

CRITICAL CARE ORDER SET

Neutropenic Fever

An oncologic emergency. Draw blood cultures from every lumen and a peripheral site, then give the first broad-spectrum antibiotic within 60 minutes — never wait on the workup. Layer MRSA and double Gram-negative coverage onto an antipseudomonal backbone by risk, support the failing circulation, and engage ID and Heme/Onc early. Adapt every dose to your institutional antibiogram, prior colonization, and the patient in front of you.

PATIENT -----	MRN -----	ANC / CRCL -----	MALIGNANCY / REGIMEN -----
ALLERGIES -----	FIRST ANTIBIOTIC TIME -----	ORDERING CLINICIAN -----	CODE STATUS -----

DEFINITION ANC < 500/ μ L (or expected < 500 within 48 h) **AND** a single temperature ≥ 38.3 °C or ≥ 38.0 °C sustained ≥ 1 h.

01 Triage, isolation & initial orders

DIAGNOSIS / PROBLEM LIST

- ☐ Neutropenic fever (per definition above)
- ☐ Underlying malignancy / chemotherapy-induced neutropenia

NURSING / MONITORING

- ☐ Vital signs q15 min $\times$ 1 h, then q1h $\times$ 4 h, then q4h if stable (ICU-standard if unstable)

- ☐ Continuous cardiac & pulse oximetry monitoring if hypotensive, tachycardic, or septic
- ☐ Strict intake & output **q1h** (q4h if on the floor and stable); daily weight

Neutropenic precautions

- ☐ Private room; mask for patient during transport
- ☐ Neutropenic / low-bacteria diet per institutional policy
- ☐ Contact precautions if diarrhea or suspicion for *C. difficile*

Isolation

- ☐ Droplet + contact if respiratory symptoms or known viral pathogen
- ☐ Airborne if suspected TB, measles, varicella, etc., per clinical context

02 Immediate sepsis workup — STAT, within 1st hour

DO NOT DELAY Draw cultures **but give the first antibiotic dose within 60 min** — the workup must not postpone therapy.

Cultures

- ☐ **Blood cultures:** 2 sets (aerobic + anaerobic) from 2 separate peripheral sites **PLUS 1 set from each lumen** of any central venous catheter
- ☐ Urine culture (clean catch / straight cath) if urinary device, symptoms, or no clear source
- ☐ Sputum / tracheal aspirate / BAL if respiratory symptoms or infiltrate
- ☐ Wound / line-site cultures if erythema, tenderness, or drainage
- ☐ Stool: *C. difficile* toxin / PCR if diarrhea; bacterial stool panel if indicated

Laboratory studies

- | | |
|---|--|
| ☐ CBC with differential | ☐ CMP (incl. LFTs, creatinine) |
| ☐ Lactate (initial + repeat 2–4 h if elevated) | ☐ Coags: PT/INR, aPTT, fibrinogen, D-dimer if DIC concern |
| ☐ CRP and/or procalcitonin (per protocol) | ☐ Peripheral smear if blasts / TLS / MAHA concern |

☐ Type & screen (crossmatch if indicated)

☐ If TLS risk: uric acid, LDH, phosphorus, calcium

Imaging

☐ Chest x-ray for all with respiratory symptoms or no obvious source

CT CHEST / ABDOMEN / PELVIS WITH CONTRAST IF

☐ Persistent fever ≥ 72 h on appropriate therapy, or concern for deep-seated infection (typhlitis, abscess, fungal)

☐ Additional imaging as indicated (sinus CT, MRI brain, etc.)

03 Source control & supportive procedures

☐ Assess central line sites; remove CVC if high suspicion of CLABSI and patient stable enough to tolerate (per ID / Onc input)

IF MUCOSITIS

☐ Oral exam q shift; saline or sodium-bicarbonate swish-and-spit rinses

IF DIARRHEA & ABDOMINAL PAIN

☐ Surgical consultation if concern for neutropenic enterocolitis (typhlitis) or perforation

04 Empiric antimicrobial therapy

≤ 60 MIN Start broad-spectrum IV antibiotics **within 60 minutes** of recognition. Tailor to local antibiogram and prior colonization / infection history.

4.1 First-line · no immediate MRSA or resistant GN risk

PICK ONE ANTIPSEUDOMONAL B-LACTAM

☐ Piperacillin–tazobactam 4.5 g IV q6h

OR

☐ Cefepime 2 g IV q8h

OR

☐ Meropenem 1 g IV q8h (or imipenem–cilastatin 500 mg IV q6h)

ADJUST Renal dose adjustment per CrCl; use extended-infusion strategies per institutional policy.

