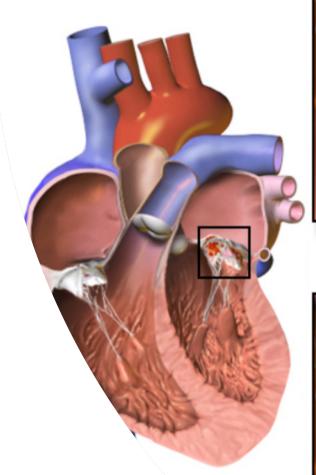
#### Infective Endocarditis

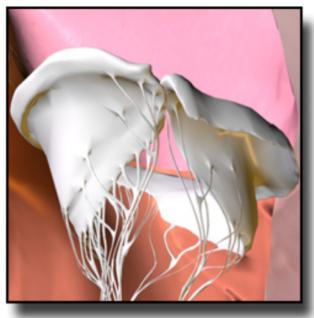
#### Nouf Alanazi

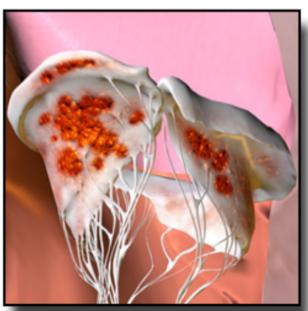
Assistant Professor & Consultant Interventional Cardiologist

King Saud University

Oct 1st 2019







#### Objectives

- Understand Infective Endocarditis definition
- Pathophysiology of endocarditis
- Diagnostic criteria of infective endocarditis
- Recognize the risk factors, signs, and symptoms of infectious endocarditis.
- Anticipate possible complications of infective endocarditis
- Treatment of endocarditis and appreciation of the necessity of rapid treatment.
- Endocarditis prophylaxis

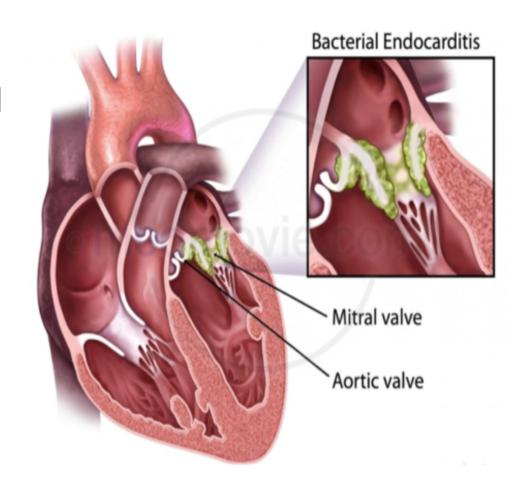
### Agenda

- What is IE?
- Epidemiology
- Microbiology
- Pathogensis
- Risk Factors
- Clinical presentation
- Diagnostic criteria
- Complications
- Investigations
- Management
- Prophylaxis



#### What is infective Endocarditis?

- An infection of the endocardial surface of the heart, which may include one or more heart valves (native or prosthetic), the mural endocardium, a septal defect or an intracardiac device.
- This leads to formation of bulky friable vegetations composed of thrombotic debris and organisms.
- Often associated with destruction of the underlying cardiac tissue



### Epidemiology

Heavily influenced by the epidemiology of the infection.

 Developing countries (endemic RF), Subacute course, viridans group streptococci.

• Developed countries, acute illness, *Staphylococcus aureus*, with numerous anatomic sites of metastatic foci of infection and worse outcomes.

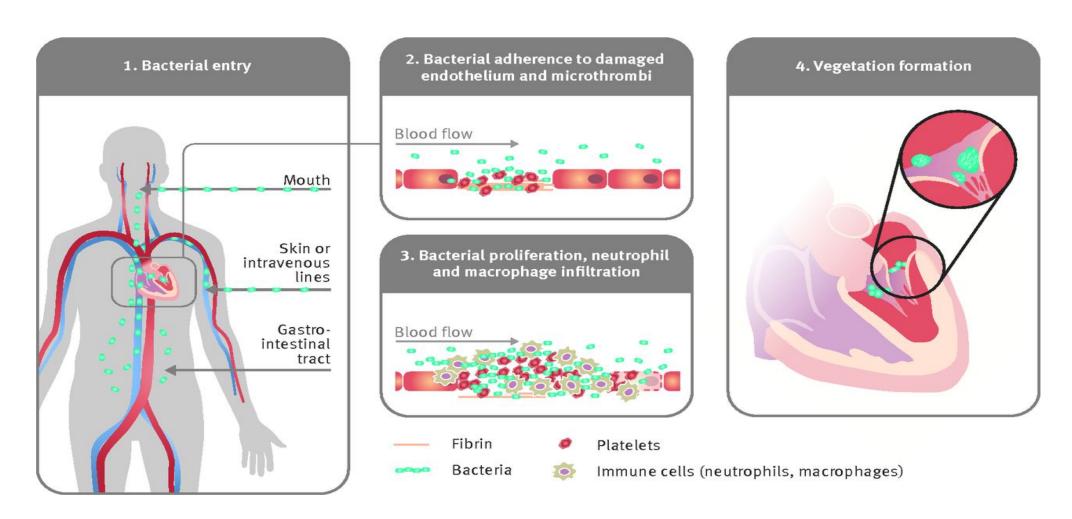
### Epidemiology

• IE develops most commonly on the mitral valve, closely followed in descending order of frequency by the aortic valve, combined mitral & aortic valve, tricuspid valve & rarely, the pulmonic valve.

 Mechanical prosthetic & bioprosthetic valves exhibit equal rates of infection.

