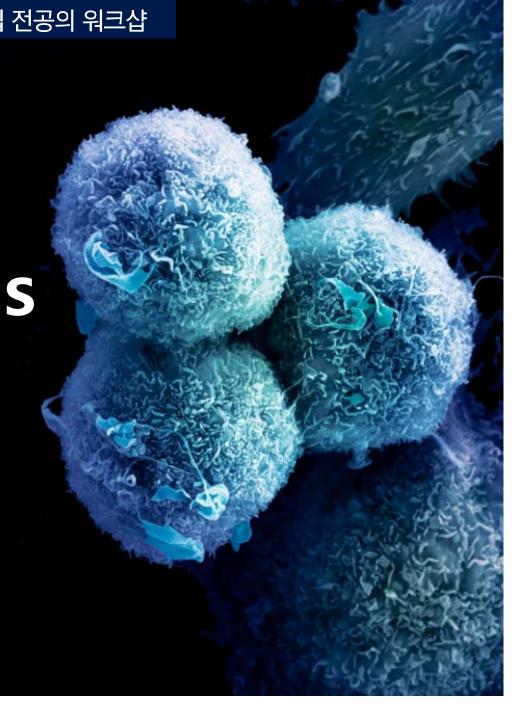
2022년 제 15차 전공의 연수교육 및 신입 전공의 워크샵

Infective endocarditis

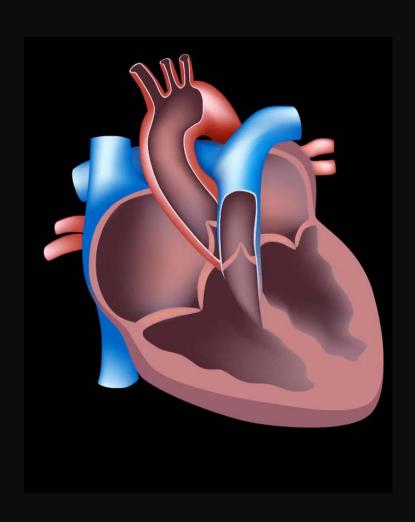
성균관 대학교 의과대학 삼성서울병원 심장외과

정수련



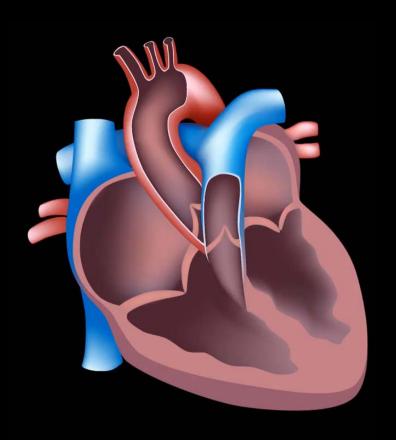






Infection of the endocardium

- Caused by bacteria/fungi in blood stream



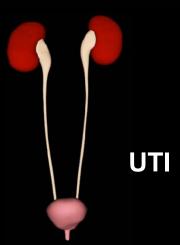
Infection of the endocardium

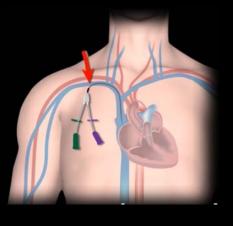
- Caused by bacteria/fungi in blood stream
- Abnormality of endocardium is required

Source of Infective Endocarditis



Skin abscess





Contaminated central line

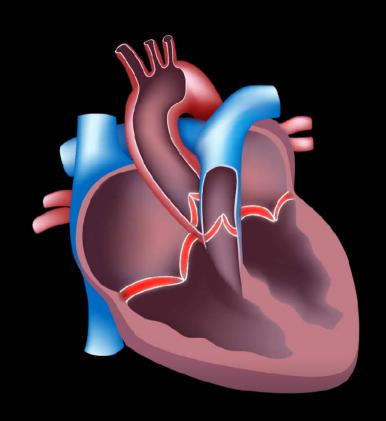




Dental procedures Or brushing

Drug injection or acupuncture

Source of Infective Endocarditis



Causative organisms:

Viridians streptococci (20%)

Streptococcus gallolyticus

Staphylococcus aureus(38%)

Coagulase-negative staphylococci

HACEK*

Enterococci

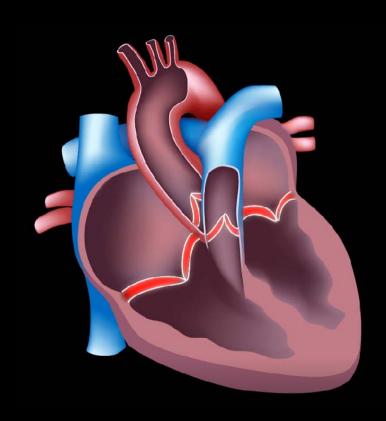
Pneumococci

Gram-negative bacillic

Fungus (Candida, Aspergillus etc)

*HACEK (Haemophilus, Aggregatibacter, Cardiobacterium, Eikenella, Kingella)

Source of Infective Endocarditis



Causative organisms:

Viridians streptococci

Streptococcus gallolyticus

Staphylococcus aureus (MRSA)

Coagulase-negative staphylococci

HACEK*

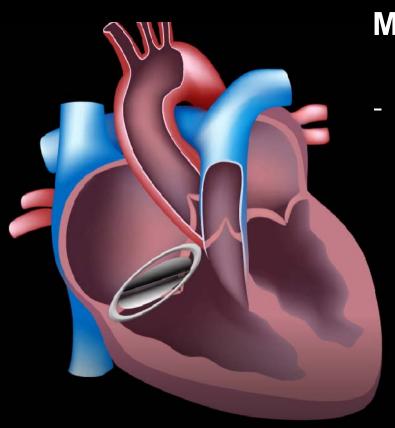
Enterococci

Pneumococci

Gram-negative bacillic

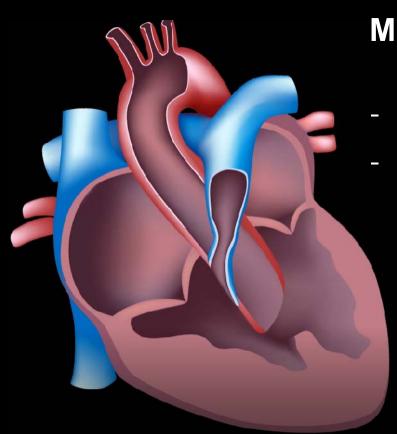
Fungus (Candida, Aspergillus etc)

*HACEK (Haemophilus, Aggregatibacter, Cardiobacterium, Eikenella, Kingella)

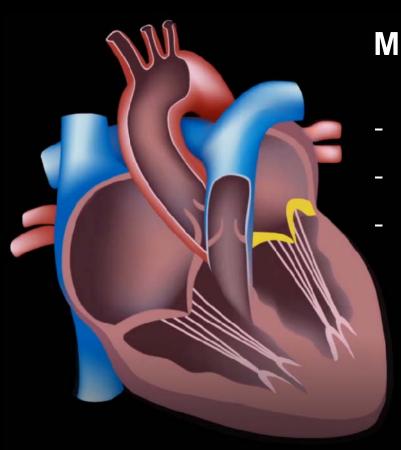


Major risk factors:

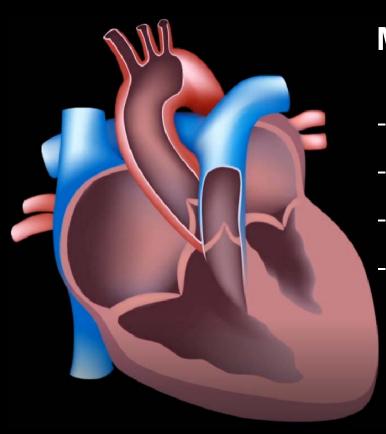
- Prosthetic valve/devices (highest risk)



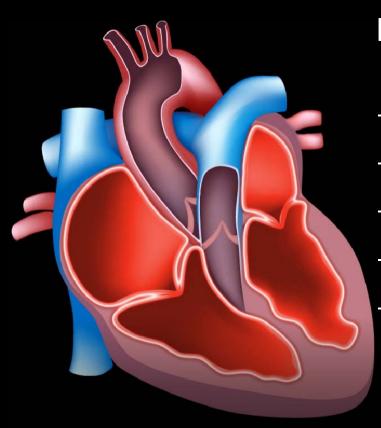
- Prosthetic valve/devices (highest risk)
- Congenital heart defect (VSD etc)



- Prosthetic valve/devices (highest risk)
- Congenital heart defect (VSD etc)
- Heart valve disorder

