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On suspicion of sepsis every second counts

B·R·A·H·M·S PCT: Early sepsis diagnosis and monitoring in critically ill patients



Procalcitonin (PCT) in INTENSIVE CARE



B·R·A·H·M·S PCT facilitates early sepsis diagnosis

PCT – the best in early diagnosis of bacterial infection and sepsis

Early detection and specific clinical intervention is crucial for the improved outcome of patients with sepsis. Every hour delay in receiving antibiotics is associated with 7.6% decrease in survival in adults with septic shock.¹

PCT levels increase within 3-6 hours after bacterial challenge and make a significant contribution to the clinical diagnosis of sepsis. In patients with >0.5 μ g/L PCT a bacterial infection is likely and the administration of antibiotics is recommended.²⁻⁴

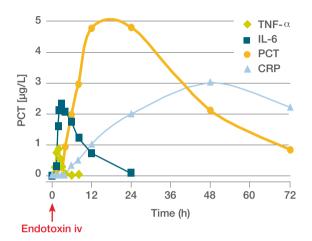


Figure 1 Kinetics of PCT compared to other inflammatory markers upon infection 2,3,5,6

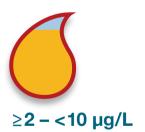
PCT levels rise with increasing severity of infection²



Systemic infection not likely



Significant, but moderate systemic inflammatory response, sepsis likely



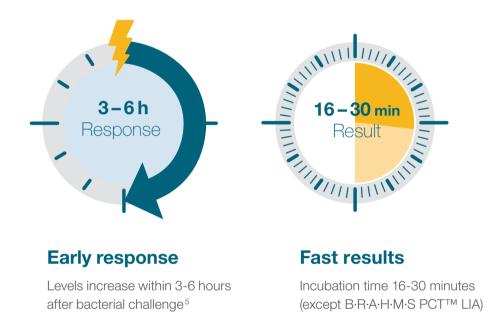
Severe sepsis: Severe systemic inflammatory response (SIRS), most likely due to infection (sepsis), high risk of developing organ dysfunction



Important systemic inflammatory response almost exclusively due to severe bacterial sepsis or septic shock

Figure 2 PCT reference ranges and their correlation with the patient's clinical condition.² As an expression of individually different immune responses and different clinical situations, the same focus of infection may be associated with varying individual elevations in PCT concentrations. Therefore, PCT results should be interpreted in context of the patient's clinical condition and other laboratory findings.

Thermo Scientific[™] B·R·A·H·M·S PCT[™] meets the needs of sepsis diagnosis





Highly sensitive and specific

A mean sensitivity of 0.77 and a mean specificity of 0.79 to discriminate sepsis from SIRS of non-infectious origin – based on meta-analysis of 3244 patients⁷

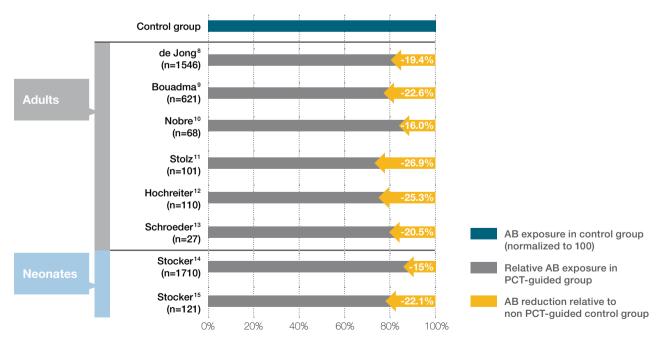
B·R·A·H·M·S PCT – effective and safe

PCT-guidance reduces antibiotic exposure

Strong evidence supports safe reduction of antibiotics using PCT-guided antibiotic stewardship protocols

- Reproducible, randomized clinical trials with more than 10,000 patients
- Proven utility across diverse clinical settings: ICU, ED, Pediatrics, Neonatology, Surgery





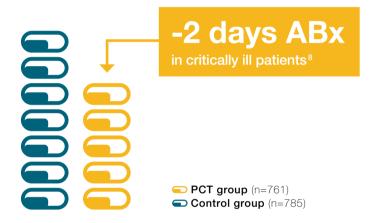


AB exposure in control group is normalized to 100, shown by a blue bar at the top. The gray bar depicts the relative exposure in PCT group and the orange bar shows the relative AB exposure reduction. All studies reported significant reduction in AB exposure.

