

I EDIZIONE

USO RAZIONALE DEGLI ANTIBIOTICI NELL'ERA DELLE RESISTENZE BATTERICHE

17 MAGGIO
2024

Sala Congressi
Ospedale Di Sarno

Infezioni
nel paziente
oncoematologico

Biagio Pinchera



Fatal Infections Among Cancer Patients: A Population-Based Study in the United States

Original Research | [Open access](#) | Published: 24 March 2021

Volume 10, pages 871–895, (2021) [Cite this article](#)

Infections are one of the leading causes of mortality in cancer patients and the rate of fatal infections is approximately three times higher than that of the general population

The incidence of sepsis is 16.4 cases per 1000 cancer patients per year, with in-hospital mortality of 37.8% in severe cases

Research | [Open access](#) | Published: 05 July 2004

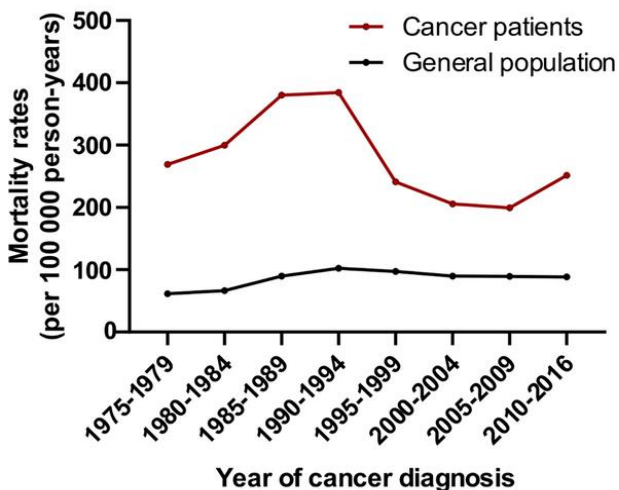
Hospitalized cancer patients with severe sepsis: analysis of incidence, mortality, and associated costs of care

[Mark D Williams](#) , [Lee Ann Braun](#), [Liesl M Cooper](#), [Joseph Johnston](#), [Richard V Weiss](#), [Rebecca L Qualy](#) & [Walter Linde-Zwirble](#)

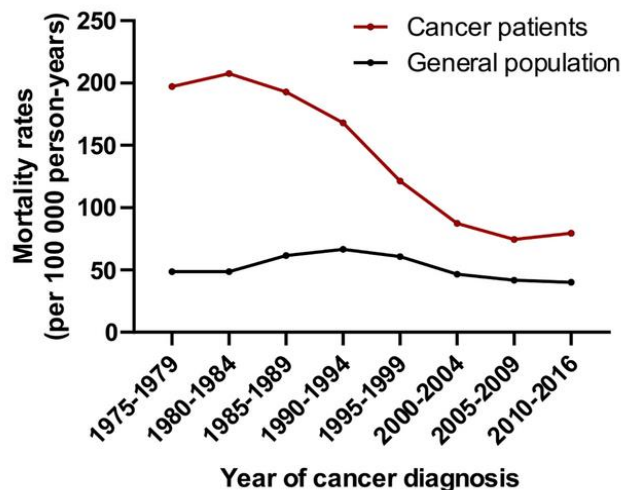
[Critical Care](#) **8**, Article number: R291 (2004)

Mortality rates of fatal infection among cancer patients by year of cancer diagnosis

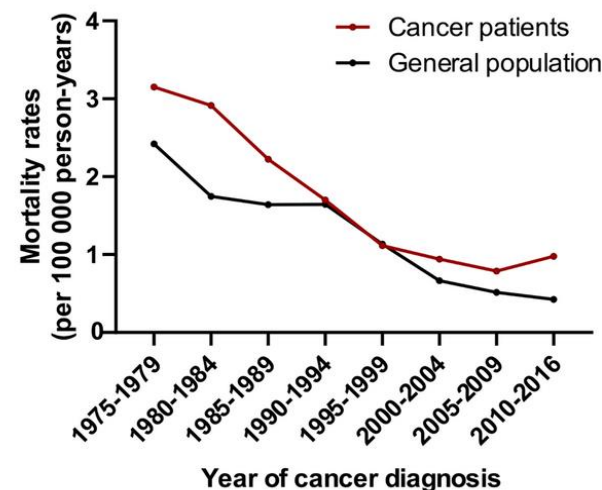
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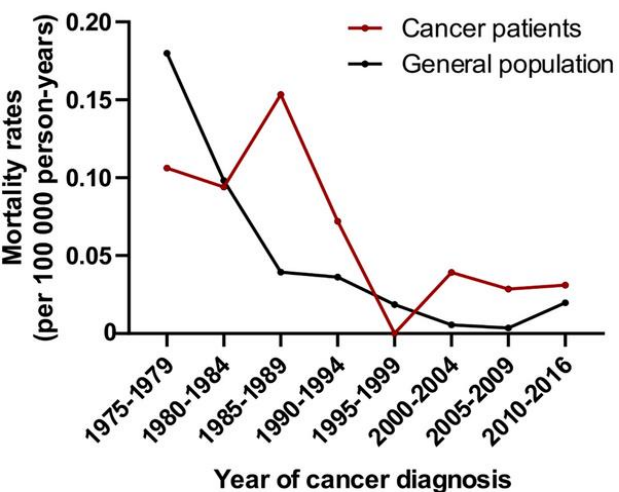
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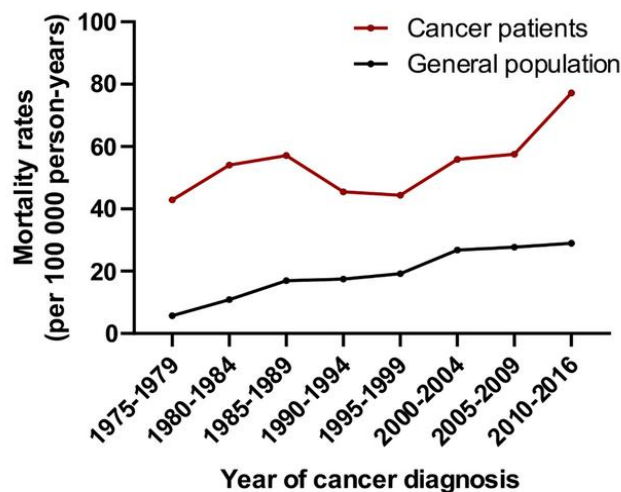
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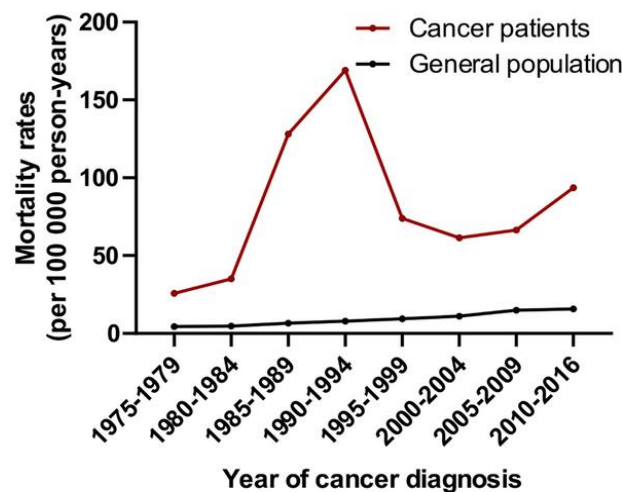
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E



F





Bloodstream infections are significantly more common in patients with hematologic malignancies than in those with solid tumors.

Interestingly, while there is a significant downward trend in bloodstream infection in hematologic patients over time, the incidence remains stable in patients with solid tumors

This underscores the importance of appropriate management of infections in cancer patients.

There are a number of differences between patients with solid tumors and those with hematologic malignancies.

the neoplastic process of those with solid tumors does not involve the effectors of the host immune system

[Home](#) > [Infectious Diseases and Therapy](#) > [Article](#)

Infections in Cancer Patients with Solid Tumors: A Review

Review | [Open access](#) | Published: 03 February 2017

the therapy of solid tumors usually does not lead to prolonged neutropenia

National Comprehensive Cancer Network NCCN Clinical Practice Guidelines in Oncology. Prevention and Treatment of Cancer-Related Infections 1. 2021

Longitudinal surveillance of bacteraemia in haematology and oncology patients at a UK cancer centre and the impact of ciprofloxacin use on antimicrobial resistance FREE

S. Schelenz ✉, D. Nwaka, P. R. Hunter

Journal of Antimicrobial Chemotherapy, Volume 68, Issue 6, June 2013, Pages 1431–1438,

Consequently, the incidence of infections in patients with solid tumors is significantly lower compared to hematologic patients

Highlights

- Bacteraemia in neutropenic cancer patients was mainly due to gram-negative bacilli.
- MDR gram-negatives were more frequent in patients with haematological malignancies.
- Pneumonia due to *Pseudomonas aeruginosa* was more common in patients with solid tumours.



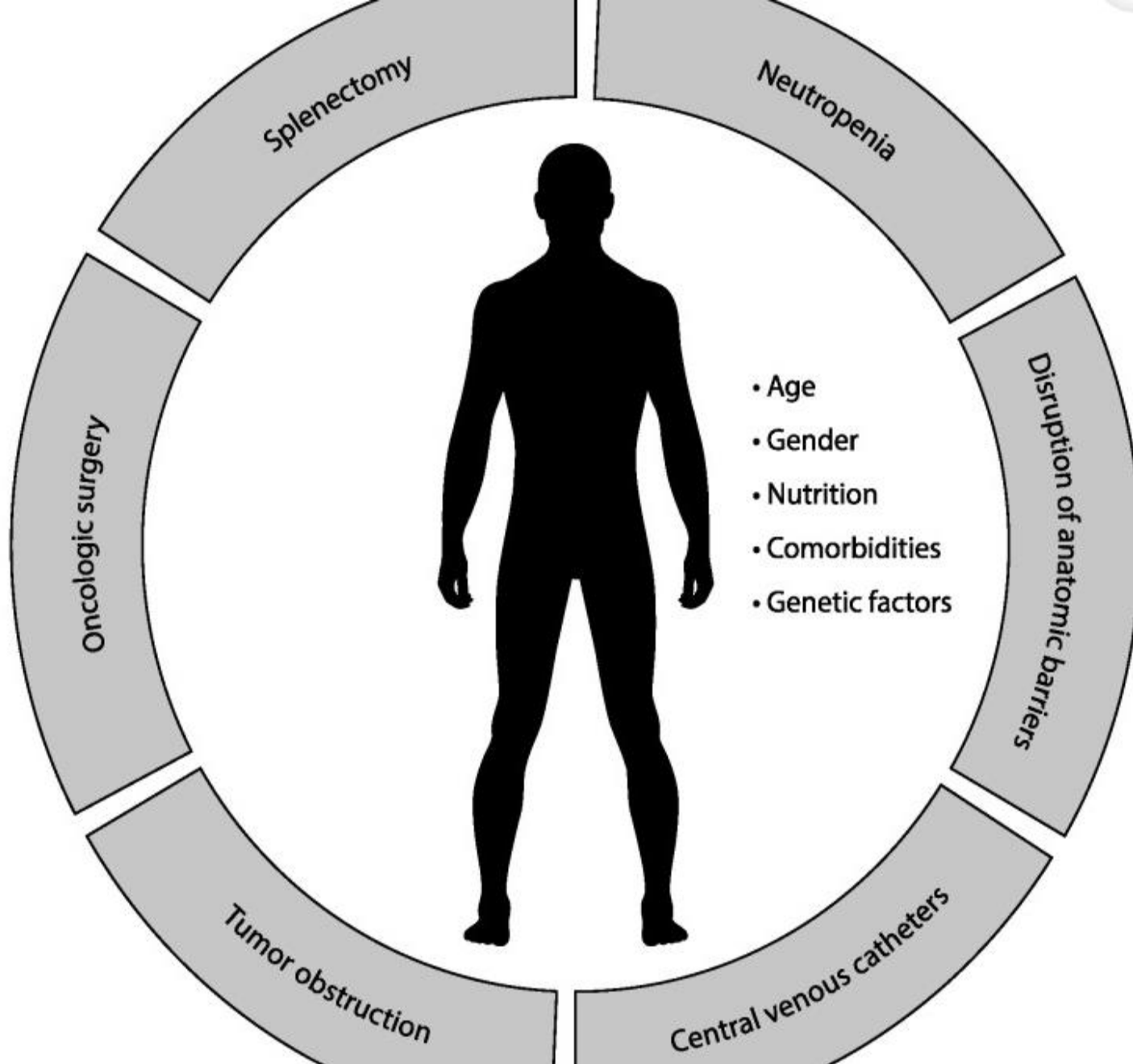
Risk Factors for Infections, Antibiotic Therapy, and Its Impact on Cancer Therapy Outcomes for Patients with Solid Tumors

[Ondřej Kubeček](#)¹, [Pavla Paterová](#)^{2,*} and [Martina Novosadová](#)³

Prophylactic use of antibiotics is, therefore, rarely used for patients with solid tumors

A possible exception is the use of trimethoprim/sulfamethoxazole for the prevention of *Pneumocystis jirovecii* pneumonia in patients receiving temozolomide together with radiation therapy, and moderate-to-high-dose corticosteroid therapy (prednisone equivalents ≥ 20 mg for ≥ 4 weeks)

National Comprehensive Cancer Network NCCN Clinical Practice Guidelines in Oncology. Prevention and Treatment of Cancer-Related Infections 1. 2024



Risk Factors for Infections in Patients



Overall Infection Risk in Patients with Cancer ^a	Disease/Therapy Examples	Antimicrobial Prophylaxis
Low	<ul style="list-style-type: none"> • Standard chemotherapy regimens for most solid tumors • Anticipated neutropenia* <7 days 	<ul style="list-style-type: none"> • Bacterial - None • Fungal - None • Viral - None unless prior HSV episode
Intermediate	<ul style="list-style-type: none"> • Autologous HCT • Lymphoma^c • Multiple myeloma^c • CLL^c • Purine analog therapy (ie, fludarabine, clofarabine, nelarabine, cladribine) • Anticipated neutropenia* 7–10 days • CAR T-cell therapy 	<ul style="list-style-type: none"> • Bacterial - Consider fluoroquinolone prophylaxis during neutropenia^d • Fungal - Consider prophylaxis during neutropenia and for anticipated mucositis (INF-2); consider PJP prophylaxis (NE-6) • Viral - During neutropenia and longer depending on risk (INF-3, INF-4, INF-5) • CAR T-cell therapy (INF-A 10 of 12)
High ^b	<ul style="list-style-type: none"> • Allogeneic HCT including cord blood • Acute leukemia <ul style="list-style-type: none"> › Induction › Consolidation/maintenance • Alemtuzumab therapy • Moderate to severe GVHD • Anticipated neutropenia* >10 days 	<ul style="list-style-type: none"> • Bacterial - Consider fluoroquinolone prophylaxis during neutropenia^d • Fungal - Consider prophylaxis during neutropenia (INF-2); consider PJP prophylaxis (INF-6) • Viral - During neutropenia and longer depending on risk (INF-3, INF-4, INF-5) • Length of prophylaxis depends on immune reconstitution.

*Neutropenia: ≤500 neutrophils/mcL or ≤1000 neutrophils/mcL and a predicted decline to ≤500/ mcL over the next 48 hours.



Disruption of Anatomic Barriers

Mucositis is another important factor increasing the risk of bloodstream infections.

Mucosal barriers in the gastrointestinal, urogenital, and respiratory tract constitute the first line of host defense against various pathogens.

Coexisting neutropenia allows for the rapid development of severe infections

Chemotherapy-induced mucositis is associated with an increased risk of infections caused by viridans group streptococci, gram-negative rods, and Candida spp.

Although the risk and grade of mucositis is higher in hematologic patients receiving high-dose chemotherapy regimens with autologous hematopoietic stem cell transplantation, some of the cytotoxic drugs used in the therapy of solid tumors can induce mucositis. These include 5-fluorouracil, capecitabine, cyclophosphamide, ifosfamide, cisplatin, carboplatin, docetaxel, paclitaxel, and vinorelbine

Management of oral and gastrointestinal mucosal injury: ESMO Clinical Practice Guidelines for diagnosis, treatment, and follow-up

D.E. Peterson • C.B. Boers-Doets • R.J. Bensadoun • J. Herrstedt •

on behalf of the ESMO Guidelines Committee ✉ • [Show footnotes](#)

Neutropenia

Severe neutropenia is defined as the ANC of <500 cells/mm³ according to the Common Terminology Criteria for Adverse Events (CTCAE)

A drop of ANC below this threshold is associated with an increased risk of infections

The incidence and severity of infections are inversely related to ANC, with the highest risk when ANC drops below 100 cells/mm³

Duration of neutropenia is another important risk factor

Chemotherapy regimens resulting in neutropenia lasting >10 days are considered high risk

However, most regimens used in the therapy of solid tumors result in neutropenia lasting <7 days and are therefore considered low risk

Primary prophylaxis of bacterial infections and *Pneumocystis jirovecii* pneumonia in patients with hematologic malignancies and solid tumors: 2020 updated guidelines of the Infectious Diseases Working Party of the German Society of Hematology and Medical Oncology (AGIHO/DGHO)

[Annika Y. Classen](#)^{1,2}, [Larissa Henze](#)³, [Marie von Lilienfeld-Toal](#)⁴, [Georg Maschmeyer](#)⁵, [Michael Sandherr](#)⁶, [Luisa Durán Graeff](#)^{1,2}, [Nael Alakei](#)⁷, [Maximilian Christopeit](#)⁸, [Stefan W. Krause](#)⁹, [Karin Mayer](#)¹⁰, [Silke Neumann](#)¹¹, [Oliver A. Cornely](#)^{1,2,12,13}, [Olaf Penack](#)¹⁴, [Florian Weißinger](#)¹⁵, [Hans-Heinrich Wolf](#)¹⁶ and [Jörg Janne Vehreschild](#)^{1,2,17}

Neutropenia is defined as an absolute neutrophil count (ANC) < 500 cells/mm³ or an expected ANC of < 500 cells/mm³ within the next 48 hours.

Chemotherapy-induced neutropenia and mucositis can lead to invasive infection most often with bacteria and fungi

Neutropenic fever occurs in 10-50% with solid tumors and >80% of patients with hematological malignancies and those receiving hematopoietic stem cell transplant (HCT).

Risk factors for bacterial infection during neutropenia

Clinical situation	Intention/recommendation	Intervention
Patients with prolonged neutropenia (> 7 days) ^a	Identify patients at risk for FN	Consider as high-risk patients
Patients with neutropenia > 0 and ≤ 7 days ^a and significant additional risk factors ^b		Consider as high-risk patients
Patients with neutropenia ≤ 7 days without additional risk factors		Consider as low-risk patients

Patient-related risk factors



Patient-related risk factors^a

Prolonged neutropenia (> 7 days)

Type and stage of underlying malignancy

Administered type and dosage of chemotherapy

First chemotherapy cycle

Cardiac insufficiency

Low baseline creatinine clearance

Low baseline leukocyte count

Elevated baseline levels of alkaline phosphatase and bilirubin

Factors that were found independently associated with development of fever during neutropenia in multivariable analyses of either the majority of studies or the majority of patients in studies testing that risk factor

[Ann Hematol.](#) 2021; 100(6): 1603–1620.

