

Draft Guidance on Alvimopan

This draft guidance, once finalized, will represent the Food and Drug Administration's (FDA's) current thinking on this topic.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

Form/Route: Capsule; Oral

Recommended studies: 2 studies

1. Type of study: Fasting
Design: Single-dose, two-way crossover in vivo
Strength: 12mg
Subjects: Healthy males and non-pregnant females, general population
Additional comments: Entereg® (alvimopan) Capsules was approved with a Risk Evaluation and Mitigation Strategy (REMS), which restricts its use to short-term (15 doses) use in hospitalized patients because of the risk of myocardial infarction observed with longer use. All pertinent elements of the REMS must be incorporated into the protocol and informed consent.

2. Type of study: Fed
Design: Single-dose, two-way crossover in vivo
Strength: 12mg
Subjects: Healthy males and non-pregnant females, general population
Additional comments: (1) Please see comment above. (2) The meal should be completed within 30 minutes and the test and reference products should be administered 30 minutes after the start of the meal.

Analytes to measure (in appropriate biological fluid): Alvimopan and the metabolite ADL-08-0011 in plasma

Bioequivalence based on (90% CI): Alvimopan

Waiver request of in vivo testing: Not Applicable

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.accessdata.fda.gov/scripts/cder/dissolution/>. Please find the dissolution information for this product at this website. Please 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 application.

Risk Evaluation and Mitigation Strategy (REMS): This drug product has an innovator developed REMS. The innovator has named the REMS 'Entereg Access Support and Education' (E.A.S.E.). Any ANDA citing the innovator drug product will also be required to have a REMS. Please refer to 73 FR 16313: March 27, 2008 and Section 505-1 of the Food, Drug and Cosmetic Act.