



UMBERTO I  
POLICLINICO DI ROMA



SAPIENZA  
UNIVERSITÀ DI ROMA

# Le infezioni associate a devices intravascolari

Dr Marco Falcone

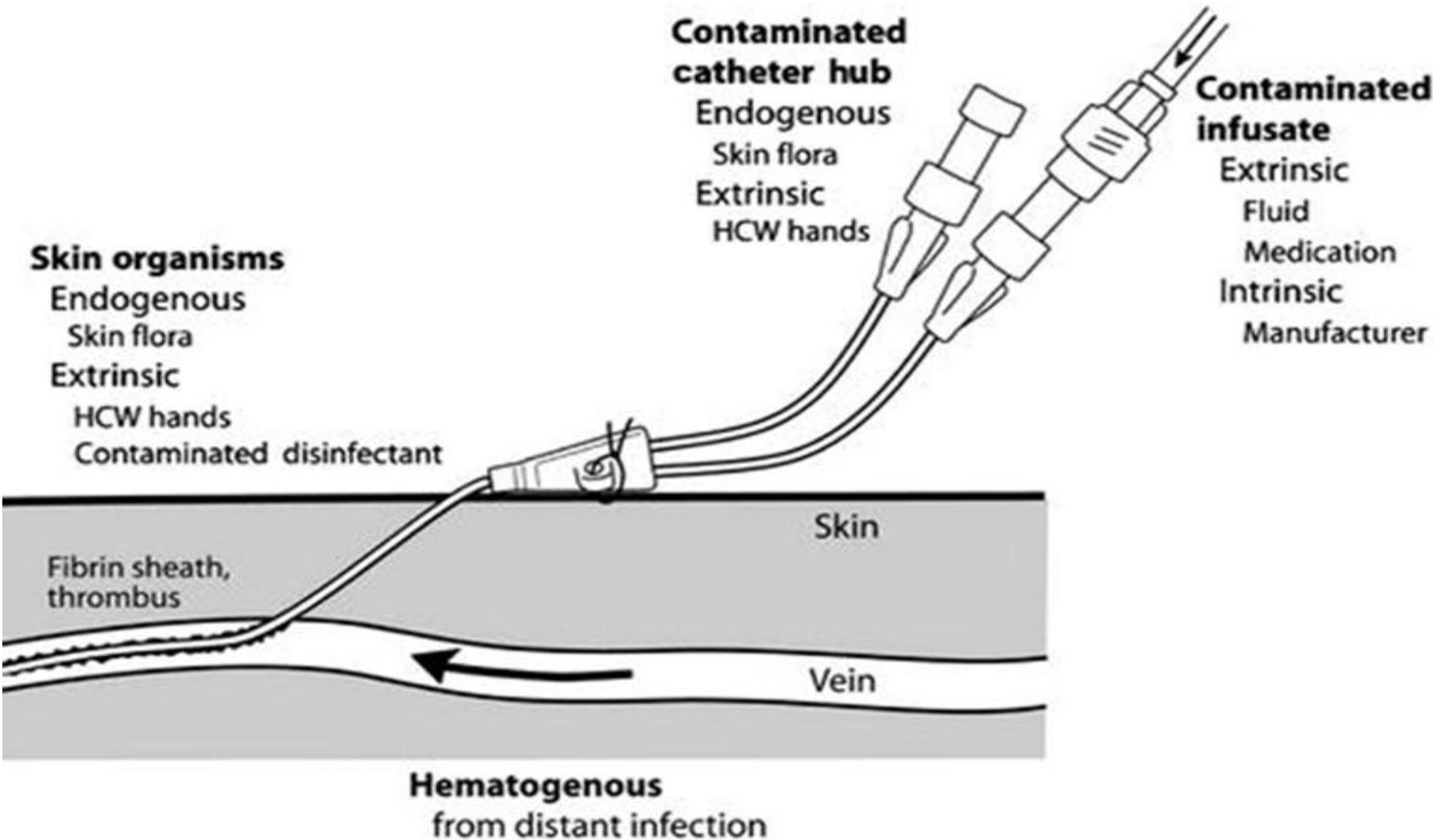
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# Types of intravascular devices

Type of intravascular device	Comments
Peripheral venous catheter	Usually inserted into the veins of the forearm or the hand; most commonly used short-term intravascular device; rarely associated with bloodstream infection
Peripheral arterial catheter	For short-term use; commonly used to monitor hemodynamic status and to determine blood gas levels of critically ill patients; risk of bloodstream infection may approach that of CVCs
Midline catheter	Peripheral catheter (size, 7.6–20.3 cm) is inserted via the antecubital fossa into the proximal basilic or cephalic veins, but it does not enter central veins; is associated with lower rates of phlebitis and infection than are CVCs
Nontunneled CVC	Most commonly used CVC; accounts for an estimated 90% of all catheter-related bloodstream infections; increased risk of infection with internal jugular vein site of insertion
Pulmonary artery catheter	Inserted through a Teflon introducer and typically remains in place for an average duration of only 3 days; most catheters are heparin bonded to reduce catheter thrombosis and microbial adherence to the catheter
Pressure-monitoring system	Used in conjunction with arterial catheter; associated with both epidemic and endemic nosocomial bloodstream infections; source is often the fluid column in the tubing between the patient's intravascular catheter and the pressure-monitoring apparatus, contaminated infusate, or nondisposable transducers
Peripherally inserted central catheter	Provides an alternative to subclavian or jugular vein catheterization; is inserted via the peripheral vein into the superior vena cava, usually by way of cephalic and basilar veins; is easier to maintain and is associated with fewer mechanical complications (e.g., hemothorax) than are nontunneled CVCs
Tunneled CVC	Surgically implanted CVC (e.g., Hickman, Broviac, Groshong, or Quinton catheter) with the tunneled portion exiting the skin and a Dacron cuff just inside the exit site; the cuff inhibits migration of organisms into the catheter tract by stimulating growth of surrounding tissue, thus the sealing catheter tract; used to provide vascular access to patients who require prolonged iv chemotherapy, home-infusion therapy, or hemodialysis (figure 4)
Totally implantable device	A subcutaneous port or reservoir with self-sealing septum is tunneled beneath the skin and is accessed by a needle through intact skin; low rates of infection

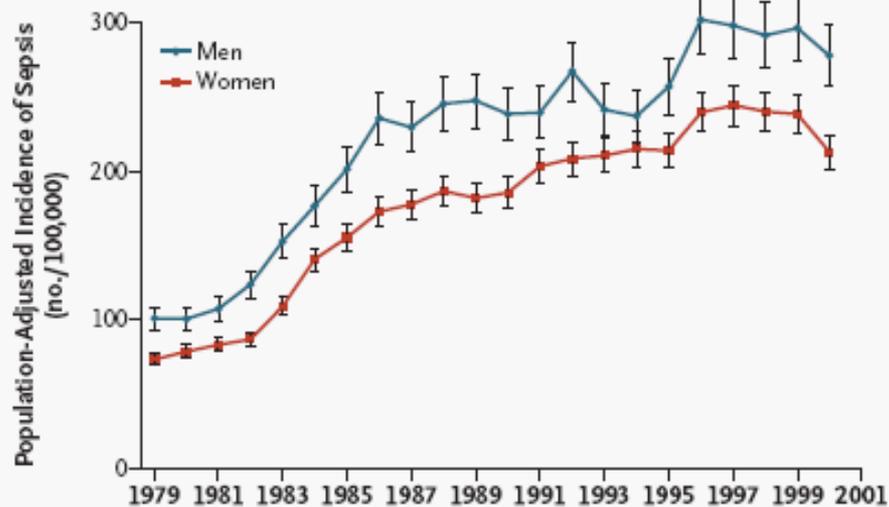
**NOTE.** CVC, central venous catheter.

# Types of intravascular devices

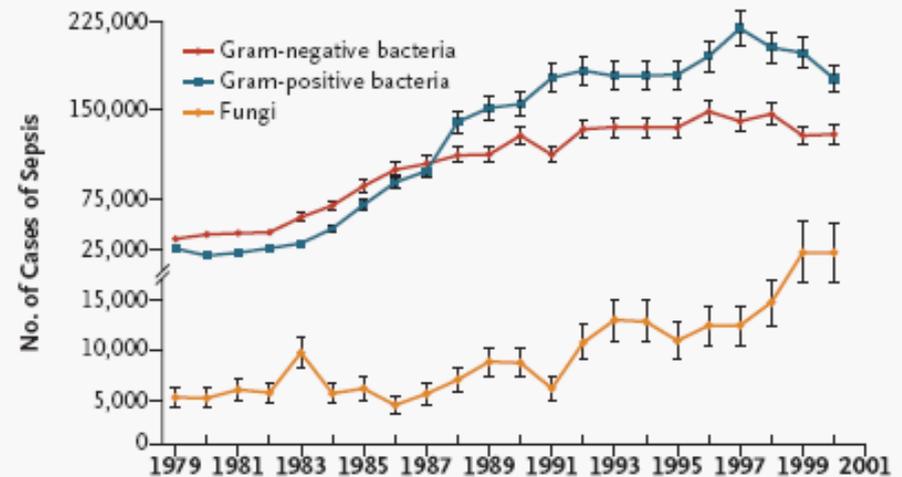


# The Epidemiology of Sepsis in the United States from 1979 through 2000

Greg S. Martin, M.D., David M. Mannino, M.D., Stephanie Eaton, M.D.,  
and Marc Moss, M.D.



**Figure 1.** Population-Adjusted Incidence of Sepsis, According to Sex, 1979–2000. Points represent the annual incidence rate, and I bars the standard error.



**Figure 3.** Numbers of Cases of Sepsis in the United States, According to the Causative Organism, 1979–2000.

Points represent the number of cases for the given year, and I bars the standard error.

# Implications of CVC use

- Increase of bacteremia due to methicillin-resistant CNS
- Increase of MRSA bacteremia
- Community spread of MRSA
- Increase of candidemia
- emergence of “non albicans” *Candida*
- Emergence of low-virulence and opportunistic bacteria and fungi

# ***Staphylococcus* species non *S. aureus***

- *S. epidermidis*
- *S. hominis*
- *S. warneri*
- *S. haemolyticus*
- *S. caprae*
- *S. pasteurii*
- *S. auriculari*
- *S. lugdunensis*
- *S. saprophyticus*
- *S. saccharolyticus*
- *S. xilosus*
- *S. capitis*
- *S. cohnii*
- *S. simulans*
- *S. schleiferi*
- *S. intermedius*

# Teicoplanin use and emergence of *Staphylococcus haemolyticus*: is there a link?

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ANNO	REPARTO	OSSERVAZIONE
1992-	Farmacia	consumo annuale di teico ca. 17.000 fl
2000		vs vanco ca. 7000 fl (500 mg)
2000-	Lab. Centrale	<i>S.haemolyticus</i> seconda specie CNS
2003	Microbiologia	22-24% isolati dal sangue. Teico-R: 11-29%
6/2000	Ematologia tutte	MR <i>S.haemolyticus</i> identificato in 18% di le batteriemie stafilococciche.

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Falcone M & Venditti M. Clin Microbiol Infect. 2006

Falcone M, Stefani S & Venditti M Diagn Microbiol Infect Dis , 2007

## *In vitro* activity of daptomycin against methicillin- and multi-resistant *Staphylococcus haemolyticus* invasive isolates carrying different *mec* complexes<sup>☆</sup>

Floriana Campanile<sup>a</sup>, Dafne Bongiorno<sup>a</sup>, Sonia Borbone<sup>a</sup>, Marco Falcone<sup>b</sup>, Maddalena Giannella<sup>b</sup>, Mario Venditti<sup>b</sup>, Stefania Stefani<sup>a,\*</sup>

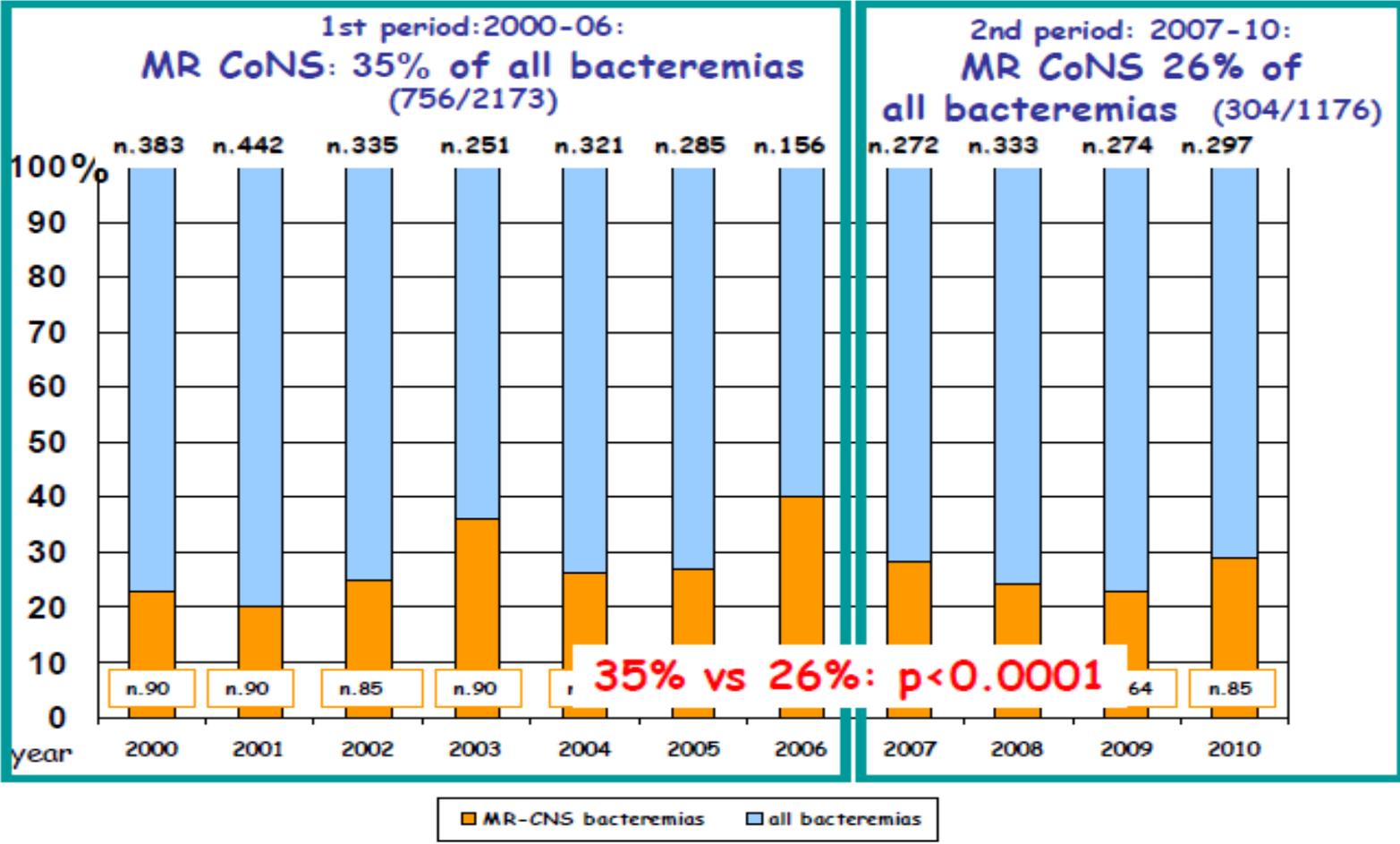
*In vitro* activity of daptomycin compared with other antibiotics versus *Staphylococcus haemolyticus* isolates

