

Antibiothérapie suppressive (dans les infections ostéo- articulaires)

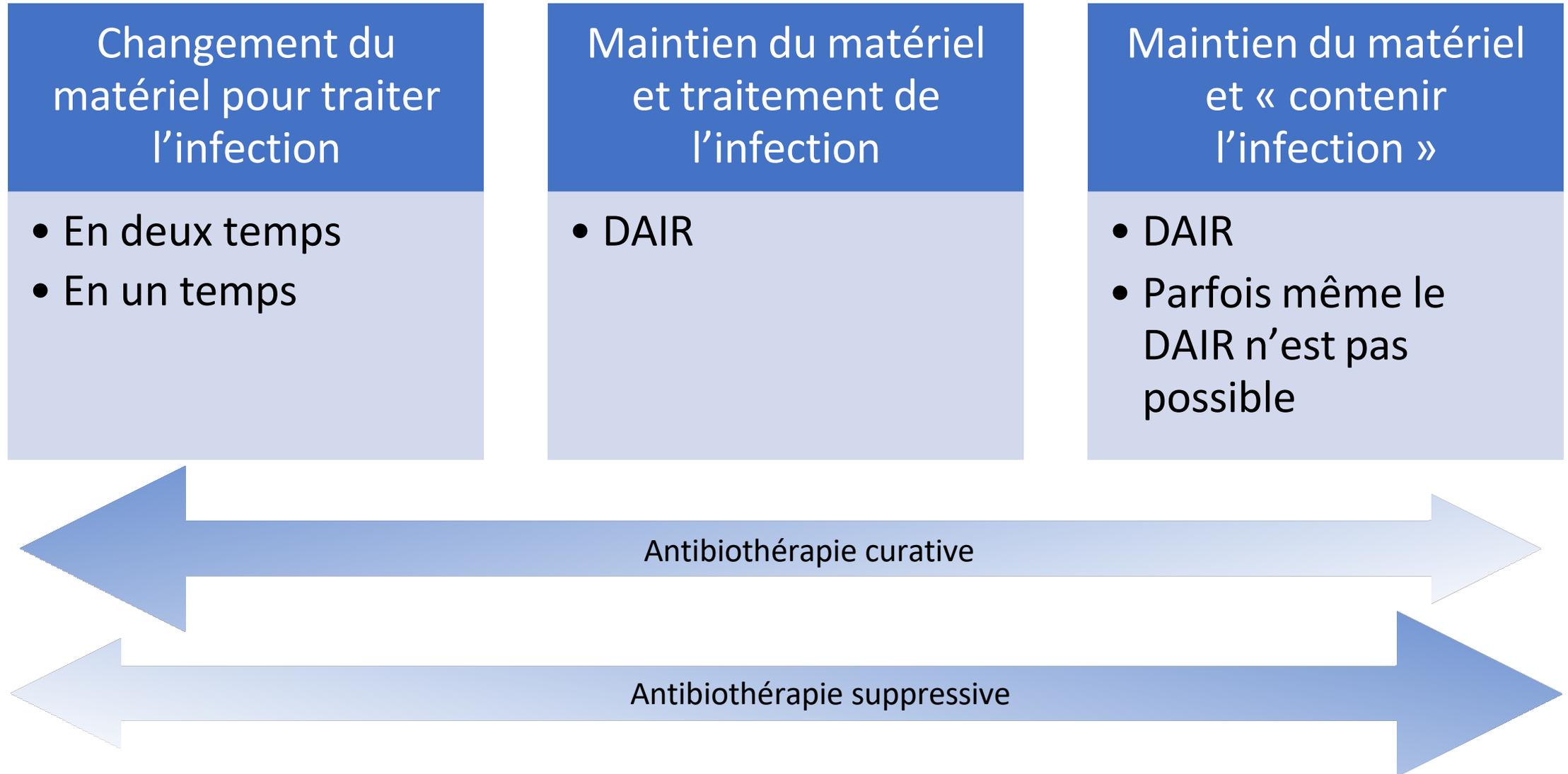
Pr Olivier Robineau

Service Universitaire des Maladies Infectieuses et du Voyageur du Pr
Senneville, Tourcoing

Quelques définitions

- Palliatif: Qui atténue les symptômes d'une maladie sans agir sur la cause et ou sans la guérir
- Suppressif: fait de faire cesser d'être, de mettre fin à l'existence
- Antibiothérapie suppressive: alternative à un traitement curateur médicochirurgical.
- Raison?
- Modalité?
- Durée?

prise en charge des IOAP



prise en charge des IOA

Changement du matériel pour traiter l'infection

Maintien du matériel et traitement de l'infection

Maintien du matériel et « contenir l'infection »

- Stratégie thérapeutique dépend:
- de la cinétique de la gravité des éléments cliniques et paracliniques
 - de l'histoire orthopédique
 - Des comorbidités

Antibiothérapie curative

Antibiothérapie suppressive

Duration of symptoms <3 weeks
OR
Joint age <30 days

*Antimicrobial agents that are recommended for prolonged use for chronic suppression or treatment of biofilm bacteria (see text for details)
**See Figure 3 and recommendation 18 and accompanying Evidence Summary for possible exceptions

YES

NO

- Well fixed prosthesis
- Absence of sinus tract
- Susceptible to oral antimicrobial agents*

YES

NO

Debridement and retention

Removal of prosthesis**

oratory monitoring for efficacy and toxicity is advisable. The decision to offer chronic suppressive therapy must take into account the individual circumstances of the patient including the ability to use rifampin in the initial phase of treatment, the potential for progressive implant loosening and loss of bone stock, and the hazards of prolonged antibiotic therapy; it is therefore generally reserved for patients who are unsuitable for, or refuse, further exchange revision, excision arthroplasty, or amputation.

Duration of symptoms <3 weeks
OR
Joint age <30 days

*Antimicrobial agents that are recommended for prolonged use for chronic suppression or treatment of biofilm bacteria (see text for details)

**See Figure 3 and recommendation 18 and accompanying Evidence Summary for possible exceptions

YES

NO

- Well fixed prosthesis
- Absence of sinus tract
- Susceptible to oral antimicrobial agents*

YES

NO

Debridement and retention

Removal of prosthesis**

oratory monitoring for efficacy and toxicity is advisable. The decision to offer chronic suppressive therapy must take into account the individual circumstances of the patient including the ability to use rifampin in the initial phase of treatment, the potential for progressive implant loosening and loss of bone stock, and the hazards of prolonged antibiotic therapy; it is therefore generally reserved for patients who are unsuitable for, or refuse, further exchange revision, excision arthroplasty, or amputation.

Duration of symptoms <3 weeks
OR
Joint age <30 days

*Antimicrobial agents that are recommended for prolonged use for chronic suppression or treatment of biofilm bacteria (see text for details)

**See Figure 3 and recommendation 18 and accompanying Evidence Summary for possible exceptions

YES

NO

Indication du traitement suppressif possible :
Pour les prises en charges non optimale
Lorsque les molécules actives sur le biofilm ne sont pas utilisables

YES

Debridement
and retention

NO

Removal of
prosthesis**

or amputation.

Antibiothérapie suppressive avant les années 2000

- Travaux rétrospectifs
- Indications du suppressif non évidente
- Haut taux d'échec:
 - sauf Segreti et col
- PEC chirurgical hétérogène (sauf Segreti (DAIR))

Johnson et col, JBJS, 1986

Goulet et col, JA, 1988

Tsoukayama Orthopedics, 1991

Brandt et col, CID, 1997

Segretti et col, CID, 1998

Patient no.	Suppressive antibiotic(s) (dosage)	Duration (mo) of therapy	Antibiotic therapy continued?	Outcome	Complications
1	Cephalexin (500 mg q.i.d.)	103	Yes	Success	None
2	Penicillin (500 mg po q.i.d.)	69	Yes	Success	None
3	Cefadroxil (500 mg po b.i.d.)	96	Yes	Success	None
4	Oxacillin (500 mg po q.i.d.)	51	Yes	Success	None
5	Minocycline/rifampin (100 mg b.i.d./600 mg q.d. po)	56	Yes	Success	None
6	Minocycline/rifampin (100 mg/600 mg po q.d.)	69	Yes	Success	Diarrhea
7	Ampicillin (500 mg po q.i.d.)	12	No	Success	None
8	Minocycline/rifampin (100 mg/600 mg po q.i.d.)	59	Yes	Success	None
9	Dicloxacillin (500 mg po q.i.d.)	22	No	Failure*	None
10	Amoxicillin/clavulanate (500 mg po t.i.d.)	103	Yes	Success	None
11	Minocycline/rifampin (100 mg/600 mg po q.d.)	16	No	Success	Diarrhea
12	Dicloxacillin (500 mg po q.i.d.)	28	No	Success	None
13	Penicillin (500 mg po q.i.d.)	49	Yes	Success	None
14	Minocycline/rifampin (100 mg/600 mg po q.d.)	50	Yes	Success	None
15	TMP-SMZ/rifampin (one DS po q.d./600 mg po q.d.)	71	Yes	Success	None
16	Clindamycin (300 mg po t.i.d.)	4	No	Failure	None
17	Dicloxacillin (500 mg po q.i.d.)	9	No	Failure	Diarrhea
18	Dicloxacillin (500 mg po q.i.d.)	13	No	Failure	Diarrhea, drug rash

NOTE. DS = double strength; TMP-SMZ = trimethoprim-sulfamethoxazole.

