

Epidémiologie de l'endocardite infectieuse

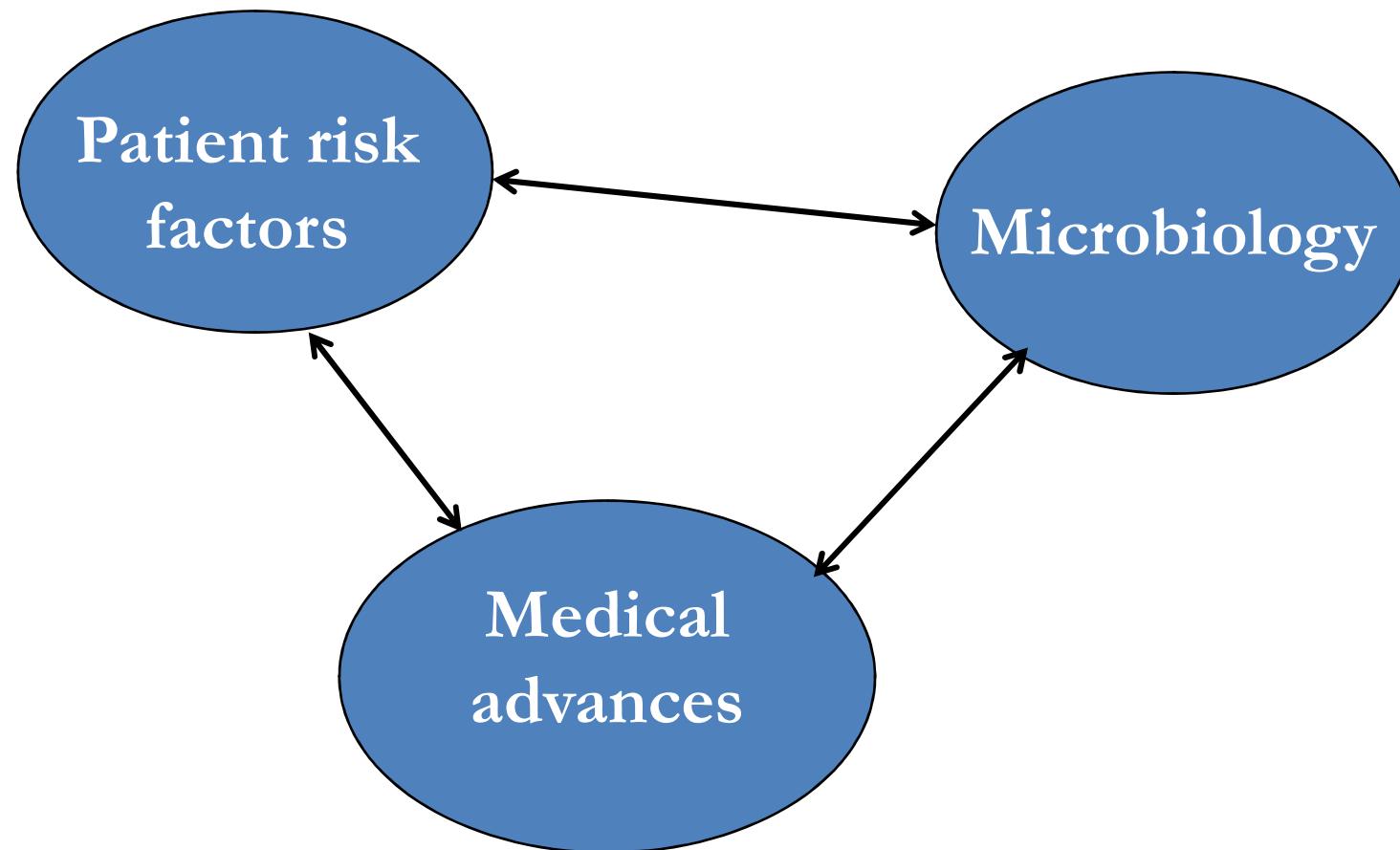
Pr AMEL OMEZZINE LETAIEF
CHU Farhat Hached Sousse

cours de collège infectiologie Sousse le27/02/2013

ENDOCARDITIS

- There is a change in epidemiology
- Clinical features of IE remains classical
- Diagnosis of IE is difficult to confirm
- IE Treatment has been optimized
- IE Prophylaxis will have no/exceptional Indications

Changing Epidemiology of Endocarditis



Clinical Presentation, Etiology, and Outcome of Infective Endocarditis in the 21st Century

The International Collaboration on Endocarditis—Prospective Cohort Study

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for the International Collaboration on Endocarditis—Prospective Cohort Study (ICE-PCS) Investigators

Background: We sought to provide a contemporary picture of the presentation, etiology, and outcome of infective endocarditis (IE) in a large patient cohort from multiple locations worldwide.

Methods: Prospective cohort study of 2781 adults with definite IE who were admitted to 58 hospitals in 25 countries from June 1, 2000, through September 1, 2005.

Results: The median age of the cohort was 57.9 (interquartile range, 43.2-71.8) years, and 72.1% had native valve IE. Most patients (77.0%) presented early in the disease (<30 days) with few of the classic clinical hallmarks of IE. Recent health care exposure was found in one-quarter of patients. *Staphylococcus aureus* was the most common pathogen (31.2%). The mitral (41.1%) and aortic (37.6%) valves were infected most commonly. The following complications were common: stroke (16.9%), embolization other than stroke (22.6%), heart failure

(32.3%), and intracardiac abscess (14.4%). Surgical therapy was common (48.2%), and in-hospital mortality remained high (17.7%). Prosthetic valve involvement (odds ratio, 1.47; 95% confidence interval, 1.13-1.90), increasing age (1.30; 1.17-1.46 per 10-year interval), pulmonary edema (1.79; 1.39-2.30), *S aureus* infection (1.54; 1.14-2.08), coagulase-negative staphylococcal infection (1.50; 1.07-2.10), mitral valve vegetation (1.34; 1.06-1.68), and paravalvular complications (2.25; 1.64-3.09) were associated with an increased risk of in-hospital death, whereas viridans streptococcal infection (0.52; 0.33-0.81) and surgery (0.61; 0.44-0.83) were associated with a decreased risk.

Conclusions: In the early 21st century, IE is more often an acute disease, characterized by a high rate of *S aureus* infection. Mortality remains relatively high.

Arch Intern Med. 2009;169(5):463-473

What is the Burden of Disease?

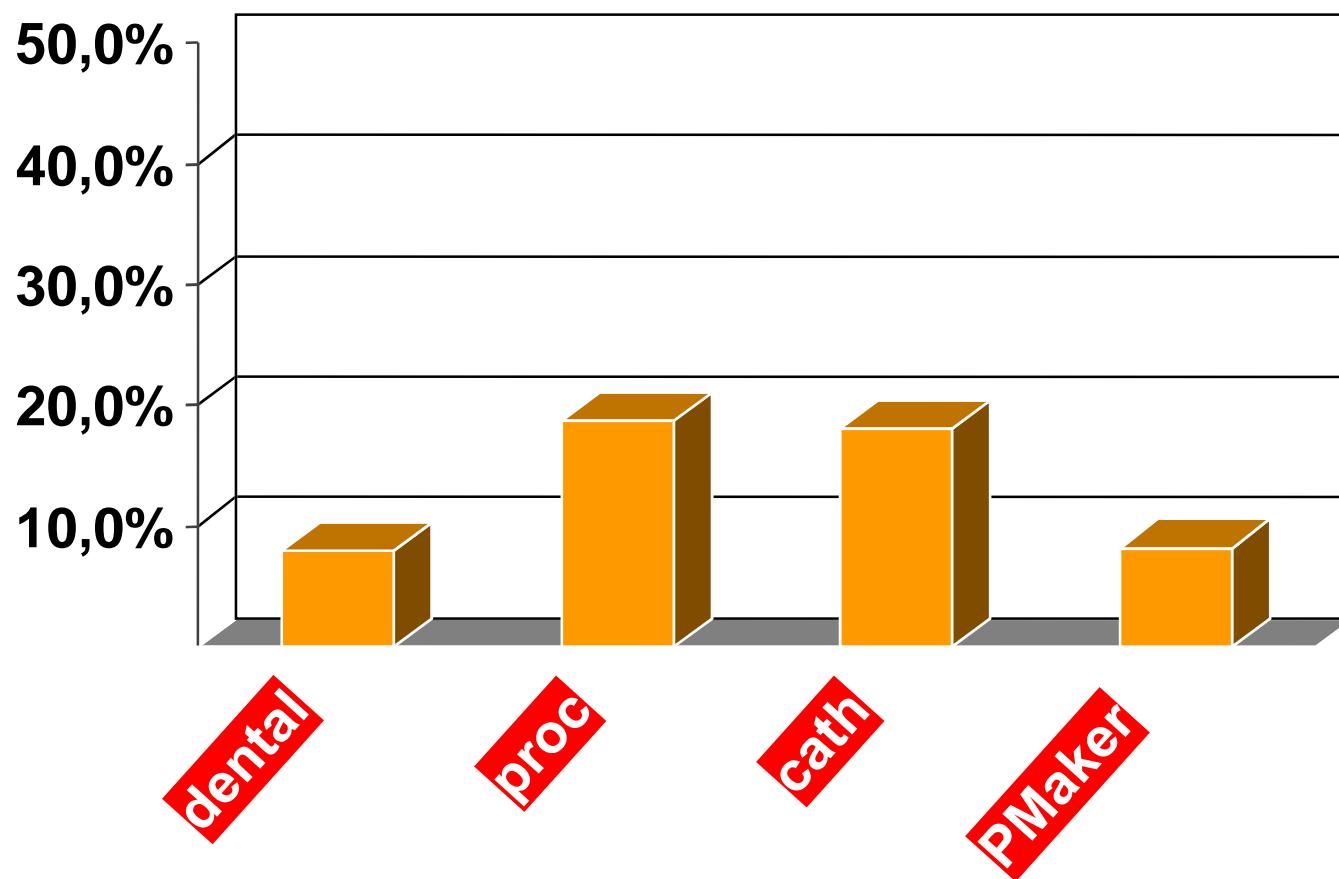
- Usual incidence 2-7 cases/100,000 person-years
 - Diagnostic criteria and reporting variable
 - Only 20% of clinically diagnosed cases definite IE
- 10-20,000 new cases/year in US
- No significant change in overall incidence last 30 years

