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The hidden face of IE: clinical characteristics and prognosis in community centres

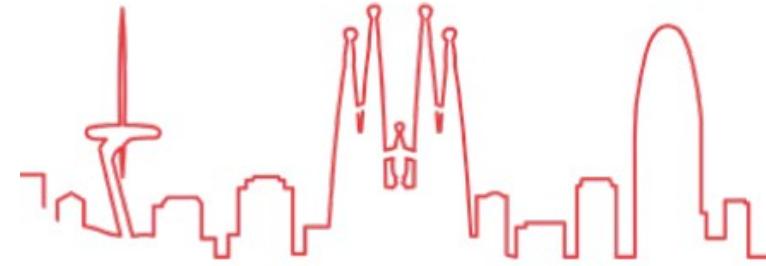

Parc Taulí Sabadell
Hospital Universitari

 I³PT
INSTITUT D'INVESTIGACIÓ
I INNOVACIÓ PARC TAULÍ

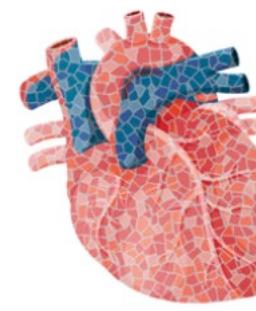
UAB
Universitat Autònoma
de Barcelona

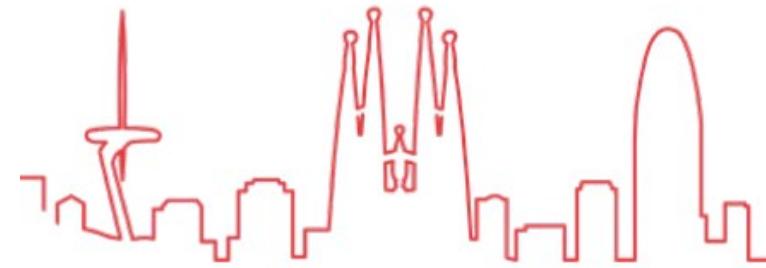
Oriol Gasch Blasi
Infectious Diseases Department
Parc Taulí University Hospital. Sabadell.

No conflicts of interest

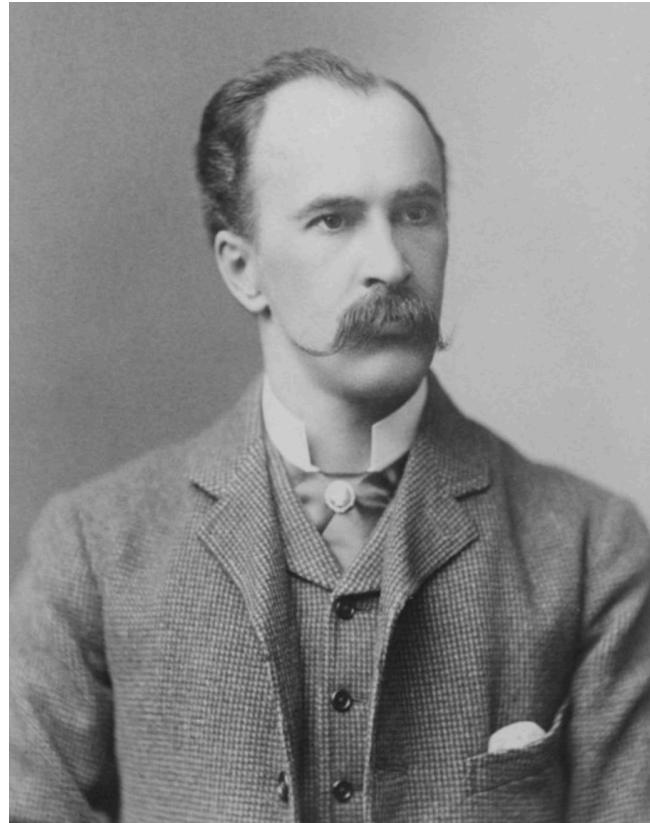


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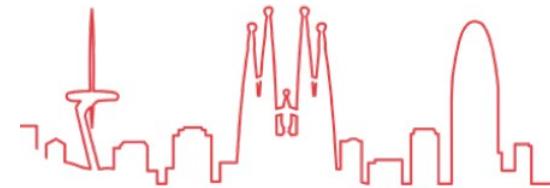
March 7, 1885.]

THE BRITISH MEDICAL JOURNAL.

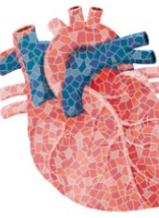
**THE GULSTONIAN LECTURES,
ON
MALIGNANT ENDOCARDITIS.**

Delivered at the Royal College of Physicians of London, March, 1885.
BY WILLIAM OSLER, M.D.,
Professor of Clinical Medicine at the University of Pennsylvania, Philadelphia.

Based on his personal experience with more than 200 patients



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Current knowledge on Infective Endocarditis

Clinical Presentation, Etiology and Outcome of Infective Endocarditis in the 21st Century: The International Collaboration on Endocarditis-Prospective Cohort Study

Murdoch DR. *Arch Intern Med.* 2009

2,781 adult patients with IE
58 Hospitals from 25 countries
Jun. 2000 – Sep. 2005

Minimum enrollment of 12 cases per year in centres with access to cardiac surgery

Clinical presentation, aetiology and outcome of infective endocarditis. Results of the ESC-EORP EURO-ENDO (European infective endocarditis) registry: a prospective cohort study

Habib. G *European Heart Journal* 2019

3,116 adult patients with IE
156 Hospitals from 40 countries
Jan. 2016 - Mar. 2018

Low-volume centres 7,6%
High-volume centres 92,4%

Current Epidemiology and Outcome of Infective Endocarditis
A Multicenter, Prospective, Cohort Study

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José María Miro, MD, PhD, Javier Bermejo, MD, PhD, Hugo Rodríguez-Abella, MD,
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Paola Tarabini, MD, and Emilio Bouza, MD, PhD, on behalf of the Spanish Collaboration on Endocarditis-
Grupo de Apoyo al Manejo de la Endocarditis Infecciosa en España (GAMES) (see Acknowledgment)

Muñoz P. *Medicine* 2015

1,804 adult patients with IE
25 Spanish Hospitals
Jan. 2008 - Dec. 2012



International cohorts

	ICE	EURO-ENDO
Study period	2000-2005	2016-18



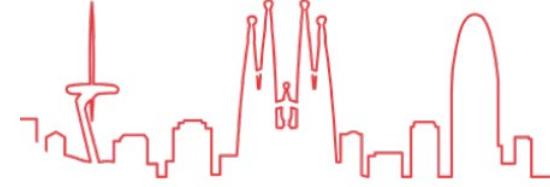
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International cohorts

