

Dose Adjusted R-EPOCH Therapy

INDICATIONS FOR USE:

INDICATION	ICD10	Regimen Code	Reimbursement Status
Treatment of patients with CD20 positive diffuse large B-cell Non Hodgkins lymphoma (NHL)	C83	00355a	Hospital

TREATMENT: (Please see treatment table on next page)

The starting dose of the drugs detailed below may be adjusted downward by the prescribing clinician, using their independent medical judgement, to consider each patients individual clinical circumstances.

Treatment is administered every 21 days for up to 6 cycles or until disease progression or unacceptable toxicity develops.

Facilities to treat anaphylaxis MUST be present when the chemotherapy is administered.

ELIGIBILITY:

- Previously untreated high or intermediate risk diffuse large B-cell NHL
- ECOG status 0-2

EXCLUSIONS:

- Hypersensitivity to cyclophosphamide, DOXOrubicin, etoposide, vinCRIStine, riTUXimab or any of the excipients
- A cumulative life-long dose of 450mg/m² of DOXOrubicin should only be exceeded with extreme caution as there is as risk of irreversible congestive heart failure
- Severe liver impairment (etoposide)
- Breast feeding
- Pregnancy

PRESCRIPTIVE AUTHORITY:

The treatment plan must be initiated by a Consultant Medical Oncologist or Consultant Haematologist working in the area of haematological malignancies.

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 1 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

Drug	Dose Level								Administration	Day
	- 2	- 1	Dose Level 1 Initial dose	2	3	4	5	6		
riTUXimab	375mg/m ²	375mg/m ²	375mg/m ²	375mg/m ²	375mg/m ²	375mg/m ²	375m ²	375mg/m ²	IV infusion in 250ml NaCl 0.9% ^a	1
^b Etoposide	50mg/m ²	50mg/m ²	50mg/m ²	60mg/m ²	72mg/m ²	86.4mg/m ²	103.7mg/m ²	124.4mg/m ²	Dilute in 500ml NaCl 0.9% & infuse over 24 hours	1,2,3,4
^{b,c} DOXOrubicin	10mg/m ²	10mg/m ²	10mg/m ²	12mg/m ²	14.4mg/m ²	17.3mg/m ²	20.7mg/m ²	24.8mg/m ²	Concomitantly in 1000ml NaCl 0.9% & infuse IV over 24 hours	1,2,3,4
^{b,d} VinCRiStine	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²	0.4mg/m ²		1,2,3,4
Cyclophosphamide	480mg/m ²	600mg/m ²	750mg/m ²	900mg/m ²	1080mg/m ²	1296mg/m ²	^e 1555mg/m ² (Requires mesna)	^e 1866mg/m ² (Requires mesna)	IV infusion in 250ml NaCl 0.9% over 30 minutes	5
^f G-CSF (Round to nearest whole syringe)	5mcg/kg	5mcg/kg	5mcg/kg	5mcg/kg	5mcg/kg	5mcg/kg	5mcg/kg	5mcg/kg	SC – No sooner than 24 hrs after the end of cyclophosphamide infusion	^g From day 6, until ANC ≥ 10 ⁹ /L
Prednisolone	120mg/m ²	120mg/m ²	120mg/m ²	120mg/m ²	120mg/m ²	120mg/m ²	120mg/m ²	120mg/m ²	PO in two divided doses (i.e. 6am and 12noon)	1-5
^a See Table 1: Guidance for administration of riTUXimab										
^b There are various ways of administering DOXOrubicin, vinCRiStine and etoposide . See local hospital policy recommendations regarding preferred combinations to be administered concomitantly.										
^c Lifetime cumulative dose of DOXOrubicin is 450mg/m ² . In establishing the maximal cumulative dose of an anthracycline, consideration should be given to the risk factors outlined below ⁱ and to the age of the patient.										
^d vinCRiStine is a neurotoxic chemotherapeutic agent. Refer to NCCP Guidance on the Safe Use of Neurotoxic drugs (including Vinca Alkaloids) in the treatment of cancer										
^e At doses of cyclophosphamide above 1500mg/m ² , pre-treatment is required with mesna. Give mesna dose equivalent to 20% of cyclophosphamide dose IV immediately before cyclophosphamide dose (T0) and 40% of the cyclophosphamide dose orally 2 and 6 hours (T2, T6) after the end of the cyclophosphamide infusion.										
^f Pegylated G-CSF should NOT be substituted for standard G-CSF in this regimen.										
<i>Prophylactic Central Nervous System Intrathecal Methotrexate should be administered with this regimen as clinically indicated (Refer to local policy)</i>										
^g Ensure G-CSF is continued until beyond nadir even if WCC is raised (typically beyond Day 10)										