4.2 Add MRSA coverage · if ANY trigger

TRIGGERS

- ☐ Hemodynamic instability / severe sepsis / shock
- ☐ Suspected catheter-related infection
- ☐ Known MRSA colonization / prior MRSA
- ☐ Pneumonia (esp. cavitary / MRSA risk)
- ☐ Skin / soft-tissue infection

ADD ONE

- ☐ **Vancomycin IV** — load 20–25 mg/kg, then 15–20 mg/kg q8–12h; trough / AUC- & renal-guided

IF VANCOMYCIN UNSUITABLE

- ☐ Daptomycin IV (**not for pneumonia**)
- ☐ Linezolid IV if lung source with high MRSA suspicion and vancomycin contraindicated

4.3 Double Gram-negative coverage · consider for MDR risk

ADD A SECOND AGENT IF

- ☐ Septic shock / high-risk instability; known ESBL / KPC / MDR GN; high-resistance environment or recent resistant infection

EXAMPLES

- ☐ Add amikacin 15–20 mg/kg IV q24h (or gentamicin)
- ☐ Add ciprofloxacin 400 mg IV q8–12h (if no prior FQ prophylaxis & local susceptibility acceptable)
- ☐ Use carbapenem (meropenem) instead of pip-tazo / cefepime for ESBL risk

05 Antifungal & antiviral therapy

5.1 Empiric antifungal · not routine at presentation

CONSIDER IF

- ☐ Persistent fever ≥ 4–7 days despite appropriate broad-spectrum antibiotics

- ☐ High-risk prolonged neutropenia (ANC < 500 for > 7 days, AML induction, allo-HCT)
- ☐ New pulmonary infiltrates / sinus disease suspicious for mold
- ☐ Positive galactomannan / β -D-glucan or other invasive fungal evidence

CHOICES — PER ID / ONC & LOCAL PRACTICE

- ☐ Voriconazole **6 mg/kg IV q12h × 2, then 4 mg/kg IV q12h**
- ☐ Liposomal amphotericin B **3–5 mg/kg IV daily**
- ☐ Echinocandin — caspofungin **70 mg IV load, then 50 mg IV daily**

5.2 Antiviral

- ☐ Influenza season / compatible symptoms: oseltamivir **75 mg PO q12h** (adjust for CrCl)
- ☐ HSV / VZV mucocutaneous disease: acyclovir **5 mg/kg IV q8h** (renal-adjusted)

06 Hemodynamic & organ support

FLUIDS — IF HYPOTENSIVE OR LACTATE $\geq$ 2 MMOL/L

- ☐ Crystalloid bolus — balanced crystalloid (or NS) **30 mL/kg IV within first 3 h**
- ☐ Maintenance — balanced crystalloid **75–100 mL/h**; adjust for HF / CKD

VASOPRESSORS — MAP < 65 DESPITE FLUIDS / FLUID-LIMITED

- ☐ **Norepinephrine** via central line — start **0.02–0.05 μ g/kg/min**, titrate to MAP $\geq$ 65 mmHg
- ☐ Add vasopressin or epinephrine per sepsis protocol if needed
- ☐ Arterial line for continuous BP monitoring in ICU-level care

RESPIRATORY SUPPORT

- ☐ O₂ to keep SpO₂ $\geq$ **92%** (or per chronic baseline); low threshold for HFNC / intubation in septic shock

07 Hematologic & oncologic management

G-CSF (filgrastim)

- ☐ Filgrastim **5 μ g/kg/day SC or IV**

INDICATIONS — PER ONC / ID

- ☐ High-risk neutropenia with sepsis / shock; expected prolonged neutropenia (> 7–10 days); elderly / comorbid / pneumonia / hypoxia

Transfusions

RBC	Hgb < 7 g/dL (higher if symptomatic ischemia / CAD)
Platelets — routine	< 10,000/μL
Platelets — fever / mucositis	< 20,000/μL
Platelets — procedure / bleeding	< 50,000/μL

- ☐ Irradiated, leukoreduced products for oncology / transplant patients as indicated

08 Prophylaxis & adjunctive orders

GI prophylaxis

- ☐ PPI or H2 blocker per protocol if additional indications present

VTE prophylaxis

- ☐ Enoxaparin **40 mg SC daily** (30 mg if CrCl < 30)
- ☐ Hold if platelets < 50,000/μL, active bleeding, high risk → use SCDs

Electrolytes & nutrition

- ☐ Daily BMP, Mg, Phos (more often if unstable); correct K / Mg / Phos aggressively
- ☐ Early nutrition consult; enteral feeding if prolonged NPO

Oral care

- ☐ Saline / sodium-bicarb rinses **q4h** while awake; avoid alcohol-based mouthwash; topical anesthetics for severe mucositis

09 De-escalation & duration

REASSESS AT 48–72 H

- ☐ Narrow based on cultures & susceptibilities
- ☐ Stop vancomycin if no MRSA source and cultures negative

If no source, cultures negative, patient stable

Continue broad-spectrum agent until ALL of:

- ☐ Afebrile $\geq$ 48 h, clinically stable, and neutrophil recovery (ANC $>$ 500/ μ L and rising)
- ☐ In high-risk / prolonged neutropenia, may continue until ANC recovery — individualize with Heme / ID

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Consults

- ☐ **Infectious Diseases** — all high-risk / complicated cases
- ☐ **Hematology / Oncology** — neutropenia, chemo timing, G-CSF
- ☐ **Critical Care** — evolving organ dysfunction or shock
- ☐ **Surgery** — typhlitis, perforation, abscess, necrotizing infection

EDUCATIONAL USE ONLY

Not a substitute for clinical judgement. Verify doses, allergies, renal function, and contraindications. Adapt all agents, targets, and thresholds to your institutional protocols, formulary, and the individual patient.

Spreading knowledge
Improving outcomes