Table 2. Baseline Characteristics and Predisposing Conditions in 2781 Patients With Definite Endocarditis^a

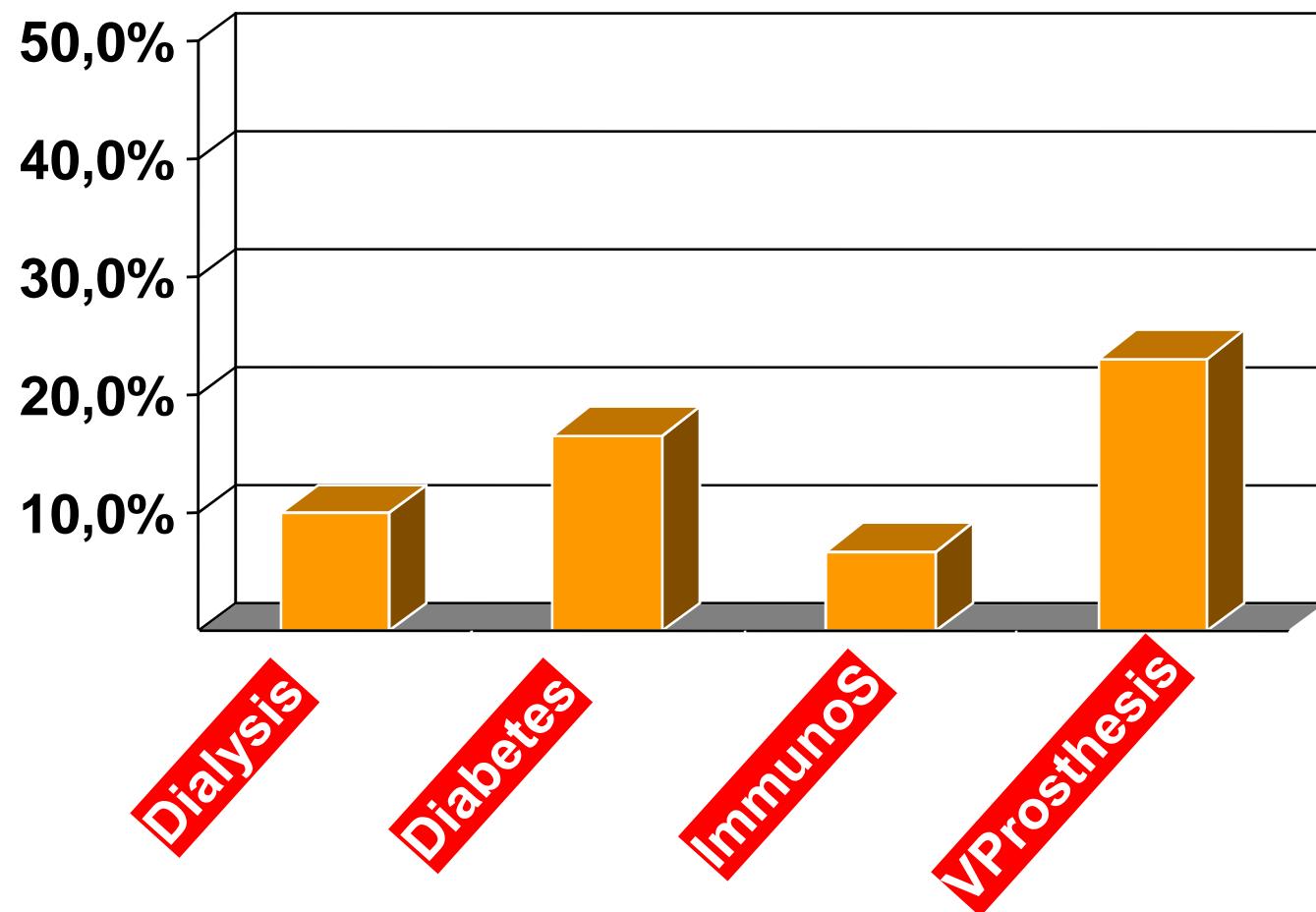
	Total Cohort	Patients Admitted Directly to Study Sites Only ^b	Region				<i>P</i> Value for the Difference in Regions
			North America	South America	Europe	Other	
Baseline characteristics							
Age, median (IQR), y	57.9 (43.2-71.8)	59.8 (44.2-73.1)	52.9 (44.1-66.4)	56.8 (40.3-70.4)	61.4 (45.1-72.7)	58.0 (40.5-72.9)	<.001
Male	1889/2777 (68)	1045/1556 (67)	388/596 (65)	179/254 (70)	873/1212 (72)	449/715 (63)	<.001
First sign to admission <1 mo	2088/2711 (77)	1201/1529 (79)	496/582 (85)	166/244 (68)	896/1174 (76)	530/711 (75)	<.001
Hemodialysis	220/2777 (8)	130/1556 (8)	124/596 (21)	20/254 (8)	49/1210 (4)	27/717 (4)	<.001
Diabetes mellitus	44/7264 (16)	261/1550 (17)	158/592 (27)	25/253 (10)	169/1207 (14)	95/712 (13)	<.001
HIV positive	58/2748 (2)	41/1540 (3)	16/594 (3)	4/236 (2)	33/1211 (3)	5/707 (0.7)	.02
Cancer	230/2772 (8)	160/1553 (10)	52/596 (9)	15/251 (6)	101/1210 (8)	62/715 (9)	.56
IE type							.05
Native valve	1901/2636 (72)	1048/1471 (71)	411/573 (72)	167/246 (68)	860/1166 (74)	463/651 (71)	
Prosthetic valve	563/2636 (21)	321/1471 (22)	116/573 (20)	66/246 (27)	227/1166 (20)	154/651 (24)	
Pacemaker/ICD	172/2636 (7)	102/1471 (7)	46/573 (8)	13/246 (5)	79/1166 (7)	34/651 (5)	
Predisposing conditions							
Current IV drug use	268/2746 (10)	157/1540 (10)	93/587 (16)	1/249 (0.4)	113/1203 (9)	61/707 (9)	<.001
Previous IE	222/2780 (8)	138/1557 (9)	66/596 (11)	26/254 (10)	84/1213 (7)	46/717 (6)	.003
Invasive procedure within 60 d	690/2581 (27)	392/1463 (27)	162/508 (32)	64/247 (26)	289/1145 (25)	175/681 (26)	.03
Chronic IV access	244/2763 (9)	142/1548 (9)	148/595 (25)	12/251 (5)	56/1205 (5)	28/712 (4)	<.001
Endovascular device							
Pacemaker	262/2752 (10)	146/1540 (9)	55/595 (9)	23/252 (9)	137/1191 (12)	47/714 (7)	.005
ICD	27/2720 (1)	15/1521 (1)	16/593 (3)	0/249 (0)	8/1172 (0.7)	3/706 (0.4)	<.001
Congenital heart disease	311/2656 (12)	167/1481 (11)	62/582 (11)	53/244 (22)	111/1156 (10)	85/674 (13)	<.001
Native valve predisposition	884/2761 (32)	538/1547 (35)	147/596 (25)	93/252 (37)	370/1201 (31)	274/712 (38)	<.001

