

Guidance on Lamotrigine

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Active ingredient: Lamotrigine

Form/Route: Tablet /Oral

Recommended studies: 2 studies

1. Type of study: Fasting
Design: Single-dose, two-treatment, two-period crossover *in-vivo*
Strength: Single-dose of 50 mg (2 x 25 mg)
Subjects: Normal healthy males and females, general population
Additional Comments: Due to safety concerns, studies on the highest strength are not recommended.

2. Type of study: Fed
Design: Single-dose, two-treatment, two-period crossover *in-vivo*
Strength: Single-dose of 50 mg (2 x 25 mg)
Subjects: Normal healthy males and females, general population
Additional comments: See comment above.

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

* Please utilize a validated analytical method such as LC-MS/MS to reliably measure plasma lamotrigine concentrations. A lower limit of quantitation (LOQ) of 10 ng/mL is recommended to adequately characterize the pharmacokinetics at 50 mg study dose.

Bioequivalence based on (90% CI): Lamotrigine

Waiver request of in-vivo testing: 100 mg, 150 mg and 200 mg based on (i) acceptable bioequivalence studies on the 25 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 and conduct comparative dissolution testing on 12 dosage units of all strengths of the test and reference products. Specifications will be determined upon review of the application.