

PROPHYLAXIS GUIDELINES FOR THE ADULT HEMATOLOGY PATIENT

_	Indication	Antibacterial	Antifungal	PJP prophylaxis	Antiviral	Duration of Prophylaxis
MDS	Receiving chemotherapy	No routine prophylaxis	Fluconazole 200 mg PO daily	No routine prophylaxis	Acyclovir 400 mg PO BID	Antifungal: Beginning when ANC ≤500 and continuing throughout neutropenia Antiviral: Throughout all chemotherapy cycles
	APL Induction	No routine prophylaxis		No routine prophylaxis	Acyclovir 400 mg PO BID	Antibacterial/Antifungal: If indicated begin when ANC ≤500 and continuing throughout neutropenia. In consolidation send Rx for patient to start at discharge continue throughout neutropenia
		If differentiating on steroids: Levofloxacin 500 mg PO daily	Micafungin 100 mg IV q24h			
AML	AML Intensive Induction	No routine prophylaxis	Voriconazole 200 mg PO BID (trough level after 5-7 days)	For patients receiving purine analogue ^{1:} TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	
	HMA + Venetoclax	Levofloxacin 500 mg PO daily	Posaconazole 300 mg tab PO daily			
	Relapsed/Refractory or >70 years Induction	Levofloxacin 500 mg PO daily	For relapsed/refractory patients unlikely to recover ANC: Posaconazole 300 mg tab PO daily			
	Consolidation	Levofloxacin 500 mg PO daily	Fluconazole 200 mg PO daily			<u>PCP</u> : Throughout all chemotherapy cycles. Continued for 6 mo following last dose of purine analogue.
	ALL Induction	Levofloxacin 500 mg PO daily	Micafungin 50 mg IV q24h	TMP-SMX (Bactrim) DS 3 times weekly (hold through methotrexate admission until level <0.1 μM)	Acyclovir 400 mg PO BID	Antiviral: Throughout all chemotherapy cycles
	Beyond Induction	No routine prophylaxis	For patients receiving HyperCVAD: Fluconazole 200 mg PO daily			
ALL		For patients receiving HyperCVAD: Levofloxacin 500 mg PO daily				
1	Blinatumomab	No routine prophylaxis	No routine prophylaxis			
		If prolonged neutropenia: Levofloxacin 500 mg PO daily	If prolonged neutropenia: Posaconazole 300 mg tab PO daily			
	Inotuzumab	Levofloxacin 500 mg PO daily	Fluconazole 200 mg PO daily			
	Hairy Cell Leukemia	Levofloxacin 500 mg PO daily	If no G-CSF support being used Fluconazole 200 mg PO daily	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Antibacterial/Antifungal: Beginning when ANC ≤500 and continuing throughout neutropenia PCP: Throughout all chemotherapy cycles. Continued for 6 mo following last dose of purine analogue. Antiviral: Throughout all chemotherapy cycles
	Aplastic Anemia	If neutropenic on discharge: Levofloxacin 500 mg PO daily	If neutropenic on discharge: Voriconazole 200 mg PO BID ² (trough level after 5-7 days)	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID EBV/CMV monitoring	Antibacterial/Antifungal: Beginning when ANC ≤500 and continuing throughout neutropenia PCP: Beginning with therapy and continuing for at least 6 mo Antiviral: Throughout all therapy



	Indication	Antibacterial	Antifungal	PJP prophylaxis	Antiviral	Duration of Prophylaxis
	High dose steroids ³	No routine prophylaxis	No routine prophylaxis	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles
la Pa	Proteasome inhibitors (e.g., bortezomib, carfilzomib, ixazomib)	No routine prophylaxis	No routine prophylaxis	No routine prophylaxis	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles and continuing for at least 3 months post last dose
Myeloma	Monoclonal antibodies (e.g., Elotuzumab, isatuximab, daratumumab)	No routine prophylaxis	No routine prophylaxis	No routine prophylaxis	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles and continuing for at least 3 months post last dose
2	VDT-PACE or DCEP	Levofloxacin 500mg daily	Fluconazole 200 mg PO daily	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Antifungal/Antibacterial: Send Rx for patient to start at discharge and continue throughout neutropenia PCP/Antiviral: Throughout all chemotherapy cycles
	BEACOPP	No routine prophylaxis	No routine prophylaxis	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles
	DA-R-EPOCH HIV Negative	No routine prophylaxis	No routine prophylaxis	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles
	DA-R-EPOCH HIV Positive ⁴	Levofloxacin 500 mg PO daily	Fluconazole 200 mg PO daily	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	Antifungal/Antibacterial: Send Rx for patient to start at discharge and continue throughout neutropenia. For outpatient EPOCH, start on day 6. PCP/Antiviral: Throughout all chemotherapy cycles
Lymphoma	HyperCVAD CODOX-M/IVAC	Levofloxacin 500 mg PO daily	Fluconazole 200 mg PO daily	TMP-SMX (Bactrim) DS 3 times weekly (hold through methotrexate admission until level <0.1 μM)	Acyclovir 400 mg PO BID	Antibacterial: Beginning with Part B of regimen and continued throughout all chemotherapy cycles Antifungal: Send Rx for patient to start at discharge and continue throughout neutropenia PCP/Antiviral: Throughout all chemotherapy cycles
	R-ICE, R-ESHAP, R-DHAP Nordic	No routine prophylaxis	No routine prophylaxis	No routine prophylaxis	Acyclovir 400 mg PO BID	Throughout all chemotherapy cycles
	PI3K inhibitor (e.g., idelalisib, copanlisib, duvelisib)	No routine prophylaxis	No routine prophylaxis	TMP-SMX (Bactrim) DS 3 times weekly	No routine prophylaxis CMV monitoring	PCP: Through duration of treatment
	Purine analogues (cladribine, fludarabine, nelarabine, pentostatin, bendamustine)	No routine prophylaxis	No routine prophylaxis	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID	PCP: Beginning with chemotherapy and continued at least 6 months after treatment and until normalization of ALC (≥1.2 k/uL) Antiviral: Throughout all chemotherapy cycles
	Alemtuzumab	No routine prophylaxis	Voriconazole 200 mg PO BID	TMP-SMX (Bactrim) DS 3 times weekly	Acyclovir 400 mg PO BID EBV/CMV monitoring	Antifungal: Beginning when ANC ≤500 and continuing throughout neutropenia PCP/Antiviral: Beginning with therapy and continued 6 mo after therapy or until normalization of ALC (≥1.2 k/uL)
	Maintenance Anti-CD20 (e.g., rituximab, obinutuzumab)	No routine prophylaxis	No routine prophylaxis	No routine prophylaxis	Acyclovir 400 mg BID Hepatitis B screen prior to initiation	Throughout all chemotherapy cycles



