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When renally cleared drugs are administered to patients with reduced renal function, drug accumulation leading to supratherapeutic concentrations and drug toxicities is a primary concern. However, clearance is only one of the pharmacokinetic parameters that affect a drug's disposition. The volume of distribution of a drug also can be altered in patients with reduced renal function. Furthermore, some patients with HIV or diabetes mellitus can have reduced oral absorption of certain drugs. Therefore, although a drug may require a dose reduction in renal failure based on reduced clearance (i.e., increased concentrations), other factors—such as an increased volume of distribution or reduced oral absorption—may decrease concentrations.

Therapeutic drug monitoring (TDM), if available and appropriate, may facilitate dose adjustments in these complicated patients. TDM allows the clinician to make informed, individualized decisions about dose adjustments that are more precise than standardized dose adjustments based on estimated creatinine clearance. Drugs that are marked with an asterisk (*) in the table below are known to have assays (for clinical and/or research purposes) available within the United States and typically in Europe as well. When TDM is appropriate, clinicians should contact their clinical laboratory to determine assay availability and turnaround time for their institution.

	Usual Dose	Dosage Adjustment in Renal Insufficiency		
Drug(s)		CrCI [^] or eGFR [#] (mL/min)	Dose	
Acyclovir [*]	IV Dose	26–50	100% of dose IV every 12 hours	
	Serious HSV	10–25	100% of dose IV every 24 hours	
	 5 mg/kg IV every 8 hours VZV Infections or HSV encephalitis 10 mg/kg IV every 8 hours 	<10	50% of dose IV every 24 hours	
		HD	50% of dose every 24 hours; administer dose after HD on days of dialysis.	
	PO Dose for Herpes	10–25	800 mg PO every 8 hours	
	Zoster: 800 mg PO five times per day	<10	800 mg PO every 12 hours	
	HD	800 mg PO every 12 hours; administer dose after HD on days of dialysis		
Adefovir	10 mg PO every 24 hours	30–49	10 mg PO every 48 hours	
		10–29	10 mg PO every 72 hours	

		Dosage Adjustment in Renal Insufficiency		
Drug(s)	Drug(s) Usual Dose		Dose	
		HD	10 mg PO weekly; administer dose after HD	
Amikacin* For mycobacterial infections	IV 15 mg/kg per day or 25 mg/kg three times per week	Use with caution in patients with renal insufficiency and family history of ototoxicity.	15 mg/kg two to three times per week Perform TDM to adjust dose, with target peak concentration 35–45 mcg/mL and trough concentration <4 mcg/mL. Administer dose after HD on days of dialysis.	
Amphotericin B [*]	3–6 mg/kg IV per day (lipid formulation) or 0.7–1.0 mg/kg IV per day (amphotericin B deoxycholate)	N/A	No dosage adjustment necessary; consider alternative antifungals if renal insufficiency occurs during therapy despite adequate hydration.	
Cidofovir	5 mg/kg IV on Day 0, repeat 5 mg/kg IV dose on Day 7, then 5 mg/kg IV every 2 weeks Give each dose with probenecid and saline hydration (see <u>Table 2</u> for dosing instructions).	Pretreatment SCr >1.5 mg/dL or CrCl ≤55 mL/min or Proteinuria ≥100 mg/dL (≥2 +)	Cidofovir is not recommended unless benefits outweigh risks. See <u>"Pharmacokinetics of cidofovir</u> in renal insufficiency and in continuous ambulatory peritoneal dialysis or high-flux hemodialysis" for recommendations on renal dose adjustments.	
		If SCr increases by 0.3–0.4 mg/dL above baseline	Decrease to 3 mg/kg IV per dose.	
		If SCr increases >0.5 mg/dL above baseline or Proteinuria ≥3 +	Discontinue therapy.	
Ciprofloxacin	500–750 mg PO every 12 hours or	30–50	500–750 mg PO every 12 hours	
	400 mg IV every 8–12 hours	<30	400 mg IV every 12 hours 250–500 mg PO every 24 hours	

