

Note: These patients should NOT be on electrolyte replacement protocols. Use of sodium bicarbonate for alkalinization of urine is currently not recommended for prevention and treatment of TLS.

¹ See Appendix A for stratification based on disease type

² If calcium-phosphorus product $\geq 50 \text{ mg}^2/dL^2$, ensure hydration is maintained and alkalinization is discontinued. Consider consulting Nephrology service, especially if the calcium-phosphorus product continues to rise $\geq 60 \text{ mg}^2/dL^2$. ³ Blood specimens for uric acid levels should kept on ice after collection and prior to testing, and processed immediately

⁴Allopurinol dose needs to be adjusted in renal failure. Maximum daily dose of allopurinol is 800 mg/day. Dose adjustments may be necessary if allopurinol is used with other drugs (e.g., 6- mercaptopurine, azathioprine,

cyclophosphamide, thiazide and loop diuretics, and warfarin) - Refer to MD Anderson Formulary for a complete list of interactions. Allopurinol should be initiated 24-48 hours prior to chemotherapy when possible.

⁵ Rasburicase must be given 4 hours prior to chemotherapy. For adult patients, rasburicase is to be given at a fixed dose of 3 mg per institutional formulary restrictions; repeat doses are permitted if patient meets restrictions based on repeat lab values prior to each dose.

⁶Rasburicase is contraindicated in glucose-6 phosphate dehydrogenase deficient patients, known hypersensitivity reactions, hemolytic anemia or methemoglobinemia. Allopurinol should be substituted in these patients.

⁷ Patients with established TLS or high risk and/or renal insufficiency should be closely monitored and have access to Nephrology service and Intensive Care Unit (ICU) in the event that dialysis is required

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APPENDIX A: Risk Assessment Based on Disease Type¹

LOW RISK	INTERMEDIATE RISK	HIGH RISK
 Leukemia CLL receiving only alkylating agents CML (excluding blast crisis) Lymphoma Anaplastic large-cell lymphoma DLBCL with LDH within normal limits (WNL) Mantle cell lymphoma (blastoid variants) with LDH WNL 	 Leukemia AML with WBC < 25 K/microliter CLL receiving targeted and/or biological therapies Lymphoma DLBCL with LDH greater than upper limit of normal (non-bulky²) Mantle cell lymphoma (blastoid variants) with LDH greater 	 Leukemia ALL AML with WBC ≥ 25 K/microliter Burkitt's leukemia CML-BC CLL treated with venetoclax and ALC ≥ 25 K/microliter or bulky lymph nodes
 Peripheral T-cell lymphoma with LDH WNL T-cell lymphoma with LDH WNL Transformed lymphoma with LDH WNL Cutaneous T-cell lymphoma Follicular lymphoma Hodgkin lymphoma Mantle cell lymphoma (non-blastoid variants) Marginal zone B-cell lymphoma Small lymphocytic lymphoma 	 than upper limit of normal (non-bulky²) Peripheral T-cell lymphoma with LDH greater than upper limit of normal (non-bulky²) T-cell lymphoma with LDH greater than upper limit of normal (non-bulky²) Transformed lymphoma with LDH greater than upper limit of normal (non-bulky²) Early stage lymphoblastic lymphoma with LDH less than 2 times upper limit of normal 	 Lymphoma Advanced Stage lymphoblastic lymphoma Burkitt's lymphoma DLBCL with LDH greater than upper limit of normal (bulky²) Mantle cell lymphoma (blastoid variants) with LDH greater than upper limit of normal (bulky²) Peripheral T-cell lymphoma with LDH greater than upper limit of normal (bulky²) Transformed lymphoma with LDH greater than upper
 Other Solid tumors (excluding neuroblastomas, germ-cell tumors, and small cell lung cancer) Multiple myeloma MDS 	Other • Neuroblastoma • Germ-cell tumors • Small cell lung cancer moblastic leukemia AML = acute myeloid leukemia CLL = chronic lymp	 limit of normal (bulky²) Other Myeloma with extramedullary disease and LDH greater than upper limit of normal Myelofibrosis - Intermediate-2 risk or High-risk disease Plasma cell leukemia