More common in males

#### Pathogenesis of infective endocarditis



#### **Risk Factors**

Patient Factors	Comorbid Conditions
• Age >60 years	Structural heart disease
Male sex	Valvular disease
Injection drug use	Congenital heart disease
Poor dentition or dental infection	<ul> <li>Prosthetic heart valve(s)/TAVR</li> </ul>
	History of infective endocarditis
	Intravascular device
	Cardiac implantable electronic device
	Chronic hemodialysis
	HIV infection

#### Microbiology

- Vary depending on the population e.g IV drug abuser, pts with PHV, hospital acquired vs community acquired.
- The three most common causes of IE worldwide are staphylococci, streptococci & enterococci.
- In the US & most developed countries, *Staph. aureus* is the most common cause of IE
- Staphylococcal IE is a common cause of health care-associated IE
- Streptococcal IE is a common cause of community-acquired IE.

# Microbiology

Staph aureus (Including MRSA)	<ul> <li>Aggressive acute IE</li> <li>Metastatic infection, valve destruction</li> <li>Mortality 25-40% (left heart)</li> </ul>
Coagulase negative staph e.g (staph. Epidermidis)	<ul><li>Foreign body infection/prosthesis</li><li>Nosocomial infection</li></ul>
Strep. bovis (gallolyticus)	<ul> <li>GI flora associated with polyps and colon cancer</li> <li>Subacute endocarditis</li> <li>Highly sensitive to penicillin</li> </ul>
Beta- hemolytic stept group (A-B-C-G)	Frequent intracardiac & extracardiac complications, abcesses
Enterococci (fecalis, faecium)	<ul> <li>GI flora</li> <li>Associated with UTI/ nosocomial infection</li> </ul>

# Microbiology

<ul> <li>HACEK:</li> <li>Hemophilus sp</li> <li>Aggregatibacter sp ( actinobacillus)</li> <li>Cardiobacterium hominis</li> <li>Eikenella corrodens</li> <li>Kingella sp.</li> </ul>	<ul> <li>Fastidious gram-negative bacilli</li> <li>URTI – oropharyngeal flora</li> <li>Positive blood culture after 5 days of incubation &amp; maybe longer</li> <li>Large vegetation</li> </ul>
Other micro-organisms (can cause culture negative endocarditis)	<ul> <li>Coxiella burnetti (Q fever): subacute endocarditis, elevate Igg titer</li> <li>Bartonella: culture negative endocardiris (perform serology or specific culture technique), cat scratch disease</li> <li>Fungi (candida) risk factors immunosuppression, prosthesis, central line, IVDA, invasive endocarditis</li> <li>Others: Brucella, tropheryma whipplei, Mycoplasma, legionella</li> </ul>

# Epidemiological Clues That May be Helpful in Defining the Etiological Dx of CNE

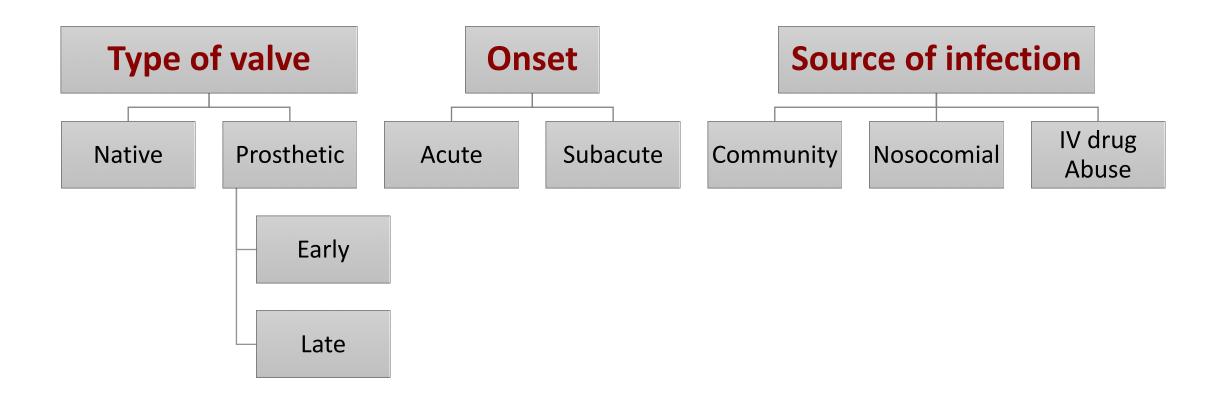
Epidemiological Feature	Common Microorganism	Alcoholism, cirrhosis	Bartonella sp
IDU	S aureus, including community-acquired		<i>Aeromonas</i> sp
	oxacillin-resistant strains		<i>Listeria</i> sp
	Coagulase-negative staphylococci		S pneumoniae
	β-Hemolytic streptococci		β-Hemolytic streptococci
	Fungi	Burn	S aureus
	Aerobic Gram-negative bacilli, including Pseudomonas aeruginosa		Aerobic Gram-negative bacilli, including
	Polymicrobial		P aeruginosa
Indwelling cardiovascular medical	S aureus		Fungi
devices	Coagulase-negative staphylococci	Diabetes mellitus	S aureus
	Fungi		β-Hemolytic streptococci
	Aerobic Gram-negative bacilli		S pneumoniae
	Corynebacterium sp	Early (≤1 y) prosthetic valve	Coagulase-negative staphylococci
Genitourinary disorders, infection,	Enterococcus sp	placement	S aureus
and manipulation, including pregnancy, delivery, and abortion	Group B streptococci (S agalactiae)		Aerobic Gram-negative bacilli
pregnancy, delivery, and abortion	Listeria monocytogenes		Fungi
	Aerobic Gram-negative bacilli		Corynebacterium sp
	Neisseria gonorrhoeae		<i>Legionella</i> sp
Chronic skin disorders, including	S aureus	Late (>1 y) prosthetic valve	Coagulase-negative staphylococci
recurrent infections	β-Hemolytic streptococci	placement	
Poor dental health, dental procedures	VGS		S aureus
	Nutritionally variant streptococci		Viridans group streptococci
	Abiotrophia defectiva		Enterococcus species
	<i>Granulicatella</i> sp		Fungi
	Gemella sp		Corynebacterium sp
	HACEK organisms		(Continued)

