

# VIÊM NỘI TÂM MẠC NHIỄM TRÙNG

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# Dịch tễ

- Mỹ 2009: 12.7 ca/ 100.000/năm  
50 năm trước: 2-4 ca/100.000 người/năm
- Tần suất mới mắc các nước khác: tương đương USA
- 50% b/n > 50 tuổi
- Nam > 3 nữ

# Phân loại

- Theo cơ địa người bệnh
  - VNTMNT trên van tự nhiên
  - VNTMNT trên van nhân tạo
  - VNTMNT trên người tiêm ma túy
- Theo tiến triển
  - VNTMNT cấp
  - VNTMNT bán cấp
- Theo tác nhân gây bệnh

TD : VNTMNT do staphylococcus aureus

# Tần suất VNTMNT theo bệnh tim chính ở Mỹ

Tổn thương	Trẻ < 2t %	Trẻ 2-5t %	Người lớn 15-50t %	Người lớn > 50t %	Người lớn tiêm ma túy %
Không bệnh tim	50-70	10-15	10-20	10	50-60
BTBS	30-50	70-80	25-35	15-25	10
Thấp tim	Hiếm	10	10-15	10-15	10
Bệnh tim thoái hoá	0	0	Hiếm	10-20	Hiếm
Tiền sử phẫu thuật tim	5	10-15	10-20	10-20	10-20
Tiền sử VNTMNT	Hiếm	5	5-10	5-10	10-20

BTBS: Bệnh tim bẩm sinh

VNTMNT: Viêm nội tâm mạc nhiễm trùng

TL : Hurst's The Heart, Mc Graw-Hill 10<sup>th</sup> ed 2001, 2089



# Nguy cơ tương đối VNTMNT thay đổi theo loại tổn thương tim

Nguy cơ cao	Nguy cơ trung bình	Nguy cơ rất thấp (không đáng kể)
Van nhân tạo	Sa van hai lá có hở	Sa 2 lá không hở
Tiền sử VNTMNT	Hẹp 2 lá	Hở van 3 lá nhẹ phát hiện bằng siêu âm
BTBS tím	Bệnh van 3 lá	Thông liên nhĩ lỗ thứ 2
Bệnh van ĐMC	Bệnh van ĐMP	Mảng xơ vữa ĐM
Hở van 2 lá	Phì đại vách thất không đối xứng	Bệnh ĐMV
Hở hẹp 2 lá	Đường truyền đến nhĩ phải	Viêm ĐMC giang mai
Còn ống động mạch	Cấy vật nhân tạo trong tim	Máy tạo nhịp
Hẹp eo ĐMC	(không là van)	
Thông liên thất	Bệnh van thoái hoá người cao tuổi	Tổn thương tim đã mổ (không cấy vật nhân tạo, trên 6 tháng sau mổ)

**Cardiac conditions at highest risk of infective endocarditis for which prophylaxis should be considered when a high-risk procedure is performed**

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>
<p>Antibiotic prophylaxis should be considered for patients at highest risk for IE</p> <p>(1) Patients with any <b>prosthetic valve</b>, including a <b>transcatheter valve</b>, or those in whom any prosthetic material was used for cardiac valve repair.</p> <p>(2) Patients with a previous episode of IE</p> <p>(3) Patients with CHD:</p> <p>(a) Any type of <b>cyanotic CHD</b></p> <p>(b) Any type of <b>CHD repaired</b> with a prosthetic material, whether placed surgically or by percutaneous techniques, up to 6 months after the procedure or lifelong if residual shunt or valvular regurgitation remains.</p>	IIa	C
<p>Antibiotic prophylaxis is <b>not recommended</b> in other forms of valvular or CHD.</p>	III	C

CHD = congenital heart disease; IE = infective endocarditis.

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

<sup>c</sup>Reference(s) supporting recommendations.

# Vi trùng gây bệnh/ VNTMNT ở người tiêm ma túy



Number of Cases (%) of Endocarditis in Drug Addicts\*

Organisms	Right-Sided N = 346	Left-Sided N = 204	Total N = 675	Spain (1977-1993) <sup>†</sup> N = 1529
Streptococci <sup>†</sup>	17 (5)	31 (15)	80 (12)	131 (8.5)
Enterococci	7 (2)	49 (24)	59 (9)	21 (1)
<i>Staphylococcus aureus</i>	267 (77)	47 (23)	396 (57)	1138 (74)
Coagulase-negative staphylococci	–	–	–	44 (3)
Gram-negative bacilli <sup>†</sup>	7 (5)	26 (13)	45 (7)	23 (1.5)
Fungi (predominantly <i>Candida</i> species)	–	25 (12)	26 (4)	18 (1)
Polymicrobia/miscellaneous	28 (8)	20 (10)	49 (7)	48 (3)
Culture negative	10 (3)	6 (3)	20 (3)	106 (7)

# Vi trùng gây bệnh trên VNTMNT van nhân tạo 1970- 2003

Organisms	Number of Cases (%) <sup>*</sup> with Time of Onset After Valve Surgery	
	Early (N = 218) <12 mo	Late (N = 272) >12 mo
Streptococci <sup>†</sup>	5 (2)	77 (28)
Pneumococci	—	—
Enterococci	20 (9)	36 (13)
<i>Staphylococcus aureus</i>	42 (19)	45 (17)
Coagulase-negative staphylococci	72 (33)	31 (11)
Fastidious gram-negative coccobacilli (HACEK group) <sup>‡</sup>	—	11 (4)
Gram-negative bacilli	34 (16)	17 (6)
Fungi, <i>Candida</i> species	16 (7)	5 (2)
Polymicrobial/miscellaneous	6 (3)	15 (6)
Diphtheroids	10 (5)	7 (3)
<i>Coxiella burnetii</i>	—	5 (2)
Culture negative	13 (6)	23 (8)



# VNTM do nhiễm trùng bệnh viện

- Đường vào của vi trùng
  - 75% do các đường mạch máu
  - Các đường khác: vết thương, chỗ đặt máy tạo nhịp, ống nội khí quản, ống thông tiểu...
- 2/3 b/n không có bệnh tim trước đó
- Vi trùng thường gặp: staphylococcus; Gram(-); enterococci; candida

# VNTM cây âm nghiệm: nguyên nhân (1)

- B/n đã sử dụng kháng sinh. TD: uống ampicillin
- Vi trùng thuộc loại mọc chậm. TD: nhóm HACEK, vài loại Streptococci
- Một số vi trùng cần môi trường cấy đặc biệt. TD: C. burnetti (sốt Q), Chlamydia, Mycoplasma, Bartonella, Legionella, vài loại Streptococci
- Vi trùng kỵ khí