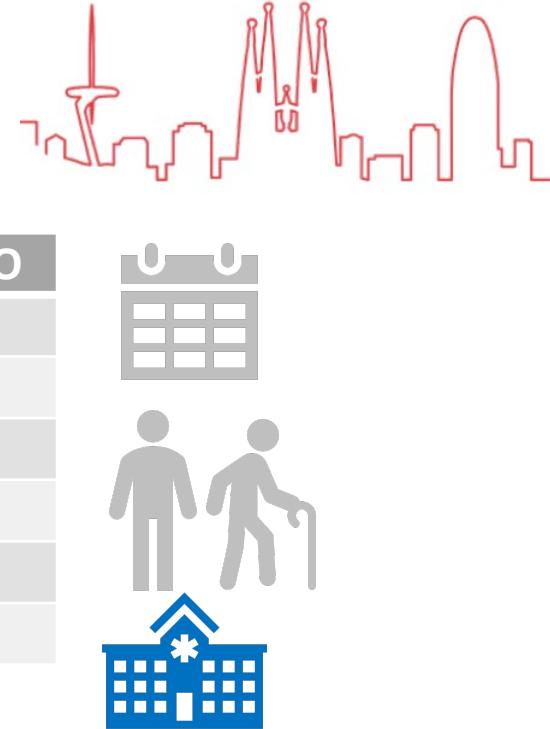
	ICE	EURO-ENDO
Study period	2000-2005	2016-18
Age, median (IQR)	58 (43-72)	63 (46-73)





International cohorts

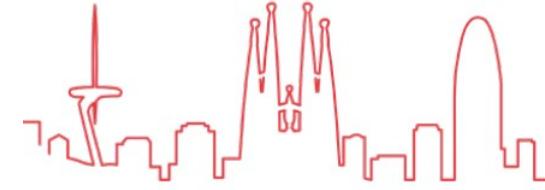
	ICE	EURO-ENDO
Study period	2000-2005	2016-18
Age, median (IQR)	58 (43-72)	63 (46-73)
Place of acquisition		
Community-acquired IE	72%	66%
Nosocomial HCA IE	14%	20%
Non-nosocomial HCA IE	9%	13%





International cohorts

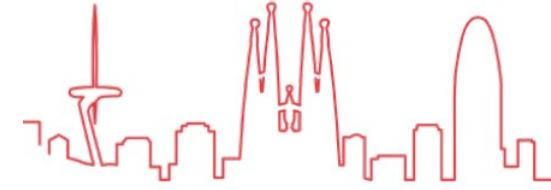
	ICE	EURO-ENDO
Study period	2000-2005	2016-18
Age, median (IQR)	58 (43-72)	63 (46-73)
Place of acquisition		
Community-acquired IE	71.5%	66%
Nosocomial HCA IE	14%	20%
Non-nosocomial HCA IE	9%	13%
IE Type		
Native	72%	59%
Prosthetic	21%	31%
CIED endocarditis	7%	10%





International cohorts

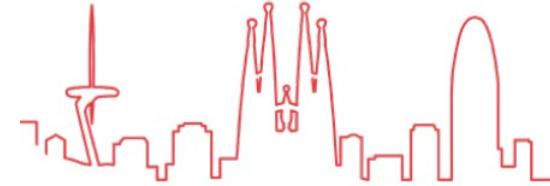
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Community-acquired IE	71.5%	66%
Nosocomial HCA IE	14%	20%
Non-nosocomial HCA IE	9%	13%
IE Type		
Native	72%	59%
Prosthetic	21%	31%
CIED endocarditis	7%	10%
Microbiology		
Viridans group Streptococci	17%	12%
<i>Enterococcus spp.</i>	10%	16%
<i>Staphylococcus aureus</i>	31%	31%
CNS	11%	13%
<i>Streptococcus gallolyticus</i>	6%	7%





International cohorts

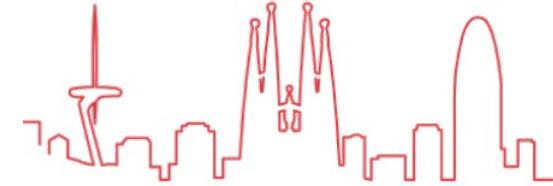
	ICE	EURO-ENDO
Complications		
Stroke	17%	27%
Embolization non-stroke	23%	25%
CHF	32%	41%
Persistent positive BC	9%	13%
Intracardiac abscess	14%	18%
New conduction abnormality	8%	16%





International cohorts

	ICE	EURO-ENDO
Complications		
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Embolization non-stroke	23%	25%
CHF	32%	41%
Persistent positive BC	9%	13%
Intracardiac abscess	14%	18%
New conduction abnormality	8%	16%
Indication surgery	NA	69%
Cardiac surgery	48%	51%

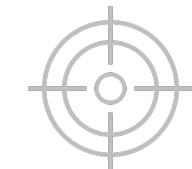
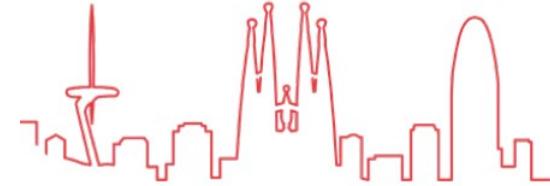


↑ 18% patients with surgical indication
were not operated



International cohorts

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Persistent positive BC	9%	13%
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New conduction abnormality	8%	16%
Indication surgery	ND	69%
Cardiac surgery	48%	51%
In-hospital mortality	18%	17%





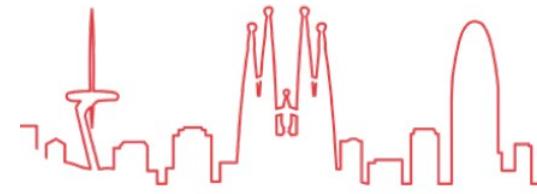
International cohorts. Risk factors for in-hospital death

Clinical Presentation, Etiology and Outcome of Infective Endocarditis in the 21st Century: The International Collaboration on Endocarditis-Prospective Cohort Study

