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Draft Guidance on Baloxavir Marboxil

May 2023

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Active Ingredient: Baloxavir marboxil

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: 80 mg
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: Ensure an adequate washout period between treatments in the crossover study due to the long elimination half-life of baloxavir. Alternatively, a parallel study design may be considered (refer to FDA approved reference product labeling with regard to the pharmacokinetic differences between Asians and non-Asians).

2. Type of study: Fed
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 80 mg
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: See comments above.

Analyte to measure: Baloxavir in plasma

Bioequivalence based on (90% CI): Baloxavir

Waiver request of in vivo testing: 40 mg strength based on (i) acceptable bioequivalence studies on the 80 mg strength, (ii) acceptable in vitro dissolution testing of both strengths, and (iii) proportional similarity of the formulations between both 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 both strengths of the test and reference products. Specifications will be determined upon evaluation of the abbreviated new drug application.

Revision History: Recommended September 2019; Revised May 2023

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