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Draft Guidance on Nateglinide

October 2024

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Active Ingredient: Nateglinide

Dosage Form: Tablet

Route: Oral

Strengths: 60 mg, 120 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: 120 mg
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: All subjects should fast overnight for at least 10 hours prior to dosing and for 4 hours after dosing. A single oral dose (120 mg) should be administered with 240 mL of 20% glucose solution. Since, multiple plasma concentration peaks were often observed under fasting conditions, please ensure that the same sampling schedule is followed during the study for both test and reference drug administration. Females of reproductive potential should use effective contraception during the study.

Analyte to measure: Nateglinide in plasma

Bioequivalence based on (90% CI): Nateglinide

Waiver request of in-vivo testing: 60 mg strength based on (i) acceptable bioequivalence study on the 120 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 product and reference listed drug (RLD.[1] Specifications will be determined upon review of the abbreviated new drug application.

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[1] If the RLD is not available, refer to the most recent version of the FDA guidance for industry on *Referencing Approved Drug Products in ANDA Submissions.*