* After antibiotic therapy was discontinued.

Patient no.	Suppressive antibiotic(s) (dosage)	Duration (mo) of therapy	Antibiotic therapy continued?	Outcome	Complications
1	Cephalexin (500 mg q.i.d.)	103	Yes	Success	None
2	Penicillin (500 mg po q.i.d.)	69	Yes	Success	None
3	Cefadroxil (500 mg po b.i.d.)	96	Yes	Success	None
4					
5					
6					
7					
8					
9					
10					
11					
12					
13					
14					
15					
16					
17	Dicloxacillin (500 mg po q.i.d.)	9	No	Failure	Diarrhea
18	Dicloxacillin (500 mg po q.i.d.)	13	No	Failure	Diarrhea, drug rash

Quelques arrêts de traitement sans échec (suivi médian de 5 ans)
Plus d'échec chez les patients présentant une infection « retardée »
Au total: 83% de succès sous traitement

NOTE. DS = double strength; TMP-SMZ = trimethoprim-sulfamethoxazole.

* After antibiotic therapy was discontinued.

Années 2000: PEC « moderne »
de l'infection « aigue »
(rifampicine+++)

Long-Term Suppression of Infection in Total Joint Arthroplasty

*Nalini Rao, MD**,**; Lawrence S. Crossett, MD**;
Raj K. Sinha, MD, PhD**; and Jack L. Le Frock, MD, PhD*

- Etude prospective!
- 37 patient traités par:
 - DAIR
 - 6 semaines d'ATB curatif
 - ATB suppressif
- 86% de succès

TABLE 1. Demographic and Clinical Characteristics of the Patients

Characteristics	Number of Patients (%)
Age (years)	
Median	77
Range	62–96
Age group	
55–65	1 (3)
66–75	15 (42)
76–85	13 (36)
86–96	7 (19)
Gender	
Male	19 (53)
Female	17 (47)
Prosthetic joint	
Hips	15 (42)
Knees	19 (53)
Elbow	2 (5.5)
Onset	
Early	13 (36)
Late	23 (64)
Symptoms	
Acute	17 (47)
Chronic	19 (53)
Outcome	
Good	31 (86)
Failed	5 (14)
Duration of treatment (months)	
Mean	52.6
Range	6–128
Duration of followup (months)	
Mean	61.5
Range	16–128

Long-Term Suppression of Infection in Total Joint Arthroplasty

*Nalini Rao, MD**,**; Lawrence S. Crossett, MD**;
Raj K. Sinha, MD, PhD**; and Jack L. Le Frock, MD, PhD*

- Etude prospective!
- 37 patient traités par:
 - DAIR
 - 6 semaines d'ATB curatif
 - ATB suppressif
- 86% de succès

TABLE 1. Demographic and Clinical Characteristics of the Patients

Characteristics	Number of Patients (%)
Age (years)	
Median	77
Range	62–96
Age group	
55–65	1 (3)
66–75	15 (42)
76–85	13 (36)
86–96	7 (19)
Gender	
Male	19 (53)
Female	17 (47)
Prosthetic joint	
Hips	15 (42)
Knees	19 (53)
Elbow	2 (5.5)
Onset	
Early	13 (36)
Late	23 (64)
Symptoms	
Acute	17 (47)
Chronic	19 (53)
Outcome	
Good	31 (86)
Failed	5 (14)
Duration of treatment (months)	
Mean	52.6
Range	6–128
Duration of followup (months)	
Mean	61.5
Range	16–128

Long-Term Suppression of Infection in Total Joint Arthroplasty

*Nalini Rao, MD**,**; Lawrence S. Crossett, MD**;
Raj K. Sinha, MD, PhD**; and Jack L. Le Frock, MD, PhD*

- Etude prospective!
- 37 patient traités par:
 - DAIR
 - 6 semaines d'ATB curatif
 - ATB suppressif
- 86% de succès

TABLE 1. Demographic and Clinical Characteristics of the Patients

Characteristics	Number of Patients (%)
Age (years)	
Median	77
Range	62–96
Age group	
55–65	1 (3)
66–75	15 (42)
76–85	13 (36)
86–96	7 (19)
Gender	
Male	19 (53)
Female	17 (47)
Prosthetic joint	
Hips	15 (42)
Knees	19 (53)
Elbow	2 (5.5)
Onset	
Early	13 (36)
Late	23 (64)
Symptoms	
Acute	17 (47)
Chronic	19 (53)
Outcome	
Good	31 (86)
Failed	5 (14)
Duration of treatment (months)	
Mean	52.6
Range	6–128
Duration of followup (months)	
Mean	61.5
Range	16–128

Long-Term Suppression of Infection in Total Joint Arthroplasty

*Nalini Rao, MD**,**; Lawrence S. Crossett, MD**;
Raj K. Sinha, MD, PhD**; and Jack L. Le Frock, MD, PhD*

- Etude prospective!
- 37 patient traités par:
 - DAIR
 - 6 semaines d'ATB curatif
 - ATB suppressif
- 86% de succès

TABLE 1. Demographic and Clinical Characteristics of the Patients

Characteristics	Number of Patients (%)
Age (years)	
Median	77
Range	62–96
Age group	
55–65	1 (3)
66–75	15 (42)
76–85	13 (36)
86–96	7 (19)
Gender	
Male	19 (53)
Female	17 (47)
Prosthetic joint	
Hips	15 (42)
Knees	19 (53)
Elbow	2 (5.5)
Onset	
Early	13 (36)
Late	23 (64)
Symptoms	
Acute	17 (47)
Chronic	19 (53)
Outcome	
Good	31 (86)
Failed	5 (14)
Duration of treatment (months)	
Mean	52.6
Range	6–128
Duration of followup (months)	
Mean	61.5
Range	16–128

Pathogènes et choix thérapeutiques

TABLE 2. Frequency of Pathogens in Infected Joint Arthroplasty

Pathogens	*Number (%)	Suppressive Antibiotics	Dosage	*Number of Patients	**Pathogen	Success Rate (%)	Followup (months) Median/Range
Methicillin-sensitive Staphylococcus aureus	7 (19)	Minocycline/Rifampin	100 mg/600 mg qd	11	MRCNS, MRSA	82%	66/(12–114)
Methicillin-resistant Staphylococcus aureus	6 (17)	Sulfamethoxazole-Trimethoprim/Rifampin	1 DS*** bid/600 mg qd	1	MRCNS	0%	16
Methicillin-sensitive coagulase negative staphylococcus species	5 (14)	Cephalexin/Rifampin	500 mg bid/600 mg qd	1	MSCNS	100%	6
Methicillin-resistant coagulase negative staphylococcus species	13 (36)	Minocycline	100 mg qd	2	MRCNS	100%	40/(33–47)
Group B streptococcus	3 (8)	TMP/Sulfa	1 DS*** bid	2	MRCNS, GBS	100%	24/(16–32)
Enterococcus	1 (3)	Cephalexin	500 mg bid	4	MSSA	75%	57.5/(22–128)
Escherichia coli	1 (3)	Dicloxacillin	500 mg bid	3	MSSA, MSCNS	100%	96/(96–120)
Candida albicans	1 (3)	Oxacillin	500 mg bid	2	MSCNS	100%	104/(88–120)
		Levofloxacin	500 mg qd	5	MRCNS, E. coli	100%	21/(12–43)
		Penicillin	500 mg bid	2	GBS	100%	82/(72–92)
		Clindamycin	300 mg bid	1	MSSA	0%	18
		Amoxicillin/Doxycycline	500 mg tid/100 mg bid	1	MRCNS, Enterococcus	100%	12
		Fluconazole	400 mg qd	1	C. albicans	100%	35
		Linezolid	600 mg qd	1	MRSA	100%	9

*One patient had enterococcus and methicillin-resistant coagulase-negative staphylococcus species

