red blood cells (~10-15 ml/kg) For _____ units or _____ ml of platelets (~5-10 ml/kg)

Note:

- Use only leukoreduced AND irradiated products, if available, unless it is known with certainty that the patient was exposed to allow dose of radiation, e.g. less than 100 cGy.
- If radiation whole body dose is not known with certainty, leukoreduced AND irradiated products are preferred, if available.
- See <u>REMM blood use page</u> for additional information.

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15. Imaging

___ Chest x-ray Urgency:_____ ___ PA/Lateral Urgency:_____ ___ Portable Urgency: _____ __ Other imaging studies Specify: _____ Urgency: _____ 16. Electrocardiogram ___ Electrocardiogram ___ STAT Electrocardiogram for chest pain, notify physician 17. IV fluid management: ___ IV Fluids: _____ @ _____ cc/hr, with additive _____ ___ IV Fluids: _____ @ _____ cc/hr, with additive _____ 18. __ Foley catheter management (specify) _____ ____ Use radiation precautions for urine and feces for patients with internal radiation contamination. 19. __ Monitor I / O Frequency _____ ___Use radiation precautions for urine and feces for patients with internal radiation contamination. 20. Deep Venous Thrombosis (DVT) prophylaxis: ____ TED hose to Bilateral Lower-Extremities ___ Sequential Compression Devices (SCD) ___ Anticoagulation regimen _____

__ Other

Note: The potential benefit of any anticoagulation regimen (e.g. **heparin**) should be balanced against the risk of excessive bleeding in patients with severe thrombocytopenia or significant gastrointestinal toxicity.

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21. Respiratory Therapy:

- ____ Use radiation precautions for personnel, equipment, and waste if patient has internal radiation contamination.
- ___ Room air ___ Chest tube care (Specify)_____
- ____ Titrate oxygen supplementation for Oxygen saturation > _____%
- ___ Nebulizer treatment (Specify) _____

22. Wound care: (see also item 25)

- ____ Decontaminate external wounds if there is external radiation contamination. See REMM radiation <u>contaminated wound</u> care recommendations.
- ___ Sterile dressing to wounds daily
- ___ Monitor waste
 - Use medical facility procedures for discarding biological/radioactive/physical waste and linens/towels/trash/personal protective equipment.
 - ___ Radiation precautions (needed if patient has radiation contamination)
- ____ Silvadene (Silver Sulfadiazine) cream topically to burns
- ___ Bacitracin topically to burns
- ___ Plastic Surgery Consultation
- ___ Other wound management per Burn team/Dermatology/Surgery: Pager ______ Phone _____

23. Orthopedic care:

- ___ Splint/brace/cast/crutches
- Other orthopedic management procedure per orthopedics:
 Pager _____ Phone _____

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24. Radiation Dose Assessment

A. Biodosimetry and Bioassay assays

- Difference between Biodosimetry and Bioassay
- <u>Define biodosimetry</u>
- More about biodosimetry
- <u>Dicentric chromosome assay</u>

B. Biodosimetry assays for radiation exposure

- See REMM information on
 - Dose Estimator for Exposure: 3 biodosimetry tools
 - Dose Reconstruction
- Estimated whole body dose from exposure: _____ (Gray)
 - Using which tool(s) e.g., vomiting, lymphocyte depletion kinetics, dicentric chromosome assay Note: if different assays give different results
- METREPOL Scores: Heme____ GI____ Neuro____Cutaneous_____
- Response Category (RC score) ______ <u>Explain METREPOL</u> <u>Consider Response Category in clinical triage</u> (Interactive tool for ARS)
- Date of exposure: ______
- Time of exposure: ______
- Location of patient at time of exposure:_____
- Estimated whole body/partial body dose, specify _____ (dose)
- Dose unknown: _____

Dicentric Chromosome Assay Instructions:

- Draw extra green top tube and provide: date _____ time _____
- See REMM for location of approved US laboratories that perform this test.
- Send this tube **ON ICE** for outside lab study

To the attention of: ______

- Name of lab:_____
- Address of lab:
- C. Radiation bioassay for evaluating/managing internal decontamination
 - Collect ≥ 70 mL spot urine for _____ (name of radioactive isotope)
 - See directions for sample collection, labeling, packaging and shipping bioassay specimen to CDC bioassay lab: <u>https://emergency.cdc.gov/radiation/labinfo.asp</u>

Note: Consult senior radiation event medical managers for name and location of other laboratories that may become available to perform this test in a large mass casualty incident. Routine labs generally cannot perform this test, although in large incidents, senior managers may announce special arrangements.

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25. General Medications:

- Drug names are generally listed as follows Generic (Brand) names
- Some drugs with **bold blue font** have **DailyMed** hyperlinks with additional information.