Published online 2021 Apr 13. doi: [10.1007/s00277-021-04452-9](https://doi.org/10.1007/s00277-021-04452-9)

PMCID: PMC8116237

PMID: [33846857](https://pubmed.ncbi.nlm.nih.gov/33846857/)

Primary prophylaxis of bacterial infections and *Pneumocystis jirovecii* pneumonia in patients with hematologic malignancies and solid tumors: 2020 updated guidelines of the Infectious Diseases Working Party of the German Society of Hematology and Medical Oncology (AGIHO/DGHO)

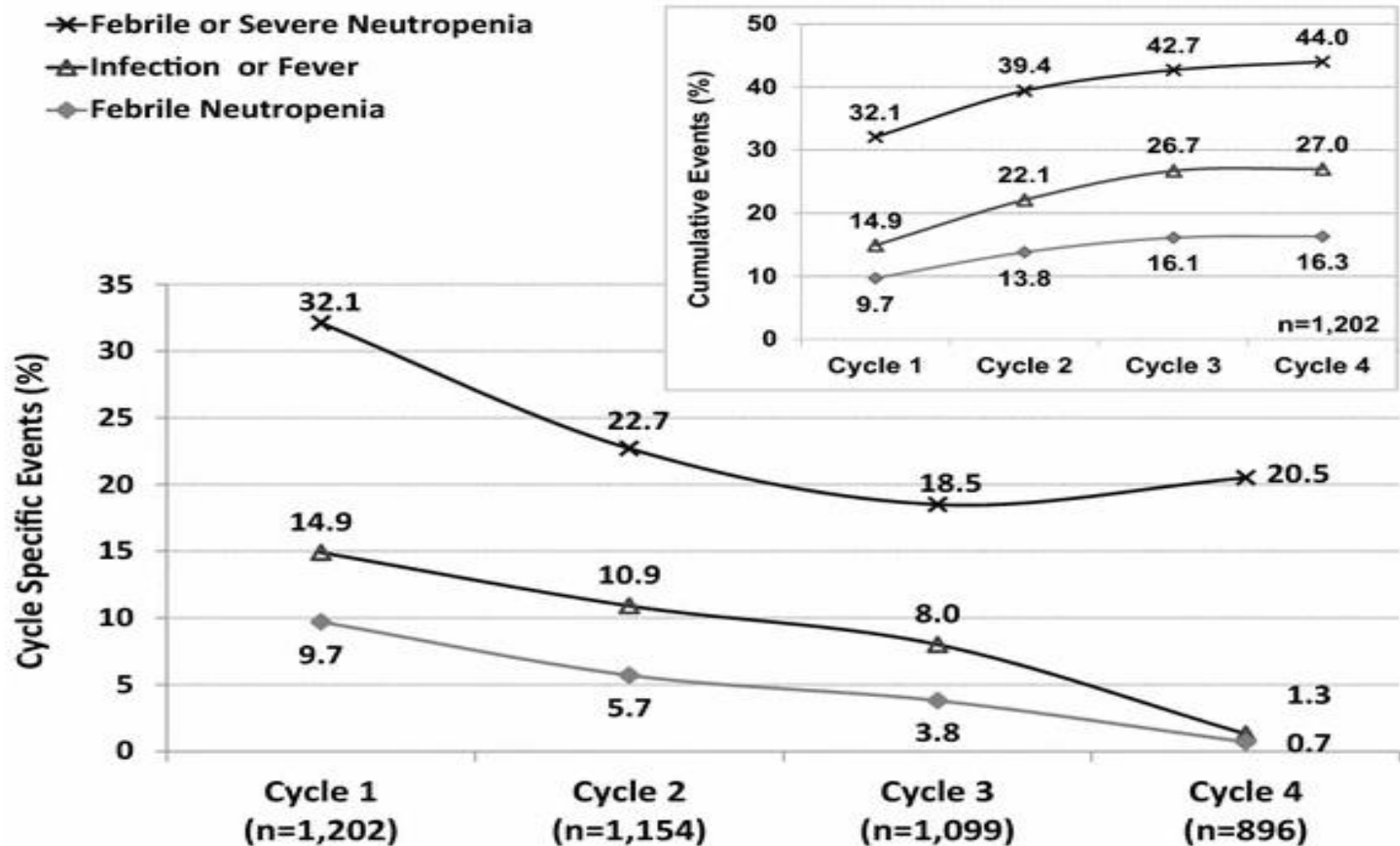
[Annika Y. Classen](#),^{1,2} [Larissa Henze](#),³ [Marie von Lilienfeld-Toal](#),⁴ [Georg Maschmeyer](#),⁵ [Michael Sandherr](#),⁶

[Luisa Durán Graeff](#),^{1,2} [Nael Alakel](#),⁷ [Maximilian Christopeit](#),⁸ [Stefan W. Krause](#),⁹ [Karin Mayer](#),¹⁰ [Silke Neumann](#),¹¹

[Oliver A. Cornely](#),^{1,2,12,13} [Olaf Penack](#),¹⁴ [Florian Weißinger](#),¹⁵ [Hans-Heinrich Wolf](#),¹⁶ and [Jörg Janne Vehreschild](#)^{21,2,17}

Incidence of febrile neutropenia in early stage breast cancer patients receiving adjuvant FEC-D treatment

Original Article | Published: 05 July 2014

Volume 22, pages 3227–3234, (2014) [Cite this article](#)

Neutropenic and infectious events during first 4 cycles. Cycle specific events and cumulative events are presented.

Predisposing factors for bloodstream infections in neutropenic and non-neutropenic patients within 48 h before bloodstream infection



Factor	No. (%) of patients			OR (95% CI)	P ^a
	ANC, neutrophils/ μ L		Total (n = 2340)		
	<1000 (n = 696)	\geq 1000 (n = 1644)			
Central venous line present	626 (89.9)	1319 (80.2)	1945 (83.1)	2.2 (1.66–2.93)	<.001
Peripheral intravenous line present	97 (13.9)	378 (23.0)	475 (20.3)	1.8 (1.44–2.37)	<.001
Arterial line present	18 (2.6)	60 (3.6)	78 (3.3)	0.7 (0.39–1.23)	.24
Urinary catheter present	83 (11.9)	381 (23.2)	464 (19.8)	2.2 (1.71–2.90)	<.001
Receipt of total parenteral nutrition	112 (16.1)	316 (19.2)	428 (18.3)	0.8 (0.63–1.03)	.08
Dialysis performed	5 (0.7)	43 (2.6)	48 (2.1)	3.7 (1.40–10.69)	.005
Stay in intensive care unit	48 (6.9)	223 (13.6)	271 (11.6)	2.1 (1.51–2.97)	<.001
Receipt of ventilator support	27 (3.9)	118 (7.2)	145 (6.2)	1.9 (1.23–3.01)	.003

NOTE. ANC, absolute neutrophil count.

^a Patients with an ANC of <1000 neutrophils/ μ L vs. those with an ANC of \geq 1000 neutrophils/ μ L.

JOURNAL ARTICLE

Current Trends in the Epidemiology of Nosocomial Bloodstream Infections in Patients with Hematological Malignancies and Solid Neoplasms in Hospitals in the United States ^{FREE}

Hilmar Wisplinghoff ✉, Harald Seifert, Richard P. Wenzel, Michael B. Edmond

Clinical Infectious Diseases, Volume 36, Issue 9, 1 May 2003, Pages 1103–1110,



Prophylaxis

G-CSF

- Risk of febrile neutropenia > 20%
- Risk of febrile neutropenia 10 – 20% with associated risk factor such as
 - ≥ 65 years
 - coexisting illness (renal, hepatic or cardiac dysfunction)
 - preexisting condition (infection, open wound or recent surgery)
 - compromised marrow reserve

CLINICAL THERAPEUTICS



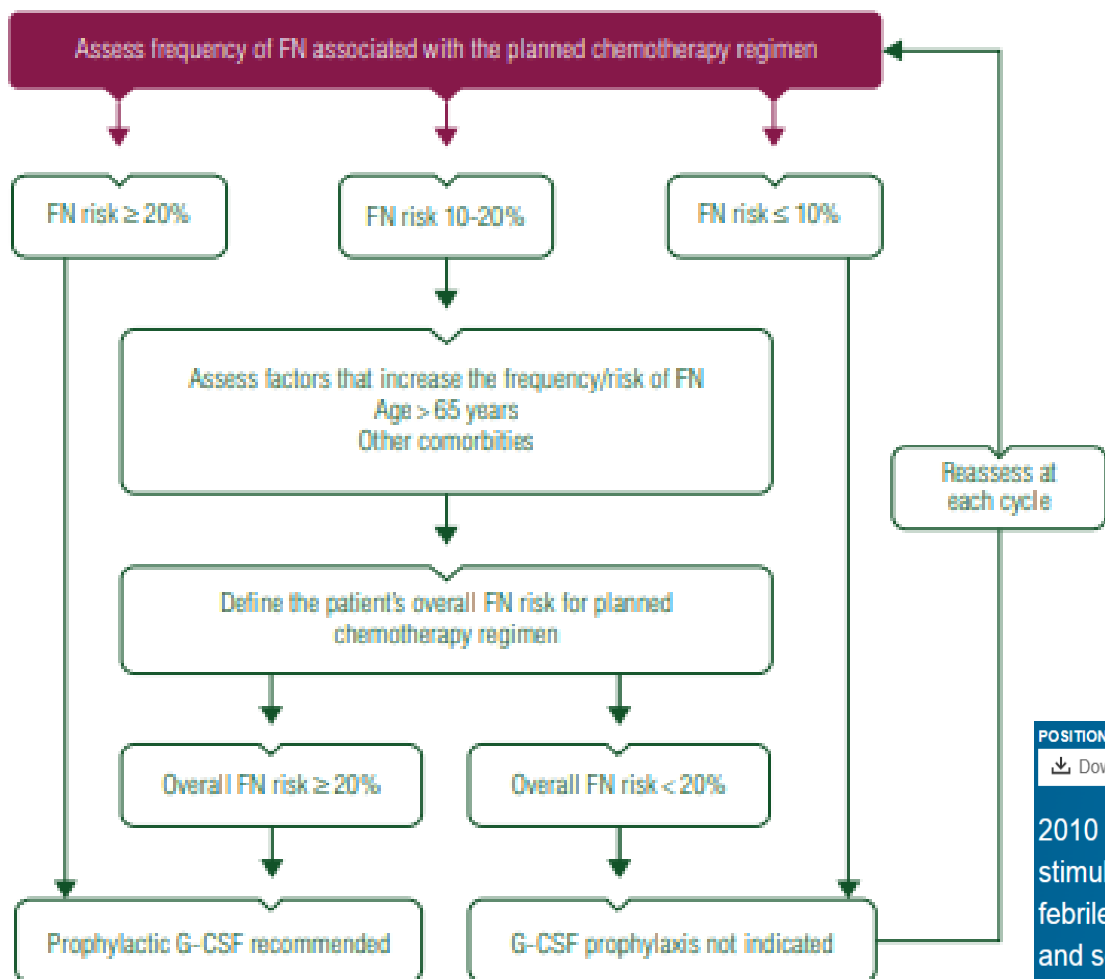
Colony-Stimulating Factors for Febrile Neutropenia during Cancer Therapy

This article has been corrected. [VIEW THE CORRECTION](#)

Authors: Charles L. Bennett, M.D., Ph.D., Benjamin Djulbegovic, M.D., Ph.D., LeAnn B. Norris, Pharm.D., and James O. Armitage, M.D. [Author Info & Affiliations](#)

indications for primary prophylaxis of FN with G-CSF

Several meta-analyses indicate that primary prophylaxis with G-CSF (i.e. G-CSF administered immediately after cycle 1 of ChT) reduces the risk of FN by at least 50%



Most guidelines recommend that G-CSF be administered prophylactically if the risk of FN is >20% for all planned cycles of treatment

Algorithm to decide primary prophylactic granulocyte colony-stimulating factor usage, adapted from European Organisation for Research and Treatment of Cancer guidelines. FN, febrile neutropaenia; G-CSF, granulocyte colony-stimulating factor.

POSITION PAPER | VOLUME 47, ISSUE 1, P8-32, JANUARY 2011

[Download Full Issue](#)

2010 update of EORTC guidelines for the use of granulocyte-colony stimulating factor to reduce the incidence of chemotherapy-induced febrile neutropenia in adult patients with lymphoproliferative disorders and solid tumours

M.S. Aapro ¹, J. Bohlius ², D.A. Cameron ³, J. Walewski ⁴, Damien C. Weber ⁵, C. Zielinski ⁶

EJC
EUROPEAN JOURNAL OF CANCER



Prophylaxis

G-CSF

Filgrastim 5 µg/kg/day per subcutaneous injection starting 24-72 hours after chemotherapy until count recovery

Pegfilgrastim 6 mg subcutaneous once 24 hours after discontinuation of chemotherapy

CLINICAL THERAPEUTICS



Colony-Stimulating Factors for Febrile Neutropenia during Cancer Therapy

This article has been corrected. [VIEW THE CORRECTION](#)

Authors: Charles L. Bennett, M.D., Ph.D., Benjamin Djulbegovic, M.D., Ph.D., LeAnn B. Norris, Pharm.D., and James O. Armitage, M.D. [Author Info & Affiliations](#)

Published March 21, 2013 | N Engl J Med 2013;368:1131-1139 | DOI: 10.1056/NEJMct1210890 | VOL. 368 NO. 12

Antimicrobial Prophylaxis and Outpatient Management of Fever and Neutropenia in Adults Treated for Malignancy: American Society of Clinical Oncology Clinical Practice Guideline



Authors: Christopher R. Flowers, Jerome Seidenfeld , Eric J. Bow, Clare Korten, Charise Gleason, Douglas K. Hawley, Nicole M. Kuderer, Amelia A. Langston, Kieren A. Marr, Kenneth V.I. Rolston, and Scott D. Ramsey | [AUTHORS INFO & AFFILIATIONS](#)

Publication: Journal of Clinical Oncology • Volume 31, Number 6 • <https://doi.org/10.1200/JCO.2012.45.8661>

Antibacterial Prophylaxis Indications

- *Anticipate ANC ≤ 100 cells/mm³ for > 7 days
(most commonly HCT recipients and patients undergoing induction therapy for acute leukemia)*

Levofloxacin 500 – 750 mg po/iv q24h


or

Ciprofloxacin 500-750 mg po/iv x 2/day

ORIGINAL ARTICLE

f X in

Levofloxacin to Prevent Bacterial Infection in Patients with Cancer and Neutropenia

Authors: Giampaolo Bucaneve, M.D., Alessandra Micozzi, M.D., Francesco Menichetti, M.D., Pietro Martino, M.D., M. Stella Dionisi, M.D., Giovanni Martinelli, M.D., Bernardino Allione, M.D., , for the Gruppo Italiano Malattie Ematologiche dell'Adulto (GIMEMA) Infection Program* | [Author Info & Affiliations](#)

Published September 8, 2005 | N Engl J Med 2005;353:977-987 | DOI: 10.1056/NEJMoa044097

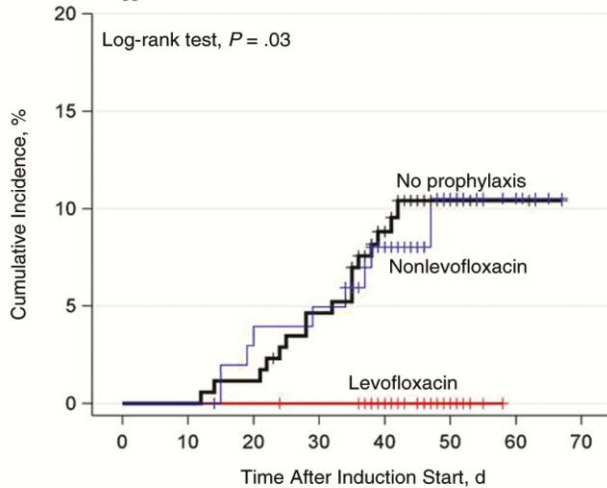


The NEW ENGLAND
JOURNAL of MEDICINE

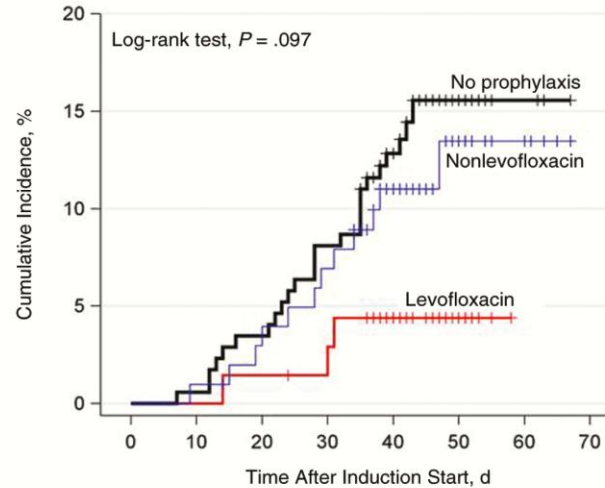
Kaplan-Meier analysis of time to infectious complications during induction for each antibiotic prophylaxis group.