¹Renal dysfunction elevates the patient to the next risk level

²Bulky disease is defined as any mass \geq 7.5 cm

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APPENDIX B: Rasburicase Criteria for Use¹

Criteria for Use	Risk Factors
 Uric acid > 7.5 mg/dL plus at least two risk factors <u>or</u> Uric acid ≤ 7.5 mg/dL plus at least three risk factors 	 High risk disease (see Appendix A) Creatinine > 1.3 mg/dL or > 50% increase from baseline WBC > 50 K/microliter Lactate dehydrogenase greater than 2 times the upper limit of normal (ULN)

¹ Criteria based on MD Anderson Formulary Restriction

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APPENDIX C: Suggested Guide for Management of Electrolyte Abnormalities

Abnormality	Management Recommendations	
Hyperphosphatemia		
Moderate (phosphorus ≥ 6 mg/dL)	 Restrict phosphorus intake (avoid IV and PO phosphorus; limit dietary sources) If tolerating oral intake, administer phosphate binder (select one): Sevelamer (Renagel[®], Renvela[®]) 800-1,600 mg PO three times a day with meals Aluminum hydroxide 300-600 mg PO three times a day with meals (avoid with renal dysfunction) 	
Severe	Dialysis may be needed in severe cases	
Hypocalcemia (calcium \leq 7 mg/dL or ionized calcium \leq 0.8 mmol/L)		
Asymptomatic	 No therapy To avoid calcium phosphate precipitation, asymptomatic patients with acute hypocalcemia and hyperphosphatemia should not be given calcium repletion until phosphorous level has normalized 	
Symptomatic	Calcium gluconate 1 gram via slow IV infusion with EKG monitoring	
Uremia (elevated BUN with altered mental status)	 Fluid and electrolyte management Uric acid and phosphate management Adjust doses for renally excreted medications Dialysis 	

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APPENDIX C: Suggested Guide for Management of Electrolyte Abnormalities - continued

Abnormality	Management Recommendations	
Hyperkalemia		
Moderate (potassium 5-6 mmol/L) <u>and</u> asymptomatic	 Restrict potassium intake (avoid IV and PO potassium; limit dietary intake) EKG and cardiac rhythm monitoring If tolerating oral intake, administer potassium binder¹ (select one): Sodium polystyrene sulfonate (Kayexalate[®]) 15-30 grams PO; repeat every 4 or 6 hours depending upon follow-up potassium levels Patiromer sorbitex calcium (Veltassa[®]) - 8.4 grams once daily PO; may increase frequency based on potassium levels; adjust dose at ≥ 1-week intervals in increments of 8.4 grams (maximum dose: 25.2 grams/day) Sodium zirconium cyclosilicate (Lokelma[®])² - 10 grams 3 times daily PO for up to 48 hours; adjust dose by 5 grams daily at 1-week intervals as needed based on potassium (maximum maintenance dose: 15 grams/day) 	
Severe (potassium > 6 mmol/L) <u>and/or</u> symptomatic	 Same as moderate, plus: Concurrent EKG changes: calcium gluconate 1 gram via slow IV infusion; may be repeated after 5-10 minutes if EKG changes persist To temporarily shift potassium intracellularly IV insulin and dextrose 0.1 units/kg of regular insulin IV followed by 25-50 grams of D50W IV based on pre-treatment glucose. Hold D50W if glucose > 250 mg/dL. See IP Hyperkalemia order set. Monitor blood glucose closely Sodium bicarbonate 50 mEq via slow IV infusion Can be used if patient is acidemic (arterial pH < 7.35); however sodium bicarbonate and calcium should not be administered through the same lumen Albuterol 10-20 mg in 4 mL saline via nebulizer over 20 minutes or 10-20 puffs via MDI over 10-20 minutes Avoid in patients with acute coronary disease 	

¹Risk for use includes intestinal necrosis. High risk patient populations for intestinal necrosis include those with post-operative bowel motility disorders or with ileus, small or large bowel obstruction, or ulcerative colitis.

² Avoid sodium zirconium cyclosilicate (Lokelma[®]) in patients with volume overload; may worsen heart failure and/or pulmonary edema

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SUGGESTED READINGS

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