Variable ^a	Original Model		
	OR ^b	95% CI	p-value
Age in ten year intervals	1.30	1.17–1.46	<0.001
Male gender	0.99	0.74–1.34	0.97
Transferred from another health care facility	0.97	0.74–1.29	0.85
Prosthetic valve endocarditis	1.47	1.13–1.90	0.004
Hemodialysis	1.06	0.73–1.53	0.76
Diabetes	1.28	0.88–1.86	0.20
Intravenous drug use	0.93	0.51–1.70	0.82
Cancer	1.04	0.65–1.67	0.86
Other chronic illness	1.36	0.95–1.95	0.10
Invasive procedure	0.96	0.66–1.39	0.82
Congenital heart disease	1.22	0.74–2.02	0.44
Elevated erythrocyte sedimentation rate	0.57	0.44–0.73	<0.001
Radiographic pulmonary edema	1.79	1.39–2.30	<0.001
Health care-associated acquisition	1.30	0.85–1.98	0.23
<i>S. aureus</i> IE	1.54	1.14–2.08	0.005
Coagulase-negative staphylococci IE	1.50	1.07–2.10	0.02
Viridans group streptococci IE	0.52	0.33–0.81	0.004
Mitral valve vegetation	1.34	1.06–1.68	0.01
Paravalvular complications	2.25	1.64–3.09	<0.001
Surgery during this episode	0.61	0.44–0.83	0.002

Clinical presentation, aetiology and outcome of infective endocarditis. Results of the ESC-EORP EURO-ENDO (European infective endocarditis) registry: a prospective cohort study

	Hazard ratio	95% CI	P-value*
Charlson index	1.07	[1.04–1.11]	<0.0001
Creatinine >2 mg/dL	1.58	[1.19–2.11]	<0.0017
Congestive heart failure	2.09	[1.58–2.77]	<0.0001
Vegetation length > 10 mm	2.12	[1.64–2.73]	<0.0001
Cerebral complication	2.21	[1.61–3.04]	<0.0001
Abscess	1.50	[1.07–2.10]	0.0186
Indication—surgery not performed	2.84	[2.00–4.03]	<0.001
Indication—surgery performed	0.63	[0.43–0.92]	0.0169



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TABLE 4. Independent Risk Factors for In-Hospital Mortality

Factor	OR	95% CI	P
Age	1.02	1.01–1.03	<0.01
Immunosuppressive therapy	2.61	1.68–4.04	<0.01
Previous heart surgery (previous to the episode of IE)	1.53	1.17–2.00	.002
CNS event	2.47	1.91–3.19	<0.01
Atrial fibrillation	1.45	1.09–1.93	.011
<i>S. aureus</i>	2.34	1.75–3.12	<0.01
Fungi	3.12	1.50–6.49	.002
Intracardiac complication	1.67	1.30–2.14	<0.01
Heart failure	2.97	2.30–3.83	<0.01
Septic shock	5.18	3.62–7.40	<0.01

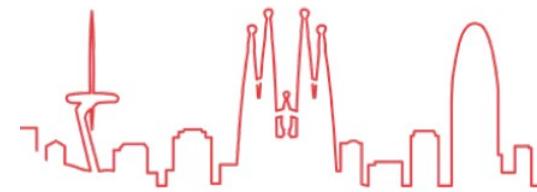


International cohorts. Risk factors for in-hospital death

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Clinical presentation, aetiology and outcome of infective endocarditis. Results of the ESC-EORP EURO-ENDO (European infective endocarditis) registry: a prospective cohort study



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Intracardiac complication	1.67	1.30–2.14	<0.01
Heart failure	2.97	2.30–3.83	<0.01
Septic shock	5.18	3.62–7.40	<0.01

Patient's background

Murdoch DR. Arch Intern Med. 2009, Habib. G European Heart Journal 2019, Muñoz P. Medicine 2015

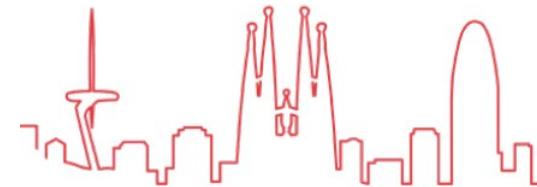


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	Hazard ratio	95% CI	P-value*
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Type of affected valve

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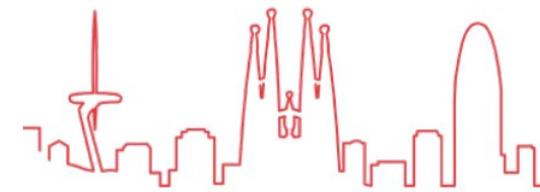


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Vegetation length > 10 mm	2.12	[1.64–2.73]	<0.0001
Cerebral complication	2.21	[1.61–3.04]	<0.0001
Abscess	1.50	[1.07–2.10]	0.0186
Indication—surgery not performed	2.84	[2.00–4.03]	<0.001
Indication—surgery performed	0.63	[0.43–0.92]	0.0169

TABLE 4. Independent Risk Factors for In-Hospital Mortality

Factor	OR	95% CI	P
Age	1.02	1.01–1.03	<0.01
Immunosuppressive therapy	2.61	1.68–4.04	<0.01
Previous heart surgery (previous to the episode of IE)	1.53	1.17–2.00	.002
CNS event	2.47	1.91–3.19	<0.01
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Heart failure	2.97	2.30–3.83	<0.01
Septic shock	5.18	3.62–7.40	<0.01

Local complications/Echocardiographic findings

Murdoch DR. Arch Intern Med. 2009, Habib. G European Heart Journal 2019, Muñoz P. Medicine 2015

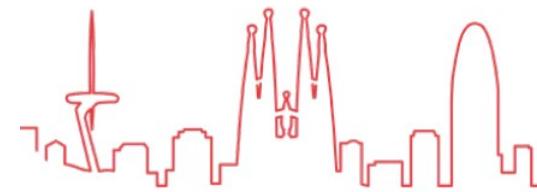


International cohorts. Risk factors for in-hospital death

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Variable ^a	Original Model		
	OR ^b	95% CI	p-value
Age in ten year intervals	1.30	1.17–1.46	<0.001
Male gender	0.99	0.74–1.34	0.97
Transferred from another health care facility	0.97	0.74–1.29	0.85
Prosthetic valve endocarditis	1.47	1.13–1.90	0.004
Hemodialysis	1.06	0.73–1.53	0.76
Diabetes	1.28	0.88–1.86	0.20
Intravenous drug use	0.93	0.51–1.70	0.82
Cancer	1.04	0.65–1.67	0.86
Other chronic illness	1.36	0.95–1.95	0.10
Invasive procedure	0.96	0.66–1.39	0.82
Congenital heart disease	1.22	0.74–2.02	0.44
Elevated erythrocyte sedimentation rate	0.57	0.44–0.73	<0.001
Radiographic pulmonary edema	1.79	1.39–2.30	<0.001
Health care-associated acquisition	1.30	0.85–1.98	0.23
<i>S. aureus</i> IE	1.54	1.14–2.08	0.005
Coagulase-negative staphylococci IE	1.50	1.07–2.10	0.02
Viridans group streptococci IE	0.52	0.33–0.81	0.004
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Systemic manifestations/complications

