

Drug-Drug Interaction Guide: From HIV Prevention to Treatment

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Table 33: Sleep Medications (also see prescribing information)

→ Non-benzodiazepine “Z-drugs,” melatonin, ramelteon, suvorexant

| Class or Drug | Mechanism of Action | Clinical Comments |
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| <ul style="list-style-type: none">• NRTIs• Dolutegravir (DTG)• Bictegravir (BIC)• Cabotegravir (CAB)• Raltegravir (RAL)• Rilpivirine (RPV)• Doravirine (DOR)• Fostemsavir (FTR) | No significant interactions reported. | No dose adjustments are necessary. |
| Elvitegravir (EVG), boosted | <ul style="list-style-type: none">• Suvorexant is a CYP3A substrate.• COBI inhibits CYP3A.• Zolpidem, eszopiclone: These drugs are CYP3A substrates and may be increased by strong inhibitors of this enzyme. | <ul style="list-style-type: none">• Suvorexant: Avoid concomitant use or use lowest effective dose (may increase somnolence, dizziness, and risk of sleep hangover).• Zolpidem: Administer lowest possible dose of zolpidem; monitor for adverse effects.• Eszopiclone: Start with 1 mg of eszopiclone at bedtime and titrate slowly to achieve clinical effect. |
| Boosted PIs | <ul style="list-style-type: none">• Zolpidem, suvorexant: Boosted PIs may increase zolpidem and suvorexant concentrations.• Ramelteon: RTV-boosted PIs may reduce ramelteon efficacy. | <ul style="list-style-type: none">• Zolpidem: Administer lowest effective dose; monitor for adverse effects, including excess sedation.• Eszopiclone: Start with 1 mg per day and titrate slowly to achieve clinical effect; monitor for adverse effects, including excess sedation.• Suvorexant: Coadministration is not recommended (may increase somnolence, dizziness, and risk of sleep hangover); use alternative sleep medication or ARV.• Ramelteon: Monitor for efficacy in cigarette smokers. |
| <ul style="list-style-type: none">• Efavirenz (EFV)• Etravirine (ETR) | Zolpidem: EFV and ETR may reduce zolpidem concentrations. | <ul style="list-style-type: none">• Zolpidem, eszopiclone: Monitor for efficacy; no dose adjustments are recommended.• Suvorexant: Monitor for efficacy; do not exceed 20 mg per day. |

Abbreviations: ARV, antiretroviral; NRTI, nucleoside reverse transcriptase inhibitor; PI, protease inhibitor; RTV, ritonavir.