		Dosage	e Adjustment in Renal I	nsufficiency
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Do	se
			or 400 mg IV every 24 hours	
		HD or PD	250–500 mg PO every 24	hours
			200–400 mg IV every 24 h or PD on days of dialysis.	ours; administer after HD
Clarithromycin*	500 mg PO every 12 hours	30–60	Usual dose unless used wi COBI, then reduce dose by	
		<30	250 mg PO twice daily	
			or	
			500 mg PO once daily	
			If used with an HIV PI or C (or consider using azithron	
Cycloserine*	10–15 mg/kg/day PO in two divided doses (maximum	30–80	Usual dose; consider TDM	and monitor for toxicities.
	1,000 mg/day); start at	<30 (not on HD) or HD	250 mg once daily or 500 r	ng three times per week
	250 mg once daily and increase dose per tolerability.	עח	Perform TDM and adjust d for toxicities.	ose accordingly. Monitor
	Target peak concentration 20–35 mcg/mL		Use with caution in patients on dialysis.	s with ESRD who are not
Emtricitabine ^{*a} (FTC)	One 200-mg capsule PO once daily	CrCl [^] or eGFR [#] (mL/min)	Oral Capsules	Oral Solution
	or	15–29	200 mg every 72 hours	80 mg every 24 hours
	240-mg solution PO once	<15 and not on HD	200 mg every 96 hours	60 mg every 24 hours
	daily	HD (administer dose after HD on days of dialysis)	200 mg every 24 hours	240 mg every 24 hours

		Dosage Adjustment in Renal Insufficiency		
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Do	se
Emtricitabine'/ Tenofovir' Alafenamide (FTC/TAF) (FDC Trade Name: Descovy) Note: Please refer to product labels for	One tablet (FTC 200 mg/TAF 25 mg) PO once daily	<30 and not on HD	Coformulated tablet is not	recommended.
dosing recommendations for other ARV FDC products containing FTC/TAF.		HD	One tablet daily. Administe of dialysis.	r dose after HD on days
Emtricitabine'/ Tenofovir [*] Disoproxil Fumarate (FTC/TDF)	One (FTC 200 mg/TDF 300 mg) tablet PO daily	30-49	One tablet PO every 48 ho worsening renal function or TAF)	
(FDC Trade Name: Truvada) Note: Please refer to product labels for dosing recommendations for other ARV FDC products containing FTC/TDF.		<30 or HD	Do not use coformulated to Use formulation for each co dose according to recommo individual drugs.	omponent drug and adjust
Entecavir	Usual Dose: 0.5 mg PO once daily For Treatment of 3TC-	CrCl [^] or eGFR [#] (mL/min)	Usual Renal Dose Adjustment	3TC-Refractory or Decompensated Liver Disease
	Refractory HBV or for Patients with Decompensated Liver Disease: 1 mg PO once daily	30 to <50	 0.25 mg PO every 24 hours, <i>or</i> 0.5 mg PO every 48 hours 	 0.5 mg PO every 24 hours, <i>or</i> 1 mg PO every 48 hours
		10 to <30	 0.15 mg PO every 24 hours, <i>or</i> 0.5 mg PO every 72 hours 	 0.3 mg PO every 24 hours, <i>or</i> 1 mg PO every 72 hours

