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Draft Guidance on Zanubrutinib

May 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 zanubrutinib.

Active Ingredient: Zanubrutinib

Dosage Form; Route: Capsule; oral

Recommended Studies: Two studies

1. Type of study: Fasting
   Design: Single-dose, two-treatment, two-period crossover in vivo
   Strength: 80 mg
   Subjects: Males and non-pregnant, non-lactating females, general population
   Additional comments: Advise females of reproductive potential and males with female partners of reproductive potential to use effective contraception methods during and for one week after the study. Exclude immunocompromised subjects and subjects with abnormal complete blood count tests. Exclude subjects who test positive for human immunodeficiency virus, hepatitis B virus, and hepatitis C virus using serologic tests. Exclude subjects who are taking antiplatelet or anticoagulant medications. Exclude subjects who have a surgical treatment history or plan within one week prior to the first

dose or after the last dose. Exclude subjects who have abnormal electrocardiogram, hypertension, or risk factors for cardiac arrhythmias.

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

Analyte to measure: Zanubrutinib in plasma

Bioequivalence based on (90% CI): Zanubrutinib

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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