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 2 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

Table 1: Guidance for administration of ritUXimab

<p>The recommended initial rate for infusion is 50 mg/hr; after the first 30 minutes, it can be escalated in 50 mg/hr increments every 30 minutes, to a maximum of 400 mg/hr. Subsequent infusions can be infused at an initial rate of 100 mg/hr, and increased by 100 mg/hr increments at 30 minute intervals, to a maximum of 400 mg/hr. Development of an allergic reaction may require a slower infusion rate. See Hypersensitivity/Infusion reactions under Adverse Effects/Regimen Specific Complications below. Any deviation from the advised infusion rate should be noted in local policies.</p>
<p>Recommended Observation period: Patients should be observed for at least six hours after the start of the first infusion and for two hours after the start of the subsequent infusions for symptoms like fever and chills or other infusion-related symptoms. Any deviation should be noted in local policies</p>
<p>ritUXimab should be diluted to a final concentration of 1-4mg/ml.</p>
<p>Rapid rate infusion scheduleⁱⁱ See NCCP guidance here. If patients did not experience a serious infusion related reaction with their first or subsequent infusions of a dose of ritUXimab administered over the standard infusion schedule, a more rapid infusion can be administered for second and subsequent infusions using the same concentration as in previous infusions. Initiate at a rate of 20% of the total dose for the first 30 minutes and then 80% of the dose for the next 60 minutes (total infusion time of 90 minutes). If the more rapid infusion is tolerated, this infusion schedule can be used when administering subsequent infusions. Patients who have clinically significant cardiovascular disease, including arrhythmias, or previous serious infusion reactions to any prior biologic therapy or to ritUXimab, should not be administered the more rapid infusion.</p>

TESTS:

Baseline tests:

- FBC, renal and liver profile, LDH, Uric acid, SPEP
- ECG
- Cardiac function using MUGA or ECHO (LVEF > 50% to administer doxorubicin) if >65 years or if clinically indicated (e.g. smoking history, hypertension).
- Virology screen - Hepatitis B (HBsAg, HBcoreAb) & C, HIV.

*See Adverse Effects/Regimen Specific Complications re Hepatitis B Reactivation

Regular tests:

- FBC (including ANC) twice weekly during treatment, three days apart
- Renal and liver profile prior to each cycle
- Assessment of peripheral neuropathy status prior to each cycle
- Cardiac function if clinically indicated

Disease monitoring:

Disease monitoring should be in line with the patient’s treatment plan and any other test/s as directed by the supervising Consultant.

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 3 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</p>		

DOSE MODIFICATIONS:

- Any dose modification should be discussed with a Consultant.
- Drug doses for subsequent cycles are based on previous cycle ANC nadir (Please see Table 2).
- Measurement of ANC nadir based on twice weekly FBC only which must be taken at least 3 days apart i.e. Low neutrophil count on 3 consecutive days does NOT require decreasing a level

Haematological:

Table 2: Dose adjustment of R-EPOCH based on previous cycle ANC nadir

FBC result	Action
If Nadir ANC $\geq 0.5 \times 10^9/L$ on all measurements	Increase 1 dose level above last cycle
If Nadir ANC $< 0.5 \times 10^9/L$ on 1 or 2 measurements	Dose at same level as last cycle
If the nadir ANC $< 0.5 \times 10^9/L$ on three separate measurements <u>OR</u> if the nadir platelet count is $< 25 \times 10^9/L$ on one measurement	Decrease to one level below the last cycle

