

A propos d'un cas d'endocardite

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Cours d'Automne 2023

Déclaration des liens d'intérêts 2018 - 2023

- ▶ **Intervenante au titre d'orateur**

GSK vaccins, MSD, Moderna, Novartis, Pfizer, Sanofi, ViiVhealth Care

- ▶ **Participation à des groupes de travail**

Gilead, GSK, Janssen, Mundipharma

- ▶ **Invitation congrès/journées scientifiques**

Eumedica, Novartis, Pfizer, Sanofi

- ▶ **Vice-présidente de la SPILF**

- ▶ **Vice-présidente du Conseil National Professionnel des maladies infectieuses et tropicales**

- ▶ **Vice-présidente de la Cs MIME du HCSP**

Cas clinique (1)

- ▶ Femme de 70 ans, Guyane
- ▶ ATCD : HTA, abcès de cornée à pneumocoque post cataracte (2020)
- ▶ Admission 08/2023 : fièvre depuis 48H + dyspnée + épigastralgies + vomissement
- ▶ Ex clinique initial : 40,6 °C, FR 18/mn, FC 99/mn sans souffle perçu (?), pas de foyer de crépitants, douleur HCD
- ▶ Sat O2 98% AA

Cas clinique (2)

- ▶ Biologie :
 - ▶ NFS : leucocytes 16 000/mm³ dont 85% PNN; plaquettes 168 000 /mm³ et Hb 12,6 g /dl
 - ▶ ASAT 143 UI/l (N<35), ALAT 208 UI/l,
 - ▶ Gamma GT 180 UI/l (N< 50); Ph alcalines 120 UI/L, bili totale 20 mmol/l
 - ▶ CRP 170 mg/l, clairance créat 97 ml/mn (MDRD)
 - ▶ 4 hémocultures positives à *S. pneumoniae*

Bilan complémentaire ?

Cas clinique (3)

- ▶ Sérologies VIH, VHB, VHC, HTLV1 négatives
- ▶ EPP normale
- ▶ Dosage pondéral des Ig : IgM < 0,20 g/l
- ▶ Immunophénotypage lymphocytaire normal
- ▶ RxP normale
- ▶ Echographie abdominale normale

Cas clinique (4)

▶ ETT à J4 :

- Hypertrophie septale globale, FEVG préservée (77 %)
- fuite mitrale modérée, par dysfonction du feuillet postérieur;
- **image mobile suspecte de végétation** dans le contexte, appendue à la face auriculaire de l'anneau mitral postérieur (12 X 2.5 mm)
- OG dilatée (44 ml/m²)

Endocardite à pneumocoque ?

- ▶ Possible car fréquence = 10% des IE sur valve native ?
- ▶ Peu probable car pas de FRD d'infection invasive à pneumocoque ?
- ▶ Peu probable car pas de valve prothétique ?
- ▶ Peu probable car pas de foyer de pneumonie ?

Endocardite à pneumocoque :

- ▶ < 2% des EI (Selton-Sury C et al, CID 2012)
- ▶ 0,86% cohorte 111 cas (de Egea V et al, Medicine 2015)
- ▶ (15% des EI avant 1980)
- ▶ 0.3% des infections à pneumocoque (Marrie TJ et al, Eur J Clin Microbiol Infect Dis 2018)

Table 3. Distribution of Causative Microorganisms in Patients With Infective Endocarditis

| Microorganisms | No. (%) of Patients (n = 497) | |
|-------------------------------------|----------------------------------|--------|
| Streptococcaceae | 240 | (48.3) |
| Streptococci | 180 | (36.2) |
| Oral streptococci ^a | 93 | (18.7) |
| Group D streptococci ^b | 62 | (12.5) |
| Pyogenic streptococci | 25 | (5.0) |
| Enterococci | 52 | (10.5) |
| Other Streptococcaceae ^c | 8 | (1.6) |
| Staphylococcaceae | 180 | (36.2) |
| <i>Staphylococcus aureus</i> | 132 | (26.6) |
| Coagulase-negative staphylococci | 48 | (9.7) |
| Other microorganisms ^d | 42 | (8.5) |
| HACEK group | 6 | ... |
| Enterobacteriaceae | 4 | ... |
| <i>Propionibacterium acnes</i> | 4 | ... |
| <i>Pseudomonas aeruginosa</i> | 3 | ... |
| <i>Lactobacillus</i> species | 2 | ... |
| <i>Corynebacterium</i> species | 2 | ... |
| <i>Coxiella burnetii</i> | 2 | ... |
| <i>Bartonella quintana</i> | 1 | ... |
| <i>Tropheryma whipplei</i> | 1 | ... |
| <i>Candida</i> species | 6 | ... |
| Miscellaneous ^e | 11 | ... |
| ≥2 Microorganisms ^f | 9 | (1.8) |
| No microorganism identified | 26 | (5.2) |

Clinique

Demographic and clinical characteristics of 50 patients with definite pneumococcal endocarditis.

| Characteristics ^a | N = 50 |
|-------------------------------------------------------|----------|
| Baseline | |
| Age at diagnosis (years), mean ± SD | 60 ± 14 |
| Male gender | 38 (76) |
| Clinical and biological manifestations | |
| Fever | |
| Vascular phenomena ^b | |
| Immunological phenomena ^c | |
| Pneumonia | |
| Meningitis | |
| Arthritis | |
| Ocular infection | |
| Other infectious localizations ^d | |
| Blood cultures positive for <i>S. pneumoniae</i> | |
| Positive histopathological examination and/or culture | |
| Affected valve | |
| Native | |
| Prosthesis | |
| Mitral | |
| Aortic | |
| Tricuspid | |
| Transthoracic echocardiographic findings | |
| TTE positive for IE | 50 (100) |
| Vegetations | 44 (88) |
| Abscess | 14 (30) |
| Valve regurgitation | 38 (79) |
| Dehiscence of prosthesis | 0 |

TABLE 2. Demographic and Clinical Characteristics of 111 Patients With *Streptococcus pneumoniae* Endocarditis

| Characteristics | Total Cohort N = 111 (%) | Spanish Cohort n = 24 | Literature Review n = 87 |
|--------------------------|--------------------------|-----------------------|--------------------------|
| Age (median, IQR) | 51 (26–63) | 57 (50–69) | 47 (15–61) |
| Sex | | | |
| Male | 71 (64) | 18 (75) | 53 (60.9) |
| Congenital heart disease | 11 (0.0) | 0 (0) | 11 (12.9) |
| | | 6 (27.3) | 2 (2.4) |
| | | 1 (4.2) | 5 (6.4) |
| | | 21 (87.5) | 58/61 (95,1) |
| | | 17 (77.3) | 44/55 (80.0) |
| | | 9 (37.5) | 42/84 (50.0) |
| | | 7 (29.2) | 38/84 (45.2) |
| | | 3 (12.5) | 26/84 (31.0) |
| | | 0 (0) | 5/62 (8.1) |
| | | 3 (13) | 0 (0) |
| | | 20 (83.3) | 84 (96.6) |
| | | 4 (16.7) | 3 (3.4) |
| | | 14 (58.3) | 45 (51.7) |
| | | 10 (41.7) | 35 (40.2) |
| | | 3 (12.5) | 11 (12.6) |
| | | 3 (12.5) | 12 (13.8) |

Âge
Localisations associées +++
Pneumonie
Méningite
Arthrite
Valve native (93,7%), aorte (53,2%)



Facteurs de risque ?

