

V CONGRESSO NAZIONALE ORTHOPEA



**Qualità ed appropriatezza
Better and Faster**

**Paolo Viganò
Infettivologo
HMD - Castellanza**

**TRATTAMENTO
DELLE INFEZIONI PROTESICHE:
STATO DELL'ARTE**

Milano 16-17 settembre 2021

CAPITOLO 1 | Il RIAP nel contesto del RIPI: aggiornamento e attività 2018-2019






Figura 1.2. Partecipanti al RIAP (novembre 2019)

riap

Registro Italiano ArtroProtesi

Report
Annuale 2019
Compendio



-  Raccolta dati attiva
-  Raccolta dati in fase di avvio
-  Registro regionale/provinciale istituito per legge
-  Ospedale singolo
-  ASL



INFEZIONE PROTESI ORTOPEDICHE

Continuo incremento della chirurgia protesica
(Italia: oltre 175.000 protesi /anno)

→ Incremento delle protesi impiantate
in soggetti in età avanzata
e con fattori di rischio per infezioni



**DIAGNOSIS AND PREVENTION OF PERIPROSTHETIC
JOINT INFECTIONS
CLINICAL PRACTICE GUIDELINE**

Adopted by the American Academy of Orthopaedic Surgeons
Board of Directors

March 11, 2019

Please cite this guideline as:

American Academy of Orthopaedic Surgeons. Diagnosis and Prevention of Periprosthetic Joint Infections
Clinical Practice Guideline. <https://www.aaos.org/specialty/orthopaedic/periprostheticinfection/>.
Published March 11, 2019.

View the background material via the [JCI CPOG Appendix 1](#)
View data extension via the [JCI CPOG Appendix 2](#)

**The American Academy of Orthopaedic Surgeons
2019 Clinical Practice Guideline
on the Diagnosis and Prevention of Periprosthetic
Joint Infections**

Creighton C. Tubb, MD; Gregory G. Polkowski, MD; Wayne E. Moschetti, MD, MS; Bryan J. Pack, MD; Kathleen G. Beavis, MD; Robin Patel, MD; James D. Slover, MD; Matthew J. Kraay, MD; Christopher J. Palestro, MD; Stefan Riedel, MD; Mihra S. Taljanovic, MD; Karl Roberts, MD; Antonia Chen, MD, MBA; AAOS Staff: Jayson N. Murray, MA; Kyle Mullen, MPH; Patrick Donnelly, MA; Nicole Nelson, MPH; Mary DeMars; Kaitlyn Sevarino, MBA; Anne Woznica, MLIS, AHIP; Peter Shores, MPH

AVOIDING ANTIMICROBIALS TWO WEEKS PRIOR TO OBTAINING INTRA-ARTICULAR CULTURE

- Limited evidence supports withholding antimicrobials for a minimum of two weeks prior to obtaining intra-articular culture to establish the diagnosis of PJI.

Strength of Recommendation: Limited





INITIATING ANTIMICROBIALS PRIOR TO OBTAINING INTRA-ARTICULAR CULTURE

- Moderate evidence supports avoiding administration of antimicrobials in patients suspected of having a periprosthetic joint infection until cultures have been obtained and a diagnosis has been established.

Strength of Recommendation: Moderate





Linee Guida SIOT

Giornale Italiano di Ortopedia e Traumatologia
2019;45:187-213; doi: 10.32050/0390-0134-203

*Coordinamento del progetto
ed elaborazione del documento*

Giuseppe Sessa
Carlo L. Romanò

Gruppo di Lavoro multidisciplinare

Linea Guida SIOT
Diagnosi di infezione peri-protesica
articolare ritardata o tardiva (tempo
trascorso dall'intervento > 90 giorni)



**SOCIETÀ ITALIANA
DI ORTOPEDIA
E TRAUMATOLOGIA**

Definizione di Infezione Periprotesica Articolare

Infectious Disease Society of America
(IDSA, Osmon 2013)

International Consensus Meetings on musculoskeletal Infection
(Parvizi J. 2018)

European Bone and Joint Infection Society
(Renz N. EBJIS 2018)

Worldwide Association against Infection in Orthopaedics and Trauma
(Romanò CL WAIOT 2019)



SOCIETÀ ITALIANA
DI ORTOPEDIA
E TRAUMATOLOGIA

Definizione di Infezione Periprotetica Articolare

Condiviso l'assunto di base ...

la definizione di Infezione Periprotetica Articolare

richiede una **valutazione combinata di più dati clinici e di diversi test diagnostici,**

ogni definizione è diversa:

- numero e tipologia dei diversi criteri diagnostici da considerare
- valori di riferimento differenti per lo stesso esame



SOCIETÀ ITALIANA
DI ORTOPEDIA
E TRAUMATOLOGIA

Definizione di Infezione Periprotetica Articolare

Limiti intrinseci propri di tutte le definizioni

tutti gli Autori segnalano

“sebbene l’utilizzo nella pratica clinica delle attuali definizioni sia utile o addirittura necessario, esistono casi nei quali le stesse definizioni non sono attendibili e, quindi, in un dato paziente, è sempre necessario utilizzare al meglio il proprio discernimento clinico per porre la diagnosi finale.”

Classificazione “temporale”

(non disponibile classificazione accettata a livello internazionale)

EARLY	- precoci:	primi 3 mesi (per alcuni prime 6 sett)
DELAYED	- ritardate:	tra 3 mesi e 2 anni
LATE	- tardive:	oltre 2 anni dall'intervento

Gli antibiotici dove arrivano?