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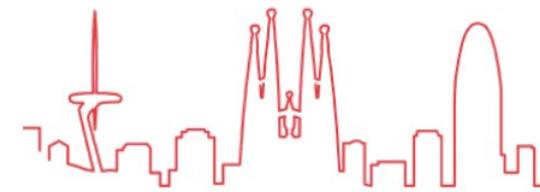


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Microbiology

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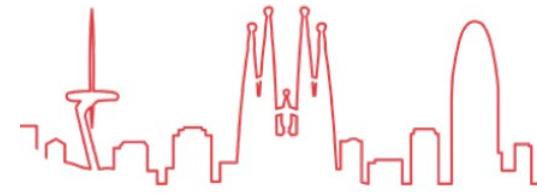


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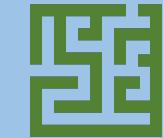
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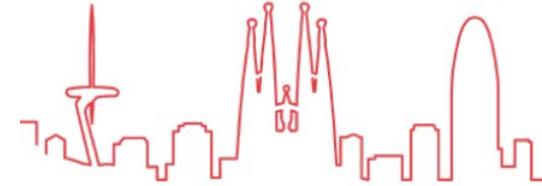
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Immunosuppressive therapy	2.61	1.68–4.04	<0.01
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Surgery

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Limitations of International cohorts



Non-cardiac surgery centres and community hospitals under-represented.

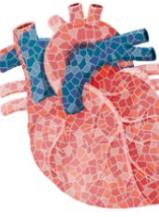
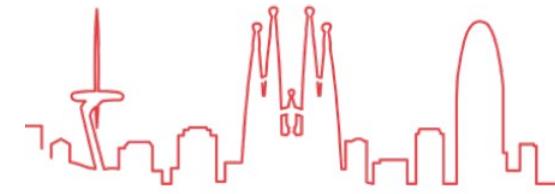
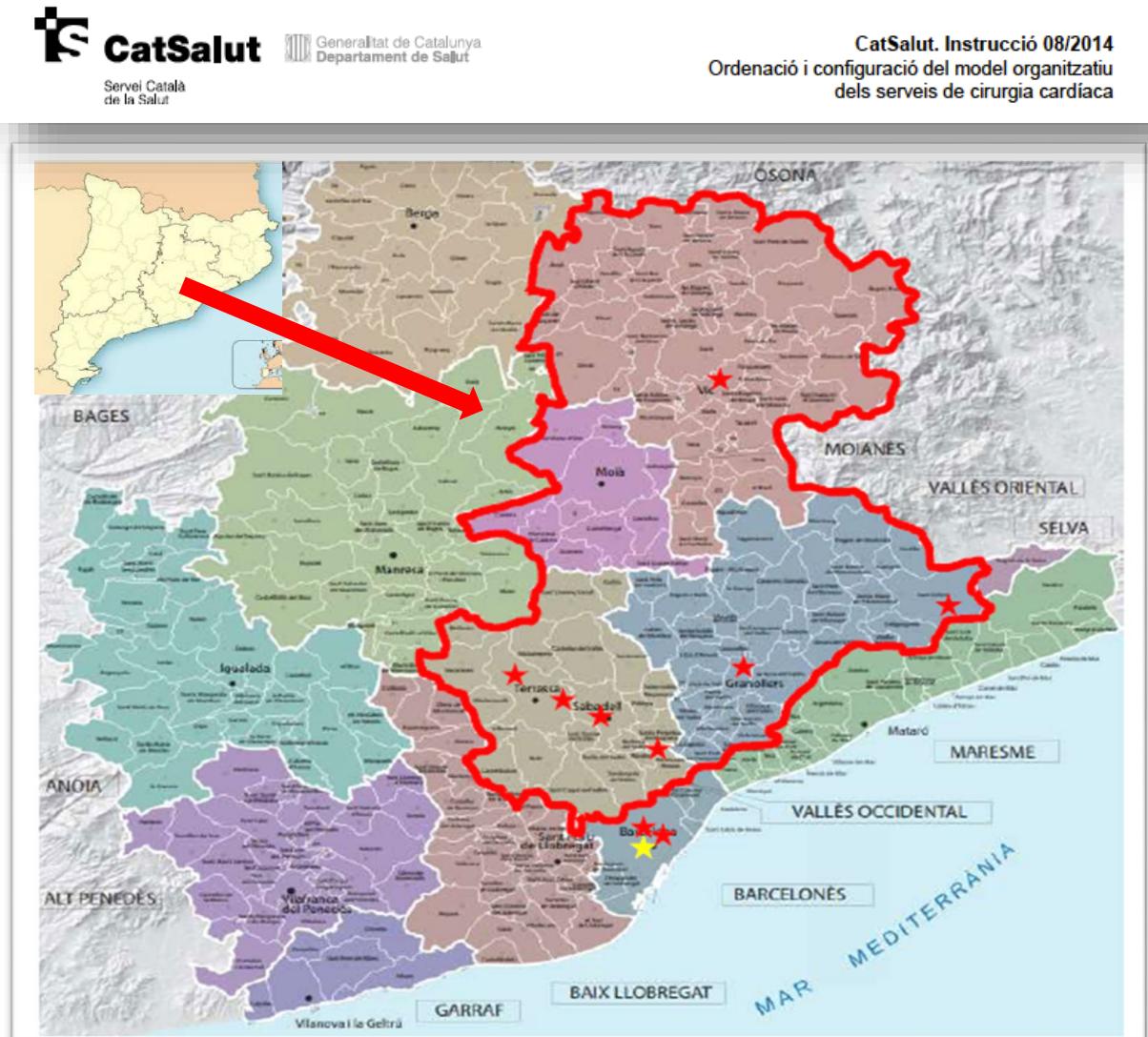
- ICE: 42% Transferred from another health facility

- EURO-ENDO: Low-volume centres 7,6%
High-volume centres 92,4%

- Demographics, clinical presentation, aetiology
- Frequency of episodes diagnosed
- Diagnostic tools (i.e [18F]FDG PET/CT), cardiac CT...)
- Direct availability of cardiac surgeon
- Prognostic factors