For gastric acid suppression:

____ Lansoprazole (Prevacid) 15-30 mg PO daily

For radiation-induced nausea & vomiting:

- __ Ondansetron (Zofran) 4-8 mg IV/PO q 8h PRN nausea/emesis
- Lorazepam (<u>Ativan</u>) 0.5 mg 1 mg PO q 6-8h PRN anxiety/insomnia/breakthrough nausea
- <u>Prochlorperazine</u> 10 mg PO/IV/IM (if adequate platelets) q 6-8h PRN anxiety/insomnia/breakthrough nausea

See REMM bibliography on treatment of nausea and vomiting

For fever:

____ <u>Acetaminophen_</u>650 mg PO q 6 – 8h PRN temperature> 38 °C

For diarrhea:

__ Loperamide hydrochloride (Imodium):

- Recommended initial dose is 4 mg (2 capsules) followed by 2 mg (1 capsule) after each unformed stool.
- Daily dose should not exceed 16 mg (8 capsules)

For rash:

- ____ Topical sterile dressing
- Diphenhydramine hydrochloride (Benadryl) 25-50 mg PO q 4-6 hours for pruritis, not to exceed 300 mg/24 hours

For pain:

- ____ Morphine sulphate _____ mg ____ route _____ frequency
- ____ Other pain medication (specify): name, dose, route, frequency

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For skin burns: (see also item 18: wound care)

Burn topical regimen
Replace body fluid
Other burn therapy

For oral mucositis:

Mouth care regimen _____

26. Radioisotope decorporation or blocking agents:

- Note: Only FDA approved radiation countermeasures are listed in table below.
- See <u>REMM Table</u> for longer list of countermeasures which have been recommended by some experts but are not FDA approved as radiation countermeasures.

Medical	Administered for	Route of Administration	Dosage	Duration
Countermeasure				
Ca-DTPA ^{1,3}	Americium	IV ¹ :	IV:	 Ca-DTPA for
Zn-DTPA ^{1,3}	(Am-241) ¹	Give once daily as a	1 g in 5 cc 5%	the first dose
		bolus or as a single		 Give Zn-DTPA
	Californium		water (D5W) or	for any follow-
information.	(Cf—252) ²	fractionate the dose.	0.9% sodium	up doses (i.e.,
			chloride (normal	
	Cobalt		saline, NS) slow	as indicated)
DTPA drug label.	(Co-60) ²	for intravenous Rx of	IV push over 3-	
			4 minutes	therapy
	Curium	internal contamination		depends on
DTPA drug label.	(Cm-244) ¹		OR	total body
		only.		burden and
	Plutonium		1 g in 100-250	response to
	(Pu-238 and		cc D5W or NS as	treatment
	Pu-239) ¹		an infusion over	
		Nebulized	30 minutes	
	Yttrium	inhalation ¹ :		
			Nebulized	
		for nebulized inhalation		
		in adults only, and if	1 g in 1:1	
		the route of	dilution with	
			sterile water or	
		through inhalation.	NS over 15-20	
			min	

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Medical	Administered	Route of	Dosage	Duration
Countermeasure	for	Administration		
Potassium iodide ¹ See REMM's KI summary information. See FDA's KI information.	Iodine (I-131)	PO	Adults >40 years: 130 mg/day (for projected thyroid exposure ≥500 cGy) Adults 18-40 years: 130 mg/day (for projected thyroid exposure ≥ 10 cGy) Pregnant or lactating women of any age: 130 mg/day (for projected thyroid exposure ≥ 5 cGy)	 Some incidents will require only a single dose of KI. Incident managers may recommend additional doses if ongoing radioactive iodine ingestion or inhalation represents a continuing threat. See <u>REMM</u> <u>page about</u> <u>duration</u>.
Prussian blue, insoluble ¹ See REMM page on Prussian Blue See FDA Prussian Blue information page. See FDA's Prussian Blue drug label.	(Cs-137) Thallium (TI-201)	PO	Adults: 3 g PO tid (See FDA package insert) OR 1 - 3 g PO tid with 100-200 mL water, up to 10-12 g/day (based on Goiânia accident data)	 Minimum 30 days course per FDA Obtain bioassay and whole body counting to assess treatment of efficacy Duration of therapy depends on total body burden and response to treatment

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27. Neutropenia therapy ± antimicrobials

Neutropenia definition:

Total count of neutrophils + bands in the peripheral blood <1,000 /microL

- The 2 drugs listed below have been approved by the FDA for the indication of acute exposure to myelosuppressive doses of radiation
- See <u>**REMM cytokines page</u>** for much more detailed information, especially potential need for <u>dose alterations during large mass casualty incidents when medical</u> <u>countermeasures may be scarce</u>.</u>

Myeloid cytokines approved by the FDA for the indication of acute exposure to myelosuppressive doses of radiation

