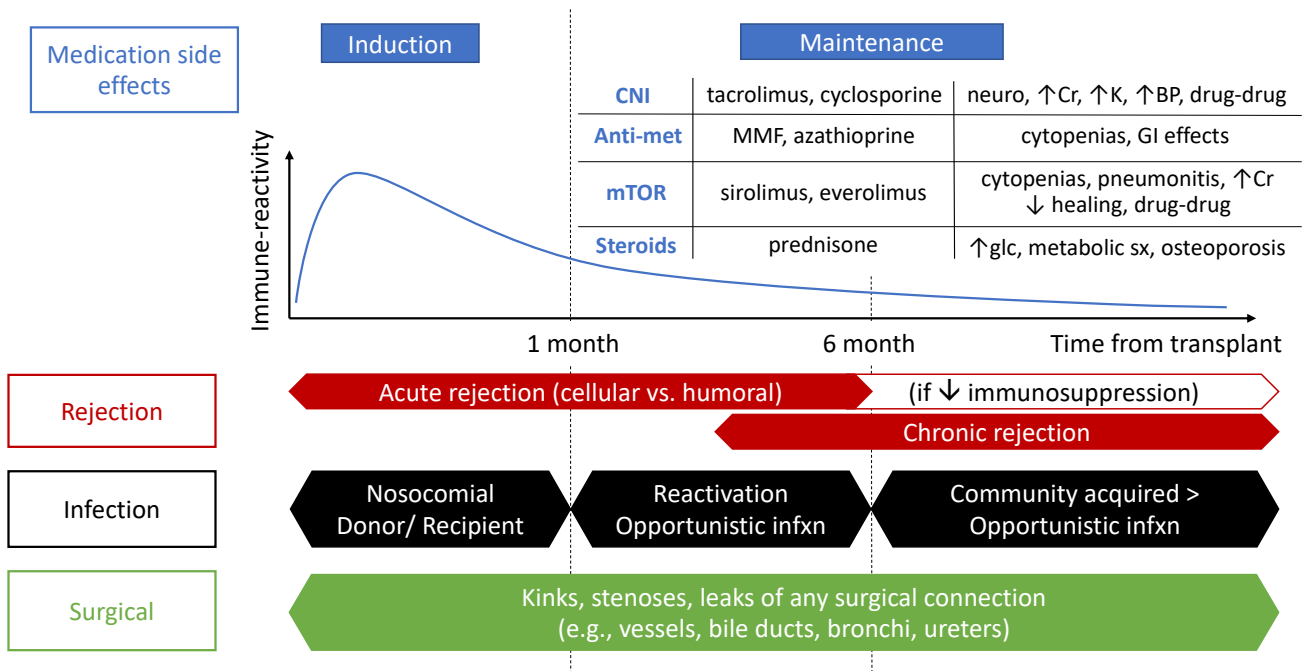
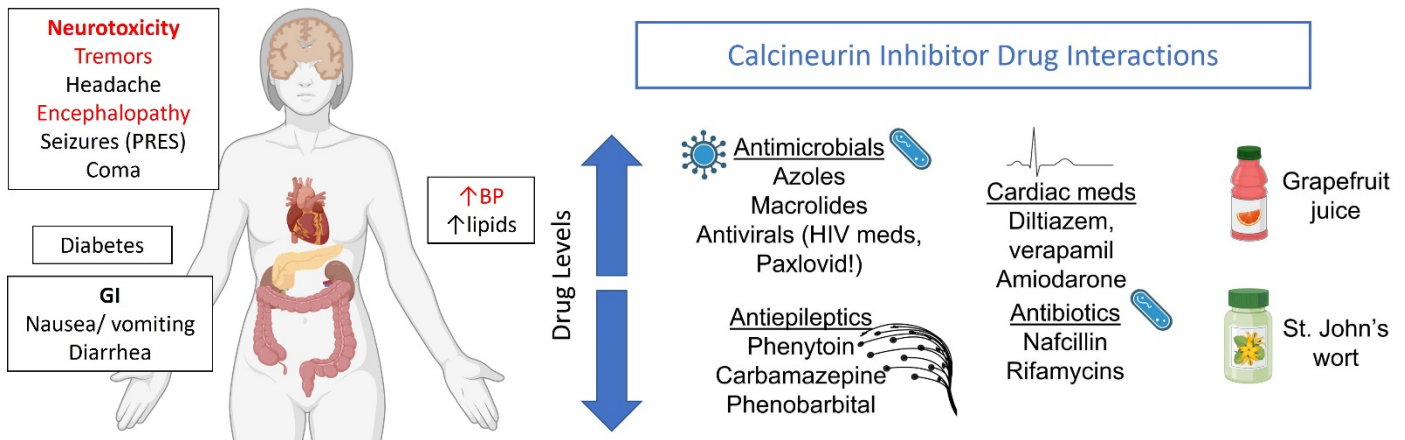