Only 3.3% had rheumatic valve disease

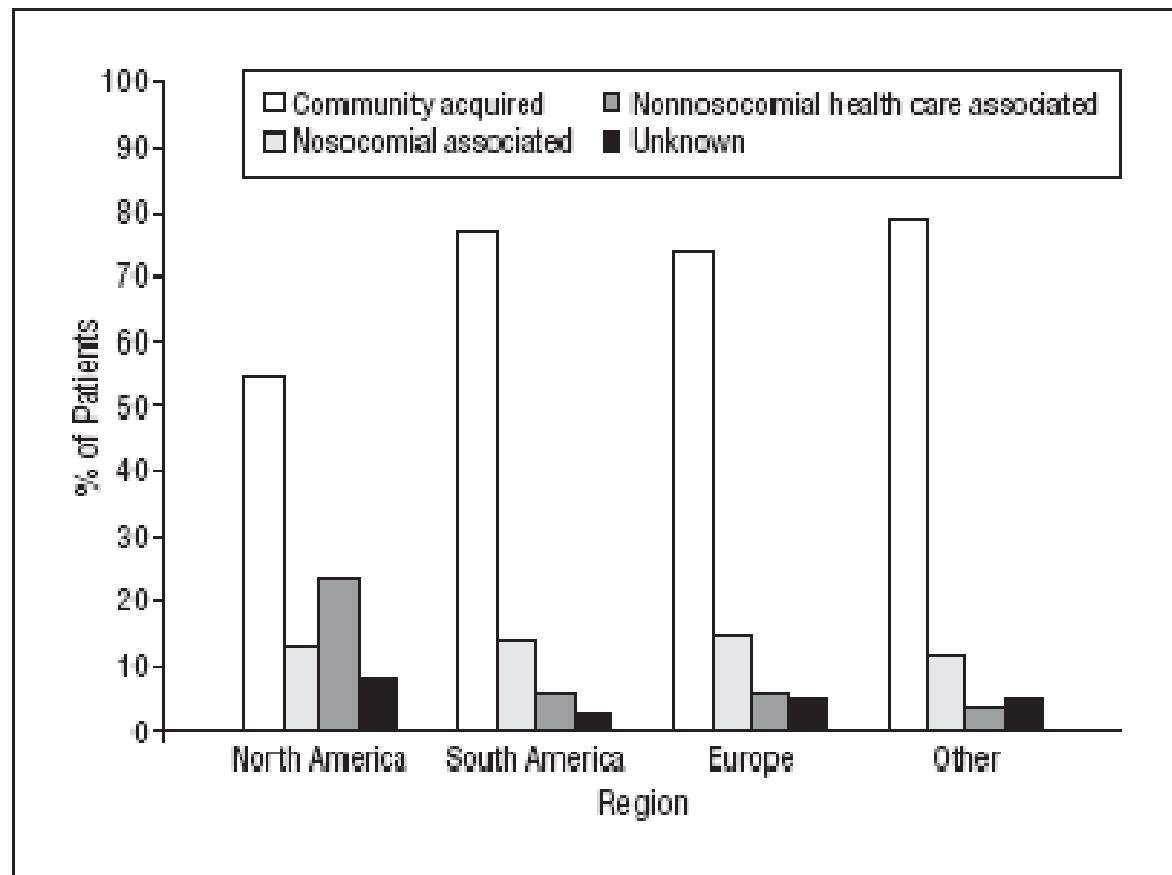
Exposures - Now: ICE/ PCS



Host Factors / Exposures - ICE PCS



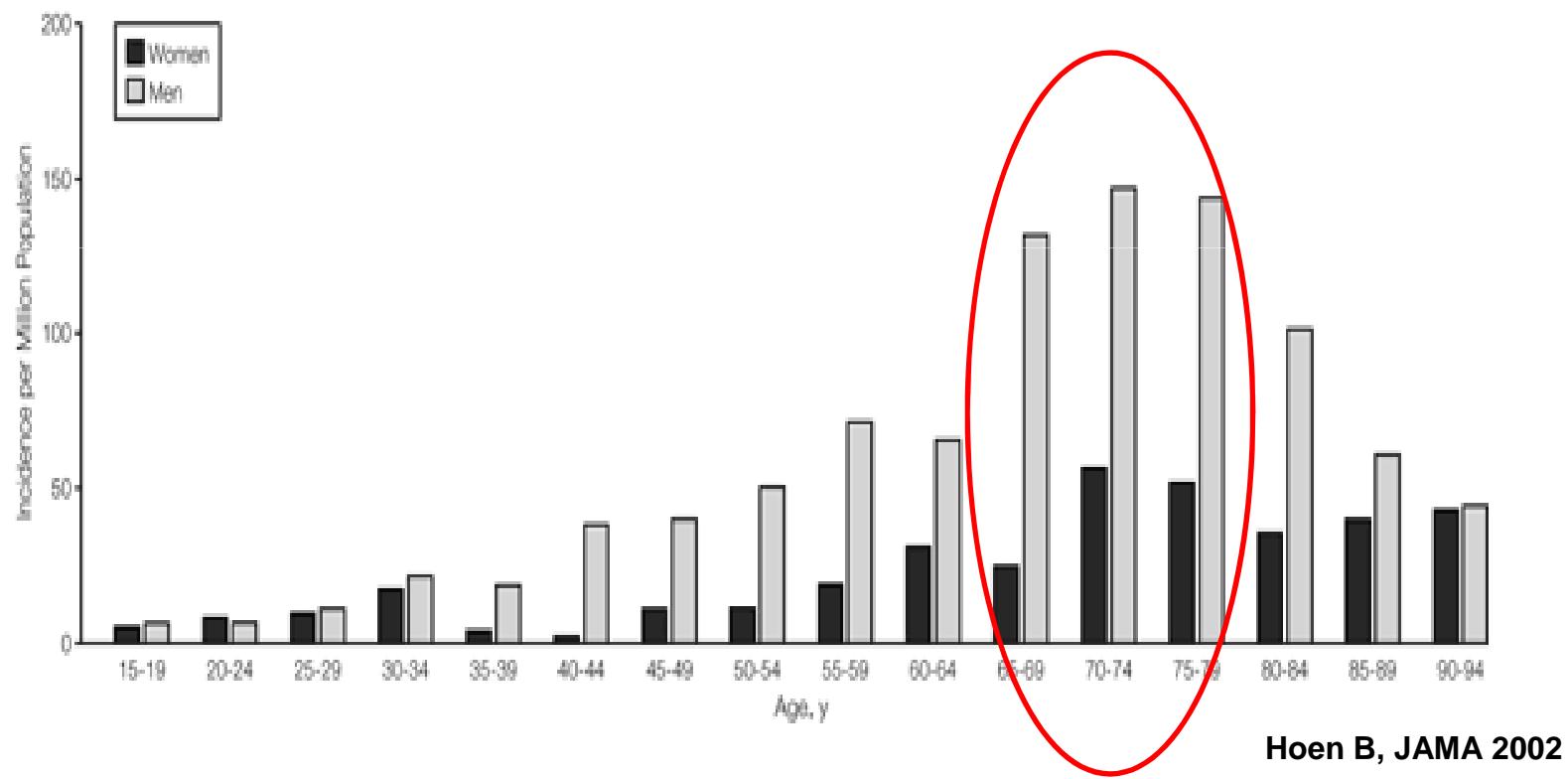
Health-Care Associated IE



Distribution of clinical and laboratory findings of 2781 patients with definite endocarditis by location of acquisition of 2781 patients with definite endocarditis

Murdoch. Arch Intern Med. 2009;169

Incidence of Infective Endocarditis by Age and Sex in the Study Population.