Cytokine	Adult dose
G-CSF or filgrastim (<u>Neupogen</u> ® drug label)	 10 mcg/kg/day as a single daily subcutaneous injection in adults and children Continue administration daily until absolute neutrophil count remains greater than 1,000/mm³ (= 1.0 x 10⁹ cells/L) for 3 consecutive (daily) CBCs or exceeds 10,000/mm³ (= 10 x 10⁹ cells/L) after a radiation-induced nadir. See <u>REMM cytokines page</u> for more information about potential dose alterations during large mass casualty incidents when medical countermeasures may be scarce.
Pegylated G-CSF or pegfilgrastim (<u>Neulasta</u> ® drug label)	 Two doses, 6 mg each, administered subcutaneously one week apart. A CBC should be obtained prior to administration of the second dose of Neulasta®. Subject matter experts recommend not administering the second dose if absolute neutrophil count is greater than 5,000/mm³ (= 5.0 x 10⁹ cells/L). See <u>REMM cytokines page</u> for more information about potential dose alterations during large mass casualty incidents when medical countermeasures may be scarce.
GM-CSF or sargramostim (<u>Leukine</u> ® drug label)	 This drug is in clinical use for various indications but is NOT approved by the FDA for the specific indication of acute exposure to myelosuppressive doses of radiation. Although Leukine® has not been approved for this indication, CDC has filed a pre-EUA with the FDA to support the issuance of an EUA under a declared emergency. Leukine® has been added to the SNS as noted on the REMM web site. See drug label for prescribing information. See <u>REMM cytokines page</u> for more information about potential dose alterations during large mass casualty incidents when medical countermeasures may be scarce.

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See Clinical Practice Guidelines for Myeloid Cytokines (Adults)

- Smith TJ, Bohlke K, Lyman GH, Carson KR, Crawford J, Cross SJ, Goldberg JM, Khatcheressian JL, Leighl NB, Perkins CL, Somlo G, Wade JL, Wozniak AJ, Armitage JO. <u>Recommendations for the Use of WBC Growth Factors: American</u> <u>Society of Clinical Oncology Clinical Practice Guideline Update</u>. (2015 ASCO guideline) J Clin Oncol. 2015 Oct 1;33(28):3199-212. [PubMed Citation] (This 2015 ASCO guideline updates the <u>2006 myeloid cytokine guideline</u>)
- <u>NCCN Clinical Practice Guidelines in Oncology, Myeloid Growth Factors, Version</u> <u>2.2016</u>. See section entitled "NCCN Guidelines for Supportive Care" > "Myeloid Growth Factors". (Registration required.)
- Dainiak N, Gent RN, et al. <u>First Global Consensus for Evidence-Based</u> <u>Management of the Hematopoietic Syndrome Resulting From Exposure to</u> <u>Ionizing Radiation</u>. Disaster Med Public Health Prep. 2011 Oct;5(3):202-212. [PubMed Citation] (<u>Full text</u>)

For Antimicrobial prophylaxis (no fever) with neutropenia:

- For patients with neutropenia who have NOT HAD NEUTROPENIC FEVER.
- Use as appropriate for each patient.
- Drugs listed are examples only.

Anti-bacterial prophylaxis:

___ Levofloxacin (Levaquin) 500 mg PO/IV daily

Anti-viral prophylaxis (neutropenia without fever)

- __ Acyclovir (Zovirax) 400 mg PO q12h, or
- __ Acyclovir (Zovirax) 250 mg/m² IV q12h

Anti-fungal prophylaxis (neutropenia without fever)

Fluconazole (<u>Diflucan</u>) 400 mg PO/IV daily – beginning when absolute neutrophil Count (ANC) becomes < 1000</p>

or

 Posaconazole (Noxafil) extended release tablets – 300 mg – one tablet twice daily day 1, then one tablet daily thereafter.
 Suspension is 200 mg TID– beginning when Absolute Neutrophil Count (ANC) becomes < 1000.

For treatment of neutropenia AND fever (defined as T>38 °C while neutropenic)

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Anti-microbial work-up and therapy

__ Blood cultures __ Uri

___ Urinalysis w/culture

- ___ Sputum culture + sensitivity ___ Chest x-ray
- __ Cefepime (Maxipime) 2gm IV q 8h
- Vancomycin (Vancocin) 1gm IV q 12h Consider if: suspected catheter-related infection, skin or soft tissue infection, pneumonia or hemodynamic instability.

Consider trough level before 4th dose.

Antifungal therapy

Consider one of the following if: fever >72 hours on antibacterial therapy, evidence of fungal infection or hemodynamic instability.

___ Voriconazole (Vfend) 6mg/kg IV q12h for two doses, then 4 mg/kg IV q12h

Maintenance oral dose: Weight <40 kg: 100 mg PO every 12 hours

Weight \geq 40 kg: 200 mg PO every 12 hours

___ Caspofungin (Cancidas) 70 mg IV once then 50 mg IV daily

____ Liposomal amphotericin B (<u>Ambisome</u>) 3 mg/kg/day IV over 1-4h

___ Amphotericin B lipid complex (Abelcet) 3 mg/kg/day IV over 1-4h

See REMM page on peer-reviewed Fever and Neutropenia Guidelines

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NOTES

1. FDA approved for this indication

2. This drug is not approved by the FDA for this indication. If used, this would be an "off label use", and physician discretion is strongly advised.

3. Ca-DTPA and Zn-DTPA have not been approved by FDA for treating internal contamination with californium, thorium, and yttrium. For initial treatment, Ca-DTPA is recommended, if available, within the first 24 hours after internal contamination. Zn-DTPA is preferred for maintenance after the first 24 hours, if available, due to safety concerns associated with prolonged use of Ca-DTPA.

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Body Chart for Recording Results of Radiation Survey