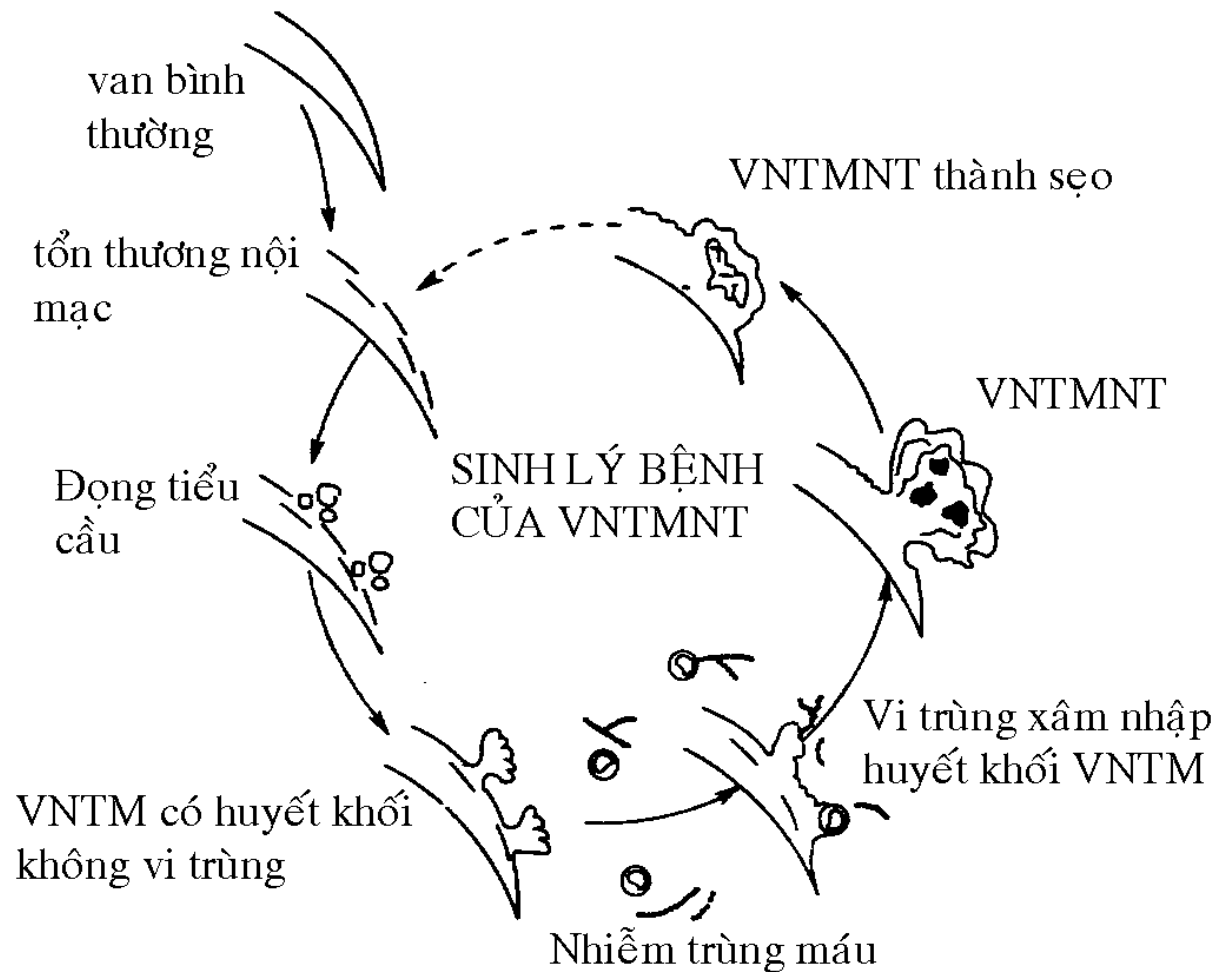
# VNTM cấy âm nghiệm: nguyên nhân (2)

- Aspergillus hoặc nấm khác
- Vi trùng không cấy được, chẩn đoán bằng PCR trên mô cơ tim khi phẫu thuật
- Bệnh giống VNTMNT. TD: thấp tim
- Bệnh VNTM huyết khối không có vi trùng/bn lao hoặc ung thư
- VNTM Libman-Sacks/Lupus ban đỏ

# Sinh bệnh học

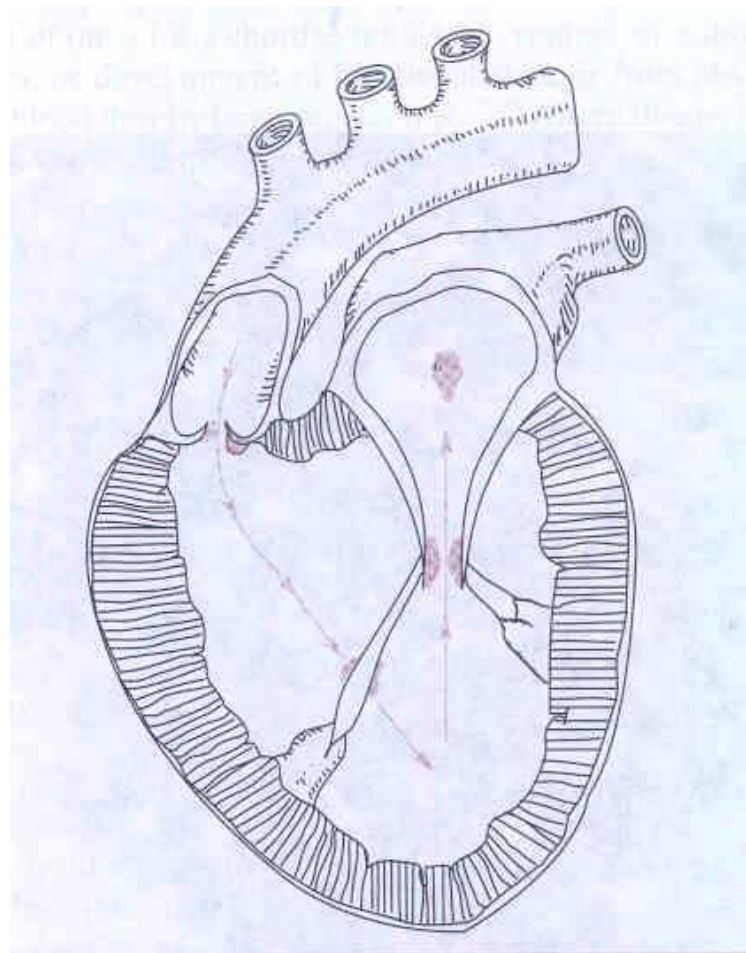
- Các điều kiện:
  - Tình trạng nội mạc
  - Sự lưu thông của máu
  - Sự hiện diện của vi trùng trong máu
- Hiệu quả Venturi
- Mảnh sùi vô trùng → mảnh sùi VNTMNT
- Cấu tạo mảnh sùi VNTMNT:
  - Vi trùng
  - Fibrin
  - Hồng cầu
  - Bạch cầu

# Các biến cố chính trong sinh bệnh học VNTMNT



TL : *Hurst's The Heart, Mc Graw-Hill 10<sup>th</sup> ed 2001, 2096*

# Vị trí mảnh sùi trên bệnh nhân hở van ĐMC và Hở 2 lá



# Triệu chứng cơ năng

- Trong vòng 2 tuần sau nhiễm bệnh
- Mệt mỏi
- Sốt nhẹ hay sốt cao
- Nhức khớp
- Viêm khớp



# Triệu chứng thực thể

- Âm thổi ở tim: mới xuất hiện hay biến đổi một âm thổi cũ  
Đôi khi: không âm thổi
- Lách
- Lấm tẩm xuất huyết dưới da (pétéchia)
- Mảnh vụn xuất huyết
- Vết Roth
- Nốt Osler
- Thương tổn Janeway
- Ngón tay dùi trống
- Dấu thuyên tắc: phổi, não, chi dưới, thận, bụng
- Biểu hiện thần kinh: do thuyên tắc hay áp xe não
- Suy tim