Penetration of Antibacterials into Bone

Pharmacokinetic, Pharmacodynamic and Bioanalytical Considerations

Cornelia B. Landersdorfer,¹ Jürgen B. Bulitta,¹ Martina Kinzig,¹ Ulrike Holzgrabe² and Fritz Sörgel^{1,3}

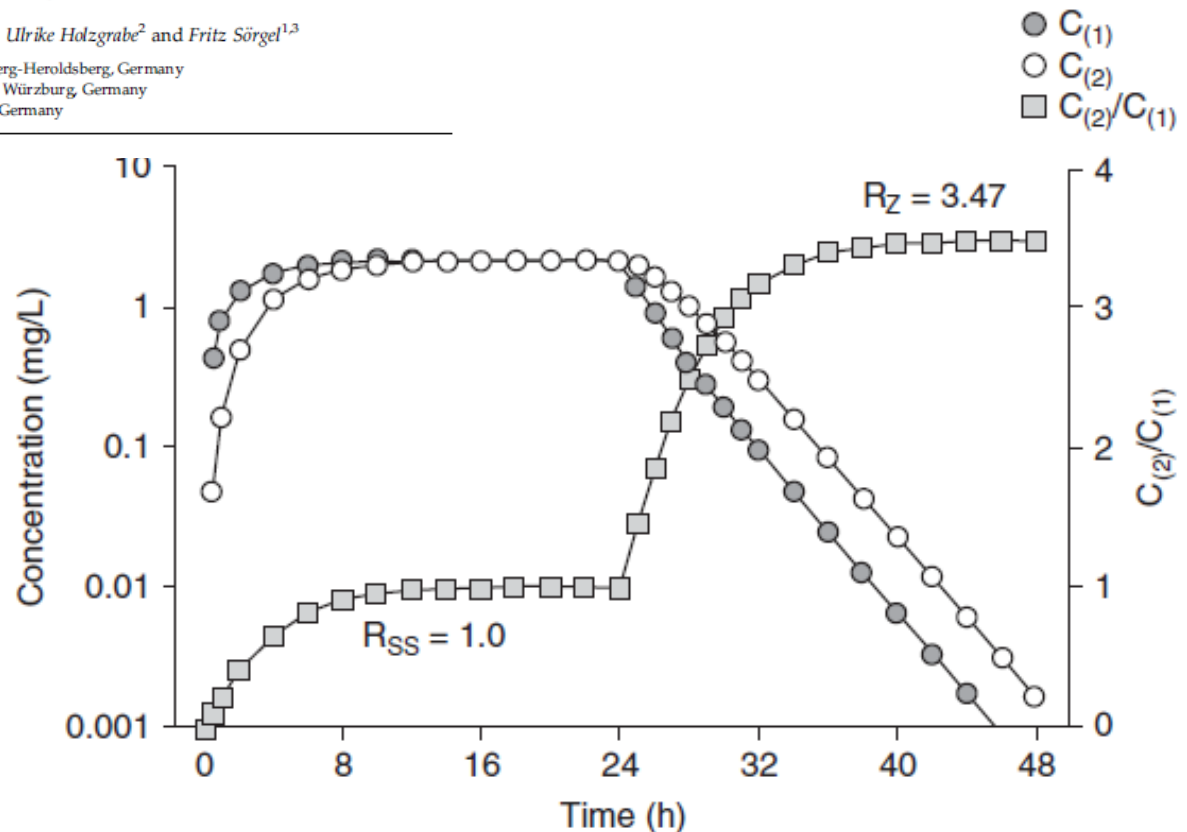
- 1 IBMP-Institute for Biomedical and Pharmaceutical Research, Nürnberg-Heroldsberg, Germany
- 2 Institute of Pharmacy and Food Chemistry, University of Würzburg, Würzburg, Germany
- 3 Department of Pharmacology, University of Duisburg-Essen, Essen, Germany

Concentrazione
 in plasma e osso dopo
 infusione costante 24/h

C1: concentrazione
 plasmatica

C2: concentrazione
 ossea

R2 concentrazione
 osso/plasma
 in fase finale



Gli antibiotici dove arrivano?

Penetration of Antibacterials into Bone

Pharmacokinetic, Pharmacodynamic and Bioanalytical Considerations

Cornelia B. Landersdorfer,¹ Jürgen B. Bulitta,¹ Martina Kinzig,¹ Ulrike Holzgrabe² and Fritz Sörgel^{1,3}

¹ IBMP-Institute for Biomedical and Pharmaceutical Research, Nürnberg-Heroldsberg, Germany

² Institute of Pharmacy and Food Chemistry, University of Würzburg, Würzburg, Germany

³ Department of Pharmacology, University of Duisburg-Essen, Essen, Germany

Rapporti di concentrazione media osso/plasma:
(concentrazione ossea in mg/kg di osso totale)

Azitromicina: tra 0,3 e 1,2

Cefalosporine e Glicopeptidi: 0,15 e 0,3

Penicilline : tra 0,1 e 0,3

Gli antibiotici dove arrivano?

Penetration of Antibacterials into Bone

Pharmacokinetic, Pharmacodynamic and Bioanalytical Considerations

Cornelia B. Landersdorfer,¹ Jürgen B. Bulitta,¹ Martina Kinzig,¹ Ulrike Holzgrabe² and Fritz Sörgel^{1,3}

¹ IBMP-Institute for Biomedical and Pharmaceutical Research, Nürnberg-Heroldsberg, Germany

² Institute of Pharmacy and Food Chemistry, University of Würzburg, Würzburg, Germany

³ Department of Pharmacology, University of Duisburg-Essen, Essen, Germany

Cefalosporine e penicilline: rapporti di concentrazione significativamente più bassi ($p < 0,05$) rispetto a linezolid.