## Complications after transplant



## Calcineurin Inhibitor Toxicity and Drug Interactions



## Take Home Points

1. Have a low threshold to call the Transplant Center early!
2. Evaluate graft dysfunction like native organ dysfunction. Additionally, consider 4 main complications: immunosuppressant side effects, rejection, infection, and surgical complications.
3. Time since transplant (a surrogate for level of immunosuppression) impacts the risk of certain infections.
4. Maintain a high level of suspicion for infection. Immunosuppression can blunt classic symptoms of infection.
5. Do not routinely hold immunosuppression. Dose CNIs consistently and time troughs (~12 hours after last dose).
6. Many drugs affect CNI levels. CNI toxicity may present with neurotoxicity and nephrotoxicity.

## Clinical Pearls

### Immunosuppressant Medications

- **Calcineurin inhibitor (CNI) levels can ↑ in acute illness and can cause neurotoxicity and nephrotoxicity.**
  - Calcium channel blockers and azoles (among others) ↑ CNI levels. Antiepileptics ↓ CNI levels.
  - **Avoid nirmatrelvir-ritonavir (Paxlovid) in patients on calcineurin inhibitors!**
  - mTOR inhibitors also have many drug-drug interactions, and they impair wound healing.
  - Myelosuppression is greatest with valganciclovir > MMF >> TMP/SMX > CNI.
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### Infectious Disease

- Consider donor-derived infections in the early post-transplant period.
  - **Note Donor/Recipient CMV status!** CMV can reactivate and cause multi-organ dysfunction.
  - **Treatment for recent rejection (any ↑ immunosuppression) shifts risk profile for infections and malignancies.**
  - ↑ immunosuppression increases risk of infection-related malignancies (Post-Transplant Lymphoproliferative Disease [PTLD], squamous cell cancer, etc.)
  - Skin lesions can also be due to opportunistic infections – Nocardia, Cryptococcus, other fungi, etc.
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### Liver Transplant

- **Obtain a liver duplex (vascular/Doppler) ultrasound in the evaluation of new liver dysfunction.**
  - Increased vascular resistance on duplex may be an early sign of rejection.
  - Acute rejection is most common in the first 6 months but can occur later with underimmunosuppression.
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### Renal Transplant

- Treat symptomatic UTI as complicated UTI. Remember to assess for (lower quadrant) graft tenderness.
  - **Routinely treating asymptomatic bacteriuria is no longer recommended.**
  - Obtain a duplex of the transplanted kidney in the evaluation of acute kidney injury.
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### Lung Transplant

- **Have a low threshold to check an extended respiratory viral panel.**
  - Lung transplants are the most likely to reject of all solid organ transplants.
  - Dry lungs are happy lungs! Lung transplants are prone to pulmonary edema due to lack of lymphatics. **Manage volume judiciously! Have a low threshold to diurese.**
  - Signs and symptoms of acute rejection are nonspecific and can mimic infection.
  - Consider prior colonization of lungs and sinuses to guide antibiotic choice.
- 

### Heart Transplant

- Transplanted hearts are denervated. Patients have chronic tachycardia.
- Ischemia does not always produce chest pain (due to denervation).
- Heart transplant patients are typically managed by Transplant Cardiologists (even at UW). **Involve the Transplant Team early!**