

Guidance on Hydrocortisone

This guidance represents 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: Hydrocortisone

Dosage Form; Route: Tablet; oral

Recommended Studies: Two studies

1. Type of study: Fasting
Design: Single-dose, two-way, crossover *in-vivo*
Strength: 20 mg
Subjects: Healthy males and nonpregnant females, general population
Additional Comments: A 4 mg dose of dexamethasone should be administered 10 hours prior to drug administration as a pre-treatment to lower endogenous hydrocortisone levels.

2. Type of study: Fed
Design: Single-dose, two-way, crossover *in-vivo*
Strength: 20 mg
Subjects: Healthy males and nonpregnant females, general population
Additional comments: See comment above.

Analytes to measure (in appropriate biological fluid): Hydrocortisone in plasma

Bioequivalence based on (90% CI): Hydrocortisone

Waiver request of in-vivo testing: 5 mg and 10 mg based on (i) acceptable bioequivalence studies on the 20 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 website, 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).