	Usual Dose	Dosage Adjustment in Renal Insufficiency		
Drug(s)		CrCl [^] or eGFR [#] (mL/min)	Do	se
		<10 or HD or CAPD (administer after HD on days of dialysis)	 0.05 mg PO every 24 hours, <i>or</i> 0.5 mg PO once every 7 days 	 0.1 mg PO every 24 hours, <i>or</i> 1 mg PO once every 7 days
Ethambutol	For MAI: 15 mg/kg PO daily For MTB: 15–25 mg/kg PO daily (See the Dosing Recommendations table in	<30 or HD	Usual dose PO three times HD, give dose after dialysis	
	the <u>Mycobacterium</u> <u>tuberculosis section</u> for additional MTB dosing recommendations.)	PD	Do not use in patients on P MAI or MTB treatment (e.g Perform TDM to guide optin	., moxifloxacin).
Ethionamide [*]	15–20 mg/kg PO daily (usually 250–500 mg PO once or twice daily)	<30 or HD	250–500 mg PO once daily Consider TDM.	/
Famciclovir*	For Herpes Zoster: 500 mg PO every 8 hours	40–59	500 mg PO every 12 hours	
	For HSV: 500 mg PO every 12 hours	20–39 <20	500 mg PO every 24 hours 250 mg PO every 24 hours	
		HD	250 mg PO only on HD day	ys, administer after HD
Fluconazole	200–1,200 mg PO or IV every 24 hours (dose and route of administration depends on type of OI)	≤50	Administer 100% of the ind dose, then adjust maintena dose every 24 hours.	
	depends on type of Of	HD	Administer 100% of the ind dose, then adjust maintena three times per week after	ince doses to full dose
Flucytosine*	25 mg/kg PO every 6 hours	21–40	25 mg/kg PO every 12 hou	rs
	TDM is recommended for patients to guide optimal	10–20	25 mg/kg PO every 24 hou	rs
	dosing (target peak serum concentration 2 hours after	<10	25 mg/kg PO every 48 hou	rs
	dose: 25-100 mcg/mL). If TDM is not possible, monitor CBC twice weekly.	HD	25–50 mg/kg PO every 48- dose after HD.	-72 hours; administer

		Dosage Adjustment in Renal Insufficiency		
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Dose	
Foscarnet	Induction Therapy for CMV Infection: 180 mg/kg/day IV in two divided doses Maintenance Therapy for CMV Infection or for Treatment of HSV Infections: 90–120 mg/kg IV once daily	Dosage adjustment needed according to calculated CrCI/kg; consult product label for dosing table.	Dosage adjustment needed according to calculated CrCI/kg; consult product label for dosing table.	
Ganciclovir*	Induction Therapy: 5 mg/kg IV every 12 hours	50–69	2.5 mg/kg IV every 12 hours	
		25-49	2.5 mg/kg IV every 24 hours	
		10–24	1.25 mg/kg IV every 24 hours	
		<10 or HD	1.25 mg/kg IV three times per week; administer dose after HD.	
	Maintenance Therapy: 5 mg/kg IV every 24 hours	50–69	2.5 mg/kg IV every 24 hours	
		25–49	1.25 mg/kg IV every 24 hours	
		10–24	0.625 mg/kg IV every 24 hours	
		<10 or HD	0.625 mg/kg IV three times per week; administer dose after HD.	
Lamivudine ^b (3TC)	300 mg PO every 24 hours	15–29	150 mg PO once, then 100 mg PO every 24 hours	
		5–14	150 mg PO once, then 50 mg PO every 24 hours	
		<5 or HD	50 mg PO once, then 25 mg PO every 24 hours; administer dose after HD on days of dialysis.	
Lamivudine/ Tenofovir Disoproxil Fumarate (3TC/TDF) (FDC Trade Names: Cimduo or Temixys) Note: Please refer to product information for dosing recommendations for other ARV FDC	One (3TC 300 mg/TDF 300 mg) tablet PO every 24 hours	<50	Coformulated tablet is not recommended.	