Pathogènes et choix thérapeutiques

TABLE 2. Frequency of Pathogens in Infected Joint Arthroplasty

Pathogens	*Number (%)	Suppressive Antibiotics	Dosage	*Number of Patients	**Pathogen	Success Rate (%)	Followup (months) Median/Range
Methicillin-sensitive Staphylococcus aureus	7 (19)	Minocycline/Rifampin	100 mg/600 mg qd	11	MRCNS, MRSA	82%	66/(12–114)
Methicillin-resistant Staphylococcus aureus	6 (17)	Sulfamethoxazole-Trimethoprim/Rifampin	1 DS*** bid/600 mg qd	1	MRCNS	0%	16
Methicillin-sensitive coagulase negative staphylococcus species	5 (14)	Cephalexin/Rifampin	500 mg bid/600 mg qd	1	MSCNS	100%	6
Methicillin-resistant coagulase negative staphylococcus species	13 (36)	Minocycline	100 mg qd	2	MRCNS	100%	40/(33–47)
Group B streptococcus	3 (8)	TMP/Sulfa	1 DS*** bid	2	MRCNS, GBS	100%	24/(16–32)
Enterococcus	1 (3)	Cephalexin	500 mg bid	4	MSSA	75%	57.5/(22–128)
Escherichia coli	1 (3)	Dicloxacillin	500 mg bid	3	MSSA, MSCNS	100%	96/(96–120)
Candida albicans	1 (3)	Oxacillin	500 mg bid	2	MSCNS	100%	104/(88–120)
		Levofloxacin	500 mg qd	5	MRCNS, E. coli	100%	21/(12–43)
		Penicillin	500 mg bid	2	GBS	100%	82/(72–92)
		Clindamycin	300 mg bid	1	MSSA	0%	18
		Amoxicillin/Doxycycline	500 mg tid/100 mg bid	1	MRCNS, Enterococcus	100%	12
		Fluconazole	400 mg qd	1	C. albicans	100%	35
		Linezolid	600 mg qd	1	MRSA	100%	9

*One patient had enterococcus and methicillin-resistant coagulase-negative staphylococcus species

Pathogènes et choix thérapeutiques

TABLE 2. Frequency of Pathogens in Infected Joint Arthroplasty

Pathogen	Number	Treatment	Dose	*Number	Organism	Success Rate	Followup
Escherichia coli	1 (3)	Penicillin	500 mg bid	2	GBS	100%	82/(72–92)
Candida albicans	1 (3)	Clindamycin	300 mg bid	1	MSSA	0%	18
		Amoxicillin/Doxycycline	500 mg tid/100 mg bid	1	MRCNS, Enterococcus	100%	12
		Fluconazole	400 mg qd	1	C. albicans	100%	35
		Linezolid	600 mg qd	1	MRSA	100%	9

*One patient had enterococcus and methicillin-resistant coagulase-negative staphylococcus species

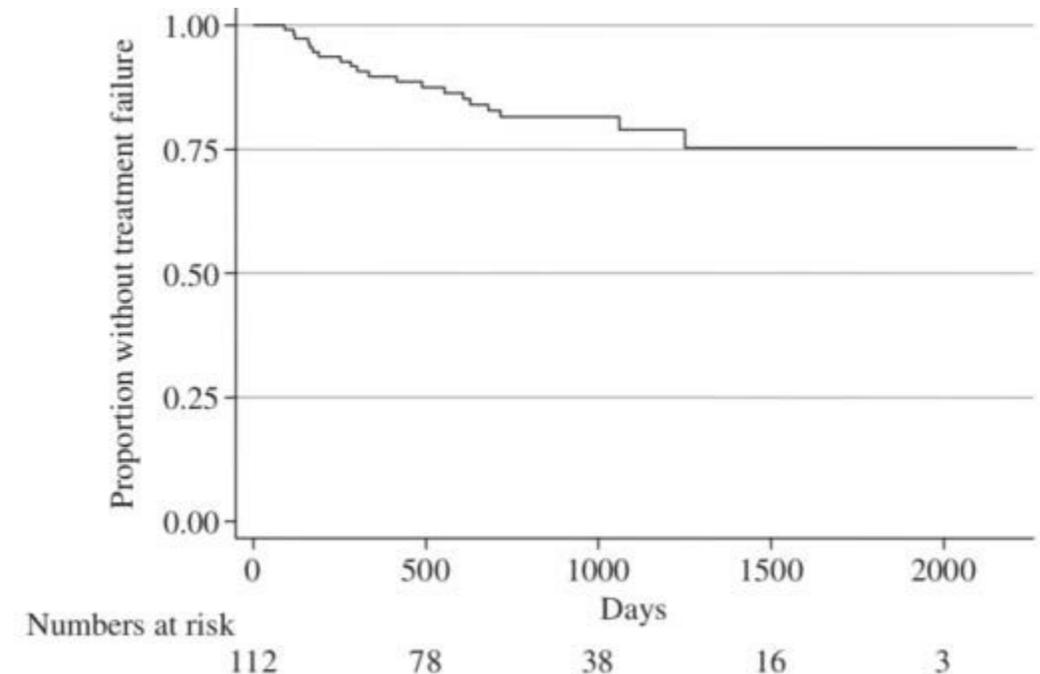
Pas de différence entre les patients à la symptomatologie aiguë ou chronique
 Pb de la définition du traitement suppressif parfois trop proche du curatif

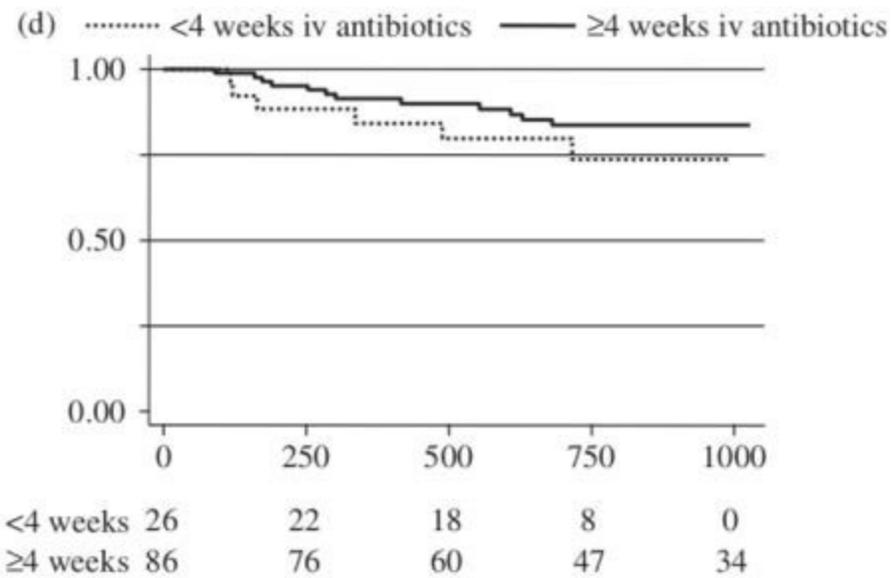
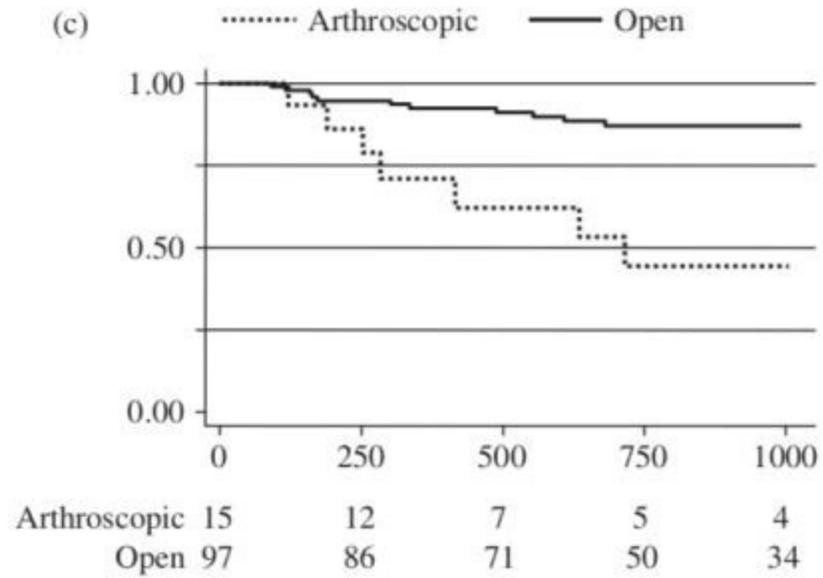
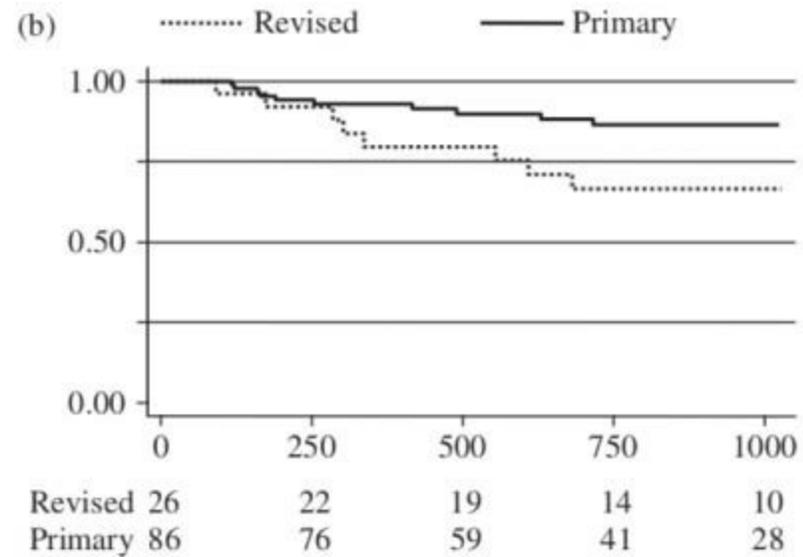
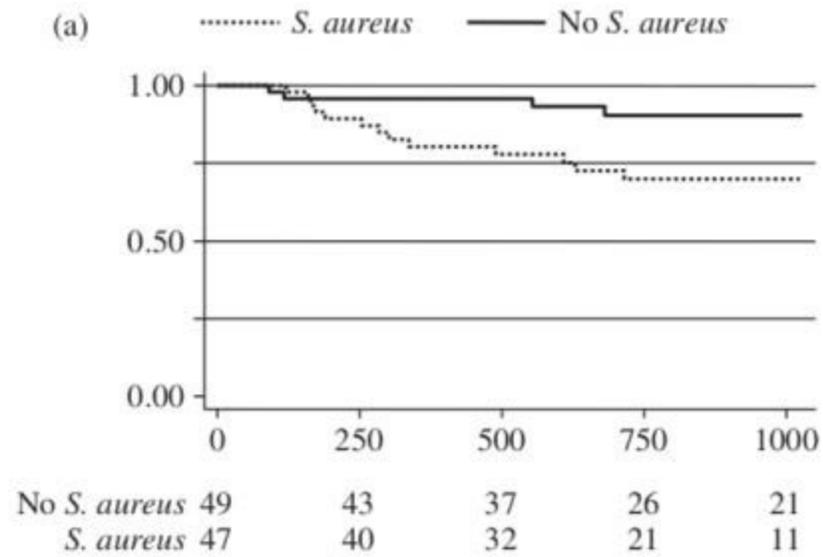
One hundred and twelve infected arthroplasties treated with 'DAIR' (debridement, antibiotics and implant retention): antibiotic duration and outcome

