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Draft Guidance on Avapritinib

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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 avapritinib.

Active Ingredient: Avapritinib

Dosage Form; Route: Tablet; oral

Recommended Study: One study

1. Type of study: Fasting
   Design: Single-dose, two- treatment, two-period crossover in vivo
   Strength: 300 mg
   Subjects: Males and females not of reproductive potential, general population
   Additional comments: Exclude subjects with a risk or history of intracranial hemorrhage. Females of reproductive potential should use effective contraception methods during the study and for six weeks after the study. Males with female partners of reproductive potential should use effective contraception during the study and for six weeks after the study. Ensure an adequate washout period between treatments in the crossover study due to the long elimination half-life of avapritinib tablets. Alternatively, a parallel study design may be considered.

Analyte to measure: Avapritinib in plasma

Bioequivalence based on (90% CI): Avapritinib

Waiver request of in vivo testing: The 100 mg and 200 mg strengths based on (i) acceptable bioequivalence study on the 300 mg strength, (ii) proportional similarity of the formulations across all strengths, and (iii) acceptable in vitro dissolution testing of all 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 all strengths of the test and reference products. Specifications will be determined upon evaluation of the abbreviated new drug application.

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