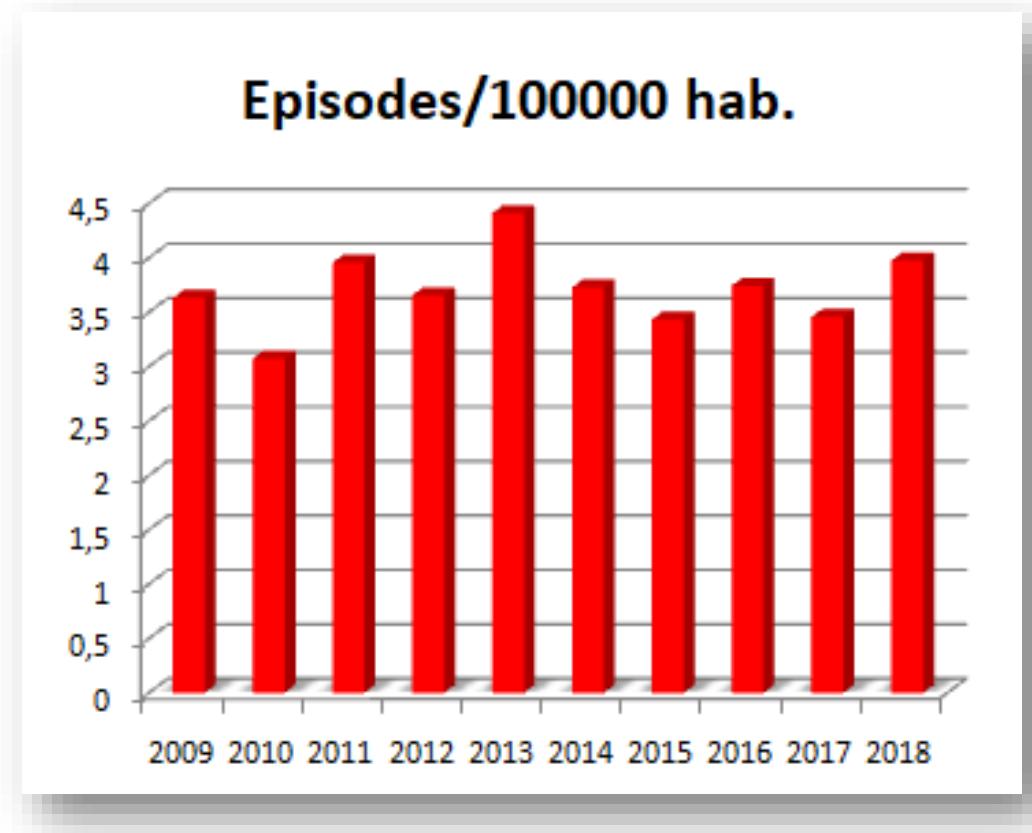
Central Catalonia 10 Endocarditis Teams (CC10ET)





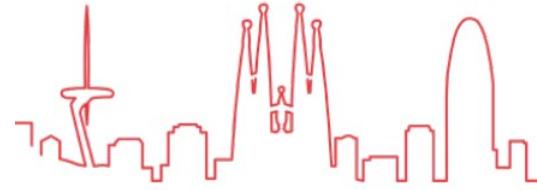
Central Catalonia 10 Endocarditis Teams (CC10ET)

502 episodes of infective endocarditis (2009-2018)



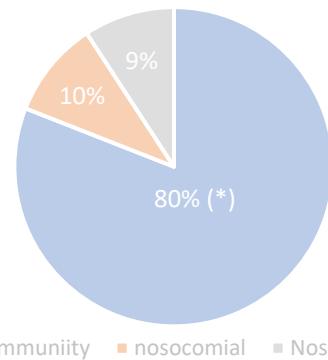
3.5 episodes/100.000 inhabitants CI 95% (3.45-3.56)





Central Catalonia 10 Endocarditis Teams (CC10ET)

DEMOGRAPHICS	
Age (P_{25} - P_{75})	72 (60-80)
Charlson (P_{25} - P_{75})	5 (3-7)



MICROBIOLOGY n (%)	
<i>S. aureus</i>	109 (21.7)
SASM	93 (17.5)
GVS	101 (20.1)
<i>E. faecalis</i>	64 (12.7)
CNS	56 (11.2)
<i>S. gallolyticus</i>	50 (10)

CC10ET > EURO-ENDO > ICE

(*) CC10ET > ICE > EURO-ENDO

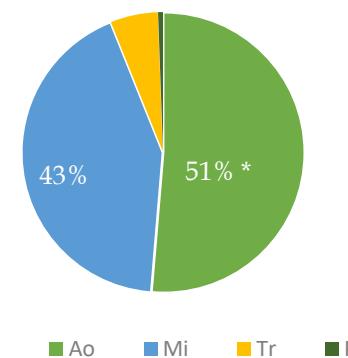
ECHOCARDIOGRAPHY n (%)	
Native	362 (72.1)
Biological Pr.	70 (13.9)
Mechanical Pr	44 (8.8)
CIEDs	37 (7.4)
Severe valve insufficiency	156 (31)
Intracardiac abscess	60 (12)
Fistula	28 (5.6)
Vegetation	375 (75)
>10 mm	43.5

CC10ET = ICE > EURO-ENDO

ICE > CC10ET

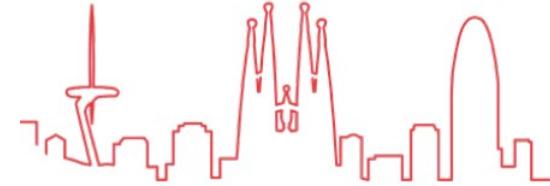
EURO-ENDO > ICE > CC10ET

ICE > EURO-ENDO = CC10ET



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Central Catalonia 10 Endocarditis Teams (CC10ET)

COMPLICATIONS n (%)	
Acute Pulmonary edema	28 (6)
Cardiogenic shock	22 (4)
Persistent bacteraemia	29 (6)
Ischaemic stroke	61 (12)
Haemorrhagic stroke	13 (3)
Systemic embolisms	136 (27)

SURGERY	
Surgical indication	240 (48)
Transferred to referral centre	183 (37)
Surgery	139 (28)