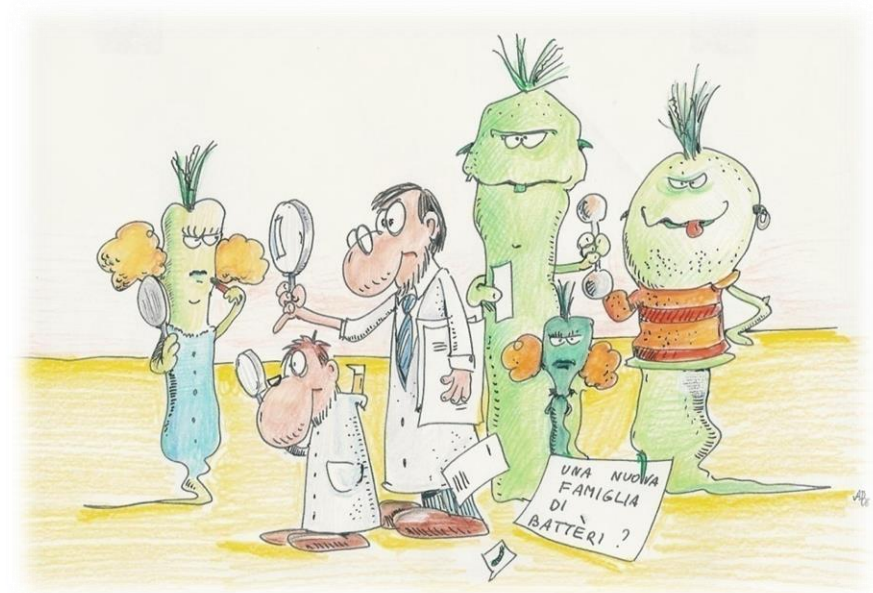
Per 20 di 25 diversi farmaci, i rapporti erano più alti per l'osso spugnoso che per l'osso corticale.

Maggiore estensione della penetrazione ossea per chinoloni, macrolidi e linezolid rispetto a β -lattamici.

... e non solo

Antibioticoresistenza

Biofilm



Resistenza agli antibiotici



... MDR – XDR – PDR
(multi, estesa, pan)

Storicamente: prerogativa degli isolati nosocomiali
Attualmente: diffuse anche in comunità

→ estrema difficoltà in “terapia empirica”

... e non solo!

Aspetti microbiologici: cosa è cambiato?



Produzione di biofilm

Vasta gamma di patogeni

(più o meno virulenti, ma spesso poliantibiotico resistenti)

Bassa predittività

della diagnostica microbiologica tradizionale

Microorganismi e biofilm

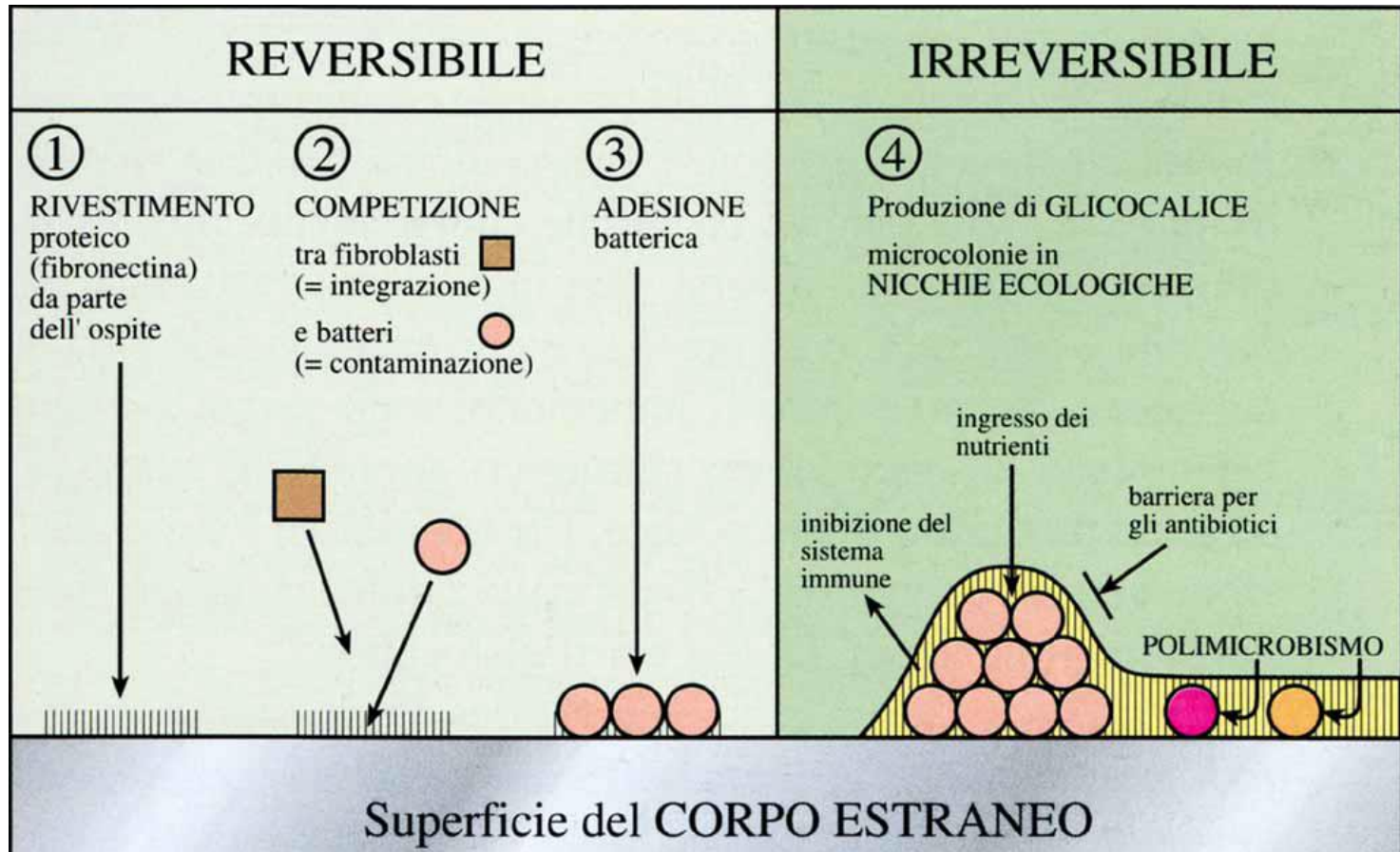
adattamento della comunità
batterica a condizioni ostili