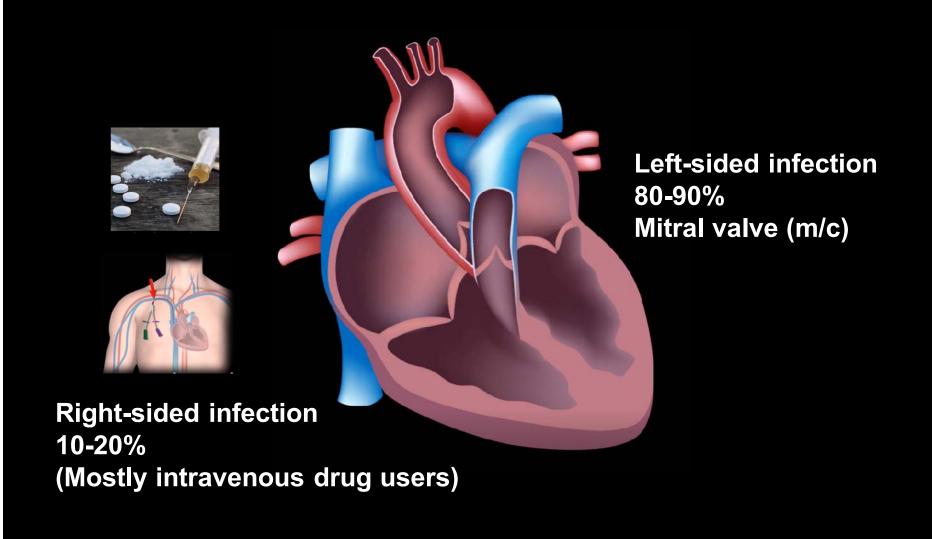


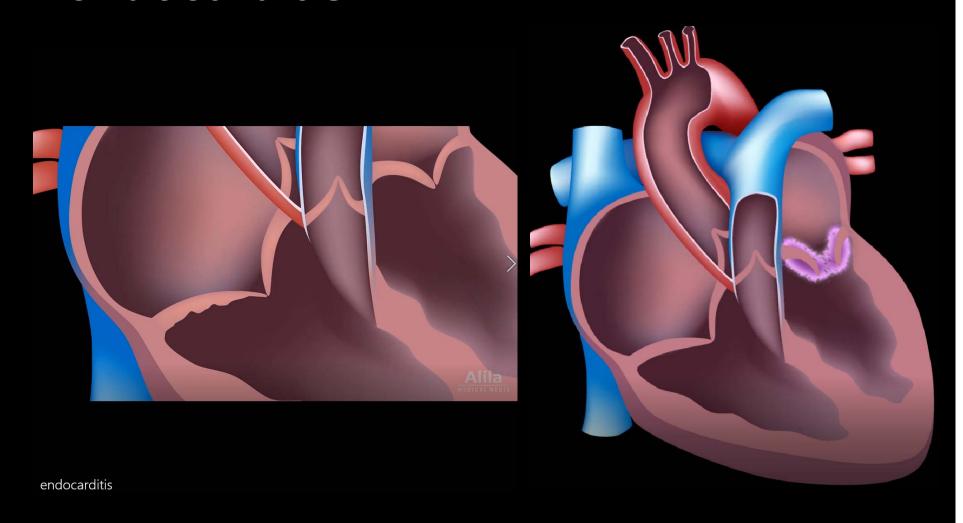
- Prosthetic valve/devices (highest risk)
- Congenital heart defect (VSD etc)
- Heart valve disorder
- Hypertrophic cardiomyopathy



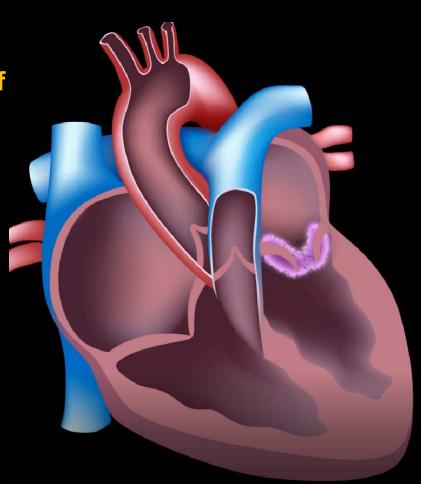
- Prosthetic valve/devices (highest risk)
- Congenital heart defect (VSD etc)
- Heart valve disorder
- Hypertrophic cardiomyopathy
- Previous endocarditis

Prevalence of Infective Endocarditis



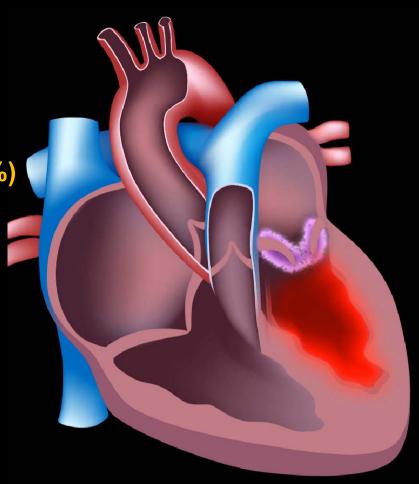


Heart valve dysfunction (in 85% of patients)



Heart valve dysfunction (in 85% of patients)

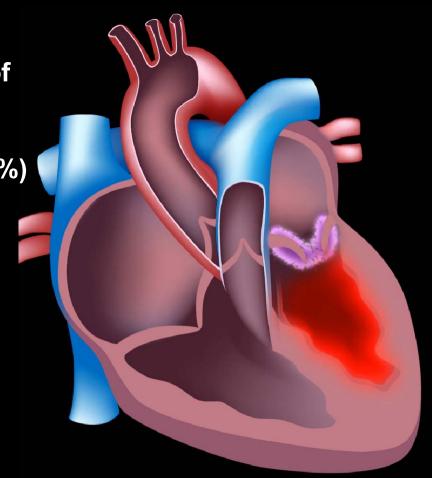
- Congestive heart failure (in 30-40%)



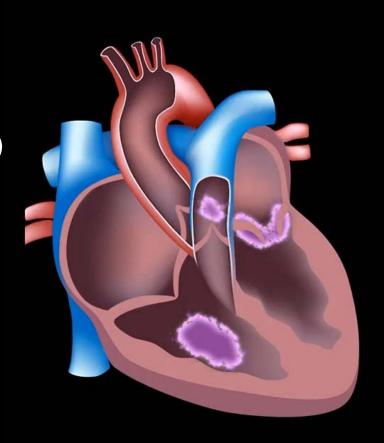
Heart valve dysfunction (in 85% of patients)

- Congestive heart failure (in 30-40%)

Conduction problemscomplete heart block



- Heart valve dysfunction (in 85% of patients)
- Congestive heart failure (in 30-40%)
- Conduction problems
- Embolism (blocked arteries)
 - Stoke
 - Pulmonary embolism
 - Acute myocardial infarct



Symptom of Infective endocarditis





Fever

Chills

Fatigues

Characteristic spot

Dyspnea or DOE

Pitting edema



Diagnosis of Infective endocarditis

1. Fever + Heart valve disorder

2. Characteristic symptoms





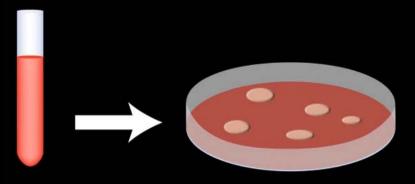


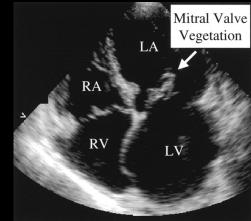




3. Blood culture for evidence of blood infection

4. Imaging to show vegetation





MAJOF CRITER		minor criteria
1. Blood positive	culture	

- 1. Typical microorganisms consistent with IE form 2 separate blood culture: Viridans streptococci, Streptococcus bovis, HACEK groups, S.aureus or community-acquired enterococci
- 2. Persistently positive blood culture result
- ; At least 2 positive culture result of blood sample drawn 12hr
- All of 3 or most of >4 separate culture samples of blood
- Single positive blood culture result for Coxiella burnetii or

	MAJOR CRITERIA	minor cr	iteria
1.	Blood culture positive		
1.	Evidence of endocardial involvement		
1	Cabacardiagram data	oto vogototion	ahaaaaa

- 1. Echocardiogram detects vegetation, abscess, or new partial dehiscence of prosthetic valve.
- 1. New valvular regurgitation or stenosis.
- 1. Oscillating intracardiac mass on valve or supporting structures

	MAJOR CRITERIA	minor criteria
1.	Blood culture positive	 Predisposing factor Temperature >38
1.	Evidence of endocardial involvement	degree 3. Vascular phenomena 4. Immunologic phenomena 5. Microbiologic evidence
1.	Intravenous drug use condition	e or a predisposing heart

Major arterial emboli, septic emboli, pulmonary infarcts, mycotic Aneurysm, intracranial