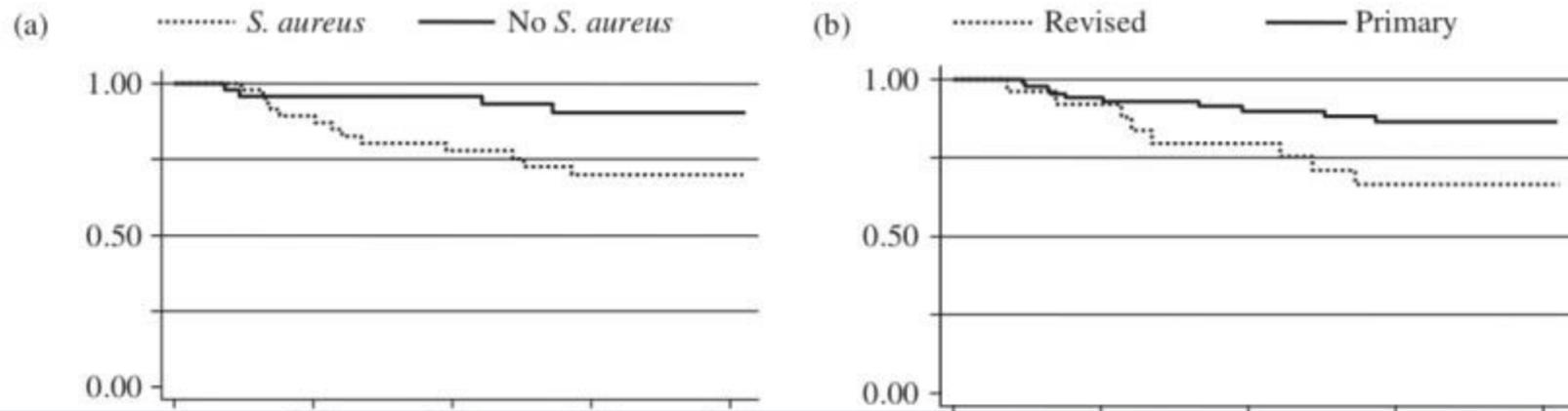
I. Byren^{1,2*†}, P. Bejon^{1,2†}, B. L. Atkins¹⁻³, B. Angus², S. Masters¹, P. McLardy-Smith¹, R. Gundle¹ and A. Berendt¹

- âgés, nombreuses comorbidités
- prothèses posées < 3 mois chez 69% des patients
- DAIR
- Durée AB = 1,5 ans
- 18% d'échecs seulement

	Hazard ratio	95% CI	<i>P</i>
Implant to debridement ≥ 90 days	1.1	0.31–3.8	0.89
Intravenous antibiotics ≥ 28 days	0.49	0.18–1.37	0.18
Arthroscopy versus open	4.2	1.5–12.5	0.008
<i>S. aureus</i>	2.9	1.0–8.4	0.050
Revised versus primary arthroplasty	3.1	1.2–8.3	0.008
Presence of co-morbidity	1.81	0.55–5.9	0.32



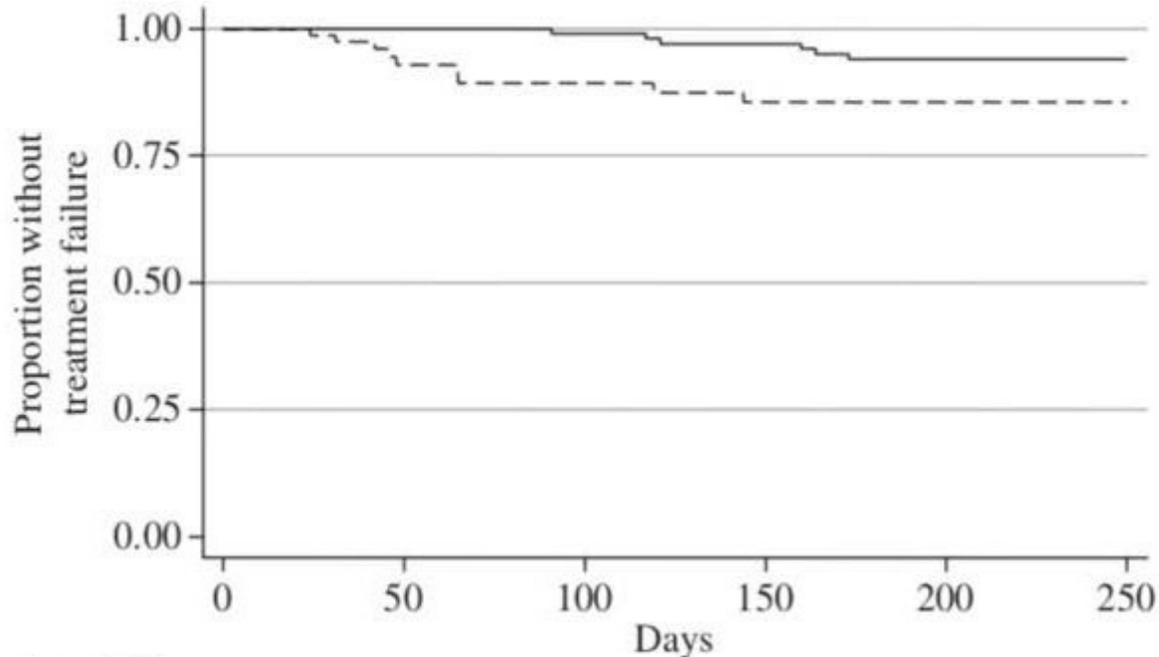




- Confirme le plus haut taux d'échec en cas:
- d'infection à *S. aureus*
 - d'infection sur prothèse changée/révisée
 - De lavage sous arthroscopie

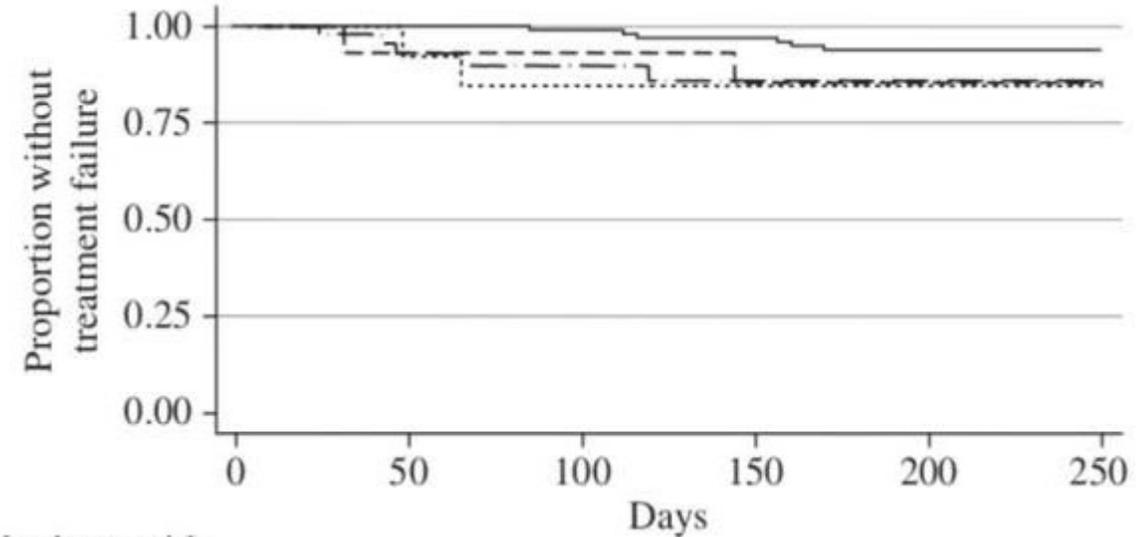
	0	250	500	750	1000		0	250	500	750	1000
Arthroscopic	15	12	7	5	4	<4 weeks	26	22	18	8	0
Open	97	86	71	50	34	≥4 weeks	86	76	60	47	34

Suppressif: reculer pour mieux sauter?