- ▶ Endocardite à pneumocoque ?
- ▶ Etude multicentrique / 15 ans
- ▶ Âge 60 ± 14 ans, 38 H (76%)
 - ▶ 51 ans (IQR, 26-63), 71 H (64%)
- ▶ Peu de FDR EI
- ▶ 78% FDR infection à pneumocoque

Prognosis of *Streptococcus pneumoniae* endocarditis in France, a multicenter observational study (2000–2015)☆

Amandine Périer ^{a,b}, Mathieu Puyade ^{a,b}, Matthieu Revest ^{c,d}, Pierre Tattevin ^{c,d}, Louis Bernard ^{e,f}, Adrien Lemaigen ^{e,f}, David Boutoille ^{g,h}, Joseph Allal ^{a,i}, France Roblot ^{a,j,k}, Blandine Rammaert ^{a,j,k,*}

Table 1

Demographic and clinical characteristics of 50 patients with definite pneumococcal endocarditis.

| Characteristics ^a | N = 50 |
|-----------------------------------------|-------------|
| Baseline | |
| Age at diagnosis (years), mean \pm SD | 60 \pm 14 |
| Male gender | 38 (76) |
| IPD predisposing conditions | |
| Asplenia | 2 |
| >65 years | 18 (36) |
| Chronic pulmonary disease | 12 (24) |
| Repeated ENT infections | 3 |
| Cardiopathy | 12 (24) |
| Alcoholism | 17 (34) |
| Malignant blood disease | 6 |
| Renal failure | 3 |
| Kidney transplantation | 1 |
| IPD risk factors | |
| None | 11 (22) |
| 1 | 12 (24) |
| ≥ 2 | 27 (54) |
| No vaccine uptake before IE | 48 (96) |
| IE predisposing conditions | |
| Valvular prosthesis | 5 |
| Previous IE | 1 |
| Congenital cardiopathy | 2 |
| Valvular disease | 10 (22) |
| Intravenous drug use | 1 |
| Pacemaker | 4 |
| IE risk factors | |
| None | 38 (76) |
| 1 | 6 |
| ≥ 2 | 6 |

Clinique et FDR ?

- ▶ 3251 adultes + IIP 2000-2014, Canada
- ▶ 28 EI (0.3%)
- ▶ UDIV
- ▶ Présentation clinique initiale + sévère (confusion mentale, USI; $p < 0,005$)
- ▶ Souffle valvulaire nouveau 39.3% vs 2.2% si pas EI ($p < 0.001$)
- ▶ **Mortalité 39.3%** vs 14.7% IIP sans EI
- ▶ Pas de sérotype spécifique

Quel bilan pour Mme L... ?

- ▶ ETO ?
- ▶ IRM cérébrale ?
- ▶ TEP/TDM ?
- ▶ TDM abdominale ?

2023 ESC Guidelines for the management of endocarditis

▶ ETO si ETT contributive ?



Section 5. Recommendation Table 5 — Recommendations for the role of echocardiography in infective endocarditis

TOE is recommended when the patient is stable before switching from intravenous to oral antibiotic therapy.

I

B

Recommendation Table 5 — Recommendations for the role of echocardiography in infective endocarditis

| Recommendations | Class ^a | Level ^b |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------|--------------------|
| A. Diagnosis | | |
| TTE is recommended as the first-line imaging modality in suspected IE. ^{166,179} | I | B |
| TOE is recommended in all patients with clinical suspicion of IE and a negative or non-diagnostic TTE. ^{166,178,179} | I | B |
| TOE is recommended in patients with clinical suspicion of IE, when a prosthetic heart valve or an intracardiac device is present. ^{166,178,179} | I | B |
| Repeating TTE and/or TOE within 5–7 days is recommended in cases of initially negative or inconclusive examination when clinical suspicion of IE remains high. ¹⁷⁸ | I | C |
| TOE is recommended in patients with suspected IE, even in cases with positive TTE, except in isolated right-sided native valve IE with good quality TTE examination and unequivocal echocardiographic findings. ^{165,166,179} | I | C |
| Performing an echocardiography should be considered in <i>S. aureus</i> , <i>E. faecalis</i> , and some <i>Streptococcus</i> spp. bacteraemia. ^{19,149,174} | IIa | B |

Intérêt ETO + ETT ?

Prevalence of infective endocarditis in patients with *Staphylococcus aureus* bacteraemia: the value of screening with echocardiography

Rasmus V. Rasmussen^{1*}, Ulla Høst², Magnus Arpi³, Christian Hassager⁴, Helle K. Johansen⁵, Eva Korup⁶, Henrik C. Schönheyder⁷, Jens Berning⁸, Sabine Gill⁹, Flemming S. Rosenvinge¹⁰, Vance G. Fowler Jr¹¹, Jacob E. Møller⁴, Robert L. Skov¹², Carsten T. Larsen¹, Thomas F. Hansen¹, Shan Mard², Jesper Smit⁷, Paal S. Andersen¹², and Niels E. Bruun¹

- ▶ 244 Bactériémies à *S. aureus*, 2009 - 2010
 - ▶ 53 EI certaines (22%)
 - ▶ Prévalence sur valve native 19%
 - ▶ Prévalence sur valve prothétique ou matériel intracardiaque 38%

- ▶ 92 patients ETT sans ETO
- ▶ 8 diagnostics d'EI sur ETT
- ▶ suivi à J30 : pas de nouveau diagnostic d'EI
- ▶ ETT de qualité « suffisante » dans un contexte où prévalence = 19%

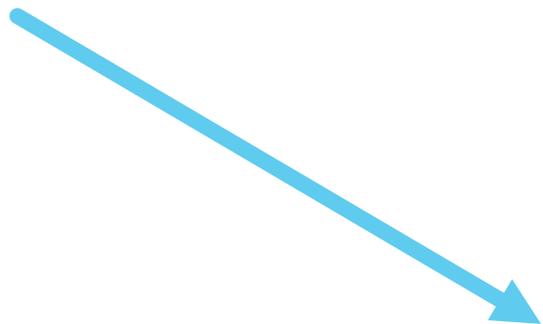
Intérêt ETO + ETT ?

TABLE 3. Diagnostic Characteristics and Complications of 111 Patients With Pneumococcal Endocarditis

| Variable | Total Cohort N = 111 | Spanish Cohort n = 24* | Literature Review n = 87* | P |
|----------------------------|----------------------|------------------------|---------------------------|------|
| Diagnostic characteristics | | | | |
| Positive blood cultures | 94 (84.7) | 22 (91.7) | 72/82 (87.8)* | 0.7 |
| Other positive cultures | 66/90 (73.3)* | 12 (57.1) | 54/69 (78.3)* | 0.08 |
| CSF | 30 (33.3) | 5 (25.0) | 17 (31.5) | |
| Heart valves | 14 (21.2) | 5 (41.7) | 9 (16.7) | |
| Respiratory tract | 11 (12.2) | 1 (8.3) | 3 (5.6) | |
| Synovial fluid | 4 (6.1) | 0 (0) | 4 (7.4) | |
| Other | 6 (9.1) | 2 (16.7) | 5 (9.3) | |
| Antimicrobial resistance | | | | |
| Nonpenicillin susceptible | 15/74 (20.3)* | 1 (4.2) | 14/54 (25.9)* | 0.05 |
| Noncefotaxime susceptible | 3/36 (8.3)* | 0 (0) | 3/22 (13.6)* | 0.2 |
| Echocardiography | 109 (98.2) | 24 (100) | 85 (97.7) | 1 |
| TTE only | 27 (24.3) | 6 (25.0) | 21 (36.8) | |
| TEE only | 20 (18.0) | 1 (4.2) | 19 (33.3) | |
| Both | 32 (28.8) | 17 (70.8) | 15 (26.3) | |
| Technique not specified | 30 (27%) | 0 (0%) | 32 (36.8%) | |

Quel Bilan ?