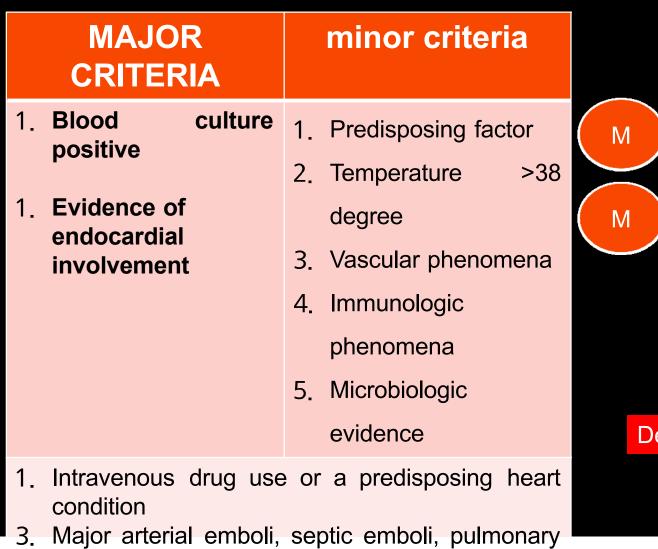
Janeway lesion



Osler's node (painful)



Roth's spot (retinal)

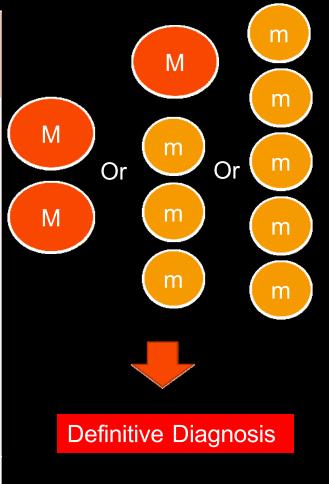


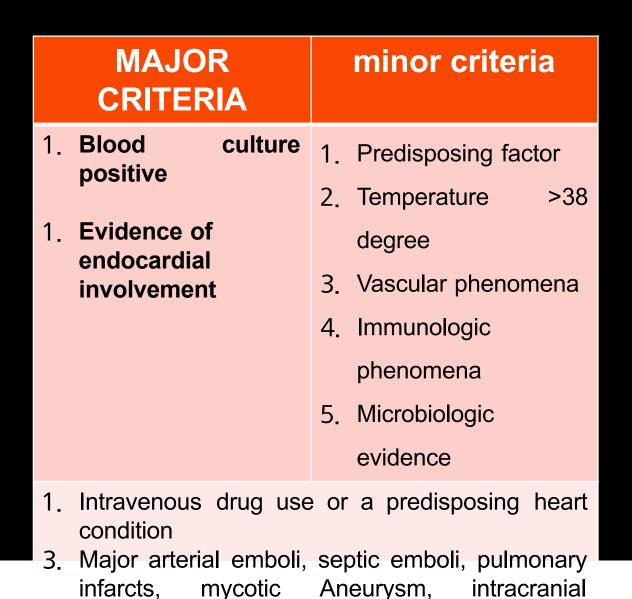
Aneurysm,

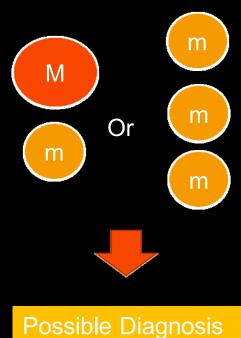
intracranial

infarcts,

mycotic







Echocardiography (TTE or TEE)

Clinical and Echocardiographic Features That Suggest Potential Need for Surgical intervention

Vegetation

Persistent vegetation after systemic embolization

Anterior mitral leaflet vegetation, particularly with size > 10mm*

≥1 Embolic events during first 2 weeks of antimicrobial therapy *

Increase vegetation size despite appropriate antimicrobial therapy *

Valvular dysfunction

Acute Aortic or mitral insufficiency with signs of ventricular failure

Heart failure unresponsive to medical therapy

Valve perforation or rupture

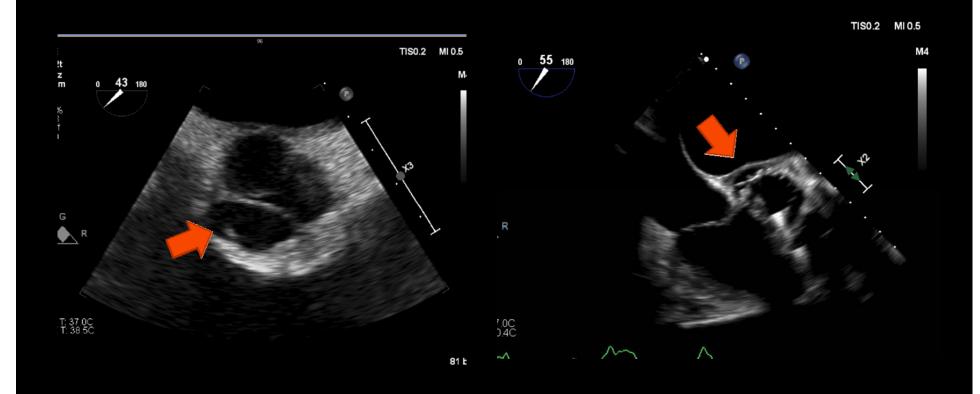
Perivalvular extension

Valvular dehiscence, rupture or fistula

New heart block

Large abscess or extension of abscess despite appropriate antimicrobial therpy

Echocardiography (TTE or TEE)



Native valve endocarditis(AV) Vegetation at leaflet Prosthetic valve endocarditis(AV)
Dehiscence

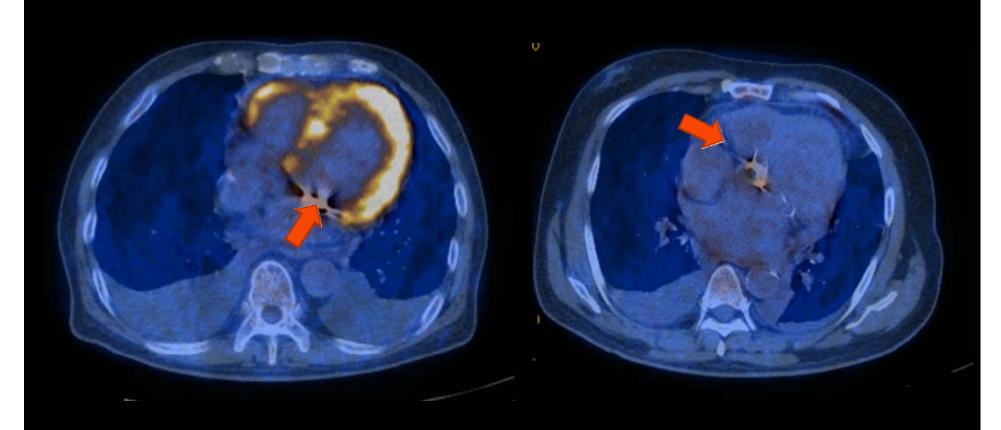
Cardiac CT for IE



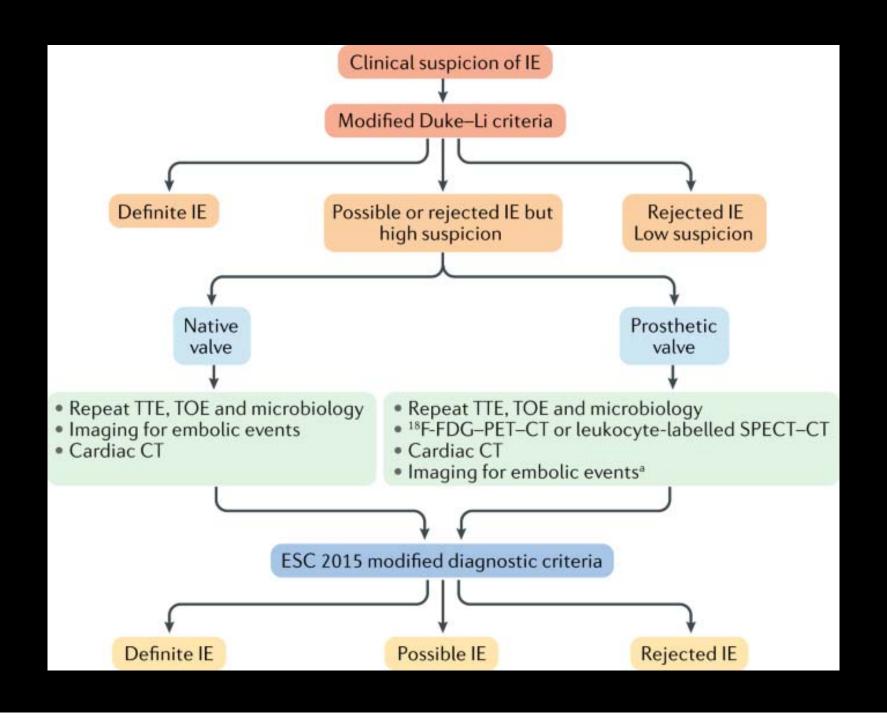
Pseudoaneurysm around prosthetic valve (AV)