Numbers at risk		0	50	100	150	200	250
On antibiotics	112	107	103	97	89	81	
After stopping	91	55	49	44	41	36	

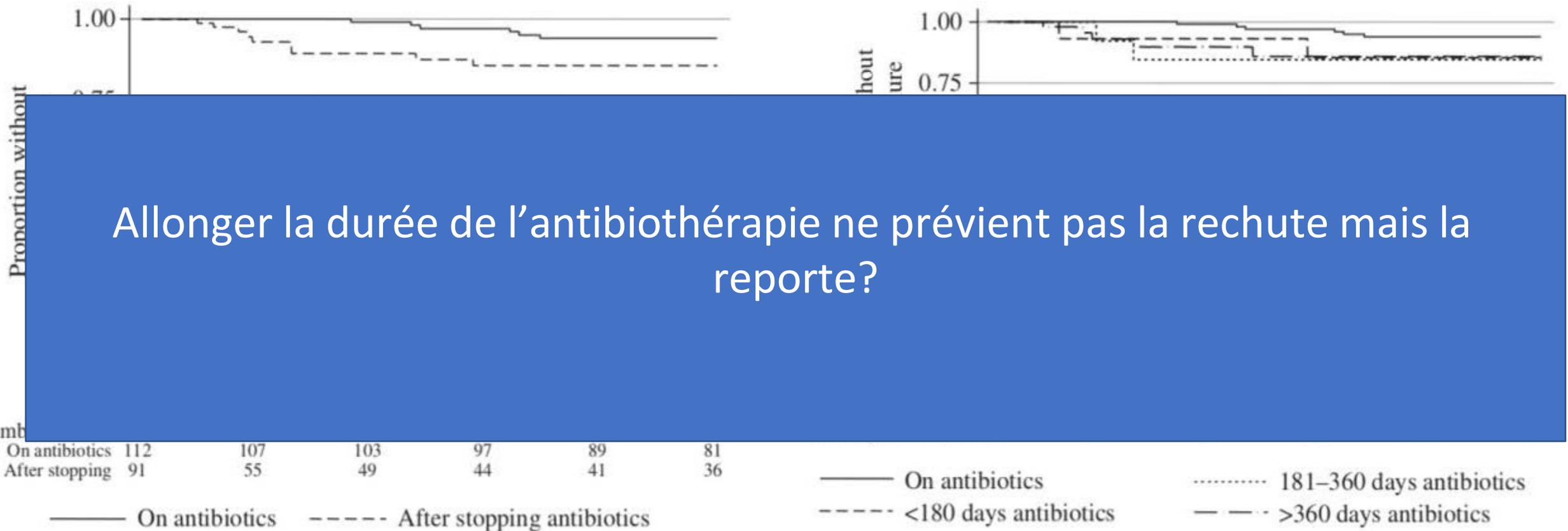
— On antibiotics - - - - - After stopping antibiotics



Numbers at risk		0	50	100	150	200	250
On antibiotics	112	107	103	97	89	81	
<180 days antibiotics	15	12	12	11	10	10	
181-360 days antibiotics	21	12	11	10	10	10	
>360 days antibiotics	55	31	26	23	21	16	

— On antibiotics 181-360 days antibiotics
 - - - - - <180 days antibiotics - · - · - · >360 days antibiotics

Suppressif: reculer pour mieux sauter?



Chronic Suppression of Periprosthetic Joint Infections with Oral Antibiotics Increases Infection-Free Survivorship

Marcelo B.P. Siqueira, MD, Anas Saleh, MD, Alison K. Klika, MS, Colin O'Rourke, MS, Steven Schmitt, MD, Carlos A. Higuera, MD, and Wael K. Barsoum, MD

Investigation performed at the Orthopaedic and Rheumatologic Institute, Cleveland Clinic Foundation, Cleveland, Ohio

- première étude comparative
- Methodologie: Entre la cohorte et le cas-témoins
- Score de propension sur:
 - L'âge
 - sexe,
 - IMC
 - Score de Charlson
 - Nb e PEC chirurgicale
 - Type de chirurgie septique
 - Traitement curatif
 - Micro-organisme (staphylococcus aureus vs les autres)

Résultats

TABLE I Results of Univariate Analyses Comparing Baseline Characteristics Between Suppression and Non-Suppression Groups

Variable	Suppression Group (N = 92)	Non-Suppression Group (N = 276)	P Value
Charlson comorbidity index*	4 [3, 5]	4 [2, 5]	0.34
Age† (yr)	63.7 ± 11.7	64.2 ± 11.5	0.72
BMI† (kg/m ²)	33.6 ± 9.2	33.2 ± 8.6	0.71
Sex†			0.90
Female	36 (39.1)	112 (40.6)	
Male	56 (60.9)	164 (59.4)	
Index surgery†			0.63
Irrigation and debridement with polyethylene exchange	54 (58.7)	152 (55.1)	
2-stage revision	38 (41.3)	124 (44.9)	
No. of previous revisions*	1 [0, 3]	1 [0, 2]	0.37
Pathogen†			0.33
<i>S. aureus</i>	44 (47.8)	114 (41.3)	
Non- <i>S. aureus</i>	48 (52.2)	162 (58.7)	
Joint†			0.94
Knee	71 (77.2)	210 (76.1)	
Hip	21 (22.8)	66 (23.9)	
Duration of symptoms* (days)	30 [7, 90]	14 [5, 45]	0.02
Duration of intravenous antibiotic therapy* (wk)	6 [6, 6]	6 [6, 6]	0.17
Previous joint infection anywhere†	41 (44.6)	130 (47.1)	0.76
Infecting organism class†			0.21
Virulent§	54 (58.7)	147 (53.2)	
Indolent#	31 (33.7)	55 (20.0)	
Fungal and acid-fast bacilli	0	1 (0.3)	
Miscellaneous and contaminants	5 (5.4)	22 (7.2)	
Multiple organisms†	18 (19.6)	35 (12.7)	0.13

*The values are given as the median with the 25th and 75th percentiles in brackets. †The values are given as the mean and standard deviation. ‡The values are given as the number of patients with the percentage in parentheses. §Includes *S. aureus*, Enterococcus, and gram-negative organisms. #Includes coagulase-negative Staphylococcus and Propionibacterium species.