# Tần suất xuất hiện các triệu chứng cơ năng bệnh nhân VNTMNT

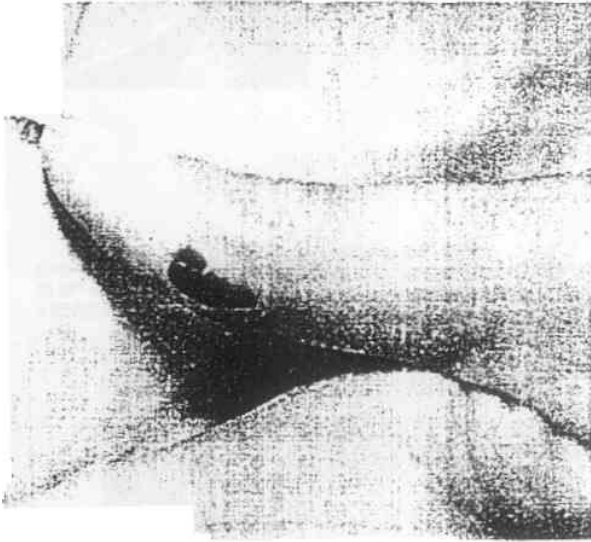
SYMPTOM	PATIENTS AFFECTED (%)
Fever	80-95
Chills	40-70
Weakness	40-50
Malaise	20-40
Sweats	20-40
Anorexia	20-40
Headache	20-40
Dyspnea	20-40
Cough	20-30
Weight loss	20-30
Myalgia/arthralgia	10-30
Stroke	10-20
Confusion/delirium	10-20
Nausea/vomiting	10-20
Edema	5-15
Chest pain	5-15
Abdominal pain	5-15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

TL: Baddour LM et al. Braunwald's Heart Disease 2015, 10<sup>th</sup> ed, Elsevier, p. 1524-1543

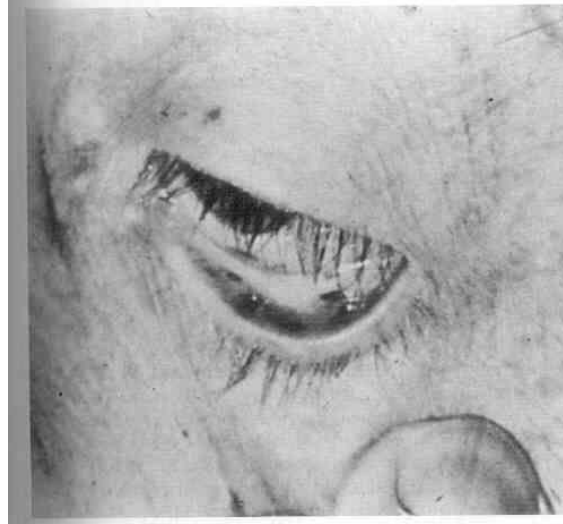
# Tần suất xuất hiện các triệu chứng thực thể bệnh nhân VNTMNT