Hoen et al, JAMA 2002;288:75-81

Aspects épidémiologiques

Étude multicentrique en 1999 par l'AEPEI

6 régions françaises: 26% de la population

16 millions d'habitants

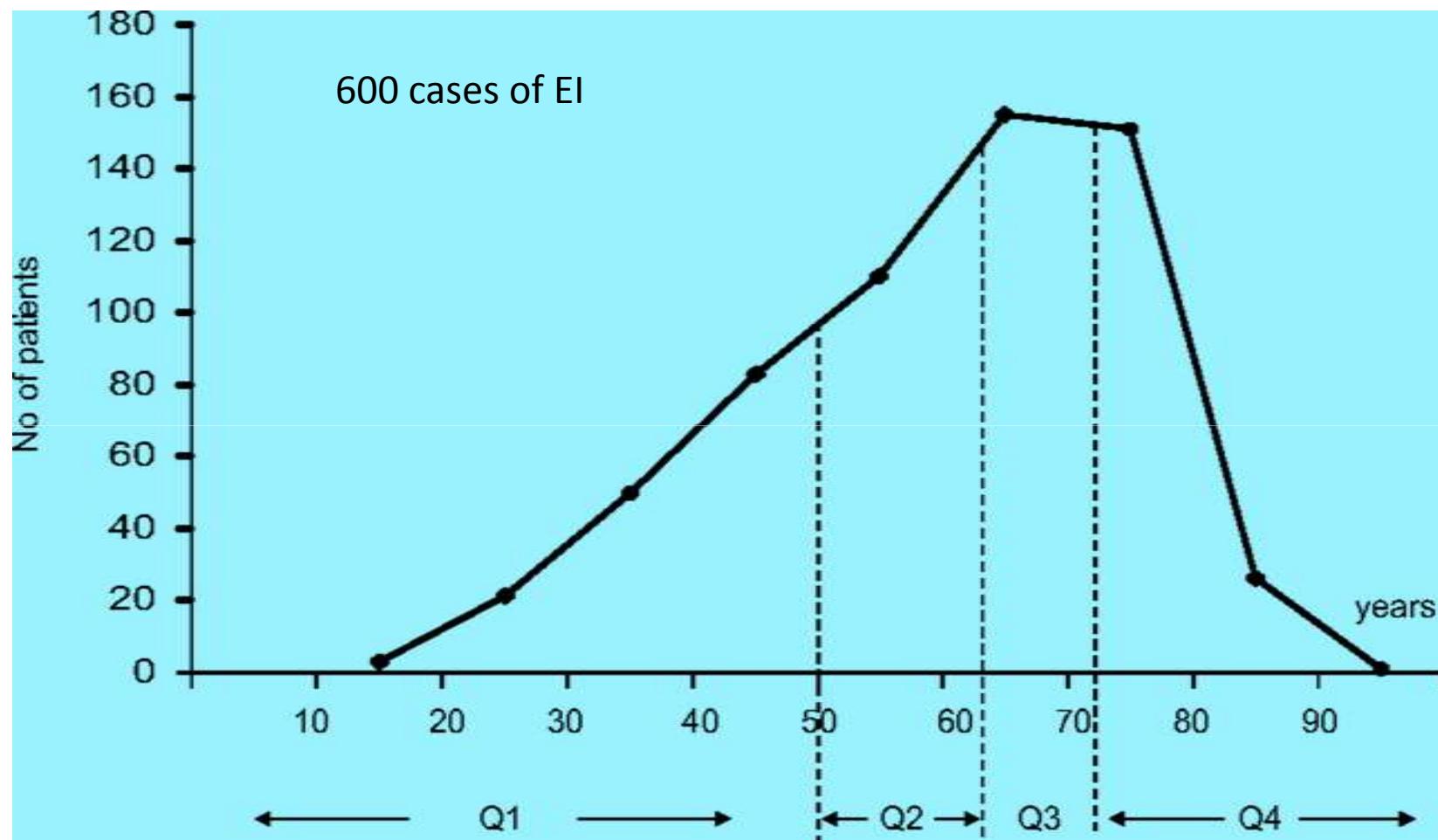
817 pts avec endocardite présumée

390 endocardites certaines selon la classification de Duke

277 hommes/113 femmes: **59.5 ± 17.2 ans**

Incidence: **31 cas/million d'habitants**

Figure 1. Patient distribution by age-specific quartiles (Q).



López J et al. Circulation 2010;121:892-897

American Heart Association 
Learn and Live

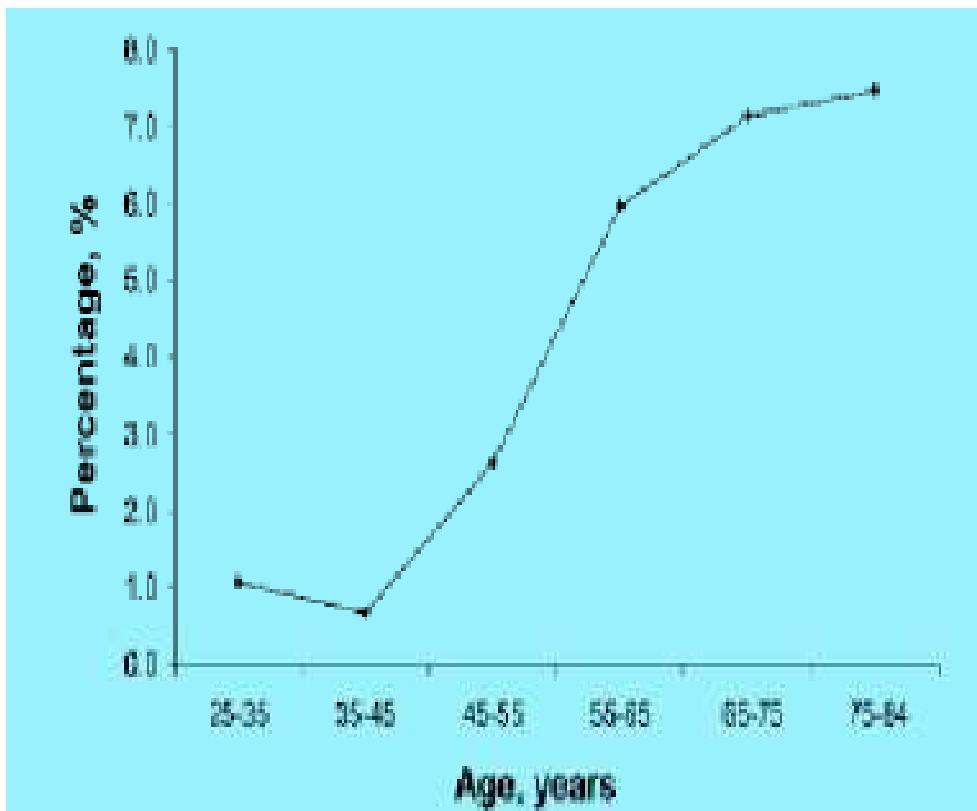


Figure 1. Age-specific prevalence of a predisposing cardiac condition among French adults (age, 25–84 years).

X. Duval Clin Infect Dis 2006 15;42:102-7

EPIDEMIOLOGY · Multicentric study France 1999

Table 1. Distribution of Underlying Heart Disease in Patients (N = 390) With Infective Endocarditis

	No. (%)
Native valve disease*	119 (31)
Left heart	112 (91)
Right heart	6 (5)
Both sides	1 (1)
Prosthetic valve	63 (16)
Congenital heart disease	4 (1)
Unspecified heart murmur	19 (5)
No previously known underlying heart disease	185 (47)

Hoen et al, JAMA 2002;288:75-81

Infective Endocarditis

Temporal Trends in Infective Endocarditis in the Context of Prophylaxis Guideline Modifications

Three Successive Population-Based Surveys

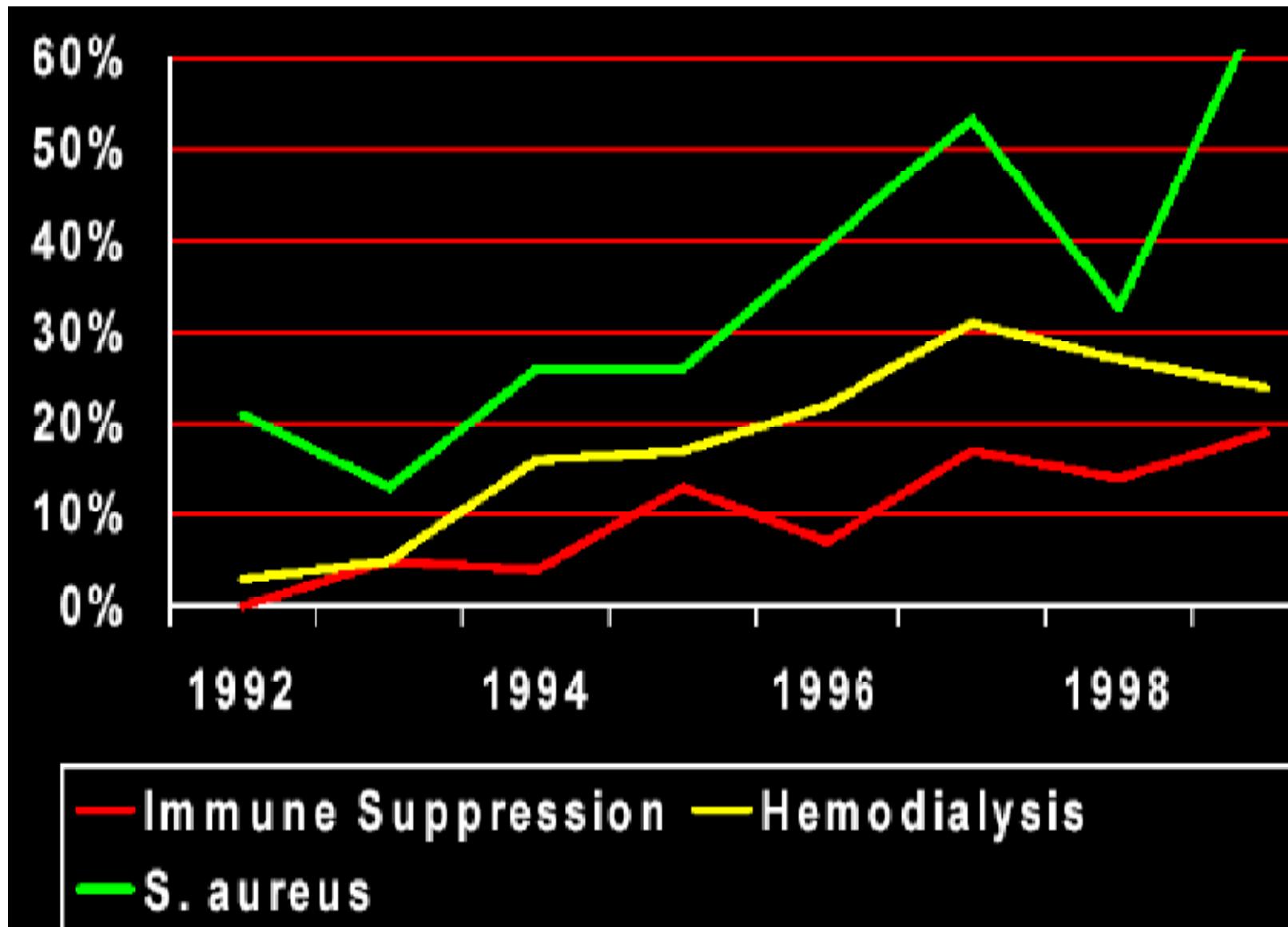
Xavier Duval, MD, PhD,*†‡ François Delahaye, MD, PhD,§|| François Alla, MD, PhD,¶#