# Epidemiological Clues That May be Helpful in Defining the Etiological Dx of CNE

Epidemiological Feature	Common Microorganism
Dog or cat exposure	Bartonella sp
	Pasteurella sp
	Capnocytophaga sp
Contact with contaminated milk or	
infected farm animals	<i>Brucella</i> sp
	Coxiella burnetii
	<i>Erysipelothrix</i> sp
Homeless, body lice	Bartonella sp
AIDS	Salmonella sp
	S pneumoniae
	S aureus
Pneumonia, meningitis	S pneumoniae
Solid organ transplantation	S aureus
	Aspergillus fumigatus
	Enterococcus sp
	<i>Candida</i> sp
Gastrointestinal lesions	S gallolyticus (bovis)
	Enterococcus sp
	Clostridium septicum

HACEK indicates *Haemophilus* species, *Aggregatibacter* species, *Cardiobacterium hominis*, *Eikenella corrodens*, and *Kingella* species; IDU, injection drug use; and VGS, viridans group streptococci.

### Types of infective endocarditis



# Symptoms

SYMPTOM	PATIENTS AFFECTED (%)
Fever	80-95
Chills	40-70
Weakness	40-50
Malaise	20-40
Sweats	20-40
Anorexia	20-40
Headache	20-40
Dyspnea	20-40
Cough	20-30
Weight loss	20-30
Myalgia/arthralgia	10-30
Stroke	10-20
Confusion/delirium	10-20
Nausea/vomiting	10-20
Edema	5-15
Chest pain	5-15
Abdominal pain	5-15
Hemoptysis	5-10
Back pain	5-10

# Signs

FINDING	PATIENTS AFFECTED (%)
Fever	80-90
Heart murmur	75-85
New murmur	10-50
Changing murmur	5-20
Central neurologic abnormality	20-40
Splenomegaly	10-40
Petechiae/conjunctival hemorrhage	10-40
Splinter hemorrhages	5-15
Janeway lesions	5-10
Osler nodes	3-10
Retinal lesion or Roth spot	2-10

### Janeway Lesions

#### Nontender maculae on the palms and soles





# Janeway Lesions





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### Subungual (splinter) hemorrhages

Dark-red, linear lesions in the nail beds



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#### Osler nodes

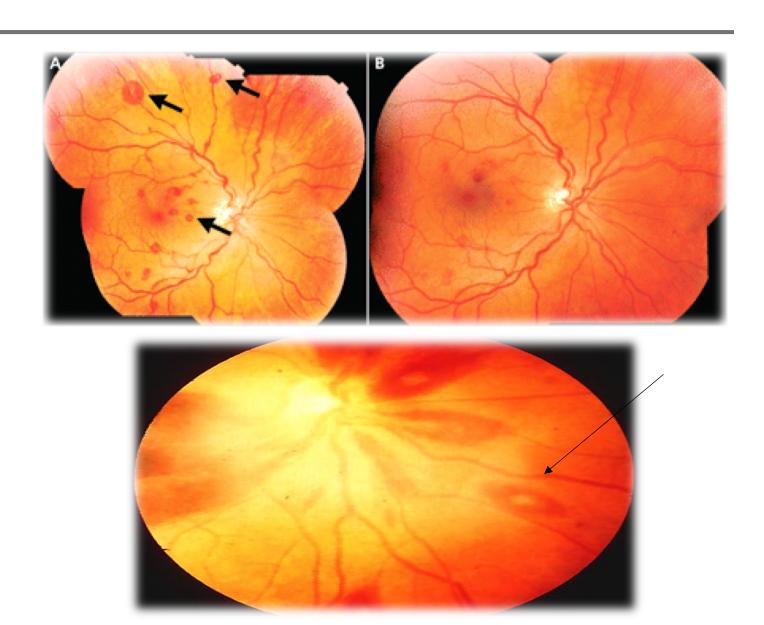
Tender subcutaneous nodules usually found on the distal pads of the digits



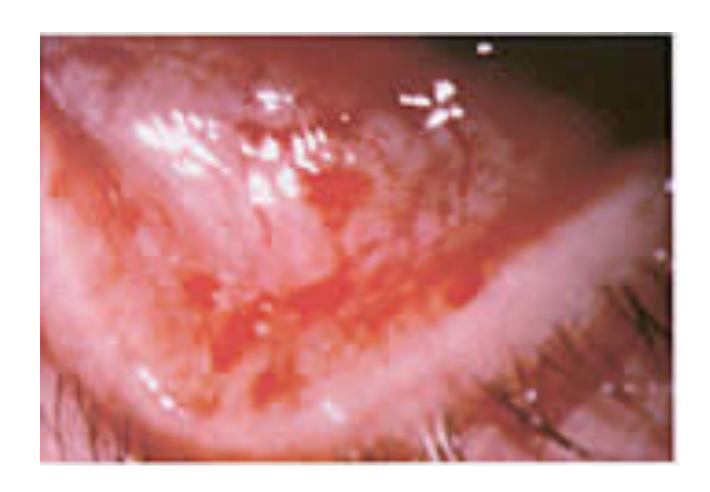
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## Roth Spots

Retinal red spots with pale (white) center



### Subconjunctival Hemorrhages



#### Petechia

Nonblanching, pinpoint reddish brown macules



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#### Other signs of IE

- Splenomegaly
- Delirium
- Conjunctival hemorrhage
- Pallor
- Cardiac arrhythmia
- Pericardial rub
- Pleural friction rub

