#### THE UNIVERSITY OF TEXAS **MDAnderson Cancer** Center Making Cancer History®

Patient presents with

neutropenia and fever

suspected or proven  $\rightarrow$ 

# **Neutropenic Fever<sup>1</sup> Outpatient Treatment For** Solid Tumor Patients (18 years and older)

Disclaimer: This algorithm has been developed for MD Anderson using a multidisciplinary approach considering circumstances particular to MD Anderson's specific patient population, services and structure, and clinical information. This is not intended to replace the independent medical or professional judgment of physicians or other health care providers in the context of individual clinical circumstances to determine a patient's care. Local microbiology and susceptibility/resistance patterns should be taken into consideration when selecting antibiotics. This algorithm should not be used to treat pregnant women.

- Complete history and physical exam
- Start IV fluids
- CBC with differential, BMP, lactic acid
- Blood cultures (with a set collected from each lumen simultaneously if CVC present and 1 peripheral site); other cultures (i.e., sputum culture, urinalysis with culture and sensitivity; respiratory PCR multiplex panel) only if clinically indicated
- Chest x-ray or other tests as clinically indicated
- Calculate MASCC Risk Index score (see Appendix A)

- Assess if patient is considered low risk: (*i.e.*, MASCC Risk Index score  $\geq 21$  and no other complicating factors present) and meets all of the following criteria for outpatient treatment:
- Solid tumor
- Able to tolerate oral medications
- Able to tolerate fluids
- Does not use feeding tube as primary route for nutrition and medications
- No confirmed focus of infection
- Resides within 1 hour travel time of MD Anderson No history of non-compliance
- Has a 24-hour caregiver

- Has access to transportation and telephone at residence
- Age  $\geq$  18 years old
- No quinolone allergy for oral regimens
- No colonization with fluoroquinolone-resistant or multi-drug resistant organisms
- Not currently on antibiotics



<sup>1</sup>ANC < 1 K/microliter and temperature  $\geq$  38.3°C or equal to 38°C for 1 hour or longer

<sup>2</sup>Doses indicated are for patients with normal renal/hepatic function

<sup>3</sup>See Appendix B: Outpatient Follow up

BMP = basic metabolic panel MASCC = Multinational Association of Supportive Care in Cancer NF = neutropenic fever

Department of Clinical Effectiveness V7 Approved by The Executive Committee of the Medical Staff on 09/17/2019

Copyright 2019 The University of Texas MD Anderson Cancer Center



### Neutropenic Fever Outpatient Treatment For Solid Tumor Patients (18 years and older)

Page 2 of 5

Disclaimer: This algorithm has been developed for MD Anderson using a multidisciplinary approach considering circumstances particular to MD Anderson's specific patient population, services and structure, and clinical information. This is not intended to replace the independent medical or professional judgment of physicians or other health care providers in the context of individual clinical circumstances to determine a patient's care. Local microbiology and susceptibility/resistance patterns should be taken into consideration when selecting antibiotics. This algorithm should not be used to treat pregnant women.

#### APPENDIX A: Multinational Association for Supportive Care in Cancer (MASCC) Risk Index Score

	MASCC Score
Characteristic	Weight
Burden of illness: no or mild symptoms	5
No hypotension	5
No chronic obstructive pulmonary disease	e 4
Solid tumor	4
No dehydration	3
Burden of illness: moderate symptoms	3
Outpatient status	3
Age < 60 years	2
<ul> <li>"Burden of illness" not cumulative</li> <li>Patients with score ≥ 21 are considered left</li> </ul>	ow risk

#### **APPENDIX B: Outpatient Follow Up**

- Schedule outpatient visit for Days 2, 3 and 7; and phone follow-up for Days 4, 5 and 6
- Day 2: CBC with differential; repeat creatinine if baseline greater than 1.2 mg/dL
- Day 3: CBC with differential, repeat creatinine
- Day 7: CBC with differential, repeat creatinine <u>or</u> phone follow-up if NF has resolved



## Neutropenic Fever Outpatient Treatment For Solid Tumor Patients (18 years and older)

Disclaimer: This algorithm has been developed for MD Anderson using a multidisciplinary approach considering circumstances particular to MD Anderson's specific patient population, services and structure, and clinical information. This is not intended to replace the independent medical or professional judgment of physicians or other health care providers in the context of individual clinical circumstances to determine a patient's care. Local microbiology and susceptibility/resistance patterns should be taken into consideration when selecting antibiotics. This algorithm should not be used to treat pregnant women.

