Lamivudine (Epivir)

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Drug Summary

Lamivudine has been extensively used for treatment of HIV infection since the mid-1990s, in both initial therapy and salvage therapy. Currently it is most frequently used as a component of the combination tablets dolutegravir-abacavir-lamivudine or abacavir-lamivudine. Two other coformulations with lamivudine (lamivudine-zidovudine and lamivudine-zidovudine-abacavir) have fallen out of favor due to toxicity from the zidovudine component of these combinations. Lamivudine is well-tolerated, rarely causes adverse effects, and has few drug interactions. Lamivudine is nearly identical to emtricitabine and the two generally are considered to be equivalent; they are used interchangeably but never together (since these two drugs have no significant additive potency and have equivalent resistance profiles). Either lamivudine or emtricitabine is included as one of two nucleoside analogues in all antiretroviral regimens recommended for initial therapy. Lamivudine has a relatively low barrier to resistance and if a patient develops virologic failure while taking a regimen that includes lamivudine, the first NRTI resistance mutation to occur is generally a lamivudine resistance mutation. Lamivudine has activity against hepatitis B, though it should only be used in combination with another anti-hepatitis B antiretroviral for treatment of hepatitis B because lamivudine monotherapy leads to high rates of hepatitis B resistance and treatment failure. The lamivudine dose should be reduced in the setting of renal insufficiency.

Key Clinical Trials

In the early era of antiretroviral therapy, before initial 3-drug ART became the standard of care, lamivudine was added to other NRTIs, such a zidovudine or stavudine, for dual therapy and this strategy demonstrated immunologic benefit as compared to continued zidovudine or stavudine monotherapy [NUCA 3001, NUCA 3002, and ACTG 306]. Subsequently, lamivudine was studied in combination with another NRTI plus an early protease inhibitor or NNRTI for 3-drug therapy and found to be effective at achieving virologic suppression; for example, lamivudine was studied in combination with zidovudine and efavirenz [GS-934], as well as with abacavir and either ritonavir-boosted fosamprenavir [KLEAN] or lopinavir [HEAT]. More recently, lamivudine was shown to be safe and effective in combination therapy with abacavir and the integrase inhibitors raltegravir [SHIELD] or dolutegravir [SINGLE]. It is also being studied as part of 2-drug initial or simplification regimens, such as lamivudine plus dolutegravir [ACTG 5353, PADDLE, GEMINI-1, and GEMINI-2] or lamivudine plus a boosted protease inhibitor [GARDEL, OLE, SALT, DUAL-GESIDA, and AtLaS].
Adverse Effects

Overall, lamivudine tends to be well tolerated. It may rarely cause side effects such as headache, diarrhea, nausea, or rash. It may be difficult to distinguish the cause of these side effects, since they are more commonly caused by other antiretrovirals in a regimen.

Resistance

For a listing of the most common clinically significant mutations associated with lamivudine (3TC) resistance, see the NRTI Resistance Notes on the Stanford University HIV Drug Resistance Database.

Key Drug Interactions

For complete information on lamivudine-related drug interactions, see the Drug Interactions section in the Lamivudine (Epivir) Prescribing Information.