

# Antimicrobial Prophylaxis for Allogeneic and Autologous Hematopoietic Stem Cell Transplant Recipients: To be initiated at the time of admission for transplantation

*Developed by: Immunocompromised ID consultative, Antimicrobial Stewardship Program, Pharmacy and Bone Marrow Transplant services*

## Allogeneic stem cell transplantation

<b>Antibacterial</b>	<p><b>Pre-engraftment: Levofloxacin PO 500mg daily*<sup>1-4</sup></b></p> <ul style="list-style-type: none"> <li>Begin at Day 0 (or sooner if ANC &lt;500) and continue until ANC &gt;500 on 2 consecutive samples or patient develops neutropenic fever</li> <li>Adjust dosage for CrCl &lt;50</li> </ul> <p><b>Chronic GVHD:</b></p> <ul style="list-style-type: none"> <li><b>Penicillin VK 250mg twice daily<sup>3</sup></b> for pneumococcal prophylaxis (not necessary if patient is still receiving levofloxacin)</li> <li>Ensure patient up to date with pneumococcal vaccine series beginning 6 months post-transplantation<sup>3</sup></li> </ul>
<b>Antifungal</b>	<p><b>Low risk patients: Fluconazole 400mg daily<sup>3-5</sup></b></p> <ul style="list-style-type: none"> <li>Begin on admission and continue until day 75 or no longer receiving immunosuppression</li> <li>Adjust dosage for CrCl &lt;50</li> </ul> <p><b>High risk patients</b> (acute GVHD (grade II to IV), chronic extensive GVHD, history of IFI, alemtuzumab receipt, or high dose steroids) or history of mold infection: <b>Posaconazole DR 300mg daily<sup>6</sup>:</b></p> <ul style="list-style-type: none"> <li>Begin on admission and continue until no longer receiving immunosuppression</li> </ul> <p>May use <b>micafungin 50mg daily</b> if unable to tolerate PO or have intolerable medication interactions</p>
<b>Antiviral</b>	<p><b>HSV/VZV: Valtrex 500mg PO q12hrs or Acyclovir 400mg q8hrs</b></p> <ul style="list-style-type: none"> <li>Begin on admission and continue for 1 year post-transplant or until immunosuppression is discontinued (whichever is longer)</li> <li>Adjust dosage for CrCl &lt;30</li> <li>If unable to tolerate PO: <b>Acyclovir IV 250mg q12hrs</b></li> <li><b>Valtrex 500mg PO q12hrs</b> or <b>Acyclovir 800mg q12hrs</b> if history of herpes zoster<sup>7</sup></li> </ul> <p><b>CMV:</b> Recipient CMV IgG positive: start <b>Letermovir 480mg daily</b> (Day +5 to day +100)</p> <ul style="list-style-type: none"> <li>Recipient CMV IgG negative: Pre-emptive approach with weekly viral load screening</li> </ul>
<b>Pneumocystis</b>	<p><b>Bactrim DS M/W/F or Bactrim SS daily<sup>3</sup></b></p> <ul style="list-style-type: none"> <li>Begin at time of engraftment and continue for 1 year and immunosuppression is discontinued</li> <li>Alternative if sulfa intolerant (2<sup>nd</sup> line agents) : <b>Atovaquone 1500mg daily, Dapsone 100mg daily</b> (check for G6PD deficiency prior to initiating Dapsone therapy)</li> </ul>
<b>Toxoplasmosis</b> (if toxo IgG+)	<p><b>Bactrim DS M/W/F or DS daily</b></p> <ul style="list-style-type: none"> <li>If unable to tolerate Bactrim, must monitor serum toxoplasma PCR q2-4 weeks until day+90</li> </ul>
<b>Hep B</b> (if recipient is S Ag+ or core Ab+)	<p><b>Tenofovir alafenamide 25mg daily or Entecavir 0.5mg daily</b></p> <ul style="list-style-type: none"> <li>Begin day 0 (or with conditioning chemo) and continue x 12 mos post-transplant</li> <li>Alternative: may monitor Hep B S Ag and VL every 3 mos post-transplant and start antiviral if VL detectable</li> </ul>