- progressiva scomparsa delle sostanze nutritive
- batteri in fase a lenta crescita o di dormienza
- aumentata resistenza alla killing
di molti antibiotici attivi
sulle cellule in divisione

INFEZIONI PERSISTENTI E CRONICHE

Formazione del biofilm



Inoltre ...

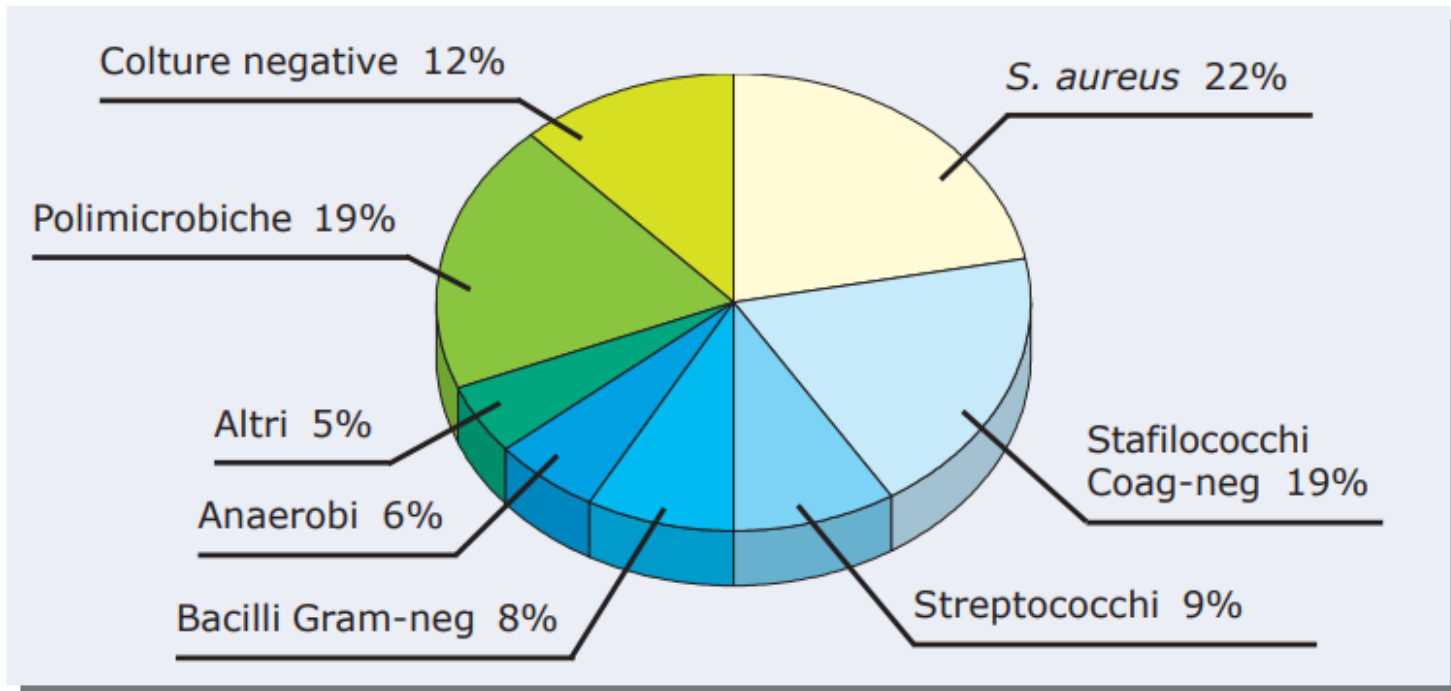
Comorbidità

Scarsa compliance

Aumentata “residenza” della protesi
→ maggior rischio di infettarsi

Cercate il colpevole!

Figura 1. Distribuzione percentuale dei patogeni isolati in 462 casi (1,8%) di infezioni protesiche occorrenti fra il 1969 ed il 1991 su un totale di 26505 interventi di artroplastica totale dell'anca o del ginocchio. (Dati adattati da Berbari *et al* 1998¹¹).



Prosthetic Joint Infection

Colture negative

Aaron J. Tande,^a Robin Patel^{a,b}

PJIs diagnosticate con metodi non microbiologici

7-15% dei casi di infezione

- precedente terapia antibiotica
- inappropriati metodi microbiologici

Diagnosi a 3 anni e mezzo dopo impianto

Terapia antibiotica mirata

Microrganismi	Antibiotici
<i>Staphylococcus aureus</i> meticillino-sensibile*	Oxacillina ± rifampicina Amoxicillina/acido clavulanico ± rifampicina Ciprofloxacina, levofloxacina oppure moxifloxacina ± rifampicina Cotrimossazolo oppure minociclina ± rifampicina Clindamicina
<i>Staphylococcus aureus</i> meticillino-resistente	Teicoplanina oppure vancomicina ± rifampicina Linezolid ± rifampicina Daptomicina
<i>Streptococcus spp.</i>	Amoxicillina ± rifampicina Levofloxacina oppure moxifloxacina ± rifampicina Ceftriaxone ± rifampicina
<i>Enterobacteriaceae</i>	Ciprofloxacina oppure levofloxacina ± rifampicina Ceftriaxone
<i>Pseudomonas aeruginosa</i>	Cefepime oppure ceftazidime Ciprofloxacina oppure levofloxacina Piperacillina/tazobactam Meropenem oppure imipenem

*Minociclina oppure fluorochinoloni o cotrimossazolo in caso di sensibilità *in vitro*.

