

Contains Nonbinding Recommendations
Draft – Not for Implementation
Draft Guidance on Valbenazine Tosylate
February 2022

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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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In July 2018, FDA issued a draft product-specific guidance for industry on generic valbenazine tosylate. We are now issuing revised draft guidance for industry that replaces the previously issued guidance.

Active Ingredient: Valbenazine tosylate

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: EQ 80 mg Base
Subjects: Healthy males and non-pregnant, non-lactating females
Additional comments: None

2. Type of study: Fed
Design: Single-dose, two-treatment, two-period, crossover in vivo
Strength: EQ 80 mg Base
Subjects: Healthy males and non-pregnant, non-lactating females
Additional comments: None

Analytes to measure: Valbenazine and its active metabolite, α -dihydrotetrabenzazine, in plasma

Submit the metabolite data as supportive evidence of comparable therapeutic outcome. For the metabolite, the following data should be submitted: individual and mean concentrations, individual and mean pharmacokinetic parameters, and geometric means and ratios of means for AUC and C_{max}.

Bioequivalence based on (90% CI): Valbenazine

Waiver request of in vivo testing: EQ 40 mg Base and EQ 60 mg Base strengths based on (i) acceptable bioequivalence studies on the EQ 80 mg Base strength, (ii) acceptable in vitro dissolution testing of all strengths, and (iii) proportional similarity of the formulations across 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 each of all strengths of the test and reference products. Specifications will be determined upon review of the abbreviated new drug application.

Revision History: Recommended July 2018; Revised February 2022

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