Contains Nonbinding Recommendations

Guidance on Losartan Potassium

This guidance represents the Food and Drug Administration's (FDA's) current thinking on this topic. It does not create or confer any rights for or on any person and does not operate to bind FDA or the public. You can use an alternative approach if the approach satisfies the requirements of the applicable statutes and regulations. If you want to discuss an alternative approach, contact the Office of Generic Drugs.

Active ingredient: Losartan Potassium

Form/Route: Tablet/Oral

Recommended studies: 2 studies

1. Type of study: Fasting
   Design: Single-dose, two-treatment, two-period crossover *in-vivo*
   Strength: 100 mg
   Subjects: Normal healthy males and females, general population
   Additional Comments: Pregnant women should be excluded from participation in the bioequivalence studies.

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2. Type of study: Fed
   Design: Single-dose, two-treatment, two-period crossover *in-vivo*
   Strength: 100 mg
   Subjects: Normal healthy males and females, general population
   Additional comments: See comment above.

Analytes to measure (in appropriate biological fluid): Losartan and the metabolite carboxylic acid in plasma.

Bioequivalence based on (90% CI): Losartan.
Please submit the metabolite data as supportive evidence of comparable therapeutic outcome. For the carboxylic acid 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}.

Waiver request of in-vivo testing: 25 mg and 50 mg based on (i) acceptable bioequivalence studies on the 100 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:

Please note that a **Dissolution Methods Database** is available to the public at the OGD website at http://www.fda.gov/cder/ogd/index.htm. Please find the dissolution information for this product at this website. Please 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 application.