SIGN	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

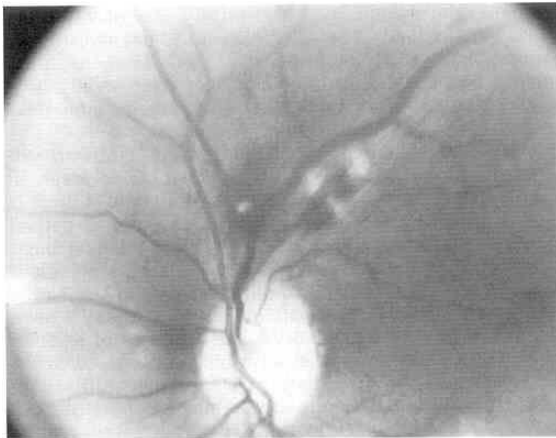
# Các triệu chứng thực thể của VNTMNT



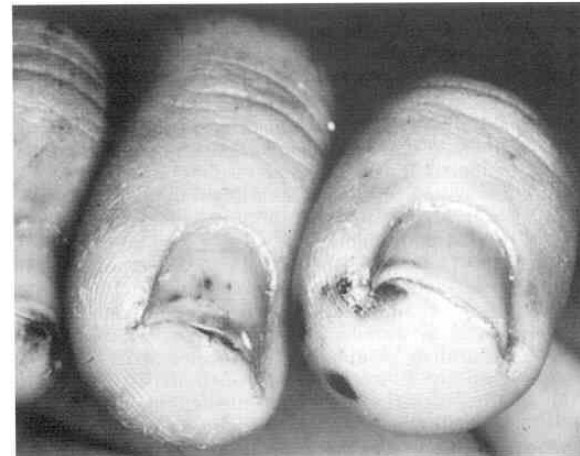
Dấu Janeway ở ngón cái



Ban xuất huyết niêm mạc mắt

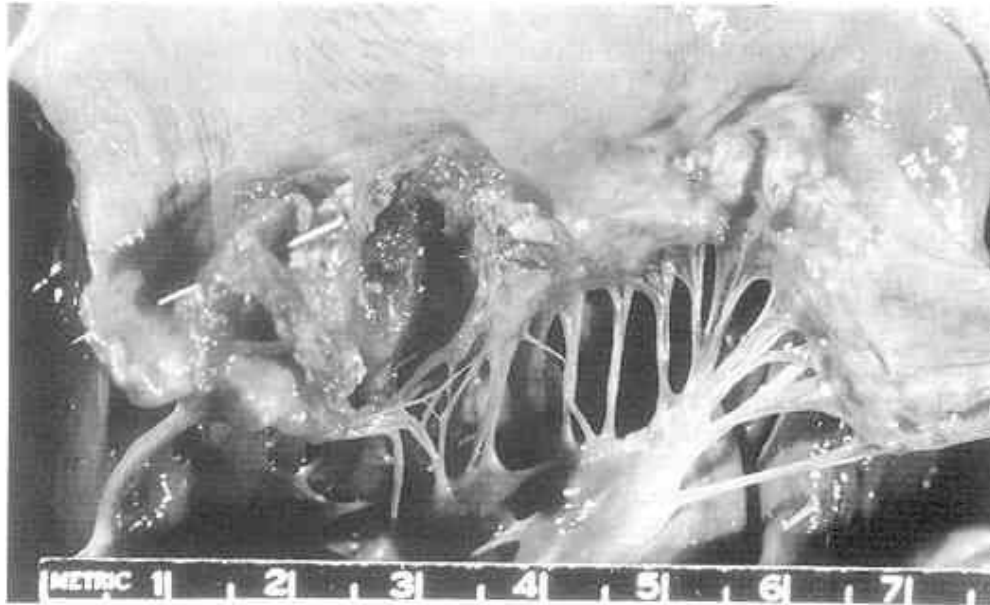


Vết Roth ở đáy mắt

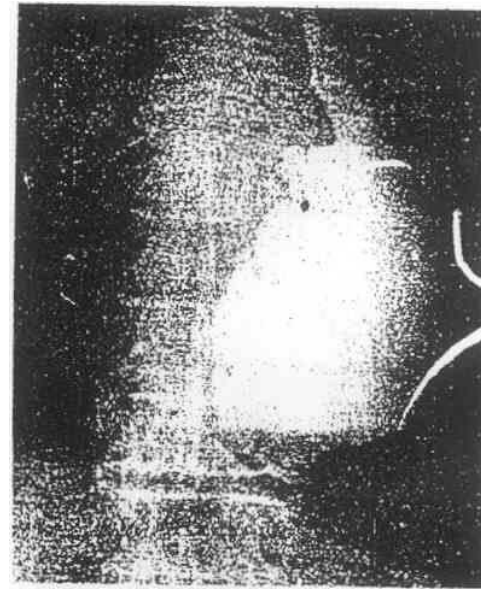


Mảnh vụn xuất huyết dưới móng tay

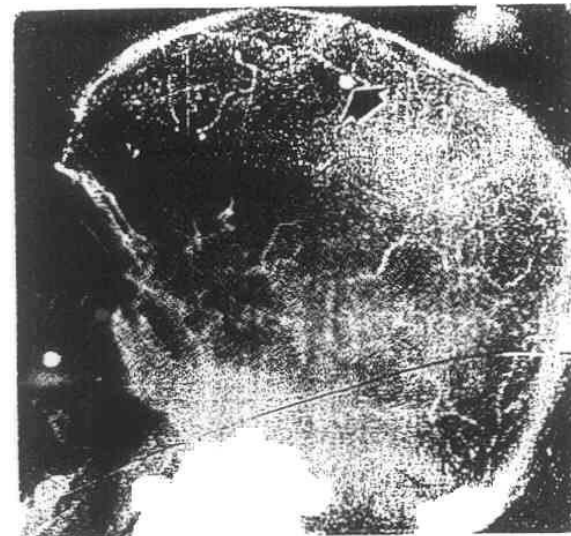
# Các triệu chứng thực thể của VNTMNT



Mảnh sùi lớn ở van 2 lá

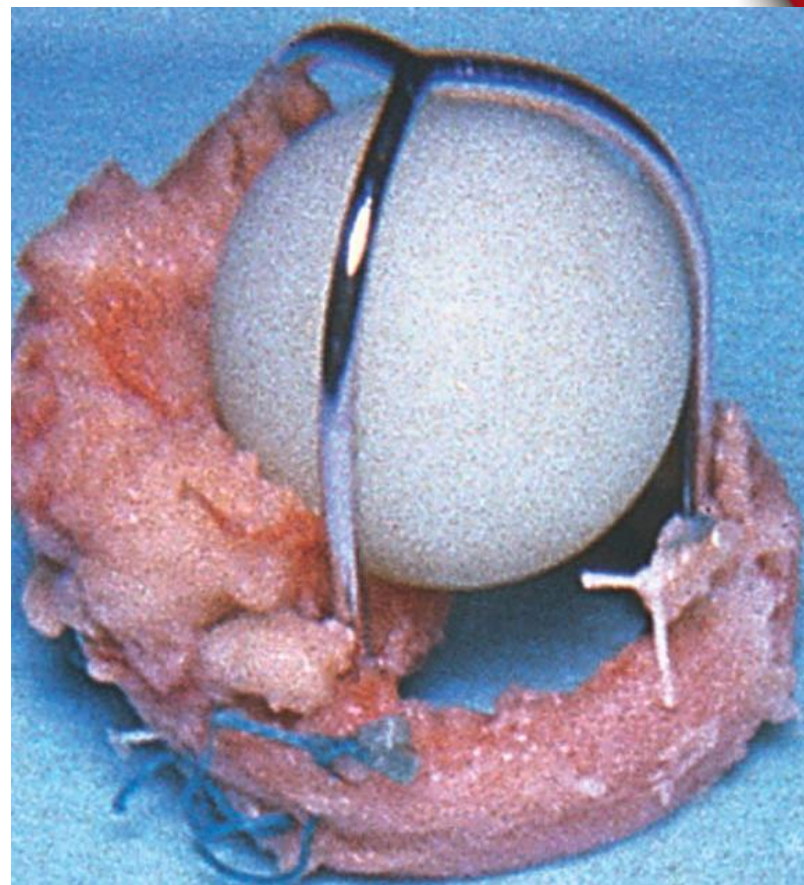
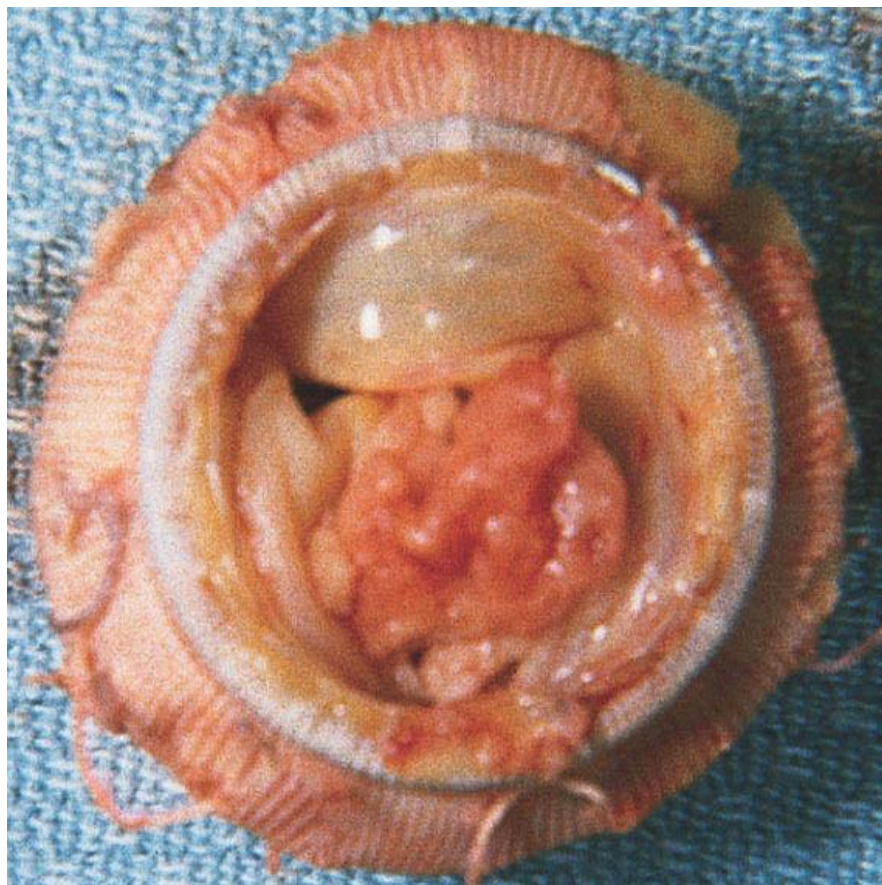


Túi phình Mycotic ở ĐMC ngực



Túi phình (mũi tên) ở ĐM não

# Mảnh sùi lớn trên van tự nhiên do *Candida albicans* (A) và mảnh sùi do nấm *Aspergillus* trên van nhân tạo Starr- Edwards