#### **IVDA IE**

- Skin is the predominant source of infection
- Most commonly affects the TV.
- 70 100% of Rt. sided IE results in pneumonia & septic emboli
- Microbiology

Staph aureus ~60%

Streptococci and Enterococci ~20%

– Gram -ve bacilli ~10%

Fungi (Candida and Aspergillus ~5%

#### **Prosthetic Valve Endocarditis**

Early (<12 months)	Late (> 12 months)
<ul> <li>Reflects perioperative contamination</li> <li>Incidence around 1%</li> <li>Microbiology: <ul> <li>✓ Staph (45 - 50%)</li> <li>○ Staph. Epiderm (~ 30%)</li> <li>○ Staph. Aureus (~ 20%)</li> <li>✓ Gram -ve aerobes (~20%)</li> <li>✓ Fungi (~ 10%)</li> <li>✓ Strep and Entero (5-10%)</li> </ul> </li> </ul>	<ul> <li>After endothelialization</li> <li>Incidence 0.2 -0.5 % / pt. year</li> <li>Transient bacteraemia from dental, GI or GU</li> <li>Microbiology:         resembles native valve endocarditis</li> </ul>

## Diagnosis

#### Modified Duke criteria for diagnosis of infective endocarditis - Table A

Definite IE is establishe	d in the presence of any of the following:
Pathologic criteria	
Pathologic lesions: vegetation	on or intracardiac abscess demonstrating active endocarditis on histology <b>OR</b>
Microorganism: demonstrate	ed by culture or histology of a vegetation or intracardiac abscess
Clinical criteria	
Using specific definitions list	ed in <b>Table B:</b>
2 major clinical criteria OR	
1 major and 3 minor clinica	I criteria OR
5 minor clinical criteria	
Possible IE*	
Presence of 1 major and 1 n	ninor clinical criteria <b>OR</b> presence of 3 minor clinical criteria
Rejected IE	
A firm alternate diagnosis is	made <b>OR</b>
Resolution of clinical manife	stations occurs after ≤4 days of antibiotic therapy <b>OR</b>
No pathologic evidence of in	fective endocarditis is found at surgery or autopsy after antibiotic therapy for four days or less
Clinical criteria for possible o	or definite IE not met

#### Modified Duke criteria for diagnosis of infective endocarditis - Table B

ajor criteria	
Positive blood cultures for IE	(one of the following):
Typical microorganisms o	onsistent with IE from two separate blood cultures:
Staphylococcus aureus	
Viridans streptococci	
Streptococcus gallolyticus	(formerly S. bovis), including nutritional variant strains (Granulicatella spp and Abiotrophia defectiva)
HACEK group: Haemophi	lus spp, Aggregatibacter (formerly Actinobacillus actinomycete comitants), Cardiobacterium hominis, Eikenella spp, and Kingella kingae
Community-acquired ent	erococci, in the absence of a primary focus; <b>OR</b>
Persistently positive bloo	d culture:
For organisms that are ty	pical causes of IE: At least two positive blood cultures from blood samples drawn >12 hours apart
For organisms that are m	ore commonly skin contaminants: Three or a majority of ≥4 separate blood cultures (with first and last drawn at least one hour apart)
Single positive blood cult	ure for <i>Coxiella burnetii</i> or phase I IgG antibody titer >1:800*
Evidence of endocardial invo	vement (one of the following):
Echocardiogram positive	for IE:
Vegetation (oscillating intantomic explanation) <b>O</b>	racardiac mass on a valve or on supporting structures, in the path of regurgitant jets, or on implanted material, in the absence of an alternative
Abscess OR	
New partial dehiscence o	prosthetic valve
New valvular regurgitation	n
Increase in or change in	preexisting murmur not sufficient

#### Minor criteria

Predisposition: Intravenous drug use or presence of a predisposing heart condition (prosthetic heart valve or a valve lesion associated with significant regurgitation or turbulence of blood flow)

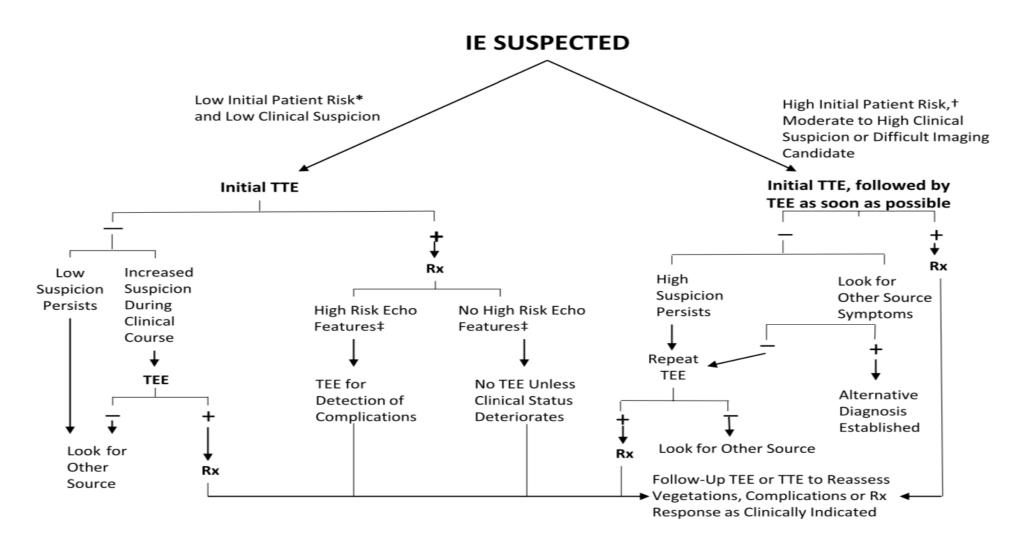
Fever: Temperature ≥38.0°C (100.4°F)