Antibiotics	Range	MIC <sub>50</sub> mg/L	MIC <sub>90</sub> mg/L	%S	%R
Daptomycin	0.12–2	0.5	1	100	0
Quinopristin/dalfopristin	0.12–1	0.25	0.5	100	0
Vancomycin *	0.5–8	1	2	100	0
Linezolid	1–2	2	2	100	0
Teicoplanin ***	0.25–≥64	8	32	62	38
Erythromycin	0.25–≥64	32	≥64	14	86
Clindamycin **	0.25–≥64	0.25	≥64	72	28
Ciprofloxacin	0.5–≥64	32	≥64	12	88
Levofloxacin	0.25–≥64	32	≥64	38	62
Gentamicin	0.25–≥64	≥64	≥64	8	92
Cotrimoxazole	0.12–≥64	8	≥64	38	62
Imipenem	0.12–≥64	32	≥64	34	66
Meropenem	0.12–≥64	32	≥64	12	88
Rifampin	≤0.06–≥64	0.12	0.25	88	12
Tetracycline	≤0.06–32	0.12	0.5	96	4

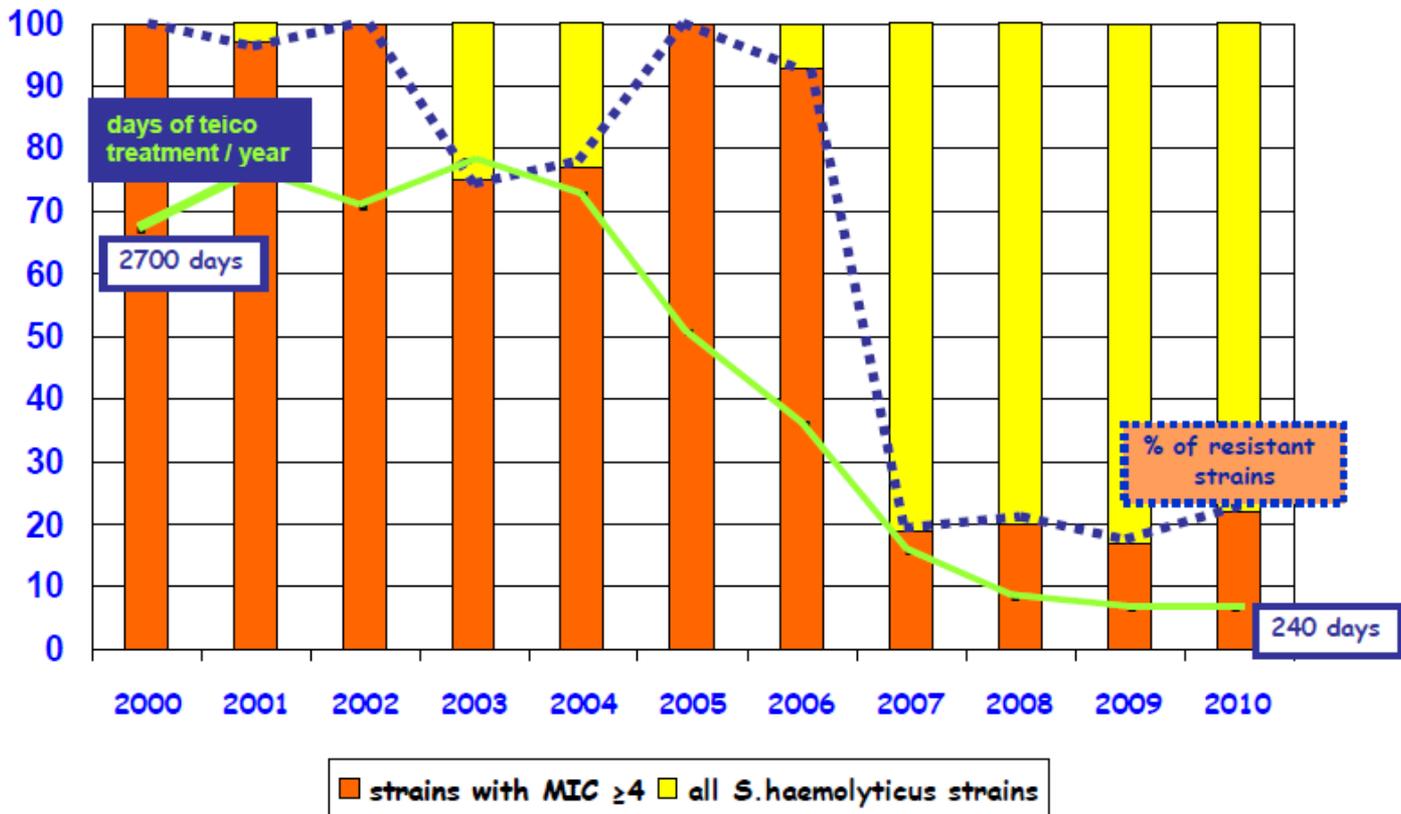
MIC susceptibility breakpoints for *Staphylococcus* spp. were according to CLSI guidelines.

\* hVISA: the two hetero-resistant strains showed daptomycin MIC values of 2 mg/L one dilution higher with respect to their wild-type strains.

# Changing of Antimicrobial Susceptibility of 175 *Staphylococcus haemolyticus* Blood Isolates from Neutropenic Patients with Hematological Malignancies: Comparison of Two Different Periods (2000-2006 vs 2007-2010)



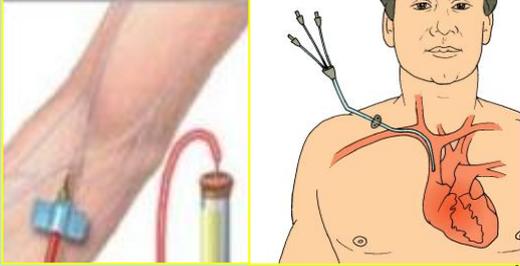
# Changing of Antimicrobial Susceptibility of 175 *Staphylococcus haemolyticus* Blood Isolates from Neutropenic Patients with Hematological Malignancies: Comparison of Two Different Periods (2000-2006 vs 2007-2010)



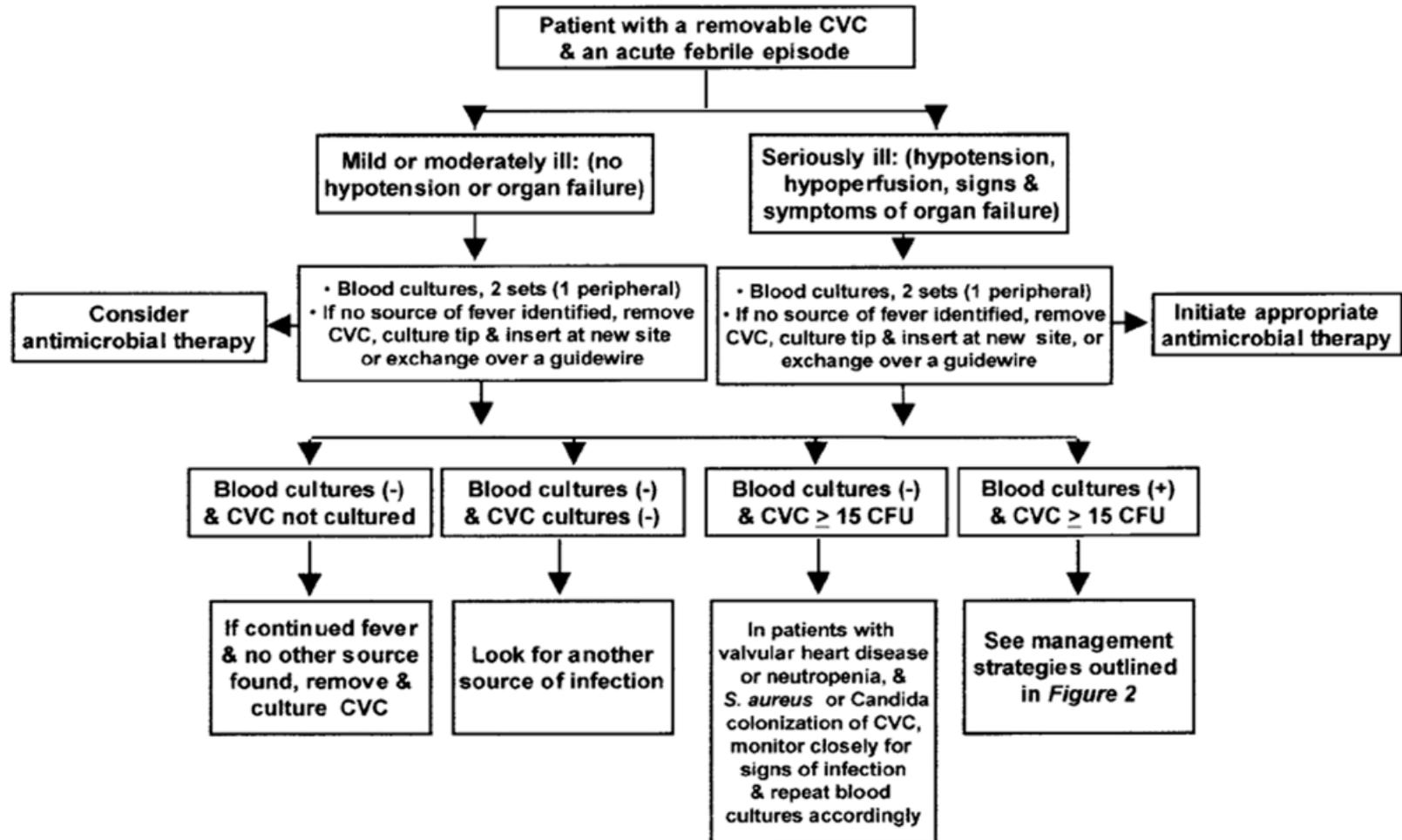
# Non conservative diagnostic methods

<b>Qualitative</b>	<b>Broth culture of distal catheter tip</b>	<b>Sensibility 95% Specificity 75% Not recommended</b>
<b>Semiquantitative “Maki technique”</b> 	<b>Culture of catheter tip on agar plate</b>  <b>Cut-off &gt; 15 CFU./plate</b>	<b>Sensibility 85% Specificity 85%</b>
<b>Quantitative</b>	<b>Flushing Vortexing Sonication</b>  <b>Cut-off <math>\geq 10^3</math> CFU./ml</b>	<b>Sensibility 83% Specificity 95%</b>

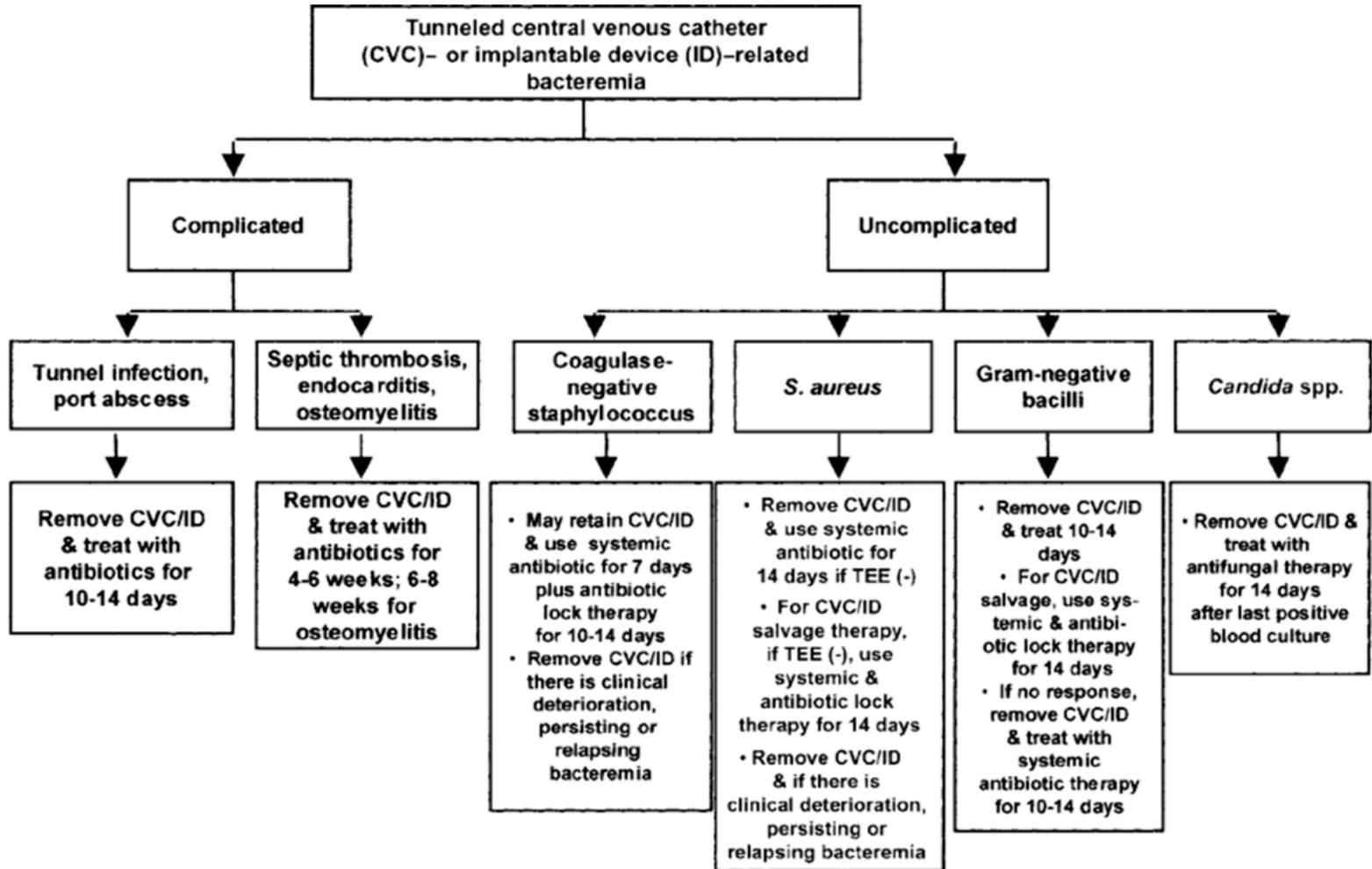
# Conservative methods

<p>Quantitative blood cultures</p> 	<p>Centrifugation/lysis of 10 ml of blood from CVC and peripheral blood</p> <p>Increase of CFU/ml from CVC respect peripheral blood (ratio 5-10:1)</p>	<p>Most accurate methodology</p> <p>Short-term CVC: Sens. 75% Spec. 97%</p> <p>Long-term CVC: Sens. 93% Spec. 100%</p> <p>Time wasting, complex</p>
<p>Time to positivity of blood cultures</p> 	<p>Automated methods</p> <p>Differential time from blood cultures drawn from CVC and peripheral blood</p> <p><b>Cut-off &gt;120 min</b></p>	<p>Sensibility 94%</p> <p>Specificity 91%</p> <p>Reduced specificity during antibiotic treatment</p>

