

*Contains Nonbinding Recommendations*

*Draft – Not for Implementation*

**Draft Guidance on Sofosbuvir**

**March 2021**

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic.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

This guidance, which interprets the Agency’s regulations on bioequivalence at 21 CFR part 320, provides product-specific recommendations on, among other things, the design of bioequivalence studies to support abbreviated new drug applications (ANDAs) for the referenced drug product. FDA is publishing this guidance to further facilitate generic drug product availability and to assist the generic pharmaceutical industry with identifying the most appropriate methodology for developing drugs and generating evidence needed to support ANDA approval for generic versions of this product.

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This is a new draft product-specific guidance for industry on generic sofosbuvir.

**Active Ingredient:** Sofosbuvir

**Dosage Form; Route:** Pellets; oral

**Recommended Studies:** Two studies

1. Type of study: Fasting  
Design: Single-dose, two-treatment, two-period crossover in vivo  
Strength: 200 mg/packet  
Subjects: Males and non-pregnant, non-lactating females, general population  
Additional comments: Measure hepatitis B surface antigen and hepatitis B core antibody and exclude subjects with evidence of current or prior hepatitis B virus infection. Due to the potential risk of serious symptomatic bradycardia, exclude subjects who are taking amiodarone.

2. Type of study: Fed  
Design: Single-dose, two-treatment, two-period crossover in vivo  
Strength: 200 mg/packet  
Subjects: Males and non-pregnant, non-lactating females, general population  
Additional comments: See comments above.
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**Analyte to measure:** Sofosbuvir in plasma

**Bioequivalence based on (90% CI):** Sofosbuvir

**Waiver request of in vivo testing:** 150 mg/packet based on (i) acceptable bioequivalence studies on the 200 mg/packet strength, (ii) proportional similarity of the formulations between both strengths, and (iii) acceptable in vitro dissolution testing of both 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 for each of both strengths of the test and reference products. Note that a dosage unit for pellets is the unit-dose packet. Specifications will be determined upon review of the abbreviated new drug application.

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