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Draft Guidance on Tucatinib

November 2021

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This guidance, which interprets the Agency's regulations on bioequivalence at 21 CFR part 320, provides product-specific recommendations on, among other things, the design of bioequivalence studies to support abbreviated new drug applications (ANDAs) for the referenced drug product. FDA is publishing this guidance to further facilitate generic drug product availability and to assist the generic pharmaceutical industry with identifying the most appropriate methodology for developing drugs and generating evidence needed to support ANDA approval for generic versions of this product.

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This is a new draft product-specific guidance for industry on generic tucatinib.

Active Ingredient: Tucatinib

Dosage Form; Route: Tablet; oral

Recommended Studies: Two studies

1. Type of study: Fasting
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 150 mg
   Subjects: Males and non-pregnant, non-lactating females, general population
   Additional comments: Exclude subjects with abnormal liver function tests and monitor liver function tests during the study. Females of reproductive potential and male subjects with female partners of reproductive potential should use effective contraception during the study and for at least one week after the final dose. Exclude subjects with concomitant use of strong cytochrome P450 2C8 inhibitors.

2. Type of study: Fed
Design: Single-dose, two-treatment, two-period crossover in vivo
Strength: 150 mg
Subjects: Males and non-pregnant, non-lactating females, general population
Additional comments: See comments above

Analyte to measure: Tucatinib in plasma

Bioequivalence based on (90% CI): Tucatinib

Waiver request of in vivo testing: 50 mg based on (i) acceptable bioequivalence studies on the 150 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.

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