# Nhồi máu ngón chân do VNTMNT bởi *Staphylococcus aureus*

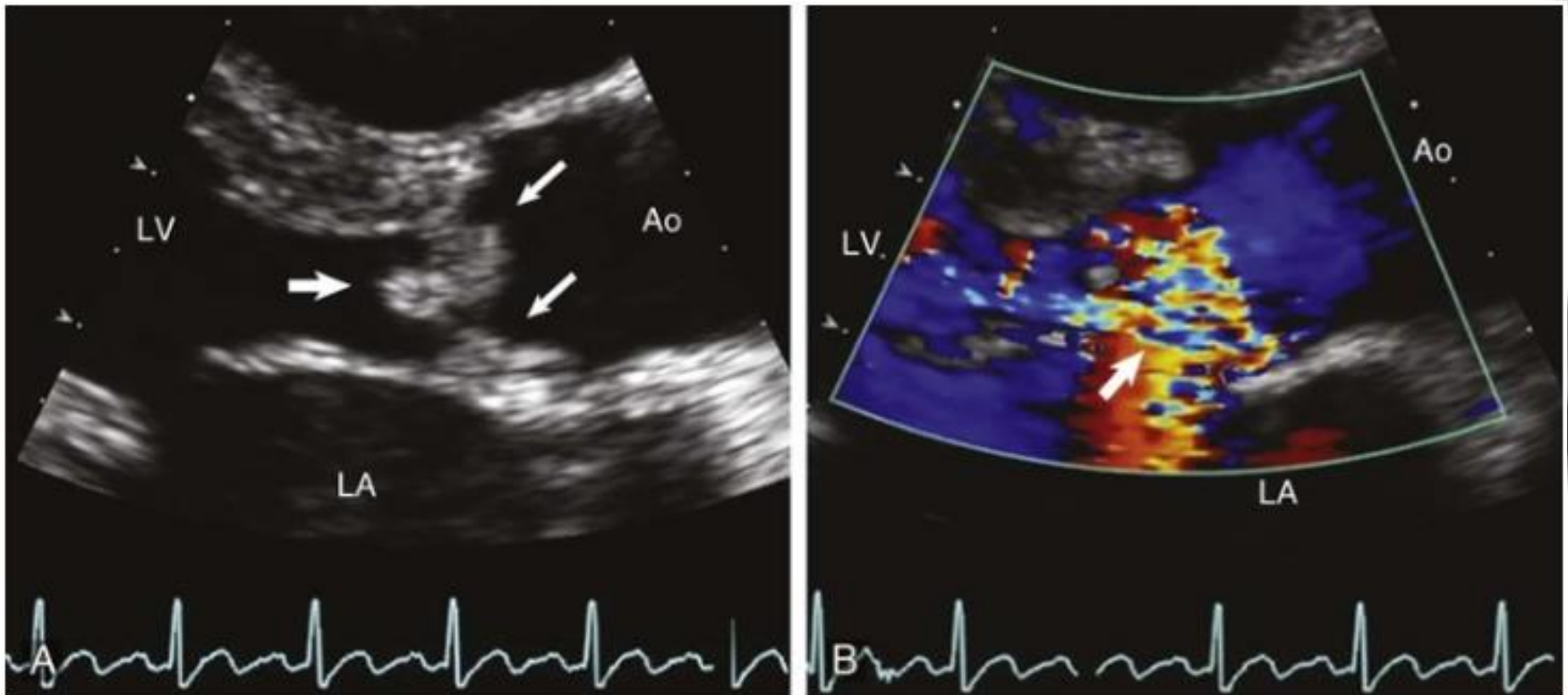


- TL: Karchmer AW. Infective Endocarditis In Braunwald's Heart Disease, ed by Libby, Bonow, Mann, Zipes. Saunders Elsevier 2008, 8th ed, p 1713-1733

# Cận lâm sàng

- Thiếu máu đẳng sắc nhẹ
- Bạch cầu có thể bình thường hay tăng
- VS tăng
- Cây máu:
  - Trước kháng sinh
  - 3 lần, lần 1,2 cách 60 phút – lần 3 cách 12 giờ kỵ khí và ái khí
- Phim lồng ngực
- Điện tâm đồ
- Siêu âm tim qua thành ngực: độ nhạy 50-80%
- Siêu âm tim qua đường thực quản: độ nhạy 90%

# Hình ảnh siêu âm tim VNTMNT trên van ĐMC tự nhiên





# Vai trò của siêu âm tim/ VNTMNT (1)

TTE: siêu âm tim qua thành ngực  
TOE: SAT qua thực quản

TL: Habib G et al. Eur. H. J 2015.  
doi/10.1093/eurheartf/ ehv 319

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
<b>A. Diagnosis</b>			
• TTE is recommended as the <u>first-line</u> imaging modality in suspected IE.	I	B	64,65
• TOE is recommended <u>in all</u> patients with clinical suspicion of IE and a negative or non-diagnostic TTE.	I	B	64, 68–71
• TOE is recommended in patients with clinical suspicion of IE, when a <u>prosthetic heart valve or an intracardiac device</u> is present.	I	B	64,71
• <u>Repeat TTE</u> and /or TOE within <u>5–7 days</u> is recommended in case of initially negative examination when clinical suspicion of IE remains high.	I	C	
• Echocardiography should be considered in <u>Staphylococcus aureus bacteraemia</u> .	IIa	B	66,67
• TOE should be considered in patients with suspected IE, even in cases with positive TTE, except in isolated right-sided native valve IE with good quality TTE examination and unequivocal echocardiographic findings.	IIa	C	

# Vai trò của siêu âm tim/ VNTMNT (2)

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
<b>B. Follow-up under medical therapy</b>			
<ul style="list-style-type: none"> <li>Repeat TTE and/or TOE are recommended as soon as a new complication of IE is suspected (new murmur, embolism, persisting fever, HF, abscess, atrioventricular block).</li> </ul>	I	B	64,72
<ul style="list-style-type: none"> <li>Repeat TTE and/or TOE should be considered during follow-up of uncomplicated IE in order to detect new silent complications and monitor vegetation size. The timing and mode (TTE or TOE) of repeat examination depend on the initial findings, type of microorganism, and initial response to therapy.</li> </ul>	IIa	B	64,72
<b>C. Intraoperative echocardiography</b>			
<ul style="list-style-type: none"> <li>Intraoperative echocardiography is recommended in all cases of IE requiring surgery.</li> </ul>	I	B	64,73
<b>D. Following completion of therapy</b>			
<ul style="list-style-type: none"> <li>TTE is recommended at completion of antibiotic therapy for evaluation of cardiac and valve morphology and function.</li> </ul>	I	C	

