

Chemotherapy Protocol
HAEMATOLOGY – HSCT ALLOGRAFT

RITUXIMAB 100mg

(Weekly)

Regimen

- HSCT Allograft - Rituximab (7 day-100mg)

Indications

- EBV reactivation, PTLD and immune mediated cytopenias

Toxicity

Drug	Adverse Effect
Rituxumab	Severe cytokine release syndrome, increased incidence of infective complications, progressive multifocal leukoencephalopathy

The adverse effects listed are not exhaustive. Please refer to the relevant Summary of Product Characteristics for full details.

Monitoring

Drugs

- FBC, LFTs and U&Es prior to day one of treatment
- Check hepatitis B status before starting treatment with rituximab

Dose Modifications

The dose modifications listed are for haematological, liver and renal function and drug specific toxicities only. Dose adjustments may be necessary for other toxicities as well.

In principle all dose reductions due to adverse drug reactions should not be re-escalated in subsequent cycles without consultant approval. It is also a general rule for chemotherapy that if a third dose reduction is necessary treatment should be stopped.

Please discuss all dose reductions / delays with the relevant consultant before prescribing, if appropriate. The approach may be different depending on the clinical circumstances.

Haematological

Rituximab is not considered myelotoxic therefore dose adjustment for haematological parameters is not required.

Hepatic Impairment

Drug	Bilirubin μmol/L		AST/ALT units/L	Dose (% of original dose)
Rituximab	N/A		N/A	No dose adjustment needed

Renal Impairment

Drug	Creatinine Clearance (ml/min)	Dose (% of original dose)
Rituximab	N/A	No dose adjustment needed

Other

Dose reductions or interruptions in therapy are not necessary for those toxicities that are considered unlikely to be serious or life threatening. For example, alopecia, altered taste or nail changes.

For all other non-haematological NCI-CTC grade 3 and above toxicities delay treatment until the adverse effect has resolved to NCI-CTC grade 1 or below. The dose should then be reduced to 75% of the original dose or discontinued as appropriate.

Rituximab

Infusion related adverse reactions have been observed in 10% of patients treated with rituximab.

Rituximab administration is associated with the onset of cytokine release syndrome. This condition is characterised by severe dyspnoea, often accompanied by bronchospasm and hypoxia, in addition to fever, chills, rigors, urticaria, and angioedema. It may be associated with some features of tumour lysis syndrome such as hyperuricaemia, hyperkalaemia, hypocalcaemia, acute renal failure, elevated lactate dehydrogenase (LDH) and can lead to acute respiratory failure and death. This effect on the lungs may be accompanied by events such as pulmonary interstitial infiltration or oedema, visible on a chest x-ray.

Cytokine release syndrome frequently occurs within one or two hours of initiating the first infusion.

Hypersensitivity reactions, including anaphylaxis, have been reported following the intravenous administration of proteins. In contrast to cytokine release syndrome, true hypersensitivity reactions typically occur within minutes of starting the infusion. Medicinal products for the treatment of allergic reactions should be available for immediate use in the event of hypersensitivity developing during the administration of rituximab.

Use of rituximab maybe associated with an increased risk of progressive multifocal leukoencephalopathy (PML). Patients must be monitored at regular intervals for any new or worsening neurological, cognitive or psychiatric symptoms that may be suggestive of PML. If PML is suspected, further dosing must be suspended until PML has been excluded. If PML is confirmed the rituximab must be permanently discontinued.

The presence of a viral upper respiratory tract infection at the time of treatment may increase the risk of rituximab associated hepatotoxicity. Patients should be assessed for any cold or flu-like symptoms prior to treatment

Regimen

7 day cycle for 6 cycles

Drug	Dose	Days	Administration
Rituximab	100mg	1	Intravenous infusion in 50ml sodium chloride 0.9% over 120 minutes for the first infusion. If this is well tolerated subsequent infusions may be given over 30-60 minutes

Dose Information

- No dose banding or rounding is necessary as this is a flat dose of rituximab.

Administration Information

Extravasation

- Rituximab - neutral

Other

- The rate of administration of rituximab varies. The first infusion should be given over 120 minutes. If this is well tolerated subsequent infusions may be given over 30-60 minutes.

Additional Therapy

- Rituximab pre-medication
 - 30 minutes prior to rituximab
 - chlorphenamine 10mg intravenous
 - paracetamol 1000mg oral
- Rituximab infusion reactions
 - hydrocortisone 100mg intravenous when required for rituximab infusion related reactions. Please discuss with a haematologist before administering.
 - salbutamol 2.5mg nebule when required for rituximab related bronchospasm
 - consider pethidine 12.5mg-25mg intravenous for rituximab related rigors that fail to respond to corticosteroids

Coding

- Procurement – X71.3
- Delivery – X72.2

REGIMEN SUMMARY

Rituximab (7 day-100mg)

Day One

1. Chlorphenamine 10mg intravenous injection
2. Paracetamol 1000mg oral
3. Rituximab 100mg intravenous infusion in 50ml sodium chloride 0.9% over 120 minutes
Administration Instructions
The first infusion (day 1, cycle 1) should be given over 120 minutes. If this is well tolerated subsequent infusions may be given over 30-60 minutes.
4. Hydrocortisone 100mg intravenous once only when required for the relief of rituximab infusion related reactions
Administration Instructions
Please confirm with a haematologist that this may be given prior to administration
5. Salbutamol 2.5mg nebule once only when required for the relief of rituximab related bronchospasm
6. Pethidine 25mg intravenous bolus when required for the relief of rituximab related rigors
Administration Instructions
Pethidine 25mg intravenous can be administered under the supervision of a doctor for the treatment of rituximab induced rigors that are not responsive to corticosteroids

DOCUMENT CONTROL

Version	Date	Amendment	Written By	Approved By
1.1	Feb 2018	Rituximab infusion volume changed to 50ml	Dr Deborah Wright Pharmacist	Harriet Launders Haematology Pharmacist
1	July 2017	None	Harriet Launders Haematology Pharmacist Dr Deborah Wright Pharmacist	Dr Deborah Richardson Consultant Haematologist Dr Kate Hill Associate Specialist Haematology

This chemotherapy protocol has been developed as part of the chemotherapy electronic prescribing project. This was and remains a collaborative project that originated from the former CSCCN. These documents have been approved on behalf of the following Trusts;

University Hospital Southampton NHS Foundation Trust

All actions have been taken to ensure these protocols are correct. However, no responsibility can be taken for errors that occur as a result of following these guidelines.