Contains Nonbinding Recommendations

Guidance on Glyburide; Metformin Hydrochloride

This guidance represenst 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: Glyburide; Metformin Hydrochloride

Form/Route: Tablets/Oral

Recommended studies: 2 studies

1. Type of study: Fasting
   Design: Single-dose, two-way crossover *in-vivo*
   Strength: 5 mg/500 mg
   Subjects: Normal healthy males and females, general population
   Additional Comments: The drug products should be administered with 240 mL of 20 % glucose solution in water, followed by 60 mL of the glucose solution administered every 15 minutes for up to 4 hours after dosing.

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2. Type of study: Fed
   Design: Single-dose, two-way crossover *in-vivo*
   Strength: 5 mg/500 mg
   Subjects: Normal healthy males and females, general population
   Additional comments: The drug products should be administered with 240 mL of 20 % glucose solution in water, followed by 60 mL of the glucose solution administered every 15 minutes for up to 4 hours after dosing.

Analytes to measure: Glyburide and Metformin

Bioequivalence based on (90% CI): Glyburide and Metformin

Waiver request of in-vivo testing: 1.25 mg/250 mg and 2.5 mg/500 mg based on (i) acceptable bioequivalence studies on the 5 mg/500 mg strength, (ii) proportionally similar 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.