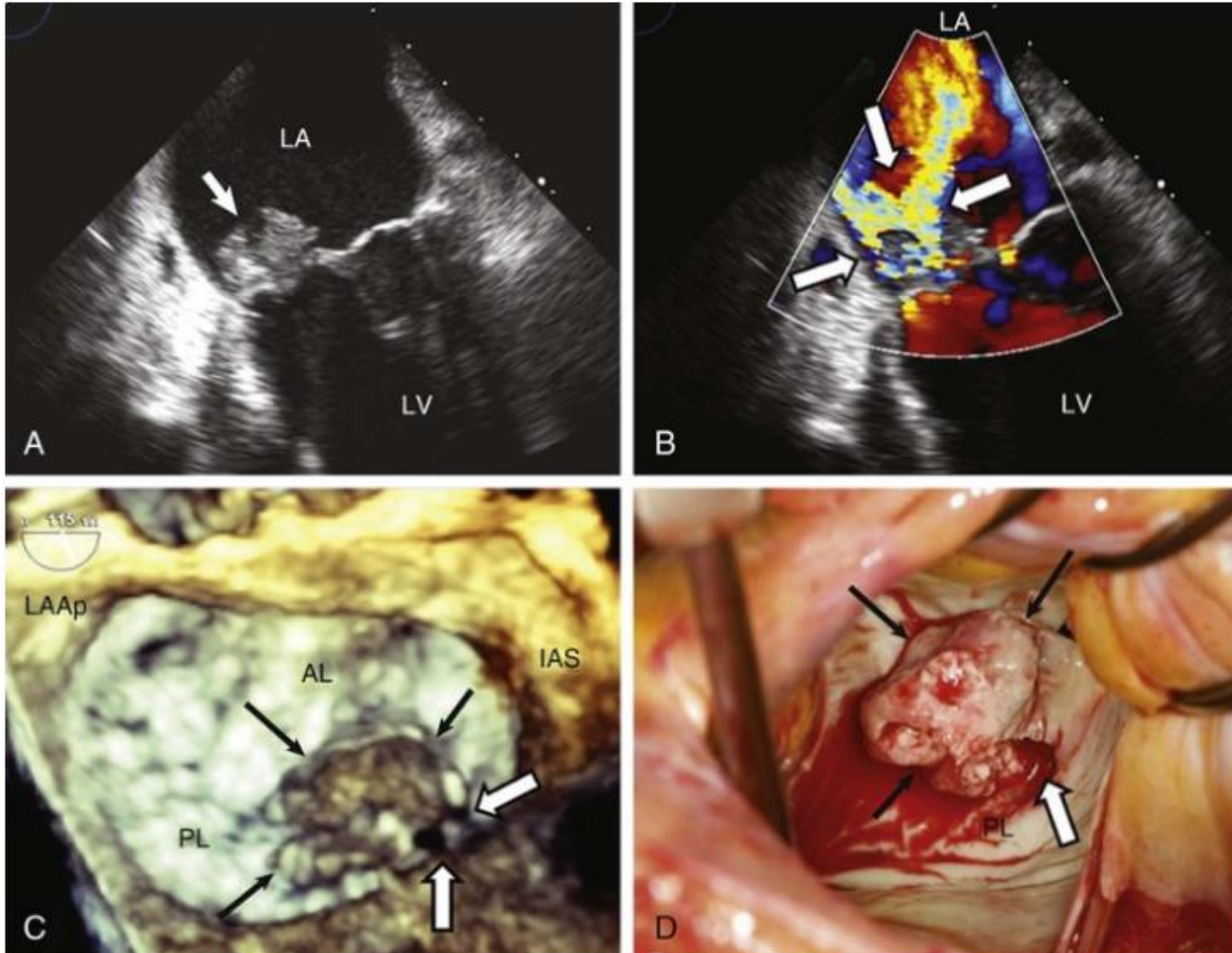
HF = heart failure; IE = infective endocarditis; TOE = transoesophageal echocardiography; TTE = transthoracic echocardiography.

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

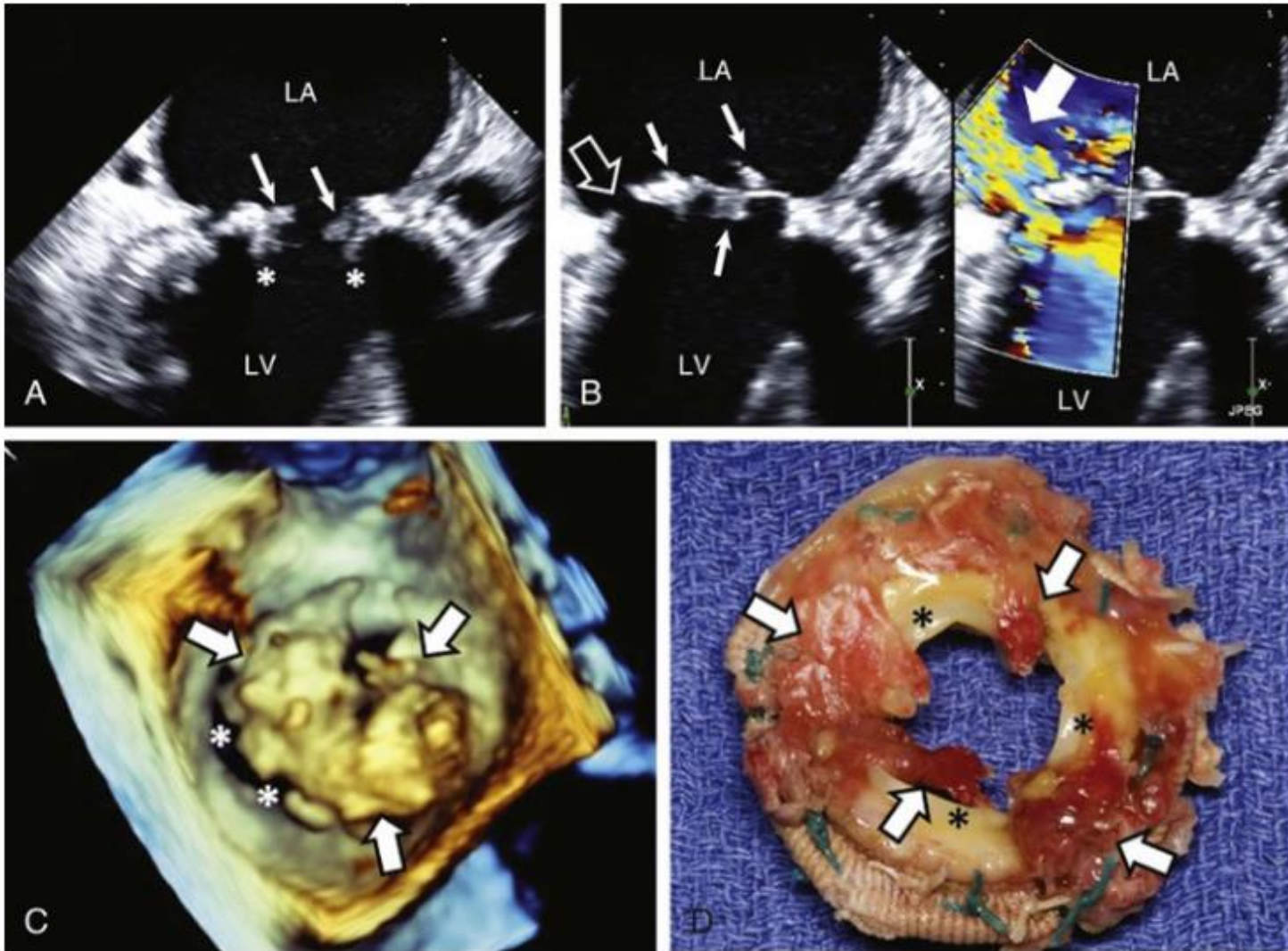
<sup>c</sup>Reference(s) supporting recommendations.