A Clostridium difficile infection



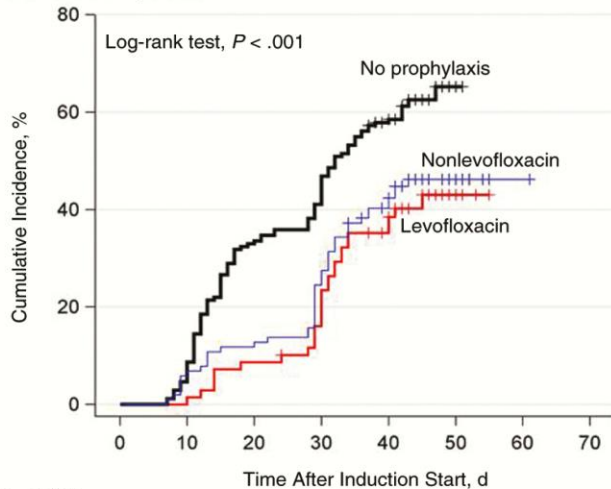
B Enterocolitis



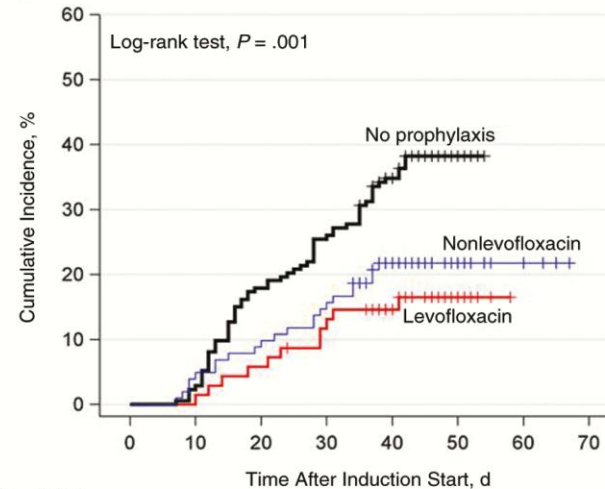
No. at risk	0	10	20	30	40	50	60	70
Levofloxacin	69	69	69	68	57	9	0	0
No prophylaxis	173	173	171	164	137	22	3	0
Nonlevofloxacin	102	102	98	96	83	25	5	0

No. at risk	0	10	20	30	40	50	60	70
Levofloxacin	69	69	68	67	55	9	0	0
No prophylaxis	173	172	167	159	132	20	3	0
Nonlevofloxacin	102	101	98	94	80	24	5	0

C Febrile neutropenia



D Likely bacterial infection



No. at risk	0	10	20	30	40	50	60	70
Levofloxacin	69	69	63	57	40	6	0	0
No prophylaxis	173	165	116	102	65	4	0	0
Nonlevofloxacin	102	96	90	77	57	11	1	0

No. at risk	0	10	20	30	40	50	60	70
Levofloxacin	69	69	65	60	51	9	0	0
No prophylaxis	173	169	142	129	98	9	0	0
Nonlevofloxacin	102	98	93	87	70	19	4	0

Anti-candidal Prophylaxis

Indications

Acute Leukemia undergoing intensive induction or reinduction

Allogenic HCT recipients until engraftment when mold-active prophylaxis is not felt to be indicated

Autologous HCT recipients with mucositis pre-engraftment

**Fluconazole 400 mg
po/iv q24h**

*until recovery of
neutropenia*

FREE ACCESS | ASCO SPECIAL ARTICLES | January 14, 2013

Journal of Clinical Oncology
An American Society of Clinical Oncology Journal

**Antimicrobial Prophylaxis and Outpatient Management of
Fever and Neutropenia in Adults Treated for Malignancy:
American Society of Clinical Oncology Clinical Practice
Guideline**

Authors: Christopher R. Flowers, Jerome Seidenfeld , Eric J. Bow, Clare Karten, Charise Gleason, Douglas K. Hawley, Nicole M. Kuderer, Amelia A. Langston, Kieren A. Marr, Kenneth V.I. Rolston, and Scott D. Ramsey | [AUTHORS INFO & AFFILIATIONS](#)

Anti-candidal Prophylaxis

Caspofungin, Micafungin or Anidulafungin

Itraconazole 200 mg po x 2/day

Posaconazole 300 mg po daily

Voriconazole 200 mg po/iv x 2/day

Liposomal Amphotericin B 3-5 mg/kg/day iv

FREE ACCESS | ASCO SPECIAL ARTICLES | January 14, 2013

Journal of Clinical Oncology
An American Society of Clinical Oncology Journal

Antimicrobial Prophylaxis and Outpatient Management of Fever and Neutropenia in Adults Treated for Malignancy: American Society of Clinical Oncology Clinical Practice Guideline

Authors: Christopher R. Flowers, Jerome Seidenfeld , Eric J. Bow, Clare Karten, Charise Gleason, Douglas K. Hawley, Nicole M. Kuderer, Amelia A. Langston, Kieren A. Marr, Kenneth V.I. Rolston, and Scott D. Ramsey | [AUTHORS INFO & AFFILIATIONS](#)

Anti-aspergillus Prophylaxis

Indications

Acute Myeloid Leukemia with neutropenia

Myelodysplastic Syndromes with neutropenia

Allogenic HCT recipients

Posaconazole 300 mg po daily

ORIGINAL ARTICLE



The NEW ENGLAND
JOURNAL of MEDICINE

Posaconazole vs. Fluconazole or Itraconazole Prophylaxis in Patients with Neutropenia

Authors: Oliver A. Cornely, M.D., Johan Maertens, M.D., Drew J. Winston, M.D., John Perfect, M.D., Andrew J. Ullmann, M.D., Thomas J. Walsh, M.D., David Helfgott, M.D., [+6](#), and David Angulo-Gonzalez, M.D.* [Author Info & Affiliations](#)

Anti-aspergillus Prophylaxis

Voriconazole 200 mg x 2/die

Itraconazole 200 mg x 2/die


Inhaled Amphotericin B 12.5 mg on 2 consecutive days/week

ORIGINAL ARTICLE



The NEW ENGLAND
JOURNAL of MEDICINE

Posaconazole vs. Fluconazole or Itraconazole Prophylaxis in Patients with Neutropenia

Authors: Oliver A. Cornely, M.D., Johan Maertens, M.D., Drew J. Winston, M.D., John Perfect, M.D., Andrew J. Ullmann, M.D., Thomas J. Walsh, M.D., David Helfgott, M.D., , and David Angulo-Gonzalez, M.D.* [Author Info & Affiliations](#)

Published January 25, 2007 | N Engl J Med 2007;356:348-359 | DOI: 10.1056/NEJMoa061094 | [VOL. 356 NO. 4](#)

Antiviral Prophylaxis

Indications

HSV – seropositive patients

undergoing allogenic HCT

or

Induction for acute leukemia

**Aciclovir 800 mg po x 2/die or Aciclovir 400 mg po x 4/die
or
Valacyclovir 500 mg po x 3/die or Famciclovir 250 mg po x 2/die**

JOURNAL ARTICLE GUIDELINES

Executive Summary: Clinical Practice Guideline for the Use of Antimicrobial Agents in Neutropenic Patients with Cancer: 2010 Update by the Infectious Diseases Society of America 

Alison G. Freifeld, Eric J. Bow, Kent A. Sepkowitz, Michael J. Boeckh, James I. Ito, Craig A. Mullen, Issam I. Raad, Kenneth V. Rolston, Jo-Anne H. Young, John R. Wingard

Clinical Infectious Diseases, Volume 52, Issue 4, 15 February 2011, Pages 427–431,

Antiviral Prophylaxis

Indications

Prevention of CMV in HSCT patients receiving an allogeneic transplant

Risk factors:

R+ D-

T-cell depleted or cord blood transplants

GVHD

Prevention strategies include both prophylaxis and preemptive therapy

Antiviral Prophylaxis

Letemovir 480 mg iv or po q24h

*Valganciclovir 900 mg po q24h
(beginning post-engraftment)*

ORIGINAL ARTICLE



The NEW ENGLAND
JOURNAL of MEDICINE

Letemovir Prophylaxis for Cytomegalovirus in Hematopoietic-Cell Transplantation

Authors: Francisco M. Marty, M.D., Per Ljungman, M.D., Ph.D., Roy F. Chemaly, M.D., M.P.H., Johan Maertens, M.D., Ph.D., Sanjeet S. Dadwal, M.D., Rafael F. Duarte, M.D., Ph.D., Shariq Haider, M.D., D.T.M.&H., [+15](#), and Cyrus Badshah, M.D., Ph.D. [Author Info & Affiliations](#)

Antiviral Prophylaxis

Letermovir was approved for the prevention of CMV infection following stem cell transplant

It is not active against HSV or VZV so addition of another agent for HSV/VZV prophylaxis is required

ORIGINAL ARTICLE



The NEW ENGLAND
JOURNAL of MEDICINE

Letermovir Prophylaxis for Cytomegalovirus in Hematopoietic-Cell Transplantation

Authors: Francisco M. Marty, M.D., Per Ljungman, M.D., Ph.D., Roy F. Chemaly, M.D., M.P.H., Johan Maertens, M.D., Ph.D., Sanjeet S. Dadwal, M.D., Rafael F. Duarte, M.D., Ph.D., Shariq Haider, M.D., D.T.M.&H., [+15](#), and Cyrus Badshah, M.D., Ph.D. [Author Info & Affiliations](#)

Antiviral Prophylaxis

Valganciclovir prophylaxis was not superior to a preemptive strategy in preventing CMV infection post-HCT

Valganciclovir prophylaxis was associated with increase in use of hematopoietic growth factors

Annals of Internal Medicine

Original Research | 6 January 2015

Valganciclovir for the Prevention of Complications of Late Cytomegalovirus Infection After Allogeneic Hematopoietic Cell Transplantation: A Randomized Trial

Authors: Michael Boeckh, MD, W. Garrett Nichols, MD, MS, Roy F. Chemaly, MD, MPH, Genovefa A. Papanicolaou, MD, John R. Wingard, MD, Hu Xie, MS, Karen L. Syrjala, PhD, Mary E.D. Flowers, MD, Terry Stevens-Ayers, MS, Keith R. Jerome, MD, PhD, and Wendy Leisenring, ScD | [AUTHOR, ARTICLE, & DISCLOSURE INFORMATION](#)

Journal of Antimicrobial Chemotherapy

JOURNAL ARTICLE

Drug-resistant cytomegalovirus in transplant recipients: a French cohort study ^{FREE}

Sébastien Hantz, Françoise Garnier-Geoffroy, Marie-Christine Mazon, Isabelle Garrigue, Pierre Merville, Catherine Mengelle, Lionel Rostaing, Franck Saint Marcoux, Marie Essig, Jean-Philippe Rerolle ... [Show more](#)

[Author Notes](#)

Journal of Antimicrobial Chemotherapy, Volume 65, Issue 12, December 2010, Pages

Resistance demonstrated in 5% of transplant recipients receiving prophylaxis

Antibiotic and Antifungal Prophylaxis

Antibiotic prophylaxis is commonly followed in patients with hematologic malignancies and solid tumors who receive myeloablative therapy and develop profound neutropenia as well as in patients in the early post-transplant period

***This approach has previously been shown to reduce
the risk of all-cause mortality
(RR, 0.52; 95% confidence interval [95% CI], 0.35-0.77)***

CA CANCER J CLIN 2018;68:340-355

Update on Infection Control Practices in Cancer Hospitals

Ella J. Ariza-Heredia, MD ¹; Roy F. Chemaly, MD, MPH²



Most standard-dose ChT regimens are associated with 6–8 days of neutropaenia,

FN is observed in ~8 cases per 1000 patients receiving cancer ChT.

FN is responsible for considerable morbidity as 20%–30% of patients present complications that require in-hospital management, with an overall in-hospital mortality of ~10%.

The mean cost per hospitalisation in Western countries is ~13 500E (15 000 US\$)

clinical practice guidelines

Annals of Oncology 27 (Supplement 5): v111–v118, 2016
doi:10.1093/annonc/mdw325

Management of febrile neutropaenia: ESMO Clinical Practice Guidelines[†]

J. Klastersky¹, J. de Naurois², K. Rolston³, B. Rapoport⁴, G. Maschmeyer⁵, M. Aapro⁶ & J. Herrstedt⁷ on behalf of the ESMO Guidelines Committee*

Table 1. MASCC febrile neutropaenia risk index

Characteristics	Score
Burden of illness: no or mild symptoms	5
Burden of illness: moderate symptoms	3
Burden of illness: severe symptoms	0
No hypotension (systolic BP > 90 mmHg)	5
No chronic obstructive pulmonary disease	4
Solid tumour/lymphoma with no previous fungal infection	4
No dehydration	3
Outpatient status (at onset of fever)	3
Age <60 years	2

The presence of a focal site of presumed infection (e.g. pneumonia, abscess, cellulitis) also makes the outcome worse.

Mortality varies according to the Multinational Association of Supportive Care in Cancer (MASCC) prognostic index: lower than 5% if the MASCC score is ≥ 21 , but possibly as high as 40% if the MASCC score is < 15

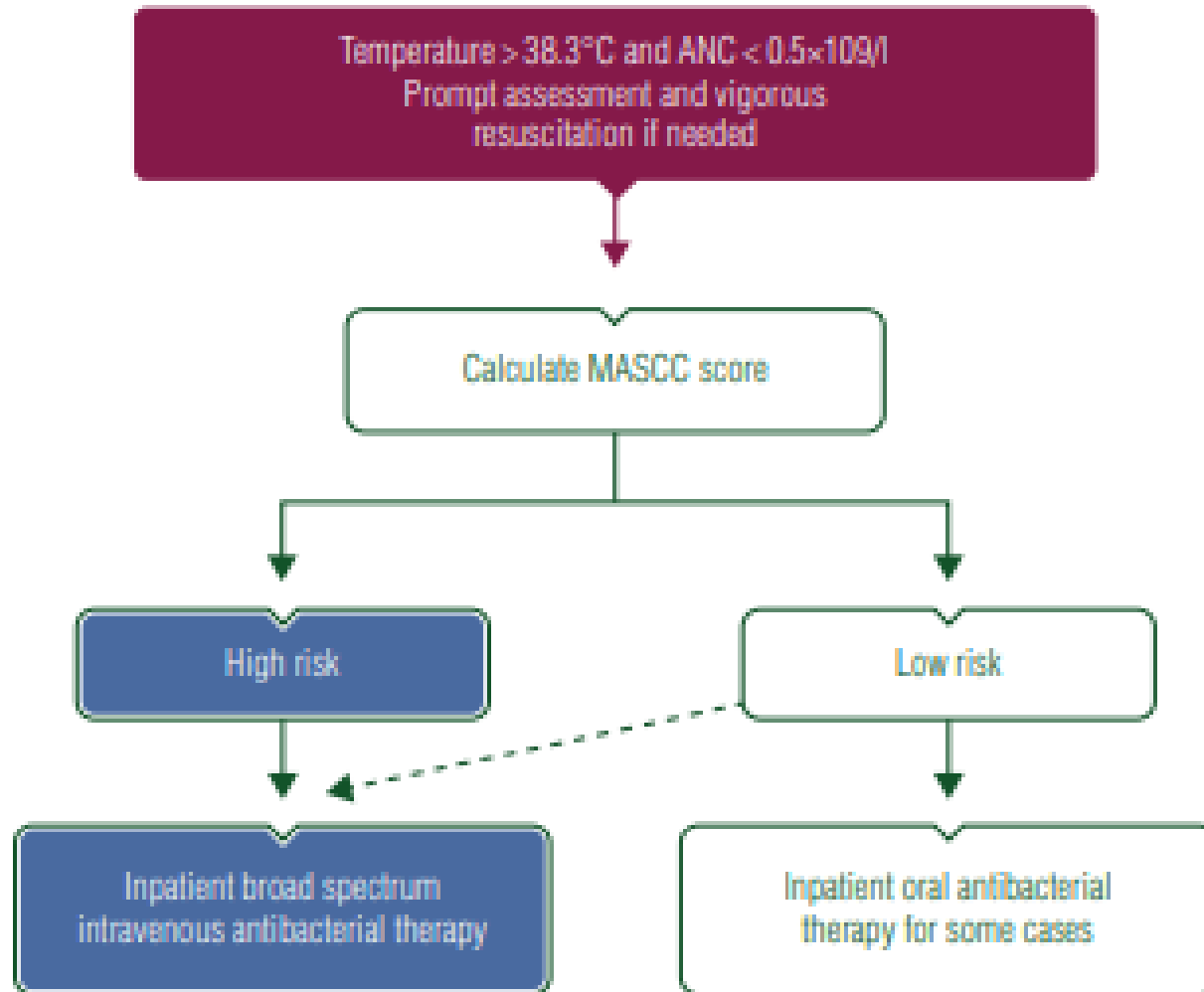
ORIGINAL REPORT | August 16, 2000

Journal of Clinical Oncology
An American Society of Clinical Oncology Journal

The Multinational Association for Supportive Care in Cancer Risk Index: A Multinational Scoring System for Identifying Low-Risk Febrile Neutropenic Cancer Patients

Authors: Jean Klastersky, Marianne Paesmans, Edward B. Rubenstein, Michael Boyer, Linda Elting, Ronald Feld, James Gallagher, Jorn Herrstedt, Bernardo Rapoport, Kenneth Rolston, and James Talcott for the Study Section on Infections of Multinational Association for Supportive Care in Cancer | [AUTHORS INFO & AFFILIATIONS](#)

Initial management of febrile neutropaenia



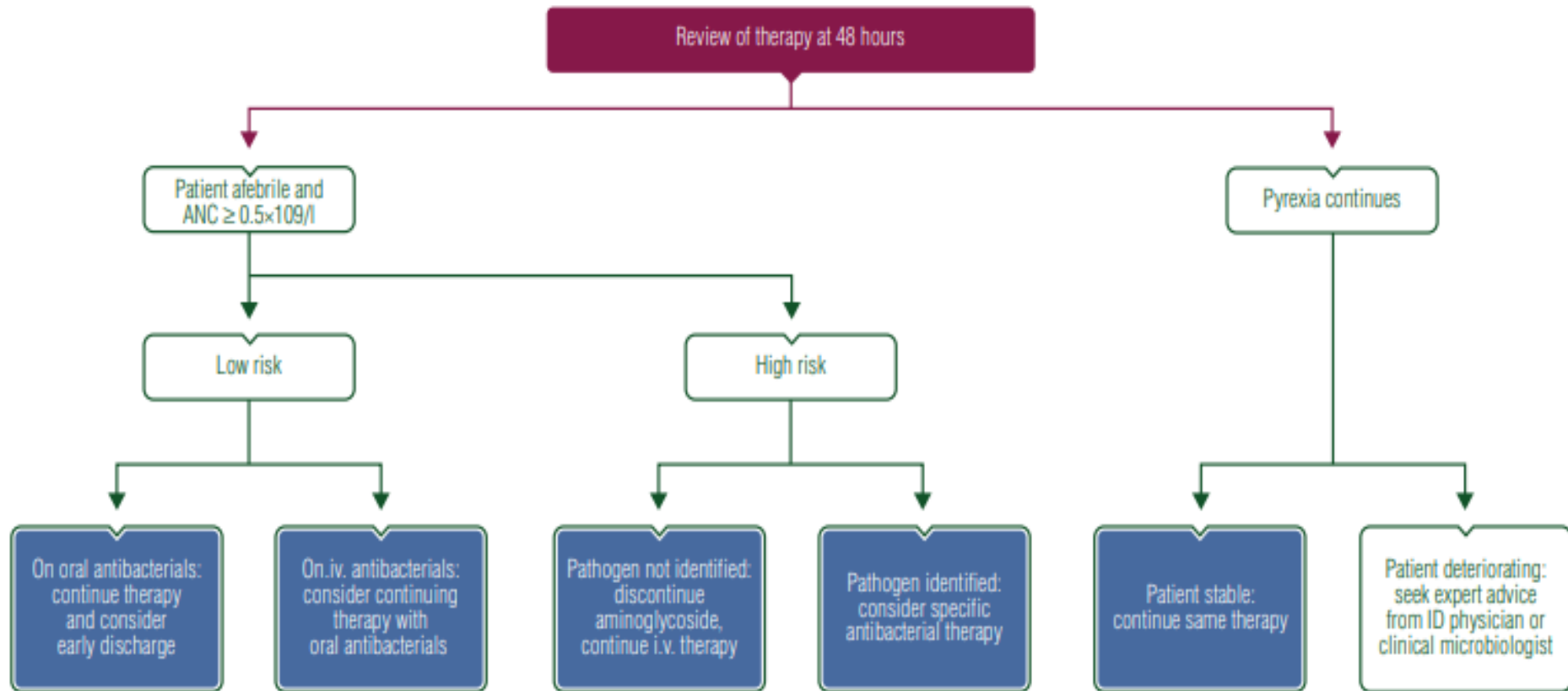
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Assessment of response and subsequent management



