

Draft Guidance on Sonidegib Phosphate

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic.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: Sonidegib Phosphate

Dosage Form; Route: Capsule; oral

Recommended Studies: One study

1. Type of study: Fasting

Design: Single-dose, two-way crossover in vivo

Strength: EQ 200 mg

Subjects: Healthy males and females (non-pregnant), general population

Additional Comments:

- All the warnings specified in sonidegib labeling should be followed and appropriately incorporated in the bioequivalence (BE) study design and informed consent. Females should not be pregnant or lactating and the pregnancy status of females of reproductive potential should be verified prior to initiating therapy. Study subjects should be informed about the potential for embryo-fetal toxicity and compromised female fertility. Appropriate safeguards, including adequate contraception, should be incorporated in the study protocol for both males and females.
- Sonidegib has a long terminal elimination half-life (>24 hours). Therefore, adequate washout periods should be ensured between treatments in the crossover studies. A parallel study design may also be considered due to its long half-life. For a crossover or parallel study, sample collection time should be adequate to ensure completion of gastrointestinal transit of the drug product and absorption of the drug substance. For long half-life drug products that demonstrate low intrasubject variability in distribution and clearance, an AUC truncated to 72 hours may be used in place of AUC_{0-t} or $AUC_{0-\infty}$.
- Sufficient blood samples should be collected in the BE studies to adequately characterize the peak concentration (C_{max}) and time to reach peak concentration (T_{max}).

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

Bioequivalence based on (90% CI): Sonidegib

Waiver request of in vivo testing: N/A

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 .