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## Autologous stem cell transplantation

<b>Antibacterial</b>	<b>Levofloxacin PO 500mg daily*<sup>1-4</sup></b> <ul style="list-style-type: none"> <li>Begin on Day 0 (or sooner if ANC &lt;500) and continue until ANC &gt;500 on 2 consecutive samples or patient develops neutropenic fever</li> <li>Adjust dosage for CrCl &lt;50</li> </ul>
<b>Antifungal</b>	<b>Fluconazole 200mg daily<sup>3,4</sup></b> <ul style="list-style-type: none"> <li>Begin Day 0 and continue to Day 30 post-transplant</li> <li>Adjust dosage for CrCl &lt;30</li> <li>If prior history of mold infection, consider <b>Posaconazole DR 300mg daily</b></li> </ul>
<b>Antiviral</b>	<b>HSV/VZV: Valtrex 500mg PO q12hrs or Acyclovir 400mg q8hrs<sup>3,4</sup></b> <ul style="list-style-type: none"> <li>Begin on Day 0 and continue until 6-12 months post-transplant</li> <li>Adjust dosage for CrCl &lt;30</li> <li>If unable to tolerate PO: <b>Acyclovir IV 5mg/kg q12hrs</b> (dosed by ideal body weight)</li> <li><b>Valtrex 500mg PO q12hrs or Acyclovir 800mg q12hrs</b> if history of herpes zoster<sup>7</sup></li> </ul>
<b>Pneumocystis</b>	<b>Bactrim DS M/W/F or Bactrim SS daily<sup>3</sup></b> <ul style="list-style-type: none"> <li>Begin at Day 30 (if WBC/platelets recovered) and continue until 6 months post-transplant</li> <li>Adjust dosage for CrCl &lt;30</li> <li>Alternative if sulfa intolerant (2<sup>nd</sup> line agents): <b>Atovaquone 1500mg daily, Dapsone 100mg daily</b> (check for G6PD deficiency prior to initiating Dapsone therapy)</li> </ul>
<b>Hep B</b> (if recipient S Ag + or core Ab+)	<b>Tenofovir alafenamide 25mg daily or Entecavir 0.5mg daily</b> <ul style="list-style-type: none"> <li>Begin day 0 (or with conditioning chemo) and continue x 12 mos post-transplant</li> <li>Alternative: may monitor Hep B S Ag and VL every 3 mos post-transplant and start antiviral if VL detectable</li> </ul>

\* If patient unable to tolerate fluoroquinolones, call Transplant ID or antibiotic stewardship for alternatives, such as 3<sup>rd</sup> generation cephalosporin

1 Bucaneve G et al. Levofloxacin to prevent bacterial infection in patients with cancer and neutropenia. New England Journal of Medicine 2005; 353:977-87

2 Gafter-Gvili A et al. Meta-analysis: antibiotic prophylaxis reduces mortality in neutropenic patients. Annals of Internal Medicine 2005; 142:979-995

3 Tomblyn M et al. Guidelines for preventing infectious complications among hematopoietic cell transplant recipients: A global perspective. Biology of Blood and Bone Marrow Transplantation 2009;15: 1143-1238

4 Freifeld AG et al. Clinical practice guideline for the use of antimicrobial agents in neutropenic patients with cancer: 2010 update by the Infectious Diseases Society of America. Clinical Infectious Diseases 2011;52:e56-e93

5 Marr KA et al. Prolonged fluconazole prophylaxis is associated with persistent protection against candidiasis-related death in allogeneic marrow transplant recipients: longterm follow up of a randomized placebo-controlled trial. Blood 2000;96:2055-61.

6 Ullmann AJ et al. Posaconazole or fluconazole for prophylaxis in severe graft-versus-host disease. New England Journal of Medicine 2007;356:335-347.

7 Erard V et al. One-year acyclovir prophylaxis for preventing varicella-zoster virus disease after hematopoietic cell transplantation: no evidence of rebound varicella-zoster virus disease after drug discontinuation. Blood 2007;110:3071-3077