

*Contains Nonbinding Recommendations*  
**Draft Guidance on Zileuton**

This draft guidance, once finalized, will represent the Food and Drug Administration's (FDA's) current thinking on this topic.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

**Form/Route:** Extended-Release Tablets/Oral

**Recommended studies:** 2 studies

1. Type of study: Fasting  
Design: Single-dose, two-way crossover *in vivo*  
Strength: 600 mg  
Subjects: Normal healthy males and females, general population.  
Additional Comments: Females should not be pregnant or lactating, and if applicable, should practice abstention or contraception during the study.

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2. Type of study: Fed  
Design: Single-dose, two-way crossover *in vivo*  
Strength: 600 mg  
Subjects: Normal healthy males and females, general population.  
Additional Comments: Please see comments above.

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**Analytes to measure (in appropriate biological fluid):** Zileuton in plasma

**Bioequivalence based on (90% CI):** Zileuton

**Waiver request of *in vivo* testing:** Not Applicable

**Dissolution test method and sampling times:** Please note that **Dissolution Method 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.

In addition to the method above, for modified release products, dissolution profiles on 12 dosage units each of test and reference products generated using USP Apparatus I at 100 rpm and/or Apparatus II at 50 rpm in at least three dissolution media (pH 1.2, 4.5 and 6.8 buffer) should be submitted in the application. Agitation speeds may have to be increased if appropriate. It is acceptable to add a small amount of surfactant, if necessary. Please include early sampling times of 1, 2, and 4 hours and continue every 2 hours until at least 80% of the drug is released, to provide assurance against premature release of drug (dose dumping) from the formulation. Specifications will be determined upon review of the data submitted in the application.