# VNTMNT/van 2 lá



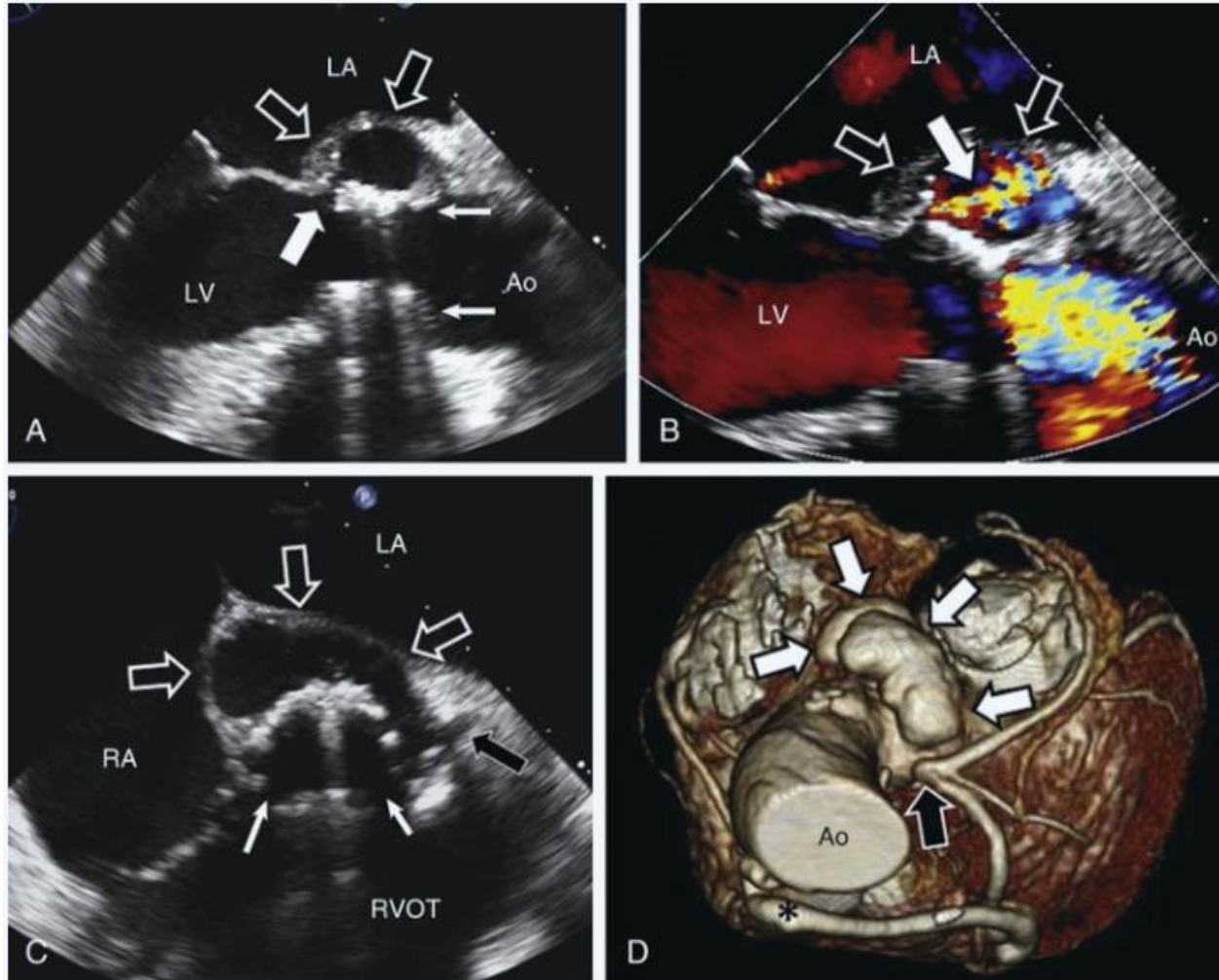
TL: Baddour LM et al. Braunwald's Heart Disease 2015, 10<sup>th</sup> ed, Elsevier, p. 1524-1543

# VNTMNT/ van sinh học nhân tạo



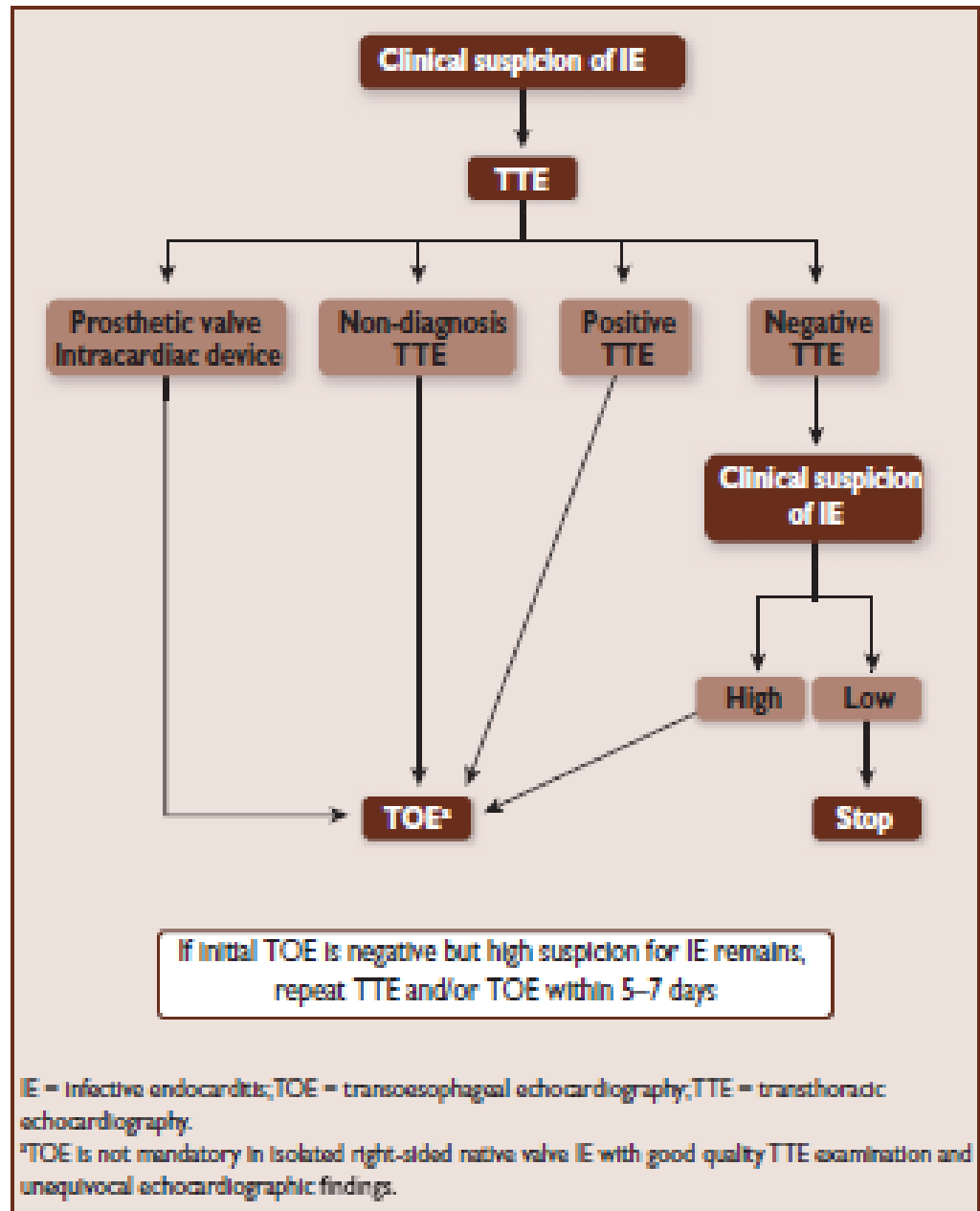
TL: Baddour LM et al. Braunwald's Heart Disease 2015, 10<sup>th</sup> ed, Elsevier, p. 1524-1543