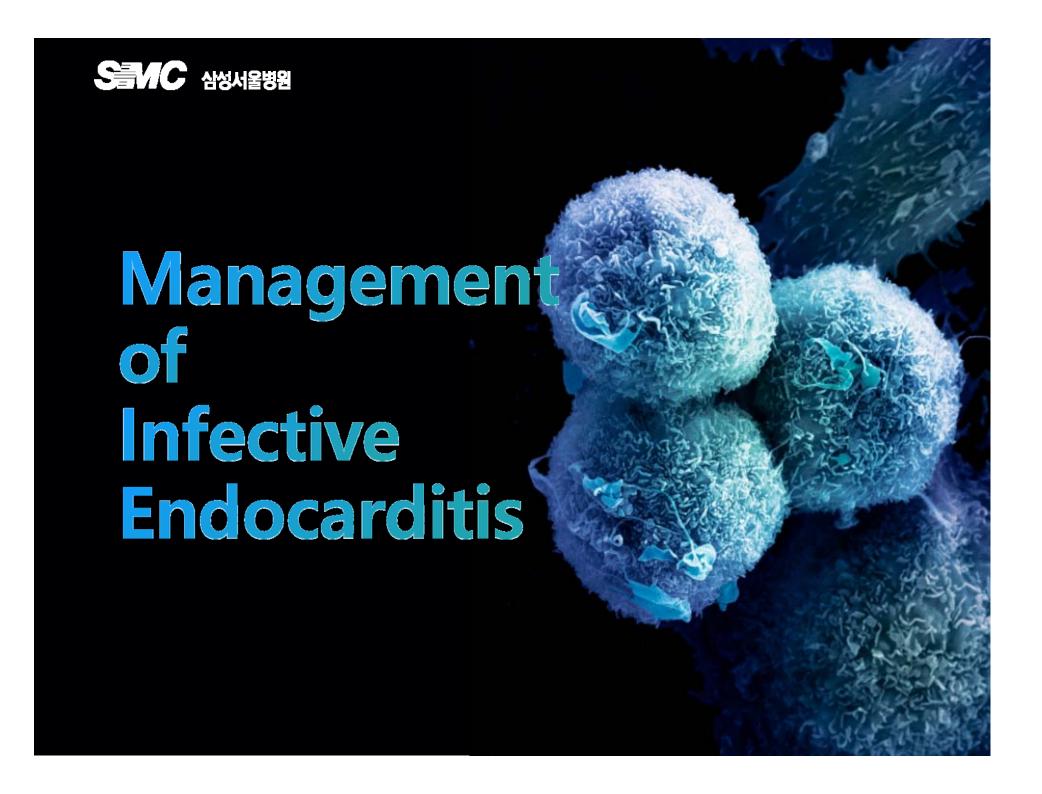
Abscess formation at aortomitral continuity

18F-fluorodeoxyglucose PET/CT



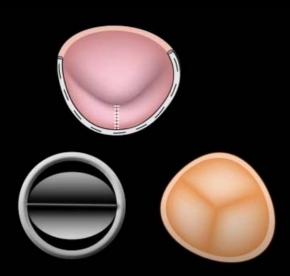
High uptake of FDG at sawing cuff of prosthetic valve





Treatment of Infection endocarditis





- Empiric antibiotics therapy
- 2 to 8 weeks of IV antibiotic therapy (after identified organism)
- Removal of potential source of infection
- Repair or replace to heart valve

Antibiotic therapy (Empirical)

Antibiotic	Dosage and route	Class	Levelc	Comments	
Community-acquired native valves or late prosthetic valves (≥12 months post surgery) endocarditis					
Ampicillin with (Flu)cloxacillin or oxacillin with Gentamicin ^d	12 g/day i.v. in 4–6 doses 12 g/day i.v. in 4–6 doses 3 mg/kg/day i.v. or i.m. in 1 dose	lla	С	Patients with BCNIE should be treated in consultation with an ID specialist.	
Vancomycin ^d with Gentamicin ^d	30–60 mg/kg/day i.v. in 2–3 doses 3 mg/kg/day i.v. or i.m. in 1 dose	Шь	С	For penicillin-allergic patients	
Early PVE (<12 m	Early PVE (<12 months post surgery) or nosocomial and non-nosocomial healthcare associated endocarditis				
Vancomycin ^d with Gentamicin ^d with Rifampin	30 mg/kg/day i.v. in 2 doses 3 mg/kg/day i.v. or i.m. in 1 dose 900–1200 mg i.v. or orally in 2 or 3 divided doses	IIb	U	Rifampin is only recommended for PVE and it should be started 3–5 days later than vancomycin and gentamicin has been suggested by some experts. In healthcare associated native valve endocarditis, some experts recommend in settings with a prevalence of MRSA infections >5% the combination of cloxacillin plus vancomycin until they have the final S. aureus identification	

Antibiotic therapy (Empiric therapy)

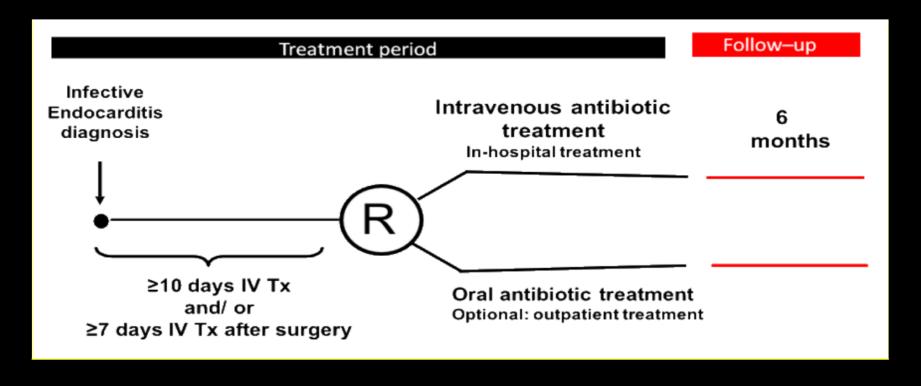
	Table 1				
Treatment of IE Caused by Viridans Group Streptococci and Streptococcus bovis*					
Duration Native-Valve IE Drug Regimen Therapy (
Penicillin-susceptible (MIC ≤0.12 mcg/mL)	Penicillin G sodium 12 to 18 million U continuous infusion over 24 hours or divided into four or six equal IV doses per 24 hours OR	4			
	Ceftriaxone 2 g in one dose every 24 hours IV/IM OR	4			
	Penicillin G sodium 12 to 18 million U continuous infusion over 24 hours or in six equal doses IV per 24 hours 0R	2			
	Ceftriaxone 2 g in one dose every 24 hours IV/IM PLUS gentamicin 3 mg/kg in one dose every 24 hours IV/IM OR	2 2			
	Vancomycin 15 mg/kg every 12 hours IV	4			
Penicillin, relatively resistant (MIC >0.12 to ≤0.5 mcg/mL)	Penicillin G sodium 24 million U continuous infusion over 24 hours or divided into four or six equal IV doses per 24 hours 0R	4			
	Ceftriaxone 2 g in one dose every 24 hours IV/IM PLUS gentamicin 3 mg/kg in one dose every 24 hours IV/IM OR	4 2			
	Vancomycin 15 mg/kg every 12 hours IV	4			
Prosthetic- Valve IE					
Penicillin-susceptible (MIC ≤0.12 mcg/mL)	Penicillin G sodium 24 million U continuous infusion over 24 hours or divided into four or six equal IV doses per 24 hours OR	6			
	Cettriaxone 2 g in one dose every 24 hours IV/IM with or without gentamicin 3 mg/kg in one dose every 24 hours IV/IM OR	6 2			
	Vancomycin 15 mg/kg every 12 hours IV	6			
Penicillin, relatively or fully resistant (MIC >0.12 mcg/mL	Penicillin G sodium 24 million U continuous infusion over 24 hours or divided into four or six equal IV doses per 24 hours OR	6			
	Ceftriaxone 2 g in one dose every 24 hours IV/IM PLUS gentamicin 3 mg/kg in one dose every 24 hours IV/IM OR	6			
	Vancomycin 15 mg/kg every 12 hours IV	6			
* Danger for normal renal function	Source: Adapted from reference 7.				

