

**Title:** *Stenotrophomonas* Smackdown: Combination vs. Monotherapy for *Stenotrophomonas maltophilia*

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**Presenter:**

Hanna Gosliak Jamison, PharmD

PGY-1 Acute Care Pharmacy Resident

UPMC Presbyterian Hospital

**Learning Objectives:**

1. Define the characteristics of *Stenotrophomonas maltophilia* infections and the specific populations it commonly affects
2. Discuss the different treatment options available for patients with *Stenotrophomonas maltophilia* infections
3. Explain the current approach to treatment based on available literature comparing combination versus monotherapy

**Abstract:**

*Stenotrophomonas maltophilia* is a gram-negative bacillus that has increasing incidence among nosocomial pathogens.<sup>1</sup> It most commonly infects immunocompromised hosts and usually manifests as pneumonia or bacteremia, which carries a significant morbidity and mortality risk.<sup>1,2</sup> However, *S. maltophilia* is often recovered as a component of polymicrobial infections and it is difficult to distinguish if *S. maltophilia* represents a colonizer or a true pathogen.<sup>1</sup> There is currently no standard of care antibiotic regimen for the treatment of *S. maltophilia* infections, specifically with debate between the additive benefit of combination therapy.<sup>1</sup> The IDSA guidelines recommend combination therapy, but UPMC Presbyterian guidelines recommend monotherapy.<sup>1</sup> CLSI has established breakpoints for four antimicrobial agents: cefiderocol, levofloxacin, minocycline, and trimethoprim-sulfamethoxazole, but there is limited pharmacokinetic/pharmacodynamic data to correlate MICs and clinical outcomes.<sup>1,3</sup> The

aim of this presentation is to review the available literature comparing combination and monotherapy regimens for *Stenotrophomonas maltophilia* infections.

## References:

1. Tamma PD, Heil EL, Justo JA, et al. Infectious diseases society of America 2024 guidance on the treatment of antimicrobial-resistant gram-negative infections. *Clin Infect Dis*. 2024;ciae403. doi: 10.1093/cid/ciae403
2. Shah MD, Coe KE, El Boghdadly Z, et al. Efficacy of combination therapy versus monotherapy in the treatment of *Stenotrophomonas maltophilia* pneumonia. *J Antimicrob Chemother*. 2019;74(7):2055–2059. doi: 10.1093/jac/dkz116
3. Kullar R, Wenzler E, Alexander J, et al. Overcoming *Stenotrophomonas maltophilia* resistance for a more rational therapeutic approach. *Open Forum Infect Dis*. 2022;9(5):ofac095. doi: 10.1093/ofid/ofac095
4. Chen L, Hua J, Hong S, et al. Assessment of the relative benefits of monotherapy and combination therapy approaches to the treatment of hospital-acquired *Stenotrophomonas maltophilia* pneumonia: a multicenter, observational, real-world study. *Ann Intensive Care*. 2023;13(1):47. doi: 10.1186/s13613-023-01144-7
5. Cefiderocol. UpToDate Lexidrug, Drug Information Lexi-Drugs Online. Waltham, MA: UpToDate, Inc.; Version 301.0. Accessed December 27, 2024
6. Ceftazidime and avibactam. UpToDate Lexidrug, Drug Information Lexi-Drugs Online. Waltham, MA: UpToDate, Inc.; Version 301.0. Accessed January 3, 2025
7. Aztreonam. UpToDate Lexidrug, Drug Information Lexi-Drugs Online. Waltham, MA: UpToDate, Inc.; Version 301.0. Accessed January 3, 2025
8. Minocycline. UpToDate Lexidrug, Drug Information Lexi-Drugs Online. Waltham, MA: UpToDate, Inc.; Version 301.0. Accessed January 3, 2025
9. Sarzynski SH, Warner S, Sun J, et al. Trimethoprim-sulfamethoxazole versus levofloxacin for *Stenotrophomonas maltophilia* infections: a retrospective comparative effectiveness study of electronic health records from 154 US hospitals. *Open Forum Infect Dis*. 2022;9(2):ofab644. doi: 10.1093/ofid/ofab644
10. Gibb J, Wong DW. Antimicrobial treatment strategies for *Stenotrophomonas maltophilia*: a focus on novel therapies. *Antibiotics*. 2021;10(10):1226. doi: 10.3390/antibiotics10101226
11. Wei C, Ni W, Cai X, et al. A Monte Carlo pharmacokinetic/pharmacodynamic simulation to evaluate the efficacy of minocycline, tigecycline, moxifloxacin, and levofloxacin in the treatment of hospital-acquired pneumonia caused by

- Stenotrophomonas maltophilia*. *Infect Dis (Lond)*. 2015;47(12):846–851. doi: 10.3109/23744235.2015.1064542
12. *Stenotrophomonas maltophilia*. *ILUM Inform*. Accessed January 12, 2025.
  13. Hase R, Sakurai A, Suzuki M, et al. Clinical characteristics and genome epidemiology of *Stenotrophomonas maltophilia* in Japan. *J Antimicrob Chemother*. 2024;79(8):1843–1855. doi:10.1093/jac/dkae168
  14. Karakostas S, Rousaki M, & Kritsotakis EI. Cefiderocol: systematic review of mechanisms of resistance, heteroresistance and in vivo emergence of resistance. *Antibiotics*. 2022;11(6):723. doi: 10.3390/antibiotics11060723
  15. Shah S, Slaven B, Clarke L, et al. Clinical and microbiologic outcomes of *Stenotrophomonas maltophilia* bloodstream infections. *Infect [accepted]*. March 2025.
  16. Sader HS, Duncan LR, Arends SJ, et al. Antimicrobial activity of aztreonam-avibactam and comparator agents when tested against a large collection of contemporary *Stenotrophomonas maltophilia* Isolates from medical centers worldwide. *Antimicrobi Agents Chemother*. 2020;64(11):e01433-20. doi: 0.1128/AAC.01433-20
  17. Fratoni, AJ, Nicolau DP, & Kuti JL.. Minocycline pharmacodynamics against *Stenotrophomonas maltophilia* in the neutropenic murine infection model: implications for susceptibility breakpoints. *J Antimicrob Chemother*. 2022;77(4);1052–1060. doi: 10.1093/jac/dkac018
  18. Petraitis V, Petraitiene R, Kavaliauskas P, et al. Efficacy of cefiderocol in experimental *Stenotrophomonas maltophilia* pneumonia in persistently neutropenic rabbits. *Antimicrob Agents Chemother*. 2022;66(10);e0061822. doi: 10.1128/aac.00618-22

### **Audience Response Questions:**

1. Which of the following patients is at the highest risk of *S. maltophilia* infection?
  - a. A 29-year-old female who presents to her primary care physician with a productive cough and sputum production for 4 days
  - b. A 36-year-old male who has leukemia and was recently treated with a course of meropenem for neutropenic fever
  - c. A 54-year-old male who stepped on a rusty nail and presents to the emergency room with concern for an SSTI
  - d. An 85-year-old female with COPD who presents to her primary care physician with fever, weakness, and confusion

2. Which of the following is an appropriate treatment option for *S. maltophilia* according to the IDSA guidelines?
  - a. Ceftazidime-avibactam + aztreonam
  - b. Trimethoprim/sulfamethoxazole
  - c. Ceftazidime + minocycline
  - d. Levofloxacin + cefepime
3. Which of the following treatment options is NOT recommended at our institution due to intrinsic resistance?
  - a. Minocycline
  - b. Levofloxacin
  - c. Trimethoprim/sulfamethoxazole
  - d. Cefepime