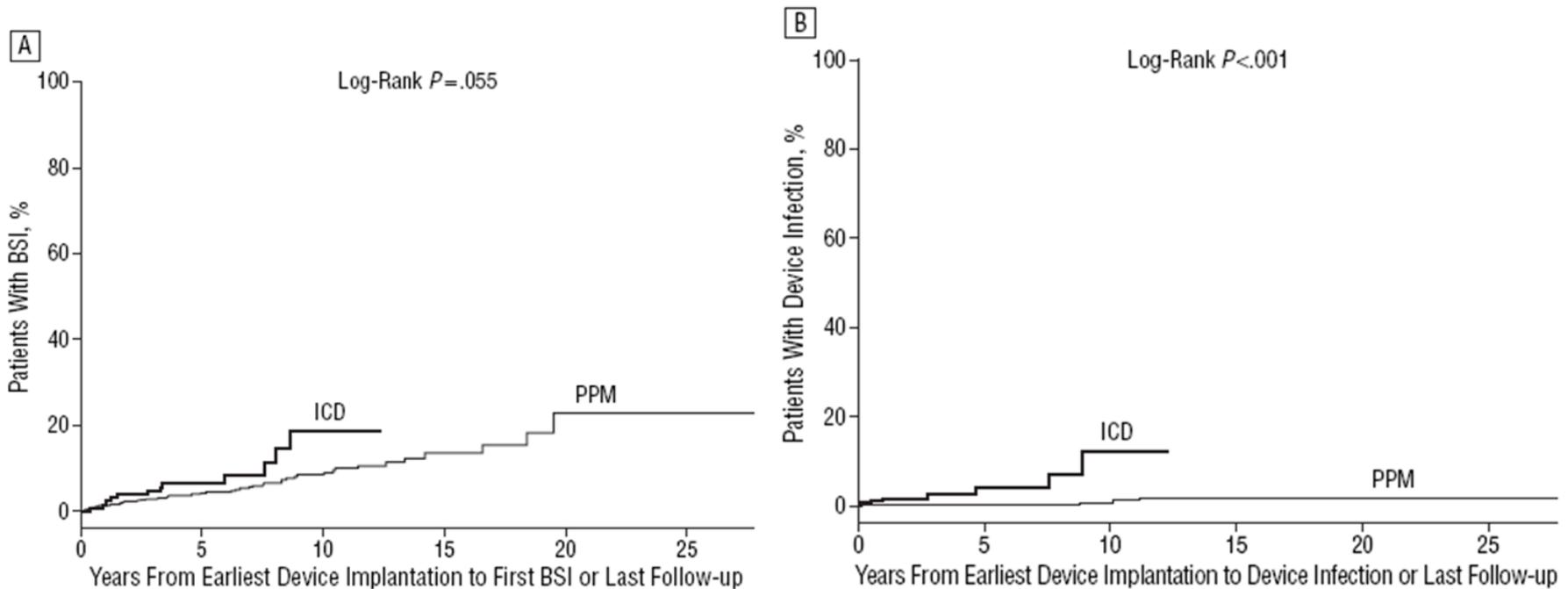
# Management of non tunneled CVC infection



# Management of tunneled CVC infection



# Pacemaker and Implantable Cardioverter Defibrillator Infection



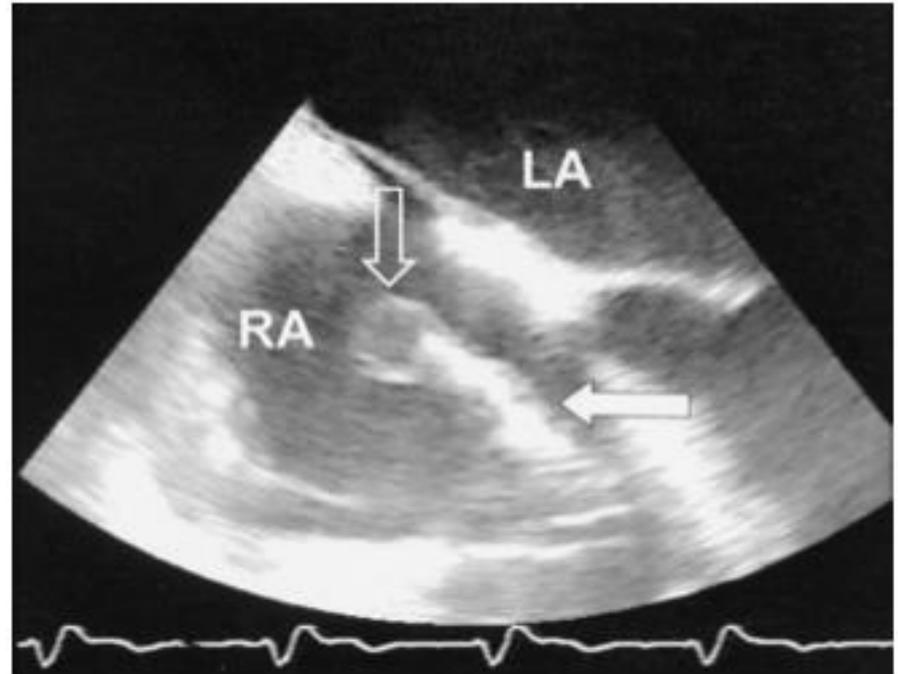
Uslan DZ, et al. Arch Intern Med 2007; 167: 670, 2007

# Pacemaker and Implantable Cardioverter Defibrillator Infection

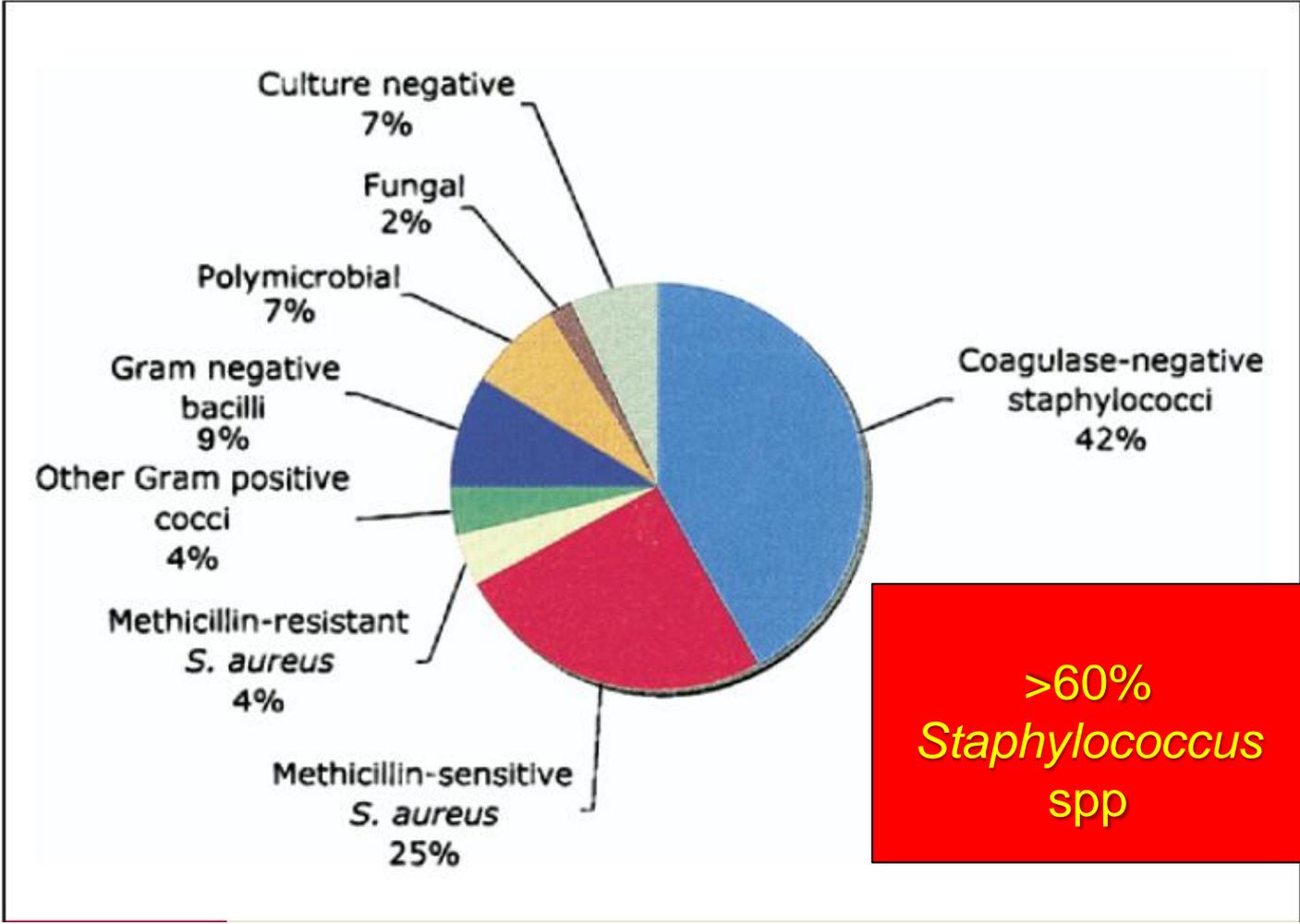
Local infection about 70-80%



Endocarditis about 15-25%



# Management and Outcome of Permanent Pacemaker and Implantable Cardioverter-Defibrillator Infections



# Sonication of Explanted Cardiac Implants Improves Microbial Detection in Cardiac Device Infections

Alessandra Oliva,<sup>a</sup> Bich Lien Nguyen,<sup>b</sup> Maria T. Mascellino,<sup>a</sup> Alessandra D'Abramo,<sup>a</sup> Marco Iannetta,<sup>a</sup> Antonio Ciccaglioni,<sup>b</sup> Vincenzo Vullo,<sup>a</sup> Claudio M. Mastroianni<sup>a,c</sup>

Sonication for Microbial Detection in Cardiac Devices

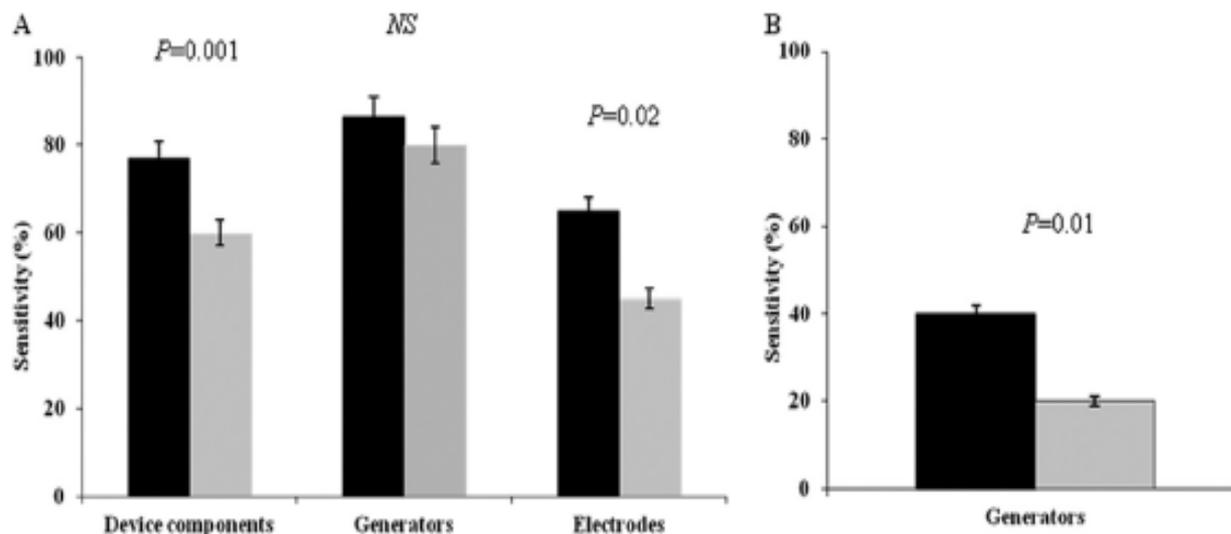


FIG 1 Sensitivity of sonication culture (black bars) and traditional culture (gray bars) for microbial detection in device components removed from infected (A) and uninfected (B) patients. Electrodes include atrial and ventricular electrodes. NS, not significant.

# Sonication of Explanted Cardiac Implants Improves Microbial Detection in Cardiac Device Infections

Alessandra Oliva,<sup>a</sup> Bich Lien Nguyen,<sup>b</sup> Maria T. Mascellino,<sup>a</sup> Alessandra D'Abramo,<sup>a</sup> Marco Iannetta,<sup>a</sup> Antonio Ciccaglioni,<sup>b</sup> Vincenzo Vullo,<sup>a</sup> Claudio M. Mastroianni<sup>a,c</sup>

TABLE 2. Pathogen detection in sonication (SC) and traditional (TC) culture

Type of infection	Components <i>n</i>	Microorganisms (n) <sup>a</sup>
Subjects with CDI ( <i>n</i> = 20)	60	
positive SC/ negative TC	10	<i>S. epidermidis</i> (6); <i>C. striatum</i> (2); <i>Brevundimonas</i> spp (1); <i>S. hominis</i> (1)
positive SC/ positive TC	36	<i>S. epidermidis</i> ( <i>n</i> = 26); <i>S. hominis</i> ( <i>n</i> = 3); <i>P. aeruginosa</i> ( <i>n</i> = 3); <i>S. aureus</i> ( <i>n</i> = 1); <i>S. haemolyticus</i> ( <i>n</i> = 1); <i>Bacillus</i> spp ( <i>n</i> = 1); <i>Klebsiella</i> spp ( <i>n</i> = 1)
negative SC/ positive TC	0	No bacterial detection
negative SC/ negative TC	14	No bacterial detection
Subjects without CDIs ( <i>n</i> = 20)	20	
positive SC/ negative TC	4	coagulase-negative staphylococci
positive SC/ positive TC	4	coagulase-positive staphylococcus ( <i>n</i> = 1); coagulase-negative staphylococci ( <i>n</i> = 3)
negative SC/ positive TC	0	No bacterial detection
negative SC/ negative TC	12	No bacterial detection

CDI: cardiac device infection.

<sup>a</sup> Polimicrobial cultures were found in 5 generators (*S. epidermidis*/*S. hominis* (*n* = 3); *S. epidermidis*/*Bacillus* spp (*n* = 1) *S. epidermidis*/*Klebsiella* spp).

