Acid Reducing Agent Treatment Selector

<table>
<thead>
<tr>
<th>DAA/PC</th>
<th>DCV</th>
<th>EBR/GZR</th>
<th>GLP/PIB</th>
<th>LED/SOF</th>
<th>OBV/PTV/r</th>
<th>OBV/PTV/r +DSV</th>
<th>RDV</th>
<th>SOF</th>
<th>SOF-VEL</th>
<th>SOF-VEL/VOX</th>
</tr>
</thead>
<tbody>
<tr>
<td>Aluminium hydroxide</td>
<td>←→</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>←→</td>
<td>←→</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Antacids</td>
<td>←→</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>←→</td>
<td>←→</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Cimetidine</td>
<td>↓</td>
<td>↑</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Famotidine</td>
<td>↓ 18%</td>
<td>↑</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Ranitidine</td>
<td>←→</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>←→</td>
<td>←→</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Esomeprazole</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Lansoprazole</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Omeprazole</td>
<td>↓ 16%</td>
<td>←→</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Pantoprazole</td>
<td>←→</td>
<td>↑</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>↓</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
<tr>
<td>Rabeprazole</td>
<td>←→</td>
<td>←→</td>
<td>←→</td>
<td>↓</td>
<td>←→</td>
<td>←→</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
<td>➔</td>
</tr>
</tbody>
</table>

**Colour Legend**
- No clinically significant interaction expected.
- These drugs should not be coadministered.
- Potential interaction which may require a dosage adjustment or close monitoring.
- Potential interaction predicted to be of weak intensity.

**Text Legend**
- ↑ Potential increased exposure of the acid reducing agent
- ↓ Potential decreased exposure of the acid reducing agent
- ↔ No significant effect

Numbers refer to increased or decreased AUC as observed in drug-drug interaction studies.

