

Epclusa[®] (sofosbuvir/velpatasvir) Coadministration with Primidone

This document is in response to your request for information regarding Epclusa[®] (sofosbuvir/velpatasvir [SOF/VEL]) and coadministration with primidone.

Some data may be outside of the US FDA-approved Prescribing Information. In providing this data, Gilead Sciences, Inc. is not making any representation as to its clinical relevance or to the use of any Gilead product(s). For information about the approved conditions of use of any Gilead drug product, please consult the FDA approved prescribing information.

The full indication, important safety information, and boxed warnings are available at: www.gilead.com/-/media/files/pdfs/medicines/liver-disease/epclusa/epclusa_pi.

PK DDI Evaluation

Drug interaction studies have not been conducted between the single-tablet regimen SOF/VEL and primidone. Based on the PK profile of each active ingredient within SOF/VEL and primidone, a PK interaction would be predicted.^{1,2} Coadministration may result in decreased SOF/VEL concentrations due to induction of P-gp by phenobarbital, an active metabolite of primidone. Coadministration is not recommended. For more information about primidone, please refer to its product labeling.²

SOF/VEL PK¹

DDI Mechanism		SOF	VEL
Drug Transporters	P-gp/BCRP	Substrate	Substrate/Inhibitor
	OATP1B1	N/A	Inhibitor
	OATP1B3	N/A	Inhibitor
	OATP2B1	N/A	Inhibitor
Drug Metabolizing Enzymes	CYP1A2	N/A	N/A
	CYP2B6	N/A	Substrate
	CYP2C8	N/A	Substrate
	CYP2C19/19	N/A	N/A
	CYP2D6	N/A	N/A
	CYP3A4	N/A	Substrate

Relevant SOF/VEL Label Information¹

There is no information in the SOF/VEL product labeling about the coadministration of SOF/VEL and primidone.

Clearance of HCV infection with direct acting antivirals may lead to changes in hepatic function, which may impact safe and effective use of concomitant medications. Frequent monitoring of relevant laboratory parameters (INR or blood glucose) and dose adjustments of certain concomitant medications may be necessary. As described in Section 7.3 of the

SOF/VEL US Prescribing Information (Established and Potentially Significant Drug Interactions), coadministration of SOF/VEL with phenobarbital is predicted to decrease the concentrations of SOF and VEL. Coadministration of these agents is not recommended.

Available Data

There are no Gilead studies evaluating the coadministration of SOF/VEL and primidone.

Additionally, a literature search was conducted in Ovid MEDLINE and Embase databases for studies published between 1946 and March 28, 2025 using search terms that included Epclusa, velpatasvir, sofosbuvir, primidone, and related search terms. No relevant citations were found.

References

1. Enclosed. Gilead Sciences Inc, EPCLUSA® (sofosbuvir and velpatasvir) tablets, for oral use. US Prescribing Information. Foster City, CA.
 2. Bausch Health US, LLC, MYSOLINE® (primidone tablets, USP). US Prescribing Information. Bridgewater, NJ.
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Abbreviations

BCRP=breast cancer
resistance protein
DDI=drug-drug interaction

OATP=organic anion
transporting polypeptide
P-gp=P-glycoprotein
PK=pharmacokinetic(s)

SOF=sofosbuvir
VEL=velpatasvir

Product Label

For the full indication, important safety information, and Boxed Warning(s), please refer to the Epclusa US Prescribing Information available at:

http://www.gilead.com/-/media/files/pdfs/medicines/liver-disease/epclusa/epclusa_pi

Follow-Up

For any additional questions, please contact Gilead Medical Information at:

☎ 1-866-MEDI-GSI (1-866-633-4474) or 🌐 www.askgileadmedical.com

Adverse Event Reporting

Please report all adverse events to:

Gilead Pharmacovigilance and Epidemiology ☎ 1-800-445-3235, option 3 or

🌐 www.gilead.com/utility/contact/report-an-adverse-event

FDA MedWatch Program by ☎ 1-800-FDA-1088 or ✉ MedWatch, FDA, 5600 Fishers Ln, Rockville, MD 20852 or 🌐 www.accessdata.fda.gov/scripts/medwatch

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