

## Draft Guidance on Abiraterone Acetate

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic. It does not establish any rights for any person and is not binding on FDA or the public. You can use an alternative approach if it satisfies the requirements of the applicable statutes and regulations. To discuss an alternative approach, contact the Office of Generic Drugs.

**Active Ingredient:** Abiraterone acetate

**Dosage Form; Route:** Tablet; oral

**Recommended Studies:** Two studies

1. Type of study: Fasting  
Design: Single-dose, two-treatment, two-period crossover in vivo  
Strength: 500 mg  
Subjects: Males, general population

**Additional comments:**

Study population should consist of males at least 18 years old.

Study participants must be in good general health as determined by no clinically significant findings on medical history, physical examination, vital signs, 12-lead electrocardiogram consistent with normal cardiac conduction and function, and clinical laboratory measurements.

Monitor vital signs and serum electrolytes during the course of the study.

Perform a 12-lead ECG at baseline and at the end of the study.

Inform study participants that it is not known whether abiraterone or its metabolites are present in semen. Therefore, study participants should use adequate contraception during the study and for at least one week after the last abiraterone acetate dose.

Applicants may consider using a reference-scaled average bioequivalence approach for this highly variable drug substance/product. (For additional information, see Progesterone Oral Capsule Guidance.) Provide evidence of high variability in the bioequivalence parameters, AUC and/or C<sub>max</sub> (i.e., within-subject variability  $\geq 30\%$ ), from the study results, when using this approach.

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2. Type of study: Fed  
Design: Single-dose, two-treatment, two-period crossover in vivo  
Strength: 500 mg  
Subjects: Males, general population

**Additional comments:**

The reference listed drug (RLD) labeling indicates that abiraterone acetate tablets must be taken on an empty stomach. However, due to the complex food effects associated with abiraterone acetate tablets, FDA recommends applicants conduct a BE study under fed conditions.

See the fasting study design above for additional comments.

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

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

**Waiver request of in vivo testing:** 250 mg based on (i) acceptable bioequivalence studies on the 500 mg strength, (ii) acceptable in-vitro dissolution testing of all strengths, and (iii) proportional similarity of the formulations across all strengths.

**Dissolution test method and sampling times:** The dissolution information for this drug product can be found on the FDA-Recommended Dissolution Methods web site, available to the public at the following location: <http://www.accessdata.fda.gov/scripts/cder/dissolution/>. 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 abbreviated new drug application (ANDA).