Treatment of IE Caused by the Staphylococcus Species*				
Native-Valve IE	Drug Regimen	Therapy (weeks)		
Oxacillin-susceptible	Nafcillin or oxacillin 12 g divided in four to six equal doses per 24 hours IV PLUS optional gentamicin 3 mg/kg in two to three equal	6		
	doses per 24 hours IV/IM	3 to 5 days		
	If allergic (nonanaphylactoid) to penicillins, consider: Cefazolin 2.g every eight hours IV PLUS optional gentamicin 3 mg/kg in two to three equal doses	6		
	per 24 hours IV/IM	3 to 5 days		
Oxacillin-resistant	Vancomycin 15 mg/kg every 12 hours IV	6		
Prosthetic- Valve IE				
Oxacillin-susceptible	Nafcillin or oxacillin 2 g every four hours IV PLUS	≥6		
	rifampin 300 mg every eight hours IV/PO PLUS gentamicin 3 mg/kg in two to three equal doses per 24 hours IV/IM	≥6 2		
Oxacillin-resistant	Vancomycin 15 mg/kg every 12 hours IV PLUS	≥6		
	rifampin 300 mg every eight hours IV/PO PLUS gentamicin 3 mg/kg in two to three equal doses	≥6		
	per 24 hours IV/IM	2		

Empirical IV antibiotics therapy during 4-6 weeks (maintained over 8weeks for Enterococci species)

Partial Oral Treatment of Endocarditis (POET) trial

- Stable, native,, left-side of heart endocarditis
- Streptococci, E.faecalis, S. aureus or coagulase-negative staphylococci
- Death, unplanned cardiac surgery, embolic event, relapse of positive blood culture after 6months



Partial Oral Treatment of Endocarditis (POET) trial

- Stable, native,, left-side of heart endocarditis
- Streptococci, E.faecalis, S. aureus or coagulase-negative staphylococci
- Death, unplanned cardiac surgery, embolic event, relapse of positive blood culture after 6months

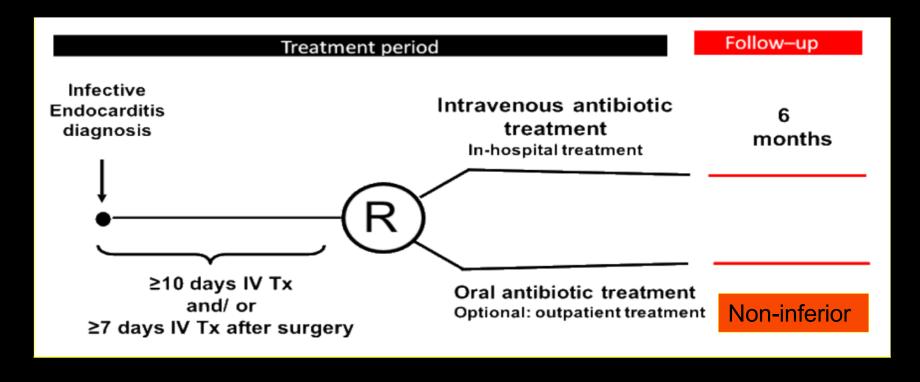
Table. Oral	Antibiotic	Regimens	Recommended	in the	POET Trial
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PENICILLIN- AND METHICILLIN-SUSCEPTIBLE STAPHYLOCOCCUS AUREUS AND COAGULASE-NEGATIVE STAPHYLOCOCCI	METHICILLIN-RESISTANT COAGULASE-NEGATIVE STAPHYLOCOCCI	ENTEROCOCCUS FAECALIS	STREPTOCOCCI WITH PENICILLIN MIC <1 MG/L	STREPTOCOCCI WITH PENICILLIN MIC >1 MG/L
Amoxicillin 1 g 4 times a day and fusidic acid 0.75 g 2 times a day	Linezolid 600 mg 2 times a day and fusidic acid 0.75 g 2 times a day	Amoxicillin 1 g 4 times a day and rifampin 600 mg 2 times a day	Amoxicillin 1 g 4 times a day and rifampin 600 mg 2 times a day	Linezolid 600 mg 2 times a day and rifampin 600 mg 2 times a day
Amoxicillin 1 g 4 times a day and rifampin 600 mg 2 times a day	Linezolid 600 mg 2 times a day and rifampin 600 mg 2 times a day	Amoxicillin 1 g 4 times a day and moxifloxacin 400 mg 1 time a day	Linezolid 600 mg 2 times a day and rifampin 600 mg 2 times a day	Moxifloxacin 400 mg 1 time a day and rifampin 600 mg 2 times a day
Linezolid 600 mg 2 times a day and fusidic acid 0.75 g 2 times a day		Linezolid 600 mg 2 times a day and rifampin 600 mg 2 times a day	Linezolid 600 mg 2 times a day and moxifloxacin 400 mg 1 time a day	Moxifloxacin 400 mg 1 time a day and clindamycin 600 mg 3 times a day
Linezolld 600 mg 2 times a day and rifampin 600 mg 2 times a day		Linezolid 600 mg 2 times a day and moxifloxacin 400 mg 1 time a day		

MIC indicates minimum inhibitory concentration.

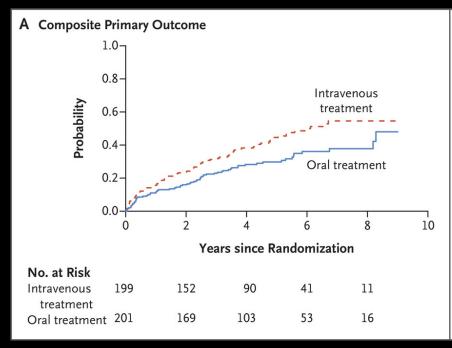
Partial Oral Treatment of Endocarditis (POET) trial

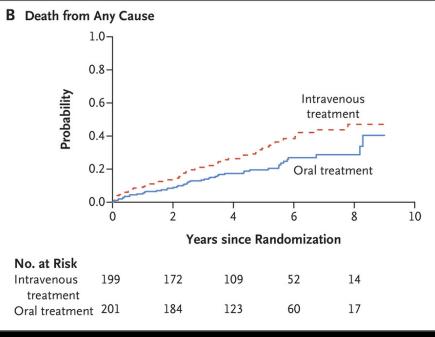
- Stable, native,, left-side of heart endocarditis
- Streptococci, E.faecalis, S. aureus or coagulase-negative staphylococci
- Death, unplanned cardiac surgery, embolic event, relapse of positive blood culture after 6months



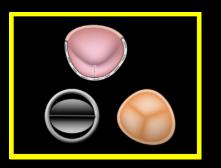
Partial Oral Treatment of Endocarditis (POET) trial

- Stable, native,, left-side of heart endocarditis
- Streptococci, E.faecalis, S. aureus or coagulase-negative staphylococci
- Randomized trial (IV antibiotics vs oral medication)





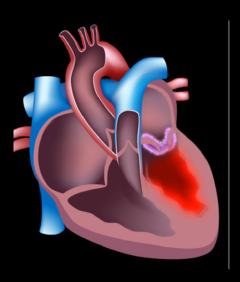
Indication for surgery



1. Heart failure

2. Uncontrolled infection

3. Prevention of embolism

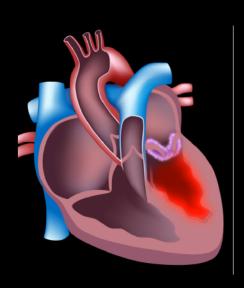


to brain, kidney, spleen, heart ...
to lungs

Valvular dysfunction Dehiscence of prosthetic valve Fistula **Failed antibiotics**

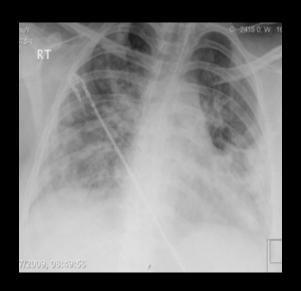
Recurrent embolism With large vegetation