- ▶ TDM cardiaque
- ▶ TEP/TDM si valve prothétique
- ▶ IRM et TDM TAP :
 - ▶ Patients symptomatiques
 - ▶ « Détection » si asymptomatique



Section 5. Recommendation Table 6 — Recommendations for the role of computed tomography, nuclear imaging, and magnetic resonance in infective endocarditis

| | | |
|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------|----------|
| Cardiac CTA is recommended in patients with possible NVE to detect valvular lesions and confirm the diagnosis of IE. | I | B |
| [18F]FDG-PET/CT(A) and cardiac CTA are recommended in possible PVE to detect valvular lesions and confirm the diagnosis of IE. | I | B |
| [18F]FDG-PET/CT(A) may be considered in possible CIED-related IE to confirm the diagnosis of IE. | IIa | B |
| Cardiac CTA is recommended in NVE and PVE to diagnose paravalvular or periprosthetic complications if echocardiography is inconclusive. | I | B |
| Brain and whole-body imaging (CT, [18F]FDG-PET/CT, and/or MRI) are recommended in symptomatic patients with NVE and PVE to detect peripheral lesions or add minor diagnostic criteria. | I | B |
| WBC SPECT/CT should be considered in patients with high clinical suspicion of PVE when echocardiography is negative or inconclusive and when PET/CT is unavailable. | IIa | C |
| Brain and whole-body imaging (CT, [18F]FDG-PET/CT, and MRI) in NVE and PVE may be considered for screening of peripheral lesions in asymptomatic patients. | IIb | B |

Quel bilan ?



SERVICE D'IMAGERIE MOLECULAIRE – TEP

Identifiants

30081953

253533



TEP au 18 FDG

Indication :

Recherche de foyer infectieux profond.

Technique :

TEP VISION 450 (Siemens N°1011) du 04/10/2019 ; Archivage PACS ; Glycémie 0.90 g/l ; Poids 66.0 kg; Taille 160 cm. Injection IV de 148 MBq de 18-FDG. Scanner : PDL = 508 mGy.cm Prescription d'un régime hypoglycémique.

Résultats :

Etage sus-diaphragmatique :

Pas d'hyperfixation suspecte de la sphère ORL, des glandes mammaires, des aires ganglionnaires cervicales, médiastino-hilaires ou axillaires.

Un foyer modérément hypermétabolique à la partie supérieure de la valve mitrale, suspect d'endocardite.

Pas de nodule pulmonaire solide hypermétabolique. Un nodule calcifié non métabolique du lobe moyen, d'allure séquellaire.

Etage sous-diaphragmatique :

Pas d'hyperfixation suspecte hépatique, splénique, surrénalienne, rénale, pancréatique, des anses digestives ou des aires ganglionnaires abdominopelviques.

Un foyer modérément hypermétabolique au contact d'une plaque calcifiée de la partie proximale de l'artère iliaque primitive gauche : plaque instable ? Greffe septique ?

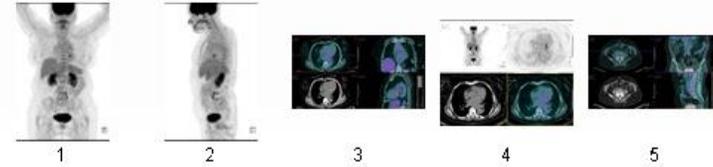
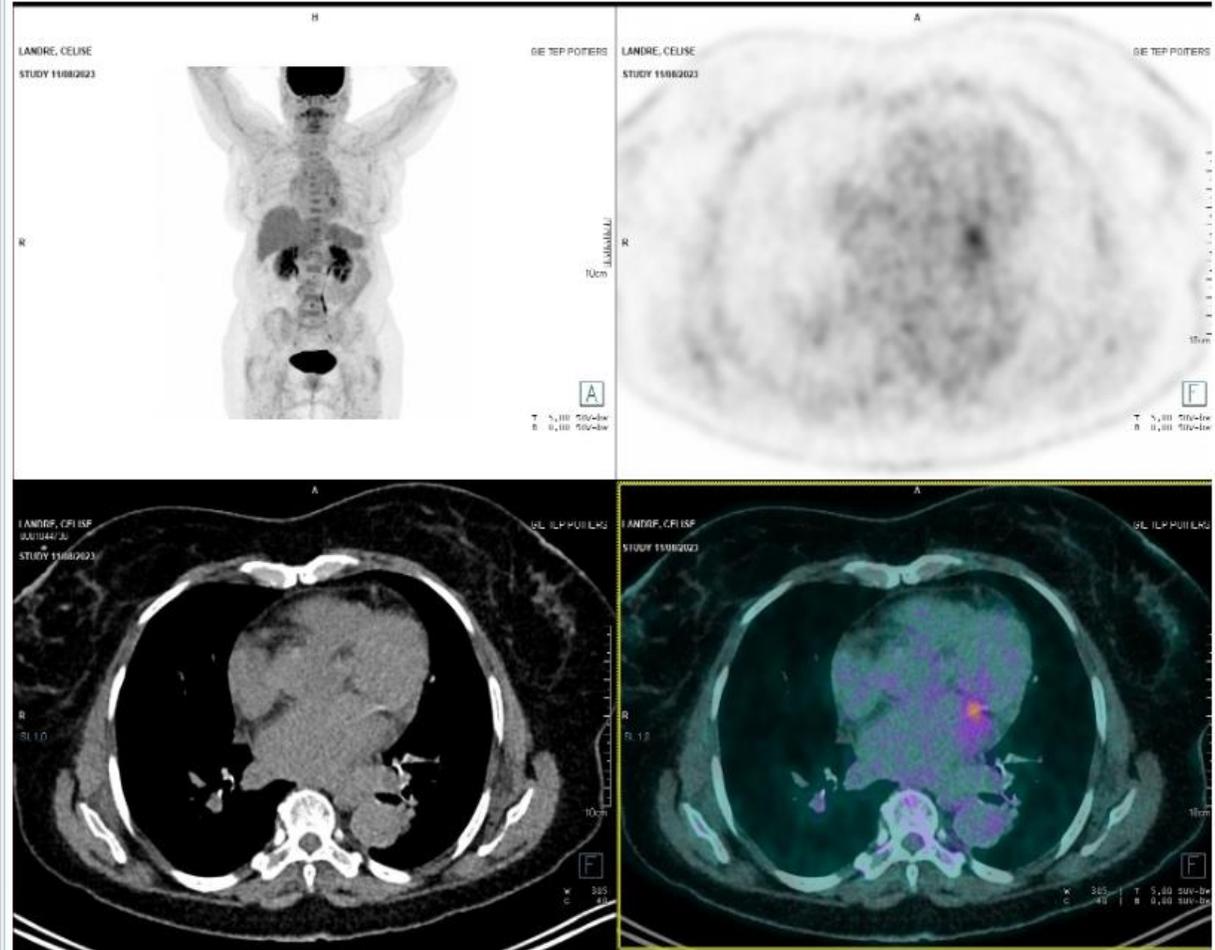


Image 4 / 5



Quel traitement ?