No routine prophylaxis is recommended for the following: R-CHOP, MR-CHOP, HD-MTX, GemOx, GELOX, immunomodulatory agents, BTK inhibitors, venetoclax, ruxolitinib, HDAC inhibitors, ABVD, Brentuximab, Tagraxafusp, selinexor, tafasitamab, belantamab mafodotin, other CML/CLL treatment (unless receiving therapy with agents in the chart above)

⁴An integrase-inhibitor based HAART regimen is recommended.

Managing Voriconazole Adverse Effects / Drug Interactions				
Visual hallucinations	Associated with high levels and often dose dependent. Obtain a level, hold dose until symptoms resolve, empiric 25% dose reduction (Tan K, Clin Pharmacol. 2006)			
Hepatotoxicity (Direct Bili > 3 or otherwise deemed clinically significant)	Hold azole and switch to micafungin 100 mg IV q24h			
Drug interactions/additive toxicity ¹	If preferred antifungal is voriconazole: Substitute with micafungin 100 mg IV q24h If preferred antifungal is fluconazole: Substitute with micafungin 50 mg IV q24h			
NPO ²	If preferred antifungal is voriconazole/posaconazole: Switch to micafungin 100 mg IV q24h If preferred antifungal is fluconazole: Switch to IV fluconazole 200 mg IV q24h			

¹Vincristine, tyrosine-kinase inhibitors, clofarabine, doxorubicin, or if mandated by clinical trial protocol (e.g., quizartinib). Micafungin should be utilized only during the period of concomitant administration of interacting chemotherapy; once interaction is no longer relevant, prophylaxis should revert to preferred agent

²Patients with feeding tubes in whom the preferred antifungal is voriconazole should continue to receive voriconazole tablets; tablets may be crushed and administered via the tube. In all scenarios, once NPO status is over, prophylaxis should revert to preferred agent.

¹Purine analogues include fludarabine, clofarabine, and cladribine

²Cyclosporine should be decreased by 50% when started concurrently with voriconazole

³High dose steroids defined as ≥20 mg prednisone equivalents continuously for ≥4 weeks



EBV/CMV Monitoring Recommendations for Patients Receiving Alemtuzumab/Anti-Thymocyte Globulin

Surveillance negative:

Continue surveillance plan

Monitor EBV and CMV PCR in the blood:

- prior to treatment
- weekly during the 1st month
- every other week during the second month
- monthly for 6 months
- every 12 months

CMV Surveillance Positive:

(>3,000 I.U./mL for two consecutive readings 7 days apart or or a single level >3,000 I.U./mL with symptoms suggestive of CMV disease or a single level >6,000 I.U./mL, measured at UMHS Labs)

Valganciclovir 900 mg PO BID*

*Consider ID consult if considering therapy, especially if therapy with valganciclovir felt to be untenable

EBV Surveillance Positive:

Antiviral therapy should NOT be initiated. Contact Hematology/Oncology service for recommendations.

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The recommendations in this guide are meant to serve as treatment guidelines for use at Michigan Medicine facilities. If you are an individual experiencing a medical emergency, call 911 immediately. These guidelines should not replace a provider's professional medical advice based on clinical judgment, or be used in lieu of an Infectious Diseases consultation when necessary. As a result of ongoing research, practice guidelines may from time to time change. The authors of these guidelines have made all attempts to ensure the accuracy based on current information, however, due to ongoing research, users of these guidelines are strongly encouraged to confirm the information contained within them through an independent source.

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