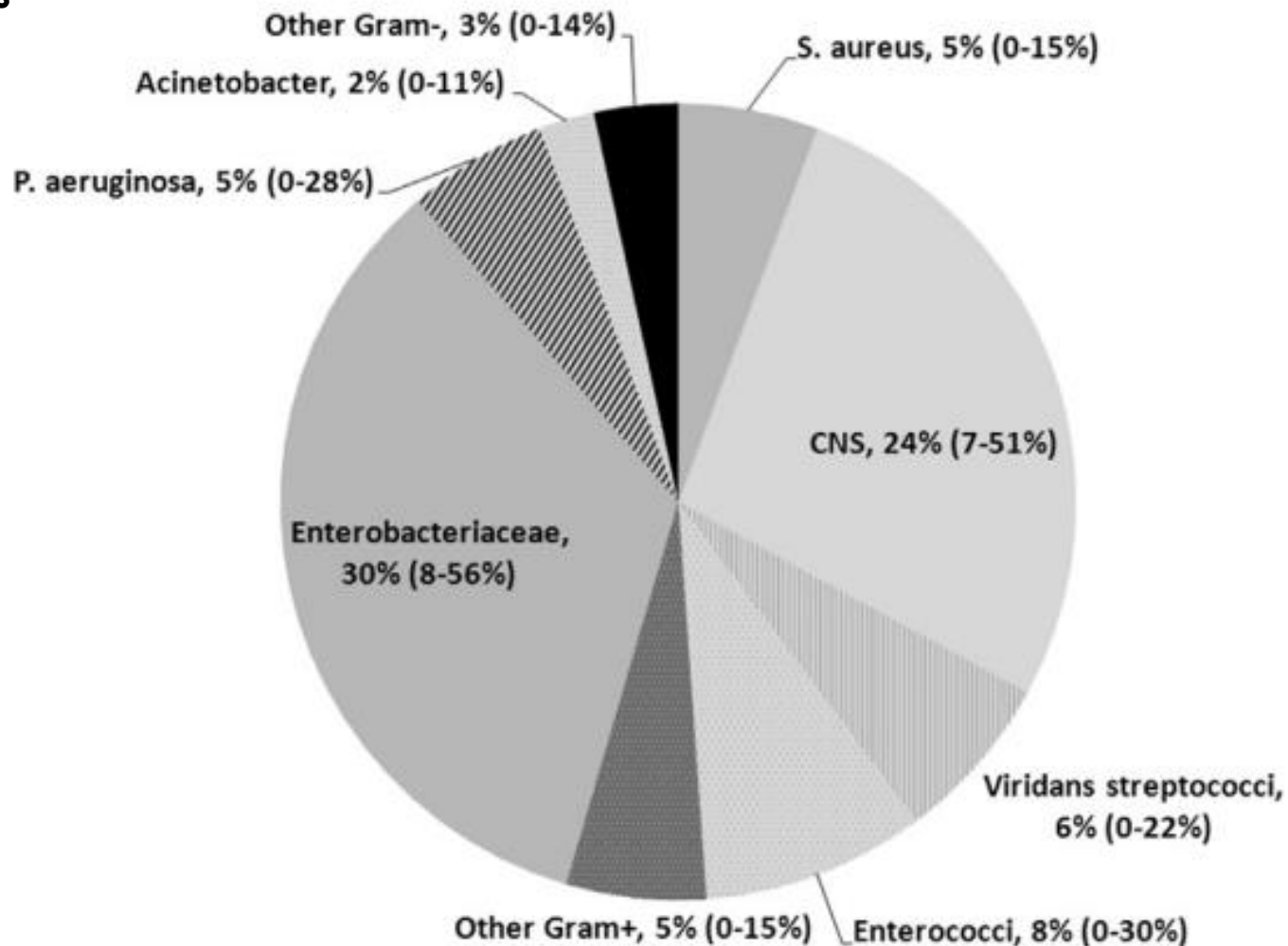
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Spectrum of pathogens in infections of neutropenic patients



RESEARCH ARTICLE | VOLUME 68, ISSUE 4, P321-331, APRIL 2014

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JOURNAL OF
INFECTION

Aetiology and resistance in bacteraemias among adult and paediatric haematology and cancer patients

Małgorzata Mikulska • Claudio Viscoli • Christina Orasch • ... Catherine Cordonnier • Murat Akova
on behalf of the Fourth European Conference on Infections in Leukemia Group (ECIL-4), a joint venture of EBMT, EORTC, ICHS, ELN and ESGICH/ESCMID •

Aetiology of bacteraemias (median prevalence with range) reported in the ECIL-4 questionnaire survey.
Notes: CNS, coagulase negative staphylococci

The current spectrum of infection in cancer patients with chemotherapy related neutropenia

Review | Published: 23 August 2013



The first administration of therapy should be given in the hospital within 1 h from the admission of a patient with FN.

Delay in antibiotic administration has been associated with significant prolongation of the hospital stay and increased mortality.


As already mentioned, the spectrum of infection in cancer patients is different from place to place and changes over time; therefore, paying attention to local epidemiology is crucial

clinical practice guidelines

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- 
- 1 Note the presence of indwelling i.v. catheters
 - 2 Symptoms or signs suggesting an infection focus:
 - Respiratory system
 - Gastrointestinal tract
 - Skin
 - Perineal region/genitourinary discharges
 - Oropharynx
 - Central nervous system
 - 3 Knowledge of previous positive microbiology results by checking clinical records
 - 4 Routine investigations:
 - Urgent blood testing to assess bone marrow, renal and liver function
 - Coagulation screen
 - C-reactive protein
 - Blood cultures (minimum of two sets) including cultures from indwelling i.v. catheter
 - Urinalysis and culture^a
 - Sputum microscopy and culture^a
 - Stool microscopy and culture^a
 - Skin lesion (aspirate/biopsy/swab)
 - Chest radiograph
 - 5 Further investigations (profound/prolonger neutropaenia/following allografts)
 - High-resolution chest CT (if pyrexial despite 72 h of appropriate antibiotics)
 - Bronchoalveolar lavage

initial assessment and investigations

clinical practice guidelines

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Spectrum of pathogens in infections of neutropenic patients

Pathogen and studies	Type of resistance	Adults median rate of resistance (range)	Children median rate of resistance (range)
<i>S. aureus</i>	MRSA	56% (18–100%) ^a	0% (0–26%) ^b
CNS	MR-CNS	80% (33–100%) ^c	38% and 39% ^d
Enterococci	VRE	23% (0–50%) ^e	0% ^f
Gram-negatives	Fluoroquinolone-resistant	41% (18–74%) ^g	7% and 32% ^{t,h}
Gram-negatives	Carbapenem-resistant	20% (11–72%) ⁱ	9% and 10% ^h
Gram-negatives	Aminoglycoside-resistant	28% (6–41%) ^j	Gentamicin-resistant 26% (25–28%) ^k
Gram-negatives	Ceftazidime-resistant	43% (17–45%) ^l	18% and 27% ^h
Enterobacteriaceae	ESBL-producing	34% (16–44%) ^m + 42% of <i>E. coli</i> ⁿ	18% ^o
Enterobacteriaceae	Fluoroquinolone-resistant	56% (28–87%) ^p + 63% of <i>E. coli</i> ⁿ	4% ^o
<i>P. aeruginosa</i>	Fluoroquinolone-resistant	53% (7–72%) ^q	18% ^r
<i>P. aeruginosa</i>	Carbapenem-resistant	44% (3–66%) ^s	25% ^r

Median rates of resistance to different antibiotics in pathogens causing bacteraemias in haematology-oncology adults and children, based upon literature reports



Pathogen	No. (%) of patients, by year					
	1995 (n = 390)	1996 (n = 556)	1997 (n = 508)	1998 (n = 451)	1999 (n = 336)	2000 (n = 411)
Gram-positive organisms	241 (61.8)	339 (61.0)	267 (52.6)	251 (55.7)	201 (59.8)	312 (75.9)
Gram-negative organisms	84 (21.5)	154 (27.7)	145 (28.5)	164 (36.4)	100 (29.8)	59 (14.4)
Anaerobic organisms	7 (1.8)	13 (2.3)	48 (9.4)	10 (2.2)	8 (2.4)	6 (1.5)
Fungi	58 (14.9)	50 (9.0)	48 (9.4)	26 (5.8)	27 (8.0)	34 (8.3)

Pathogen	No. (%) of patients			OR (95% CI)	P ^a
	ANC, neutrophils/ μ L		Total (n = 2711)		
	<1000 (n = 798)	\geq 1000 (n = 1913)			
Gram-positive organisms					
All	487 (61.0)	1152 (60.2)	1639 (60.5)	1.0 (0.87–1.23)	.7
CoNS	252 (31.6)	566 (29.6)	818 (30.2)	1.1 (0.91–1.32)	.3
<i>Staphylococcus aureus</i>	98 (12.3)	213 (11.1)	311 (11.5)	1.1 (0.86–1.45)	.4
Enterococci					
All	50 (6.3)	265 (13.9)	315 (11.6)	2.4 (1.74–3.34)	<.001
<i>Enterococcus faecalis</i>	25 (3.1)	100 (5.2)	125 (4.6)	1.7 (1.07–2.73)	.02
<i>Enterococcus faecium</i>	18 (2.3)	122 (6.4)	140 (5.2)	3.0 (1.75–5.04)	<.001
Streptococci					
All	73 (9.1)	90 (4.7)	163 (6.0)	2.0 (1.46–2.84)	<.001
VGS	23 (2.9)	15 (0.8)	38 (1.4)	1.9 (1.87–7.60)	<.001
Other	14 (1.8)	18 (0.9)	32 (1.2)	1.9 (0.88–3.99)	.1
Gram-negative organisms					
All	199 (24.9)	521 (27.2)	720 (26.6)	0.9 (0.73–1.08)	.2
<i>Escherichia coli</i>	58 (7.3)	148 (7.7)	206 (7.6)	0.9 (0.67–1.29)	.7
<i>Klebsiella</i> species	43 (5.4)	130 (6.8)	173 (6.4)	0.8 (0.54–1.13)	.2
<i>Pseudomonas aeruginosa</i>	29 (3.6)	90 (4.7)	119 (4.4)	0.8 (0.49–1.19)	.3
<i>Enterobacter</i> species	25 (3.1)	55 (2.9)	80 (3.0)	1.1 (0.66–1.81)	.8
Other Enterobacteriaceae	14 (1.8)	42 (2.2)	56 (2.1)	0.8 (0.41–1.49)	.5
Other gram-negative organisms	30 (3.8)	57 (3.0)	87 (3.2)	1.3 (0.79–2.04)	.4
Anaerobes	38 (4.8)	55 (2.9)	93 (3.4)	1.7 (1.08–2.63)	.02
Fungi					
All	74 (9.3)	185 (9.7)	259 (9.6)	0.9 (0.71–1.28)	.8
<i>Candida</i> species	69 (8.6)	161 (8.4)	230 (8.5)	1.0 (0.76–1.39)	.9
Other	5 (0.6)	23 (1.2)	28 (1.0)	0.5 (0.17–1.44)	.3

JOURNAL ARTICLE

Current Trends in the Epidemiology of Nosocomial Bloodstream Infections in Patients with Hematological Malignancies and Solid Neoplasms in Hospitals in the United States ^{FREE}

Hilmar Wisplinghoff , Harald Seifert, Richard P. Wenzel, Michael B. Edmond

NOTE. ANC, absolute neutrophil count; CoNS, coagulase-negative staphylococci; VGS, viridans group streptococci.
^a Patients with an ANC of <1000 neutrophils/ μ L vs. those with an ANC of \geq 1000 neutrophils/ μ L.

Duration of therapy

If the ANC is $\geq 0.5 \times 10^9 /l$, the patient is asymptomatic and has been afebrile for 48 h and blood cultures are negative, antibacterials can be discontinued.

If the ANC is $\leq 0.5 \times 10^9 /l$, the patient has suffered no complications and has been afebrile for 5–7 days, antibacterials can be discontinued except in certain high-risk cases with acute leukaemia and following high-dose ChT when antibacterials are often continued for up to 10 days, or until the ANC is $\geq 0.5 \times 10^9 /l$.

Patients with persistent fever despite neutrophil recovery should be assessed by an ID physician or clinical microbiologist and antifungal therapy considered.

clinical practice guidelines

Annals of Oncology 27 (Supplement 5): v111–v118, 2016
doi:10.1093/annonc/mdw325

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Fever and neutropenia: still a challenge

M J DiNubile ¹

Clinic in Oncohematologic Patients

Early detection of infections is essential in cancer patients. Clinical signs of infection might be vague, especially in neutropenic patients, but fever remains an early, although non-specific, sign of infection

Approximately 50–60% of patients who became febrile have an underlying infection

However, non-infectious causes of fever, including paraneoplastic etiology (neoplastic fever) and drug reactions, are not rare in cancer patients

[Home](#) > [Supportive Care in Cancer](#) > Article

Neoplastic fever: a neglected paraneoplastic syndrome

Review Article | Published: 29 April 2005



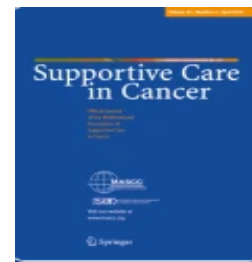
The NEW ENGLAND
JOURNAL of MEDICINE

REVIEW ARTICLE | DRUG THERAPY

Management of Fever in Patients with Cancer and Treatment-Induced Neutropenia

Author: Philip A. Pizzo [Author Info & Affiliations](#)

Published May 6, 1993 | N Engl J Med 1993;328:1323-1332 | DOI: 10.1056/NEJM199305063281808



Tumor-related leukocytosis is linked with poor prognosis in patients with lung carcinoma

I Kasuga ¹, S Makino, H Kiyokawa, H Katoh, Y Ebihara, K Ohyashiki

Moreover, laboratory markers of infection, including elevated C-reactive protein and leukocytosis, are frequently present in cancer patients

[Front Immunol](#). 2020; 11: 595835.

Published online 2020 Nov 19. doi: [10.3389/fimmu.2020.595835](https://doi.org/10.3389/fimmu.2020.595835)



C-Reactive Protein and Cancer—Diagnostic and Therapeutic Insights

[Peter C. Hart](#), ^{*} [Ibraheem M. Rajab](#), [May Alebraheem](#), and [Lawrence A. Potempa](#) ^{*}

Making the diagnosis of infection challenging.

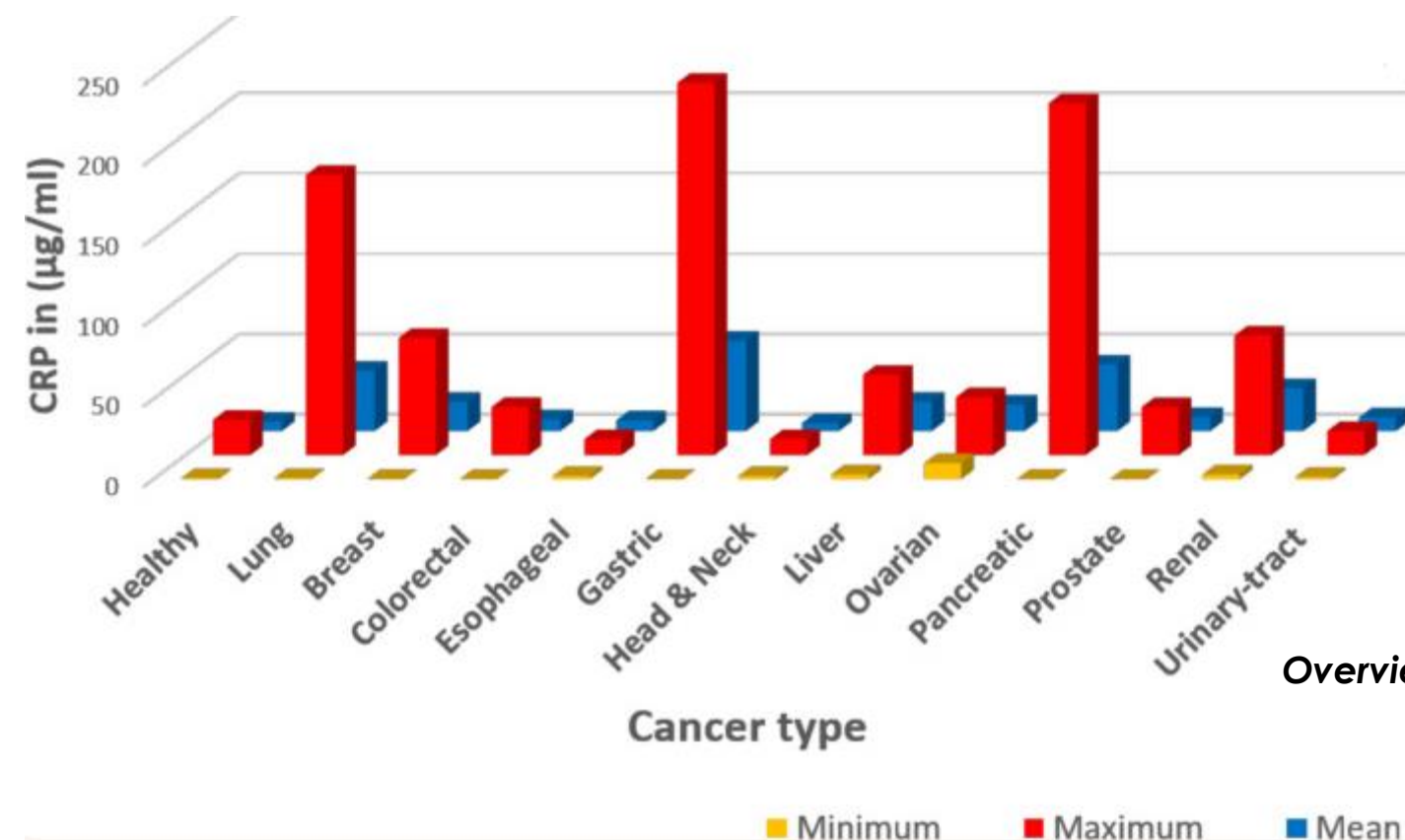
In such circumstances, additional markers such as procalcitonin can be used

C-Reactive Protein and Cancer—Diagnostic and Therapeutic Insights

Peter C. Hart, *Ibraheem M. Rajab, May Alebraheem, and Lawrence A. Potempa *



A comprehensive review of the literature on the diagnostic significance and therapeutic value of CRP blood levels in cancer proved to be problematic. Reported CRP levels varied from <1 $\mu\text{g/ml}$ to more than 175 $\mu\text{g/ml}$ and were most often reported with reference to the tissues affected with cancerous growths (e.g. lung, breast, gastrointestinal, esophageal, head and neck, sexual and reproductive organs, renal, pancreas, and blood)



Overview of CRP values reported in various cancers



It achieves a risk reduction to overestimate an infection status of 23.4%.

It supports the clinic usefulness of serum PCT dosage in febrile advanced solid tumor patients.

A PCT cut-off of 1.52 ng/dL could be helpful in the management of the antibiotic therapy preventing delays of oncologic treatments.