- ▶ Amoxicilline 100 mg/kg/j ?
- ▶ + Gentamicine 3 mg/kg pendant 2 semaines ?
- ▶ + Rifampicine pendant 2 semaines ?
- ▶ Ceftriaxone 2 G / j
- ▶ Durée totale ?
- ▶ Relais oral ?

Traitement ?

Table 2

Treatment and outcome of 50 patients with definite pneumococcal endocarditis.

| | N = 50 |
|---------------------------------------------------------------------------|---------------|
| Time from 1st symptoms onset to antibiotherapy, days (mean \pm SD) | 6.3 \pm 6.9 |
| Number of antibiotics, <i>n</i> (%) | |
| Monotherapy | 7 |
| Bitherapy | 28 (56) |
| ≥ 3 antibiotics | 15 (30) |
| Antibiotic choice, <i>n</i> (%) | |
| 3rd generation cephalosporin or amoxicillin or penicillin G or vancomycin | 50 (100) |
| Gentamicin | 39 (78) |
| Antibiotic duration (mean \pm SD) | |
| Total antibiotherapy duration, weeks | 5.5 \pm 2.3 |
| Beta-lactam or vancomycin duration, days | 37 \pm 14.7 |
| Gentamicin duration, days | 8.5 \pm 8.4 |

Quel traitement ?

- ▶ Amoxicilline 100 mg/kg perfusion continue
- ▶ 6 semaines
- ▶ Relais oral ?
- ▶ 400 EI coeur gauche
- ▶ 10 j de TTT ATBT IV
 - ▶ 199 IV
 - ▶ 201 relais oral (bithérapie)
- ▶ Nombre de *S. pneumoniae* ?

Partial Oral versus Intravenous Antibiotic Treatment of Endocarditis

Table 1. Characteristics of the Patients at Baseline.*

| Characteristic | Intravenous Treatment (N=199) | Oral Treatment (N=201) |
|-----------------------------------------------|-------------------------------|------------------------|
| Mean age — yr | 67.3±12.0 | 67.6±12.6 |
| Female sex — no. (%) | 50 (25.1) | 42 (20.9) |
| Body temperature — °C | 36.9±0.45 | 37.0±0.44 |
| Coexisting condition or risk factor — no. (%) | | |
| Diabetes | 36 (18.1) | 31 (15.4) |
| Renal failure | 25 (12.6) | 21 (10.4) |
| Dialysis | 13 (6.5) | 15 (7.5) |
| COPD | 17 (8.5) | 9 (4.5) |
| Liver disease | 7 (3.5) | 6 (3.0) |
| Cancer | 14 (7.0) | 18 (9.0) |
| Intravenous drug use | 3 (1.5) | 2 (1.0) |
| Pathogen — no. (%)† | | |
| Streptococcus | 104 (52.3) | 92 (45.8) |
| <i>Enterococcus faecalis</i> | 46 (23.1) | 51 (25.4) |
| <i>Staphylococcus aureus</i> ‡ | 40 (20.1) | 47 (23.4) |
| Coagulase-negative staphylococci | 10 (5.0) | 13 (6.5) |

Quel traitement ?

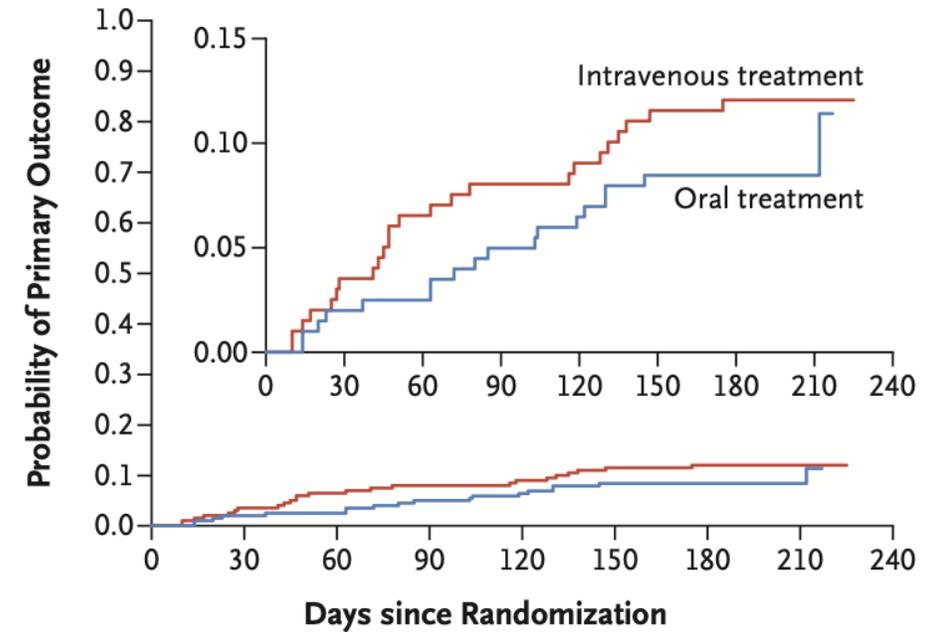
- ▶ Pas de différence à 6 mois
 - ▶ Mortalité
 - ▶ Embols
 - ▶ Chirurgie cardiaque en urgence
 - ▶ Rechute de bactériémie

Table 2. Distribution of the Four Components of the Primary Composite Outcome.*

| Component | Intravenous Treatment (N=199) | Oral Treatment (N=201) | Difference | Hazard Ratio (95% CI) |
|----------------------------------------|-------------------------------|------------------------|----------------------------|-----------------------|
| | number (percent) | number (percent) | percentage points (95% CI) | |
| All-cause mortality | 13 (6.5) | 7 (3.5) | 3.0 (-1.4 to 7.7) | 0.53 (0.21 to 1.32) |
| Unplanned cardiac surgery | 6 (3.0) | 6 (3.0) | 0 (-3.3 to 3.4) | 0.99 (0.32 to 3.07) |
| Embolic event | 3 (1.5) | 3 (1.5) | 0 (-2.4 to 2.4) | 0.97 (0.20 to 4.82) |
| Relapse of the positive blood culture† | 5 (2.5) | 5 (2.5) | 0 (-3.1 to 3.1) | 0.97 (0.28 to 3.33) |

* Six patients, three in each group, had two outcomes.

† For details about relapse of the positive blood culture, see the Supplementary Appendix.



No. at Risk

| | | | | | | | | | |
|-----------------------|-----|-----|-----|-----|-----|-----|-----|----|---|
| Intravenous treatment | 199 | 192 | 186 | 183 | 181 | 176 | 174 | 28 | 0 |
| Oral treatment | 201 | 197 | 196 | 191 | 188 | 184 | 183 | 36 | 0 |

Figure 2. Kaplan–Meier Plot of the Probability of the Primary Composite Outcome.

The primary composite outcome was all-cause mortality, unplanned cardiac surgery, embolic events, or relapse of bacteremia with the primary pathogen, from randomization until 6 months after antibiotic treatment was completed. The oral treatment group shifted from intravenously administered antibiotics to orally administered antibiotics at a median of 17 days after the start of treatment. The inset shows the same data on an enlarged y axis.