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHSC contributors: Prof Maccon Keane, Dr Cliona Grant	Page 4 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

Renal and Hepatic Impairment:

Table 3: Dose modifications based on renal and hepatic impairment

Drug	Renal Impairment		Hepatic Impairment			
riTUXimab	Probably no dose reduction necessary		Probably no dose reduction necessary			
Etoposide	CrCl(ml/min)	Dose	Bilirubin (micromol/L)		AST (Units)	Dose
	>50	100%	26-51	or	60-180	50%
	15-50	75%				
	<15	50%				
	Subsequent doses should be based on clinical response Dialysis Start at reduced dose and increase according to clinical response.		>51	or	>180	Clinical decision
DOXOrubicin	No dose reduction required. Clinical decision in severe renal impairment		Bilirubin (micromol/L)		Dose	
			20-51		50%	
			51-85		25%	
			>85		Omit	
			If AST 2-3 x normal, give 75% dose. If AST >3x ULN, give 50% dose			
Cyclophosphamide	CrCl (ml/min)	Dose	Severe impairment: Clinical decision.			
	>20	100%				
	10-20	75%				
	<10	50%				
	Clinical decision – consider whether patient is being treated with high dose treatment. Dialysis Renal Drug Handbook -dose at 50- 100% dose (CAPD, HD and HDF/High flux) Do not perform dialysis for 12 hours. Dose at 75-100%dose (CAV/VVHD). If anuric, mesna may not be required.					
vinCRiStine	No dose reduction required		Bilirubin (micromol/L)		AST/ALT (Units)	Dose
			26-51	or	60-180	50%
			>51	and	Normal	50%
			>51	and	>180	Omit

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHSC contributors: Prof Maccon Keane, Dr Cliona Grant	Page 5 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

Neurotoxicity:

Table 4: Dose modification of vinCRISTine based on neurotoxicity (CTCAE v 4.0)

Symptom	Dose of vinCRISTine
Grade 1	100%
Grade 2	Hold until recovery, then reduce dose by 50%
Grade 3,4	Omit

Table 5: Dose modification schedule of riTUXimab based on adverse events

Adverse reactions	Discontinue	Recommended dose modification
Severe infusion related reaction (e.g dyspnoea, bronchospasm, hypotension or hypoxia) First occurrence		Interrupt infusion immediately. Evaluate for cytokine release/tumour lysis syndrome (appropriate laboratory tests) and pulmonary infiltration (chest x-ray). Infusion may be restarted on resolution of all symptoms, normalisation of laboratory values and chest x-ray findings at no more than one-half the previous rate.
Second occurrence	Consider discontinuing treatment	Consider coverage with steroids for those who are not already receiving steroids.
Mild or moderate infusion-related reaction		Reduce rate of infusion. The infusion rate may be increased upon improvement of symptoms

SUPPORTIVE CARE:

EMETOGENIC POTENTIAL:

Level -2, -1, 1, 2, 3, 4 - Moderate (Refer to local policy)
 Level 5, 6 - High (Refer to local policy).

PREMEDICATIONS:

Premedication consisting of an anti-pyretic and an anti-histamine should always be administered before each infusion of riTUXimab.

Table 6: Suggested pre-medications prior to riTUXimab infusion:

Drugs	Dose	Route
Paracetamol	1g	PO 60minutes prior to riTUXimab infusion
Chlorphenamine	10mg	IV bolus 60minutes prior to riTUXimab infusion
Hydrocortisone	100mg	IV bolus 60 minutes prior to riTUXimab infusion

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 6 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

OTHER SUPPORTIVE CARE:

