

Guidance on Hydrochlorothiazide; Olmesartan Medoxomil

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Active ingredient: Hydrochlorothiazide; Olmesartan Medoxomil

Form/Route: Tablets/Oral

Recommended studies: 2 studies

1. Type of study: Fasting
Design: Single-dose, two-way crossover *in-vivo*
Strength: 25 mg/40 mg
Subjects: Normal healthy males and females, general population. Females should not be pregnant, and if applicable, should practice abstinence or contraception during the study.
Additional Comments: The labeling for this drug contains a Black Box regarding pregnancy and fetal/neonatal morbidity and mortality.

2. Type of study: Fed
Design: Single-dose, two-way crossover *in-vivo*
Strength: 25 mg/40 mg
Subjects: Normal healthy males and females, general population. Females should not be pregnant, and if applicable, should practice abstinence or contraception during the study.
Additional comments: Please see comment above.

Analytes to measure (in appropriate biological fluid): Hydrochlorothiazide and Olmesartan in plasma.

Bioequivalence based on (90% CI): Hydrochlorothiazide and Olmesartan

Waiver request of in-vivo testing: 12.5 mg/40 mg and 12.5 mg/20 mg strengths based on (i) acceptable bioequivalence studies on the 25 mg/40 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.