Et à 5 ans ?

Five-Year Outcomes of the Partial Oral Treatment of Endocarditis (POET) Trial

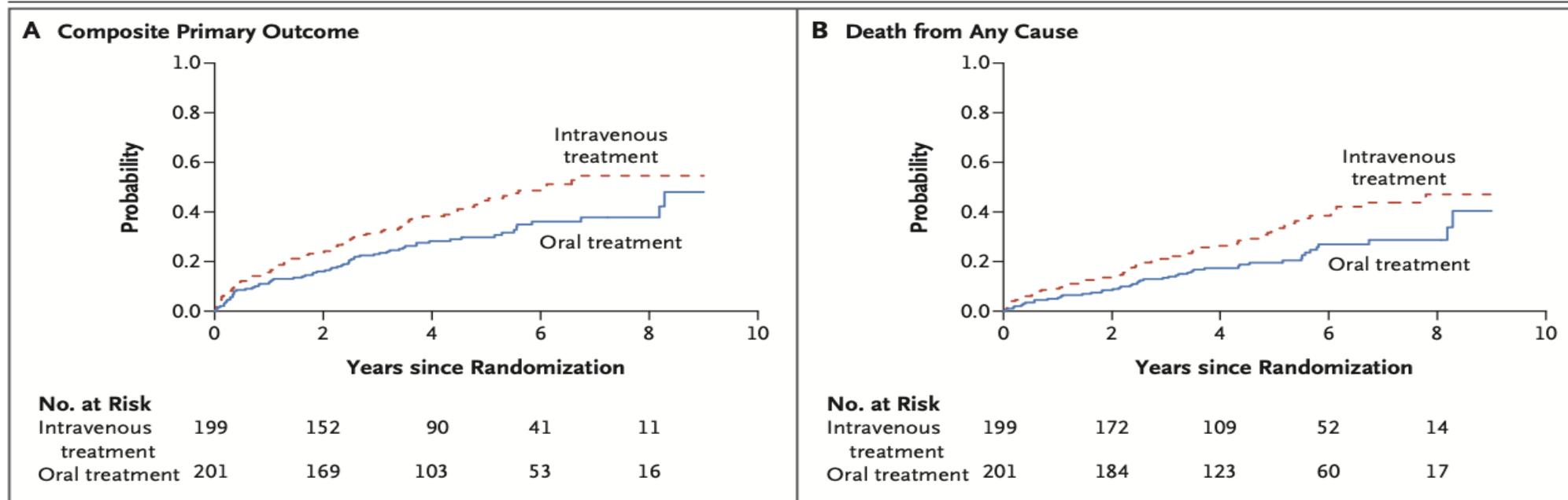


Figure 1. Cumulative Incidence of Events.

Shown are plots of the cumulative incidence of events from randomization to a median follow-up of 5.4 years. The patients assigned to the intravenous treatment group received intravenous antibiotic therapy for the entire treatment period, and the patients assigned to receive step-down treatment shifted from intravenous antibiotics to oral antibiotics after clinical stabilization was reached. The composite primary outcome consisted of death from any cause, unplanned cardiac surgery, embolic events, and relapse of a blood culture result positive for the primary pathogen.

Relais oral pour les endocardites à *S. pneumoniae* ?

Attainment of Target Antibiotic Levels by Oral Treatment of Left-Sided Infective Endocarditis: A POET Substudy

- ▶ Analyse PK/PD, POET
- ▶ Amoxicilline per os 1 g/ 6h
- ▶ Probabilité d'atteindre la cible
 - ▶ BP (n=236) et / CMI (N=74)

Table 1. Clinical Breakpoints and Pharmacokinetic/Pharmacodynamic Targets

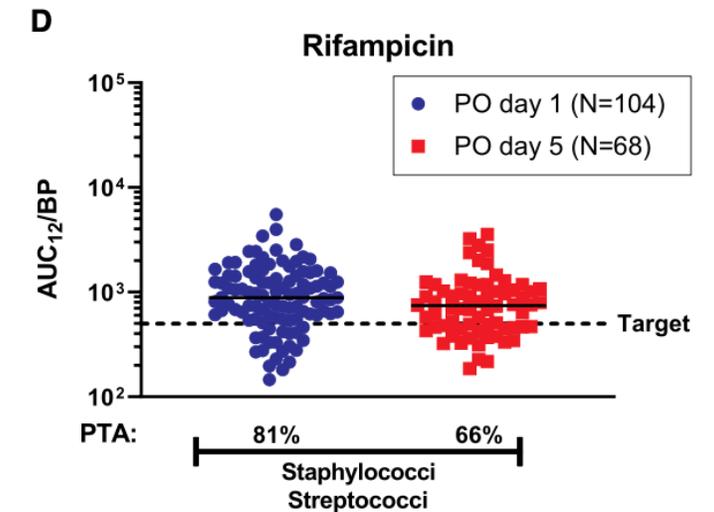
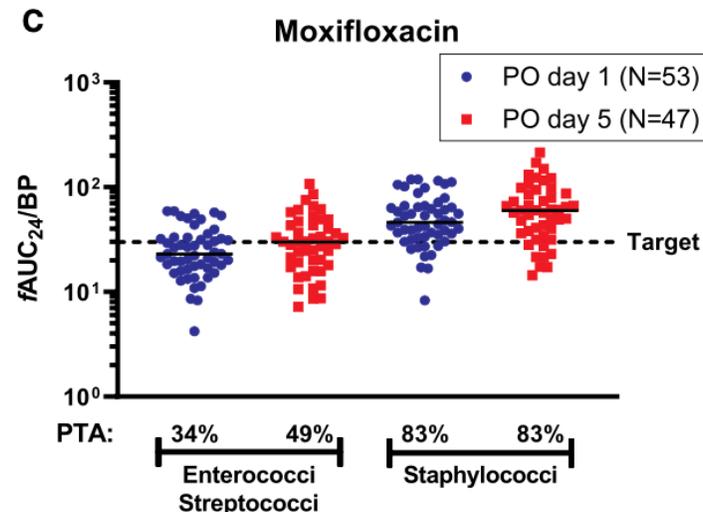
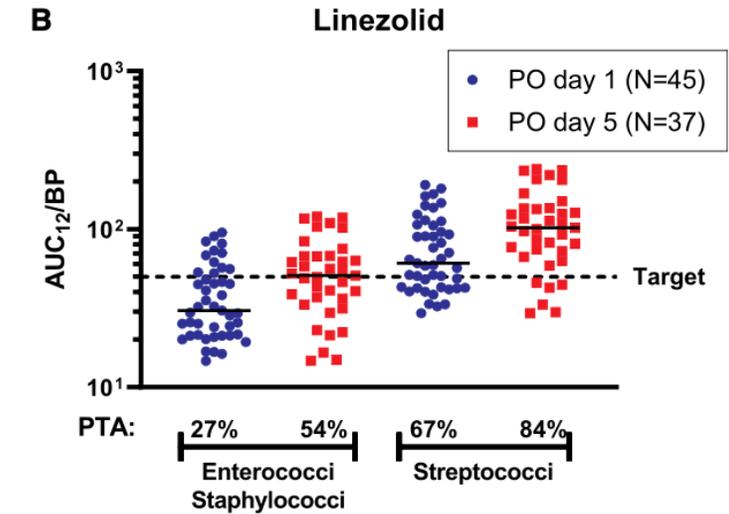
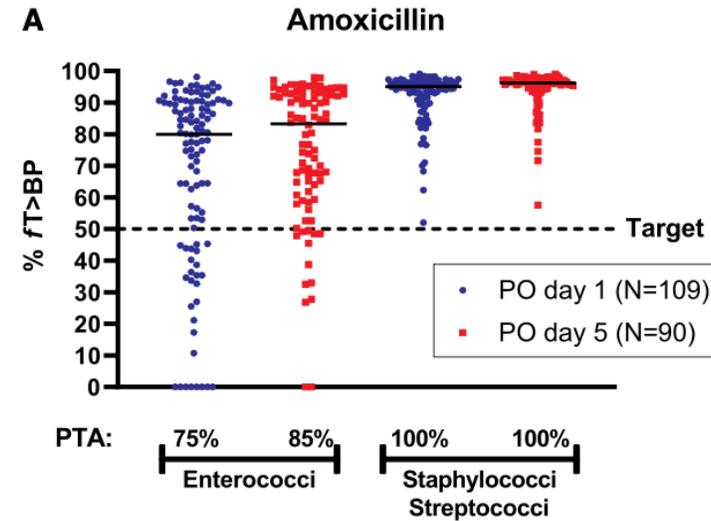
| Drug | Dose | Approximate Unbound Fraction (%) | Bacterial Species | Clinical Breakpoint (mg/L) | Pharmacokinetic/ Pharmacodynamic Target |
|-------------------------|--------------------------------------|----------------------------------|-------------------------------|----------------------------|-----------------------------------------------------------------------|
| Amoxicillin | PO: 1000 mg q6 h | 80 | Enterococci | 4 | $fT > BP$ or $fT > MIC$ more than 50% of dosing interval ^b |
| | | | Staphylococci Streptococci | 0.5 ^a | |
| Dicloxacillin | PO: 1000 mg q6 h IV: 3000 mg q6 h | 3 | Staphylococci | 0.5 ^c | $fT > BP$ more than 50% of dosing interval ^b |
| Linezolid | PO or IV: 600 mg q12 h | Not used | Enterococci Staphylococci | 4 | $AUC_{12}/BP > 50$ or $AUC_{12}/MIC > 50$ |
| | | | Streptococci | 2 ^d | |
| Moxifloxacin | PO or IV: 400 mg q24 h | 50 | Enterococci Streptococci | 0.5 ^e | $fAUC_{24}/BP > 30$ or $fAUC_{24}/MIC > 30$ ^b |
| | | | Staphylococci | 0.25 | |
| Rifampicin ^f | PO: 600 mg q12 h | Not used | Staphylococci Streptococci | 0.064 | $AUC_{12}/BP > 500$ or $AUC_{12}/MIC > 500$ |

