BVHG/BASL/BSG/BHIVA/BIA/CVN Guidelines for management of chronic HCV infection

Headline Recommendations

- 1. We recommend that NHSE considers commissioning pan-genotypic regimens for use in the community for patients who are treatment naïve and do not have cirrhosis to avoid the need for genotyping and facilitate rapid access to care.
- 2. We recommend that ribavirin be avoided whenever possible.
- 3. We recommend that 8 week regimens without ribavirin are first choice for treatment naïve non-cirrhotic patients treated in community or prison settings regardless of genotype.
- 4. We reiterate that transplantation is not contra-indicated in patients with HCV even in the presence of 'difficult' drug resistant mutations.
- 5 Drug-drug interactions should continue to be assessed and therapy should take account of potential interactions.

Genotype Specific Recommendations

Non-cirrhotic

G1a

Sofosbuvir/ledipasvir 8 weeks (treatment naïve) or 12 weeks (treatment experienced)

Grazoprevir/elbasvir 12 weeks OR 16 weeks + ribavirin for patients with viral load >800,000 and resistance associated substitutions(16 weeks + ribavirin is NOT a preferred regimen)

Paritaprevir/ritonavir/ombitasvir+dasabuvir+ribavirin 12 weeks – should be discarded when Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir 12 weeks

Sofosbuvir/velpatasvir/voxilaprevir - 8 weeks

Glecaprevir/pibrentasvir - 8 weeks

G₁b

Sofosbuvir/ledipasvir 8 weeks (treatment naïve) or 12 weeks (treatment experienced)

Grazoprevir/elbasvir 12 weeks

Paritaprevir/ritonavir/ombitasvir+dasabuvir 12 weeks- should be discarded when Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir 12 weeks

Sofosbuvir/velpatasvir/voxilaprevir 8 weeks

Glecaprevir/pibrentasvir 8 weeks

Compensated cirrhosis

G1a

Sofosbuvir/ledipasvir 12 weeks

Grazoprevir/elbasvir 12 weeks OR 16 weeks + ribavirin for patients with viral load >800,000 and resistance associated substitutions (16 weeks + ribavirin is NOT a preferred regimen)

Paritaprevir/ritonavir/ombitasvir+dasabuvir+ribavirin 12-24 weeks - should be discarded when Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir 12 weeks

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks

Glecaprevir/pibrentasvir 12 weeks

G₁b

Sofosbuvir/ledipasvir 12 weeks

Grazoprevir/elbasvir 12 weeks

Paritaprevir/ritonavir/ombitasvir+dasabuvir 12 weeks - should be discarded when Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir 12 weeks

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks

Glecaprevir/pibrentasvir 12 weeks

Decompensated cirrhosis G1a &1b

Sofosbuvir/ledipasvir +/- ribavirin 12 weeks Sofosbuvir/velpatasvir + ribavirin 12 weeks

Re-treatment for DAA failures

Requires pre-treatment virological sequencing to identify resistance associated variants whose presence/absence should be used to guide treatment decisions.

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks Glecaprevir/pibrentasvir 12 weeks (no prior NS5A) or 16 weeks (prior NS5A)

Decompensated cirrhosis – retreatment requires Sof/vel +/- riba 24 weeks

Non cirrhotic

Strongly recommend that IFN is removed and ribavirin free regimens are preferred.
Sof/Vel 12 weeks
Sof/Vel/Vox 8 weeks
Glecaprevir/pibrentasvir 8 weeks

G2

Cirrhosis

Sof/Vel 12 weeks Sof/vel/vox 12 weeks Glecaprevir/pibrentasvir 12 weeks **Decompensated cirrhosis** Sof/vel +/- riba 12 weeks

Re-treatment of DAA failures

Requires pre-treatment virological sequencing to identify resistance associated variants whose presence/absence should be used to guide treatment decisions.