Indication for surgery



Valvular dysfunction with heart failure

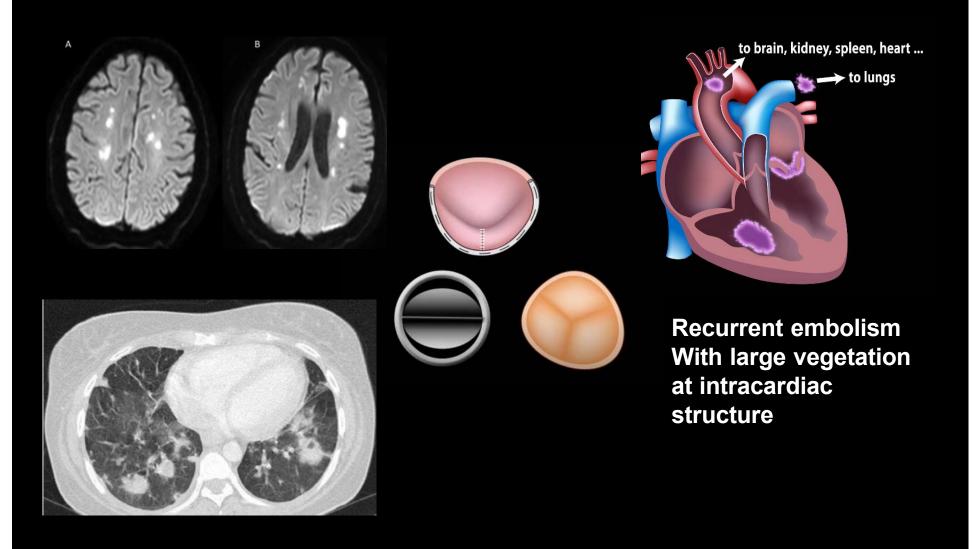
Dehiscence of prosthetic valve



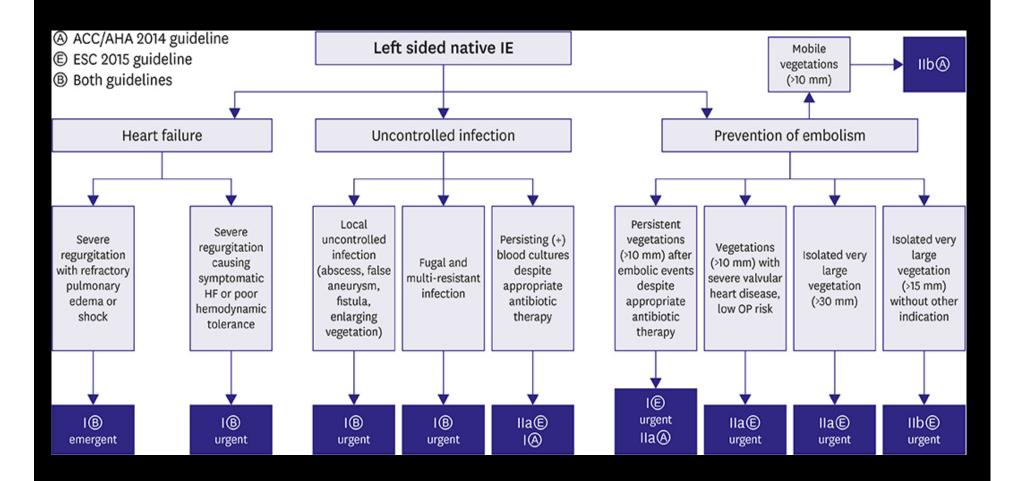




Indication for surgery



Indication for early surgery

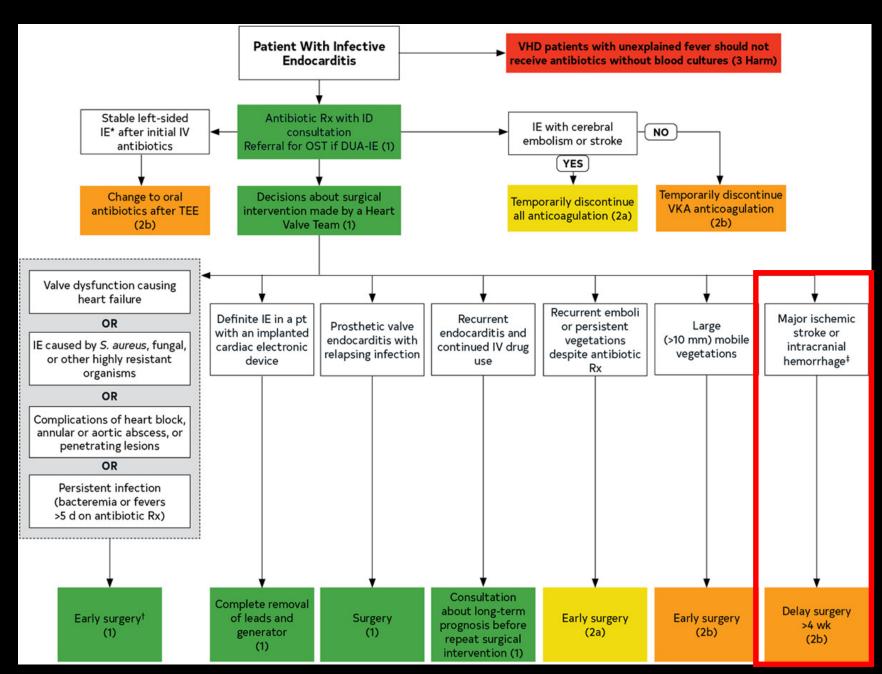


Early surgery in valvular heart disease KAMJE v48.(11)

Indication for early surgery

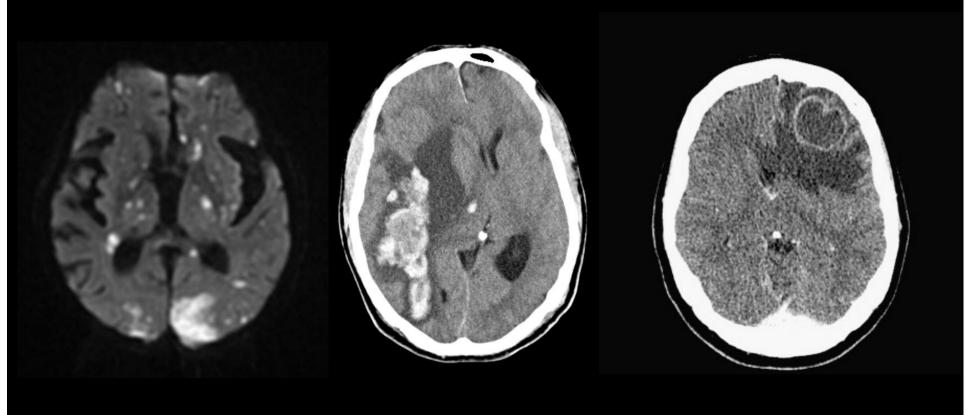
L-Sided NATIVE Valve IE	R-Sided NATIVE Valve IE	Prosthetic Valve IE
IE-associated valve dysfunction -> symptomatic left-sided heart failure. • Often MR or AR	Very large vegetations (>/= 20 mm diameter)	IE-associated symptomatic heart failure 2/2 valve dehiscence, intracardiac fistula, or severe prosthetic valve dysfunction.
Paravalvular extension of infection. Annular or Aortic Abscess Fistula Heart Block Destructive Penetrating Lesions	Recurrent septic pulmonary emboli	Paravalvular extension of infection. Annular or Aortic Abscess Fistula Heart Block Destructive Penetrating Lesions
Difficult-to-Treat Pathogens: Fungi: Candida, Aspergillus MDRO: Pseudomonas aeruginosa, Vancomycin- Resistant Enterococcus.	Presence of highly-resistant organism	 Difficult-to-Treat Pathogens: Fungi: Candida, Aspergillus MDRO: Pseudomonas aeruginosa, Vancomycin-Resistant Enterococcus.
Persistent Infection: • Persistent bacteremia or fever >/= 7 days after initiation of appropriate abx therapy, and no other etiology of infection identified.	Persistent bacteremia	Persistent Infection: Persistent bacteremia or fever >/= 7 days after initiation of appropriate abx therapy, and no other etiology of infection identified.
Recurrent emboli and persistent/enlarging vegetations despite appropriate abx.		Recurrent emboli despite appropriate abx / Relapsing PVE
Severe valve regurgitation and mobile vegetation >10 mm (esp. if anterior leaflet of mitral valve).		Mobile vegetations > 10 mm.