(Bock M et al Clin Infect Dis 2023)

Objectifs PK/PD ? (1)

Attainment of Target Antibiotic Levels by Oral Treatment of Left-Sided Infective Endocarditis: A POET Substudy

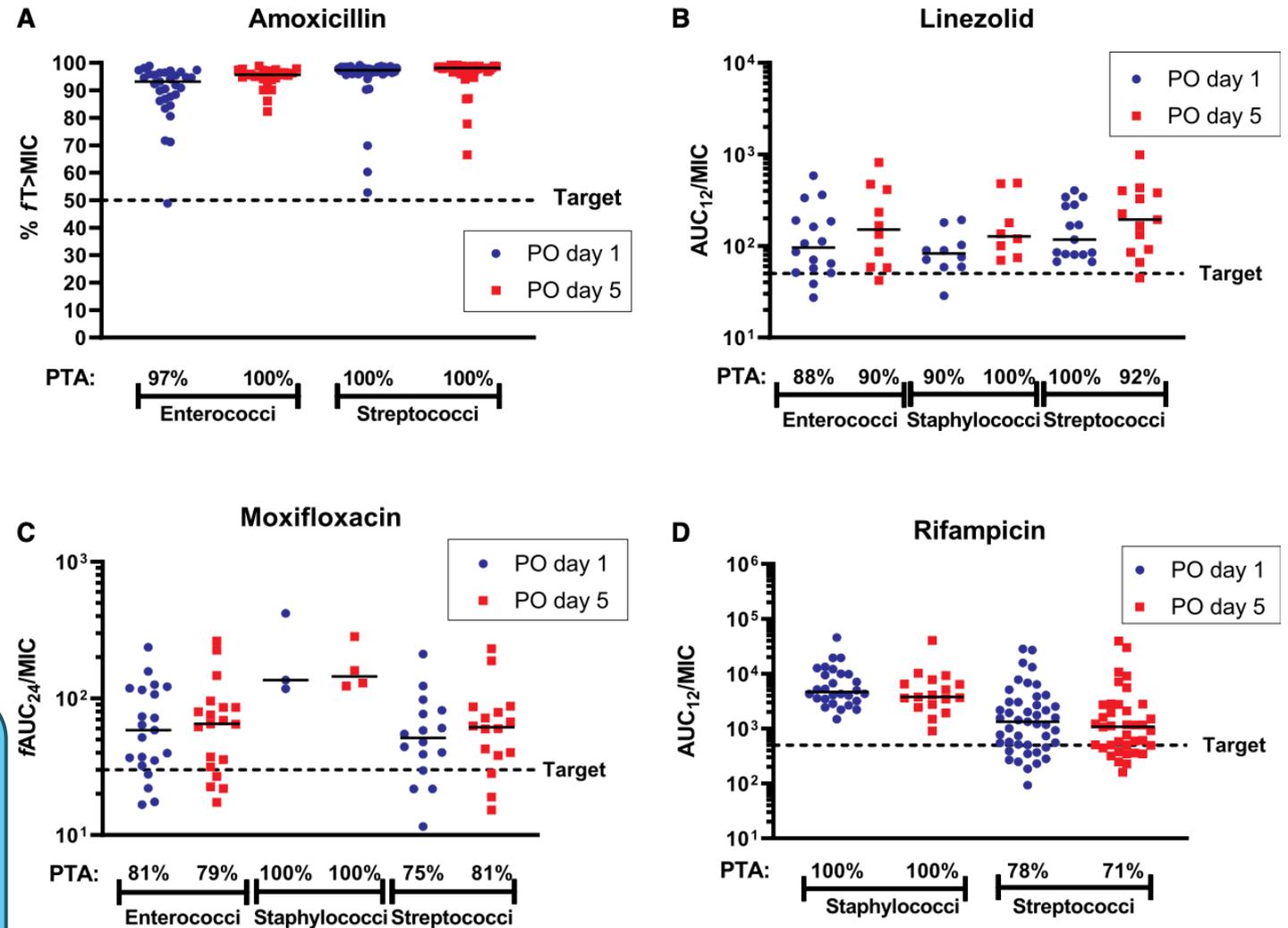
- Probabilité d'atteindre la cible / BP
- Amoxicilline et Linézolide 88 - 100%
- Moxifloxacine et rifampicine 71 - 10



Objectifs PK/PD ? (2)

- ▶ Probabilité d'atteindre la cible / CMI
- ▶ J1 : 13 patients n'atteignent la cible que pour 1 ATBT
- ▶ J5 : 14 patients n'atteignent la cible que pour 1 ATBT

Attainment of Target Antibiotic Levels by Oral Treatment of Left-Sided Infective Endocarditis: A POET Substudy



Posologie Amoxicilline per os ?
TDM

Figure 4. Target attainment of oral antibiotics in relation to minimal inhibitory concentrations. Solid black bars are median values. The letter *f* indicates the free unbound concentration; eg, $fT > MIC$ means the time above MIC of the unbound concentration. Abbreviations: AUC, area under concentration-time curve; MIC, minimal inhibitory concentration; PO, oral; PTA, probability of target attainment; T, time.