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scientific reports

Article | [Open access](#) | Published: 17 June 2016

Procalcitonin as diagnostic marker of infection in solid tumors patients with fever

[B. Vincenzi](#), [I. Fioroni](#), [F. Pantano](#), [S. Angeletti](#), [G. Dicuonzo](#), [A. Zoccoli](#), [D. Santini](#) & [G. Tonini](#)



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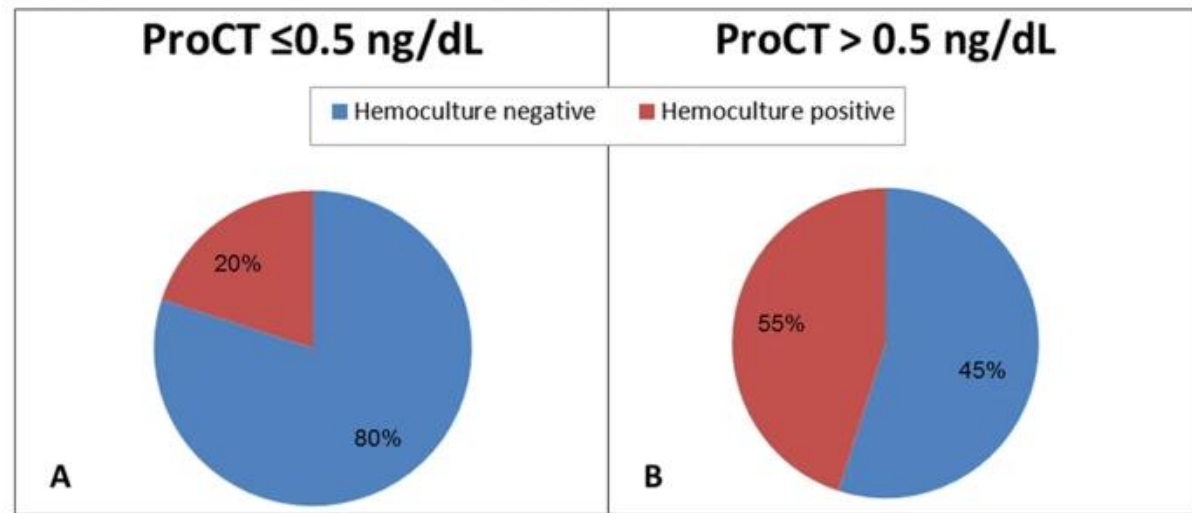
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Procalcitonin as diagnostic marker of infection in solid tumors patients with fever

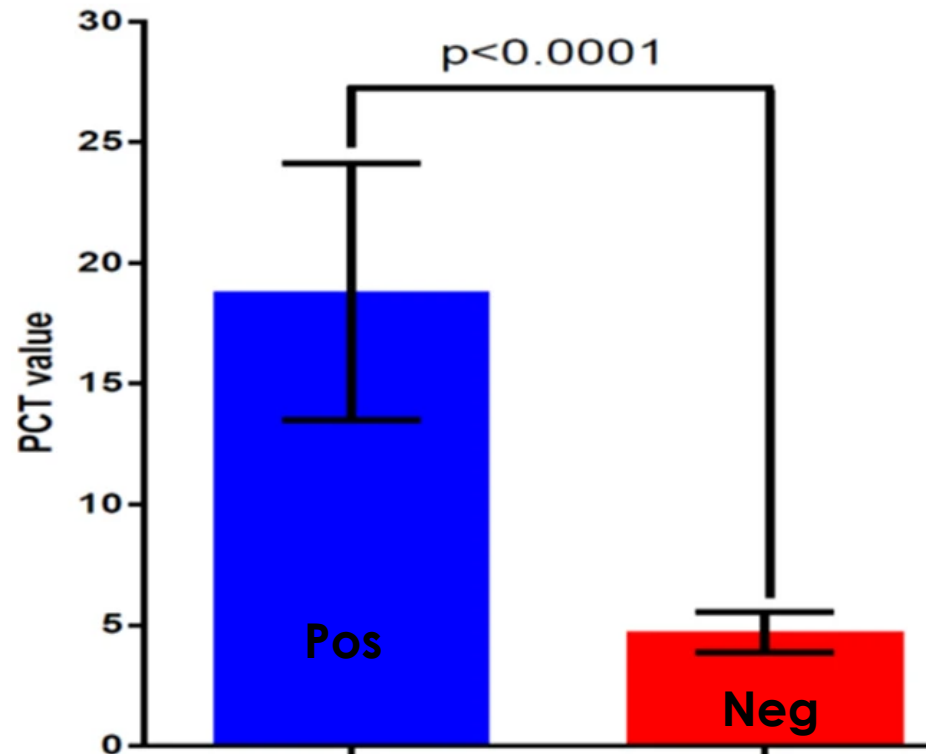
[B. Vincenzi](#), [I. Fioroni](#), [F. Pantano](#), [S. Angeletti](#), [G. Dicuonzo](#), [A. Zoccoli](#), [D. Santini](#) & [G. Tonini](#)

(A) Hemoculture stratification for PCT ≤ 0.5 ng/dL patients;

(B) Hemoculture stratification for PCT > 0.5 ng/dL patients



Non parametric Mann-Whitney test results describing difference between hemoculture positive vs. hemoculture negative population



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scientific reports

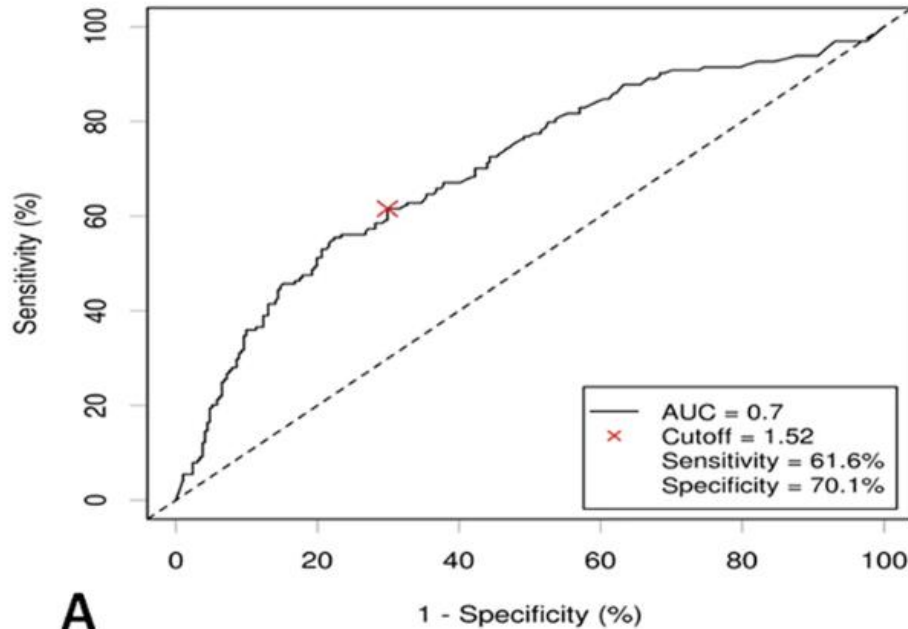
Procalcitonin as diagnostic marker of infection in solid tumors patients with fever

[B. Vincenzi](#), [I. Fioroni](#), [F. Pantano](#), [S. Angeletti](#), [G. Dicuonzo](#), [A. Zoccoli](#), [D. Santini](#) & [G. Tonini](#)

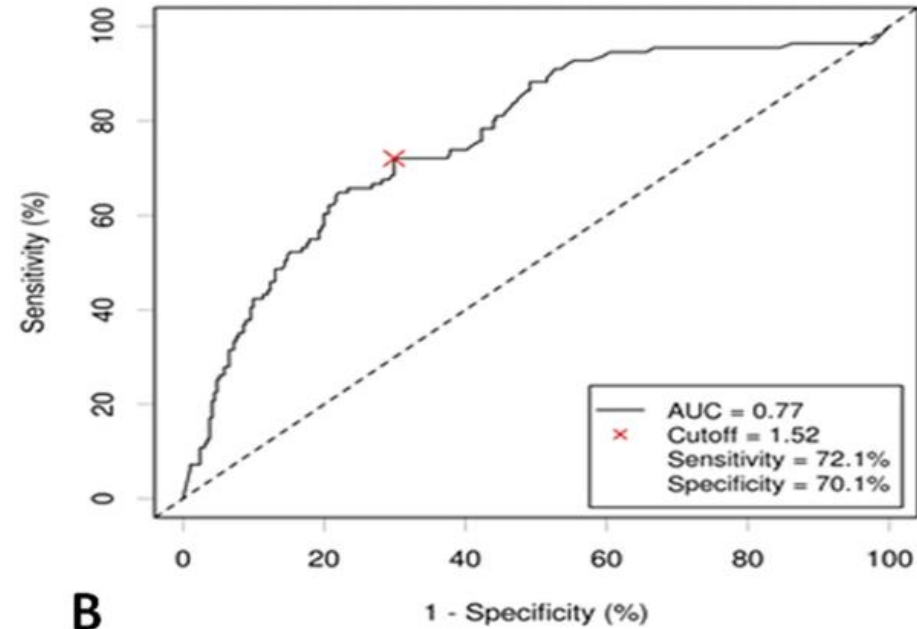
The following analysis showed that patients with a positive hemoculture mostly presented a Gram-negative bacteria infection (130 patients (72%)) and only 45 patients (24.8%) had a Gram-positive bacteria infection and only 6 patients (3.3%) showed a funginemia



PTC VALUE as positive marker for HEMOCULTURE



PCT value as positive marker for GRAM negative Hemoculture



(A) ROC curve for various cut-off levels of PCT in differentiating patients with positive and negative hemoculture;

(B) ROC curve analysis of sensitivity and specificity for the Gram-negative bacteria patients' group.

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Procalcitonin as diagnostic marker of infection in solid tumors patients with fever

[B. Vincenzi](#), [I. Fioroni](#), [F. Pantano](#), [S. Angeletti](#), [G. Dicuonzo](#), [A. Zoccoli](#), [D. Santini](#) & [G. Tonini](#)

Central Venous Catheters

Central venous catheters (CVCs) are widely used in cancer patients and offer benefits to those who receive chemotherapy.

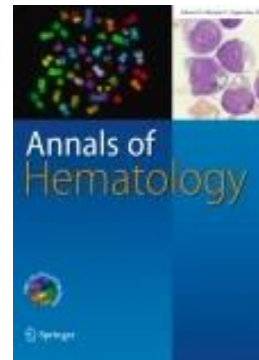
However, the presence of CVCs is considered a risk factor for infections in cancer patients and may affect the etiology of bacteremia

The incidence of CVC-associated bloodstream infections in cancer patients is estimated to be 0.5–10 per 1000 CVC days, with mortality ranging from 12% to 40%

[Home](#) > [Annals of Hematology](#) > Article

Central venous catheter–related infections in hematology and oncology: 2020 updated guidelines on diagnosis, management, and prevention by the Infectious Diseases Working Party (AGIHO) of the German Society of Hematology and Medical Oncology (DGHO)

Original Article | [Open access](#) | Published: 30 September 2020



Rates of infection for single-lumen versus multilumen central venous catheters: A meta-analysis

Dezfulian, Cameron MD; Lavelle, James MD; Nallamotheu, Brahmajee K. MD, MPH; Kaufman, Samuel R. MA; Saint, Sanjay MD, MPH

[Author Information](#)

Critical Care Medicine 31(9):p 2385-2390, September 2003. | DOI: 10.1097/01.CCM.0000084843.31852.01



Central Venous Catheters

A recent meta-analysis found a significantly higher risk of infectious complications in the PICC groups (RR 3.43; 95% CI 2.58–4.56; P < 0.05).

Both the local infections of punctures and catheter-related infections were more frequent in patients with PICCs

In addition, infection complications are more frequent in multi-lumen CVCs than in single-lumen CVCs

Complications and Costs of Peripherally Inserted Central Venous Catheters Compared With Implantable Port Catheters for Cancer Patients

A Meta-analysis

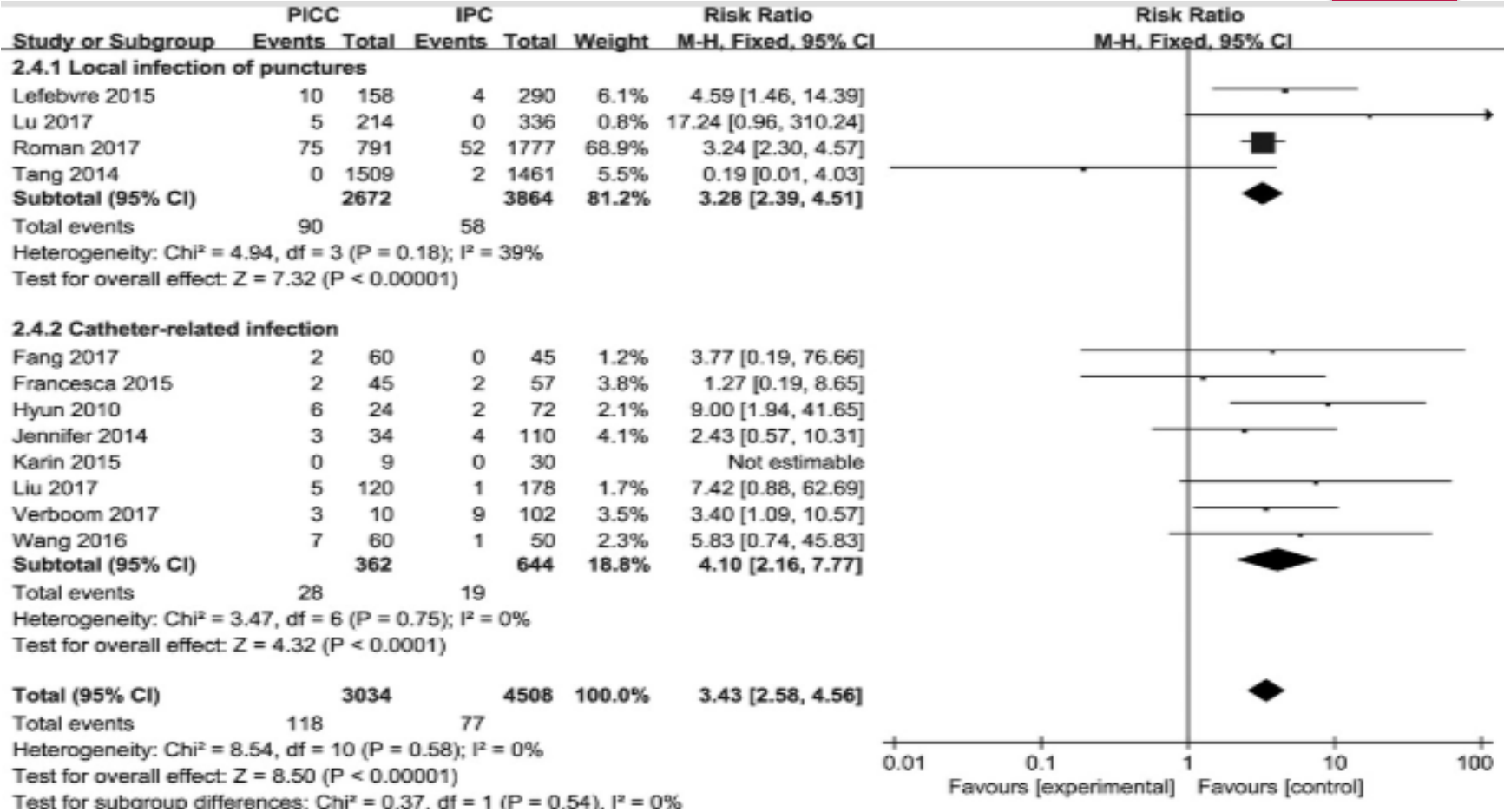
Pu, Ya-Lou MS, RN; Li, Zhuang-Shuang MS, RN; Zhi, Xiao-Xu MS, RN; Shi, Yi-An MMedSc, RN; Meng, Ai-Feng BSN, RN; Cheng, Fang MS, RN; Ali, Ali MD; Li, Cheng MS, RN; Fang, Hong MS, RN; Wang, Cheng MD

[Author Information](#)

Cancer Nursing 43(6):p 455-467, 11/12 2020. |



Forest plot of meta-analysis comparing the infection complications between PICC and IPC



Complications and Costs of Peripherally Inserted Central Venous Catheters Compared With Implantable Port Catheters for Cancer Patients

A Meta-analysis

Pu, Ya-Lou MS, RN; Li, Zhuang-Shuang MS, RN; Zhi, Xiao-Xu MS, RN; Shi, Yi-An MMedSc, RN; Meng, Ai-Feng BSN, RN; Cheng, Fang MS, RN; Ali, Ali MD; Li, Cheng MS, RN; Fang, Hong MS, RN; Wang, Cheng MD

Author Information ©

Central Venous Catheters

Catheter-related infections: diagnosis and intravascular treatment

E. Bouza   • A. Burillo • P. Muñoz

Most CVC infections originate from the skin flora (65%), catheter or catheter joints (30%), or other pathways (5%)

The most commonly detected pathogens causing CVC-related infections in cancer patients are

coagulase-negative staphylococci, followed by other gram-positive bacteria, including Staphylococcus aureus, enterococci, and streptococci

[Home](#) > [Annals of Hematology](#) > [Article](#)


Central venous catheter–related infections in hematology and oncology: 2020 updated guidelines on diagnosis, management, and prevention by the Infectious Diseases Working Party (AGIHO) of the German Society of Hematology and Medical Oncology (DGHO)

Original Article | [Open access](#) | Published: 30 September 2020

JOURNAL ARTICLE

Central Venous Catheters

Current Trends in the Epidemiology of Nosocomial Bloodstream Infections in Patients with Hematological Malignancies and Solid Neoplasms in Hospitals in the United States

Hilmar Wisplinghoff , Harald Seifert, Richard P. Wenzel, Michael B. Edmond

Clinical Infectious Diseases, Volume 36, Issue 9, 1 May 2003, Pages 1103–1110,

Consensus guidelines for the diagnosis and management of invasive candidiasis in haematology, oncology and intensive care settings, 2021

Caitlin Keighley ✉, Louise Cooley, Arthur J. Morris, David Ritchie, Julia E. Clark, Peter Boan, Leon J. Worth, the Australasian Antifungal Guidelines Steering Committee

First published: 22 December 2021 | <https://doi.org/10.1111/imj.15589> | Citations: 12

INTERNAL MEDICINE JOURNAL



Global Guideline for the Diagnosis and Management of Candidiasis: An Initiative of the ECMM in Cooperation with ISHAM