### SUGGESTED READINGS

- Blot, E., Héron, F., Lishner, M., Rubenstein, E., Rolston, K., Kim, Y., . . . Talcott, J. (2000). Oral antibiotics for febrile patients with neutropenia due to cancer chemotherapy. *The New England Journal of Medicine*, 342(1), 55-58. https://doi.org/10.1056/NEJM200001063420111
- Cooper, M., Durand, C., Beaulac, M., & Steinberg, M. (2011). Single-agent, broad-spectrum fluoroquinolones for the outpatient treatment of low-risk febrile neutropenia. *Annals of Pharmacotherapy*, 45(9), 1094-1102. https://doi.org/10.1345/aph.1Q147
- Cornely, O., Wicke, T., Seifert, H., Bethe, U., Schwonzen, M., Reichert, D., ... Fätkenheuer, G. (2004). Once-daily oral levofloxacin monotherapy versus piperacillin/tazobactam three times a day: A randomized controlled multicenter trial in patients with febrile neutropenia. *International Journal of Hematology*, 79(1), 74-78. https://doi.org/10.1007/BF02983537
- Elting, L., Lu, C., Escalante, C., Giordano, S., Trent, J., Cooksley, C., . . . Rolston, K. (2008). Outcomes and cost of outpatient or inpatient management of 712 patients with febrile neutropenia. Journal of Clinical Oncology, 26(4), 606-611. https://doi.org/10.1200/JCO.2007.13.8222
- Escalante, C., Weiser, M., Manzullo, E., Benjamin, R., Rivera, E., Lam, T., . . . Rolston, K. (2004). Outcomes of treatment pathways in outpatient treatment of low risk febrile neutropenic cancer patients. *Supportive Care in Cancer*, *12*(9), 657-662. https://doi.org/10.1007/s00520-004-0613-6
- Flowers, C., Seidenfeld, J., Bow, E., Karten, C., Gleason, C., Hawley, D., . . . Ramsey, S. (2013). Antimicrobial prophylaxis and outpatient management of fever and neutropenia in adults treated for malignancy: American Society of Clinical Oncology clinical practice guideline. *Journal of Clinical Oncology*, *31*(6), 794-810. https://doi.org/10.1200/JCO.2012.45.8661
- Freifeld, A., Bow, E., Sepkowitz, K., Boeckh, M., Ito, J., Mullen, C., . . . Wingard, J. (2011). Clinical practice guideline for the use of antimicrobial agents in neutropenic patients with cancer: 2010 update by the Infectious Diseases Society of America. *Clinical Infectious Diseases*, 52(4), 427-431. https://doi.org/10.1093/cid/ciq147
- Freifeld, A., Marchigiani, D., Walsh, T., Chanock, S., Lewis, L., Hiemenz, J., & Pizzo, P. (1999). A double-blind comparison of empirical oral and intravenous antibiotic therapy for low-risk febrile patients with neutropenia during cancer chemotherapy. *The New England Journal of Medicine*, 341(5), 305-311. https://doi.org/10.1056/NEJM199907293410501
- Johnson, T., De Jesus, Y., McMahon, L., Rolston, K., & Row, M. B. (2008). Outpatient management of febrile neutropenia: Is it safe yet? *Journal of Supportive Oncology*, 6(5), 219-220. Retrieved from http://search.proquest.com/docview/71670069/
- Kern, W., Cometta, A., de Bock, R., Langenaeken, J., Paesmans, M., Zanetti, G., . . . Gaya, H. (1999). Oral versus intravenous empirical antimicrobial therapy for fever in patients with granulocytopenia who are receiving cancer chemotherapy. *The New England Journal of Medicine*, 341(5), 312-318. https://doi.org/10.1056/NEJM199907293410502

Continued on next page



## Neutropenic Fever Outpatient Treatment For Solid Tumor Patients (18 years and older)

Disclaimer: This algorithm has been developed for MD Anderson using a multidisciplinary approach considering circumstances particular to MD Anderson's specific patient population, services and structure, and clinical information. This is not intended to replace the independent medical or professional judgment of physicians or other health care providers in the context of individual clinical circumstances to determine a patient's care. Local microbiology and susceptibility/resistance patterns should be taken into consideration when selecting antibiotics. This algorithm should not be used to treat pregnant women.