PCT-guided reduction in antibiotic exposure could also reduce mortality rates

Data from: The Stop Antibiotics on Procalcitonin Guidance Study⁸

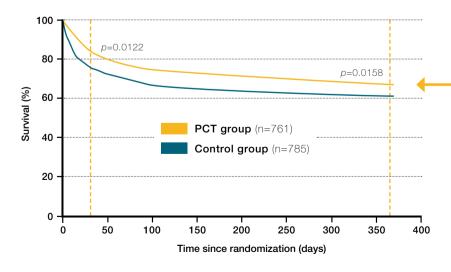
- · Largest prospective, multi-centre, randomized, controlled, open-label intervention trial with PCT in critically ill patients
- Conducted in the Netherlands a healthcare system with comparatively low use of ABx¹⁶
- 1575 critically ill patients, 15 centers



PCT could help customize the duration of antibiotics to individual patient needs.

By avoiding unnecessary antibiotics, it could also reduce the side-effects associated with them.

Figure 4 Median duration of AB treatment in PCT-guided group = 5 days, in control group = 7 days⁸



-6% mortality

at 1 year compared with control group⁸

Appropriate antibiotic use translates into survival benefit

Figure 5 Probability of survival to day 365 in the PCT-guided group vs standard of care group $^{\rm 8}$

Tailoring antibiotic therapy to individual patient needs

Monitoring with B·R·A·H·M·S PCT provides therapeutic guidance

Effective antibiotic treatment is reflected by declining PCT values,¹⁷ consistent with its half-life time of about 20-24 hours.³

Serial determinations of PCT can be used to monitor the course of infection in sepsis patients. Appropriate empiric antibiotic therapy was associated with a significant decline in PCT from day 2 to day 3 (Δ PCT \geq 30%).¹⁷

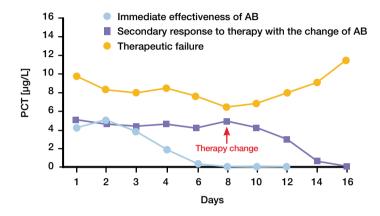
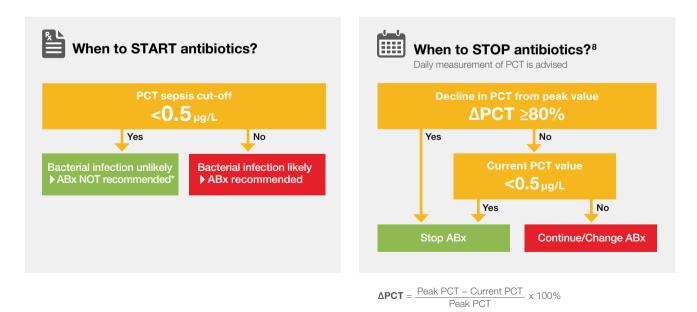


Figure 6 Typical course of PCT serum level according to patient's response to antibiotic treatment $(n=109)^{18}$

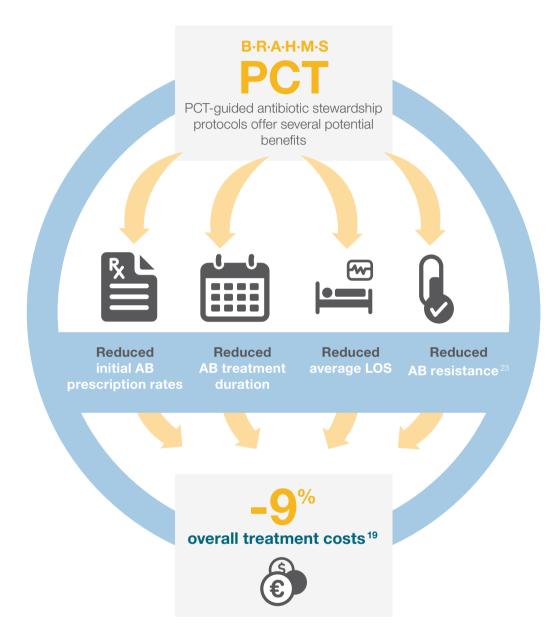
B·R·A·H·M·S PCT guided algorithms for antibiotic therapy