Clinical Infectious Diseases

JOURNAL ARTICLE GUIDELINES

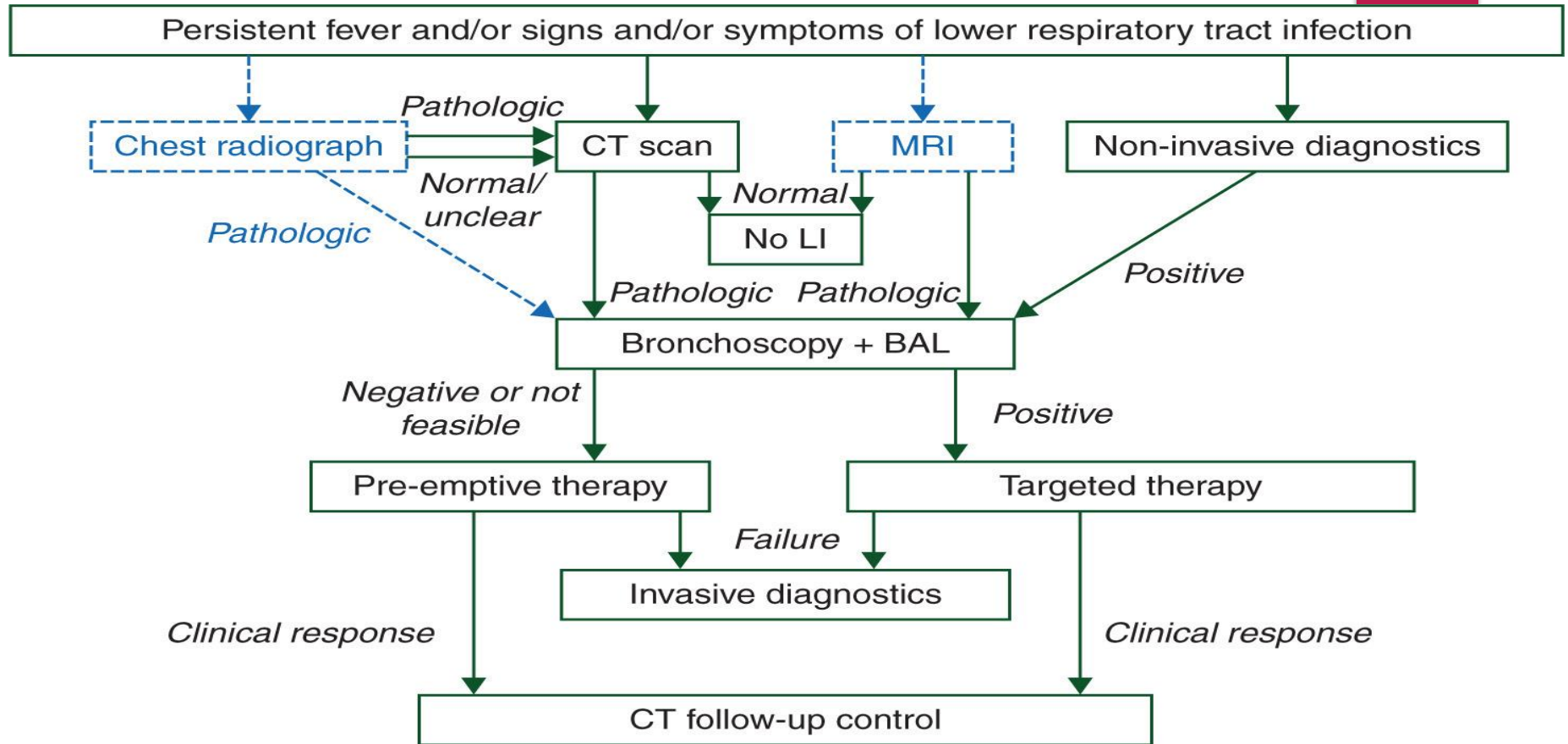
Clinical Practice Guideline for the Management of Candidiasis: 2016 Update by the Infectious Diseases Society of America □

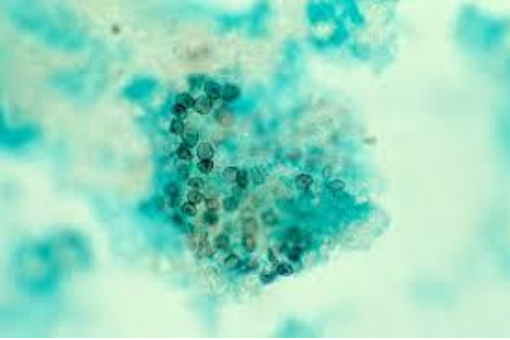
Peter G. Pappas, Carol A. Kauffman, David R. Andes, Cornelius J. Clancy, Kieren A. Marr, Luis Ostrosky-Zeichner, Annette C. Reboli, Mindy G. Schuster, Jose A. Vazquez, Thomas J. Walsh ... Show more

Clinical Infectious Diseases, Volume 62, Issue 4, 15 February 2016, Pages e1–e50, <https://doi.org/10.1093/cid/civ933>

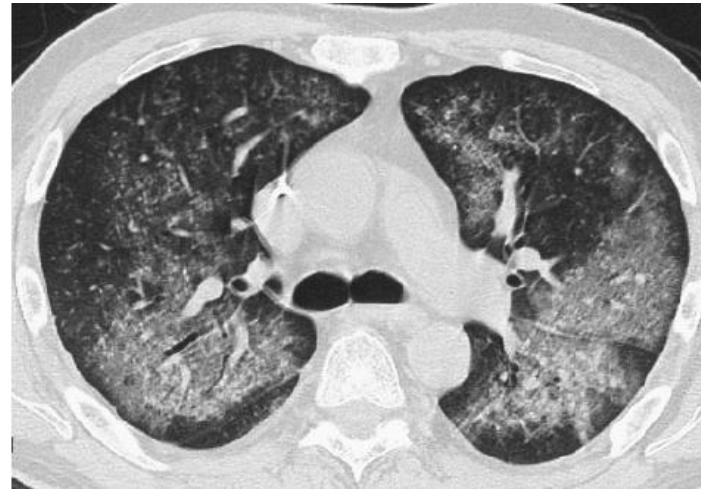
Published: 16 December 2015 Article history ▾

Diagnostic procedures and treatment of neutropenic patients with fever and suspected or proven lung infiltrates





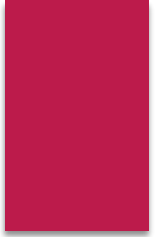
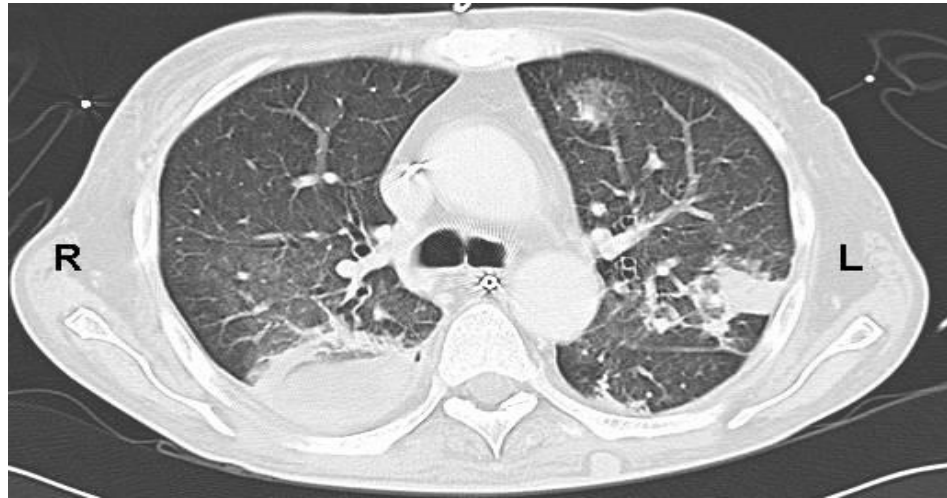
Pneumocystis jirovecii



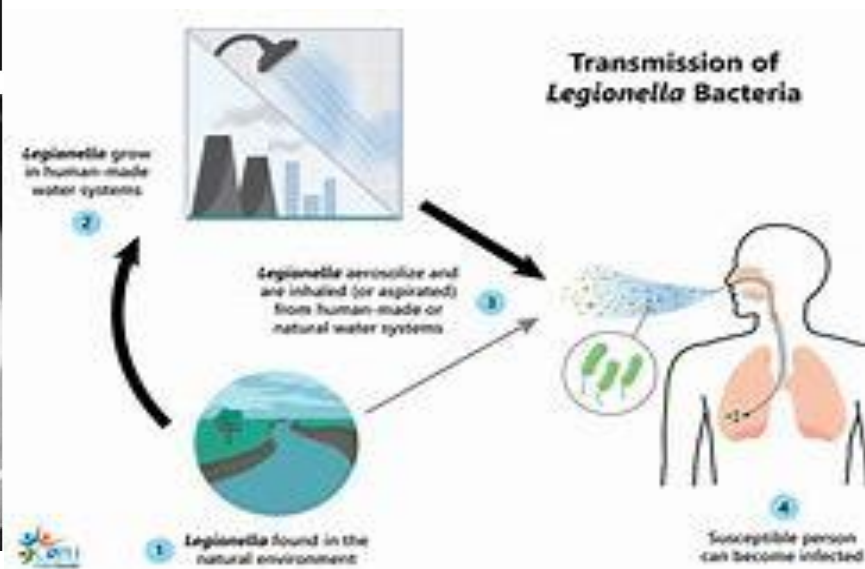
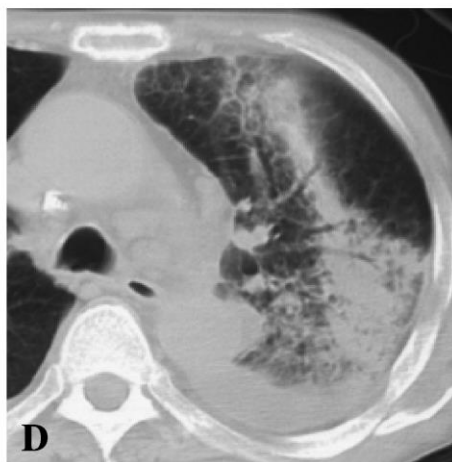
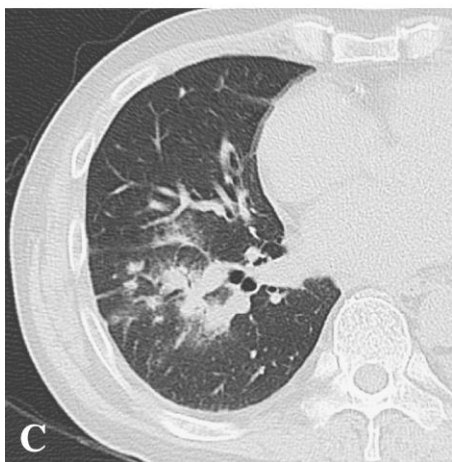
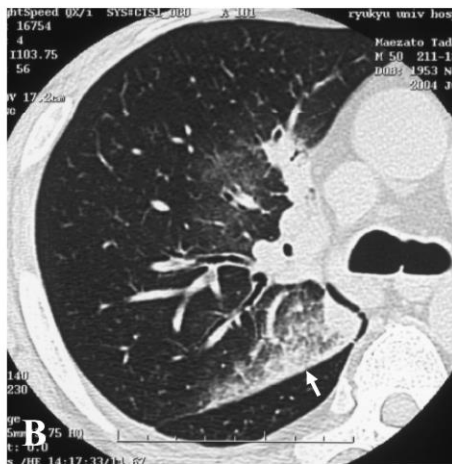
Guidelines (population)

Recommendation		CDC, NIH, HIVMA/IDSA (HIV) [12]	ECIL (hematology) [18]	American Society of Transplantation (SOT) [14]
Targeted Treatment	Population	HIV/AIDS patients with suspected/ diagnosed PcP	Hematological malignancy, solid cancer, solid organ transplant, autoimmune/inflammatory conditions with suspected/ diagnosed PcP	All SOT with suspected/ diagnosed PcP
	Duration	3 weeks	A minimum of 14 days	At least 14 days, extended to 21 days for severe cases
	Therapy ^a	Frontline: Trimethoprim/ sulfamethoxazole (15–20 mg/kg TMP; 75–100 mg/kg SMX per day) For moderate-to-severe disease (i.e. hypoxemia) adjunctive corticosteroids should be used Second line for severe disease: Primaquine and clindamycin (30 mg/ (600mgx3)) per day Pentamidine IV (4 mg/kg/day) Second line for mild/moderate disease: Dapsone (100 mg daily) + trimethoprim (15 mg daily) Atovaquone (750 mg BID)	Frontline: Trimethoprim/sulfamethoxazole (15–20 mg/ kg TMP; 75–100 mg/kg SMX per day) Second line: Primaquine and clindamycin (30 mg/(600mgx3)) per day Pentamidine IV (4 mg/kg/day)	Frontline: Trimethoprim/ sulfamethoxazole (15–20 mg/kg TMP; 75–100 mg/kg SMX per day) with TMP administered by IV every 6–8 h. For hypoxemic patients potentially in combination with 40–60 mg of prednisolone (twice daily) Second line: IV Pentamidine (Initially 4 mg/kg/day over 1–2 h) Recipients of pancreas/ islet transplants should receive an alternative second-line therapy.

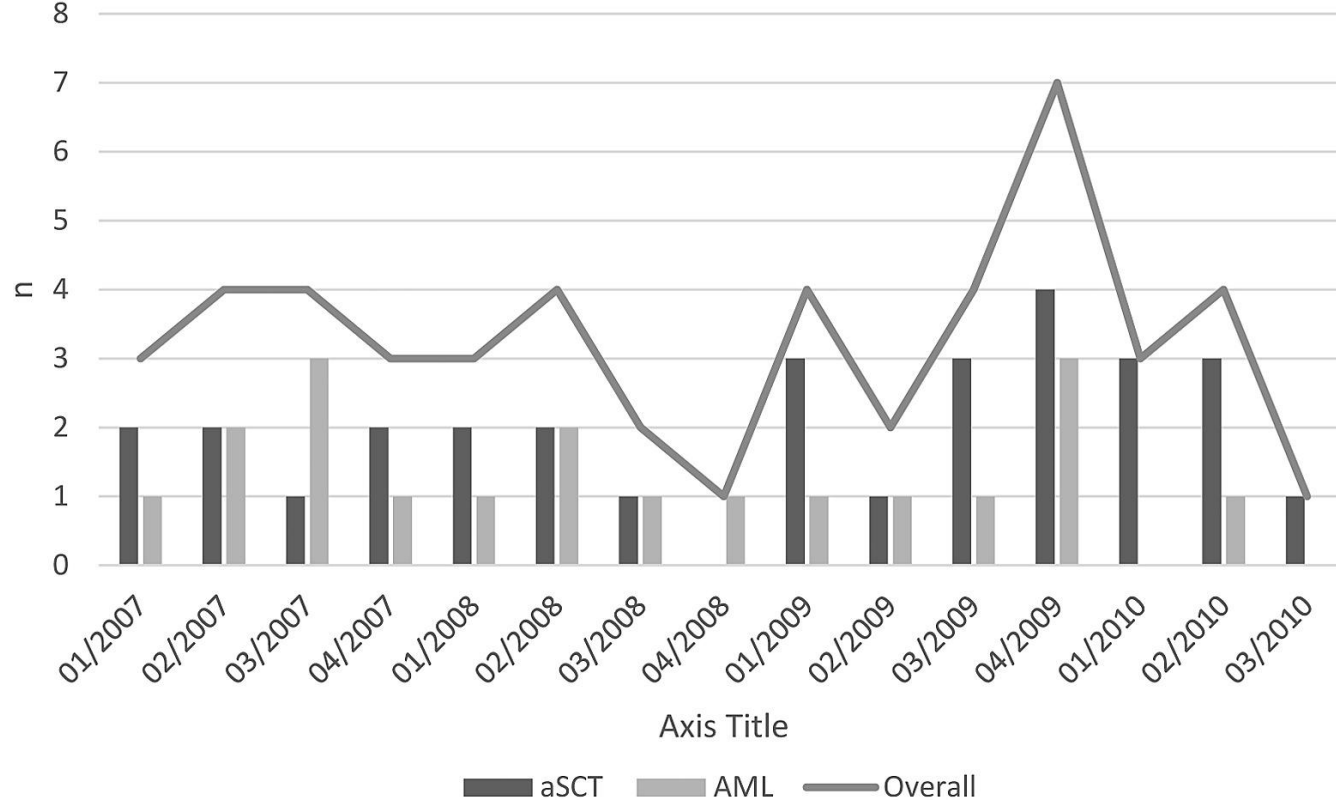
Aspergillus spp.



Legionella



Distribution of CDI cases over time



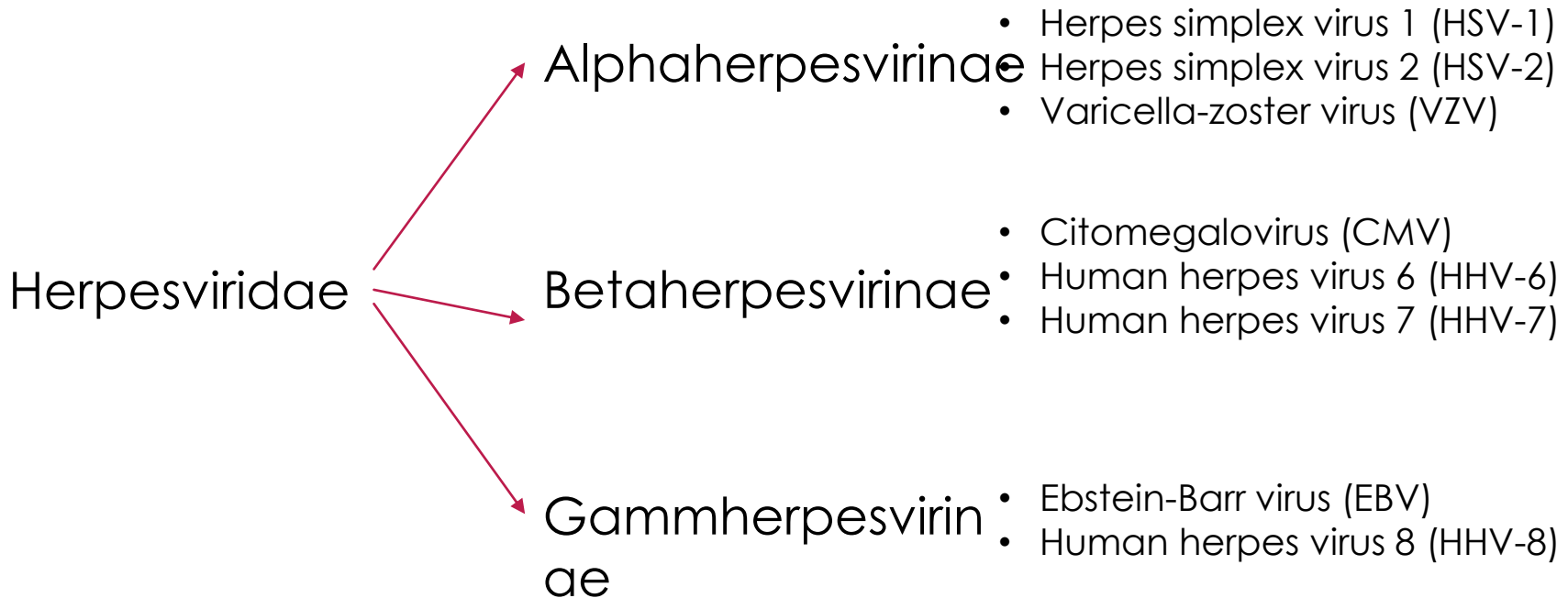
To better understand the impact of fluoroquinolone (FQ) prophylaxis on local epidemiology, continuous microbiological surveillance is highly recommended. This should not only focus on FQ-resistant pathogens but also on rates of *Clostridioides difficile* infection (CDI)

Clostridium Difficile Infection in Patients with Acute Myelogenous Leukemia and in Patients Undergoing Allogeneic Stem Cell Transplantation: Epidemiology and Risk Factor Analysis

Maria J.G.T. Vehreschild • David Weitershagen • Lena M. Biehl • ... Michael v. Bergwelt-Baildon •

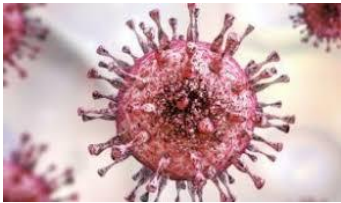
Oliver A. Cornely • Joerg J. Vehreschild • [Show all authors](#)

