

Contains Nonbinding Recommendations

Draft – Not for Implementation

Draft Guidance on Macitentan

November 2025

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Active Ingredient: Macitentan

Dosage Form: Tablet

Route: Oral

Strength: 10 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: 10 mg
Subjects: Healthy males and healthy females not of reproductive potential
Additional comments: None

Analyte to measure: Macitentan in plasma

Bioequivalence based on (90% CI): Macitentan

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 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 guidance for industry *Referencing Approved Drug Products in ANDA Submissions*.