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Draft Guidance on Ivacaftor

May 2023

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Active Ingredient: Ivacaftor

Dosage Form; Route: Granule; 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: 75 mg/packet
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: Mix granules with 1 teaspoon (5 mL) of soft food or liquid. Conduct bioequivalence study according to the reference product labeling.

2. Type of study: Fed
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 75 mg/packet
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: See comments above.

Analyte to measure: Ivacaftor in plasma

Bioequivalence based on (90% CI): Ivacaftor

Waiver request of in vivo testing: 25 mg/packet and 50 mg/packet strengths based on (i) acceptable bioequivalence studies on the 75 mg/packet strength, (ii) acceptable in vitro dissolution testing of all strengths, and (iii) proportional similarity of the formulations across 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 each of all strengths of the test and reference products. Specifications will be determined upon review of the abbreviated new drug application.

Revision History: Recommended April 2016; Revised May 2023

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