



QUICK GUIDE

to Clinical Utility of PCT*

EARLY IDENTIFICATION AND RISK ASSESSMENT

Clinical Utility of PCT	Key Message	Why	Scientific Proof
<p>PCT aids in the risk assessment of critically ill patients for progression to sepsis/septic shock.</p>	<p>Combined with clinical assessment, PCT contributes significantly to early identification and risk stratification of patients with possible sepsis:</p> <ul style="list-style-type: none"> - Low PCT levels may help to rule out sepsis risk and help physicians focus on other medical conditions. - High PCT levels help confirm that sepsis is very likely and treatment should be initiated. 	<ul style="list-style-type: none"> - PCT production is mainly stimulated in response to severe bacterial infections - Compared to C-reactive protein or lactate, PCT increases rapidly (within 4-6 hours) upon stimulation. This helps with early confirmation of sepsis risk allowing clinicians to intervene before complications like organ dysfunction set in. 	<p>Meisner M. Update on procalcitonin measurements. <i>Ann Lab Med.</i> 2014;34:263-273</p> <p>Christ-Crain M, et al. Procalcitonin in bacterial infections-hype, hope, more or less? <i>Swiss Medical Weekly</i> 2005; 135(31-32):451-460</p> <p>Harbarth S, et al. Diagnostic value of procalcitonin, interleukin-6 and interleukin 8 in critically ill patients admitted with suspicion of sepsis. <i>Am J Respir Crit Care Med.</i> 2001;164:396-402</p> <p>Muller B, et al. Calcitonin precursors are reliable markers of sepsis in medical intensive care unit. <i>Crit Care Med.</i> 2000;28:977-983</p>

MONITORING AND PROGNOSIS

Clinical Utility of PCT	Key Message	Why	Scientific Proof
<p>PCT aids in monitoring treatment response and prognosis in patients with sepsis and septic shock.</p>	<p>Serial measurements of PCT over time provide valuable information for patient response to treatment and risk of mortality.</p>	<p>Following a rapid increase in the presence of a bacterial infection, PCT levels drop by about 50% daily if patient evolution is favorable, helping to determine treatment success:</p> <ul style="list-style-type: none"> - with a 24-hour half-life, decreasing PCT levels are consistent with an improving condition - PCT levels that fail to decrease may be indicative of treatment failure - in a large study (n=646), a PCT decrease >80% between day 1 and 4 showed a mortality rate of 10% whereas a decrease ≤80% showed a mortality rate of 20%. 	<p>Schuetz P, et al. Serial procalcitonin predicts mortality in severe sepsis patients: Results from the multicenter procalcitonin Monitoring Sepsis (MOSES) study. <i>Crit Care Med.</i> 2017;45(5):781-789</p>

*PCT: Procalcitonin

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ANTIBIOTIC THERAPY GUIDANCE - SBI**/ SEPSIS

Clinical Utility of PCT	Key Message	Why	Scientific Proof
<p>PCT aids in decision-making on antibiotic therapy discontinuation for patients with suspected or confirmed sepsis, or other severe bacterial infections.</p>	<p>When supported by a clinical assessment, PCT helps in guiding decisions to stop or continue antibiotic therapy. It thereby contributes to reducing unnecessary prolonged use of antibiotics, patient toxicity, <i>C. difficile</i> infections and antimicrobial resistance.</p>	<p>PCT's unique kinetics help guide physicians in decisions for antibiotic treatment:</p> <ul style="list-style-type: none"> - many studies have demonstrated that PCT is safe and effective in an algorithm to determine continuing or stopping antibiotic treatment - decreasing PCT levels are consistent with an improving condition - PCT levels that fail to decrease may be indicative of treatment failure - when supported by an improved clinical condition, PCT levels that show a constant decrease (drop to ≤0.5 ng/mL or >80% from peak) over time (1-2 days) are supportive of discontinuing antibiotic therapy in septic patients. 	<p>Siriwardena AK, et al. A Procalcitonin-based Algorithm to Guide Antibiotic Use in Patients with Acute Pancreatitis (PROCAP): A Single-centre, Patient-blinded, Randomised Controlled Trial. <i>Lancet Gastroenterol Hepatol.</i> 2022;7(10):913-921</p> <p>Kyriazopoulou E, et al. Procalcitonin to Reduce Long-Term Infection-associated Adverse Events in Sepsis: A Randomized Trial. <i>Am J Respir Crit Care Med.</i> 2021;203(2):202-210</p> <p>De Jong E, et al. Efficacy and safety of procalcitonin guidance in reducing the duration of antibiotic treatment in critically ill patients: A randomised, controlled, open-label trial. <i>The Lancet.</i> 2016;16(7):819-827</p> <p>Drozdov D, et al. Procalcitonin and Pyuria-based Algorithm Reduces Antibiotic Use in Urinary Tract Infections: A Randomized Controlled Trial. <i>BMC Med.</i> 2015;13:104</p> <p>Bouadma L, et al. Use of procalcitonin to reduce patients' exposure to antibiotics in intensive care units (PRORATA Trial): a multicentre randomised controlled trial. <i>The Lancet</i> 2010;375(9713):463-74</p>

ANTIBIOTIC THERAPY GUIDANCE - LRTI***

Clinical Utility of PCT	Key Message	Why	Scientific Proof
<p>PCT aids in decision-making on antibiotic therapy discontinuation for patients with suspected or confirmed LRTI - defined as community-acquired pneumonia (CAP), acute bronchitis, or acute exacerbation of chronic obstructive pulmonary disease (AECOPD).</p>	<p>Most LRTI infections are of non-bacterial origin, however most patients presenting with these conditions are put on antibiotics.</p> <p>PCT can assist in determining if antibiotics are really needed in these patient groups, and once antibiotic therapy is started, PCT can help in determining when it is safe to discontinue treatment.</p>	<p>Measuring PCT levels upon patient presentation has been found to reduce the initiation of antibiotic treatment:</p> <ul style="list-style-type: none"> - in the case of a low PCT level (<0.25 ng/mL), it is unlikely that the patient has a bacterial infection - once antibiotic treatment has been initiated, serial PCT measurements are helpful to determine if treatment should be continued or if it can be safely discontinued - PCT levels that drop below 0.25 ng/ml are indicative of the resolution of an infection. If the patient condition has improved, antibiotic treatment may be discontinued. 	<p>Tsalik EL, et al. Efficacy and Safety of Azithromycin versus Placebo to Treat Lower Respiratory Tract Infections associated with Low Procalcitonin: A Randomised, Placebo-controlled, Double-blind, Non-inferiority Trial. <i>Lancet Infect Dis.</i> 2023;23(4):484-495</p> <p>Schuetz P, et al. Effect of Procalcitonin-Guided Antibiotic Treatment on Mortality in Acute Respiratory Infections: A Patient Level Meta-Analysis. <i>Lancet Infect Dis.</i> 2018;18(1):95-107</p> <p>Albrich WC, et al. Effectiveness and safety of procalcitonin-guided antibiotic therapy in lower respiratory tract infections in «Real Life»: An international, multicenter poststudy survey (ProREAL). <i>Arch Intern Med.</i> 2012;172(9):715-722</p> <p>Schuetz P, et al. Effect of procalcitonin-based guidelines vs standard guidelines on antibiotic use in lower respiratory tract infections: The ProHOSP randomized controlled trial. <i>JAMA</i> 2009;302(10):1059-1066</p>

*PCT: Procalcitonin **SBI: Severe Bacterial Infection ***LRTI: Lower Respiratory Tract Infection