7.4. *Streptococcus pneumoniae*, β -haemolytic streptococci (groups A, B, C, and G)

Traitement

- ▶ Idem Streptocoques oraux
- ▶ Pas de schéma en 2 semaines
- ▶ Si PSDP : ceftriaxone ou cefotaxime ou vancomycine
- ▶ Si méningite pas d'amoxicilline
- ▶ Relais oral possible à J10-J14 : 2 antibiotiques ?



Pronostic

Prognosis of *Streptococcus pneumoniae* endocarditis in France, a multicenter observational study (2000–2015)☆

Amandine Périer ^{a,b}, Mathieu Puyade ^{a,b}, Matthieu Revest ^{c,d}, Pierre Tattevin ^{c,d}, Louis Bernard ^{e,f}, Adrien Lemaigen ^{e,f}, David Boutoille ^{g,h}, Joseph Allal ^{a,i}, France Roblot ^{a,j,k}, Blandine Rammaert ^{a,j,k,*}

- ▶ Chirurgie dans la semaine suivant le diagnostic (28, 60%; délai 6.5 j [2.0-10.5])
- ▶ Taux de survie à J90 : 83% (n=33)
 - ▶ 7 décès dont 6 liées à l'EI
 - ▶ 5 EI sur valve prothétique: 3 décès délai 22j (21-40,5)
 - ▶ 2 rechutes 48H et 6 mois après arrêt ATBT
- ▶ Taux de survie à 2 ans : 67% (n=28)
 - ▶ 5 décès
 - ▶ Âge > 65 ans (p<0,001)
 - ▶ Chirurgie “protectrice” (15 vs 6; p = 0.012)
 - ▶ Méningite ns, Austrian syndrome ns
- ▶ 50% des décès directement liés à l'EI

Pronostic

- ▶ Etude cas/témoins (23 ans, monocentrique)
- ▶ 28 EI à pneumocoque / 56 EI autres causes (21 *S. aureus*)
- ▶ Alccolisme et tabagisme
- ▶ Absence de valvulopathie
- ▶ Gravité clinique : choc et insuffisance cardiaque
- ▶ Chirurgie cardiaque 64,3%
 - ▶ précoce (14.1 ± 18.2 versus 69.0 ± 61.1 j)

(M. Daudin et al, Clin Microbiol Infect 2016)

TABLE I. Comparison of pneumococcal endocarditis (cases), and non-pneumococcal infective endocarditis (controls^a)

| Characteristics | Pneumococcal endocarditis (n = 28) | Endocarditis due to other bacteria (n = 56) | p value |
|----------------------------------------------------|------------------------------------|---------------------------------------------|---------|
| Baseline | | | |
| Age (years), mean \pm SD | 59.1 \pm 15.3 | 60.9 \pm 15.3 | NS |
| Male gender, n (%) | 19 (67.8) | 40 (71.4) | NS |
| Alcoholism, n (%) | 11 (39.3) | 6 (10.7) | <0.01 |
| Smoking, n (%) | 17 (60.7) | 12 (21.4) | < 0.01 |
| Previously known valvular disease, n (%) | 5 (17.9) | 22 (39.3) | 0.047 |
| Valve(s) involved, n (%) | | | |
| Aortic | 19 (70.4) | 35 (62.5) | NS |
| Mitral | 10 (37.0) | 28 (50.0) | NS |
| Tricuspid | 3 (11.1) | 2 (3.6) | NS |
| Pulmonary | 1 (3.7) | 0 (0) | NS |
| Two or more valves | 4 (14.8) | 9 (16.1) | NS |
| Peri-valvular abscess | 8 (34.8) | 17 (30.4) | NS |
| Cardiac surgery, n (%) | | | |
| Time from symptoms onset to surgery, days \pm SD | 14.1 \pm 18.2 | 69.0 \pm 61.1 | <0.001 |
| Time from admission to surgery, days \pm SD | 13.3 \pm 17.1 | 34.3 \pm 43.0 | 0.02 |
| Complications, n (%) | | | |
| Shock | 15 (53.6) | 13 (23.2) | <0.01 |
| Heart failure ^b | 18 (64.3) | 13 (23.2) | <0.01 |
| Embolism | 5 (17.9) | 16 (28.6) | NS |
| Meningitis | 8 (28.6) | 3 (5.4) | <0.01 |
| In-hospital mortality | 2 (7.1) | 7 (12.5) | NS |
| 5-year mortality | 11 (39.3) | 10 (17.9) | NS |

Abbreviations: NS, not significant; SD, standard deviation.

^aControls were endocarditis due to *Staphylococcus aureus* (n = 21), non-pneumococcal *Streptococcus* spp. (n = 20), *Enterococcus* spp. (n = 8), other Gram-positive cocci (n = 4) and Gram-negative bacilli (n = 3).

