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Guideline for the Management of *Clostridium Difficile* Infection in Children and Adolescents With Cancer and Pediatric Hematopoietic Stem-Cell Transplantation Recipients

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The "Guideline for the Management of *Clostridium Difficile* Infection in Children and Adolescents with Cancer and Pediatric Hematopoietic Stem-Cell Transplantation Recipients" developed by the Pediatric Oncology Group of Ontario (POGO) was endorsed by the COG Supportive Care Guideline Committee in February 2019.

The source guideline is published (Diorio C, Robinson PD, Ammann R, et al. Guideline for the management of *Clostridium difficile* infection in children and adolescents with cancer and pediatric hematopoietic stem cell transplantation recipients. J Clin Oncol 2018; 36:31, 3162-3171.) and is available at: https://doi.org/10.1200/JCO.18.00407

The purpose of the source guideline is to create a clinical practice guideline for the prevention and treatment of *Clostridium difficile* in children and adolescents with cancer and pediatric HSCT patients. Recommendations from the endorsed clinical practice guideline are presented in the table below.

Summary of Recommendations for the Management of *Clostridium Difficile* Infection (CDI) in Children and Adolescents with Cancer and Pediatric Hematopoietic Stem-Cell Transplantation Recipients

RECOMMENDATIONS	Strength of Recommendation and Quality of Evidence*		
What interventions should be used for the prevention of Clostridium difficile infection (CDI) in			
children and adolescents with cancer and pediatric HSCT patients?			
1. We suggest that probiotics not be used routinely for the	Weak recommendation		
prevention of CDI in children and adolescents with cancer and	Low quality evidence		
pediatric HSCT patients			
What interventions should be used for the treatment of CDI in children and adolescents with cancer			
and pediatric HSCT patients?			
2. Use either oral metronidazole or oral vancomycin for the	Strong recommendation		
treatment of nonsevere CDI in children and adolescents with cancer	Low quality evidence		
and pediatric HSCT patients			
3. Use oral vancomycin for the treatment of severe CDI in children	Strong recommendation		
and adolescents with cancer and pediatric HSCT patients	Low quality evidence		
4. Consider fidaxomicin for the treatment of recurrent CDI in children	Weak recommendation		
and adolescents with cancer and pediatric HSCT patients	Low quality evidence		
5. Do not use fecal microbiota transplantation routinely for the	Strong recommendation		
treatment of CDI in children and adolescents with cancer and	Low quality evidence		
pediatric HSCT patients			
6. We suggest that monoclonal antibodies not be used routinely for	Weak recommendation		
the treatment of CDI in children and adolescents with cancer and	Low quality evidence		
pediatric HSCT patients	, ,		
7. We suggest that probiotics not be used routinely for the treatment	Weak recommendation		
of CDI in children and adolescents with cancer and pediatric HSCT	Low quality evidence		
patients.			

^{*}see Appendix 1

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Appendix 1: Systems for Classifying Recommendations and Evidence used by the Source Clinical Practice Guidelines

I. GRADE

Strength of Recommendations:

Strong Recommendation	When using GRADE, panels make strong recommendations when they are confident that the desirable effects of adherence to a recommendation outweigh the undesirable effects.
Weak Recommendation	Weak recommendations indicate that the desirable effects of adherence to a recommendation probably outweigh the undesirable effects, but the panel is less confident.

Strength of Recommendations Determinants:

Factor	Comment	
Balance between desirable	The larger the difference between the desirable and undesirable	
and undesirable effects	effects, the higher the likelihood that a strong recommendation	
	is warranted. The narrower the gradient, the higher the	
	likelihood that a weak recommendation is warranted	
Quality of evidence	The higher the quality of evidence, the higher the likelihood that	
	a strong recommendation is warranted	
Values and preferences	The more values and preferences vary, or the greater the	
	uncertainty in values and preferences, the higher the likelihood	
	that a weak recommendation is warranted	
Costs (resource allocation)	The higher the costs of an intervention—that is, the greater the	
	resources consumed—the lower the likelihood that a strong	
	recommendation is warranted	

Quality of Evidence

High Quality	Further research is very unlikely to change our confidence in the estimate of effect
Moderate Quality	Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate
Low Quality	Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate
Very Low Quality	Any estimate of effect is very uncertain

Guyatt, G.H., et al., GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. BMJ, 2008; 336: 924-926.

Guyatt, G.H., et al., GRADE: going from evidence to recommendations. BMJ, 2008; 336: 1049-1051.