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*Draft – Not for Implementation*

**Draft Guidance on Sofosbuvir**

**October 2024**

This draft guidance, when finalized, will represent the current thinking of the Food and Drug Administration (FDA, or the Agency) on this topic.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

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**Active Ingredient:** Sofosbuvir

**Dosage Form:** Tablet

**Route:** Oral

**Strengths:** 200 mg, 400 mg

**Recommended Study:** One in vivo bioequivalence study with pharmacokinetic endpoints

1. Type of study: Fasting  
Design: Single-dose, two-treatment, two-period crossover in vivo  
Strength: 400 mg  
Subjects: Healthy males and non-pregnant, non-lactating females  
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.

**Analyte to measure:** Sofosbuvir in plasma

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

**Waiver request of in vivo testing:** 200 mg strength based on (i) acceptable bioequivalence study on the 400 mg strength, (ii) acceptable in vitro dissolution testing of both strengths, and (iii) proportional similarity of the formulations between 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 product and reference listed drug (RLD).<sup>1</sup> Specifications will be determined upon review of the abbreviated new drug application.

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**Document History:** Recommended June 2015; Revised May 2021, October 2024

**Unique Agency Identifier:** PSG\_204671

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<sup>1</sup> If the RLD is not available, refer to the most recent version of the FDA guidance for industry on *Referencing Approved Drug Products in ANDA Submissions*.