Résultats

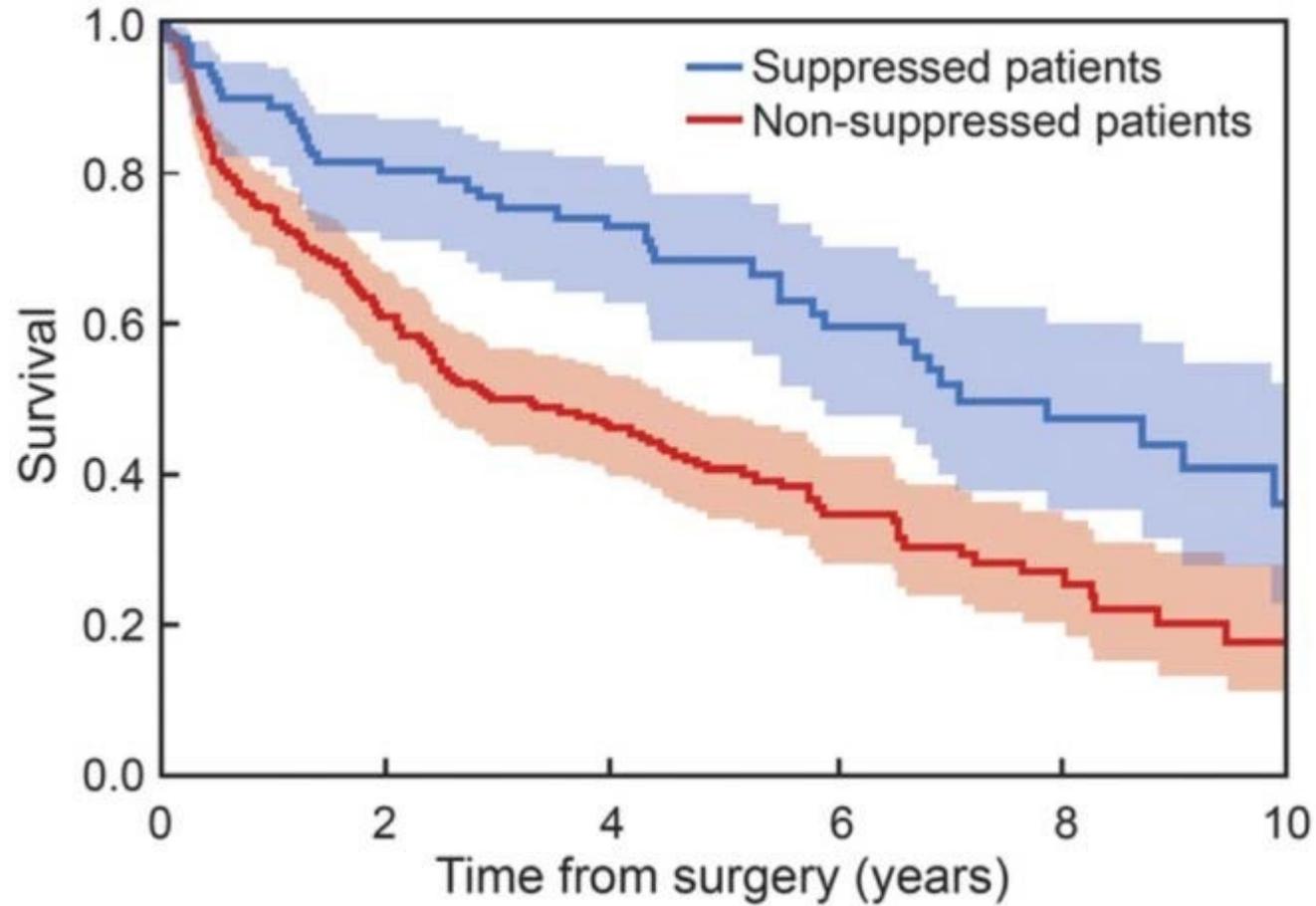
TABLE I Results of Univariate Analyses Comparing Baseline Characteristics Between Suppression and Non-Suppression Groups

Variable	Suppression Group (N = 92)	Non-Suppression Group (N = 276)	P Value
Charlson comorbidity index*	4 [3, 5]	4 [2, 5]	0.34
Age† (yr)	63.7 ± 11.7	64.2 ± 11.5	0.72
BMI† (kg/m ²)	33.6 ± 9.2	33.2 ± 8.6	0.71
Sex‡			0.90
Female	36 (39.1)	112 (40.6)	
Male	56 (60.9)	164 (59.4)	
Index surgery‡			0.63
Irrigation and debridement with polyethylene exchange	54 (58.7)	152 (55.1)	
Viridans§	54 (58.7)	147 (53.2)	0.37
Indolent#	31 (33.7)	55 (20.0)	0.33
Fungal and acid-fast bacilli	0	1 (0.3)	0.94
Miscellaneous and contaminants	5 (5.4)	22 (7.2)	0.02
Multiple organisms‡	18 (19.6)	35 (12.7)	0.17
			0.76
			0.21

Pourquoi les patients similaires ont-ils eu des PEC différentes?

*The values are given as the median with the 25th and 75th percentiles in brackets. †The values are given as the mean and standard deviation. ‡The values are given as the number of patients with the percentage in parentheses. §Includes *S. aureus*, Enterococcus, and gram-negative organisms. #Includes coagulase-negative Staphylococcus and Propionibacterium species.

Résultats



Echec du traitement et raison de pose de prothèse

Pt	Sex	Age	Indication AST	Indication prosthesis	Revised prosthesis	Affected joint	Type of prosthesis	Months after implantation	Surgeries before start AST (n)	Micro-organism(s)	Outcome
1	M	70	comorbidity	osteoarthritis	yes	hip	standard	54 (late)	lavage (3)	<i>S. pyogenes</i>	Successful
2	M	69	comorbidity/prognosis	malignancy	yes	hip	tumor	1 (early)	no	<i>S. epidermidis</i>	Successful
3	M	40	poor soft tissue	malignancy	yes	hip	tumor	1 (early)	lavage (1)	<i>S. aureus</i>	Successful
4	M	71	prognosis	malignancy	yes	hip	standard	1 (early)	no	<i>S. epidermidis, E. cloacae</i>	Successful
5	F	76	comorbidity	osteoarthritis	yes	knee	standard	30 (late)	no	<i>S. epidermidis</i>	Successful
6	F	55	comorbidity/prognosis	malignancy	yes	hip	standard	1 (early)	lavage (1)	<i>S. aureus, E. faecalis</i>	Successful
7	F	71	patient wish	osteoarthritis	yes	hip	standard	39 (late)	lavage (1)	<i>Bacteroides fragilis</i>	Successful
8	F	47	poor soft tissue/bone stock	osteoarthritis	yes	hip	standard	1 (early)	DAIR (1), lavage (1)	<i>S. epidermidis (rifampin R)</i>	Successful
9	M	80	comorbidity	fracture	no	shoulder	standard	1 (early)	lavage (1)	<i>P. acnes</i>	Successful
10	M	52	pore bonestock/prognosis	malignancy	no	knee	tumor	6 (delayed)	DAIR (1)	<i>S. aureus</i>	Successful
11	M	35	pore bonestock	osteomyelitis	yes	hip	tumor	44 (late)	no	<i>S. epidermidis</i>	Successful
12	M	21	prognosis	malignancy	no	knee	tumor	24 (late)	excision sarcoma	<i>P. acnes</i>	Successful
13	M	73	poor soft tissue/bone stock	fracture	yes	hip	standard	78 (late)	DAIR (1), lavage (2)	<i>S. aureus</i>	Successful
14	M	88	comorbidity/prognosis	osteoarthritis	no	knee	standard	1 (early)	lavage (2)	<i>S. epidermidis</i>	Successful
15	F	54	poor bonestock	RA	yes	knee	tumor	27 (late)	no	<i>S. aureus</i>	Failed ¹
16	M	70	comorbidity/prognosis	RA	yes	knee	standard	176 (late)	lavage (1)	<i>S. aureus</i>	Failed ²
17	F	59	poor soft tissue/bone stock	osteoarthritis	yes	hip	tumor	6 (delayed)	reposition (3)	<i>S. aureus, S. epidermidis</i>	Failed ¹
18	F	67	poor soft tissue	fracture	no	shoulder	standard	7 (delayed)	lavage (2)	<i>S. aureus</i>	Failed ²
19	M	58	patient wish	malignancy	no	hip	tumor	2 (early)	DAIR (2)	<i>E. coli (ESBL+), E. faecalis</i>	Failed ²
20	M	68	poor soft tissue	fracture	no	hip	tumor	43 (late)	DAIR (1), lavage (2)	<i>S. agalactiae</i>	Failed ³
21	F	65	comorbidity	RA	yes	hip	tumor	9 (delayed)	DAIR (1)	<i>S. epidermidis</i>	Failed ¹

Ce qu'il ne faut pas faire!

Table 1. Clinical Characteristics of the Cohort (N = 510)

Variable	No. (%)
Age, y, mean (SD)	70.4 (11)
Sex	
Female	294 (58)
Male	216 (42)
Body mass index, ^a mean (SD)	32.1 (7.6)
Joint	
Knee	254 (50)
Hip	256 (50)
Arthroplasty	
Primary	351 (69)
Cemented	392 (77)
Comorbidities	
Diabetes mellitus	101 (20)
Heart failure	55 (11)
Rheumatoid arthritis	48 (9)
Chronic obstructive pulmonary disease	43 (8)
Chronic kidney disease	31 (6)
Liver cirrhosis	9 (2)
Alcohol use ^b	88 (21)
Active smoking ^c	57 (14)
History of fracture	50 (10)
Infection type	
Monomicrobial	327 (64)
Polymicrobial	160 (31)
Bacteremia	78 (15)
Microbial etiology	
<i>Staphylococcus aureus</i>	193 (38)
Coagulase-negative staphylococci	146 (29)
<i>Streptococcus</i> species	97 (19)
<i>Enterococcus</i> species	59 (12)
Other gram-positive bacteria	92 (18)
Gram-negative bacteria	88 (17)
Anaerobic bacteria	12 (2)
Therapy	
Rifampin use	282 (55)
Quinolone use	221 (43)
Modular component exchange	239 (47)

^aMissing in 25 cases.

^bDenominator was 410.

^cDenominator was 411.

