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Draft Guidance on Bempedoic Acid; Ezetimibe

August 2024

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Active Ingredients: Bempedoic acid; Ezetimibe

Dosage Form: Tablet

Route: Oral

Strength: 180 mg; 10 mg

Recommended Study: One in vivo bioequivalence study with pharmacokinetic endpoints

1. Type of study: Fasting
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 180 mg; 10 mg
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: None

Analytes to measure: Bempedoic acid and its active metabolite, ESP15228; ezetimibe (unconjugated) and total ezetimibe (ezetimibe + ezetimibe glucuronide) in plasma

Bioequivalence based on (90% CI): Bempedoic acid and ezetimibe (unconjugated) and total ezetimibe (ezetimibe + ezetimibe glucuronide)

Submit the metabolite data (ESP15228) as supportive evidence of comparable therapeutic outcome. For the metabolite, the following data should be submitted: individual and mean concentrations, individual and mean pharmacokinetic parameters, and geometric means and ratios of means for area under the curve and maximum concentration.

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

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