EURO-ENDO > ICE > CC10ET

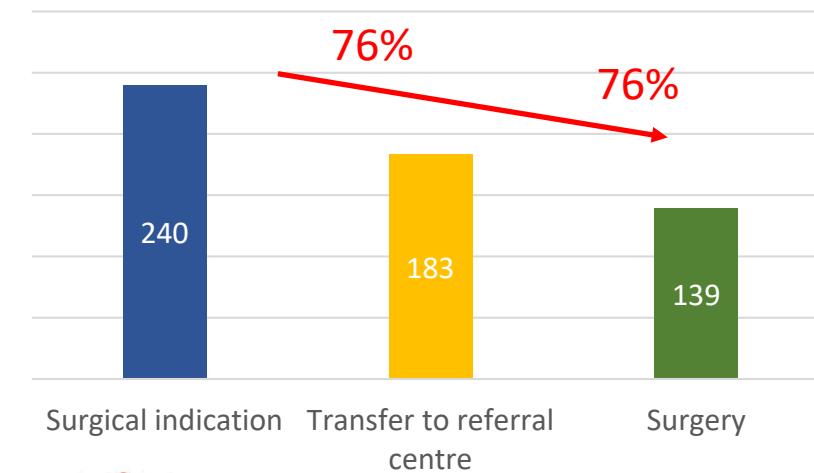
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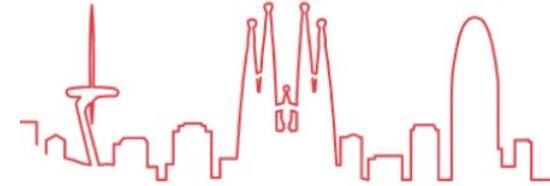
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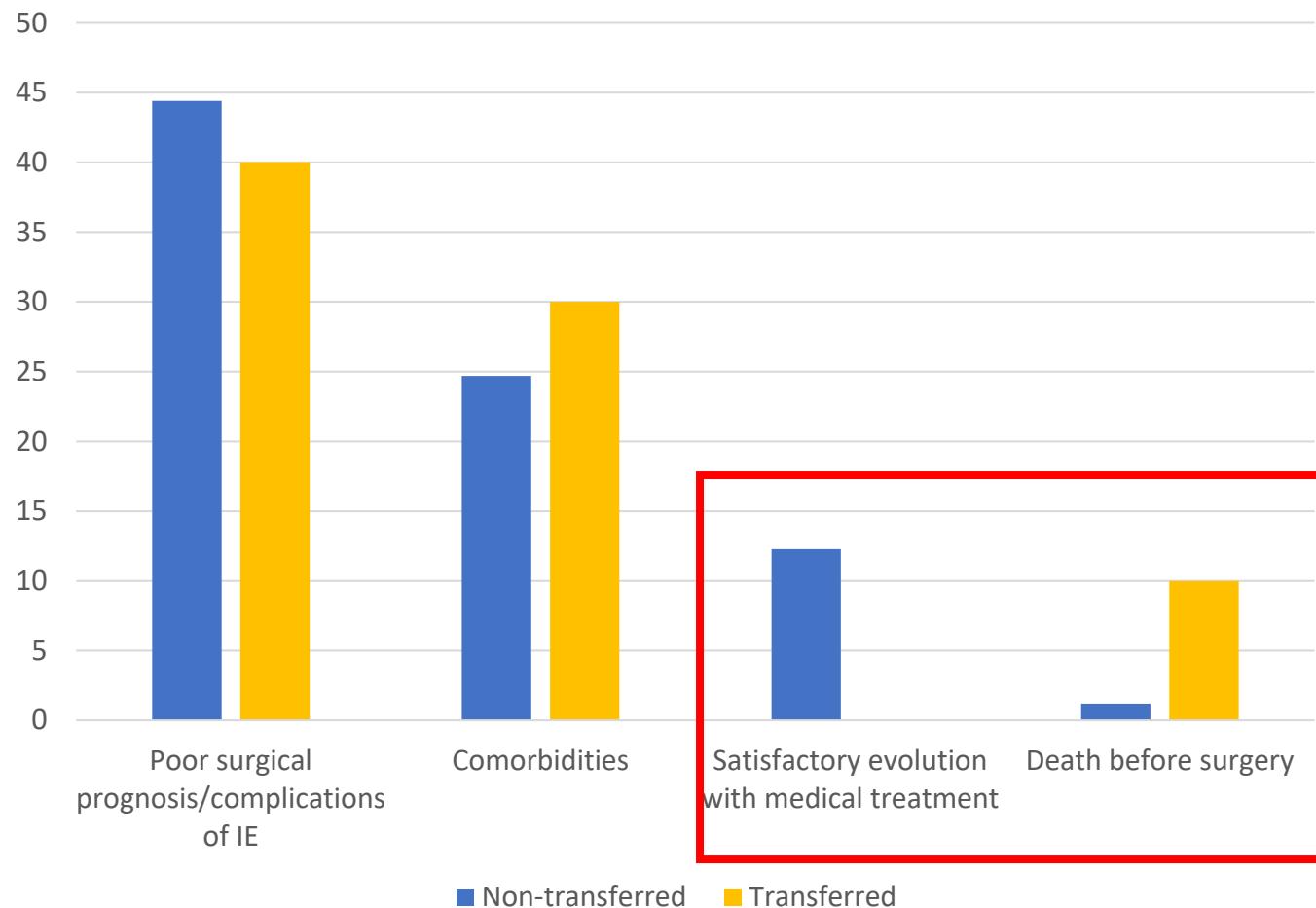


101 patients, 42% of all surgical indications **were not operated**



Central Catalonia 10 Endocarditis Teams (CC10ET)

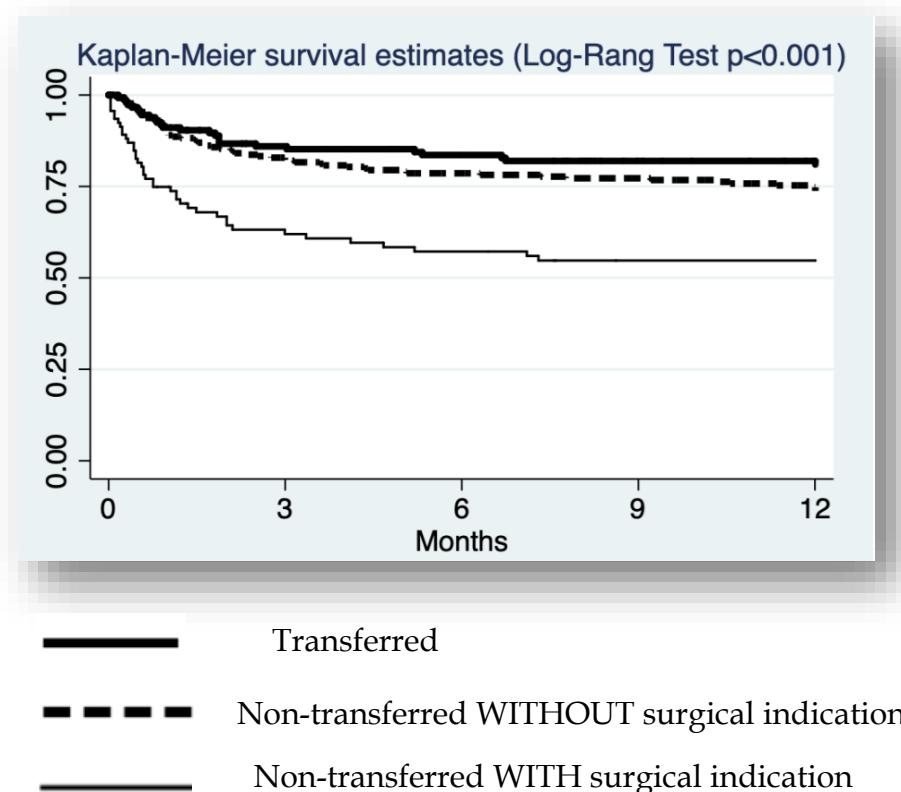
Reasons to preclude surgery

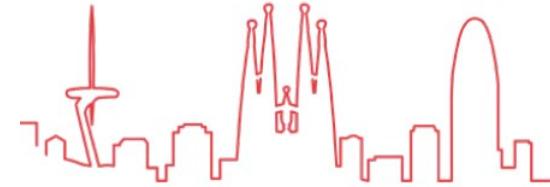


OUTCOMES

In-hospital mortality	98 (19.5)
One-year mortality	139 (30)