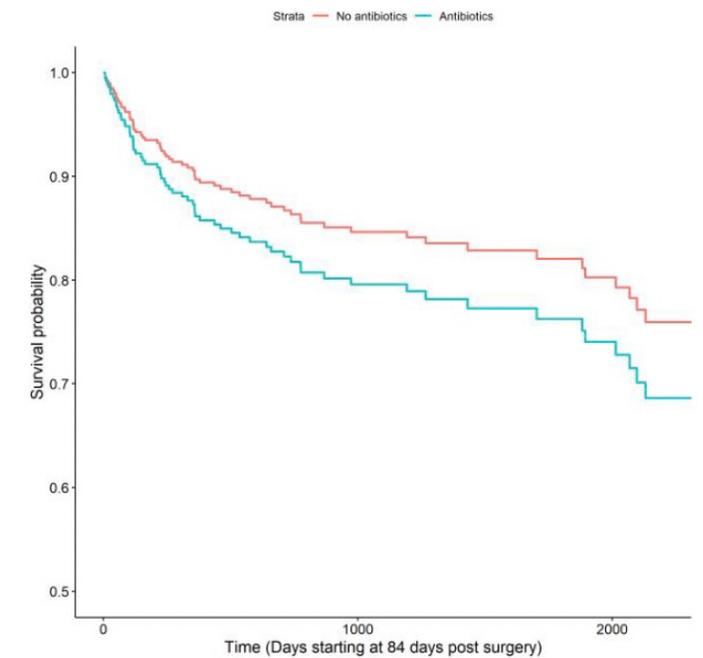


Figure 1. Estimated survival curve for treatment failure on and off Suppressive Antibiotic Therapy based on Cox model.

Table 2. Analysis of Treatment Outcomes With Suppressive Antibiotic Therapy as a Time-Varying Covariate

Group	No. of Patients	No. of Events	HR (95% CI)	P Value
Overall	510	66	1.37 (.79–2.39)	.27
PJI				
Knee	255	49	1.38 (.72–2.65)	.33
Hip	255	17	0.87 (.25–3.02)	.83
Knee after 180 d	236	37	1.02 (.50–2.10)	.95
Acute infection				
Early	367	30	1.41 (.63–3.17)	.40
Late	143	36	0.95 (.42–2.19)	.91
Cohort				
United States	184	36	0.36 (.11–1.15)	.09
Netherlands	236	16	11.46 (2.08–63.03)	.01
Spain	90	14	0.62 (.10–3.93)	.61

Auteurs	Design de l'étude	Effectif suppressif	Antibiothérapie curative	Durée moyenne d'antibiothérapie suppressive	Chirurgie	Antibiothérapie suppressive : molécule utilisée / effets indésirables	Succès	Définition Echec et délai suivi
Pavoni CMI 2004	Rétrospectif 1995-2003	N=34	IV 6 semaines Rifampicine	> 6 mois	Hétérogène (14/34 DAIR)	Hétérogène / pas d'arrêt pour EI	91%	Clinique Suivi 9-57 mois
Marculescu CID 2006	Rétrospectif 1995-1999	N=88	28j IV Absence de Rifampicine	5-2673 jours (médiane à 541 jours)	DAIR	Hétérogène / NR	52%	Clinico-biologique Suivi = 700 jours (1-2779)
Koeppe IDCP 2008	Rétrospectif	N=12	IV/PO moyenne 48jRifampicine	NR	NR	NR	83%	Clinique Suivi = 28 mois (4-48)
El Helou EJCMID 2010	IPA SA et SCN 3 cohortes : -1 prospective P (n=14) -2 rétrospective R1 (n=31) et R2 (n=56)	P : N=13 R1 : N=24 R2 : N=34	P : 3/6 mois dont 4 semaines IV avec rifampicine R1 : 4 semaines IV avec rifampicine R2 : 4 sem IV sans rifampicine	P : 192 jours (140-306) R1 : 183 jours (3-323) R2 : 322 jours (6-365)	DAIR	Hétérogène / Effets indésirables : P=14% R1=29% R2=16%	P=13/14 (93%) R1=21/31 (68%) R2 = 35/56 (63%)	Suivi à un an : clinique, radiologique, bactériologique
Prendki IJID 2014	Rétrospectif 2004 – 2008 Patients > 80 ans	N=38	59 jours (15-90) Rifampicine	> 6 mois	Hétérogène, chirurgie pour seulement 9/38 patients	Hétérogène / 10,5%	60%	Clinico-biologique Suivi médian : 24 mois

Auteurs	Design de l'étude	Effectif suppressif	Antibiothérapie curative	Durée moyenne d'antibiothérapie suppressive	Chirurgie	Antibiothérapie suppressive : molécule utilisée / effets indésirables	Succès	Définition Echec et délai suivi
Pavoni CMI 2004	Rétrospectif 1995-2003	N=34	IV 6 semaines Rifampicine	> 6 mois	Hétérogène (14/34 DAIR)	Hétérogène / pas d'arrêt pour EI	91%	Clinique Suivi 9-57 mois
Marculescu CID 2006	Rétrospectif 1995-1999	N=88	28j IV Absence de Rifampicine	5-2673 jours (médiane à 541 jours)	DAIR	Hétérogène / NR	52%	Clinico-biologique Suivi = 700 jours (1-2779)

Hétérogénéité de:
 -patients
 -traitement^s
 -suivi
Efficacité: 50 à 93%

			sans rifampicine					
Prendki IJID 2014	Rétrospectif 2004 – 2008 Patients > 80 ans	N=38	59 jours (15- 90) Rifampicine	> 6 mois	Hétérogène, chirurgie pour seulement 9/38 patients	Hétérogène / 10,5%	60%	Clinico-biologique Suivi médian : 24 mois

Quels antibiotiques?

Microorganism	Preferred Treatment	Alternative Treatment
Staphylococci, oxacillin-susceptible	Cephalexin 500 mg PO tid or qid or Cefadroxil 500 mg PO bid	Dicloxacillin 500 mg PO tid or qid Clindamycin 300 mg PO qid Amoxicillin-clavulanate 500 mg PO tid
Staphylococci, oxacillin-resistant	Cotrimoxazole 1 DS tab PO bid Minocycline or doxycycline 100 mg PO bid	
β -hemolytic streptococci	Penicillin V 500 mg PO bid to qid or Amoxicillin 500 mg PO tid	Cephalexin 500 mg PO tid or qid
<i>Enterococcus</i> spp, penicillin susceptible	Penicillin V 500 mg PO bid to qid or Amoxicillin 500 mg PO tid	
<i>Pseudomonas aeruginosa</i>	Ciprofloxacin 250–500 mg PO bid	
Enterobacteriaceae	Cotrimoxazole 1 DS tab PO bid	β -lactam oral therapy based on in vitro susceptibilities
<i>Propionibacterium</i> spp	Penicillin V 500 mg PO bid to qid or Amoxicillin 500 mg PO tid	Cephalexin 500 mg PO tid or qid Minocycline or doxycycline 100 mg PO bid

Antibiothérapie suppressive sans chirurgie = palliative?

- 21 Patients âgés (86 ans) aux multiples comorbidités
- Pas de PEC chirurgicale
- Monothérapie pour la plupart des patients
- Observations en fin de suivi (suivi médian: 17.3 mois (1.4- 56.6)):
 - 8 patients ont présenté des évènements
 - 1 effet secondaire
 - 2 décès sans lien avec l'infection
 - 3 sepsis
 - 2 évolutions défavorables locales

Antibiothérapie suppressive sans chirurgie?

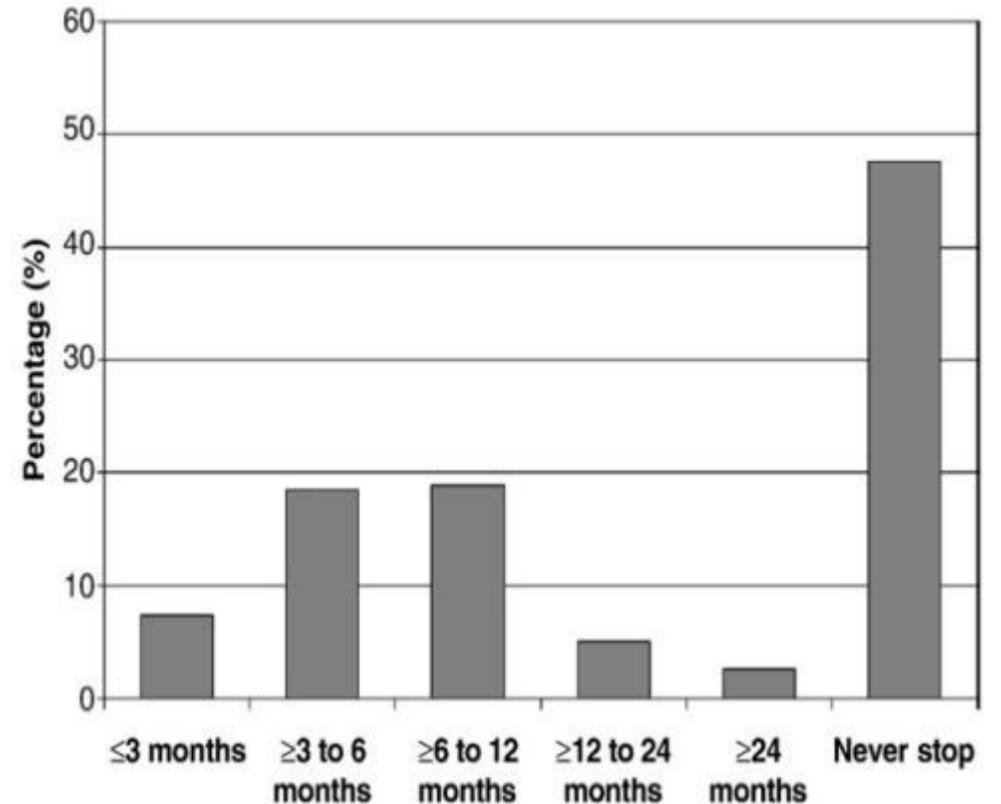
- 21 Patients âgés (86 ans) aux multiples comorbidités

Place de la fistule?