PERIPROSTHETIC JOINT INFECTION

A Review of Antibiotic Treatment

Ryan Miller, DO

Carlos A. Higuera, MD

Janet Wu, PharmD

Alison Klika, MS

Maja Babic, MD

Nicolas S. Piuze, MD

*Investigation performed at the
Cleveland Clinic, Cleveland, Ohio*

Abstract

» A team approach among orthopaedic surgeons, infectious disease specialists, and patients is of paramount importance when treating periprosthetic joint infections (PJIs). Treatment usually includes various surgical approaches along with antibiotic treatment.

» Antibiotic selection requires a multifactorial decision that depends on the organism that is identified, its antibiotic-resistance profile, the extent of the infection, and factors associated with the host.

» Antibiotic duration is dependent on surgical intervention and the type of organism. Typically, patients are treated for 6 weeks after debridement, antibiotics, and implant retention (DAIR) and for 4 to 6 weeks after single-stage and 2-stage revision arthroplasty.

TABLE I Common Organisms by Timing of Patient Presentation After Surgery and Symptoms

Timing from Surgery	Timing of Symptoms	Organisms Expected*
Early/delayed: < 12 months from prosthesis implantation	Not relevant	<i>Staphylococcus aureus</i> , CoNS, enterobacteriaceae, streptococci
Late: ≥ 12 months from prosthesis implantation	Acute: ≤ 4 weeks of symptoms	<i>S. aureus</i> , streptococci, enterobacteriaceae, CoNS
	Chronic: > 4 weeks of symptoms	CoNS, culture-negative infections, <i>Cutibacterium acnes</i> , <i>S. aureus</i> , streptococci

*CoNS = coagulase-negative Staphylococcus.

TABLE II Relative Antibiotic Concentrations and Properties*

Antibiotic	Spectrum of Activity	Average Cancellous Bone Concentration	Average Joint Concentration	Drug-Drug Interactions†	Major Adverse Effects
Flucloxacillin	MSSA	+++	NA	Mild	Nausea, vomiting, diarrhea, jaundice, and hepatitis
Cephalexin	Streptococcus spp., MSSA, enterobacteriaceae	–	+	Mild	Nausea, vomiting, diarrhea, confusion, jaundice, and arthralgia
Clindamycin	Anaerobic gram-positive and gram-negative organisms, including Bacteroides, MRSA, MSSA, <i>Streptococcus pyogenes</i> , and <i>Streptococcus pneumoniae</i>	+++	++	Mild	Severe diarrhea, metallic taste, rash, and jaundice
Doxycycline	Staphylococcus spp., <i>Streptococcus pneumoniae</i> , <i>Streptococcus pyogenes</i> , rickettsial infections, and <i>Borrelia burgdorferi</i>	+++	NA	Mild	Ultraviolet sensitivity, tooth discoloration for those who are <8 years old, and pill esophagitis

Trimethoprim-sulfamethoxazole	<i>Staphylococcus aureus</i> , <i>Staphylococcus epidermidis</i> , <i>Streptococcus pneumonia</i> , and enterobacteriaceae	++	-	Moderate: warfarin	Hypersensitivity reactions, nephrotoxicity, agranulocytosis, hemolysis, and jaundice
Fluoroquinolones	Staphylococcus spp., Streptococcus spp., enterobacteriaceae, and Pseudomonas (ciprofloxacin and levofloxacin)	+++	+++	Moderate: QT- prolonging medications	QT prolongation, confusion, and tendon inflammation and rupture
Rifampin	Staphylococcus spp.	+++	NA	High: linezolid, clindamycin, warfarin, and estrogen derivatives	Renal failure, hepatic test abnormalities, anemia, nausea, gastroesophageal reflux, and adrenocortical insufficiency
Linezolid	Staphylococcus spp., VRE, and Streptococcus spp.	+	+	Moderate: SSRIs	Diarrhea, headache, anemia, serotonin syndrome, pancytopenia, and optic neuropathy

*MSSA = methicillin-sensitive *Staphylococcus aureus*, NA = not available, MRSA = methicillin-resistant *S. aureus*, VRE = vancomycin-resistant Enterococcus, SSRIs = selective serotonin reuptake inhibitors, and MIC₉₀ = minimum inhibitory concentration at which growth is inhibited in 90% of isolates. - indicates <1 concentration-to-MIC₉₀ ratio, + indicates between 1 and 2 concentration-to-MIC₉₀ ratio, ++ indicates >2 to 5 concentration-to-MIC₉₀ ratio, and +++ indicates >5 concentration-to-MIC₉₀ ratio. MIC₉₀ for *S. aureus* from Clinical and Laboratory Standards Institute, 2018. The included fluoroquinolones are levofloxacin and moxifloxacin. †Drug-drug interactions: mild is defined as few drug interactions or mild severity of interactions that do exist, moderate is defined as some drug interactions or somewhat serious interactions, and high is defined as multiple drug interactions with serious outcomes that may occur.

