PAIN MITIGATION AND MANAGEMENT STRATEGIES FOR ANTI-GD2 INFUSIONS

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Abstract

Monoclonal antibodies (mAbs) targeting disialoganglioside 2 (GD2) are an important treatment advance for high-risk neuroblastoma, including in patients with refractory or relapsed disease. Dinutuximab and dinutuximab beta are administered for [?]8 hours (and up to 10 days for dinutuximab beta), whereas naxitamab is administered over 0.5 to 2 hours as tolerated. As acute pain is a class effect of anti-GD2 mAbs, effective pain management is crucial to successful treatment. Here, we provide an overview of current pain-management strategies for anti-GD2 mAb infusions, including discussion of opioid analgesics, ketamine, gabapentin, and other similar agents, and non-pharmacologic approaches. Potential future pain management options are also discussed, in addition to the use of sedatives to reduce the anxiety that may be associated with infusion-related pain. Specific guidance for pain management during naxitamab infusions is provided, as these infusions are administered over 0.5 to 2 hours and may not need overnight hospitalization based on the physician's assessment, and require rapid-onset analgesia options suitable for potential outpatient administration.

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Infusion conditions	Anti-GD2		Pre-infusion								Infusion*			
		Daya	-4	-3	-2	-1	0	1						
		Timing	-	-		-	•	2 hours pre- infusion	1 hour pre- infusion	30 minutes pre-infusion	Infusion	1 hour post- infusion	2 hours post- infusion	4 hours post- infusion
Inpatient / > 8 hours infusion duration	Dinutuximab	Gabapentin			10 mg/kg q.d. p.o.	10 mg b.i.d.					10 mgti.d. p.o.			
		Morphine sulfate								0.05 mg/kg i.v.	20–50 mcg/kg/hr PRN i.v.	0.02-0.05 m	g/kg PRN i.v.	
	Dinutuximab beta	Gabapentin			10 mg/kg q.d. p.o.	10 mg b.i.d.			10 mgLid p.o.					
		Morphine sulfate						0.02–0.05 mg/kg bolus infusion			0.03 mg/kg/hri.v.	Continuous infusion in response to the patient's pain perception, wean off over 5 days by decreasing its dosing rate (e.g., to 0.02-0.05 mg/kg/hour, 0.02 mg/kg/hour, 0.01 mg/kg/hour, 0.005 mg/kg/hour)		
												Daily infusion: 0.01 mg/kg/hour for 4 hours post-infusion		
Outpatient / < 2 hours infusion duration	Naxitamab	Gabapentin	5–10 mg/kg q.d. p.o.	5–10 mg/kg b.i.d. p.o.	5-10 mgl (d p.o.									
		Opioids (morphine sulfate, hydromorphone, or oxycodone)							45-60 minutes pre- infusion: 0.1-0.2 mg/kg.p.o. oxycodone (maximum dose 5 mg)	15 minutes before influsion: 0.025-0.1 morphine sulfate OR 0.00375-0.015 mg/kgi.v., hydromorphone	PRN every 15-30 minutes from onset of pain to a maximum of 4 doses: 0.025-0.1 mg/kg I.v., morphine OR 0.00376-0.015 mg/kgI.v., hydromorphone		Consider discharge if vital signs are stable	



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