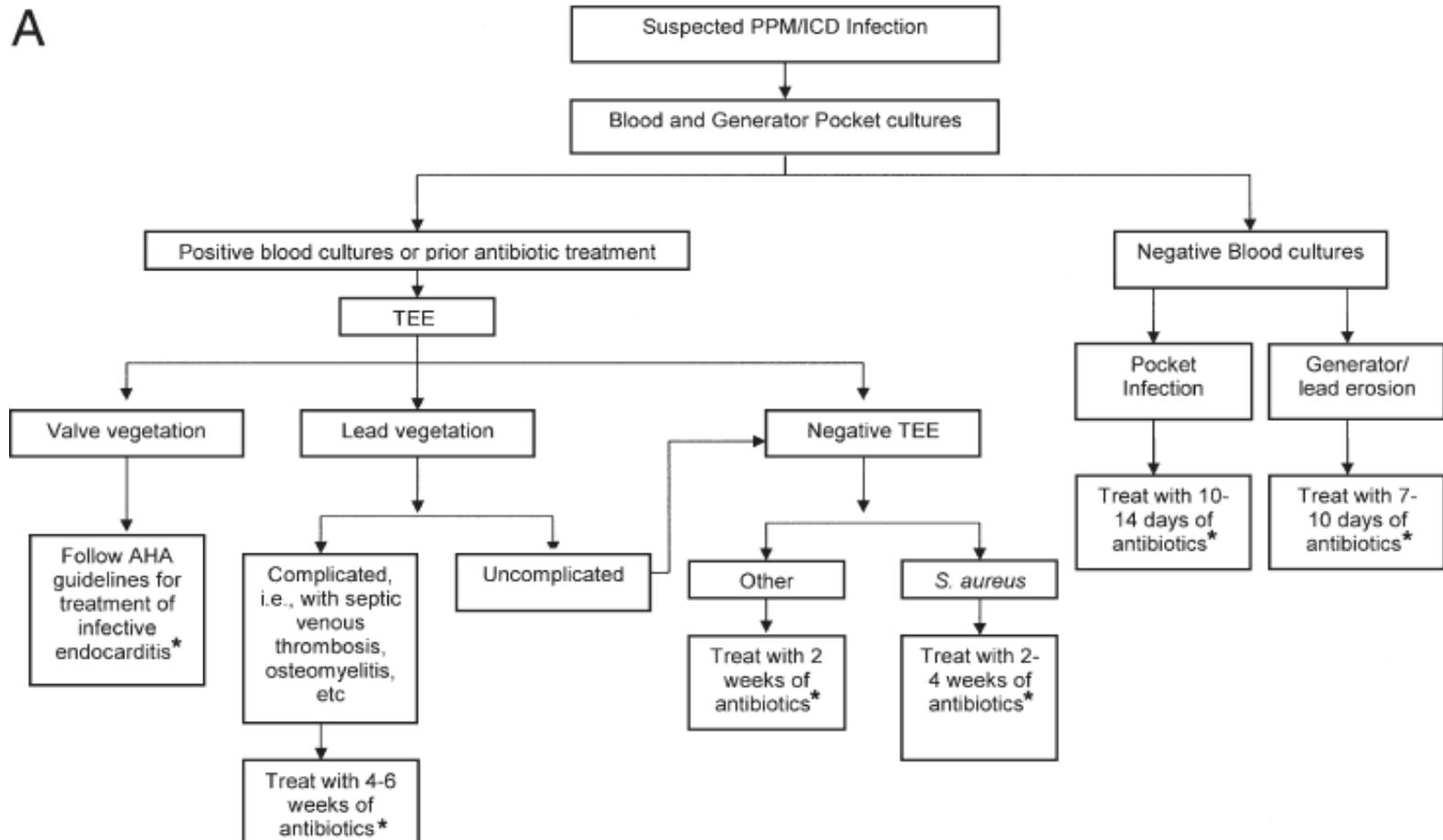
# Pacemaker Lead Endocarditis Due to Multidrug-Resistant *Corynebacterium striatum* Detected with Sonication of the Device<sup>∇</sup>

Alessandra Oliva,<sup>1</sup> Valeria Belvisi,<sup>1,2</sup> Marco Iannetta,<sup>1</sup> Carolina Andreoni,<sup>1</sup> Maria T. Mascellino,<sup>1</sup>  
Miriam Lichtner,<sup>1,2</sup> Vincenzo Vullo,<sup>1</sup> and Claudio M. Mastroianni<sup>1,2\*</sup>

*Corynebacterium striatum* is a commensal of human skin and has been recently recognized as an emerging pathogen. A case of nosocomial pacemaker lead endocarditis due to a multidrug-resistant *C. striatum* strain is described, highlighting the role of sonication as a diagnostic tool in cardiac device infections (CDIs).

# Management and Outcome of Permanent Pacemaker and Implantable Cardioverter-Defibrillator Infections

A



# Clinical Characteristics and Outcome of Infective Endocarditis Involving Implantable Cardiac Devices

**Table 2.** Characteristics of Patients With Cardiac Device Infective Endocarditis and With or Without Cardiac Device Removal During Hospitalization

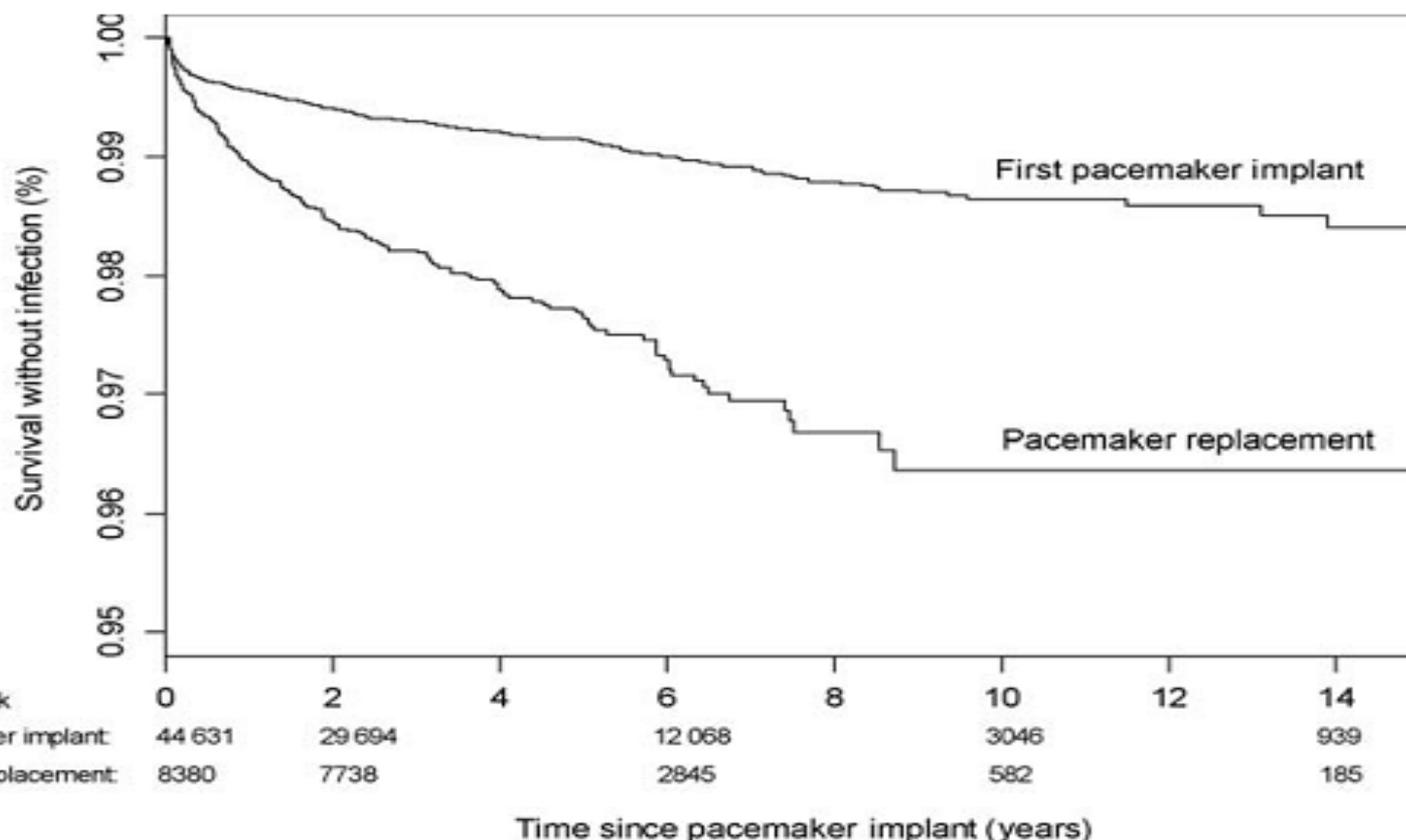
Variable	Device Removal (n = 141)	No Device Removal (n = 34)	P Value
Age, median (IQR), y	70.1 (59.8-76.3)	70.4 (68.1-78.6)	.13
Diabetes mellitus	39 (27.7)	8 (23.5)	.76
Hemodialysis	9 (6.4)	2 (5.9)	.93
History of cancer	13 (9.2)	6 (17.7)	.14
Transferred from another hospital	61 (43.3)	16 (47.1)	.61
Positive blood cultures	114 (80.9)	34 (100)	.006
<i>Staphylococcus aureus</i>	47 (33.3)	15 (44.1)	.24
Coagulase-negative staphylococci	46 (32.6)	10 (29.4)	.72
Health care-associated infection	62 (44.0)	19 (55.9)	.21
Concomitant valve vegetation	54 (38.3)	9 (26.5)	.20
Heart failure	18 (12.8)	9 (26.5)	.03
Pulmonary embolism	14 (9.9)	2 (5.9)	.46
In-hospital mortality	18 (12.8)	8 (23.5)	.12
1-y mortality	28 (19.9)	13 (38.2)	.02

Abbreviation: IQR, interquartile range.

# Infection after pacemaker implantation: infection rates and risk factors associated with infection in a population-based cohort study of 46299 consecutive patients

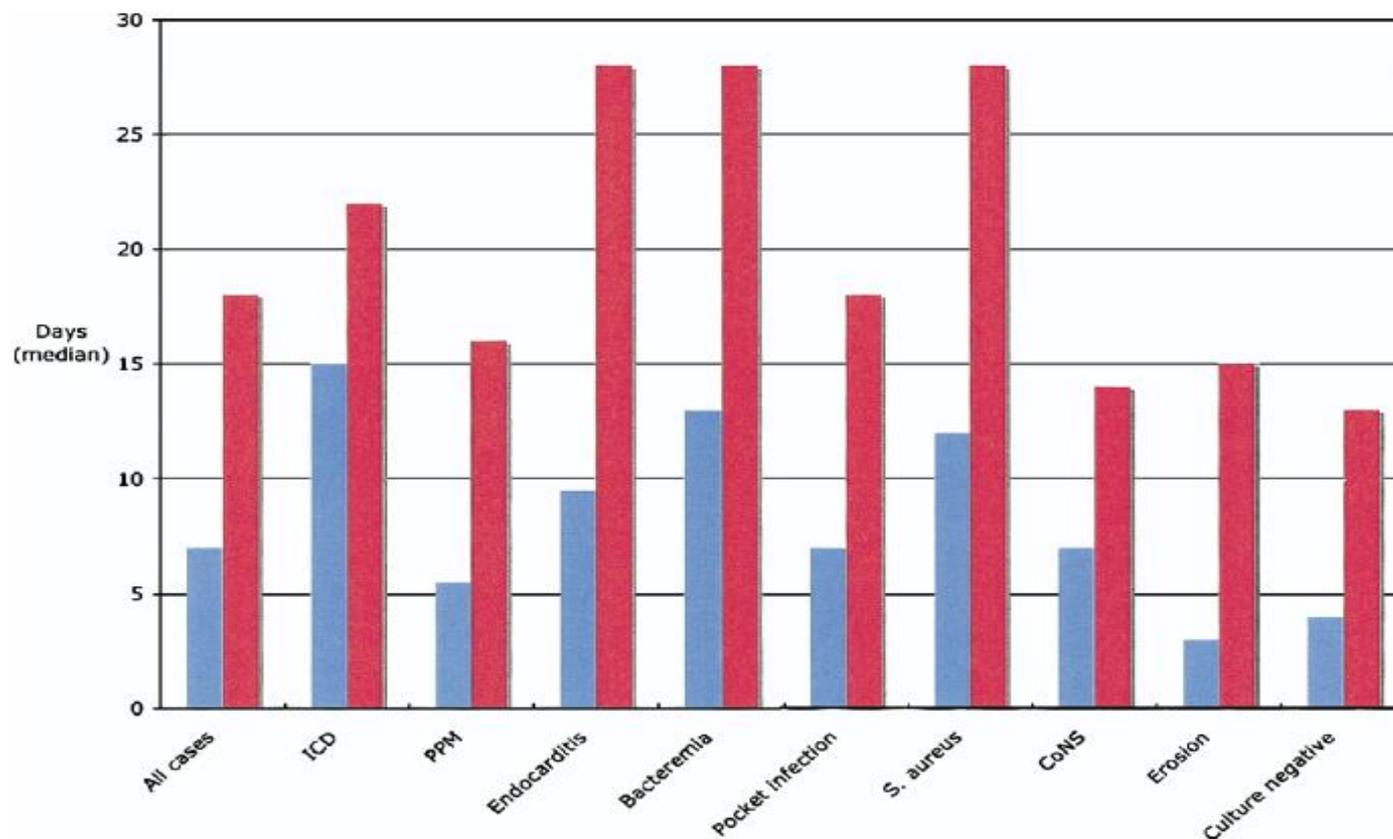


European Heart Journal (2011) 32, 991–998  
doi:10.1093/eurheartj/ehq497



# Management and Outcome of Permanent Pacemaker and Implantable Cardioverter-Defibrillator Infections

Time from removal and reimplantation of disposal (Blue) and time of antibiotic therapy (red)



# Antibiotic review: sepsis from catheter-related bloodstream infection (CRBSI)

Infection	Example antibiotic regimens
CRBSI	Daptomycin or vancomycin <sup>1</sup> + antipseudomonal $\beta$ -lactam <sup>2,3</sup> +/- aminoglycoside or rifampin <sup>4</sup>
Fungemia risk factors	+ echinocandin <sup>5</sup> or fluconazole