CC10ET >= ICE = EURO-ENDO





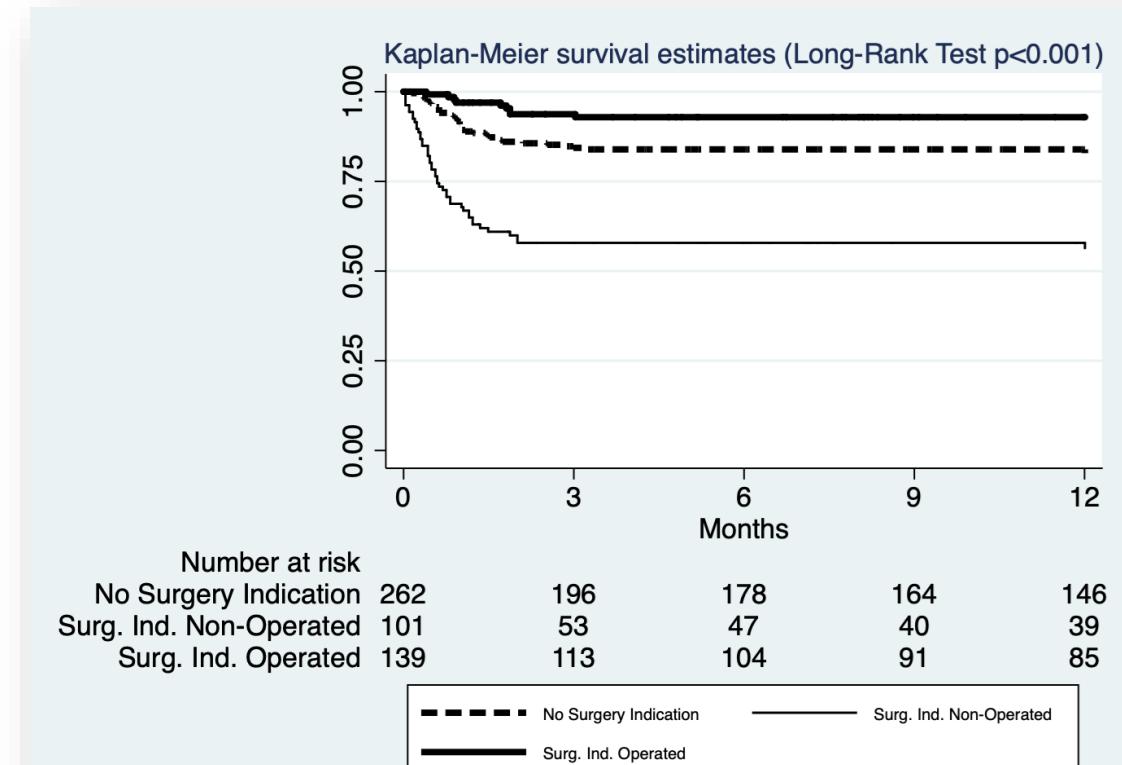
Central Catalonia 10 Endocarditis Teams (CC10ET)

Risk factors for In-hospital mortality

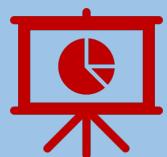
	OR	IC 95%	p-valor
Charlson score	1.19	(1.09, 1.30)	<0.001
Community acquisition	0.52	(0.29, 0.93)	0.027
<i>Staphylococcus aureus</i>	1.93	(1.08, 3.47)	0.027
Cardiac insufficiency	3.87	(2.28, 6.57)	<0.001
CNS embolism	2.95	(1.41, 5.14)	0.004
Surgery	0.42	(0.20, 0.87)	0.020

Risk factors for one-year mortality

	OR	IC 95%	p-valor
Charlson score	1.23	(1.13, 1.33)	<0.001
<i>Staphylococcus aureus</i>	1.82	(1.04, 3.18)	0.035
Cardiac insufficiency	3.74	(2.27, 6.16)	<0.001
Surgery	0.41	(0.21, 0.79)	0.008



(*) Transfer to cardiac surgery centre 1.23 (0.84-3.95)





Central Catalonia 10 Endocarditis Teams (CC10ET)

Among patients with surgical indication, are those transferred similar to not transferred?

	Transferred	Non-transferred	p-value
	183 (36.5)	88 (17.5)	
Male, n (%)	141 (77)	50 (56.8)	<0.001
Age (years), median (IQR)	67.0 (54-75)	79.0 (68.0-84.5)	<0.001
Charlson index, median (IQR)	3.0 (2.0-5.0)	6.0 (4.0-7.0)	<0.001
Community acquired, n (%)	155 (84.7)	62 (70.5)	0.004
<i>Staphylococcus aureus</i> , n (%)	29 (15.8)	25 (28.4)	0.011
Aortic Valve, n (%)	115 (62.8)	35 (39.8)	<0.001
Vegetation size (mm), median (IQR)	13.0 (9.0-17.0)	8.0 (6.0-13.0)	<0.001
Vegetation >10 mm, %	86 (66.7)	19 (33.3)	<0.001
Intracardiac abscess, n (%)	44 (24)	12 (13.6)	0.041
Intracardiac fistula, n (%)	25 (13.7)	3 (3.4)	0.008
CHE, n (%)	88 (48)	28 (29.8)	0.009



Not-transferred patients are older and have more comorbidities but have fewer echocardiographic and clinical findings associated with severity.





Central Catalonia 10 Endocarditis Teams (CC10ET)

Among unoperated patients WITH surgical indication, those transferred are different from those who were not?