Surgical management of Infective endocarditis Circulation v132, issue 15, 1435-1486



2020 AHA guideline for valvular heart disease

Management of neurologic complication



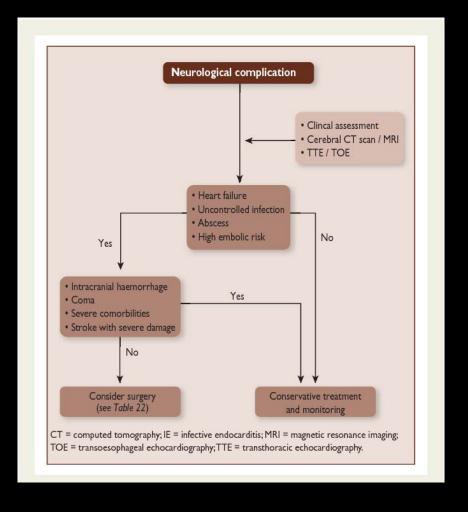
Cerebral infarct (stoke)

Cerebral hemorrhage (mycotic Aneurysm)

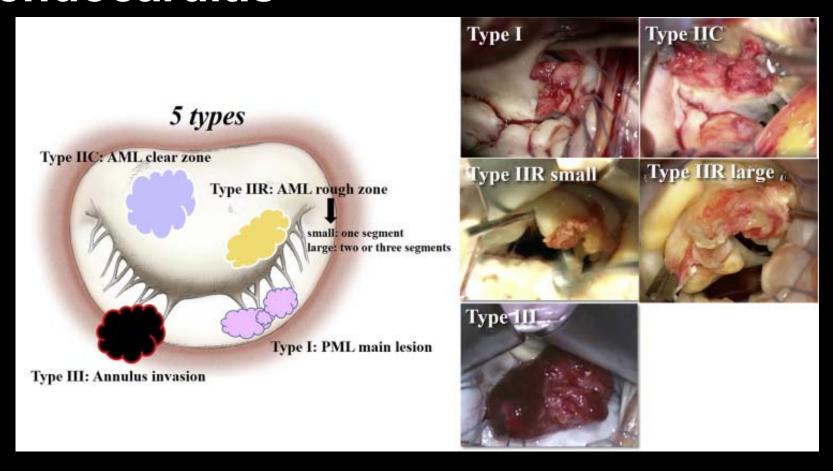
Brain abscess Meningitis

Management of neurologic complication

Recommendations	Classa	Level ^b	Ref. ^c
After a silent embolism or transient ischaemic attack, cardiac surgery, if indicated, is recommended without delay	ı	В	105, 263
Neurosurgery or endovascular therapy is recommended for very large, enlarging or ruptured intracranial infectious aneurysms	1	С	
Following intracranial haemorrhage, surgery should generally be postponed for ≥ 1 month	lla	В	264–266
After a stroke, surgery indicated for HF, uncontrolled infection, abscess, or persistent high embolic risk should be considered without any delay as long as coma is absent and the presence of cerebral haemorrhage has been excluded by cranial CT or MRI	lla	В	9,263
Intracranial infectious aneurysms should be looked for in patients with IE and neurological symptoms. CT or MR angiography should be considered for diagnosis. If non-invasive techniques are negative and the suspicion of intracranial aneurysm remains, conventional angiography should be considered	lla	В	267, 268

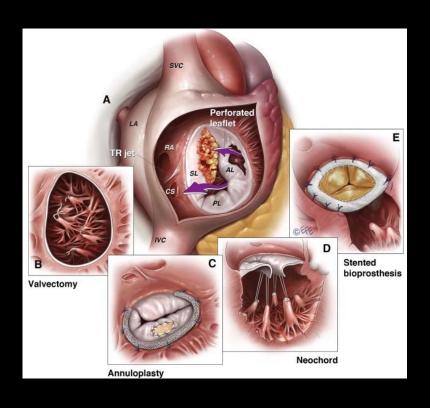


Various condition of infective endocarditis



Impact of lesion localization on durability of mitral valve in infective endocarditis, Ann of Thorac Surg 2020;109:1335-42

Goal of surgery



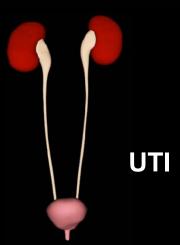
- Valve replacement
 - Mechanical valve
 - Tissue valve
 - Homograft
- Valve repair
- Patch closure etc.....
- 1. Infected source or material and Vegetation removal
- 2. Reconstruction of destroyed structure (ex. aortomitral continuity)

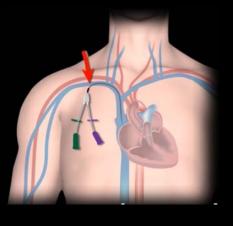


Source of Infective Endocarditis



Skin abscess





Contaminated central line





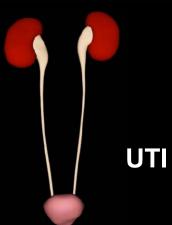
Dental procedures Or brushing

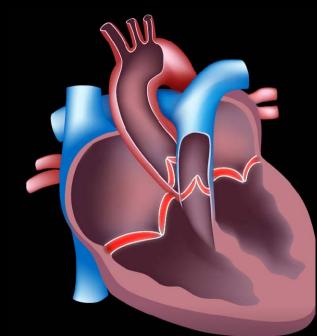
Drug injection or acupuncture

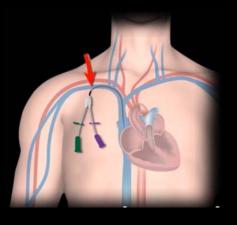
Source of Infective Endocarditis



Skin abscess







Contaminated central line







Dental procedures Or brushing (S. viridians)

Drug injection or acupuncture

Prophylactic antibiotic before Dental procedure







Who?

When?

How?

Prophylactic antibiotic before Dental procedure



- 1. Prosthetic cardiac valve or prosthetic material used for cardiac valve repair or other implantable cardiac devices
- 1. Recurrent or relapse IE
- 1. Congenital heart disease
- 1. Cardiac transplantation

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High risk population (Indication)

Table 3. AP for a Dental Procedure: Underlying Conditions for Which AP Is Suggested

Prosthetic cardiac valve or material
Presence of cardiac prosthetic valve
Transcatheter implantation of prosthetic valves
Cardiac valve repair with devices, including annuloplasty, rings, or clips
Left ventricular assist devices or implantable heart
Previous, relapse, or recurrent IE
CHD
Unrepaired cyanotic congenital CHD, including palliative shunts and conduits.
Completely repaired congenital heart defect with prosthetic material or device, whether placed by surgery or by transcatheter during the first 6 mo after the procedure
Repaired CHD with residual defects at the site of or adjacent to the site of a prosthetic patch or prosthetic device
Surgical or transcatheter pulmonary artery valve or conduit placement

such as Melody valve and Contegra conduit

Cardiac transplant recipients who develop cardiac valvulopathy

AP for a dental procedure not suggested

Implantable electronic devices such as a pacemaker or similar devices

Septal defect closure devices when complete closure is achieved

Peripheral vascular grafts and patches, including those used for hemodialysis

Coronary artery stents or other vascular stents

CNS ventriculoatrial shunts

Vena cava filters

Pledgets

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Prophylactic antibiotic before Dental procedure



Table 4. Dental Procedures and AP

AP suggested

All dental procedures that involve manipulation of gingival tissue or the periapical region of teeth or perforation of the oral mucosa

AP not suggested

Anesthetic injections through noninfected tissue, taking dental radiographs, placement of removable prosthodontic or orthodontic appliances, adjustment of orthodontic appliances, placement of orthodontic brackets, shedding of primary teeth, and bleeding from trauma to the lips or oral mucosa

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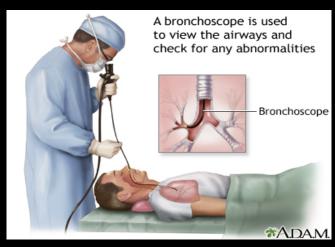
Prophylactic antibiotic before Dental procedure



Situation	Agent	Adults	Children
Oral	Amoxicillin	2 g	50 mg/kg
Unable to take oral medication	Ampicillin OR	2 g IM or IV	50 mg/kg IM or IV
	Cefazolin or ceftriaxone	1 g IM or IV	50 mg/kg IM or IV
Allergic to penicillin or	Cephalexin*† OR	2 g	50 mg/kg
ampicillin—oral	Azithromycin or clarithromycin OR	500 mg	15 mg/kg
	Doxycycline	100 mg	<45 kg, 2.2 mg/kg >45 kg, 100 mg
Allergic to penicillin or ampicillin and unable to take oral medication	Cefazolin or ceftriaxone†	1 g IM or IV	50 mg/kg IM or IV

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Prophylactic antibiotic is not recommended situation



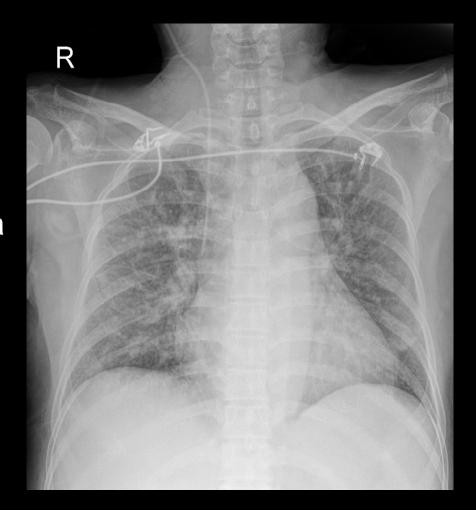




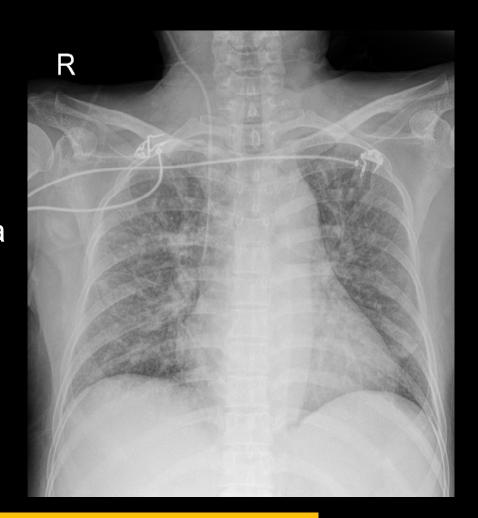




- F/60y
- Previous medial history (-)
- Fever, DOE, pitting edema
- COVID-19 (-)
- Blood culture:
 - Gram positive cocci(+)