CLINICAL REVIEW

For the full versions of these articles see bmj.com

Diagnosis and management of prosthetic joint infection

Philippa C Matthews,^{1,2,3} Anthony R Berendt,^{1,2} Martin A McNally,¹ Ivor Byren^{1,2}

¹Bone Infection Unit, Nuffield Orthopaedic Centre NHS Trust, Headington, Oxford OX3 7LD

²Department of Infectious Diseases, Oxford Radcliffe Hospitals NHS Trust, John Radcliffe Hospital, Headington, Oxford OX3 9DU

³University of Oxford, Peter Medawar Building for Pathogen Research, Oxford OX1 3SY

Correspondence to: P C Matthews
p.matthews@doctors.org.uk

Cite this as: *BMJ* 2009;338:b1773
[doi:10.1136/bmj.b1773](https://doi.org/10.1136/bmj.b1773)

Le “Revisione sistematiche”

Box 1 Definitions of prosthetic joint infection

No uniform case definitions exist for prosthetic joint infection, but widely accepted definitions include any of the following^{6 7 9-12 w6 w14}:

- Purulence around a prosthesis at arthrotomy or arthroscopy
- Presence of one or more sinus tract communicating with the joint
- Histological features of infection
- Isolation of an indistinguishable organism from at least two deep culture samples (“indistinguishable” refers to widely performed laboratory characterisation of an organism; in most cases this will be identification of the genus and species, plus antibiotic susceptibilities¹³). Isolation of a virulent organism, such as *Staphylococcus aureus*, *Escherichia coli*, or *Candida* spp, in one deep tissue sample is regarded by some as sufficient to confirm the diagnosis.⁷

Early prosthetic joint infection

Early prosthetic joint infection is most widely defined as deep infection of an arthroplasty occurring within three months of joint replacement. Subsequent presentation is divided into delayed (3-12 months after index surgery) or late (>12 months).^{8 9 11 14}

Le “Revisione sistematiche”

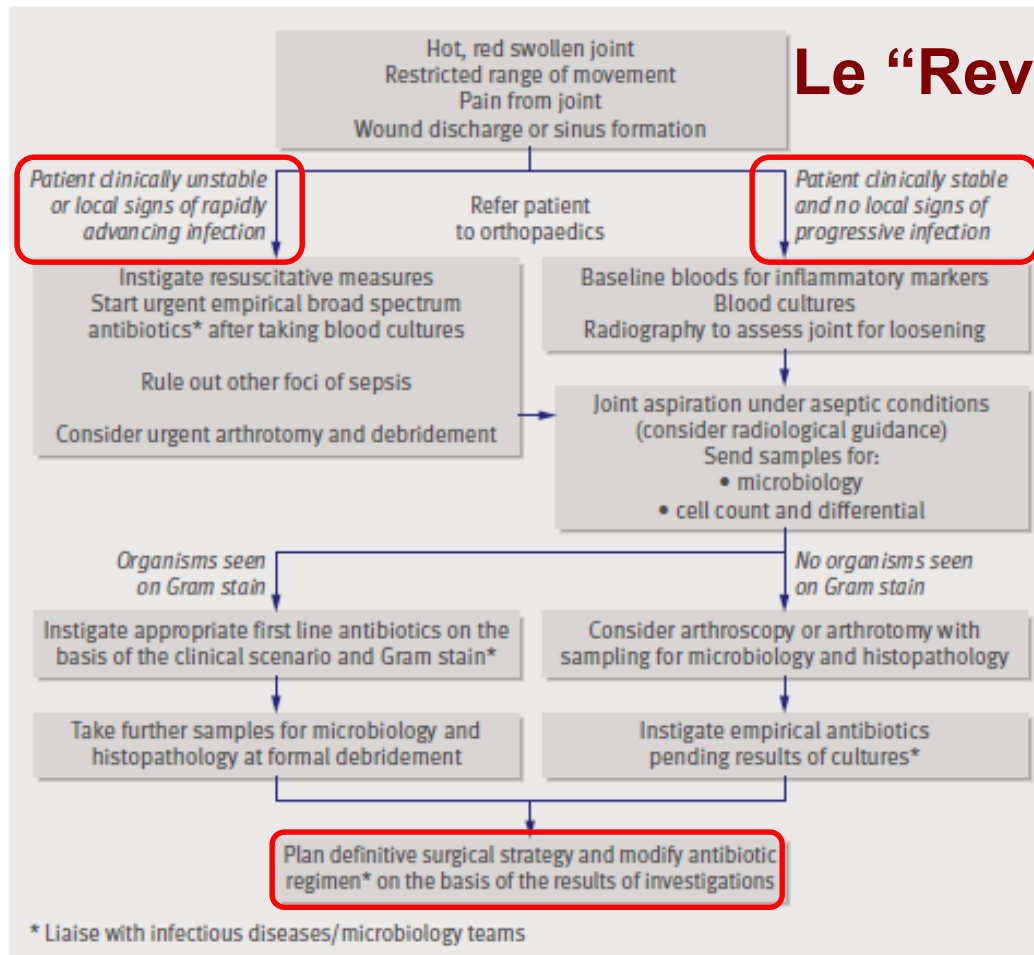


Fig 1 | Algorithm outlining diagnostic and therapeutic interventions in the management of early prosthetic joint infection

Le “Revisione sistematiche”

Box 2 Common risk factors for prosthetic joint infection

Patient factors

Systemic

Advanced age⁹

Obesity^{5 9 15 w8}

Diabetes mellitus^{9 15}

Steroids⁹

Malignancy⁶

Rheumatoid arthritis⁵

Local

Previous arthroplasty on the same joint^{5 9 w38}

Arthroplasty undertaken to treat a fracture⁵

Type of joint replaced (for example, risk is greater for the knee than the hip)²

Perioperative wound complications,⁹ including superficial wound infection,^{6 w12} haematoma, or persistent wound drainage^{1 6}

