Contains Nonbinding Recommendations

Draft Guidance on Ethinyl Estradiol; Norethindrone Acetate

Active Ingredient: Ethinyl Estradiol; Norethindrone Acetate

Dosage Form; Route: Tablet, chewable; oral

Recommended Studies: Three studies

1. Type of study: Fasting
   Design: Single-dose, two-way crossover in-vivo
   Strength: 0.01 mg/1 mg Ethinyl Estradiol and Norethindrone Acetate
   Subjects: Healthy non-pregnant females, general population.
   Additional Comments: 1. Subjects should not be taking hormonal contraceptives. 2. The drug product should be administered as specified in the labeling, i.e., chewed and swallowed with water.

---

2. Type of study: Fed
   Design: Single-dose, two-way crossover in-vivo
   Strength: 0.01 mg/ 1 mg Ethinyl Estradiol and Norethindrone Acetate
   Subjects: Healthy non-pregnant females, general population. Additional Comments: Please see comments above.

---

3. Type of study: Fasting
   Design: Single-dose, two-way crossover in-vivo
   Strength: 0.01mg Ethinyl Estradiol
   Subjects: Healthy non-pregnant females, general population.
   Additional Comments: 1. Subjects should not be taking hormonal contraceptives. 2. The drug product should be administered as specified in the labeling, i.e., swallowed whole with water.

Analytes to measure (in appropriate biological fluid): Ethinyl Estradiol and Norethindrone in plasma

Bioequivalence based on (90% CI): Ethinyl Estradiol; Norethindrone

Waiver request of in-vivo testing: Not Applicable

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).