Contains Nonbinding Recommendations

Draft – Not for Implementation

Draft Guidance on Osilodrostat Phosphate

August 2021

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic. It does not establish any rights for any person and is not binding on FDA or the public. You can use an alternative approach if it satisfies the requirements of the applicable statutes and regulations. To discuss an alternative approach, contact the Office of Generic Drugs.

This guidance, which interprets the Agency's regulations on bioequivalence at 21 CFR part 320, provides product-specific recommendations on, among other things, the design of bioequivalence studies to support abbreviated new drug applications (ANDAs) for the referenced drug product. FDA is publishing this guidance to further facilitate generic drug product availability and to assist the generic pharmaceutical industry with identifying the most appropriate methodology for developing drugs and generating evidence needed to support ANDA approval for generic versions of this product.

The contents of this document do not have the force and effect of law and are not meant to bind the public in any way, unless specifically incorporated into a contract. This document is intended only to provide clarity to the public regarding existing requirements under the law. FDA guidance documents, including this guidance, should be viewed only as recommendations, unless specific regulatory or statutory requirements are cited. The use of the word should in FDA guidances means that something is suggested or recommended, but not required.

This is a new draft product-specific guidance for industry on generic osilodrostat phosphate.

Active Ingredient: Osilodrostat phosphate

Dosage Form; Route: Tablet; oral

Recommended Studies: Two studies

1. Type of study: Fasting
   Design: Single-dose, two treatment, two-period, crossover in vivo
   Strength: EQ 10 mg Base
   Subjects: Males and non-pregnant, non-lactating females, general population
   Additional comments: Exclude subjects with risk factors for prolonged QT interval and Torsades de Pointes. Monitor subjects for the electrocardiogram changes during the study.

---

2. Type of study: Fed
   Design: Single-dose, two treatment, two-period, crossover in vivo
   Strength: EQ 10 mg Base

Subjects: Males and non-pregnant, non-lactating females, general population
Additional comments: See comments above.

Analyte to measure: Osilodrostat in plasma

Bioequivalence based on (90% CI): Osilodrostat

Waiver request of in vivo testing: EQ 1 mg Base and EQ 5 mg Base, based on (i) acceptable bioequivalence studies on the EQ 10 mg Base strength, (ii) proportionally similar across all strengths, and (iii) acceptable in vitro dissolution testing of all strengths

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 for each of all strengths of the test and reference products. Specifications will be determined upon review of the abbreviated new drug application.

Unique Agency Identifier: PSG_212801