Vascular phenomena: Major arterial emboli, septic pulmonary infarcts, mycotic aneurysm, intracranial hemorrhage, conjunctival hemorrhages, or Janeway lesions

Immunologic phenomena: Glomerulonephritis, Osler nodes, Roth spots, or rheumatoid factor

Microbiologic evidence: Positive blood cultures that do not meet major criteria OR serologic evidence of active infection with organism consistent with IE

#### An approach to the diagnostic use of echo



### Complications

- 1. Embolic
- 2. Local spread
- 3. Metastatic spread of infection
- 4. Formation of immune complexes

#### 1. Embolic complications

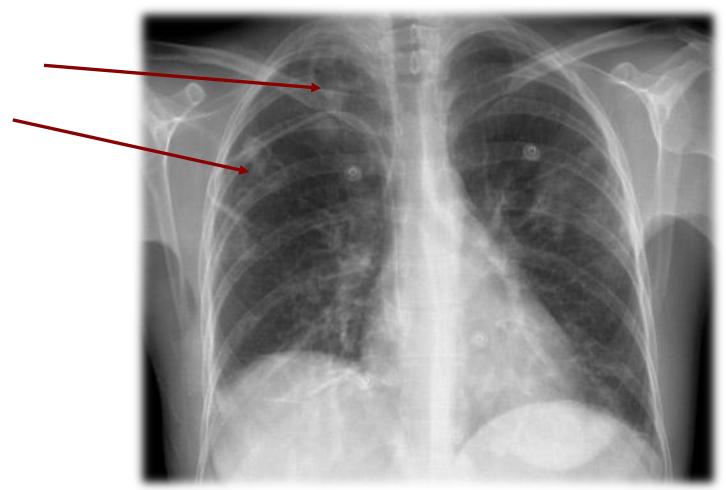
- Stroke: can either be embolic or due to ICH from a ruptured mycotic aneurysm or hemorrhagic transformation of stroke.
- MI
- Ischemic limb
- PE
- Splenic or renal infarction
- Digital infarcts
- Mesenteric ischemia

#### Uncommon after 2 weeks of effective treatment

#### **Risk Factors:**

- ✓ Size of vegetation (>10mm)
- ✓ Left side vegetations
- ✓ Staph. aureus
- ✓ Fungal pathogens

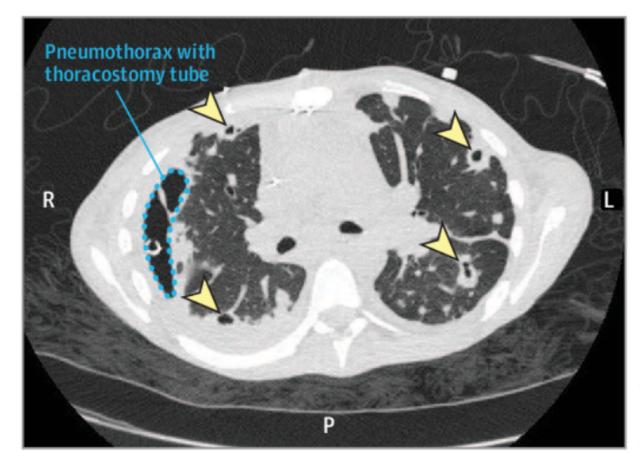
# Septic Emboli



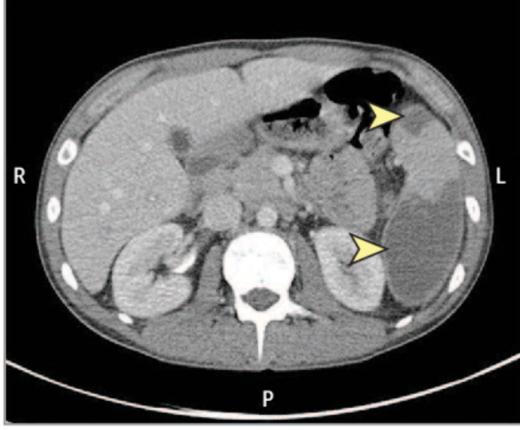
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### Septic Emboli

Pulmonary cavitation



**B** Splenic infarct



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#### 2. Local spread

#### **Heart failure:**

 Extensive valvular destruction, ruptured chordae tendinea, fistulas, valve obstruction.

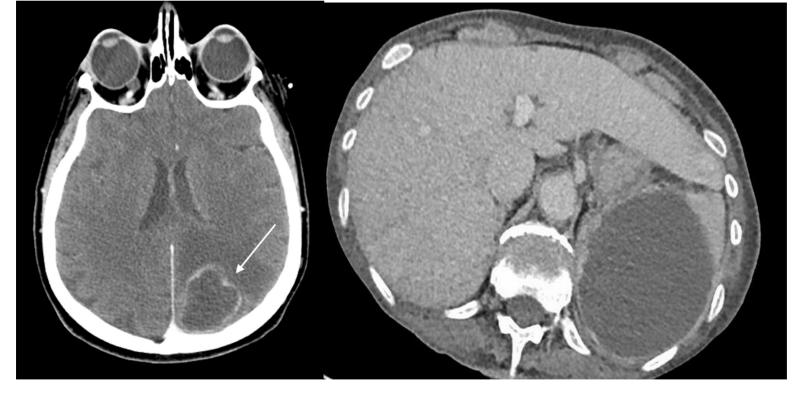
#### Paravalvular abscess:

- Most common in Aortic valve, IVDA, staph. Aureus
- AV block/conduction disorders

#### **Pericarditis**

### 3. Metastatic spread

- Meningitis
- Vertebral osteomyelitis
- Metastatic abscesses
- Septic arthritis



**Brain abscess** 

**Splenic abscess** 

# 4. Formation of immune complexes

- Immune complex glomerunephritis leading to ARF
- Arthritis

# Investigations

#### **Echocardiography:**

- TTE: sensitivity 70%
- TEE: sensitivity 96%, especially with prosthetic endocarditis, intracardiac devices or complications.

