Contains Nonbinding Recommendations

Draft – Not for Implementation

Draft Guidance on Fruquintinib

October 2025

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic. It does not establish any rights for any person and is not binding on FDA or the public. You can use an alternative approach if it satisfies the requirements of the applicable statutes and regulations. To discuss an alternative approach, contact the Office of Generic Drugs.

In general, FDA's guidance documents do not establish legally enforceable responsibilities. Instead, guidances describe the Agency's current thinking on a topic and should be viewed only as recommendations, unless specific regulatory or statutory requirements are cited. The use of the word *should* in Agency guidances means that something is suggested or recommended, but not required.

Active Ingredient: Fruquintinib

Dosage Form: Capsule

Route: Oral

Strengths: 1 mg, 5 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: 5 mg
   Subjects: Healthy males and healthy females not of reproductive potential
   Additional comments:
   - Exclude subjects with abnormal complete blood counts or liver function tests. Exclude subjects who have undergone or plan to undergo surgery for at least two weeks prior to and after the study. Male subjects with female partners of reproductive potential should use effective contraception during the study and for two weeks after the last dose.
   - Ensure an adequate washout period between treatments in the crossover study due to the long elimination half-life of fruquintinib. Alternatively, a parallel study design may be considered.

Analyte to measure: Fruquintinib in plasma

Bioequivalence based on (90% CI): Fruquintinib

Waiver request of in vivo testing: 1 mg strength based on (i) an acceptable bioequivalence study on the 5 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.

Document History: Recommended October 2025

Unique Agency Identifier: PSG_217564

[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*.