#### **SUGGESTED READINGS - continued**

- Kern, W., Marchetti, O., Drgona, L., Akan, H., Aoun, M., Akova, M., . . . Calandra, T. (2013). Oral antibiotics for fever in low-risk neutropenic patients with cancer: a double-blind, randomized, multicenter trial comparing single daily moxifloxacin with twice daily ciprofloxacin plus amoxicillin/clavulanic acid combination therapy—EORTC infectious diseases group trial XV. *Journal of Clinical Oncology*, 31(9), 1149-1156. https://doi.org/10.1200/JCO.2012.45.8109
- Rolston, K., Manzullo, E., Elting, L., Frisbee-Hume, S., McMahon, L., Theriault, R., . . . Benjamin, R. (2006). Once daily, oral, outpatient quinolone monotherapy for low-risk cancer patients with fever and neutropenia: A pilot study of 40 patients based on validated risk-prediction rules. *Cancer, 106*(11), 2489-2494. https://doi.org/10.1002/cncr.21908
- Rubenstein, E., Rolston, K., Escalante, C., Manzullo, E., Hughes, P., Fender, A., . . . Benjamin, R. (1993). Outpatient treatment of febrile episodes in low-risk neutropenic patients with cancer. *Cancer*, 71(11), 3640-3646. https://doi.org/10.1002/1097-0142(19930601)71:11<3640::AID-CNCR2820711128>3.0.CO;2-H
- Taplitz, R., Kennedy, E., Bow, E., Crews, J., Gleason, C., Hawley, D., . . . Flowers, C. (2018). Outpatient management of fever and neutropenia in adults treated for malignancy. American Society of Clinical Oncology and Infectious Diseases Society of America Clinical Practice guideline update. *Journal of Clinical Oncology*, *36*(14):1443-1453. https://doi.org/10.1200/JCO.2017.77.6211
- Vidal, L., Paul, M., Ben-Dor, I., Pokroy, E., Soares-Weiser, K., & Leibovici, L. (2004). Oral versus intravenous antibiotic treatment for febrile neutropenia in cancer patients. *Cochrane Database of Systematic Reviews*, (4), CD003992. https://doi.org/10.1002/14651858.CD003992.pub2

#### THE UNIVERSITY OF TEXAS MDAnderson Cancer Center Making Cancer History\*

### Neutropenic Fever Outpatient Treatment For Solid Tumor Patients (18 years and older)

Disclaimer: This algorithm has been developed for MD Anderson using a multidisciplinary approach considering circumstances particular to MD Anderson's specific patient population, services and structure, and clinical information. This is not intended to replace the independent medical or professional judgment of physicians or other health care providers in the context of individual clinical circumstances to determine a patient's care. Local microbiology and susceptibility/resistance patterns should be taken into consideration when selecting antibiotics. This algorithm should not be used to treat pregnant women.

### **DEVELOPMENT CREDITS**

This practice consensus statement is based on majority opinion of the Neutropenic Fever experts at the University of Texas MD Anderson Cancer Center for the patient population. These experts included:

Antimicrobial Stewardship Team<sup>∓</sup> Samuel L. Aitken, PharmD (Pharmacy Clinical Programs) Patrick Chaftari, MD (Emergency Medicine) Tami N. Johnson, PharmD (Pharmacy Clinical Programs) Victor E. Mulanovich, MD (Infectious Diseases) Loretta Nastoupil, MD (Lymphoma/Myeloma) Terry W. Rice, MD (Emergency Medicine) Frank P. Tverdek, PharmD (Pharmacy Clinical Programs) Khanh D. Vu, BS, MD (General Internal Medicine) Mary Lou Warren, DNP, APRN, CNS-CC<sup>+</sup>

<sup>†</sup> Core Development Team Lead <sup>•</sup> Clinical Effectiveness Development Team