#### **Blood cultures:**

- Key diagnostic investigation.
- At least 3 sets of samples taken from different venipuncture sites over 24h under a meticulous aseptic technique.

# Investigations

- CBC (anemia, leukocytosis)
- U/E (Glomerulonephritis)
- LFT
- Coagulation panel.
- Inflammatory markers (CRP, ESR), CRP can help in monitoring response to therapy
- Urinalysis (proteinuria/ hematuria).
- Serology for culture negative IE (organisms that don't grow in cultures e.g Coxiella, Legionella, Bartonella).

# Investigations

- Investigations that will aid in detecting complications:
  - ✓ ECG
  - ✓ CXR
  - ✓ CT brain
  - ✓ MRI

# Management

#### **Antimicrobial Therapy:**

- Requires identification of specific pathogen & its susceptibility to antimicrobials.
- Prolonged administration of intravenous antibiotics remains the mainstay of treatment for IE.
- Empirical therapy should be started as soon as possible targeting the most likely pathogen.
- Bactericidal drugs should be used.
- Resolution of fever occurs in 5-7 days (persistence of fever indicates possible complication e.g abscess, embolization etc), drug reaction
- ID consult

# Therapy of NVE Caused by Highly Penicillin-Susceptible VGS & Streptococcus gallolyticus (bovis)

Regimen	Dose* and Route	Duration, wk	Strength of Recommendation	Comments	
Aqueous crystalline penicillin G sodium	12–18 million U/24 h IV either continuously or in 4 or 6 equally divided doses	4	Class IIa; Level of Evidence B	Preferred in most patients >65 y or patients with impairment of eighth cranial nerve function or renal function.	
Or				Ampicillin 2 g IV every 4 h is a reasonable alternative to penicillin if a penicillin shortage exists.	
Ceftriaxone sodium	2 g/24 h IV/IM in 1 dose	4	Class IIa; Level of Evidence B	to personal transfer and analysis	
Aqueous crystalline penicillin G sodium Or	12–18 million U/24 h IV either continuously or in 6 equally divided doses	2	Class IIa; Level of Evidence B	2-wk regimen not intended for patients with know cardiac or extracardiac abscess or for those with creatinine clearance of <20 mL/min, impaired eight cranial nerve function, or <i>Abiotrophia, Granulicatel</i> or <i>Gemella</i> spp infection; gentamicin dose should adjusted to achieve peak serum concentration of 3 μg/mL and trough serum concentration of <1 μg/mL	
Ceftriaxone sodium Plus	2 g/24 h IV or IM in 1 dose	2	Class IIa; Level of Evidence B		
Gentamicin sulfate‡	3 mg/kg per 24 h IV or IM in 1 dose	2		when 3 divided doses are used; there are no optimal drug concentrations for single daily dosing.†	
Vancomycin hydrochloride§	30 mg/kg per 24 h IV in 2 equally divided doses	4	Class IIa; Level of Evidence B	Vancomycin therapy is reasonable only for patients unable to tolerate penicillin or ceftriaxone; vancomycin dose should be adjusted to a trough concentration range of 10–15 µg/mL.	

# Therapy for NVE Caused by Staphylococci

Evidence C   For uncomplicated right-sided IE, 2 wk (see text For penicillin-allergic (nonanaphylactoid type) patients   Cefazolin*   6 g/24 h IV in 3 equally divided doses   6   Class I; Level of Evidence B   Evidence B   anaphylactoid-type hypersensitivity to penicillin.	Regimen	Dose* and Route	Duration, wk	Strength of Recommendation	Comments	
For penicillin-allergic (nonanaphylactoid type) patients  Cefazolin*  6 g/24 h IV in 3 equally divided doses  Oxacillin-resistant strains  Vancomycin§  30 mg/kg per 24 h IV in 2 equally divided doses  Daptomycin  Daptomycin  Evidence C  Consider skin testing for oxacillin-susceptible staphylococci and questionable history of imm type hypersensitivity to penicillin.  Cefazolin*  6 g/24 h IV in 3 equally divided doses  6 Class I; Level of Evidence B  Adjust vancomycin dose to achieve trough concentration of 10–20 μg/mL (see text for vancomycin alternatives).  Daptomycin  ≥8 mg/kg/dose  6 Class Ilb; Level of Await additional study data to define optimal of the concentration of study data to define optimal of the c	Oxacillin-susceptible strains					
(nonanaphylactoid type) patients  Cefazolin* 6 g/24 h IV in 3 equally divided doses 6 Class I; Level of Evidence B  Nanaphylactoid-type hypersensitivity to penicillin.  Oxacillin-resistant strains  Vancomycin§ 30 mg/kg per 24 h IV in 2 equally divided doses  Class I; Level of Evidence C  Cephalosporins should be avoided in patients anaphylactoid-type hypersensitivity to β-lacta vancomycin should be used in these cases.  Chass I; Level of Evidence C  Class I; Level of Concentration of 10–20 μg/mL (see text for vancomycin alternatives).  Daptomycin  ≥8 mg/kg/dose 6 Class IIb; Level of Await additional study data to define optimal of the concentration o	Nafcillin or oxacillin	12 g/24 h IV in 4–6 equally divided doses	6	•	For complicated right-sided IE and for left-sided IE; for uncomplicated right-sided IE, 2 wk (see text).	
Evidence B anaphylactoid-type hypersensitivity to β-lactary vancomycin should be used in these cases.  Oxacillin-resistant strains  Vancomycin§ 30 mg/kg per 24 h IV in 2 equally divided doses  Oxacillin-resistant strains  Vancomycin§ 30 mg/kg per 24 h IV in 2 equally divided doses  Class I; Level of Evidence C concentration of 10–20 μg/mL (see text for vancomycin alternatives).  Daptomycin ≥8 mg/kg/dose  6 Class IIb; Level of Await additional study data to define optimal of the concentration of the concent	(nonanaphylactoid type)				Consider skin testing for oxacillin-susceptible staphylococci and questionable history of immediate-type hypersensitivity to penicillin.	
Vancomycin§       30 mg/kg per 24 h IV in 2 equally divided doses       6       Class I; Level of Evidence C       Adjust vancomycin dose to achieve trough concentration of 10–20 μg/mL (see text for vancomycin alternatives).         Daptomycin       ≥8 mg/kg/dose       6       Class IIb; Level of Await additional study data to define optimal of the concentration of 10–20 μg/mL (see text for vancomycin alternatives).	Cefazolin*	6 g/24 h IV in 3 equally divided doses	6	•	Cephalosporins should be avoided in patients with anaphylactoid-type hypersensitivity to $\beta$ -lactams; vancomycin should be used in these cases.	
doses  Evidence C concentration of 10–20 μg/mL (see text for vancomycin alternatives).  Daptomycin  ≥8 mg/kg/dose  6 Class Ilb; Level of Await additional study data to define optimal of the concentration of 10–20 μg/mL (see text for vancomycin alternatives).	Oxacillin-resistant strains					
	Vancomycin§		6	,	concentration of 10–20 μg/mL (see text for	
	Daptomycin	≥8 mg/kg/dose	6	Class Ilb; Level of Evidence B	Await additional study data to define optimal dosing.	

