

SUPPLEMENTARY TABLE. Lenacapavir drug interactions reported in association with selected commonly prescribed medications
— Drug Prescribing Information^{*,†,§,¶}

Drug Class	Drug Name	Comment	Effect on Concentration
Antimycobacterial	Rifampin (strong CYP3A inducer)	Supplemental LEN doses required	↓ LEN
Anticonvulsant	Carbamazepine, Phenytoin (strong CYP3A inducers)	Supplemental LEN doses required	↓ LEN
Benzodiazepine	Midazolam (Oral), Clonazepam	Use with caution. Consider alternative. Dose adjustment might be required.	↑ Midazolam (Oral), Clonazepam
Corticosteroid (systemic)	Dexamethasone, Hydrocortisone, Prednisone	Dose adjustment required. There is a risk for Cushing's syndrome and adrenal suppression.	↑ Dexamethasone, Hydrocortisone, Prednisone
Herbal Supplement	St John's Wort (strong CYP3A inducer)	Supplemental LEN doses required	↓ LEN
Hormonal Contraceptive (oral, injection)	Levonorgestrel, Ethinyl Estradiol, Depot MPA	No dose adjustment required	—
Hormones (other)	Estradiol, Testosterone	No dose adjustment required	—
Lipid Modifying	Lovastatin, Simvastatin	Dose adjustment required. Monitor for myopathy.	↑ Lovastatin, Simvastatin
Narcotic Analgesic	Fentanyl, Oxycodone, Tramadol	Careful monitoring of adverse reactions is recommended, including respiratory depression. Dose adjustment might be needed.	↑ Fentanyl, Oxycodone, Tramadol
Opioid Dependence Treatment	Buprenorphine, Methadone	Dose adjustment required	Unknown; possible ↑ Buprenorphine, Methadone
Phosphodiesterase-5 Inhibitor (Erectile Dysfunction)	Sildenafil, Tadalafil, Vardenafil	Dose adjustment required	↑ Sildenafil, Tadalafil, Vardenafil

Abbreviations: LEN = lenacapavir; MPA = medroxyprogesterone acetate

* Food and Drug Administration YEZTUGO (lenacapavir) Label:

https://www.accessdata.fda.gov/drugsatfda_docs/label/2025/220020s000lbl.pdf

† Department of Health and Human Services Guidelines for the Use of Antiretroviral Agents in Adults and Adolescents with HIV 2024:

<https://clinicalinfo.hiv.gov/en/guidelines/adult-and-adolescent-arv>

§ University of Liverpool HIV Drug Interactions Checker: <https://www.hiv-druginteractions.org/checker>

¶ Bekker LG, et al. Lenacapavir dosing in special situations: tuberculosis and beyond. 13th International AIDS Society Conference on HIV Science, Kigali, session OAC05, 2025. Available from: <https://www.askgileadmedical.com/conferences/hiv-aids-liver-diseases/IAS%202025>.