- Tumour lysis syndrome prophylaxis (**Refer to local policy**)
- Proton pump Inhibitor (**Refer to local policy**)
- PJP prophylaxis (**Refer to local policy**)
- Anti-viral prophylaxis (**Refer to local policy**)
- Anti-fungal prophylaxis (**Refer to local policy**)
- Prophylactic regimen against vinCRiStine induced constipation is recommended (**Refer to local policy**)
- Patients should have an increased fluid intake of 2-3 litres on day 5 to prevent haemorrhagic cystitis associated with cyclophosphamide.
- Consider mesna if high dose cyclophosphamide prescribed on dose escalation (>1.5g/m² at levels 5, 6)

ADVERSE EFFECTS / REGIMEN SPECIFIC COMPLICATIONS:

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

Please see NCCP Regimen 00542 riTUXimab 375mg/m² Combination Therapy -21 day for detailed information on the adverse effects/Regimen Specific Complications relating to riTUXimab Therapy

- **Hypersensitivity/Infusion Reactions:** Close monitoring is required throughout the first infusion of riTUXimab. (**Refer to local policy**). riTUXimab can cause allergic type reactions during the IV infusion such as hypotension, wheezing, rash, flushing, pruritis, sneezing, cough, fever or faintness. Monitor infusion of etoposide for the first 15 minutes for signs of hypotension.
- **Hepatitis B Reactivation:** Patients should be tested for both HBsAg and HBcoreAb as per local policy. If either test is positive, such patients should be treated with anti-viral therapy. (**Refer to local infectious disease policy**). These patients should be considered for assessment by hepatology.
- **Cardiac Toxicity:** Patients with a history of cardiac disease and/or cardiotoxic chemotherapy should be monitored closely while on riTUXimab. DOXOrubicin is cardiotoxic and must be used with caution, if at all, in patients with cardiac dysfunction.
- **Neuropathy:** vinCRiStine may cause peripheral neuropathy which is dose related and cumulative, requiring monitoring before each dose is administered. The presence of pre-existing neuropathies or previous treatment with other neurotoxic drugs may increase risk of peripheral neuropathy. Patients with mild peripheral neuropathy can usually continue to receive full doses of vinCRiStine, but when symptoms increase in severity and interfere with neurologic function, dose reduction or discontinuation of the drug may be necessary. The natural history following discontinuation of treatment is gradual improvement, which may take up to several months. A routine prophylactic regimen against constipation is recommended for all patients receiving vinCRiStine sulphate. Paralytic ileus may occur. The ileus will reverse itself upon temporary discontinuance of vinCRiStine and with symptomatic care.
- **Extravasation:** DOXOrubicin and vinCRiStine cause pain and possible tissue necrosis if extravasated (refer to local policy).

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 7 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</p>		

DRUG INTERACTIONS:

- Antihypertensives: Additive effect of hypotension during ritUXimab infusion. Consider withholding antihypertensives 12 hours before and during ritUXimab infusion.
- DOXOrubicin cardiotoxicity is enhanced by previous or concurrent use of other anthracyclines, or other potentially cardiotoxic drugs (e.g. 5-FU, cyclophosphamide or PACLitaxel) or with products affecting cardiac function (e.g. calcium antagonists).
- Current drug interaction databases should be consulted for more information including potential for interactions with CYP3AR inhibitors/ inducers.

REFERENCES:

1. Wilson WH et al. Dose-adjusted EPOCH chemotherapy for untreated large B-cell lymphomas: a pharmacodynamic approach with high efficacy. *Blood* 2002; 99:2685-93.
2. Wilson WH et al. Phase II Study of Dose-Adjusted EPOCH and Rituximab in Untreated Diffuse Large B-Cell Lymphoma with Analysis of Germinal Center and Post-Germinal Center Biomarkers. *J Clin Oncol.* 2008;26:2717-2724
3. Dunleavy K, Pittaluga S et al. Dose-Adjusted EPOCH-Rituximab Therapy in Primary Mediastinal B-Cell Lymphoma *N Engl J Med* 2013;368:1408-16.
4. Dosage Adjustment for Cytotoxics in Renal Impairment January 2009; North London Cancer Network.
5. Dosage Adjustment for Cytotoxics in Hepatic Impairment January 2009; North London Cancer Network.
6. NCCP Classification Document for Systemic Anti-Cancer Therapy (SACT) Induced Nausea and Vomiting. V3 2021. Available at: <https://www.hse.ie/eng/services/list/5/cancer/profinfo/chemoprotocols/nccp-classification-document-for-systemic-anti-cancer-therapy-sact-induced-nausea-and-vomiting.pdf>
7. ritUXimab (MabThera®) Summary of Product Characteristics. Last updated: 10/10/2019. Accessed Dec2019. Available at: https://www.ema.europa.eu/en/documents/product-information/mabthera-epar-product-information_en.pdf
8. Cyclophosphamide (Endoxana®) Injection 500mg Powder for Solution for Injection. Summary of Product Characteristics Last updated: 21/12/2018. Accessed Dec2019 Available at https://www.hpra.ie/img/uploaded/swedocuments/Licence_PA2299-027-001_21122018112107.pdf
9. DOXOrubicin Accord 2mg/ml concentrate for solution for infusion. Summary of Product Characteristics. Last updated: 27/02/2019 Accessed Dec 2019 Available at https://www.hpra.ie/img/uploaded/swedocuments/Licence_PA2315-083-001_25022019110336.pdf
10. Etoposide 20 mg/ml Concentrate for Solution for Infusion Summary of Product Characteristics. Last updated: 29/07/2019. Accessed Dec 2019. Available at https://www.hpra.ie/img/uploaded/swedocuments/Licence_PA2059-036-001_29072019103821.pdf
11. vinCRiStine Summary of Product Characteristics. Last updated: 30/10/2018. Accessed Dec 2019. Available at https://www.hpra.ie/img/uploaded/swedocuments/Licence_PA0822-232-001_30102018164938.pdf

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 8 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		

Version	Date	Amendment	Approved By
1	18/12/2017		Dr Cliona Grant, Prof Maccon Keane
2	16/10/2020	Reviewed. Standardisation of treatment table and pre-medications. Updated HepB reactivation wording. Updated recommendation for hepatic impairment	Dr Cliona Grant, Prof Maccon Keane
3	15/09/2021	Amended footnote in treatment table in relation to pegylated G-CSF.	Dr Cliona Grant, Prof Maccon Keane

Comments and feedback welcome at oncologydrugs@cancercontrol.ie.

ⁱ Cardiotoxicity is a risk associated with anthracycline therapy that may be manifested by early (acute) or late (delayed) effects.

Risk factors for developing anthracycline-induced cardiotoxicity include:

- high cumulative dose, previous therapy with other anthracyclines or anthracenediones
- prior or concomitant radiotherapy to the mediastinal/pericardial area
- pre-existing heart disease
- concomitant use of other potentially cardiotoxic drugs

In establishing the maximal cumulative dose of an anthracycline, consideration should be given to the risk factors above and to the age of the patient

ⁱⁱThe rapid infusion is an unlicensed means of administration of rituximab for the indications described above, in Ireland. Patients should be informed of this and consented to treatment in line with the hospital's policy on the use of unlicensed medication and unlicensed or "off label" indications. Prescribers should be fully aware of their responsibility in communicating any relevant information to the patient and also ensuring that the unlicensed or "off label" means of administration has been acknowledged by the hospital's Drugs and Therapeutics Committee, or equivalent, in line with hospital policy.

NCCP Regimen: Dose Adjusted R-EPOCH Therapy	Published: 18/12/2017 Review: 16/10/2025	Version number: 3
Tumour Group: Lymphoma NCCP Regimen Code: 00355	ISMO/IHS Contributors: Prof Maccon Keane, Dr Cliona Grant	Page 9 of 9
<p>The information contained in this document is a statement of consensus of NCCP and ISMO or IHS professionals regarding their views of currently accepted approaches to treatment. Any clinician seeking to apply or consult these documents is expected to use independent medical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. Use of these documents is the responsibility of the prescribing clinician and is subject to HSE's terms of use available at http://www.hse.ie/eng/Disclaimer</p> <p><i>This information is valid only on the day of printing, for any updates please check www.hse.ie/NCCPchemoregimens</i></p>		