Operative factors

A score ≥ 36 on the American Society of Anesthesiologists score^{w39}

Duration of operation >75 th centile for the procedure or longer than three hours^{6 w39}

Wound classed as contaminated or dirty^{6 w39}

No systemic antibiotic prophylaxis

No antibiotic cement⁵

Patients with the first three operative factors are at greatest risk of developing infection at the surgical site

Le “Revisione sistematiche”

Table 1 | Organisms that commonly cause prosthetic joint infection

Infesting organism	Frequency of pathogen	Suggested intravenous antibiotic*	Suggested oral antibiotic*
Gram positive			
Coagulase negative staphylococci	13-37% ^{2 7 10} 15 17 19 w2 w28	Glycopeptide ^{9 10 w27}	Rifampicin plus another agent, such as ciprofloxacin, fusidic acid, trimethoprim, or doxycycline, according to sensitivities ^{8 9 15 18 20 w4}
Meticillin sensitive <i>Staphylococcus aureus</i>	20-62% ^{7 9 10} 17-19 w2 w28	Flucloxacillin or ceftriaxone ^{8 10 12}	Rifampicin combinations as above, according to sensitivities ^{8 9 15 18 20 w4}
Meticillin resistant <i>S aureus</i>	2-49% ^{4 7 9 10}	Glycopeptide ^{9 10 w27}	Rifampicin combinations as above, according to sensitivities Linezolid ^{w36 w37†}
<i>Streptococcus</i> spp	4-27% ^{4 7 10 17} w2	Penicillin or ceftriaxone ^{8 10 w19}	Amoxicillin or cefalexin ^{10 19 w19}
<i>Enterococcus</i> spp	6-13% ^{7 9 10 w3}	Penicillin or glycopeptide with or without an aminoglycoside ^{10 w40}	Amoxicillin ^{8 10} or linezolid ^{w36 w37†}
Diphtheroids (<i>Corynebacteria</i> spp, <i>Propionibacteria</i> spp)	6-20% ^{7 10}	Glycopeptide ¹⁰	Rifampicin combinations as above, or ampicillin ¹⁰
Gram negative			
Enteric Gram negative bacilli	2-15% ^{4 7 10 17}	Ceftriaxone ¹⁰ ; carbapenem with or without aminoglycoside ¹⁰	Ciprofloxacin ^{8 10}
<i>Pseudomonas</i> spp	1-4% ^{7 12}	Ceftazidime, ^{8 10} carbapenem, or ticarcillin with or without aminoglycoside	Ciprofloxacin ^{8 10}

Le “Revisione sistematiche”

Table 1 | Organisms that commonly cause prosthetic joint infection

Infesting organism	Frequency of pathogen	Suggested intravenous antibiotic*	Suggested oral antibiotic*
Others			
Anaerobes	1-8% ^{10 17}	Carbapenem ¹⁰	Clindamycin ⁸ or metronidazole ¹⁰
Mycobacteria	<1%-6% ^{14 w7}	A combination of agents (usually including rifampicin) on the basis of the antimicrobial susceptibility profile of the organism; seek microbiology advice	
Fungi	<1% ¹⁰	Common approaches include amphotericin or fluconazole, but seek microbiology advice	Common approaches include fluconazole or itraconazole, but seek microbiology advice
Polymicrobial infections			
≥2 organisms	4-56% ^{9 10 17 w2 w25}	Consider co-amoxiclav ⁸ ; select treatment on the basis of the combination of pathogens and sensitivities ^{w25}	
Culture negative infections			
No pathogen identified	11-26% ^{8 9 18 w2}	Glycopeptide with or without carbapenem or cephalosporin ²¹	Consider standard combination, such as rifampicin plus ciprofloxacin

*All suggested regimens should be tailored according to sensitivities obtained from the microbiology laboratory and should be prescribed after consideration of comorbidities (such as renal or hepatic dysfunction), potential hypersensitivity or side effects (such as rash, nausea, nephrotoxicity), and drug interactions (such as rifampicin and warfarin). Oral therapy generally requires combination regimens; rifampicin should never be used as monotherapy, owing to a high rate of selection for resistance.

†Patients receiving linezolid should be monitored with regular full blood counts and advised about the risk of peripheral or optic neuropathy.

Le “Revisione sistematiche”

Table 2 | Characteristics of biofilms relevant to prosthetic joint infection

Characteristic	Relevance to pathophysiology	Clinical correlates
Genetic diversity: small colony variants ^{w17}	Subpopulations of organisms exist as slow growing phenotypic variants	Organisms may look atypical in vitro, so can confound laboratory diagnosis Slow growth increases the risk of antibiotic failure and justifies the use of prolonged treatment ¹¹
Polysaccharide matrix	Extracellular slime; distributes nutrients and facilitates communication between cells	Microenvironment favours persistence of organisms and may inhibit bacterial killing and phagocytosis; justifies use of prolonged antibiotic regimens incorporating rifampicin ^{12 20}
Adherence to surfaces	Whole population of infecting organisms may be adherent to prosthetic material	Can significantly reduce yield from bacterial culture of synovial fluid; therefore, periprosthetic and tissue samples are also needed to confirm diagnosis ^{7 w14}
Quorum sensing ^{w16}	Genetic interactions between populations of cells	Antibiotic resistance is conferred by sharing of genetic resistance determinants; this characteristic is one reason for using combined oral antibiotic regimens ⁸

Le “Revisione sistematiche”