- 3 sepsis
- 2 évolutions défavorables locales

Quelles sont les pratiques?

- Enquête portant sur 545 médecins sur la PEC des IOAP
- A propos de l'indication du DAIR et le traitement suppressif:
 - Risque opératoire: 76%
 - Délai long par rapport à la chirurgie initiale 50%
 - Refus de la chirurgie 1,3%
 - Comorbidités, âge avancé, espérance de vie courte: 2,1%
 - micro-organisme sensible, « faible virulence » 37,9%



How We Approach Suppressive Antibiotic Therapy (SAT) Following Debridement, Antibiotics, and Implant Retention for Prosthetic Joint Infection

Nicolas Cortes-Penfield, et al. CID. 2023 August

Proposition indications SAT

Table 3. Factors We Suggest Guide Prescription of Suppressive Antibiotic Therapy Following DAIR

Factors strongly suggesting benefit from SAT after DAIR (the authors would offer SAT to most patients with at least 1 of these factors)

- Limited options for arthroplasty revision (ie, recurrent infection would require amputation, arthrodesis, or difficult wound coverage and likely result in a substantially worse functional outcome)^a
- Recurrent PJI/prior PJI treatment failure^b
- Infection with difficult-to-treat pathogens (*S. aureus*^b and possibly others^a, eg *Pseudomonas aeruginosa* or *Candida*)
- Severe immunocompromise (ie, solid-organ or stem cell transplant, active chemotherapy, chronic systemic steroid therapy, TNF-inhibitor therapy, advanced HIV)^a
- Underwent arthroscopy instead of open DAIR, or polyethylene liner was not exchanged^b

Factors that may suggest benefit from SAT after DAIR (the authors would consider SAT in patients with at least 1 of these factors)

- Major end-organ disease predisposing to poor outcome (ie, cirrhosis, ESRD, or heart failure)^b
- Age >75 y ^b or estimated life expectancy <10 ya
- Late hematogenous infection (onset >2 y after initial arthroplasty), particularly if associated with active bacteremia^b
- Gram-negative infection that cannot be treated with a fluoroquinolone^b
- Patient strongly values the potential benefits of SAT over the potential risks after both have been explained in an informed shared-decision-making conversation^a

Factors suggesting little benefit from SAT (the authors would not offer SAT to most patients with these factors)

- Completion of >6 wk of adjunctive rifampin for susceptible, monomicrobial coagulase-negative *Staphylococcus* spp. infection (as part of a minimum 3–6 mo total antibiotics)^b
- Completion of a fluoroquinolone-based regimen for a gram-negative infection^b
- Culture-negative infection^b

Abbreviations: DAIR, debridement, antibiotics, and implant retention; ESRD, end-stage renal disease; HIV, human immunodeficiency virus; PJI, prosthetic joint infection; SAT, suppressive antibiotic therapy; TNF, tumor necrosis factor.

^aBased on the consensus opinion of the authors, but not directly supported by the data identified in this review.

^bBased on retrospective data identifying this as a risk factor for treatment failure.

How We Approach Suppressive Antibiotic Therapy (SAT) Following Debridement, Antibiotics, and Implant Retention for Prosthetic Joint Infection

Nicolas Cortes-Penfield, et al. CID. 2023 August

Proposition indications SAT

Table 3. Factors We Suggest Guide Prescription of Suppressive Antibiotic Therapy Following DAIR

Factors strongly suggesting benefit from SAT after DAIR (the authors would offer SAT to most patients with at least 1 of these factors)

- Limited options for arthroplasty revision (ie, recurrent infection would require amputation, arthrodesis, or difficult wound coverage and likely result in a substantially worse functional outcome)^a
- Recurrent PJI/prior PJI treatment failure^b
- Infection with difficult-to-treat pathogens (*S. aureus*^b and possibly others^a, eg *Pseudomonas aeruginosa* or *Candida*)
- Severe immunocompromise (ie, solid-organ or stem cell transplant, active chemotherapy, chronic systemic steroid therapy, TNF-inhibitor therapy, advanced HIV)^a
- Underwent arthroscopy instead of open DAIR^b or polyethylene liner was not exchanged^b

Factors that may suggest benefit from SAT after DAIR (the authors would consider SAT in patients with at least 1 of these factors)

- Major end-organ disease predisposing to poor outcome (ie, cirrhosis, ESRD, or heart failure)^b
- Age >75 y^b or estimated life expectancy <10 ya
- Late hematogenous infection (onset >2 y after initial arthroplasty), particularly if associated with active bacteremia^b
- Gram-negative infection that cannot be treated with a fluoroquinolone^b
- Patient strongly values the potential benefits of SAT over the potential risks after both have been explained in an informed shared-decision-making conversation^a

Factors suggesting little benefit from SAT (the authors would not offer SAT to most patients with these factors)

- Completion of >6 wk of adjunctive rifampin for susceptible, monomicrobial coagulase-negative *Staphylococcus* spp. infection (as part of a minimum 3–6 mo total antibiotics)^b
- Completion of a fluoroquinolone-based regimen for a gram-negative infection^b
- Culture-negative infection^b

Abbreviations: DAIR, debridement, antibiotics, and implant retention; ESRD, end-stage renal disease; HIV, human immunodeficiency virus; PJI, prosthetic joint infection; SAT, suppressive antibiotic therapy; TNF, tumor necrosis factor.

^aBased on the consensus opinion of the authors, but not directly supported by the data identified in this review.

^bBased on retrospective data identifying this as a risk factor for treatment failure.

Conclusions

Quels patients?

- Ablation du matériel impossible:
 - Comorbidités
 - Raréfaction osseuse
 - Age
- Refus du patient
- Echecs multiples
- Micro-organisme résistant aux molécules de choix?
- Prise en charge initiale inappropriée?

Quels traitements?

- Pas de conclusion possible sur le traitement optimal:
 - Molécules?
 - Posologies?
- Cyclines au long court sont bien tolérées
- Rifampicine à éviter
- Place des nouvelles molécules
 - Tedizolide
 - Dalbavancine
- Durée: le plus longtemps possible

Quelle efficacité?

- 50 à 80% de réussite
- Suivi souvent trop court
- Reculer pour mieux sauter: quelle importance?

Le traitement
suppressif at-
il un avenir?

Pose primaire de prothèse X6
d'ici 2030

Poses de plus en plus jeune
(obésité et genou)

Changement de prothèse plus
fréquent

Vieillessement/comorbidités

Merci de votre attention

Quelles recommandations?



3.3.2.2.6 Antibiothérapie suppressive

Elle consiste à maintenir une antibiothérapie orale dans la grande majorité des cas pour une durée indéterminée dans le but d'inhiber la multiplication bactérienne autour de la prothèse.

Elle ne s'applique qu'aux situations pour lesquelles la documentation bactérienne est connue et pour lesquelles l'infection persiste chez un malade inopérable ayant une prothèse non descellée. Elle ne se conçoit qu'avec des molécules bien supportées, d'administration aisée (voie orale) et pour lesquelles une monothérapie est possible [131, 242-244] (**grade C**).



Contents lists available at [ScienceDirect](#)

The Journal of Arthroplasty

journal homepage: www.arthroplastyjournal.org



Hip and Knee Section, Treatment, Antimicrobial Suppression: Proceedings of International Consensus on Orthopedic Infections

Federico Calabrò ¹, Massimo Coen ¹, Massimo Franceschini ¹,
Rafael Franco-Cendejas ¹, Angela Hewlett ², John Segreti ², Eric Senneville ³

- For patients in whom surgery is contraindicated because of the patient's general condition,
- when surgery is not expected to improve the functional outcome for patient, such as those with multiple prior failures
- for patients who refuse surgery.