		Dosage	e Adjustment in Renal I	nsufficiency
Drug(s)	Drug(s) Usual Dose		Do	ose
products containing 3TC/TDF.				
Levofloxacin	500 mg (low dose) or 750– 1,000 mg (high dose) IV or	CrCl [^] or eGFR [#] (mL/min)	Low Dose	High Dose
	PO daily	20–49	500 mg once, then 250 mg every 24 hours, IV or PO	750 mg every 48 hours IV or PO
		<20 or CAPD or HD (administer dose after HD on days of dialysis)	500 mg once, then 250 mg every 48 hours, IV or PO Dose can be adjusted based on serum concentrations.	750 mg once, then 500 mg every 48 hours, IV or PO
Paromomycin	500 mg PO every 6 hours	<10	Minimal systemic absorption necessary but monitor for and ototoxicity in patients w	worsening renal function
Peginterferon Alfa- 2a	°	<30	135 mcg SQ once weekly	
24		HD	135 mcg SQ once weekly May reduce to 90 mcg onc adverse effects or laborato	
Penicillin G (Potassium or Sodium)	Neurosyphilis, Ocular Syphilis, or Otosyphilis	10–50	2–3 million units every 4 he as continuous infusion	ours <i>or</i> 12–18 million units
or Sourcery	3–4 million units IV every 4 hours, or	<10	2 million units every 4–6 hours, <i>or</i> 8–12 million units as continuous infusion	
	• 18–24 million units IV daily as continuous infusion	HD or CAPD	2 million units every 4–6 he continuous infusion	ours, <i>or</i> 8 million units as
Pentamidine	4 mg/kg IV every 24 hours May reduce dose to 3 mg/kg IV daily in the event of toxicities	<10	4 mg/kg IV every 48 hours	
Posaconazole [*]	 IV: 300 mg twice daily on Day 1; then 300 mg once daily Delayed-Release Tablet: 300 mg PO once daily 	<50	No dosage adjustment of or renal insufficiency. Higher concentrations observed ir <20 mL/min. Perform posaconazole TD concentration at least >1.2	variability in serum patients with CrCl M (target trough

		Dosage Adjustment in Renal Insufficiency		
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Dose	
	Oral Suspension: 400 mg PO twice daily		IV posaconazole is not recommended by the manufacturer because of potential toxicity due to accumulation of SBCD (vehicle of IV product). However, an observational study did not find worsening in renal function in patients with CrCl <50 ml/min given SBCD. Switch patients with CrCl <50 mL/min to oral	
Pyrazinamide [*]	See the <u>Mycobacterium</u> <u>tuberculosis section</u> for weight-based dosing guidelines.	<30 or HD	posaconazole when feasible. 25–35 mg/kg/dose three times per week; administer dose after HD.	
Quinine Sulfate [*]	650 mg salt (524 mg base) PO every 8 hours	<10 or HD	650 mg once, then 325 mg PO every 12 hours	
Rifabutin	5 mg/kg PO daily (usually 300 mg PO daily) See the <u>Mycobacterium</u> <u>tuberculosis section</u> and <u>Drug–Drug Interactions</u> in the Adult and Adolescent Antiretroviral Guidelines for dosage adjustment based on interactions with ARVs.	<30	If toxicity is suspected, consider 50% of dose once daily and perform rifabutin TDM.	
Sofosbuvir [*]	400 mg PO daily	<30	Not recommended. Up to 20-fold higher sofosbuvir metabolite observed in patients with this level of renal impairment.	
Streptomycin	15 mg/kg IM or IV every 24 hours <i>or</i> 25 mg/kg IM or IV three times per week	Use with caution in patients with renal insufficiency.	TDM is no longer available. Consider an alternative aminoglycoside, as clinically appropriate. If used: 15 mg/kg two to three times weekly. Administer dose after HD.	
Sulfadiazine	1,000–1,500 mg PO every 6 hours (1,500 mg every 6 hours for patients >60 kg)	≤ 50	No data. Use alternative anti-toxoplasma therapy.	
Tecovirimat	IV: <i>35 to <120 kg</i> : 200 mg every 12 hours	30–89	No dosage adjustment necessary Use with caution due to potential accumulation of hydroxypropyl-β-cyclodextrin.	