Table 3 | Characteristics of patients appropriate for different management approaches to prosthetic joint infection

Surgical strategy	Patient characteristics
Debridement, antibiotics, and implant retention	<p>No loosening of prosthesis; adequately functioning joint^{10 12 w19}</p> <p>Healthy soft tissue envelope^{8 19 w18}</p> <p>Short duration of symptoms^{8 19 24 w19} (although recent studies also support this approach in some patients with more chronic disease)^{w18}</p> <p>Clearly characterised bacteriology, and highly antibiotic-susceptible organism(s), such as penicillin susceptible <i>Streptococcus</i> spp^{w19}</p>
One stage revision arthroplasty	<p>Unstable implant but intact soft tissue⁸</p> <p>Organism susceptible to antibiotics^{8 w19}</p> <p>More complex surgery contraindicated because of comorbidity</p>

Le “Revisione sistematiche”

Table 3 | Characteristics of patients appropriate for different management approaches to prosthetic joint infection

Surgical strategy	Patient characteristics
Two stage revision arthroplasty	Unstable implant ⁸ Considerable damage to soft tissue ^{8 19} Resistant or difficult to treat organism, ^{8 22} such as methicillin resistant <i>Staphylococcus aureus</i> , vancomycin resistant enterococci, and fungi Long established infection Failure of previous attempt at debridement and retention ³
Removal of prosthesis or arthrodesis	Serious comorbidity Repeat surgery unacceptable to the patient or deemed unsafe ⁸
Amputation	Last resort in the context of uncontrolled symptoms from the joint, including intractable pain or profuse discharge from sinuses that does not respond to antibiotic suppression Uncontrolled systemic sepsis Mechanical joint failure not amenable to salvage, ^{w41} or severe soft tissue damage not amenable to reconstruction Other options declined by the patient

Consigli da fornire al Medico di Medicina Generale

- secrezione dalla ferita
- eritema sopra l'articolazione
- gonfiore, dolore o limitazione funzionale

prontamente indagati per l'infezione dell'articolazione protesica

Inviare immediatamente a un'équipe ortopedica

- marcatori di flogosi
- emocolture
- artrocentesi

Le radiografie standard possono essere normali,
ma l'allentamento della protesi pone il sospetto di infezione

Consigli da fornire al Medico di Medicina Generale

Immediata collaborazione
con **Microbiologo e Infettivologo**

No antibiotici empirici fino al referto delle colture

Terapia antibiotica, immediatamente dopo emocolture,
se presenti segni di compromissione sistemica

Tentativo di mantenere la protesi:

sbrigliamento chirurgico dei tessuti molli infetti
intorno all'articolazione più 3-6 mesi di antibiotici,
con due antibiotici tra cui Rifampicina

La terapia antibiotica “soppressiva”

The Journal of Arthroplasty 35 (2020) 1154–1160



Contents lists available at [ScienceDirect](#)

The Journal of Arthroplasty

journal homepage: www.arthroplastyjournal.org



Systematic Review & Meta-Analysis

The Role of Long-Term Antibiotic Suppression in the Management of Peri-Prosthetic Joint Infections Treated With Debridement, Antibiotics, and Implant Retention: A Systematic Review



Michael-Alexander Malahias, MD, PhD ^a, Alex Gu, BS ^a, Evan C. Harris, BS ^b, Marco Adriani, MD ^a, Andy O. Miller, MD ^c, Geoffrey H. Westrich, MD ^a, Peter K. Sculco, MD ^{a,*}

^a The Stavros Niarchos Foundation Complex Joint Reconstruction Center, Hospital for Special Surgery, New York, NY

^b George Washington University School of Medicine and Health Sciences, Washington, DC

^c Infectious Diseases & Internal Medicine, Hospital for Special Surgery, New York, NY

La terapia antibiotica “soppressiva”

7 articoli di bassa qualità (livello III o IV)

437 casi di PJI trattati

- chirurgicamente con DAIR (Debridement, Antibiotics, Implant retention)
- poi con SAT (Suppressive Antibiotic Treatment)
 - assenza di infezione nel 75% (318/424) dei pazienti
 - reintervento per qualsiasi causa nel 6,7%
- Effetti avversi: 15,4%
- Effetti avversi che hanno portato all'interruzione di SAT: 4,3%.

Non differenze significative dei tassi di fallimento di DAIR con SAT.

tra infezioni acute (post-operatoria o ematogena, con insorgenza dei sintomi 4 settimane)
e infezioni croniche (insorgenza dei sintomi >4 settimane)

La terapia antibiotica “soppressiva”

Influenza di:

posizione anatomica (anca vs ginocchio)
microrganismi.

sulla percentuale di successo di DAIR con SAT
Nessun dato conclusivo dalla letteratura

Solo prove di bassa qualità per quanto riguarda l'effetto terapeutico di DAIR combinato con SAT: **impossibili conclusioni definitive.**

Sono necessari **studi prospettici di alta qualità** per comprendere

- efficacia
- sicurezza di SAT

La terapia antibiotica “soppressiva”

Interruzioni del trattamento antibiotico per effetti collaterali:
poche 4,3%

Effetti collaterali dopo DAIR con SAT:
molti 15,4%

Per molti soggetti con Infezione di Protesi Articolare, evidente:

- Fragilità di fondo e complessità
- Terapie disponibili non ottimali

S. aureus sembra un importante fattore per il rischio di fallimento

Non dati sufficienti per stabilire quali pazienti trarrebbero beneficio da DAIR con SAT post-operatorio

Necessari
studi epidemiologici policentrici

Epidemiologia locale
puntualmente aggiornata

Stretta collaborazione
clinici - microbiologi





BUON LAVORO!