Draft Guidance on Metformin Hydrochloride; Repaglinide

This draft guidance, once finalized, will represent the Food and Drug Administration's (FDA's) current thinking on this topic.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:** Metformin Hydrochloride; Repaglinide

**Form/Route:** Tablet/Oral

**Recommended studies:** 1 study

- **Type of study:** Fed
- **Design:** Single-dose, two-way, crossover in-vivo
- **Strength:** 500 mg/2 mg
- **Subjects:** Healthy males and nonpregnant females, general population
- **Additional Comments:** The drug products should be administered with 240 mL of a 20% glucose solution in water, followed by 60 mL of the glucose solution administered every 15 minutes for up to 4 hours after dosing. Please refer to the Amantadine Hydrochloride Tablet Draft Guidance for additional information regarding fed studies.

**Analytes to measure:** Metformin and repaglinide in plasma.

**Bioequivalence based on (90% CI):** Metformin and repaglinide

**Waiver request of in-vivo testing:** 500 mg/1 mg based on (i) acceptable bioequivalence studies on the 500 mg/2 mg strength, (ii) acceptable dissolution testing across all strengths, and (iii) proportional similarity in the formulations across all strengths. Please refer to the Mirtazapine Tablet Draft Guidance for additional information regarding waivers of in-vivo testing.

**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.accessdata.fda.gov/scripts/cder/dissolution/](http://www.accessdata.fda.gov/scripts/cder/dissolution/). 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.

Recommended Oct 2009, Revised Dec 2010