		Dosage Adjustment in Renal Insufficiency		
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Dose	
	≥120 kg: 300 mg every 12 hours	<30	Contraindicated due to potential accumulation of hydroxypropyl-β-cyclodextrin. Note: IV formulation may be considered in patients with CrCl <30 only if drug absorption via enteral administration is expected to be problematic based on an individual risk-benefit assessment in consultation with CDC. In these circumstances, use with caution and monitor renal function continuously. Switch to the oral formulation as soon as possible.	
	PO: 40 to <120 kg: 600 mg every 12 hours ≥120 kg: 600 mg every 8 hours	Any eGFR	No dosage adjustment necessary	
Tenofovir [*] Alafenamide (TAF)	25 mg PO daily	<15	Not recommended	
Note: Please refer to product labels for dosing recommendations for other ARV FDC products containing FTC/TAF.		<15 on HD	No dosage adjustment required. Administer dose after HD on days of dialysis.	
Tenofovir [*] Disoproxil Fumarate (TDF)	300 mg PO daily	30-49	300 mg PO every 48 hours (consider switching to TAF for treatment of HBV)	
Note: Please refer to product labels for		10–29	300 mg PO every 72–96 hours (consider switching to alternative agent for treatment of HBV)	
dosing recommendations for other ARV FDC		<10 and not on dialysis	Not recommended	
products containing TDF.		HD	300 mg PO once weekly; administer dose after dialysis	
Trimethoprim [*] / Sulfamethoxazole (TMP-SMX)	For PCP Treatment 5 mg/kg (of TMP	15–30	5 mg/kg (TMP) IV every 12 hours, or two TMP-SMX DS tablets PO every 12 hours	
(TMP-SMX)	component) IV every 6–8 hours, <i>or</i>	<15	5 mg/kg (TMP) IV every 24 hours, or one TMP-SMX DS tablet PO every 12 hours (or two TMP-SMX DS tablets every 24 hours)	

		Dosage	e Adjustment in Renal Insufficiency
Drug(s)	Usual Dose	CrCl [^] or eGFR [#] (mL/min)	Dose
	Two TMP-SMX DS tablets PO every 8 hours	HD	5 mg/kg/day (TMP) IV, or two TMP-SMX DS tablets PO daily; administer dose after HD on days of dialysis.
			Consider TDM to optimize therapy (target TMP concentrations: 5–8 mcg/mL).
	For PCP Prophylaxis	15–30	Reduce dose by 50% (e.g., 1 SS tablet PO daily).
	One TMP-SMX DS tablet PO daily,	<15	Reduce dose by 50% or use alternative agent.
	One TMP-SMX DS tablet PO three times per week, or		
	One TMP-SMX SS tablet PO daily		
	For Toxoplasmosis Encephalitis (TE) Treatment: 5 mg/kg (TMP component) IV or PO every 12 hours	15–30	5 mg/kg (TMP component) IV or PO every 24 hours
		<15	5 mg/kg (TMP component) IV or PO every 24 hours or use alternative agent
	 For TE Chronic Maintenance Therapy One TMP-SMX DS tablet twice daily, or 	15–30	Reduce dose by 50%.
		<15	Reduce dose by 50% or use alternative agent.
	One TMP-SMX DS tablet daily		
	For Toxoplasmosis	15–30	Reduce dose by 50%.
	Primary Prophylaxis: One TMP-SMX DS tablet PO daily	<15	Reduce dose by 50% or use alternative agent.
Valacyclovir*	For Herpes Zoster: 1 g PO	30–49	1 g PO every 12 hours
	three times daily	10–29	1 g PO every 24 hours
		<10	500 mg PO every 24 hours
		HD	500 mg PO every 24 hours; administer dose after HD on days of dialysis.
		30–49	No dosage adjustment
		10–29	For Treatment: 1 g PO every 24 hours

		Dosage Adjustment in Renal Insufficiency					
Drug(s)	Drug(s) Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Do	se			
	For Herpes Simplex Virus Treatment: 1 g PO twice daily		For Suppressive Therapy hours	r: 500 mg PO every 24			
	For Herpes Simplex	<10	500 mg PO every 24 hours				
	Chronic Suppressive Therapy: 500 mg PO twice daily	HD	500 mg PO every 24 hours HD on days of dialysis.	; administer dose after			
Valganciclovir	Induction Therapy: 900 mg PO twice daily	CrCl [^] or eGFR [#] (mL/min)	Induction	Maintenance			
	Maintenance Therapy: 900 mg PO once daily	40–59	450 mg PO twice daily	450 mg PO daily			
		26–39	450 mg PO daily	450 mg PO every 48 hours			
		10–25	450 mg PO every 48 hours	450 mg PO twice weekly			
	<10 and not on	Not recommended	Not recommended				
		dialysis	Use IV ganciclovir.	Use IV ganciclovir.			
			May consider:200 mg (oral powder for solution) PO three times per week	May consider:100 mg (oral powder for solution) PO three times per week			
			If oral powder formulation is not available, consider: • 450 mg (tablet) PO three times weekly	If oral powder formulation is not available, consider: • 450 mg (tablet) PO twice weekly			
		HD	Not recommended	Not recommended			
			Use IV ganciclovir.	Use IV ganciclovir.			
			 May consider: 200 mg (oral powder for solution) PO three times per week after HD 	 May consider: 100 mg (oral powder for solution) PO three times per week after HD 			
							If oral powder formulation is not available, may consider: • 450 mg (tablet) PO three times per week after HD

