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Draft Guidance on Brincidofovir

May 2023

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In general, FDA's guidance documents do not establish legally enforceable responsibilities. Instead, guidances describe the Agency's current thinking on a topic and should be viewed only as recommendations, unless specific regulatory or statutory requirements are cited. The use of the word *should* in Agency guidances means that something is suggested or recommended, but not required.

Active Ingredient: Brincidofovir

Dosage Form; Route: Tablet; Oral

Recommended Studies: Two in vivo bioequivalence studies with pharmacokinetic endpoints

1. Type of study: Fasting
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 100 mg
   Subjects: Healthy males not of reproductive potential (i.e., surgically sterile) and females not of reproductive potential
   Additional comments: Exclude subjects with abnormal liver function tests. Monitor liver function tests prior to dosing, one to two weeks post-dosing, and at end of study. Monitor subjects until resolution of adverse events. Ensure an adequate washout period between treatments in the crossover study due to the long elimination half-life of cidofovir-diphosphate.

2. Type of study: Fed
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 100 mg
   Subjects: Healthy males not of reproductive potential (i.e., surgically sterile) and females not of reproductive potential
   Additional comments: See comments above.

Analyte to measure: Brincidofovir in plasma

Bioequivalence based on (90% CI): Brincidofovir

Waiver request of in vivo testing: Not applicable

Dissolution test method and sampling times: The dissolution information for this drug product can be found in the FDA's Dissolution Methods database, http://www.accessdata.fda.gov/scripts/cder/dissolution/. Conduct comparative dissolution testing on 12 dosage units each of the test and reference products. Specifications will be determined upon review of the abbreviated new drug application.

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