EPIDEMIOLOGY : Multicentric study France 1999

Table 2. Location of Infective Endocarditis (IE) in Patients (N = 390)

Location	No. (%)
Aortic valve	136 (35)
Mitral valve	112 (29)
Aortic and mitral valves	55 (14)
Tricuspid valve	37 (9)
Pulmonic valve	2 (1)
Bilateral IE	6 (2)
Pacemaker	18 (5)
With left-sided IE	5 (1)
Without left-sided IE	13 (4)
Undetermined*	24 (6)

Hoën et al, JAMA 2002;288:75-81



Duke University Medical Center, Durham, NC, from 1993 to 1999

Cabell, Arch Internal Med 2002

Clinical /Laboratory findings in 2781 Patients with Definite IE

Findings	No (%) of pts
Fever, temperature 38°C	2322/2428 (96)
Splinter hemorrhages	213/2655 (8)
Osler nodes	77/2648 (3)
Janeway lesions	123/2650 (5)
Roth spots	50/2649 (2)
Vascular embolic event	456/2665 (17)
Conjunctival hemorrhage	122/2655 (5)
Splenomegaly	284/2662 (11)
New murmur	1068/2232 (48)
Worsening of old murmur	359/1787 (20)
Elevated ESR	1611/2645 (61)
Elevated C-reactive protein level	1632/2650 (62)
Elevated rheumatoid factor	138/2549 (5)
Hematuria	666/2587 (26)

Clinical presentation/complications

Changing +++

Cardiac complications 30-50%

- Valvular regurgitation Heart failure+++
- Perivalvular abscesses
(30-40% at surgery/autopsy)

Omari, B, Chest 1998

Neurologic complications 30%

Renal complications

Vertebral osteomyelitis

...

Microbiologic etiology of IE

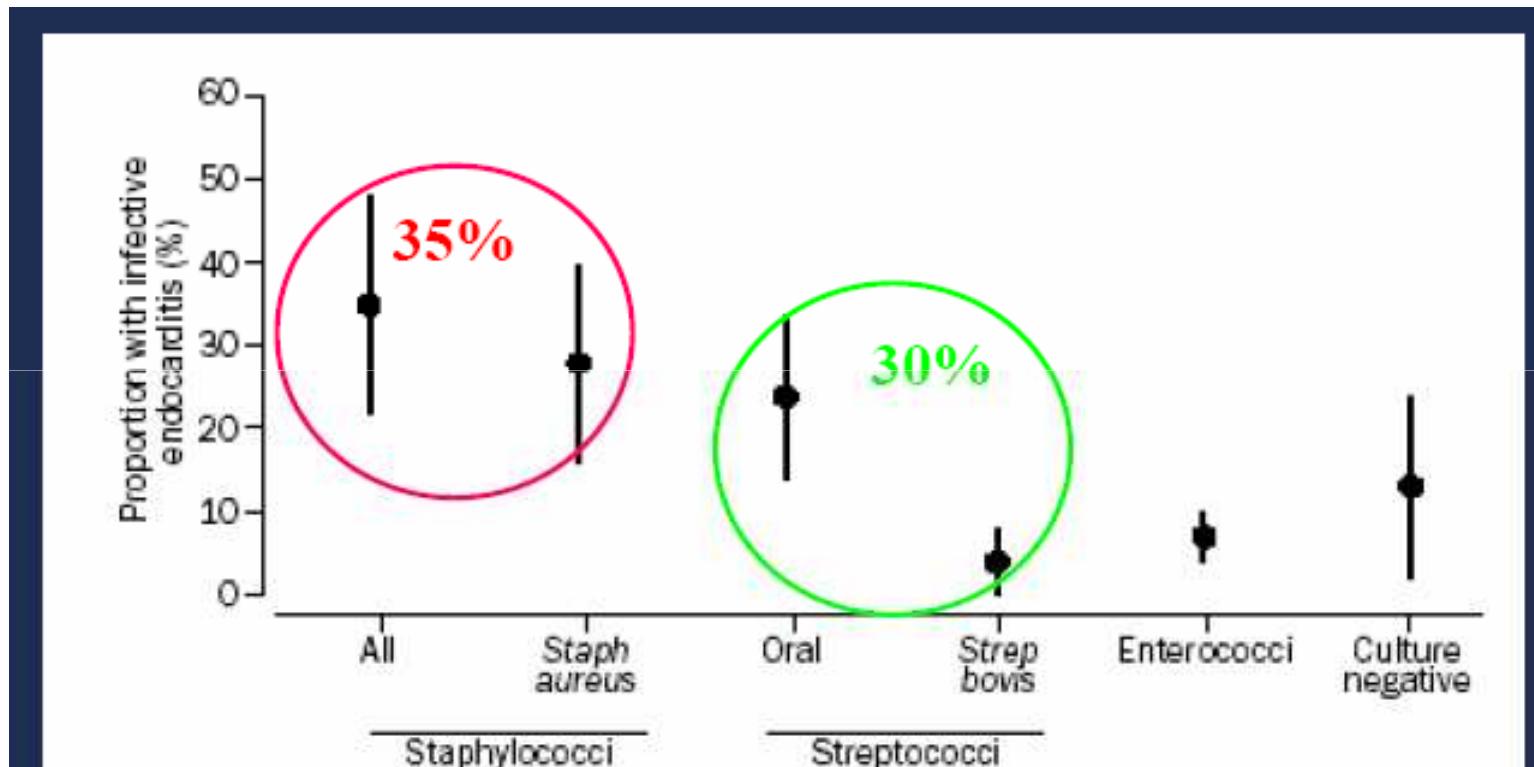
Cause of Endocarditis	No. (%) of Patients ^a						P Value for the Difference Between Regions	
	Total Cohort (N=2781)	Patients Admitted Directly to Study Sites Only ^b (n=1558)	Region					
			North America (n=597)	South America (n=254)	Europe (n=1213)	Other (n=717)		
<i>Staphylococcus aureus</i>	869 (31)	487 (31)	256 (43)	43 (17)	339 (28)	231 (32)	<.001	
Coagulase-negative staphylococcus	304 (11)	161 (10)	69 (12)	18 (7)	156 (13)	61 (9)	.005	
Viridans group streptococci	483 (17)	288 (19)	54 (9)	66 (26)	198 (16)	165 (23)	<.001	
<i>Streptococcus bovis</i>	165 (6)	101 (7)	9 (2)	17 (7)	116 (10)	23 (3)	<.001	
Other streptococci	162 (6)	101 (7)	38 (6)	16 (6)	66 (5)	42 (6)	.86	
<i>Enterococcus</i> species	283 (10)	158 (10)	78 (13)	21 (8)	111 (9)	73 (10)	.05	
HACEK	44 (2)	26 (2)	2 (0.3)	6 (2)	19 (2)	17 (2)	.02	
Fungi/yeast	45 (2)	25 (2)	20 (3)	3 (1)	13 (1)	9 (1)	.002	
Polymicrobial	28 (1)	23 (2)	8 (1)	1 (0.4)	13 (1)	6 (0.8)	.60	
Negative culture findings	277 (10)	122 (8)	41 (7)	51 (20)	123 (10)	62 (9)	<.001	
Other	121 (4)	66 (4)	22 (4)	12 (5)	59 (5)	28 (4)	.61	

Murdoch. Arch Intern Med. 2009;169.