1 high rates of vancomycin MIC  $\geq$  2  $\mu$ g/mL

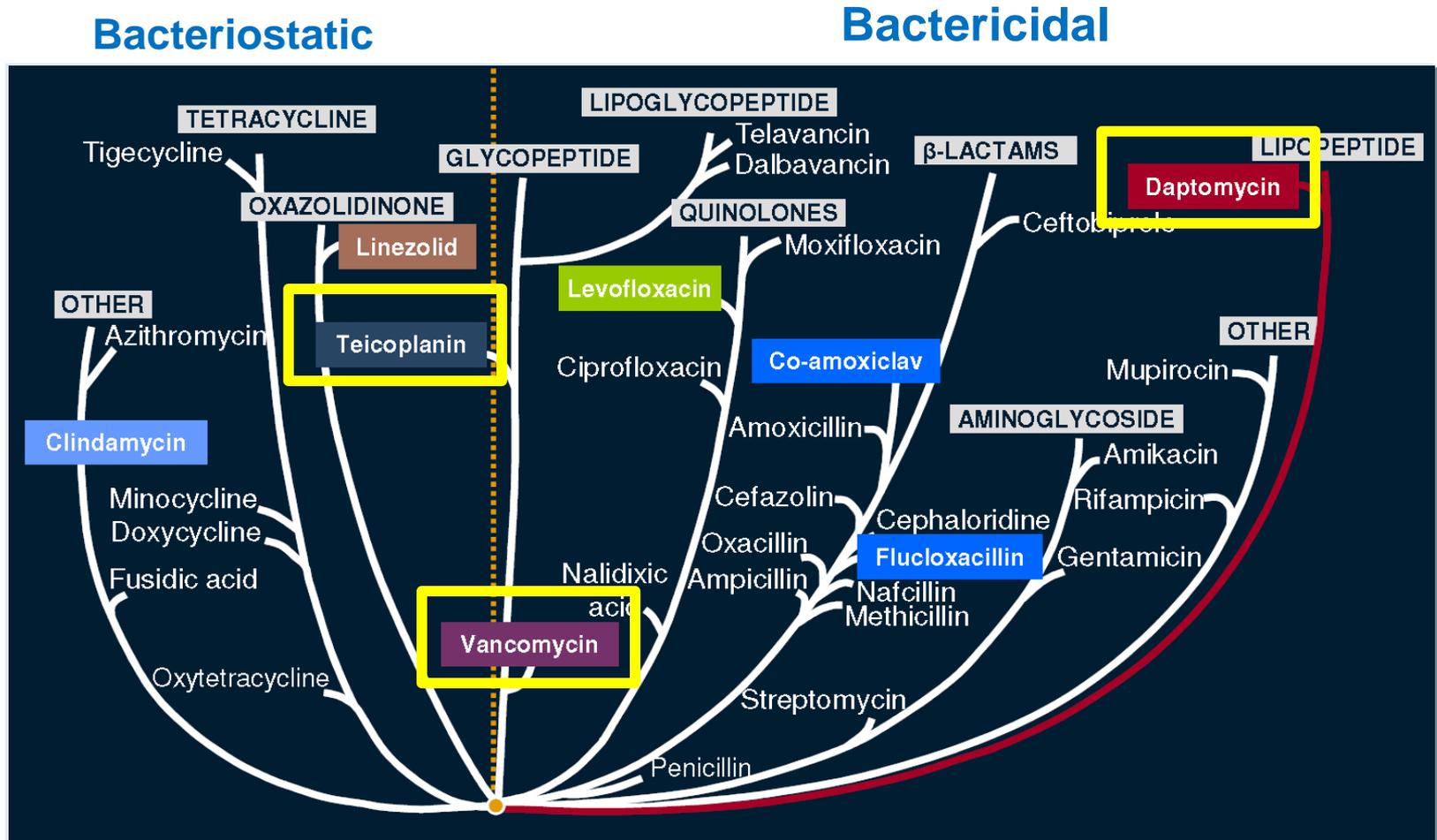
2 piperacillin/tazobactam, cefepime

3 meropenem, imipenem, doripenem

4 gentamicin, tobramycin, amikacin

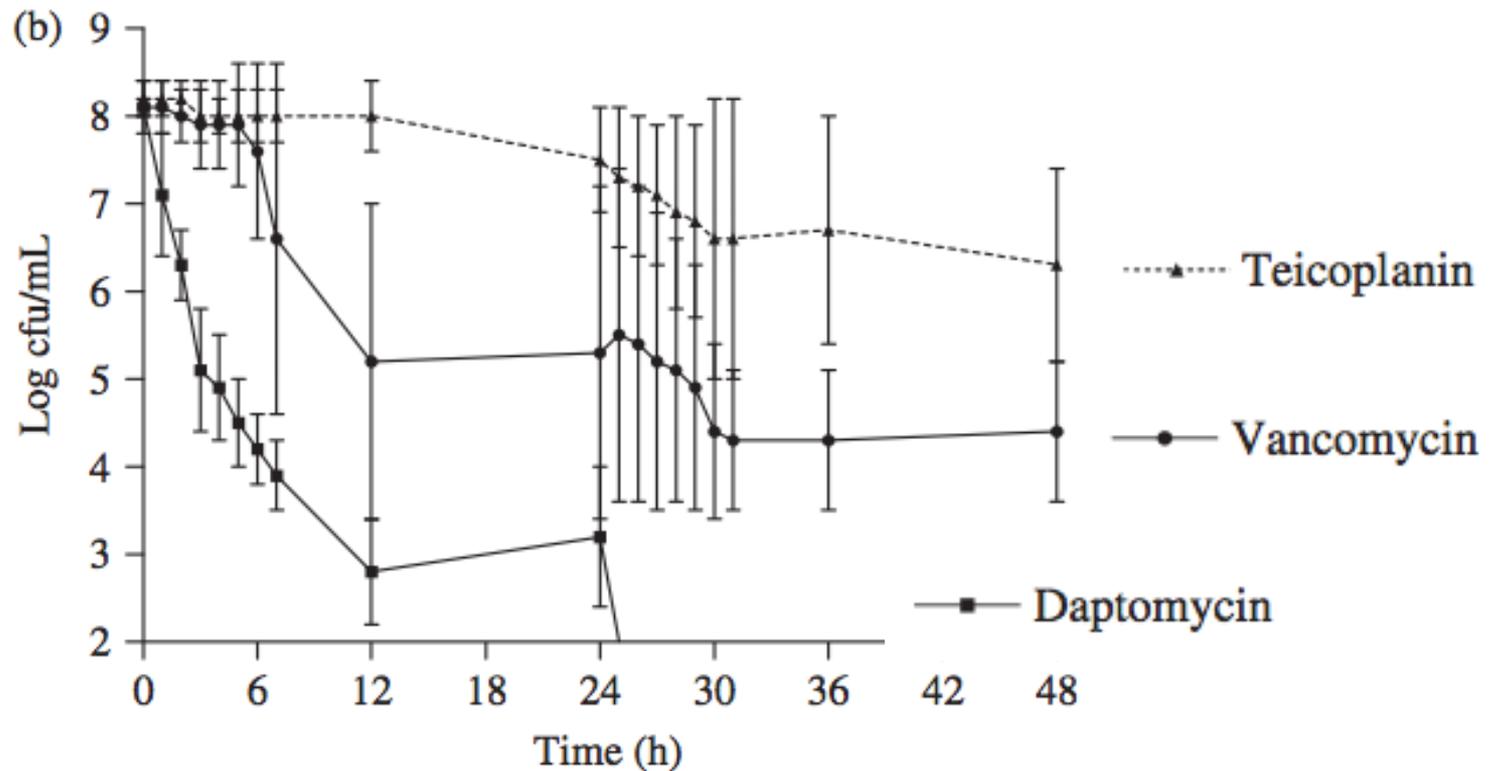
5 caspofungin, micafungin, anidulafungin

# Bacteriostatic and bactericidal antibacterials active against Gram positive

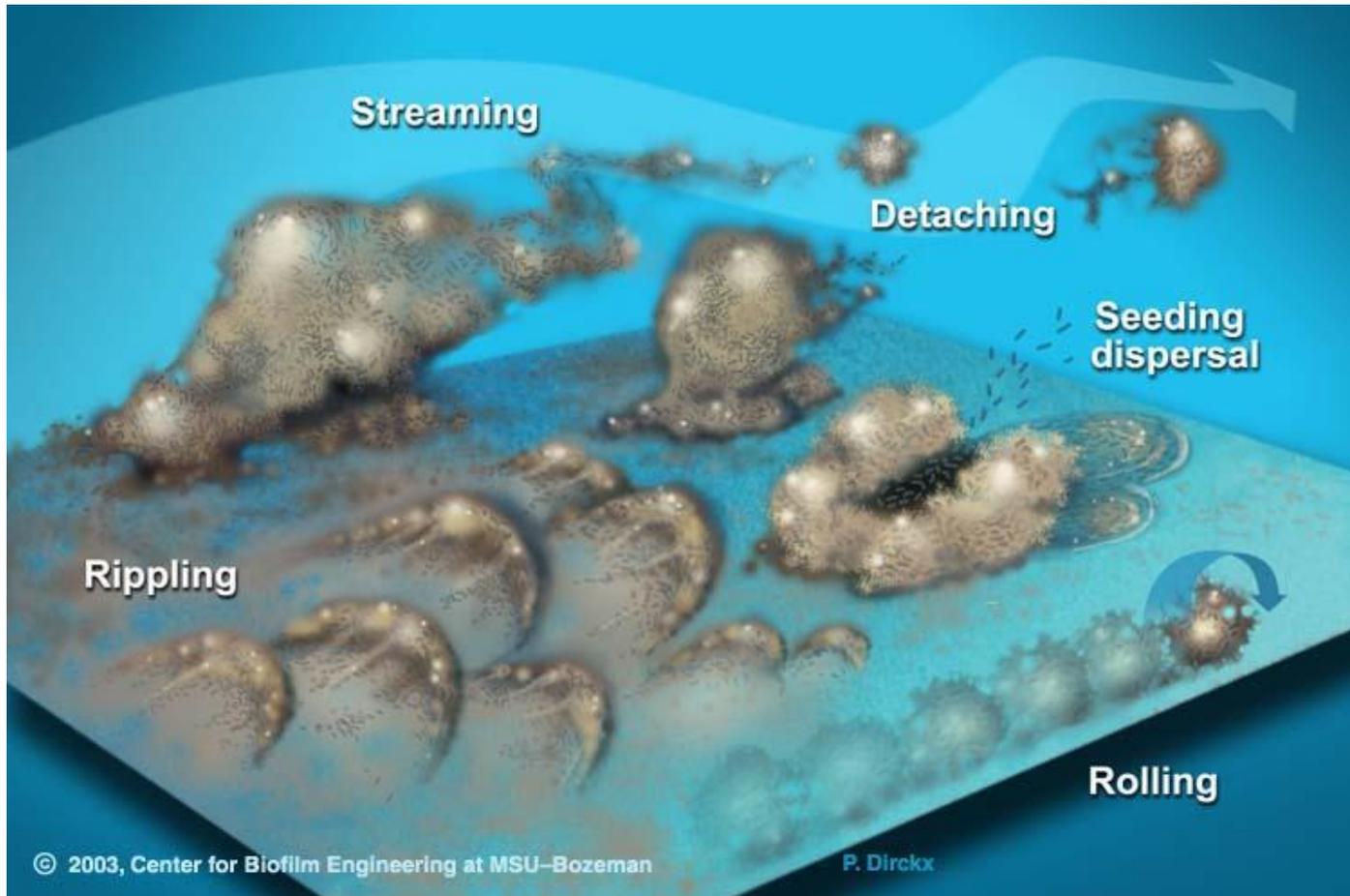


# Comparative antibacterial effects of daptomycin, vancomycin and teicoplanin studied in an *in vitro* pharmacokinetic model of infection

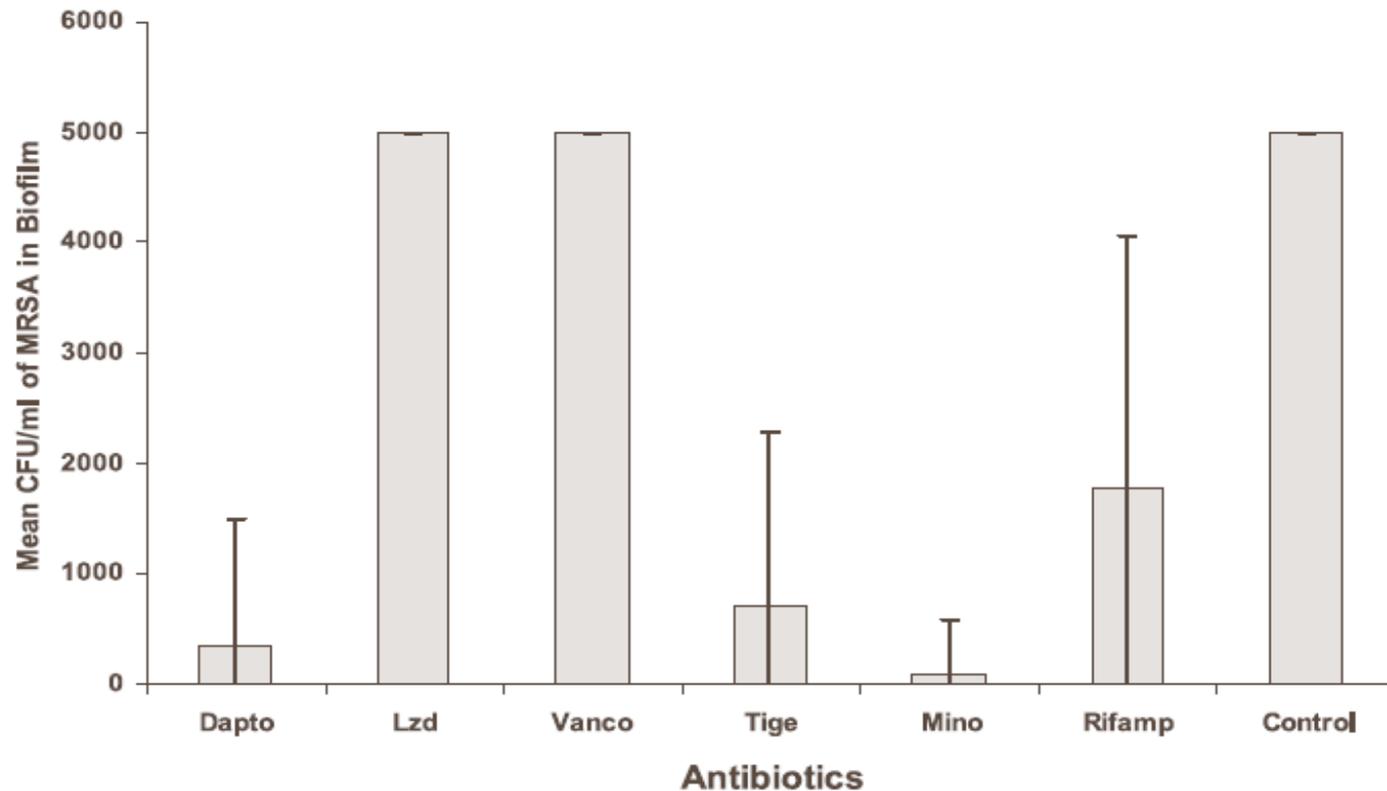
Karen E. Bowker\*, Alan R. Noel and Alasdair P. MacGowan



# The biofilm problem



# Comparative Activities of Daptomycin, Linezolid, and Tigecycline against Catheter-Related Methicillin-Resistant *Staphylococcus* Bacteremic Isolates Embedded in Biofilm<sup>▽</sup>



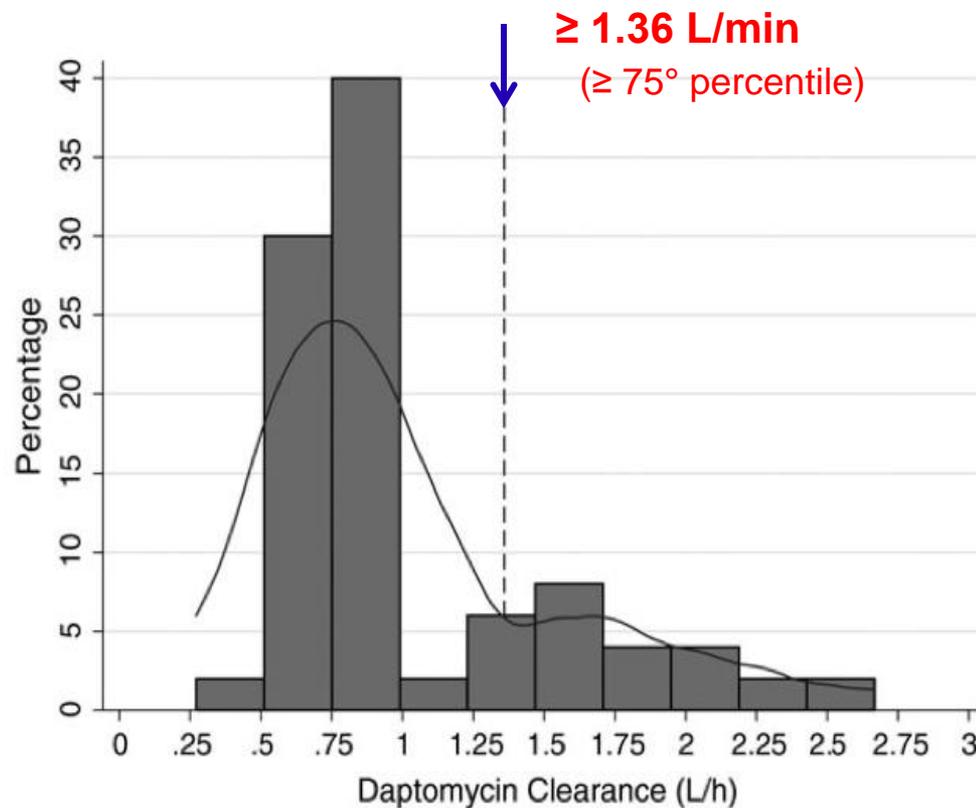
# A retrospective case control analysis of patients with staphylococcal infections receiving daptomycin or glycopeptides therapy

