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Draft Guidance on Niraparib Tosylate

February 2024

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Active Ingredient:	Niraparib tosylate
Dosage Form:	Tablet
Route:	Oral
Strengths:	EQ 100 mg Base, EQ 200 mg Base, EQ 300 mg Base
Recommended Study:	One in vivo bioequivalence study with pharmacokinetic endpoints

1. Type of study: Steady-state
Design: Multiple-dose, two-treatment, two-period crossover
Strength: Either EQ 200 mg Base only or EQ 300 mg Base only
Subjects: Female patients with advanced epithelial ovarian, fallopian tube, or primary peritoneal cancer who are in a complete or partial response to first-line platinum-based chemotherapy or female patients with deleterious or suspected deleterious germline BRCA-mutated recurrent epithelial ovarian, fallopian tube, or primary peritoneal cancer who are in complete or partial response to platinum-based chemotherapy.
Additional comments: Exclude patients who require dosage modification or with expected changes in concomitant medications that may potentially affect the pharmacokinetics of niraparib during the study. Females of reproductive potential should use effective contraception during treatment and 6 months after the last dose of niraparib. Implement safety precautions and monitoring including complete blood count during treatment as recommended in the labeling. Niraparib can be administered with or without food per the labeling. For the purpose of a bioequivalence study, patients should be instructed to take the drug under similar food conditions during both periods of the study. Submission of an investigational new drug application is required prior to the conduct of a bioequivalence study for a cytotoxic drug of niraparib pursuant to 21 CFR § 320.31.

Analyte to measure: Niraparib in plasma

Bioequivalence based on (90% CI): Niraparib

Waiver request of in vivo testing: EQ 100 mg Base and EQ 200 mg Base strengths based on (i) acceptable bioequivalence study on the EQ 300 mg Base strength, (ii) acceptable in vitro dissolution testing across all the strengths, and (iii) proportional similarity of the formulations across all the strengths

Alternatively, EQ 100 mg Base and EQ 300 mg Base strengths based on (i) acceptable bioequivalence study on the EQ 200 mg Base strength, (ii) acceptable in vitro dissolution testing across all the strengths, and (iii) proportional similarity of the formulations across all the strengths

Dissolution test method and sampling times: The dissolution information for this drug product can be found in the FDA's Dissolution Methods database, <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.

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