Changing Microbiology of IE

- *Staphylococcus aureus* now the most common cause worldwide, 31% of patients
- Other gram positives important
 - Viridans strep, coagulase-negative staph, *Enterococcus*
- Fastidious organisms
 - HACEK 2% (0.3% in N. America)
 - *Haemophilus, Aggregatibacter, Cardiobacterium, Eikenella, Kingella*
 - Fungi/yeast 2%
- Culture negative 10%

Microorganism: Meta-analysis 3784 cases Endocarditis



Moreillon et al Lancet 2004

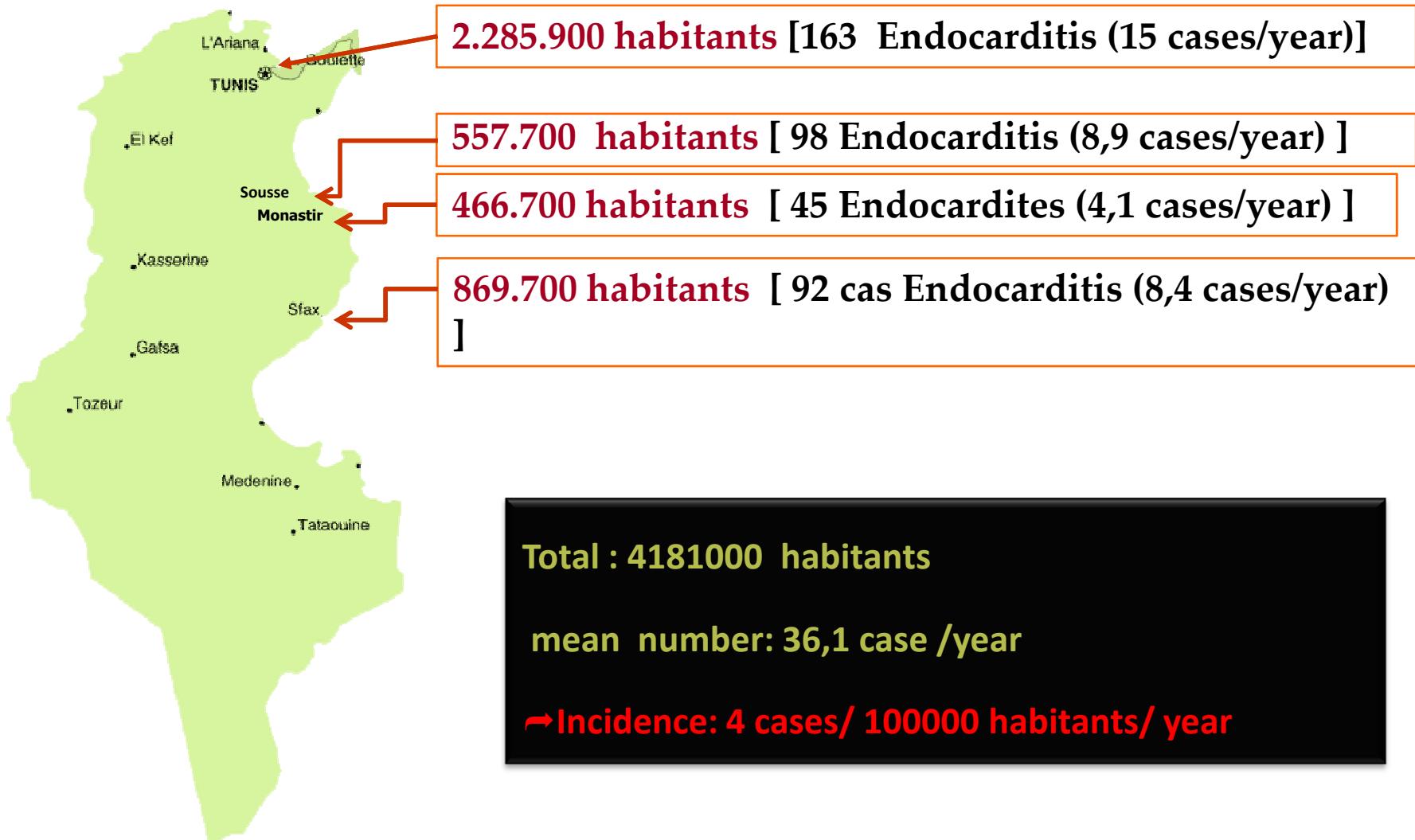
Table 5. Surgery and Mortality During the Initial Hospital Stay as a Function of Different Variables in Patients (N = 390) With Infective Endocarditis*

Variable	Total No. (%)	Surgery		Mortality	
		No. (%)	P Value	No. (%)	P Value
All	390 (100)	191 (49)	...	62 (16)	...
Sex					
Women	113 (29)	42 (37)	.03	20 (18)	.53
Men	277 (71)	150 (54)		42 (15)	
Location					
Only mitral valve	112 (29)	52 (46)		20 (18)	
Only aortic valve	136 (35)	82 (60)		22 (16)	
Aortic and mitral	55 (14)	40 (73)	<.001	11 (20)	
Right-sided or bilateral	45 (12)	14 (31)		4 (9)	.67
Pacemaker	18 (5)	5 (28)		2 (11)	
Unknown	24 (6)	0		3 (13)	
Previous heart disease					
Native valve disease	119 (31)	67 (56)		14 (12)	
Prosthetic valve	63 (16)	29 (46)	.29	15 (24)	
Miscellaneous	23 (6)	11 (48)		3 (13)	
No known heart disease	185 (47)	83 (45)		30 (16)	.20
Microorganisms					
Streptococci	196 (50)	106 (54)		22 (11)	
Enterococci	29 (7)	15 (52)		5 (17)	
Staphylococci	115 (29)	43 (37)	.02	29 (25)	.02
Others or ≥2	31 (8)	20 (63)		5 (17)	
No microorganism	19 (5)	9 (45)		2 (10)	
Valve surgery					
Yes	191 (49)	...		11%	
No	199 (51)	...		20%	.02

