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Draft – Not for Implementation

Draft Guidance on Amlodipine Besylate; Olmesartan Medoxomil

October 2024

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Active Ingredients: Amlodipine besylate; Olmesartan medoxomil

Dosage Form: Tablet

Route: Oral

Strengths: EQ 5 mg Base; 20 mg, EQ 5 mg Base; 40 mg, EQ 10 mg Base; 20 mg, EQ 10 mg Base; 40 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: EQ 10 mg Base; 40 mg
   Subjects: Healthy males and non-pregnant, non-lactating females
   Additional comments: Females of reproductive potential should use effective contraception during the study. Ensure an adequate washout period between treatments in the crossover study due to the long elimination half-life of amlodipine. Alternatively, a parallel study design may be considered.

Analytes to measure: Amlodipine and olmesartan in plasma

Bioequivalence based on (90% CI): Amlodipine and olmesartan

Waiver request of in vivo testing: EQ 5 mg Base; 20 mg, EQ 5 mg Base; 40 mg and EQ 10 mg Base; 20 mg strengths, based on (i) acceptable bioequivalence study on the EQ 10 mg Base; 40 mg 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 for each of all strengths 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 FDA guidance for industry on *Referencing Approved Drug Products in ANDA Submissions.*