Variable	Daptomycin (n = 43)	Vancomycin or teicoplanin (n=63)	p =
Presence of at least 2 comorbidities	30 (69.8%)	44 (69.8%)	1.0
Renal replacement therapy	8 (18.6%)	5 (7.9%)	0.13
Creatinine clearance < 35 ml/h	6 (14%)	14 (22.2%)	0.32
Recent surgery (previous 30 days)	16 (37.2%)	9 (14.3%)	<b>0.008</b>
Removable intravascular device	34 (79.1%)	30 (47.6%)	<b>0.001</b>
CVC	24 (55.8%)	23 (36.5%)	
Pace-maker	10 (23.3%)	7 (11.1%)	
Fever	23 (53.5%)	25 (39.7%)	0.17
Immunosuppressive therapy	4 (9.3%)	3 (4.8%)	0.64
Severe sepsis or septic shock	19 (44.2%)	25 (39.7%)	0.69
SOFA score (mean)	2.65	2.04	0.15
Mean duration of antibiotic therapy (DAYS)	16.4	22.2	<b>&lt;0.001</b>
Mean length of hospital stay (days)	26.5	31.8	<b>0.04</b>

**Daptomycin: which dosages to  
what patients?**

# Considerations for Higher Doses of Daptomycin in Critically Ill Patients With Methicillin-Resistant *Staphylococcus aureus* Bacteremia

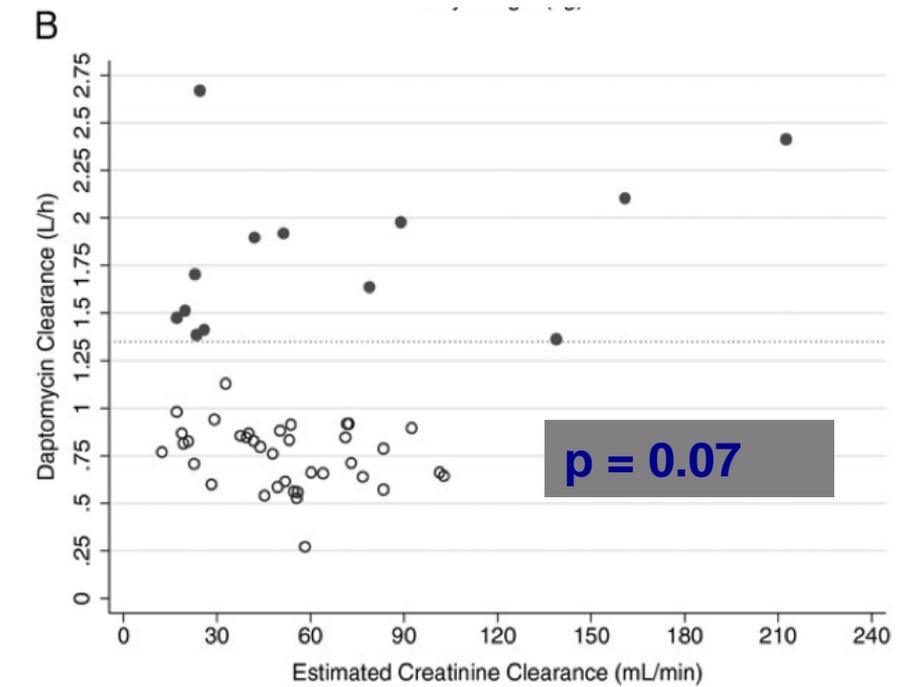
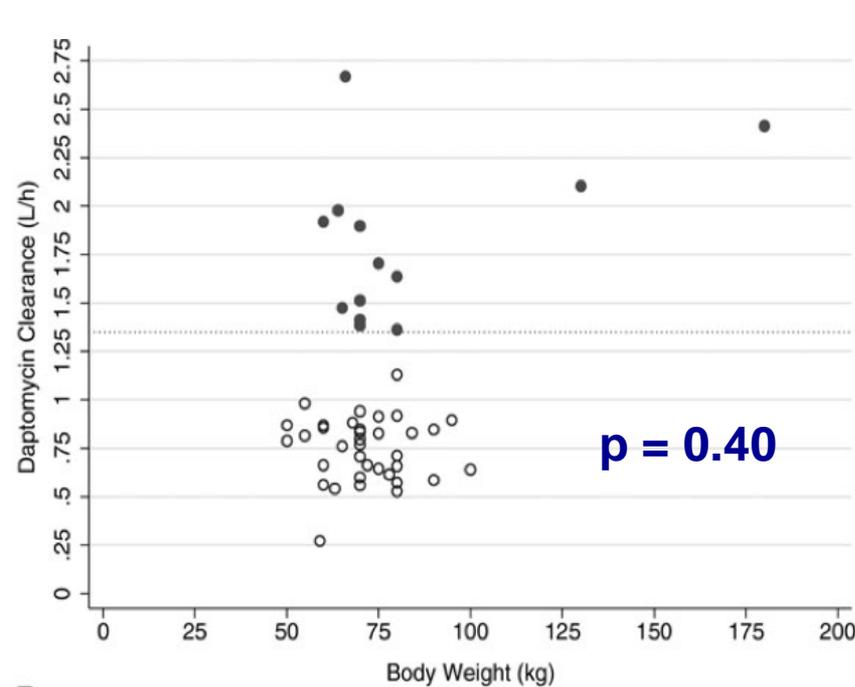
Marco Falcone,<sup>1</sup> Alessandro Russo,<sup>1</sup> Mario Venditti,<sup>1</sup> Andrea Novelli,<sup>2</sup> and Manjunath P. Pai<sup>3</sup>



# Considerations for Higher Doses of Daptomycin in Critically Ill Patients With Methicillin-Resistant *Staphylococcus aureus* Bacteremia

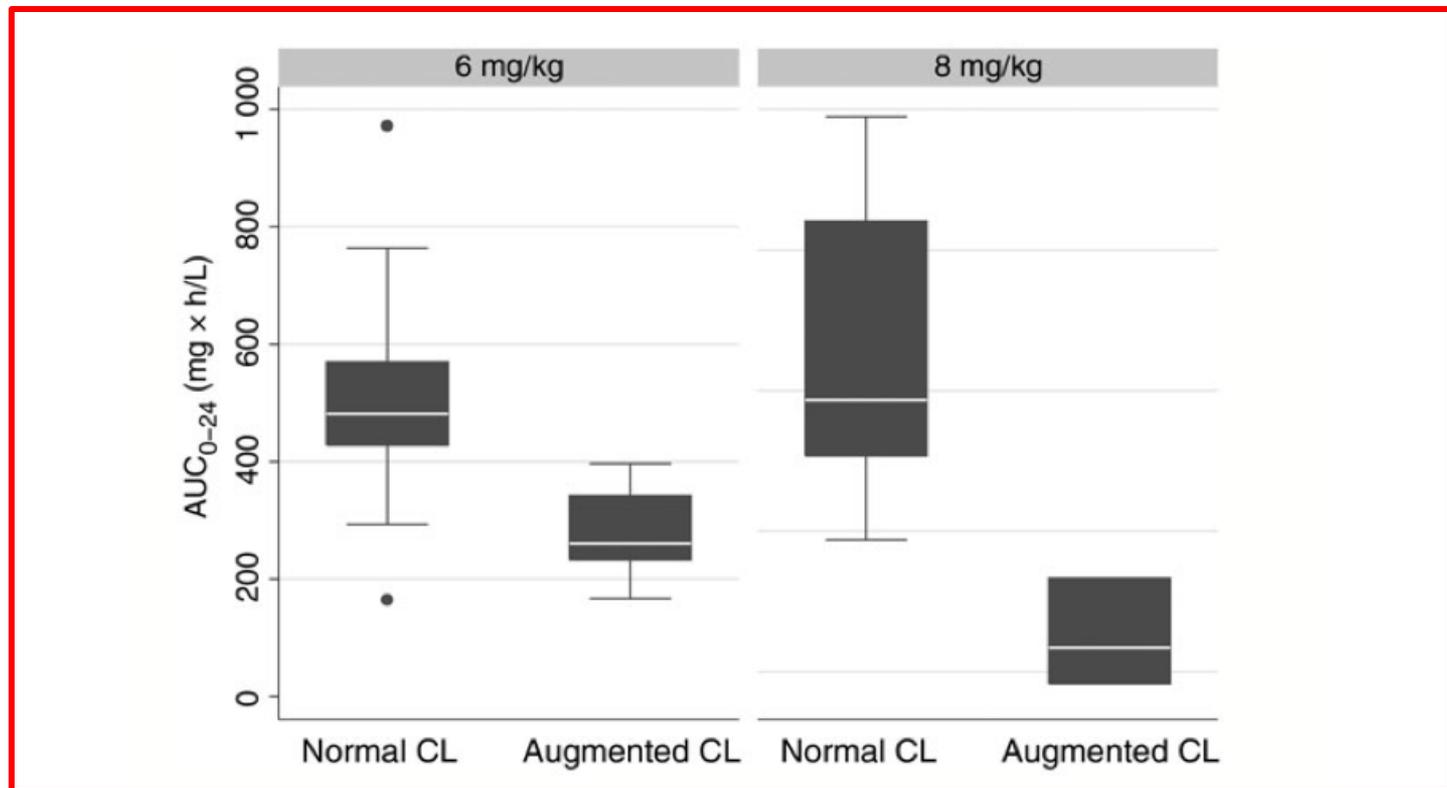
Marco Falcone,<sup>1</sup> Alessandro Russo,<sup>1</sup> Mario Venditti,<sup>1</sup> Andrea Novelli,<sup>2</sup> and Manjunath P. Pai<sup>3</sup>

## Correlation between daptomycin clearance, body weight, and CRCl



# Considerations for Higher Doses of Daptomycin in Critically Ill Patients With Methicillin-Resistant *Staphylococcus aureus* Bacteremia

Marco Falcone,<sup>1</sup> Alessandro Russo,<sup>1</sup> Mario Venditti,<sup>1</sup> Andrea Novelli,<sup>2</sup> and Manjunath P. Pai<sup>3</sup>



# Considerations for Higher Doses of Daptomycin in Critically Ill Patients With Methicillin-Resistant *Staphylococcus aureus* Bacteremia

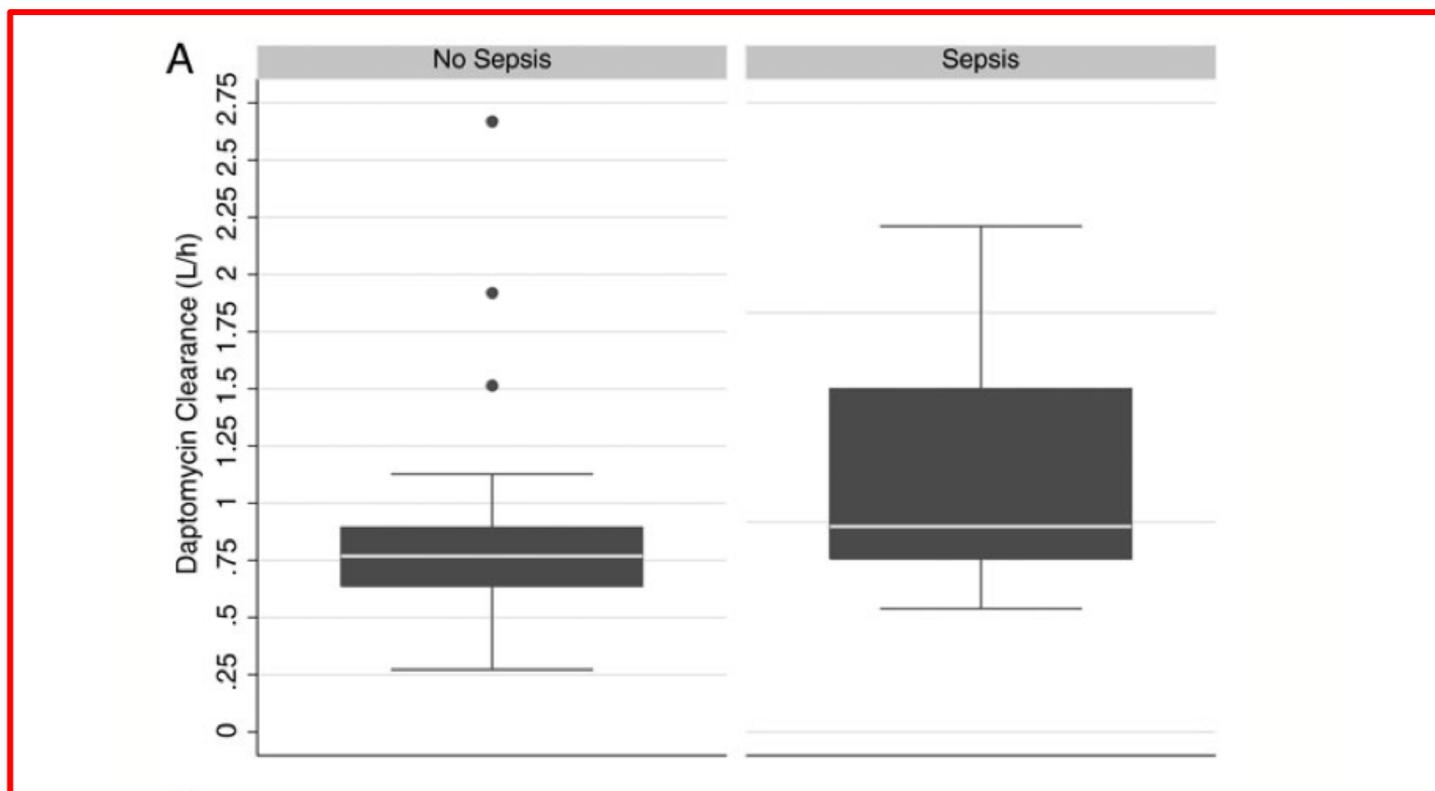
Marco Falcone,<sup>1</sup> Alessandro Russo,<sup>1</sup> Mario Venditti,<sup>1</sup> Andrea Novelli,<sup>2</sup> and Manjunath P. Pai<sup>3</sup>

Variable	Augmented CL (n = 13)	Normal CL (n = 37)	P Value
<b>Type of infection</b>			
Bacteremia— endocarditis	13 (100%)	8 (21.6%)	<.001
Skin and soft tissue infection	0	20 (54.1%)	<.001
Prosthetic joint infection	0	6 (16.2%)	<.001
Osteomyelitis	0	4 (10.8%)	<.001
<b>Causative pathogen</b>			
MRSA	11 (84.6%)	2 (5.2%)	<.001
MRSE	0	8 (21%)	<.001
MRSH	0	7 (18.4%)	<.001
Other	2 (15.4%)	14 (36.8%)	<.001
<b>Comorbidities and outcomes</b>			
COPD	8 (61.5%)	21 (55.2%)	.71
Heart failure	7 (53.8%)	19 (50%)	.29
Diabetes mellitus	7 (53.8%)	18 (47.3%)	.18
Neoplasm	2 (15.3%)	4 (10.5%)	.68
Chronic liver disease	3 (23%)	7 (18.4%)	.08
Presence of at least 2 comorbidities	9 (69.2%)	22 (57.8%)	.09
Recent surgery, previous 30 d	6 (46.1%)	9 (24.3%)	.07
ICU acquisition of infection	8 (61.5%)	12 (31.5%)	.04
Severe sepsis or septic shock <sup>a</sup>	13 (100%)	9 (24.3%)	.01
SOFA score, mean (SD)	3.31 (1.03)	2.11 (0.84)	<.001
Mean length of hospital stay, days	36.8	22.5	<.001
In-hospital mortality	4 (30.7%)	4 (10.8%)	<.001

**Clinical features of patients with augmented daptomycin clearance**

# Considerations for Higher Doses of Daptomycin in Critically Ill Patients With Methicillin-Resistant *Staphylococcus aureus* Bacteremia

Marco Falcone,<sup>1</sup> Alessandro Russo,<sup>1</sup> Mario Venditti,<sup>1</sup> Andrea Novelli,<sup>2</sup> and Manjunath P. Pai<sup>3</sup>