		Dosage	Adjustment in Renal Insufficiency	
Drug(s)	Usual Dose	CrCI [^] or eGFR [#] (mL/min)	Dose	
Voriconazole [*]	6 mg/kg IV every 12 hours for two doses, then 4 mg/kg IV every 12 hours <i>or</i> 200–300 mg PO every 12 hours	<50	IV voriconazole is not recommended by the manufacturer because of potential toxicity due to accumulation of SBCD (vehicle of IV product). An observational study did not find worsening in renal function in patients with CrCl <50 ml/min. Switch patients with CrCl <50 ml/min to oral voriconazole when feasible. No need for dosage adjustment when the oral dose is used. Perform TDM to adjust dose.	

* Drugs marked with asterisk (*) are those known to have assays available (for clinical and/or research purposes) within the United States and typically in Europe. When TDM is appropriate, clinicians should contact their clinical laboratory to determine assay availability and turnaround time for their institution.

^a The prescribing information for emtricitabine (Emtriva) recommends adjusting doses for patients with CrCl 30-49 and for patients on hemodialysis. However, the prescribing information for several FDC products that contain emtricitabine (including Descovy, Biktaryy, Genvoya, and Odefsey) recommends that the standard dose (emtricitabine 200 mg) can be given once daily in these patients (on days of hemodialysis, give after completion of dialysis). The recommendations in this table incorporate the dosing guidance from the FDC products.

^b The prescribing information for lamivudine (Epivir) recommends dosage adjustment from 300 mg once daily to 150 mg once daily for patients with CrCl 30–49 mL/min. However, the prescribing information for several FDC products that contain lamivudine (including Epzicom, Dovato, and Triumeq) recommends no dose adjustment for CrCl 30–49 mL/min. The recommendation in this table incorporates the dosing guidance from the FDC products.

[^] Creatinine Clearance Calculation	
Male:	Female:
$\frac{(140 - age in years) \times weight in kg}{72 \times serum creatinine}$	$\frac{(140 - age in years) \times weight in kg \times 0.85}{72 \times serum creatinine}$

[#]When estimating kidney function to facilitate drug dosing in patients with renal insufficiency, please refer to the drug's prescribing information and to the National Institute of Diabetes and Digestive and Kidney Diseases' <u>Determining Drug Dosing in</u> <u>Adults with Chronic Kidney Disease</u> page for a discussion on using CrCl based on the Cockcroft-Gault equation versus eGFR.

Key: 3TC = lamivudine; ARV = antiretroviral; CAPD = continuous ambulatory peritoneal dialysis; CBC = complete blood count; CMV = cytomegalovirus; COBI = cobicistat; CrCI = creatinine clearance; DS = double strength; eGFR = estimated glomerular filtration rate; ESRD = end-stage renal disease; FDC = fixed-dose combination; FTC = emtricitabine; HBV = hepatitis B virus; HD = hemodialysis; HSV = herpes simplex virus; IM = intramuscular; IV = intravenous; MAI = *Mycobacterium avium intracellulare;* MTB = *Mycobacterium tuberculosis;* N/A = not applicable; OI = opportunistic infection; PCP = *Pneumocystis* pneumonia; PD = peritoneal dialysis; PI = protease inhibitor; PO = orally; SCr = serum creatinine; SQ = subcutaneous; SBCD = sulfobutylether cyclodextrin; SS = single strength; TAF = tenofovir alafenamide; TDF = tenofovir disoproxil fumarate; TDM = therapeutic drug monitoring; TMP-SMX = trimethoprim-sulfamethoxazole; VZV = varicella zoster virus