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Draft Guidance on Promethazine Hydrochloride

August 2024

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Active Ingredient: Promethazine hydrochloride

Dosage Form: Suppository

Route: Rectal

Strengths: 12.5 mg, 25 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: 25 mg
Subjects: Healthy males and non-pregnant, non-lactating females
Additional comments: None

Analyte to measure: Promethazine in plasma

Bioequivalence based on (90% CI): Promethazine

Waiver request of in vivo testing: 12.5 mg strength based on (i) an acceptable bioequivalence study on the 25 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 each of all strengths of the test product and reference products. Specifications will be determined upon review of the abbreviated new drug application.

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