# Therapy IE Involving a Prosthetic Valve or Other Prosthetic Material Caused by & Streptococcus gallolyticus (bovis)

			Strength of		
Regimen	Dose* and Route	Duration, wk	Recommendation	Comments	
Penicillin-susceptible strain (≤0.12 µg/mL)					
Aqueous crystalline penicillin G sodium	24 million U/24 h IV either continuously or in 4–6 equally divided doses	6	Class IIa; Level of Evidence B	Penicillin or ceftriaxone together with gentamicin has not demonstrated superior cure rates compared with	
Or				monotherapy with penicillin or ceftriaxone for patients with highly susceptible strain; gentamicin therapy	
Ceftriaxone	2 g/24 h IV or IM in 1 dose	6	Class IIa; Level of Evidence B	should not be administered to patients with creatinine clearance <30 mL/min.	
With or without					
Gentamicin sulfate†	3 mg/kg per 24 h IV or IM in 1 dose	2		Ampicillin 2 g IV every 4 h is a reasonable alternative to penicillin if a penicillin shortage exists.	
Vancomycin hydrochloride‡	30 mg/kg per 24 h IV in 2 equally divided doses	6	Class IIa; Level of Evidence B	Vancomycin is reasonable only for patients unable to tolerate penicillin or ceftriaxone.	
Penicillin relatively or fully resistant strain (MIC >0.12 µg/mL)					
Aqueous crystalline penicillin sodium	24 million U/24 h IV either continuously or in 4–6 equally divided doses	6	Class IIa; Level of Evidence B	Ampicillin 2 g IV every 4 h is a reasonable alternative to penicillin if a penicillin shortage exists.	
Or					
Ceftriaxone	2 g/24 h IV/IM in 1 dose	6	Class Ila; Level of Evidence B		
Plus					
Gentamicin sulfate	3 mg/kg per 24 h IV/IM in 1 dose	6			
Vancomycin hydrochloride	30 mg/kg per 24 h IV in 2 equally divided doses	6	Class Ila; Level of Evidence B	Vancomycin is reasonable only for patients unable to tolerate penicillin or ceftriaxone.	

# Therapy of IE Involving a Prosthetic Valve or Other Prosthetic Material Caused by Staphylococci

Regimen	Dose* and Route	Duration, wk	Strength of Recommendation	Comments	
	Dose and noute	Duration, wk	necommendation	Comments	
Penicillin-susceptible strain (≤0.12 µg/mL)					
Aqueous crystalline penicillin G sodium	24 million U/24 h IV either continuously or in 4–6 equally divided doses	6	Class IIa; Level of Evidence B	not demonstrated superior cure rates compared with	
Or				monotherapy with penicillin or ceftriaxone for patients	
Ceftriaxone	2 g/24 h IV or IM in 1 dose	6	Class IIa; Level of Evidence B	with highly susceptible strain; gentamicin therapy should not be administered to patients with creating clearance <30 mL/min.	
With or without					
Gentamicin sulfate†	3 mg/kg per 24 h IV or IM in 1 dose	2		Ampicillin 2 g IV every 4 h is a reasonable alternative to penicillin if a penicillin shortage exists.	
Vancomycin hydrochloride‡	30 mg/kg per 24 h IV in 2 equally divided doses	6	Class IIa; Level of Evidence B	Vancomycin is reasonable only for patients unable to tolerate penicillin or ceftriaxone.	
Penicillin relatively or fully resistant strain (MIC >0.12 µg/mL)					
Aqueous crystalline penicillin sodium	24 million U/24 h IV either continuously or in 4–6 equally divided doses	6	Class IIa; Level of Evidence B	Ampicillin 2 g IV every 4 h is a reasonable alternative to penicillin if a penicillin shortage exists.	
Or					
Ceftriaxone	2 g/24 h IV/IM in 1 dose	6	Class Ila; Level of Evidence B		
Plus					
Gentamicin sulfate	3 mg/kg per 24 h IV/IM in 1 dose	6			
Vancomycin hydrochloride	30 mg/kg per 24 h IV in 2 equally divided doses	6	Class IIa; Level of Evidence B	Vancomycin is reasonable only for patients unable to tolerate penicillin or ceftriaxone.	

