



Resource: ART Drug-Drug Interactions

April 2023

| Table 31: Benzodiazepines [a] (also see drug package inserts) | | |
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| Class or Drug | Mechanism of Action | Clinical Comments |
| <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"> Boosting with cobicistat: May increase benzodiazepine concentrations via CYP3A4 inhibition. Midazolam, triazolam: Levels likely to be increased by COBI-boosted EVG | <ul style="list-style-type: none"> Alprazolam, clonazepam, diazepam: Consider alternative benzodiazepine (e.g., lorazepam, oxazepam, temazepam); if used, administer lowest effective dose; monitor closely for adverse effects. Midazolam: <ul style="list-style-type: none"> Oral: Concomitant use is contraindicated. Parenteral: Administer in closely monitored setting. Consider dose reduction, especially if >1 dose is administered. Triazolam: Concomitant use is contraindicated. |
| Boosted PIs | <ul style="list-style-type: none"> Alprazolam: Boosted ARVs may increase alprazolam concentrations via CYP3A4 inhibition. Diazepam: CYP3A4 inhibition may reduce metabolism of diazepam. | <ul style="list-style-type: none"> Consider alternative benzodiazepine (e.g., lorazepam, oxazepam, temazepam). If used, administer lowest effective dose; monitor closely for adverse effects. Diazepam: Monitor for excess sedation. |
| Efavirenz (EFV) | Alprazolam, diazepam: EFV may reduce alprazolam and diazepam concentrations. | <ul style="list-style-type: none"> Alprazolam: Monitor for benzodiazepine withdrawal if patient starts taking EFV. Alprazolam, clonazepam, diazepam: Titrate slowly to achieve clinical effect; monitor for benzodiazepine efficacy. |
| Etravirine (ETR) | Alprazolam, diazepam: ETR may reduce alprazolam and diazepam concentrations. | Alprazolam: Monitor for benzodiazepine withdrawal. |
| Lenacapavir (LEN) | Midazolam (oral), triazolam: Moderate inhibition of CYP3A4 and P-gP potentially increases sedative levels. | Midazolam (oral), triazolam: Use with caution; monitor for excess sedation. |
| <p>Abbreviations: ARV, antiretroviral; COBI, cobicistat; CYP, cytochrome P450; NRTI, nucleoside reverse transcriptase inhibitor; P-gP, P-glycoprotein; PI, protease inhibitor.</p> <p>Note:</p> <p>a. Lorazepam, oxazepam, and temazepam do not interact clinically with and do not require dose adjustments when coadministered with ARVs.</p> | | |