-2 Etudes rétrospectives multicentriques

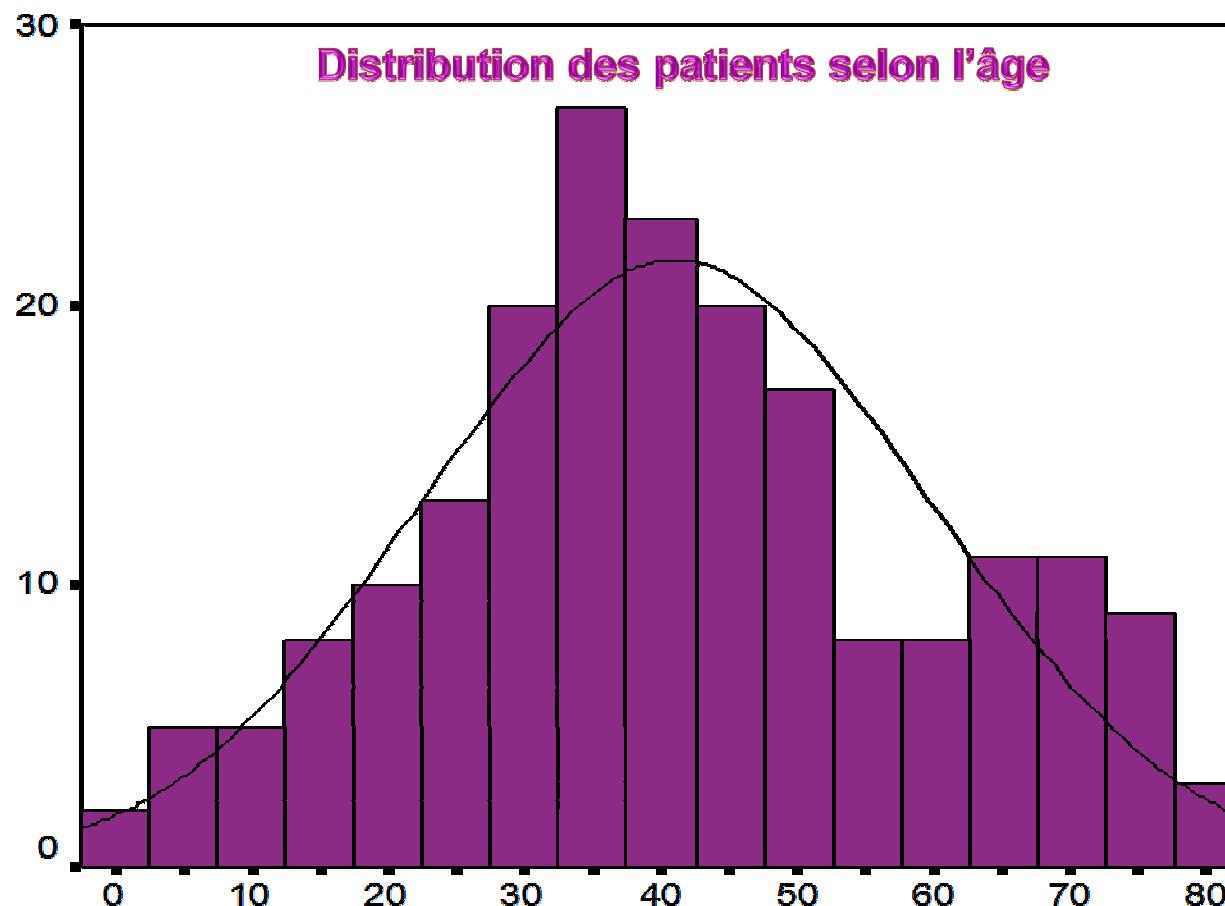
- 1991 - 2000
- 2001 to 2011

Results



Epidemiological characteristics

- Age



30% : <30 ans

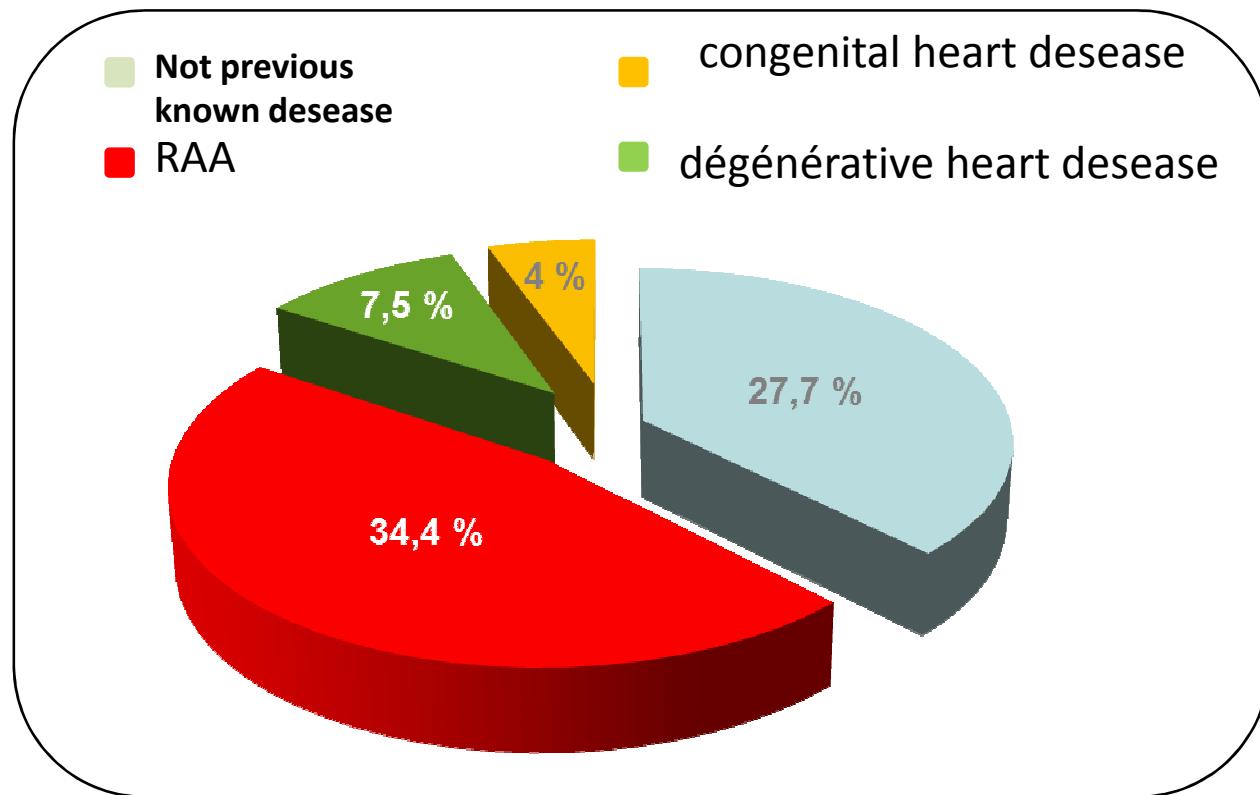
Mean age : $42 \pm 18,44$

Results

Epidemiological Characteristics

- Sex-ratio : 234/164 (1.42)
- native valve endocarditis : 292 (73,36 %)
- prosthetic valve endocarditis : 92 (23%)
 - Early Endocarditis: 24 cas (26 %)
 - Late Endocarditis : 69 cas (74.19 %)
- pace maker Endocarditis : 16 (4 %)

- **Underlying Heart disease : 277 (72,3%)**



- **Les facteurs de co-morbidité**

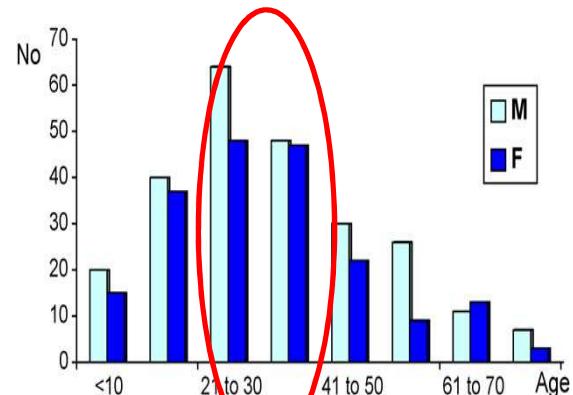
- Diabetes: 33 cas (8.3%)
- IRC hémodialysis : 12 cas (3%)
- Toxicomania : 5 cas (1.3%)



Epidemiology of infective endocarditis in Tunisia: a 10-year multicenter retrospective study

Letaief A, Int J of Inf Dis 2007

mean (SD) age 32.4 (16.8) years



Incidence of IE by age and sex
in 450 patients.

Table 1 Distribution of underlying heart disease in 440 cases of infective endocarditis

Underlying heart disease	No. (%)
Native valve disease	364 (82.7)
Rheumatic valve disease	199 (45.2)
Not previously known	100 (22.7)
Congenital heart disease	39
Degenerative	24
Other	2
Prosthetic valve	76 (17.3)

*Epidemiology of infective endocarditis in Tunisia:
a 10-year multicenter retrospective study 1991-2000*

1991-2000

2000/2011

Table 1 Distribution of underlying heart disease in 440 cases of infective endocarditis

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Degenerative	24
Other	2
Prosthetic valve	76 (17.3)

Native valve
73,26%
RAA 34,4%

Prosthetic valve
23%

Results

clinical-epidemiological features

	Europe 2001 N=159	France 1999 N= 390	Tunisia 1991-2000 N=440	Tunisia 2001-2011 N=398
mean age	57	59.5	32.4	42,07
Sex ratio	-	2	1.25	1.42
prosthetic valve EI	41 (26%)	63 (16%)	76 (17.3%)	92 (23 %)
Underlying heart disease	53 (45%)	142 (37%)	264 (60%)	277 (72.3%)
Rheumatic valve disease	-	-	199 (45.2%)	137 (34,4%)
congenital heart disease	-	4 (1%)	39 (8.9%)	40 (10%)
Right heart EI	8 (5%)	39 (10%)	20 (4.5%)	16 (4%)
pace maker EI	4 (2.5%)	18 (5%)	-	16 (4%)

Underling heart desease

			Valve native		Prothèse		
	Année	Nbre	RAA	Dégénér.	Congénit	C.sain*	
B. Ismail-Tunis(1)	66-85	210	142 (67%)		8 (4%)	2 (1%)	58 (27%)
Cetinkaya-Ankara(2)	74-99	228	147 (64%)		11 (5%)	17 (7,5%)	
Delahaye-France(3)	90-91	415		139 (33%)	21 (5%)	141 (34%)	90 (22%)
Série Tunisienne	90-00	440	201 (45,7%)	24 (5,4%)	39 (10%)	100 (22%)	76 (17,3%)
	20 01-2011	398	34,4%		7,5% 4%	23%	27,7%

(1) L'information cardiologique, Dec 1986 ; (2) Cetinkaya, International J Antimicrobial Agents 18 (2001):1-7 ;

(3) Delahaye, Eur Heart J 1995; 16 : 394-401 ; (4) Habib Ann. Cardiol. Angéol. 1998,47 (10) : 735-8.

suspected Portal of entry ➤ 129 cases (32,4 %)

	Total (n=398)	VN (n=292)	VP (n=93)	p
Dental procedure	96 (27%)	81 (30.2%)	15 (17.2%)	0.5
Endoscopy	1 (0.3%)	—	—	—
Cardiac Chirurgy	24 (6.4%)	5 (2%)	19 (23%)	0.001
Skin	8 (4%)	5 (2.5%)	3 (1.5%)	NS
Other	11 (5.4%)			

Results

Clinical characteristics

	Total (n=398)	VN (n=292)	VP (n=93)	p
Delay consultation(day)	23.8	27.54	8.37	0.003
Fever>38°	334 (83.9%)	133 (83.6%)	40 (95.2%)	0.05
Heart murmur	249 (62,6%)	210 (59,5%)	36 (44,4%)	0.001
Heart fealure	28 (14.4%)	25 (16.4%)	3 (7.3%)	NS
Cerebral emboli	32 (16%)	25 (16%)	7 (16%)	0.07
Péphérique emboli	9 (4.4%)	5 (3.1%)	4 (9.5%)	
Secondary localisations	20 (10%)	18 (11.3%)	2 (4.8%)	0.18
Purpura	3 (1.5%)	3 (1.9%)	0	0.37
lesions of Janeway	4 (2%)	4 (2.5%)	0	0.29
Osler Nodes	2 (1%)	1 (0.6%)	1 (2.4%)	0.3
Mycotic aneurysms	1(0.5%)	1 (0.6%)	0	0.6