- F/60y
- Previous medial history (-)
- Fever, DOE, pitting edema
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Heart failure 동반한 fever + blood infection (+) Infective endocarditis 의심

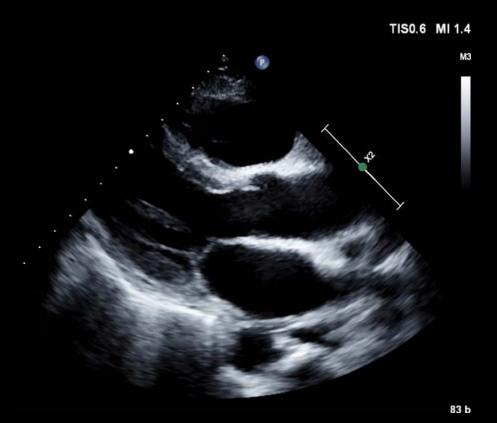


MAJOR CRITERIA	minor criteria
Blood culture positive	1. Predisposing factor
-	2. Temperature >38 degree
Evidence of endocardial	3. Vascular phenomena
involvement	4. Immunologic phenomena
	5. Microbiologic evidence



Infective endocarditis

Vegetation at native MV (2.0*1.8cm, hypermobile)



MAJOR CRITERIA			minor criteria
1,	Blood culture positive	1. 2.	Predisposing factor Temperature >38 degree
1.	Evidence of endocardial involvement	3. 4. 5.	Vascular phenomena Immunologic phenomena Microbiologic evidence



Infective endocarditis

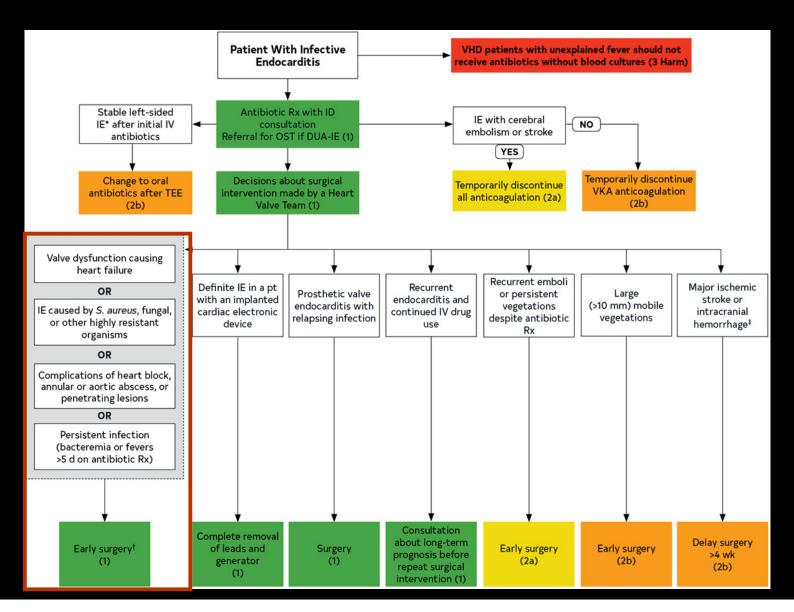


Heart failure Embolic risk

Vegetation at native MV (2.0*1.8cm, hypermobile)

Early surgery (MVR)

Case I I





Vegetation at native MV (2.0*1.8cm, hypermobile)

MAJOR CRITERIA	minor criteria
1. Blood culture positive	 Predisposing factor Temperature >38 degree
Evidence of endocardial involvement	 Vascular phenomena Immunologic phenomena Microbiologic evidence



Infective endocarditis



Case I I

- M/60y
- s/p AVR (2009)
- Bacterial meningitis
- DOE, Pitting edema
- Blood culture:
 - E. faeicum (+)

High risk of predisposing factor

Meningitis

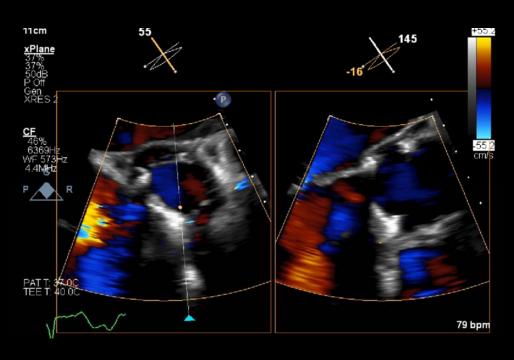
Heart failure 증상

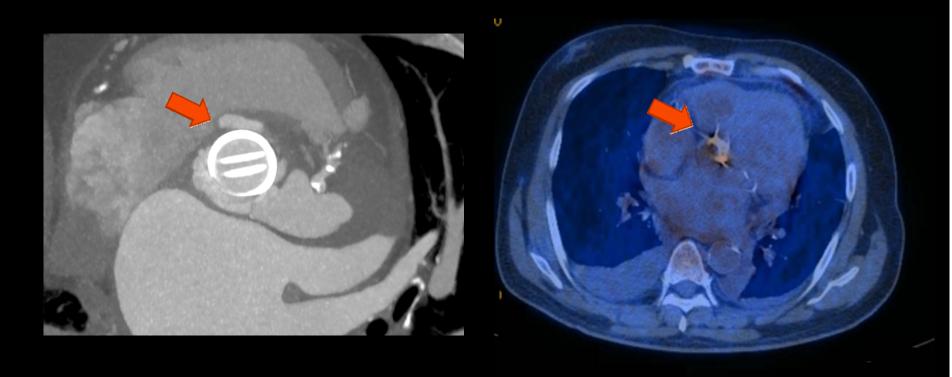
Blood infection (+)

Prosthetic infective endocarditis 의심

- M/60y
- s/p AVR (2009)
- Bacterial meningitis
- DOE, Pitting edema
- Blood culture:
 - E. faeicum (+)



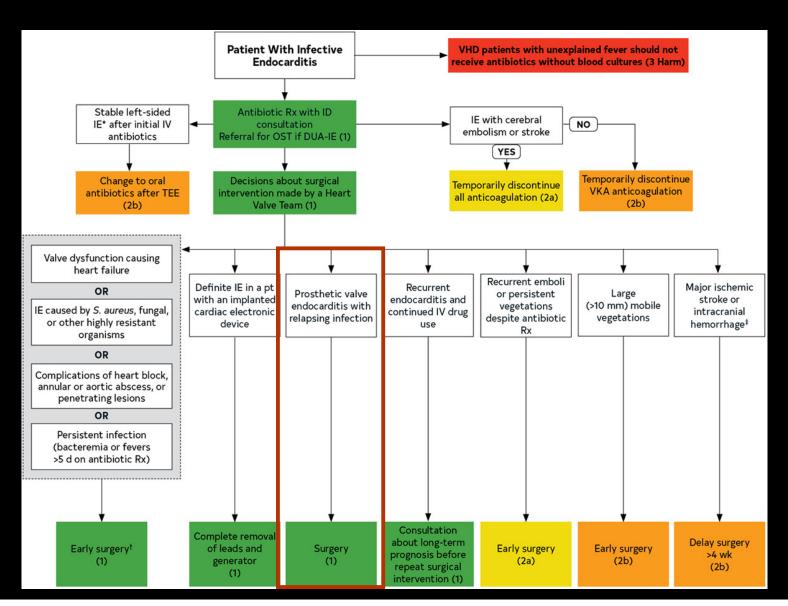




Cardia CT : Pseudoaneurysm r/o dehiscence

PET-CT : FDG uptake around prosthetic valve (SUV:3.7)

Prosthetic valve endocarditis with dehiscence



Summary

Fever with heart failure symptom Suspicious Embolic event (stroke, peripheral spot) Predisposing risk factor (+) Blood culture Diagnosis Echocardiography (TTE, TEE), Cardiac CT, PET-CT Antibiotics therapy (4-6weeks) Surgery (Heart failure, Uncontrolled infection, Management Prevent of emboli) Neurologic evaluation (huge hemorrhage, large stroke) High risk patients (prosthetic material, CHD, HT) Prophylaxis Dental procedure Amoxicillin or IV cepha before 1-2hrs