* PCT values should always be interpreted in context of the patient's clinical condition. Antibiotic treatment should be started/continued on suspicion of infection, particularly in high-risk patients.

Saving costs

The economic impact of PCT-guided treatment has been studied through health economic modeling.¹⁹⁻²¹ Treatment cost reductions ranging from 9% to 12% have been demonstrated across various countries.²²

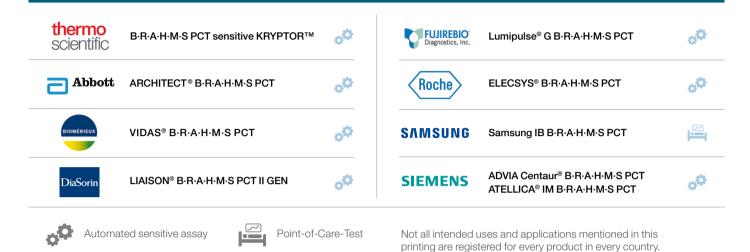


Surviving Sepsis Campaign: "We suggest that measurement of procalcitonin levels can be used to support **shortening the** duration of antimicrobial therapy in sepsis patients."

Source: International Guidelines for Management of Severe Sepsis and Septic Shock, 2016²⁴

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B·R·A·H·M·S PCT – Secured clinical decision making independent of platform



References

- Kumar et. al., Crit Care Med 2006; 34(6); 1589-96
- Kurriar et. al., Oric Carle Med 2006, 34(6), 1589-990 Harbarth et al., Am J Bespir Crit Care Med 2001; 164: 396-402 Meisner M, Procalcitonin Biochemistry and Clinical Diagnosis, ISBN 978-3-8374-1241-3, UNI-MED, Bremen 2010 Müller et al., Crit Care Med 2000; 28 (4): 977-83 3

- Brunkhorst et al., Int Care Med 1998; 24: 888-92 Dandona, J Clin Endocrinol Metab 1994; 79: 1605-8 6
- Wacker et al., Lancet Infect Dis 2013; 13: 426-35 de Jong et al., Lancet Infect Dis 2016; 3099: 1-9
- 9. Bouadma et al., Lancet 2010; 375: 463-74
- 10. Nobre et al., Am J Respir Crit Care Med 2008; 177: 498-505

- Nobre et al., Am J Respir Crit Care Med 2008; 177: 498-505
 Stolz et al., Eur Respir J 2009; 34: 1364-1375
 Hochreiter et al., Crit Care 2009; 13: R83
 Schroeder et al., Langenbecks Arch Surg 2009; 394: 221-226
 Stocker et al., Langenbecks Arch Surg 2009; 394: 221-226
 Stocker et al., Neonatology 2010; 97: 165-74
 Filippini et al., Health Policy 2006; 78(1): 77-92
 Charles et al., Orit Care 2009; 13: R38
 Stüber, 21st Int. Congress of Intensive Care and Emergency Medicine (ISICEM). Response 10: 2010 Medicine (ISICEM), Brussels 2001

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19. Kip et al., J Med Econ 2015: 1-10.

Milan, 2015

Schuetz et al., Clin Chem Lab Med 2015; 53(4): 583-92
 Maas et al., Poster, ISPOR 18th Annual European Congress,

22. Steuten, Cost effectiveness of biomarker testing in sepsis

patients; ISICEM Brussels, March 15-19, 2018 23. Hohn et al., Infection 2015; 43(4): 405-12

24. Rhodes et al., Int Care Med 2017; 43(3): 304-77

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