Les données échocardiographiques

- ETT : 398 cas (100%) / ETO : 314 cas (82.8%)
- Localisation of E I:
 - Mitral valve :144(36,2%)
 - Aortic valve :134 (33,7)
 - Tricuspid valve :16 (4%)
 - Pulmonary valve :10 (2,5%)
 - Mitro-aortic :26(6,5%)
 - Other (pace maker) : 16(4%)
- Prothèse : 92 (23%)

microbiological Data

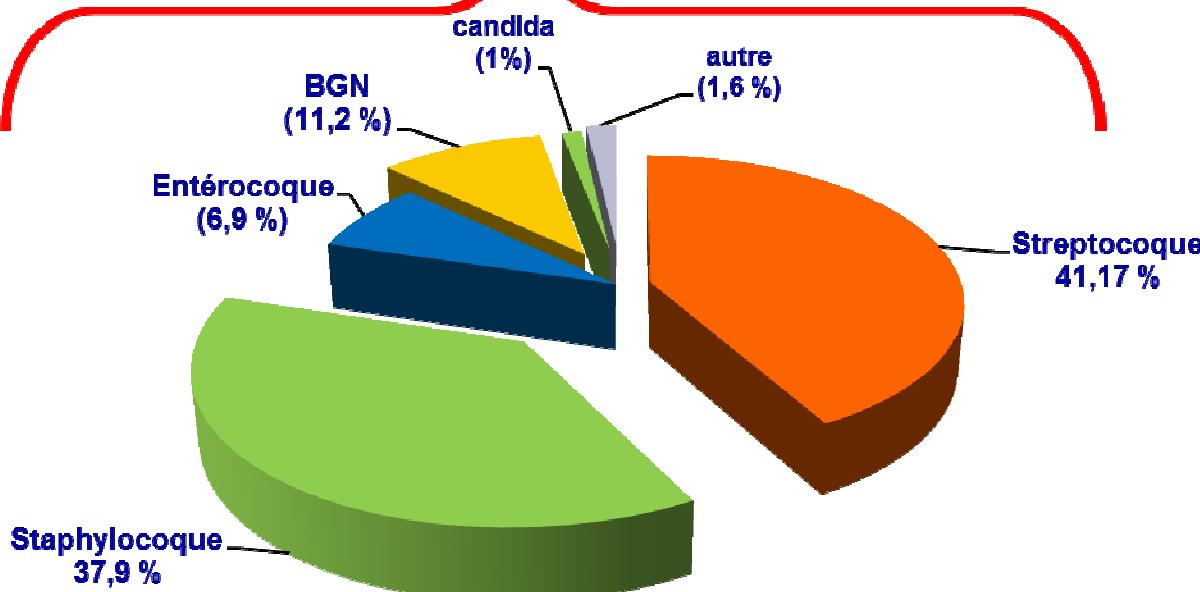
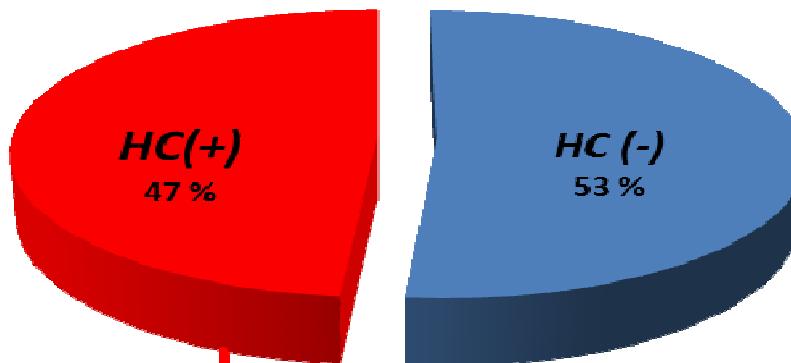
Blood culture HC positives : 172 cas 47%

	Total n=398	HC(+) n = 187	HC(-) n = 211	p
Previous Antibiotherapy	97 (24,4%)	38 (20,3 %)	59 (27,9 %)	0.32
Mean Number blood culture	5.9	5,96	5.65	0.4
Endocardite probable	134 (33,6 %)	29 (15,5 %)	102 (48,3%)	-
Endocardite certaine	184(46,23 %)	129 (68,9%)	55 (26%)	-

La sérologie

	sérologies F 81 (40%)	sérologie + 16 (4%)
<i>Chlamydia</i>	88 (20%)	0
<i>Brucella</i>	103 (25,9%)	3 (1.5%)
<i>Coxiella burnetii</i>	88 (22%)	1 (0.5%)
<i>Bartonella</i>	51 (12,8%)	2 (1%)

bacteriological Profile



186 patients Surgical therapy



Valve culture

60 (32%)

Positive

13 (21%)

- ↳ *A. Baumanii* : 1
- ↳ *B. quintana* : 1
- ↳ *E. feacalis* : 2
- ↳ *S. adjacens* : 1
- ↳ *Kytococcus* : 1
- ↳ *Lactobacillus* : 1
- ↳ SCN : 4
- ↳ *S. gordonii* : 1
- ↳ *H. parainfluenzae* : 1

Negative

47 (79%)

Distribution of causative microorganisms

microbiological profile of Endocarditis

	Europe 2001 n = 159	France 1999 n = 390	Tunisie 1991-2000 n = 440	Tunisie 2001-2011 n = 398
Streptococci	28%	225 (58%)	76 (17.3%)	77 (19,3%)
Enterococci	14%	29 (8%)	17 (3.9%)	13 (3.3%)
Staphylococci	33%	115 (29%)	79 (18%)	71(17,8%)
Other	10%	21 (8%)	49 (4.4%)	24 (06%)
No microorganism identified	14%	19 (5%)	219 (49.8%)	211 (53%)

Bacteriology

	NG	Streptocoque D+enteroc	Autres	Staphylocoque aureus	CN	Autres	HC (-)
Delahaye (3)	27	23	7	18	4	11	8
Habib (94-96, 71 cas) (4)	16,9	19,7	5,6	18,5	9,8	25*	4,2
B. Ismail (1)		8,5	26,6	11,5	3,3	4,1	46
Cetinkaya (2)	7,5	6,9	6	11,8	4,8	13	50
Série Tunisienne 1990-2000	10,7	4,3	6,1	12	6	11	49,8%
Serie Tunisienne 2001 2011	19,3%			17,3%		53%	

Surgical Therapy

	Total (n=398)	VN (n=292)	VP (92)	p
Surgery	186 (47%)	142 (48%)	44 (47%)	0.03
<u>Indications :</u>				
Hemodynamic	82 (44%)	67 (47%)	15 (34%)	0.001
Annular Abces	28 (15%)	12 (14%)	4 (16%)	0.7
Size of vegetation>10	69 (29%)	55 (38%)	14 (31%)	0.4
Mobility of vegetation	54 (29%)	44 (30%)	23 (11.8%)	0.8
Prosthetic desinsertion	11 (2,8%)			

Results

	Evolution			p
	Total (n=398)	VN (n=292)	VP (n=92)	
Hospital stay means (day)	39 ± 22,16j	40.31	40,31	0.86
Delay of d'apyrexia(day)	6,79 ± 7,7j	6,53j	6,82	0.82
Complications :				
Neurologic	51 (12,8%)	34 (14,8%)	17 (22,4%)	0.12
Secondary Localisations	39 (9,8%)	31 (13,6%)	7 (9,6%)	0.32
Recidive	13 (3.3%)	9 (4.1%)	3 (4,2%)	0.9
Mortality	49 (12,3%)	33 (14%)	21 (16%)	0.08

Results

Evolution and series

	Europe 2001 n=159	France 1999 n= 390	Tunisie 1991-2000 n=440	Tunisie 2001-2011 n=398
Complications :				
Neurologic	24 (15%)	60 (15.3%)	89 (16.3%)	51 (12,8%)
secondary Localisations	-	-	26 (6%)	39 (9,8%)
Surgery	82 (52%)	191 (49%)	223 (50.6%)	186 (47%)
Mortality	20 (12.6%)	62 (16%)	90 (20.6%)	49 (12,3%)

Results

Evolution / microorganism

	Streptocoques	SCN	S. aureus	GNI*
Delay of apyrexia (day)	4.06	6.3	5.75	4.7
Neurological complications	7 (16.7%)	5 (20.8%)	6 (40%)	11 (12.2%)
Localisations secondaires	4 (9.5%)	2 (8.3%)	3 (20%)	7 (7.7%)
Mean Duration of antibiotherapy(day)	33.7	45.2	31.9	37.1
Early Surgery	22 (51.2%)	14 (58.3%)	8 (53.3%)	44 (46.8%)
Récidives	5 (13.2%)	1 (4.2%)	0	2 (2.4%)
Décès	6 (15%)	1 (4.2%)	5 (35.7%)	15 (17%)

*GNI : aucun germe identifié

Conclusion

- ➔ Incidence des EI : **4 cas/100000 habitants/year**
- ➔ Mean age : **42 ans**
- ➔ Underlying heart disease : **RAA 40.4 %**
- ➔ Antibiotic before diagnosis : **31 %**
- ➔ Enquête microbiologic positive (HC/sérology/valve): **55.2 %**
- ➔ Staphylococci + Streptococci : **80%**
- ➔ Surgery : **47 %**
- ➔ Mortality : **12,3%**



National de survey of IE

Prophylaxis

Early recognition

Epidemiology

Risk factors

Clinical Presentation

Morbi-Mortality

Duke criteria

ECG confirmation

Blood cultures

Pho cardiology

Early treatment

Antibiotics

Optimal surgery