H2 RA  H2 receptor antagonists
PPI  Proton pump inhibitors

- a It is recommended to separate administration of the acid reducing agent and the DAA by 4 hours.
- b Consider separating administration of the acid reducing agent and sofosbuvir by 2 hours.
- c Elbasvir AUC increased by 5%; grazoprevir AUC increased by 10%.
- d H2 receptor antagonists at a dose that does not exceed doses comparable to famotidine 40 mg twice daily can be given simultaneously with or 12 hours apart from the DAA.
- e It is recommended that H2-receptor antagonists should be administered 4 hours after ravidasvir.
- f Simultaneous administration increased ledipasvir and sofosbuvir AUCs both by 11%; administration 12 h apart decreased ledipasvir and sofosbuvir AUCs by 2% and 5%, respectively. Famotidine may be administered simultaneously with or 12 hours apart from ledipasvir/sofosbuvir at a dose that does not exceed 40 mg twice daily. Simultaneous administration of the DAA decreased sofosbuvir and velpatasvir AUCs by 18% and 19%, respectively; administration 12 h apart decreased sofosbuvir and velpatasvir AUCs by 20% and 15%, respectively. Famotidine may be administered simultaneously with or 12 hours apart from sofosbuvir/velpatasvir at a dose that does not exceed 40 mg twice daily.
- g Famotidine at a dose that does not exceed 40 mg twice daily can be given simultaneously with or 12 hours apart from sofosbuvir/velpatasvir/voxilaprevir.
- h For omeprazole, the European SmPC for glecaprevir/pibrentasvir indicates that no dose adjustment is required and the US Prescribing Information indicates no clinically significant interaction and no dose adjustment required. However, it is important to note that currently there are no data with doses of omeprazole greater than 40 mg once daily.
- j Proton pump inhibitor doses comparable to omeprazole 20 mg can be administered simultaneously with ledipasvir/sofosbuvir. Proton pump inhibitors should not be taken before ledipasvir/sofosbuvir.
- k A weak potential interaction is feasible, as an increase in gastric pH may decrease ravidasvir concentrations.
- l If use of a proton pump inhibitor is considered medically necessary, the European Summary of Product Characteristics states that sofosbuvir/velpatasvir could be administered with food and taken 4 hours before a proton pump inhibitor at a dose not to exceed that comparable to omeprazole 20 mg. However, the US Prescribing Information recommends sofosbuvir/velpatasvir to be administered with food and taken 4 hours before omeprazole 20 mg but does not recommend the use of other proton pump inhibitors.
- m If use of a proton pump inhibitor is considered medically necessary, the European Summary of Product Characteristics states that sofosbuvir/velpatasvir/voxilaprevir could be administered with a proton pump inhibitor at a dose not to exceed that comparable to omeprazole 20 mg. However, the US Prescribing Information recommends sofosbuvir/velpatasvir/voxilaprevir to be administered with food and taken 4 hours before omeprazole 20 mg but does not recommend the use of other proton pump inhibitors.
- n Coadministration of omeprazole (20 mg) decreased glecaprevir AUC by 29%; coadministration of omeprazole (40 mg) decreased glecaprevir AUC by ~50% but had no effect on pibrentasvir AUC. The European SmPC for glecaprevir/pibrentasvir indicates that no dose adjustment is required and the US Prescribing Information indicates no clinically significant interaction and no dose adjustment required. However, it is important to note that currently there are no data with doses of omeprazole greater than 40 mg once daily.
- o Simultaneous administration decreased ledipasvir AUC by 4% and had no effect on sofosbuvir; administration 2 h prior to ledipasvir decreased ledipasvir AUC by 42%. Omeprazole 20 mg can be administered simultaneously with ledipasvir/sofosbuvir but should not be taken before ledipasvir/sofosbuvir.
- p Simultaneous administration decreased sofosbuvir and velpatasvir AUC by 29% and 26%; administration 4 h after sofosbuvir/velpatasvir increased sofosbuvir AUC by 5% but decreased velpatasvir AUC by 26%. Coadministration of omeprazole is not recommended. If it is considered medically necessary to coadminister, sofosbuvir/velpatasvir should be administered with food and taken 4 hours before omeprazole 20 mg.
- q Administration 2 h prior to sofosbuvir/velpatasvir/voxilaprevir decreased sofosbuvir, velpatasvir and voxilaprevir AUC by 27%, 54% and 20%, respectively; administration 4 h after sofosbuvir/velpatasvir/voxilaprevir decreased sofosbuvir, velpatasvir and voxilaprevir AUC by 16%, 51% and 5%, respectively. If use of a proton pump inhibitor is considered medically necessary, the European Summary of Product Characteristics states that sofosbuvir/velpatasvir/voxilaprevir could be administered with omeprazole at a dose not to exceed 20 mg. The US Prescribing Information recommends sofosbuvir/velpatasvir/voxilaprevir to be administered with omeprazole 20 mg but does not recommend the use of other proton pump inhibitors.
- r Elbasvir AUC increased by 5%; grazoprevir AUC increased by 12%.

Abbreviations:  DCV  Daclatasvir  EBR/GZR  Elbasvir/Grazoprevir  GLP/PIB  Glecaprevir/Pibrentasvir  LED  Ledipasvir  OBV/PTV/r  Ombitasvir/Paritaprevir/Ritonavir  +DSV  Dasabuvir  PTV  Posaconazole  VEL  Velpatasvir  VOX  Voxilaprevir

© Liverpool Drug Interactions Group, University of Liverpool, 3rd Floor William Henry Duncan Building, 6 West Derby Street, Liverpool, L7 8TX

We aim to ensure that information is accurate and consistent with current knowledge and practice. However, the University of Liverpool and its servants or agents shall not be responsible or in any way liable for the continued currency of information in this publication whether arising from negligence or otherwise however or for any consequences arising therefrom. The University of Liverpool expressly excludes liability for errors, omissions or inaccuracies to the fullest extent permitted by law.