Herpesviridae family



Clinical Manifestations

Immunocompromised Host



- **CMV Syndrome**

- **End-organ Disease**

- **Allograft Dysfunction**

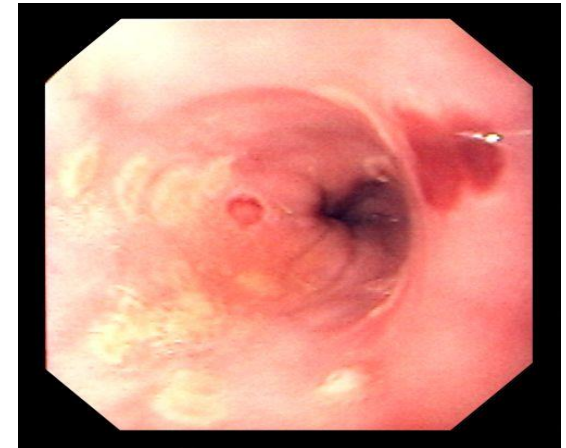
Direct effects	
Gastrointestinal	Cardiovascular
Colitis	Myocarditis
Enteritis	Venous thrombosis
Gastritis	Neurological
Hepatitis	Meningitis
Pancreatitis	Encephalitis
Cholangitis	Myelitis
Respiratory	Retinitis
Pneumonitis	Uveitis
Haematological	Urological
Thrombocytopenia	Nephritis
Leukopenia	Prostatitis
Anaemia	
Disseminated intravascular coagulation	
Myelodysplastic change	
Indirect effects	
Atherosclerosis acceleration	Accelerated AIDS progression
Graft dysfunction and rejection	Increased opportunistic infections

Clinical Manifestations

Immunocompromised Host

Gastrointestinal Disease

- Colitis
- Hepatitis
- Esophagitis



Clinical Manifestations Immunocompromised Host

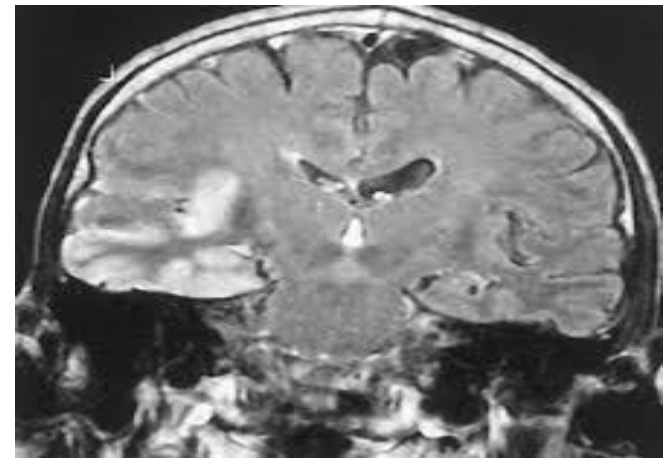
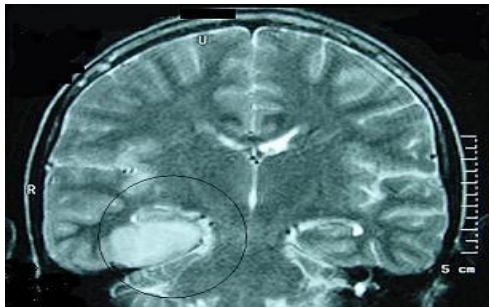
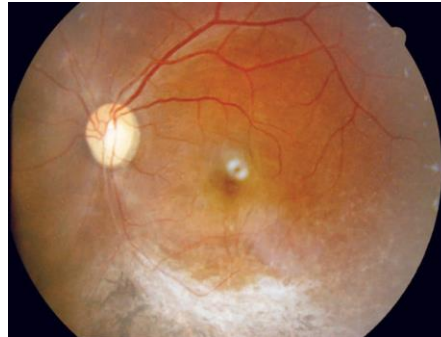
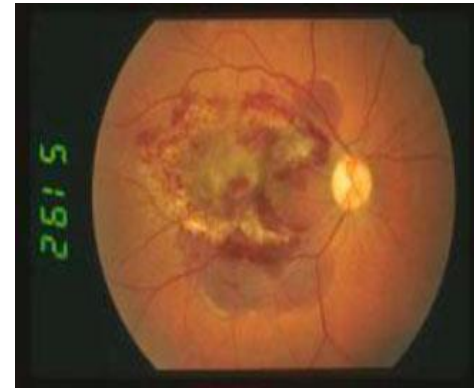
Pneumonia



Clinical Manifestations Immunocompromised Host Central Nervous System

- *Retinitis*

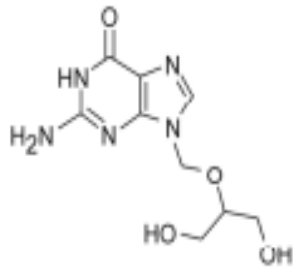
- *Encephalitis*



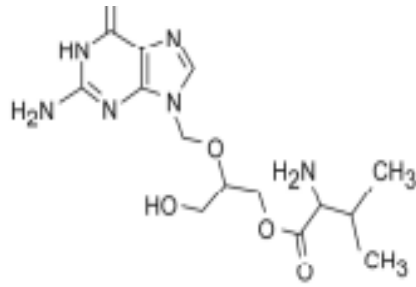
Dioverti MV et al. Cytomegalovirus. Microbiol Spectr. 2016

Therapy

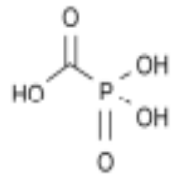
Management of Cytomegalovirus Infections in the Era of the Novel Antiviral Players, Letermovir and Maribavir



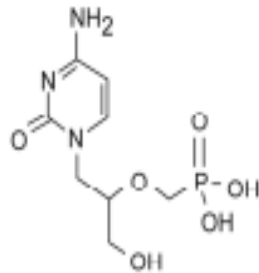
Ganciclovir
EC₅₀ = 1–4 μM



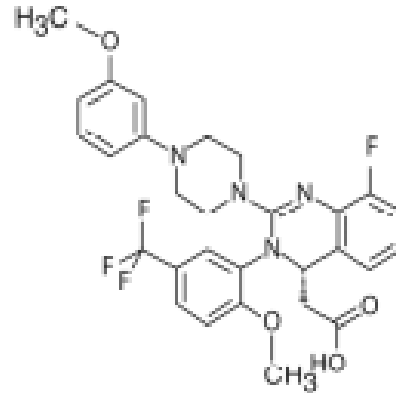
Valganciclovir



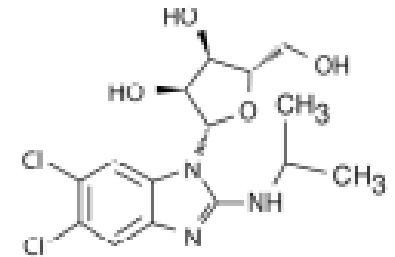
Foscarnet
EC₅₀ = 40–200 μM



Cidofovir
EC₅₀ = 0.2–0.4 μM



Letermovir
EC₂₀ = 5 nM



Maribavir
EC₅₀ = 0.1–13 μM

Antiviral agent	CMV	EBV	HHV-6	HHV-8	HSV	VZV	BK	Adenovirus
Acyclovir/valacyclovir/famciclovir ^a	High dose ±				X	X		
Ganciclovir IV/valganciclovir PO	X		X	±	X	X		
Foscarnet ^b	X		X	±	X	X		
Cidofovir ^b	X		X	±	X	X	Poor	IC ₅₀ ±
Letermovir ^a (prophylaxis only)	X							
Maribavir ^b (resistant/refractory CMV treatment only)	X	In vitro only						

Anti-CMV Immunoglobulins...therapy?

Currently indicated only for prophylaxis in patients undergoing SOT and HSCT


1 ml/kg for each administration for at least six administrations at intervals of two to three weeks.

TRANSPLANT INFECTIOUS DISEASE



ORIGINAL ARTICLE

Use of anti-CMV immunoglobulins in lung transplant recipients: The French experience

Charlotte Roy  François Parquin, Jonathan Messika, Boussaud Véronique, Olivier Brugière, Tristan Degot, Séverine Feuillet, Jérôme Lepavec, Adrien Tissot, Claire Dromer ... See all authors

First published: 01 November 2021 | <https://doi.org/10.1111/tid.13754> | Citations: 2



Transplantation Proceedings, 53, 1284–1287 (2021)

Successful Treatment of UL97 Mutation Ganciclovir-Resistant Cytomegalovirus Viremia in a Renal Transplant Recipient With Letemovir and Adjunct Hyperimmune Cytomegalovirus Immunoglobulin: A Case Report

Aaron P. Pearston^{a*}, Amanda I. Ingemi^a, Kathryn Ripley^b, Tyler J. Wilson^a, Jacqueline Gruber^a, Megan McMahon^a, Sharon Sutton^a, and Nancy Khardori^b

 frontiers
in Immunology

CASE REPORT
published: 25 January 2021
doi: 10.3389/fimmu.2020.623178



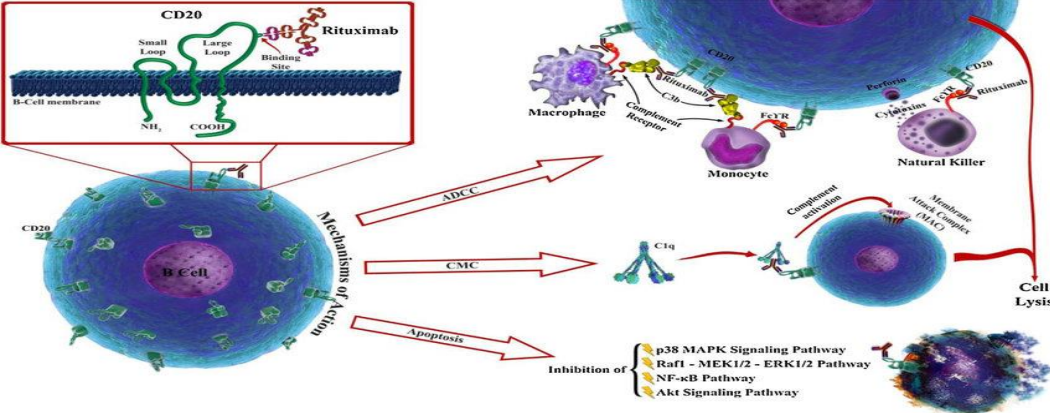
OPEN ACCESS

Edited by:
Oriol Bastard,
Bellvitge University Hospital, Spain

Reviewed by:
Lionel Couzi,
University of Bordeaux, France

Case Report: Management of a Multidrug-Resistant CMV-Strain in a Renal Transplant Recipient by High-Dose CMV-Specific Immunoglobulins, Modulation in Immunosuppression, and Induction of CMV-Specific Cellular Immunity

Vanessa Wfening¹, Tina Schmidt², Maximilian Dahmen^{1†}, Sami Slam¹, Stefan Reuter¹, Hermann-Joseph Pavenstädt¹, Martina Sester² and Barbara Suwelack^{1*}



[Cancers \(Basel\)](#). 2018 Jun; 10(6): 197.

Published online 2018 Jun 13. doi: [10.3390/cancers10060197](https://doi.org/10.3390/cancers10060197)

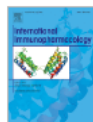
Antiviral Drugs for EBV

[Joseph S. Pagano](#)^{1,*}, [Christopher B. Whitehurst](#)² and [Graciela Andrei](#)³



International Immunopharmacology

Volume 96, July 2021, 107606

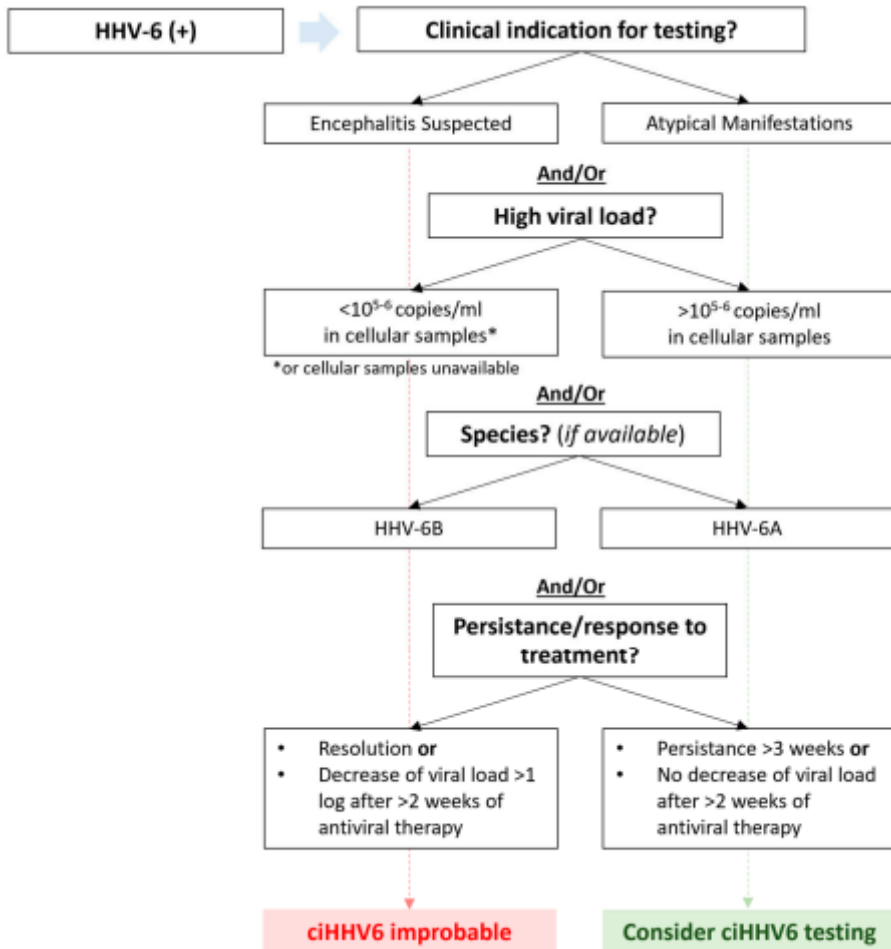


Rituximab-containing immuno-chemotherapy regimens are effective for the elimination of EBV for EBV-HLH with only and mainly B lymphocytes of EBV infection

[J Virol](#). 2013 May;87(9):5311-5. doi: [10.1128/JVI.03505-12](https://doi.org/10.1128/JVI.03505-12). Epub 2013 Feb 28.

Maribavir inhibits Epstein-Barr virus transcription through the EBV protein kinase

[Christopher B Whitehurst](#)¹, [Marcia K Sanders](#), [Mankit Law](#), [Fu-Zhang Wang](#), [Jie Xiong](#), [Dirk P Dittmer](#), [Joseph S Pagano](#)



HHV-6

Algorithm for testing for chromosomally integrated HHV-6.

Testing for chromosomally integrated HHV-6 (ciHHV6) should be considered in cases with atypical manifestations and/or high viral load $>10^{5-6}$ in cellular samples (assuming that white blood count is normal) and/or detection of species HHV-6A (based on data that HHV-6A reactivation and disease are rarely seen in this patient population and appear to be primarily detected in ciHHV6).

Testing should also be considered in the absence of a virological response despite antiviral treatment (e.g., persistent detection without a $\geq \log_{10}$ decline in viral load after ≥ 2 weeks).



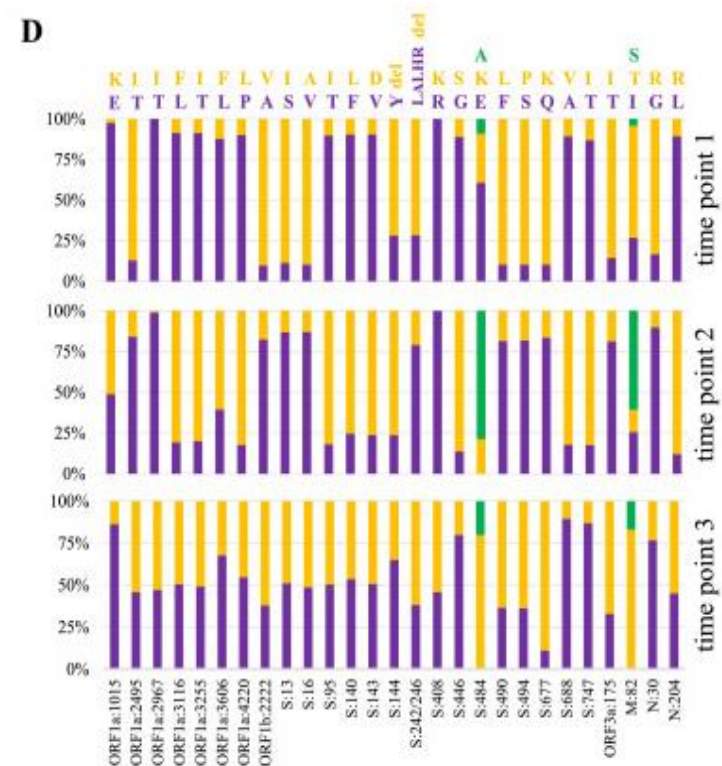
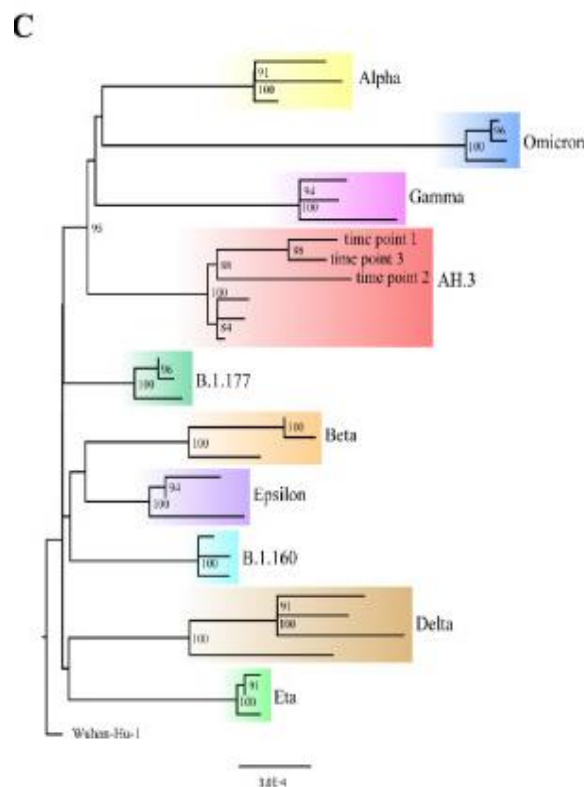
SARS-CoV-2 intra-host evolution during prolonged infection in an immunocompromised patient

Erika Giorgia Quaranta ¹, Alice Fusaro ², Edoardo Giussani ², Valeria D'Amico ², Maria Varotto ², Matteo Pagliari ², Maria Teresa Giordani ³, Maira Zoppelletto ⁴, Francesca Merola ⁴, Antonio Antico ⁴, Paola Stefanelli ⁵, Calogero Terregino ², Isabella Monne ²

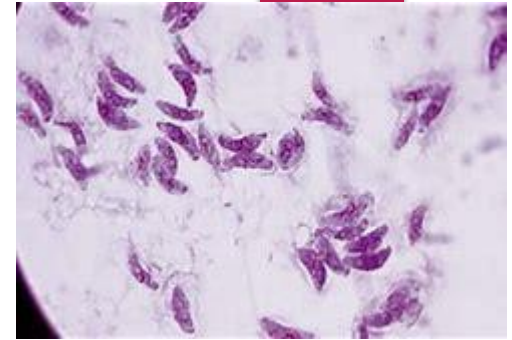
SARS-CoV-2 mutation accumulation during chronic infection in a patient with HIV over 109 days.

(C) Phylogenetic analysis of the sequences obtained from patient swabs at the three time points. The sequences were aligned to a set of representative SARS-CoV-2 genome sequences belonging to AH.3 lineage and the main lineages and VOCs identified so far. The maximum-likelihood phylogenetic tree was constructed with IQ-TREE (GTR+F+R2) and rooted on the Wuhan-Hu-1 reference genome. Ultra-fast bootstrap supports are indicated above the nodes.