### Persistence of fever

- Abscess
- Septic embolization
- An extracardiac site of infection (native or prosthetic)
- Infected indwelling catheters or devices
- Inadequate antibiotic treatment of a resistant organism
- An adverse reaction to the antibiotic therapy itself.

### Indications of Early surgery with native valve IE

- ☐ Valve dysfunction resulting in symptoms or signs of heart failure.
- ☐ IE caused by fungi or highly resistant organisms.
- ☐ IE complicated by heart block, annular abscess, or destructive perforating lesions.
- ☐ Persistent infection (bacteremia or fever) lasting >5-7 days after the start of appropriate antimicrobial therapy, assuming other sources of infection or fever have been excluded.
- ☐ Recurrent emboli or persistent/enlarging vegetations despite appropriate antimicrobial therapy.

### Indications of Early surgery with prosthetic valve IE

- 1. Symptoms or signs of heart failure resulting from valve dehiscence, intracardiac fistula, or severe prosthetic dysfunction.
- 2. Persistent bacteremia >5-7 days after the start of appropriate antimicrobial therapy.
- 3. Prosthetic valve IE complicated by heart block, annular abscess, or destructive perforating lesions.
- 4. Prosthetic valve IE caused by fungi or highly resistant organisms.
- 5. Recurrent emboli despite appropriate antimicrobial therapy.

# Prophylaxis

# Cardiac Conditions Associated With the Highest Risk of Adverse Outcome From Endocarditis for Which Prophylaxis With Dental Procedures Is Reasonable

Prosthetic cardiac valve or prosthetic material used for cardiac valve repair Previous IE

Congenital heart disease (CHD)\*

Unrepaired cyanotic CHD, including palliative shunts and conduits

Completely repaired congenital heart defect with prosthetic material or device, whether placed by surgery or by catheter intervention, during the first 6 months after the procedure†

Repaired CHD with residual defects at the site or adjacent to the site of a prosthetic patch or prosthetic device (which inhibit endothelialization)

Cardiac transplantation recipients who develop cardiac valvulopathy

†Prophylaxis is reasonable because endothelialization of prosthetic material occurs within 6 months after the procedure.

<sup>\*</sup>Except for the conditions listed above, antibiotic prophylaxis is no longer recommended for any other form of CHD.

# Dental Procedures for Which Endocarditis Prophylaxis Is Reasonable

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

\*The following procedures and events do not need prophylaxis: routine 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 deciduous teeth, and bleeding from trauma to the lips or oral mucosa.

# Regimens for a Dental Procedure

		Regimen: Single Dose 30 to 60 min Before Procedure		
Situation	Agent	Adults	Children	
Oral	Amoxicillin	2 g	50 mg/kg	
Unable to take oral medication	Ampicillin	2 g IM or IV	50 mg/kg IM or IV	
	OR			
	Cefazolin or ceftriaxone	1 g IM or IV	50 mg/kg IM or IV	
Allergic to penicillins or ampicillin—oral	Cephalexin*† <b>OR</b>	2 g	50 mg/kg	
	Clindamycin <b>OR</b>	600 mg	20 mg/kg	
	Azithromycin or clarithromycin	500 mg	15 mg/kg	
Allergic to penicillins or ampicillin and unable to take oral medication	Cefazolin or ceftriaxone† OR	1 g IM or IV	50 mg/kg IM or IV	
	Clindamycin	600 mg IM or IV	20 mg/kg IM or IV	

IM indicates intramuscular; IV, intravenous.

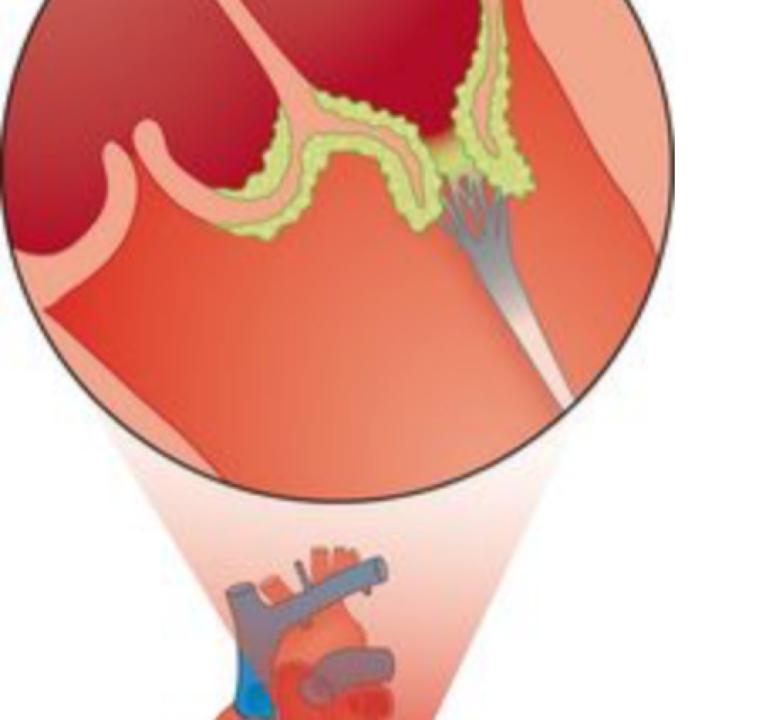
<sup>\*</sup>Or other first- or second-generation oral cephalosporin in equivalent adult or pediatric dosage.

<sup>†</sup>Cephalosporins should not be used in an individual with a history of anaphylaxis, angioedema, or urticaria with penicillins or ampicillin.

# Take home message

Infective endocarditis is a serious disease that requires a high index of clinical suspension and early management.

Recognize the exact pathophysiology, diagnostic criteria, treatment and indications of prophylaxis.



Thank You