Pronostic

► Mortalité à 5 ans 54,1%

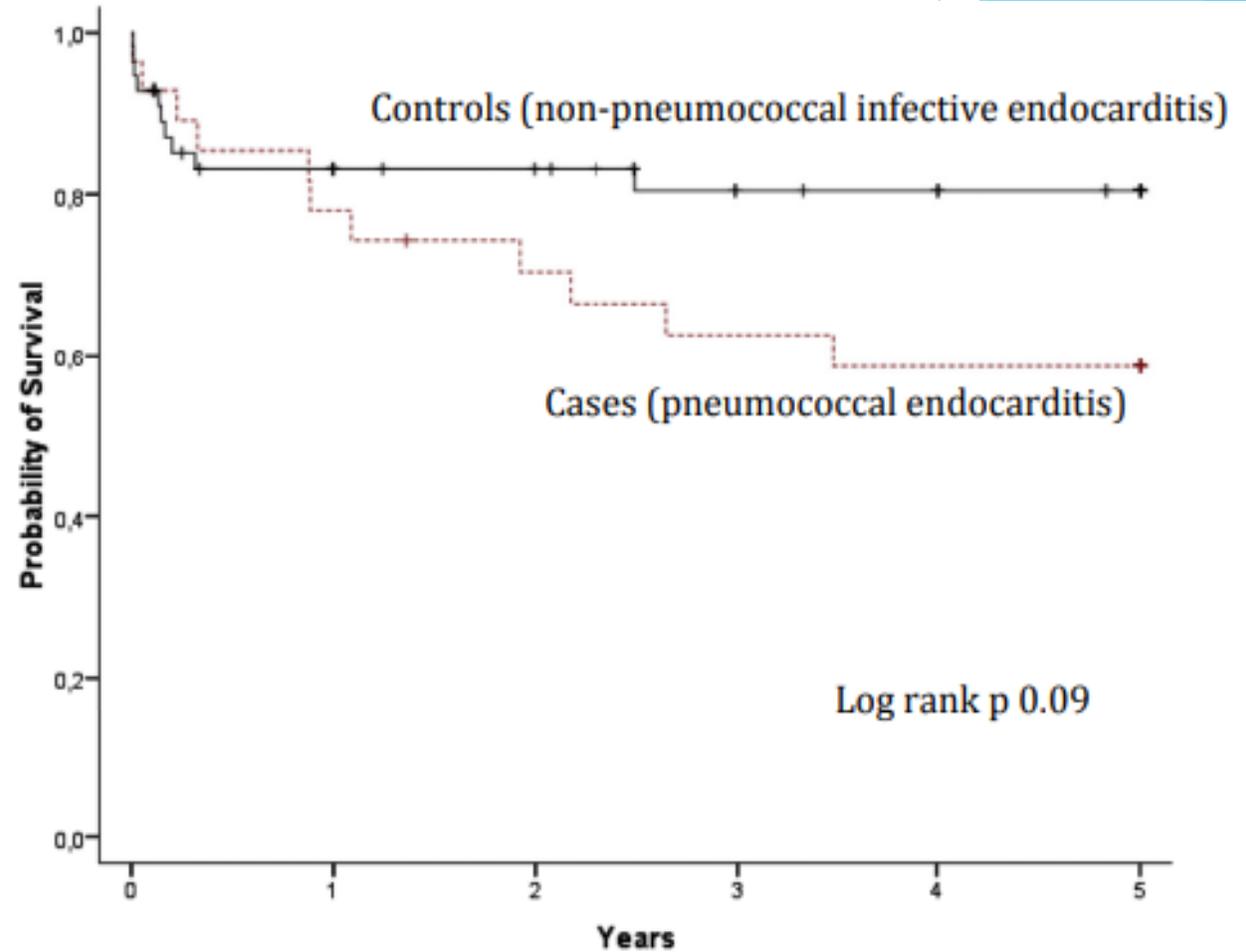


FIG. 1. Kaplan–Meier curve for cumulative survival probability.

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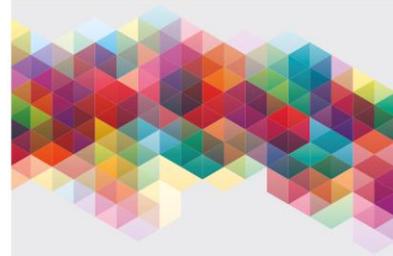
Quelle prévention ?

- ▶ Vaccination anti pneumococcique ?
- ▶ Si FDR d'infection invasive à pneumocoque

b) Patients non immunodéprimés porteurs d'une maladie sous-jacente prédisposant à la survenue d'Infection Invasive à Pneumocoque (IIP) :

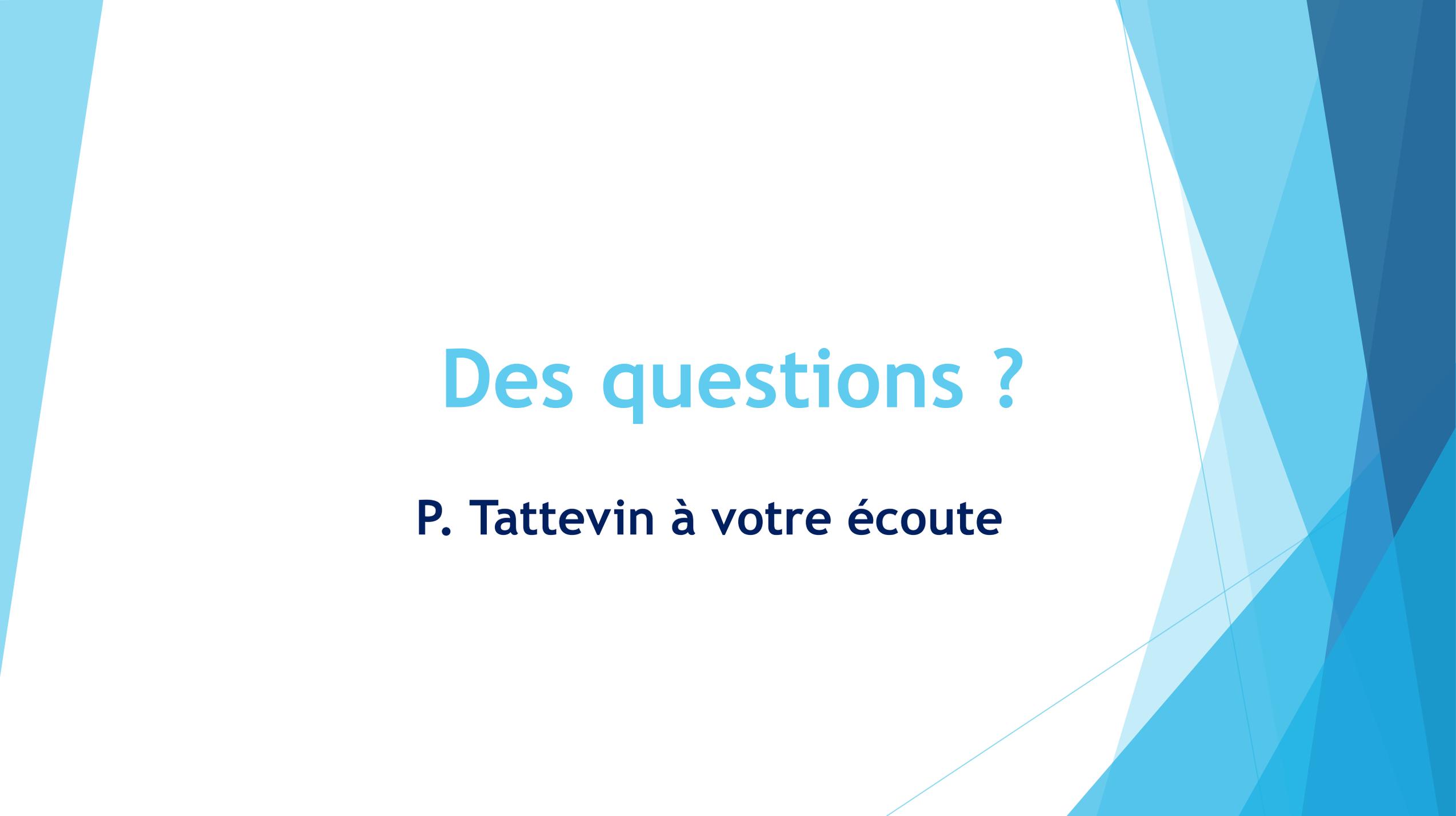
- Cardiopathie congénitale cyanogène, insuffisance cardiaque ;
- Insuffisance respiratoire chronique, bronchopneumopathie obstructive, emphysème ;
- Asthme sévère sous traitement continu ;
- Insuffisance rénale ;
- Hépatopathie chronique d'origine alcoolique ou non ;
- Diabète non équilibré par le simple régime ;
- Patients présentant une brèche ostéo-méningée, un implant cochléaire ou candidats à une implantation cochléaire.

- ▶ Pas au décours d'une IIP en l'absence de FDR



Conclusion

- ▶ Rare mais ça existe
- ▶ Grave
- ▶ Peu FDR EI
- ▶ Facteurs de virulence spécifiques des *S. pneumoniae* ?
- ▶ Relais oral ?

The background features abstract, overlapping geometric shapes in various shades of blue, ranging from light sky blue to deep navy blue. The shapes are primarily triangles and polygons, creating a dynamic, layered effect. The central area is white, providing a clean space for the text.

Des questions ?

P. Tattevin à votre écoute