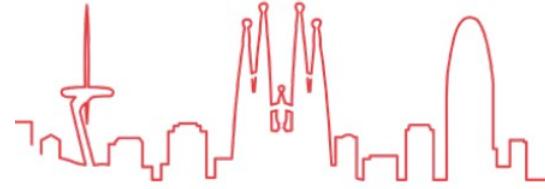


	Non-transferred (n= 81)	Transferred (n=20)	p-value
Male, n (%)	46 (56,8)	17 (85)	0,004
Severe regurgitation	27 (33,3)	13 (65)	0,009
Vegetation >10 mm, %	19 (33,3)	11 (66,7)	0,006
Vegetation size (mm), median (IQR)	8,0 (6,0-13,0)	13,5 (10,0-18,5)	0,002
Persistent bacteraemia, n (%)	5	0	0,023
One-year mortality, n (%)	44 (56,4)	11 (57,9)	0,907
One-year surgical treatment, n (%)	3 (3,7)	0 (0)	0,081

Transferred patients have more echocardiographic and clinical findings that confer severity, but are similar in terms of comorbidities.

One-year mortality in both groups is similar

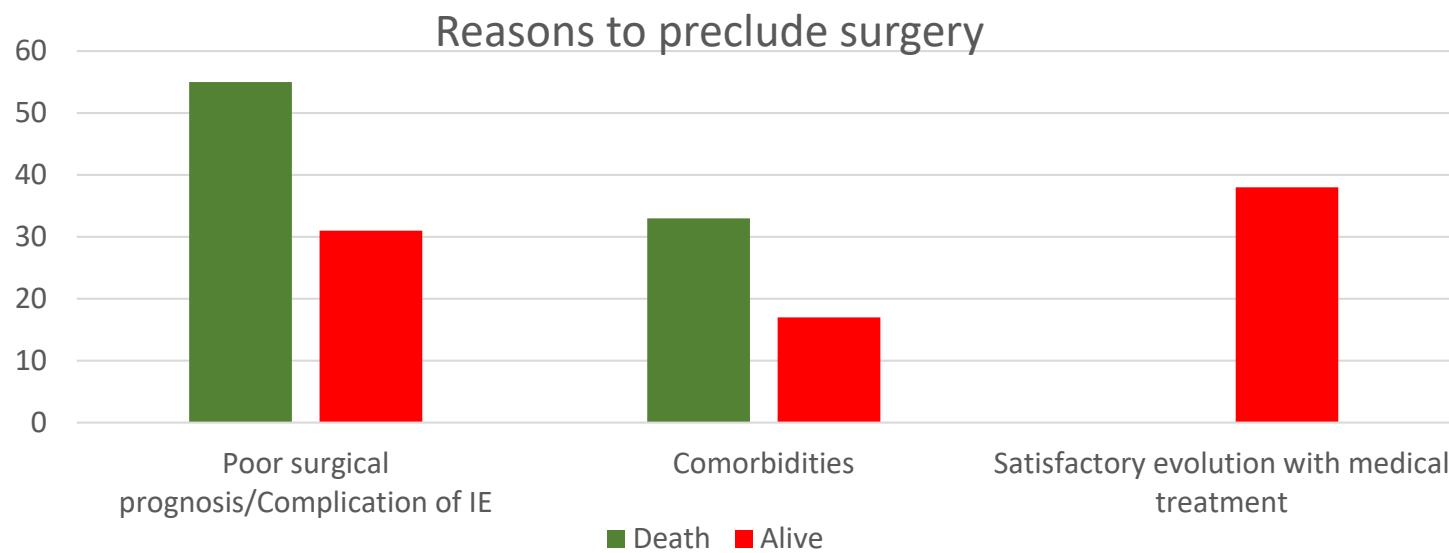


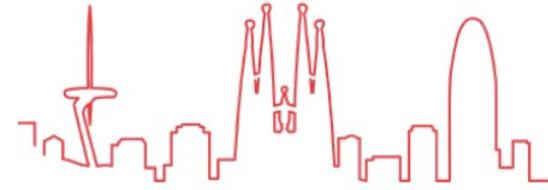


Central Catalonia 10 Endocarditis Teams (CC10ET)

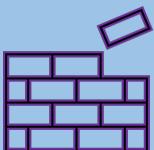
Among unoperated patients WITH surgical indication, those that died within the first year are different from those who alive?

	Death (n=55)	Alive (n=42)	p-value	OR (CI95%)
Charlson index, median (IQR)	6.0 (5.0-8.0)	4.0 (3.0-6.0)	<0.001	1,29 (1.08-1.54)
Heart Failure n (%)	26 (47.3)	9 (21.4)	0,006	3,08 (1,20-7,91)



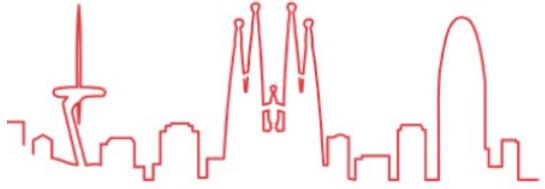


Conclusions



The hidden face of IE

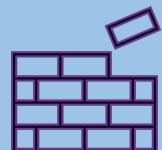




Conclusions

A significant proportion of patients diagnosed with IE in community centres will never reach the cardiac-surgery referral hospital despite their surgical indication.

The hidden face of IE



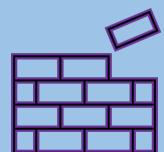


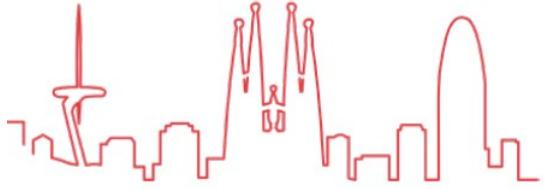
Conclusions

A significant proportion of patients diagnosed with IE in community centres will never reach the cardiac-surgery referral hospital despite their surgical indication.

These patients have differential characteristics with respect to those diagnosed at tertiary hospitals.

The hidden face of IE





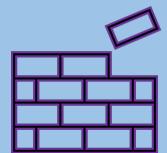
Conclusions

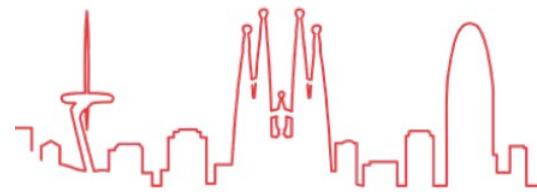
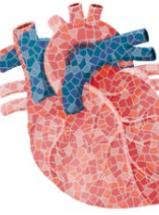
A significant proportion of patients diagnosed with IE in community centres will never reach the cardiac-surgery referral hospital despite their surgical indication.

These patients have differential characteristics with respect to those diagnosed at tertiary hospitals.

They are under-represented in large international cohorts.

The hidden face of IE





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