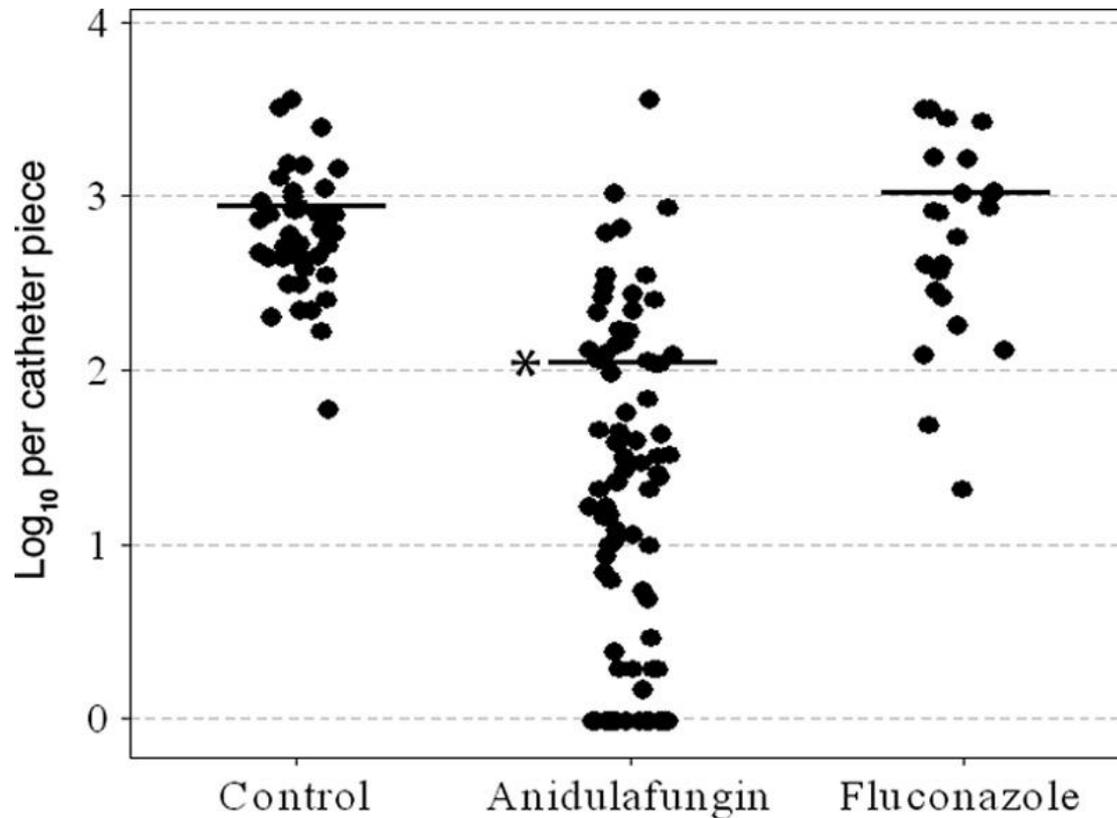
# Probability of target attainment (PTA) and toxicity in patients with severe sepsis or septic shock at Monte Carlo simulation

Daily Dose	% CFR Based on $AUC_{0-24}/MIC$			% Probability $C_{min} \geq 24.3$ mg/L
	$\geq 579$	$\geq 666$	$\geq 753$	
Weight-based dosing				
6 mg/kg/d	87.3	82.1	77.2	0.08
8 mg/kg/d	94.1	91.3	88.0	0.78
10 mg/kg/d	97.1	95.4	93.4	2.64
Fixed dosing				
500 mg/d	93.1	89.2	84.8	0.02
750 mg/d	98.4	97.3	95.6	1.26
1000 mg/d	99.5	99.1	98.5	6.20

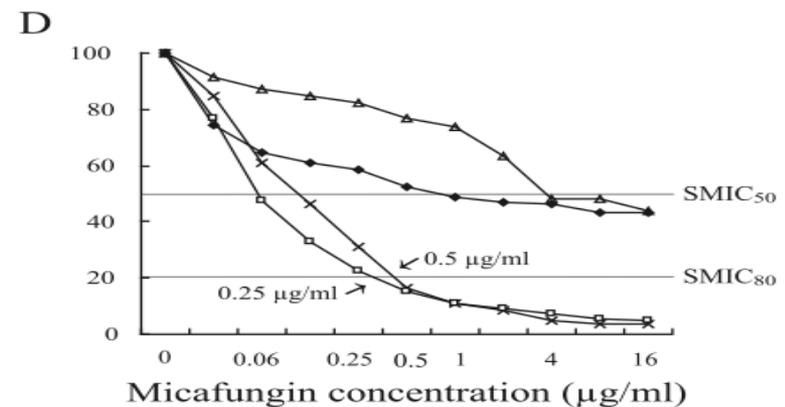
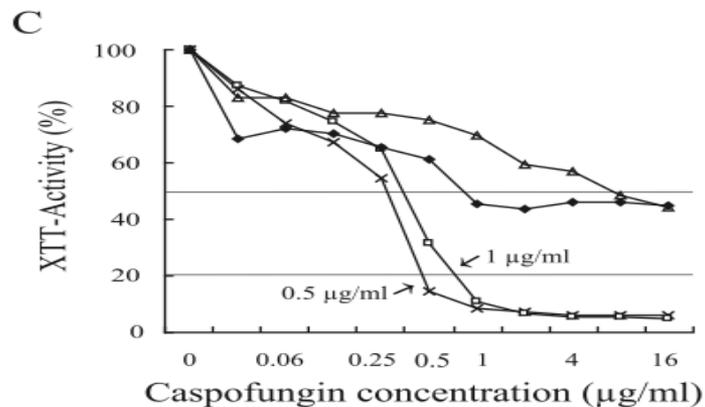
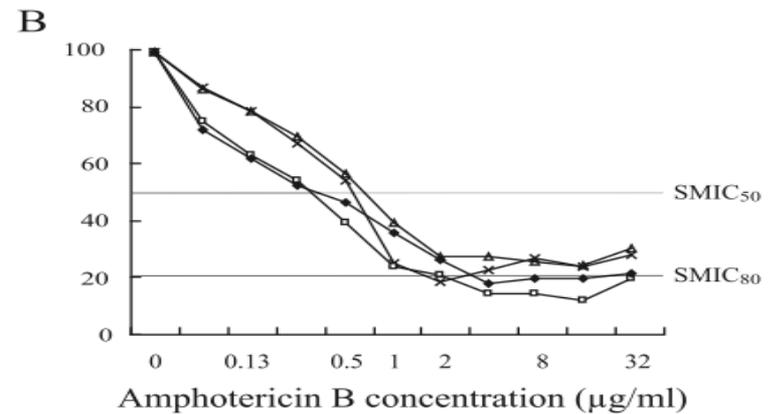
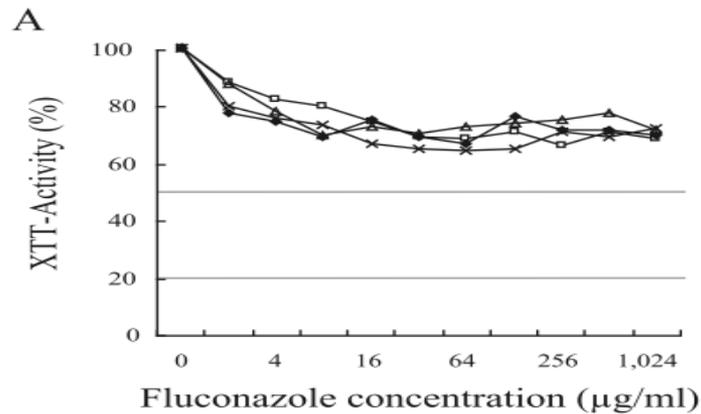
# Probability of target attainment (PTA) and toxicity in patients **without sepsis** at Monte Carlo simulation

Daily Dose	% CFR Based on AUC <sub>0-24</sub> /MIC			% Probability C <sub>min</sub> ≥ 24.3 mg/L
	≥579	≥666	≥753	
Weight-based dosing				
6 mg/kg/d	94.8	92.3	89.5	1.52
8 mg/kg/d	97.9	96.7	95.1	4.88
10 mg/kg/d	99.1	98.6	97.6	11.0
Fixed dosing				
500 mg/d	96.8	95.1	92.9	1.38
750 mg/d	99.3	98.8	98.1	7.64
1000 mg/d	99.8	99.7	99.4	19.3

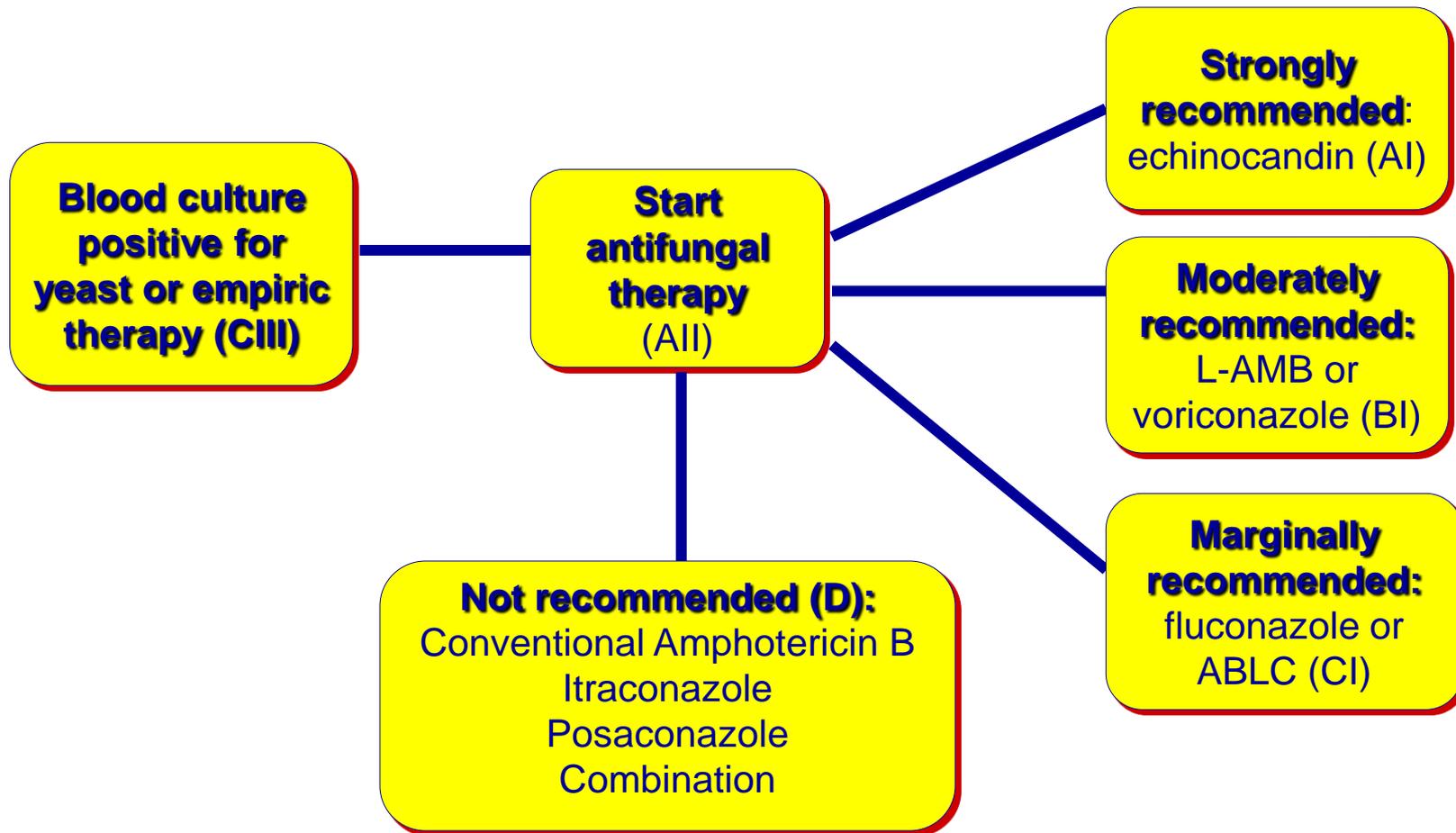
# Effect of antifungal treatment on mature *Candida* biofilms\*



# Antifungals and biofilm

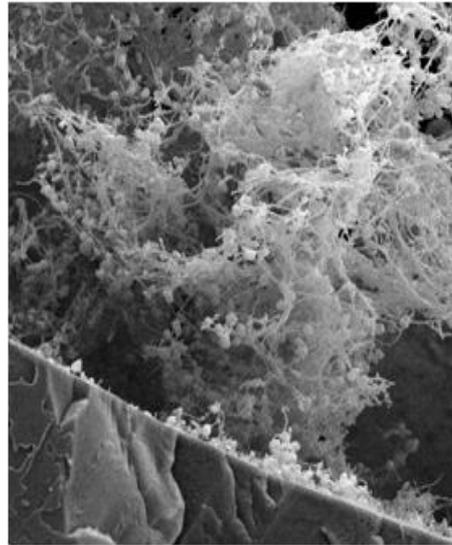


# Treatment of *Candida* in non-neutropenic patients (ESCMID guidelines 2012)

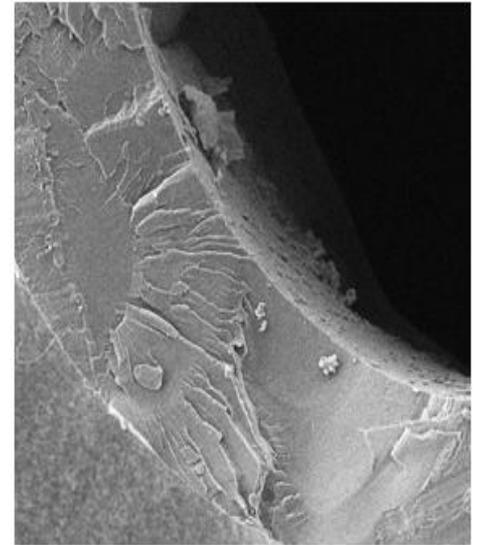


# Removal of all foreign objects correlates with better outcomes\*

*C. albicans* biofilms formed on an implanted medical device ex. CVC, urinary catheter, ETT, prosthetic heart valve, or pacemaker play a role in the persistence and proliferation of Candidiasis. Cells in biofilms are much more resistant to antifungal agents\*. The echinocandins have penetration and action in *Candida* biofilms and thus may have an advantage in this setting\*\*.



Wild-Type



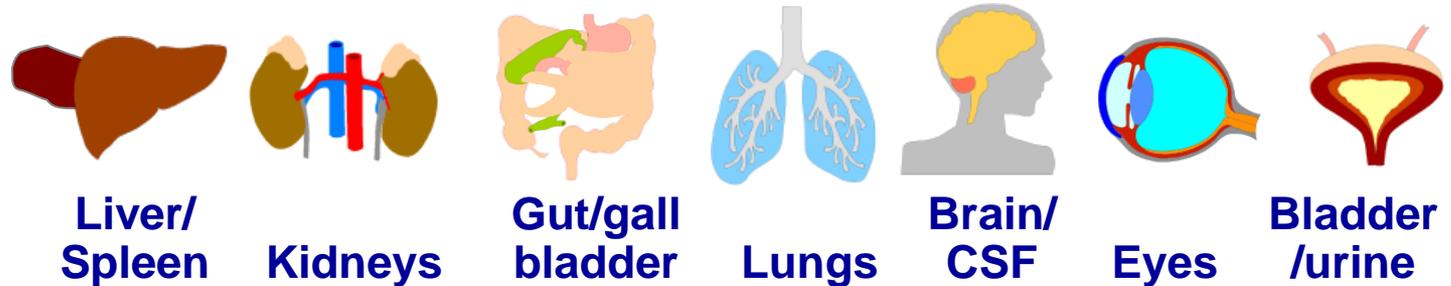
*eap1-1/eap1-2*

*C. albicans* adhesion as a virulence factor

\* Nucci M et al. 2002. *CID*; 34: 591-599.

