GILEAD | Medical Information

Epclusa® (sofosbuvir/velpatasvir) Coadministration with Naltrexone

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

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 been conducted between the single-tablet regimen SOF/VEL and naltrexone. Based on the PK profile of each active ingredient within SOF/VEL and naltrexone, a PK interaction would not be predicted.[1,2] For more information about naltrexone, please refer to its product labeling.[2]

SOF/VEL PK[1]

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 Information[1]

Based on drug interaction studies conducted with the components of SOF/VEL, no clinically significant drug interactions have been observed or are expected with naltrexone.

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. For more information, please refer to Sections 7.3 and 7.4 of the SOF/VEL US Prescribing Information (Established and

Potentially Significant Drug Interactions and Drugs without Clinically Significant Interactions with EPCLUSA).

Clinical Trial in People who Inject Drugs (PWID), Including Those on Medication-Assisted Treatment (MAT) for Opioid Use Disorder[1]

SIMPLIFY was an open-label Phase 2 clinical trial that evaluated 12 weeks of treatment with SOF/VEL in 103 HCV-infected PWID (defined as self-reported injection drug use within previous 6 months), including 58 subjects on MAT for opioid use disorder. The proportions of subjects with genotype 1, 2, 3, and 4 HCV infection were 35%, 5%, 58%, and 2%, respectively. The median age was 48 years (range: 24 to 67); 71% were male; 89% were White; and 2% were Black. At baseline, 74% and 26% of subjects reported injection drug use or daily injection drug use, respectively, in the past month; 56% had baseline HCV RNA levels at least 800,000 IU/mL; 10% had compensated cirrhosis; and all subjects were naïve to prior exposure with SOF or an HCV NS5A inhibitor. Subjects on MAT for opioid use disorder reported concomitant use of methadone (76%) and buprenorphine naloxone (17%) with SOF/VEL. The overall SVR rate was 94% (97/103). One subject completed EPCLUSA treatment and was re-infected with a phylogenetically different virus; the other 5 subjects who did not achieve SVR12 did not meet virologic failure criteria.

References

1. Enclosed. Gilead Sciences Inc, EPCLUSA® (sofosbuvir and velpatasvir) tablets, for oral use. US Prescribing Information. Foster City, CA.
2. Alkermes, Inc, VIVITROL® (naltrexone for extended-release injectable suspension) for intramuscular use. US Prescribing Information. Waltham, MA

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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