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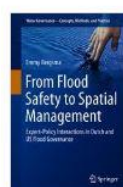
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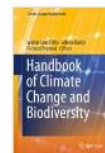


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1. Lehrnbecher T, Robinson P, Fisher B, Alexander S, Ammann RA, Beauchemin M, Carlesse F, Groll AH, Haeusler GM, Santolaya M, Steinbach WJ, Castagnola E, Davis BL, Dupuis LL, Gaur AH, Tissing WJE, Zaoutis T, Phillips R, Sung L (2017) Guideline for the management of fever and neutropenia in children with cancer and hematopoietic stem-cell transplantation recipients: 2017 update. *J Clin Oncol* 35:2082–2094.
<https://doi.org/10.1200/JCO.2016.71.7017>
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2. Stryjewski GR, Nylen ES, Bell MJ, Snider RH, Becker KL, Wu A, Lawlor C, Dalton H (2005) Interleukin-6, interleukin-8, and a rapid and sensitive assay for calcitonin precursors for the determination of bacterial sepsis in febrile neutropenic children. *Pediatr Crit Care Med* 6:129–135. <https://doi.org/10.1097/01.PCC.0000149317.15274.48>
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Predictive value of PCT and IL-6 for bacterial infection in children with cancer and febrile neutropenia

Hilde T. van der Gallen¹ · Erik A. H. Loeffen¹ · Karin G. E. Miedema¹ · Wim J. E. Tissing¹

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Abstract

Purpose Only a third of children with cancer and febrile neutropenia (FN) have a proven bacterial infection; nevertheless, most children are hospitalized and treated with intravenous antibiotics. Several biomarkers have been proposed as predictive markers for bacterial infection in this population. We aimed to evaluate the role of interleukin-6 (IL-6) and procalcitonin (PCT) in diagnosing bacterial infection in children with cancer and FN.

Methods The study population was derived from a prospective database (2006–2013, IL-8 study) comprising children with cancer who presented with FN. From stored plasma samples (taken at admission and/or at 12–24 h), we determined the PCT and IL-6 levels. Consequently, we explored their relation with the presence of bacterial infection (positive blood culture, radiologically documented infection or clinical bacterial focus). We predefined cutoff values at 60 ng/L for IL-6 and 0.25 ng/mL for PCT.

Results Seventy-seven FN episodes in 55 children with cancer were included. In 18 episodes (23.4%), a bacterial infection was documented. Both at presentation and after 12–24 h, median values of IL-6 and PCT were significantly higher in patients with a bacterial infection compared to patients without a bacterial infection. With both biomarkers above cutoff values, sensitivity was 93% (with either one, this was even 100%). The identified group at low risk for bacterial infection comprised 41% of the population.

Conclusion PCT and IL-6 are promising markers in identifying bacterial infection in children with cancer and FN. In a subsequent project, we will incorporate these biomarkers in a risk assessment model that we will test prospectively in a clinical trial.

Keywords Pediatric oncology · Febrile neutropenia · Bacterial infection · Biomarkers · Procalcitonin · Interleukin-6

Introduction

Children with cancer who develop fever after receiving chemotherapy are usually immediately referred to the hospital. In case their neutrophil count is very low, all patients used to be hospitalized and treated with broad-spectrum intravenous antibiotics until they become afebrile and neutrophil counts are starting to recover [1]. This aggressive strategy to manage febrile neutropenia has led to a great decrease in infection-related mortality and morbidity [2, 3]. On the other hand, studies have shown that only 20–30% of children with febrile

causes of fever during chemotherapy may be the effect of (chemotherapeutic) drugs, viral or fungal infections, blood transfusions, mucositis, or the malignancy itself [4]. In recent years, it has become practice to manage some patients with low risk fever and neutropenia as an outpatient with oral antibiotics. Being at low risk of systemic infection or sepsis, these patients benefit from a milder approach, preventing overtreatment [6, 7]. Yet, it is still very difficult to distinguish children with and without bacterial infection upon presentation, because at that time, fever may be the only symptom of bacterial infection [8]. Other symptoms are typically lacking,

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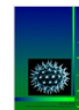
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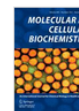
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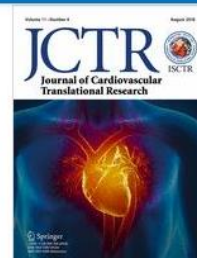
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