Regimen Reference Order – LYMP – fludarabine (oral) + riTUXimab

ARIA: LYMP – [R-fludarabine (po)]

Planned Course: Every 28 days for 6 cycles
Indication for Use: Non-Hodgkin Lymphoma

CVAD: At Provider's Discretion

Proceed with treatment if:

- ***ANC equal to or greater than 1 x 10^9/L AND Platelets equal to or greater than 75 x 10^9/L***
- ***Creatinine clearance greater than 30 mL/minute***
  - ❖ **Contact Hematologist if parameters not met**

Pre-treatment Requirements

Drug	Dose	CCMB Administration Guideline
allopurinol*	300 mg	Orally once daily for 10 days to begin 3 days prior to Cycle 1 and at provider's discretion for subsequent cycles **(Self-administered at home)** *Only patients at risk of tumor lysis syndrome will be prescribed allopurinol

Treatment Regimen – LYMP – fludarabine (oral) + riTUXimab

Establish primary solution 500 mL of: normal saline

Drug	Dose	CCMB Administration Guideline
Cycle 1		
Day 1		
acetaminophen	650 mg	Orally 30 minutes prior to riTUXimab
dexamethasone	20 mg	IV in normal saline 50 mL over 15 minutes
diphenhydrAMINE	50 mg	IV in normal saline 50 mL over 15 minutes
riTUXimab (IV brand name specific)	375 mg/m^2	**Slow infusion (if greater than 6 months since last riTUXimab dose or no previous riTUXimab):** IV made up to a final concentration of 1 mg/mL in normal saline. Start at 50 mg/hr for 60 minutes and escalate infusion rate in 50 mg/hr increments every 30 minutes to a maximum of 400 mg/hr *Nursing Alert: IV tubing is primed with riTUXimab* *Alert: Ensure brand name on prescription label (indicated in brackets on prescription label) matches prescribed order
		OR

		<u>Slow infusion</u> (if equal to or less than 6 months since last riTUXimab dose): IV made up to a final concentration of 1 mg/mL in normal saline. Start at 100 mg/hr for 30 minutes and escalate infusion rate in 100 mg/hr increments every 30 minutes to a maximum of 400 mg/hr **Nursing Alert: IV tubing is primed with riTUXimab* **Alert: Ensure brand name on prescription label (indicated in brackets on prescription label) matches prescribed order*
fludarabine	40 mg/m^2 (rounded to nearest 10 mg)	Orally once with or without food Swallow whole **(Self-administered at home)**
Days 2 to 5		
fludarabine	40 mg/m^2 (rounded to nearest 10 mg)	Orally once daily with or without food Swallow whole **(Self-administered at home)**
Cycle 2 and Onwards		
Day 1		
acetaminophen	650 mg	Orally 30 minutes prior to riTUXimab
dexamethasone	12 mg	Orally 30 minutes prior to riTUXimab
diphenhydrAMINE	50 mg	Orally 30 minutes prior to riTUXimab
riTUXimab (Subcutaneous)	1400 mg (1400 mg = 11.7 mL)	**<u>Subcutaneous</u>:** Administer over 5 minutes into abdomen Syringe should be held in hand for 5 minutes to warm up and decrease viscosity Use 25G needle **Nursing Alert: Ensure subcutaneous riTUXimab formulation is used (riTUXimab-hyaluronidase, human)*
	OR	
riTUXimab (IV brand name specific)	375 mg/m^2	**<u>Rapid infusion</u>**: IV in normal saline over 90 minutes: Infuse 50 mL of a 250 mL bag (or 100 mL of a 500 mL bag) over 30 minutes, then infuse the remaining 200 mL (or 400 mL of a 500 mL bag) over 60 minutes **Alert: Ensure brand name on prescription label (indicated in brackets on prescription label) matches prescribed order* *Concentration dependent drug. Pharmacy will adjust diluent volume to ensure drug stability*
		OR
		<u>Slow infusion</u>: IV made up to a final concentration of 1 mg/mL in normal saline. Start at 100 mg/hr for 30 minutes and escalate infusion rate in 100 mg/hr increments every 30 minutes to a maximum of 400 mg/hr **Nursing Alert: IV tubing is primed with riTUXimab* **Alert: Ensure brand name on prescription label (indicated in brackets on prescription label) matches prescribed order*

fludarabine	40 mg/m^2 (rounded to nearest 10 mg)	Orally once with or without food Swallow whole **(Self-administered at home)**
Days 2 to 5		
fludarabine	40 mg/m^2 (rounded to nearest 10 mg)	Orally once daily with or without food Swallow whole **(Self-administered at home)**
All doses will be automatically rounded that fall within the DSG Approved Dose Bands. See LYMP DSG – Dose Banding document for more information		
fludarabine (Fludara®) available dosage strength: 10 mg tablets **Classification: Cytotoxic, Hazardous**		

Flush after each medication:

- 50 mL over 6 minutes (500 mL/hr)

In the event of an infusion-related hypersensitivity reaction, refer to the '**Hypersensitivity Reaction Standing Order**'

REQUIRED MONITORING

All Cycles

Day 1

- CBC, serum creatinine, urea, electrolytes, liver enzymes, LDH, total bilirubin, uric acid and albumin as per Physician Orders

INTRAVENOUS riTUXimab

- Full vital signs (temperature, heart rate, respiratory rate, blood pressure and O_2 saturation) prior to each dose and as clinically indicated
- No observation period is required. Patient can be discharged from treatment room if stable whether they had a reaction or not

SUBCUTANEOUS riTUXimab

- Full vital signs (temperature, heart rate, respiratory rate, blood pressure and O_2 saturation) prior to each dose, at discharge and as clinically indicated
- **15 minute observation period required after each dose**

Recommended Support Medications

Drug	Dose	CCMB Administration Guideline
valACYclovir	500 mg	Orally once daily
sulfamethoxazole-trimethoprim	800/160 mg	Orally twice daily on Saturdays and Sundays only
metoclopramide	10 – 20 mg	Orally every 4 hours as needed for nausea and vomiting

DISCHARGE INSTRUCTIONS

- Patients should be instructed to contact their cancer team immediately if symptoms of hypersensitivity reactions occur after discharge
- Instruct patient to continue taking anti-emetic(s) and support medications at home
- Treatment room nurse to provide oral fludarabine on Day 1 of each cycle. fludarabine is a cancer therapy in this treatment regimen. Remind patient to take fludarabine at home
- Reinforce applicable safe handling precautions of medications, blood and body fluids for 48 hours after completion of chemotherapy

ADDITIONAL INFORMATION

- Administering nurse must document any infusion-related reactions with any dose of riTUXimab
- Ensure there were **no Grade 3 or 4** infusion-related reactions with any previous dose prior to administering riTUXimab via subcutaneous injection or rapid infusion
- Patients on fludarabine should receive irradiated blood products
- valACYlovir (shingles prophylaxis) and sulfamethoxazole-trimethoprim (*Pneumocystis jirovecii* pneumonia prophylaxis) continue while on treatment and for 6 months after discontinuation of treatment due to risk of prolonged immunosuppression
- Intravenous riTUXimab formulation is available from more than one manufacturer and uses several different brand names. Brand name will be indicated in brackets after riTUXimab. **Ensure prescription label matches the brand name on prescribed order for intravenous riTUXimab**
- Oral fludarabine is dispensed by the pharmacy site that prepares the riTUXimab