\*\* Kuhn et al. 2002. *Antimicrob Agents Chemother*; 46:1773-1780.

# Antifungal PK: Drug Distribution



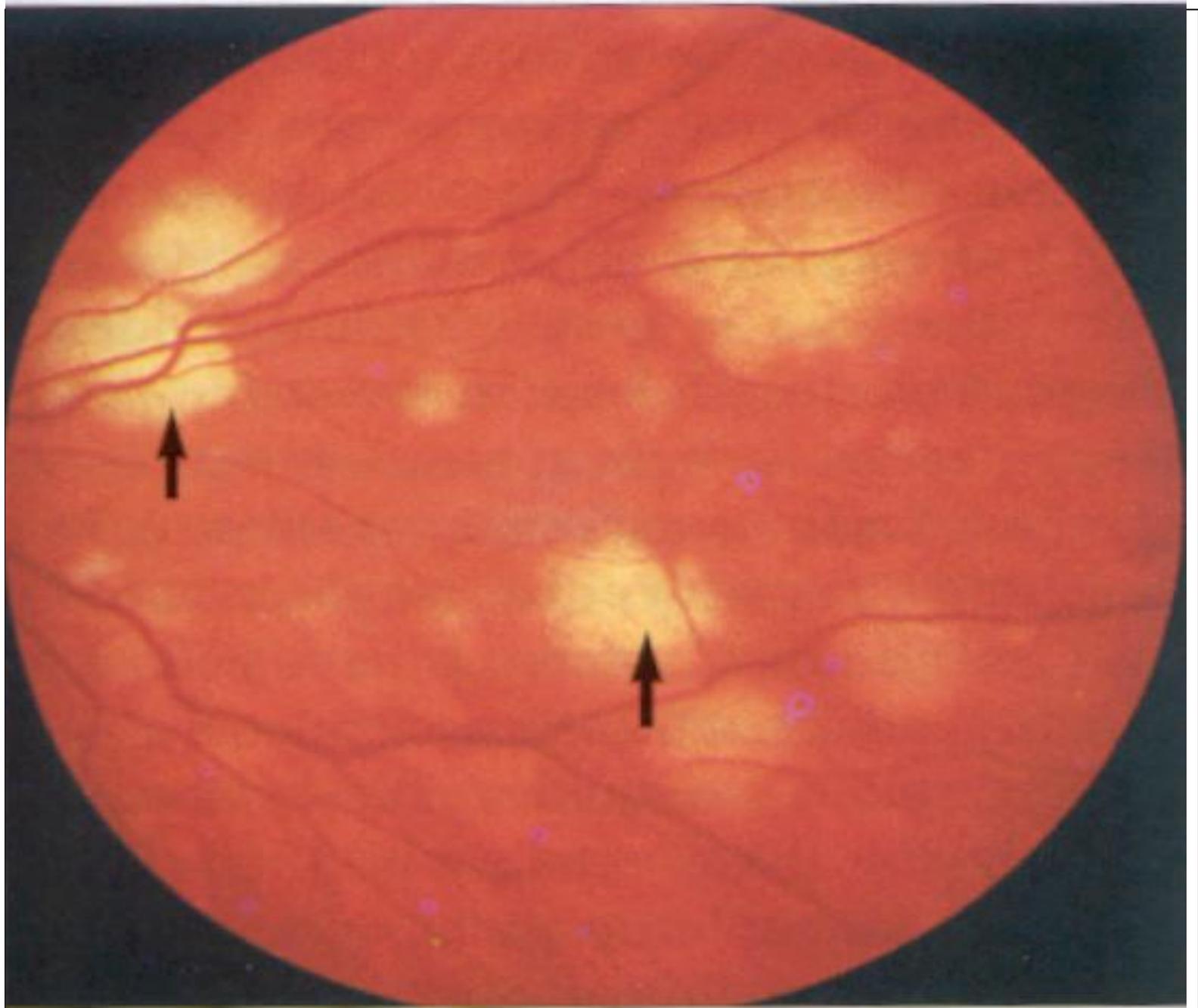
	Liver/ Spleen	Kidneys	Gut/gall bladder	Lungs	Brain/ CSF	Eyes	Bladder /urine
<b>AMB</b>	+	+	+	+	-	-	-
<b>5FC</b>	+	+	+	+	+	+	+
<b>FLU</b>	+	+	+	+	+	+	+
<b>ITR</b>	+	+	+	+	-	-	-
<b>VOR</b>	+	+	+	+	+	+	-
<b>POS*</b>	+	+	+	+	-	-	-
<b>Echino</b>	+	+	+	+	-	-	-

+, ≥50% of serum concentrations.

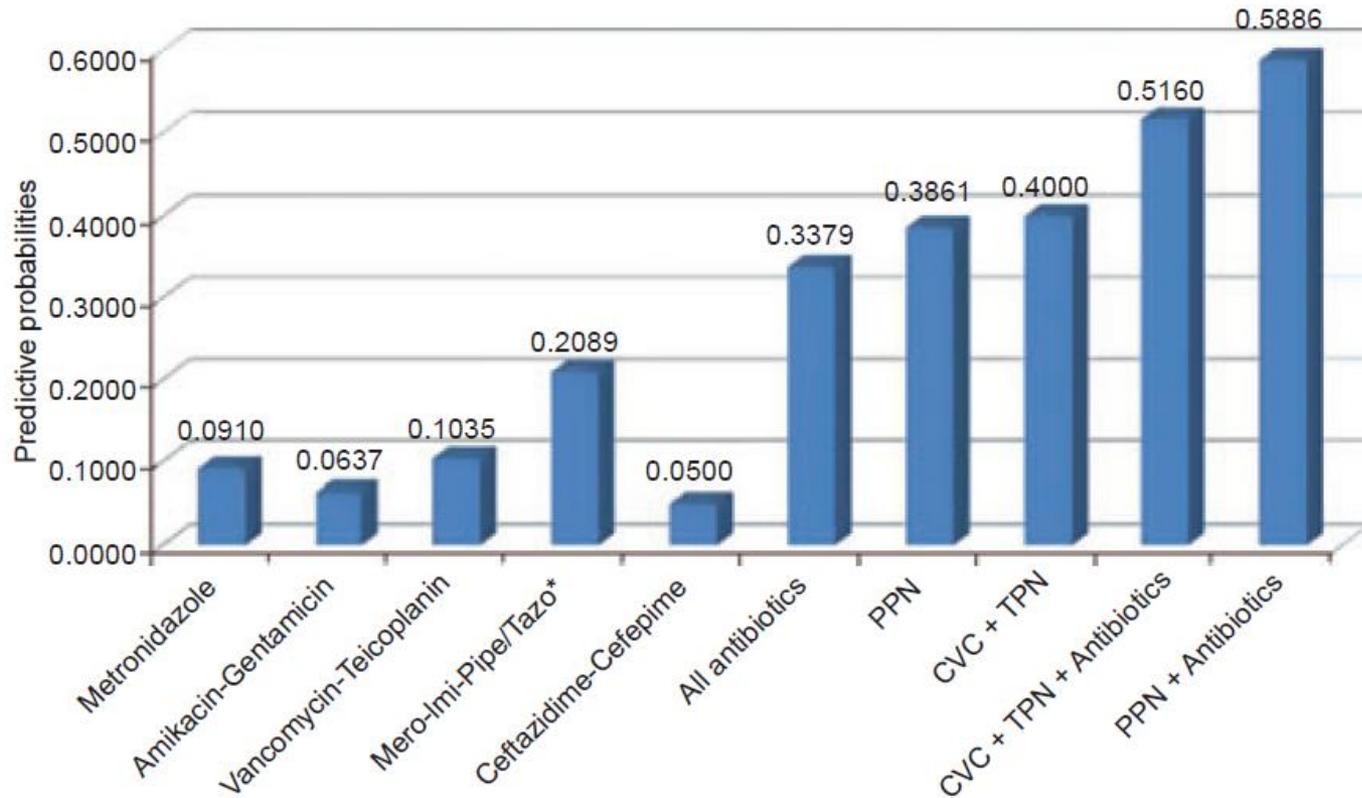
-, <10% of serum concentrations.

\*Predicted.

1. Dodds-Ashley ES, et al. *Clin Infect Dis*. 2006;43:S28-S39.
2. Groll AH, et al. *Adv Pharmacol*. 1998;44:343-500.
3. Eschenauer G, et al. *Ther Clin Risk Manage*. 2007;3:71-97.



# Peripheral and total parenteral nutrition as the strongest risk factors for nosocomial candidemia in elderly patients: a matched case-control study



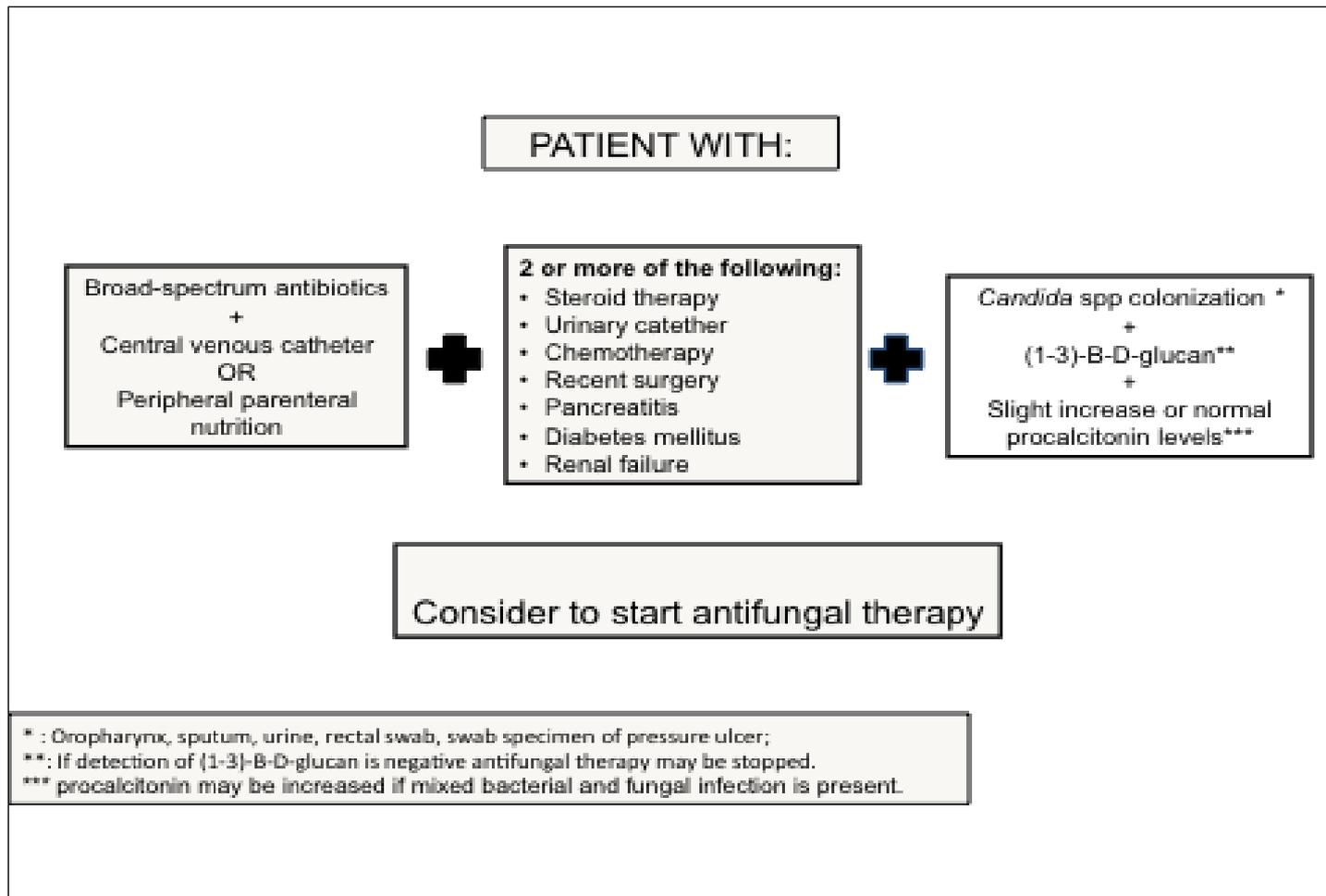
# Candidemia Subsequent to Severe Infection Due to *Clostridium difficile*: Is There a Link?

Maurizio Guastalegname,<sup>a</sup> Alessandro Russo,<sup>a</sup>  
 Marco Falcone, Simone Giuliano, and  
 Mario Venditti

Patient Number, Sex, Age, Ward of Hospitalization	Cause of Hospital Admission	Previous Antimicrobial Chemotherapy	Antimicrobial Regimen for Primary CDI	CDI Relapse (Time to Relapse <sup>a</sup> )	Antimicrobial Regimen for CDI Relapse	Time at Risk for Candidemia <sup>b</sup>	Candidemia Risk Factors	<i>Candida</i> Species and Antifungal Chemotherapy	Clinical Outcome
1, female, 78, medical ward A	HCAP (patient coming from a nursing home)	Piperacilin/tazobactam, ciprofloxacin	Oral vancomycin (125 mg qid)	Yes (24 d)	Oral vancomycin (125 mg qid) + IV metronidazole (500 mg qid)	16 d	<ul style="list-style-type: none"> <li>• PICC</li> <li>• TPN</li> <li>• Type 2 diabetes mellitus</li> </ul>	<i>Candida glabrata</i>	Survived
2, female, 84, medical ward B	Acute heart failure	Ceftriaxone	Oral vancomycin (125 mg qid)	Yes (37 d)	Oral vancomycin (125 mg qid) + IV metronidazole (500 mg qid)	44 d	<ul style="list-style-type: none"> <li>• PICC</li> <li>• TPN</li> <li>• Chronic renal failure</li> <li>• Glucocorticoids</li> </ul>	<i>Candida albicans</i>	Death at day 3 of antifungal therapy
3, male, 87, medical ward C	Pyelonephritis	Levofloxacin	Oral vancomycin (125 mg qid)	Yes (21 d)	Oral vancomycin (125 mg qid) + IV metronidazole (500 mg qid)	10 d	<ul style="list-style-type: none"> <li>• PICC</li> <li>• TPN</li> <li>• Chronic renal failure</li> </ul>	<i>Candida albicans</i>	Survived
4, male, 82, medical ward C	Hypothyroidism and hypokalemia	Levofloxacin	Oral vancomycin (125 mg qid)	Yes (32 d)	Oral vancomycin (125 mg qid) + IV metronidazole (500 mg qid)	13 d	<ul style="list-style-type: none"> <li>• PICC</li> <li>• TPN</li> </ul>	<i>Candida parapsilosis</i>	Death at day 13 of antifungal therapy

# Identification and management of invasive mycoses in Internal Medicine: a road-map for physicians

Falcone M, Concia E, Iori I, Lo Cascio G, Mazzone A, Pea F, Violi F, and Venditti M



# Conclusions

- ✓ Intravascular devices are a major cause of infection in western countries
- ✓ Use algorithm for appropriate diagnosis and therapy
- ✓ Importance of biofilm activity of antibiotics and antifungals used
- ✓ Consider the patient and individualize therapy