(D) Histograms representing the amino acid frequency at the positions that differ between the three time points. Amino acids are indicated above the columns for each position. VOCs = variants of concern.



Toxoplasmosis

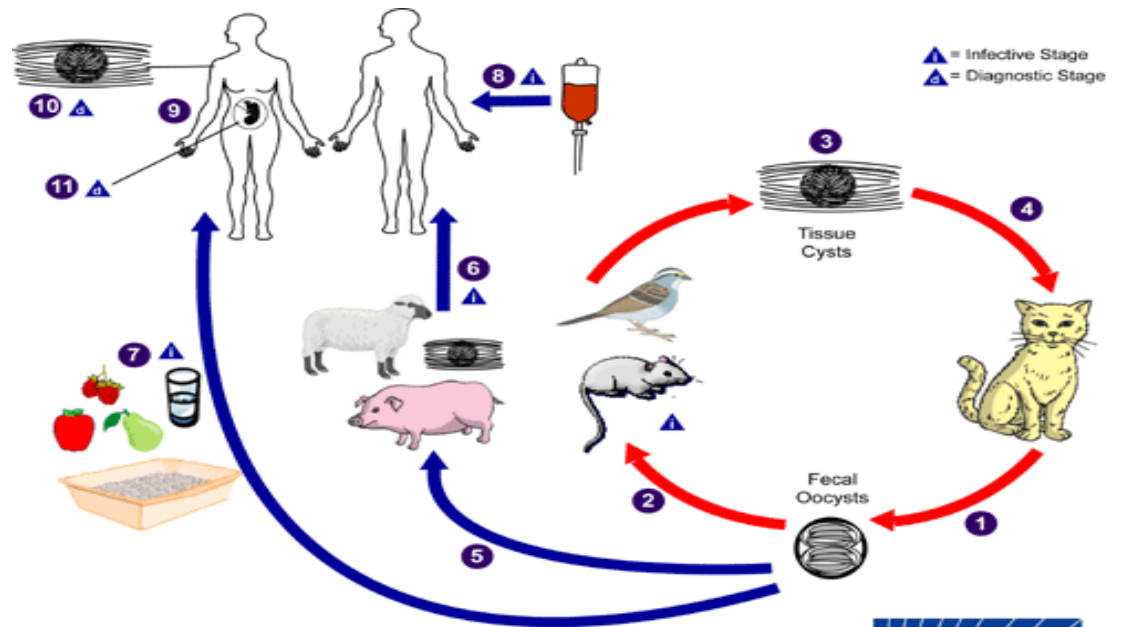
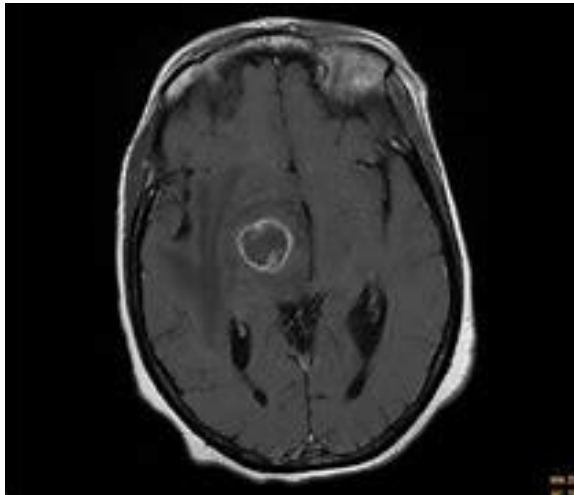


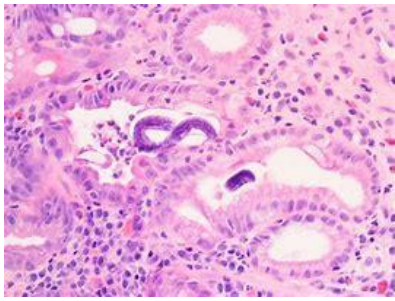
THE LANCET

Toxoplasmosis

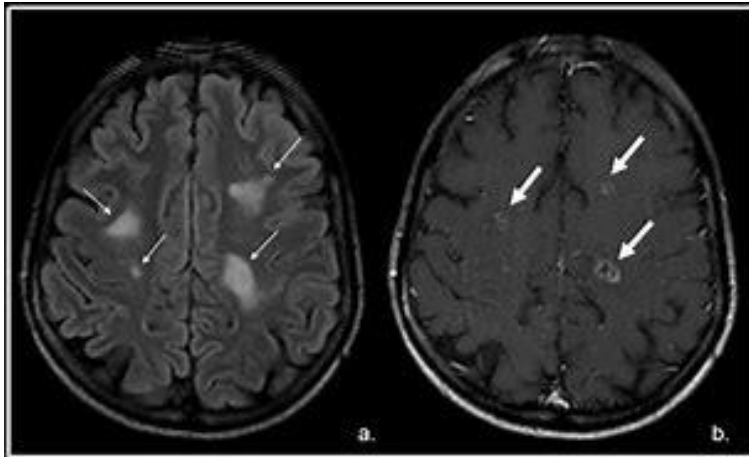
JG Montoya, MD • Prof O Liesenfeld, MD

Published: June 12, 2004 • DOI: [https://doi.org/10.1016/S0140-6736\(04\)16412-X](https://doi.org/10.1016/S0140-6736(04)16412-X)

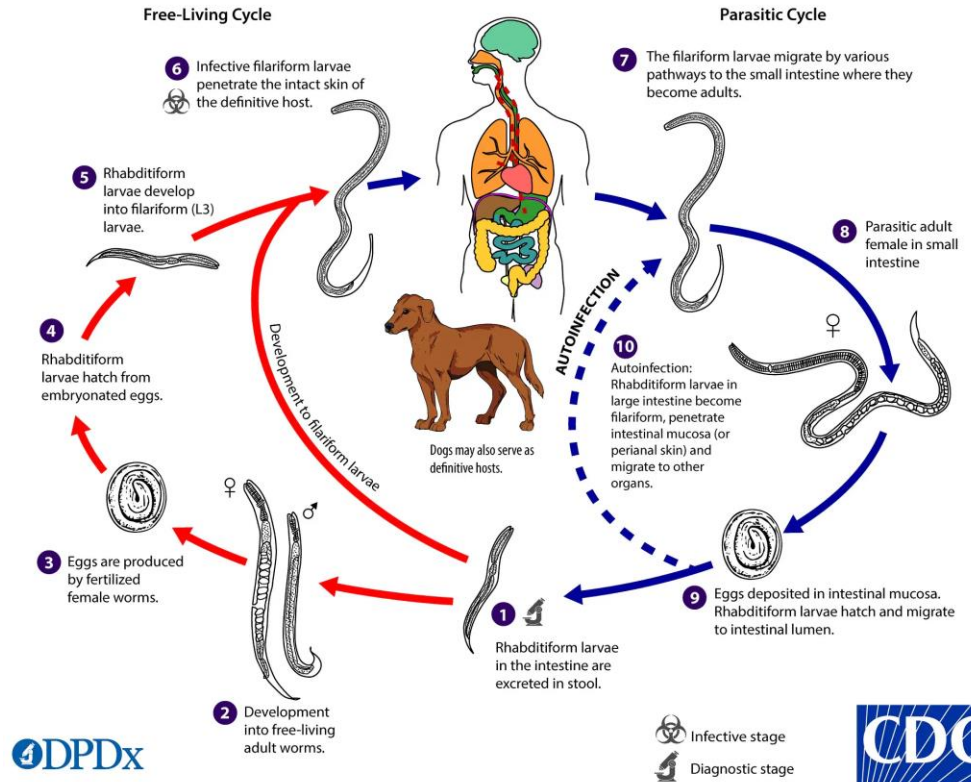




Strongyloidiasis



Strongyloides stercoralis



DPDx



Primer | Published: 25 January 2024

Strongyloidiasis

nature reviews disease primers

Catherine A. Gordon , Jürg Utzinger, Stephen Muhi, Sören L. Becker, Jennifer Keiser, Virak Khieu & Darren

J. Gray

The number of indications for splenectomy in cancer patients has declined over the years.

In the case of solid tumors, splenectomy is traditionally performed in gastric cancer to dissect the splenic hilar lymph nodes, although it seems to have no benefit in tumors located at lesser curvature

Splenectomy may also be performed in cases of oligometastatic disease from other sites, especially ovarian cancer

Additionally, intraoperative splenic injury during abdominal surgery may result in splenectomy

Besides surgical splenectomy resulting in asplenia, radiotherapy and some pathologic conditions (including graft versus host disease following allogeneic hematopoietic stem cell transplantation) lead to a decreased function of the spleen—hyposplenism (function asplenia)

Splenectomy and Function Asplenia

The spleen is a lymphoid organ that plays an important role in regulating immune homeostasis through both innate and adaptive immunity.

The function of the spleen is crucial in the elimination of encapsulated bacteria

Asplenic patients are therefore at risk of sepsis caused by **encapsulated bacteria**

most commonly *Streptococcus pneumoniae* (50–70%), but also *Haemophilus influenzae* and *Neisseria meningitidis* (15–25% each)

Other pathogens causing serious infections in asplenic patients include *Capnocytophaga canimorsus* after animal bites, *Bordetella holmesii*, *Ehrlichia* spp., and intraerythrocytic parasites such as *Babesia* spp. after tick bites

Review

Medical complications following splenectomy



ELSEVIER

Journal of Visceral Surgery

Volume 153, Issue 4, August 2016, Pages 277-286



R. Buzelé ^{a 1}, L. Barbier ^{b 2}, A. Sauvanet ^b, B. Fantin ^a

Hippocrates (circa 460-377 BCE) originally recognized the effects of our surroundings on human diseases in his treatise On Airs, Waters and Places, attributing illness to characteristics of climate, water, modes of life, and nutrition

Two thousand years later, in the 1800s, Ignaz Semmelweis (1818-1865 CE) documented the effects of environmental control through hand hygiene on clinical outcomes, achieving a dramatic decrease in puerperal mortality with the widespread use of aseptic techniques, in which practitioners cleaned their hands with chlorine solution in between patients

Provided to the PMC COVID-19 Collection by

Wiley

[CA Cancer J Clin](#). 2018 Sep-Oct; 68(5): 340–355.

Published online 2018 Jul 9. doi: [10.3322/caac.21462](https://doi.org/10.3322/caac.21462)

Update on infection control practices in cancer hospitals

[Ella J. Ariza-Heredia](#), MD, Assistant Professor ¹ and [Roy E. Chemaly](#), MD, MPH, Professor ²

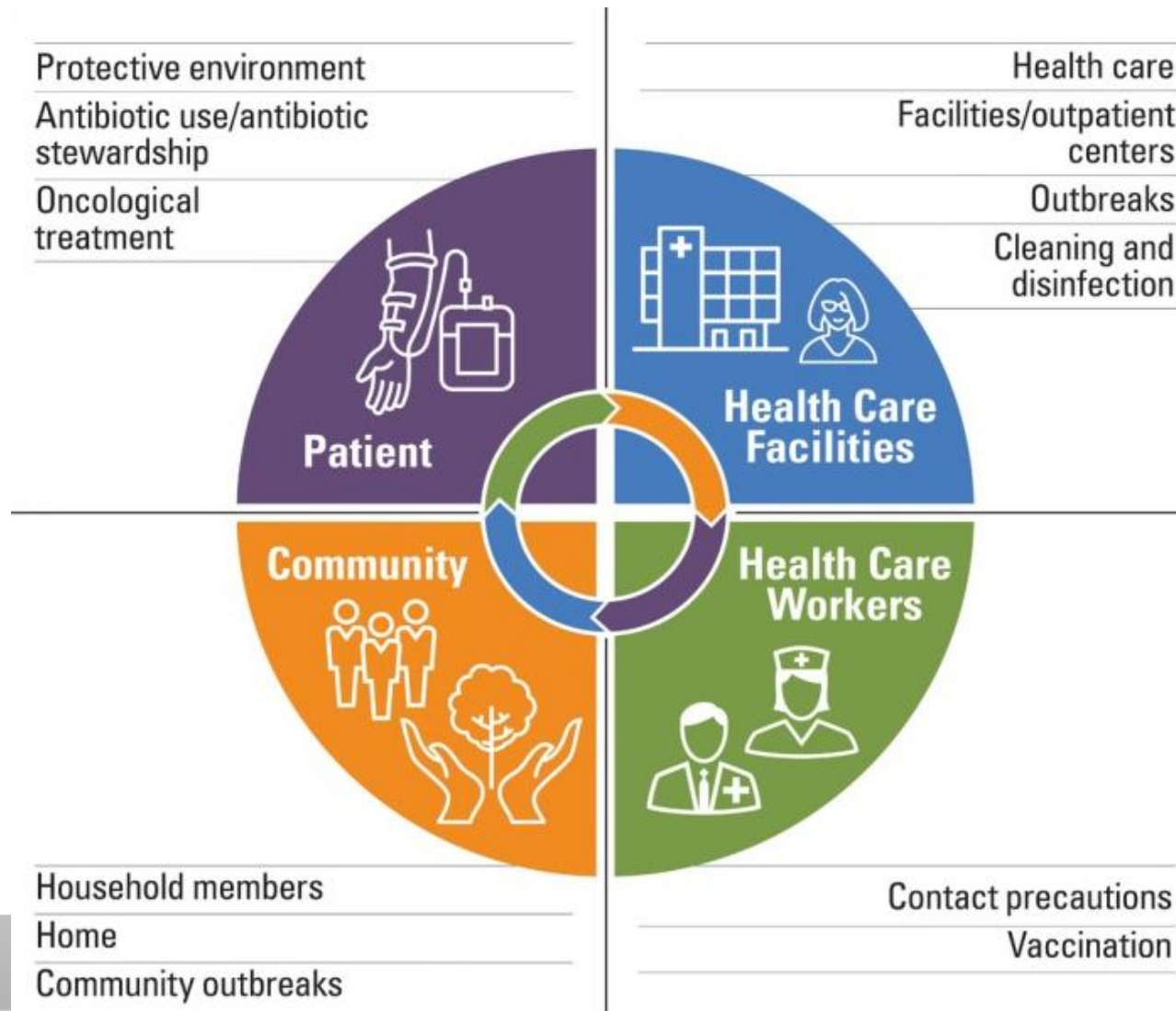
Important Aspects of Infection Control and Prevention in Patients Living With Cancer

CA Cancer J Clin. 2018 Sep-Oct; 68(5): 340–355.

Published online 2018 Jul 9. doi: [10.3322/caac.21462](https://doi.org/10.3322/caac.21462)

Update on infection control practices in cancer hospitals

[Ella J. Ariza-Heredia](#), MD, Assistant Professor¹ and [Roy E. Chemaly](#), MD, MPH, Professor²

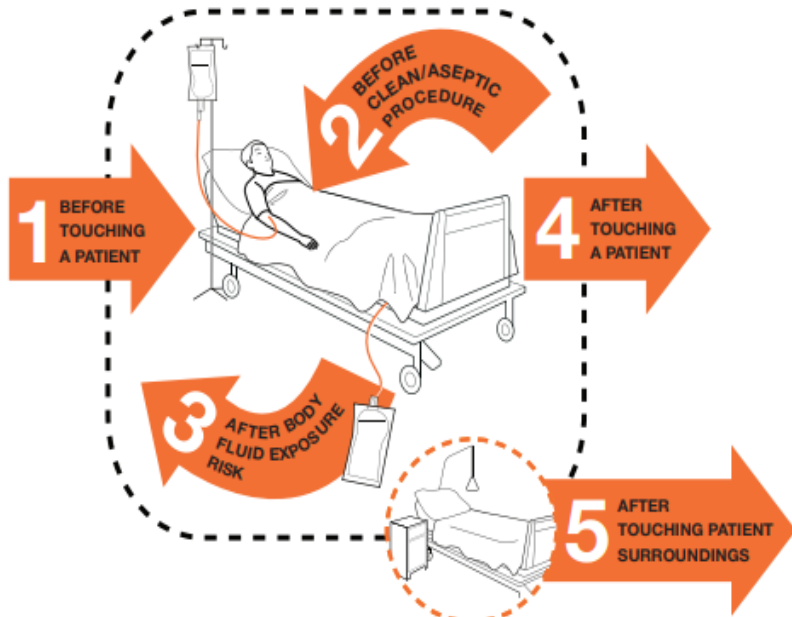


Your 5 Moments for Hand Hygiene

Preventing Infection

Hand Hygiene

“My Five Moments for Hand Hygiene”



1 BEFORE TOUCHING A PATIENT	WHEN? Clean your hands before touching a patient when approaching him/her. WHY? To protect the patient against harmful germs carried on your hands.
2 BEFORE CLEAN/ASEPTIC PROCEDURE	WHEN? Clean your hands immediately before performing a clean/aseptic procedure. WHY? To protect the patient against harmful germs, including the patient's own, from entering his/her body.
3 AFTER BODY FLUID EXPOSURE RISK	WHEN? Clean your hands immediately after an exposure risk to body fluids (and after glove removal). WHY? To protect yourself and the health-care environment from harmful patient germs.
4 AFTER TOUCHING A PATIENT	WHEN? Clean your hands after touching a patient and her/his immediate surroundings, when leaving the patient's side. WHY? To protect yourself and the health-care environment from harmful patient germs.
5 AFTER TOUCHING PATIENT SURROUNDINGS	WHEN? Clean your hands after touching any object or furniture in the patient's immediate surroundings, when leaving – even if the patient has not been touched. WHY? To protect yourself and the health-care environment from harmful patient germs.

Preventing Infection


Hand Hygiene

Siegel JH, Korniewicz DM. Keeping patients safe: an interventional hand hygiene study at an oncology center. Clin J Oncol Nurs. 2007

Regardless of the product used, hand washing is an important modality for the prevention of infection and should be performed by patients, visitors, and health care workers.

Research | [Open access](#) | Published: 19 April 2024

Ten years of hand hygiene excellence: a summary of outcomes, and a comparison of indicators, from award-winning hospitals worldwide

[Ermira Tartari](#) , [Jacopo Garlasco](#), [Marcela Hernández-de Mezerville](#), [Moi Lin Ling](#), [Hilda Márquez-Villarreal](#), [Wing-Hong Seto](#), [Anne Simon](#), [Thomas-Jörg Hennig](#) & [Didier Pittet](#)

[Antimicrobial Resistance & Infection Control](#) **13**, Article number: 45 (2024) | [Cite this article](#)



a 40% overall decrease in the rate of nosocomial infections

 **BMC** Part of Springer Nature

THE LANCET
Infectious Diseases

REVIEW | VOLUME 6, ISSUE 10, P641-652, OCTOBER 2006

Evidence-based model for hand transmission during patient care and the role of improved practices

Prof Didier Pittet, MD   • [Benedetta Allegranzi, MD](#) • [Hugo Sax, MD](#) • [Sasi Dharan, Dip HIC](#) •

[Carmem Lúcia Pessoa-Silva, MD](#) • [Liam Donaldson, MD](#) • et al. [Show all authors](#)

Published: October 2006 | DOI: [https://doi.org/10.1016/S1473-2893\(06\)70600-4](https://doi.org/10.1016/S1473-2893(06)70600-4)



Grazie per l'attenzione