Sofosbuvir/velpatasvir/voxilaprevir **12 weeks** Glecaprevir/pibrentasvir 16 weeks

G3

Non cirrhotic

Sof/Vel 12 weeks Sof/Vel/Vox 8 weeks Glecaprevir/pibrentasvir 8 weeks

Cirrhotic

Sof/Vel 12 weeks Sof/Vel/Vox 12 weeks Glecaprevir/pibrentasvir 16 weeks

Decompensated cirrhosis

12 weeks sofosbuvir/velpatasvir +ribavirin. Consideration should be given to the use of sof/vel for 24 weeks in patients deemed unlikely to respond or intolerant of ribavirin.

Re-treatment for DAA failures

Requires pre-treatment virological sequencing to identify resistance associated variants whose presence/absence should be used to guide treatment decisions.

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks

Decompensated cirrhosis – retreatment requires Sof/vel +/- riba 24 weeks

G4

Non Cirrhotic

Given the paucity of data and the availability of better-validated regimens we recommend that the use of sofosbuvir/ledipasvir for patients with Genotype 4 HCV should be discontinued.

Grazoprevir/elbasvir 12

Paritaprevir/ritonavir/ombitasvir 12 weeks - should be discarded when

Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir/voxilaprevir 8 weeks

Sofosbuvir/velpatasvir 12 weeks

Glecaprevir/pibrentasvir 8 weeks

Cirrhosis

Grazoprevir/elbasvir 12 OR 16 weeks

Sofosbuvir/velpatasvir 12 weeks

Paritaprevir/ritonavir/ombitasvir 12 weeks - should be discarded when

Glecaprevir/pibrentasvir is available.

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks

Glecaprevir/pibrentasvir 12 weeks

Decompensated cirrhosis

12 weeks sofosbuvir/velpatasvir +ribavirin.

Re-treatment for DAA failures

Requires pre-treatment virological sequencing to identify resistance associated variants whose presence/absence should be used to guide treatment decisions.

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks Glecaprevir/pibrentasvir 16 weeks

G5/6

The small number of patients G5/6 infection in trials reported to date was noted.

Non cirrhotic

Sof/Vel 12 weeks Glecaprevir/pibrentasvir 8-12weeks Sof/Vel/Vox 8 weeks

Cirrhotic

Sofosbuvir/velpatasvir 12 weeks Glecaprevir/pibrentasvir 12 weeks Sof/vel/vox 8 weeks

Decompensated cirrhosis

12 weeks sofosbuvir/velpatasvir +ribavirin.

Re-treatment for DAA failures

Requires pre-treatment virological sequencing to identify resistance associated variants whose presence/absence should be used to guide treatment decisions.

Sofosbuvir/velpatasvir/voxilaprevir 12 weeks Glecaprevir/pibrentasvir 16 weeks (note that in patients with both NS5A and NS3 resistance associated variants this regimen is likely to be inadequate)

Special Patient Categories

Patients with renal impairment

We recommend treatment as above but recommend that sofosbuvir be avoided in patients with GFR <45 ml/min.

HIV-hepatitis C coinfection

We recommend that patients with HIV-hepatitis C coinfection are treated for chronic hepatitis C with the same DAA-based treatment regimens as patients with hepatitis C mono-infection, although consideration of drug-drug interactions between DAAs and antiretrovirals should be taken into account.

We recommend that where HIV therapy cannot be switched to avoid drug-drug interactions, an appropriate alternate DAA-based regimen is identified.

Acute hepatitis C infection

We note emerging data shows public health benefits with early DAA therapy for patients with acute HCV who are at high risk of transmission. We recognize that pegylated interferon and ribavirin (the only current treatment option) is unlikely to be acceptable to patients and we therefore recommend that DAA-based treatment is made available for the treatment of acute and early hepatitis C infection, replacing pegylated-interferon +/- ribavirin 24 to 48 weeks

Re-infection following successful DAA-based hepatitis C treatment

We recommend that DAA-based treatment is made available for the treatment of hepatitis C re-infection following successful DAA-based hepatitis C treatment.

Solid Organ Transplantation

HCV infection acquired from a donor organ can be readily treated with currently available drug regimens. We recommend that patients without HCV infection should be offered an opportunity to receive an organ infected with HCV and we recommend that such recipients are offered antiviral therapy as soon as practicable post transplantation; with usual practice being to initiate treatment within the first month.