# VNTMNT và ĐMC nhân tạo biến chứng lan quanh vòng van



TL: Baddour LM et al. Braunwald's Heart Disease 2015, 10<sup>th</sup> ed, Elsevier, p. 1524-1543

# Chỉ định siêu âm tim trên b/n ngghi ngờ VNTMNT



# Các định nghĩa về cơ thể học và siêu âm tim

	Surgery/necropsy	Echocardiography
Vegetation	Infected mass attached to an endocardial structure or on implanted intracardiac material.	Oscillating or non-oscillating intracardiac mass on valve or other endocardial structures, or on implanted intracardiac material.
Abscess	Perivalvular cavity with necrosis and purulent material not communicating with the cardiovascular lumen.	Thickened, non-homogeneous perivalvular area with echodense or echolucent appearance.
Pseudoaneurysm	Perivalvular cavity communicating with the cardiovascular lumen.	Pulsatile perivalvular echo-free space, with colour-Doppler flow detected.
Perforation	Interruption of endocardial tissue continuity.	Interruption of endocardial tissue continuity traversed by colour-Doppler flow.
Fistula	Communication between two neighbouring cavities through a perforation.	Colour-Doppler communication between two neighbouring cavities through a perforation.
Valve aneurysm	Saccular outpouching of valvular tissue.	Saccular bulging of valvular tissue.
Dehiscence of a prosthetic valve	Dehiscence of the prosthesis.	Paravalvular regurgitation identified by TTE/TOE, with or without rocking motion of the prosthesis.

TL: Habib G et al. Eur. H. J 2015. doi/10.1093/eurheartf/ ehv 319

# Chẩn đoán

- ✓ Sốt trên 1 tuần + âm thổi + không tìm thấy nguyên nhân sốt: nghi ngờ VNTMNT
- ✓ Tiêm ma túy + sốt kéo dài không rõ nguyên nhân: VNTMNT?



# Chẩn đoán bệnh VNTMNT (1)

## Chẩn đoán xác định:

Hai tiêu chuẩn chính hoặc một tiêu chuẩn chính kèm 3 tiêu chuẩn phụ, hoặc 5 tiêu chuẩn phụ

## Tiêu chuẩn chính:

1. Phân lập được *Streptococcus viridans*, *S. bovis*, nhóm HACEK, hoặc (khi không thấy ổ nhiễm trùng) phân lập được *S. aureus*, *Enterococcus* trong 2 mẫu máu riêng biệt hoặc phân lập được vi trùng phù hợp với VNTM ở 2 mẫu máu cách  $\geq 12$  giờ hoặc cả 3 hay trên 3 mẫu máu, mẫu đầu và mẫu cuối cách ít nhất 1 giờ.
2. Siêu âm tim thấy mảnh sùi lác (di động) hoặc áp xe, hoặc mới hở một phần van nhân tạo hoặc hở van tự nhiên mới có.

# Chẩn đoán bệnh VNTMNT (2)

## Chẩn đoán xác định:

### Tiêu chuẩn phụ:

1. Có tổn thương dễ gây VNTMNT hoặc người nghiện ma túy
2. Sốt  $\geq 38^{\circ}\text{C}$
3. Thuyên tắc động mạch lớn, nhồi máu phổi nhiễm trùng, túi phình mycotic, xuất huyết nội sọ, xuất huyết niêm mạc mắt, tổn thương Janeway
4. Viêm vi cầu thận, nốt Osler, nốt Roth, yếu tố thấp
5. Cây máu dương nghiệm nhưng không đạt tiêu chuẩn chính (loại trừ cây máu dương nghiệm chỉ một lần các vi khuẩn không đặc hiệu của VNTMNT) hoặc phản ứng huyết thanh dương của nhiễm trùng đang xảy ra với vi khuẩn thường gây VNTMNT
6. Siêu âm phù hợp với VNTMNT nhưng không đạt tiêu chuẩn chính

# Chẩn đoán bệnh VNTMNT (3)

## Có thể VNTMNT

- Không đủ tiêu chuẩn xác định, nhưng cũng không thuộc vào nhóm loại trừ

## Loại trừ

- Có chẩn đoán khác hoặc triệu chứng biến mất hoặc không có chứng cơ của VNTMNT khi mổ hoặc phẫu nghiệm tử thi, với chỉ dưới hay bằng 4 ngày kháng sinh

# Definition of infective endocarditis according to the modified Duke criteria (adapted from Li et al<sup>87</sup>)

Definite IE
<b>Pathological criteria</b> <ul style="list-style-type: none"><li>• Microorganisms demonstrated by culture or on histological examination of a vegetation, a vegetation that has embolized, or an intracardiac abscess specimen; or</li><li>• Pathological lesions, vegetation or intracardiac abscess confirmed by histological examination showing active endocarditis</li></ul>
<b>Clinical criteria</b> <ul style="list-style-type: none"><li>• 2 major criteria; or</li><li>• 1 major criterion and 3 minor criteria; or</li><li>• 5 minor criteria</li></ul>
Possible IE
<ul style="list-style-type: none"><li>• 1 major criterion and 1 minor criterion; or</li><li>• 3 minor criteria</li></ul>
Rejected IE
<ul style="list-style-type: none"><li>• Firm alternate diagnosis; or</li><li>• Resolution of symptoms suggesting IE with antibiotic therapy for <math>\leq 4</math> days; or</li><li>• No pathological evidence of IE at surgery or autopsy, with antibiotic therapy for <math>\leq 4</math> days; or</li><li>• Does not meet criteria for possible IE, as above</li></ul>

# Definitions of the terms used in the European Society of Cardiology 2015 modified criteria for the diagnosis of infective endocarditis (1)

Major criteria
<p>1. Blood cultures positive for IE</p> <p>a. Typical microorganisms consistent with IE from 2 separate blood cultures:</p> <ul style="list-style-type: none"><li>• Viridans streptococci, <i>Streptococcus gallolyticus</i> (<i>Streptococcus bovis</i>), HACEK group, <i>Staphylococcus aureus</i>; or</li><li>• Community-acquired enterococci, in the absence of a primary focus; or</li></ul> <p>b. Microorganisms consistent with IE from persistently positive blood cultures:</p> <ul style="list-style-type: none"><li>• <math>\geq 2</math> positive blood cultures of blood samples drawn <math>&gt;12</math> h apart; or</li><li>• All of 3 or a majority of <math>\geq 4</math> separate cultures of blood (with first and last samples drawn <math>\geq 1</math> h apart); or</li></ul> <p>c. Single positive blood culture for <i>Coxiella burnetii</i> or phase I IgG antibody titre <math>&gt;1:800</math></p>
<p>2. Imaging positive for IE</p> <p>a. Echocardiogram positive for IE:</p> <ul style="list-style-type: none"><li>• Vegetation;</li><li>• Abscess, pseudoaneurysm, intracardiac fistula;</li><li>• Valvular perforation or aneurysm;</li><li>• New partial dehiscence of prosthetic valve.</li></ul> <p>b. Abnormal activity around the site of prosthetic valve implantation detected by <math>^{18}\text{F}</math>-FDG PET/CT (only if the prosthesis was implanted for <math>&gt;3</math> months) or radiolabelled leukocytes SPECT/CT.</p> <p>c. Definite paravalvular lesions by cardiac CT.</p>

# Definitions of the terms used in the European Society of Cardiology 2015 modified criteria for the diagnosis of infective endocarditis (2)

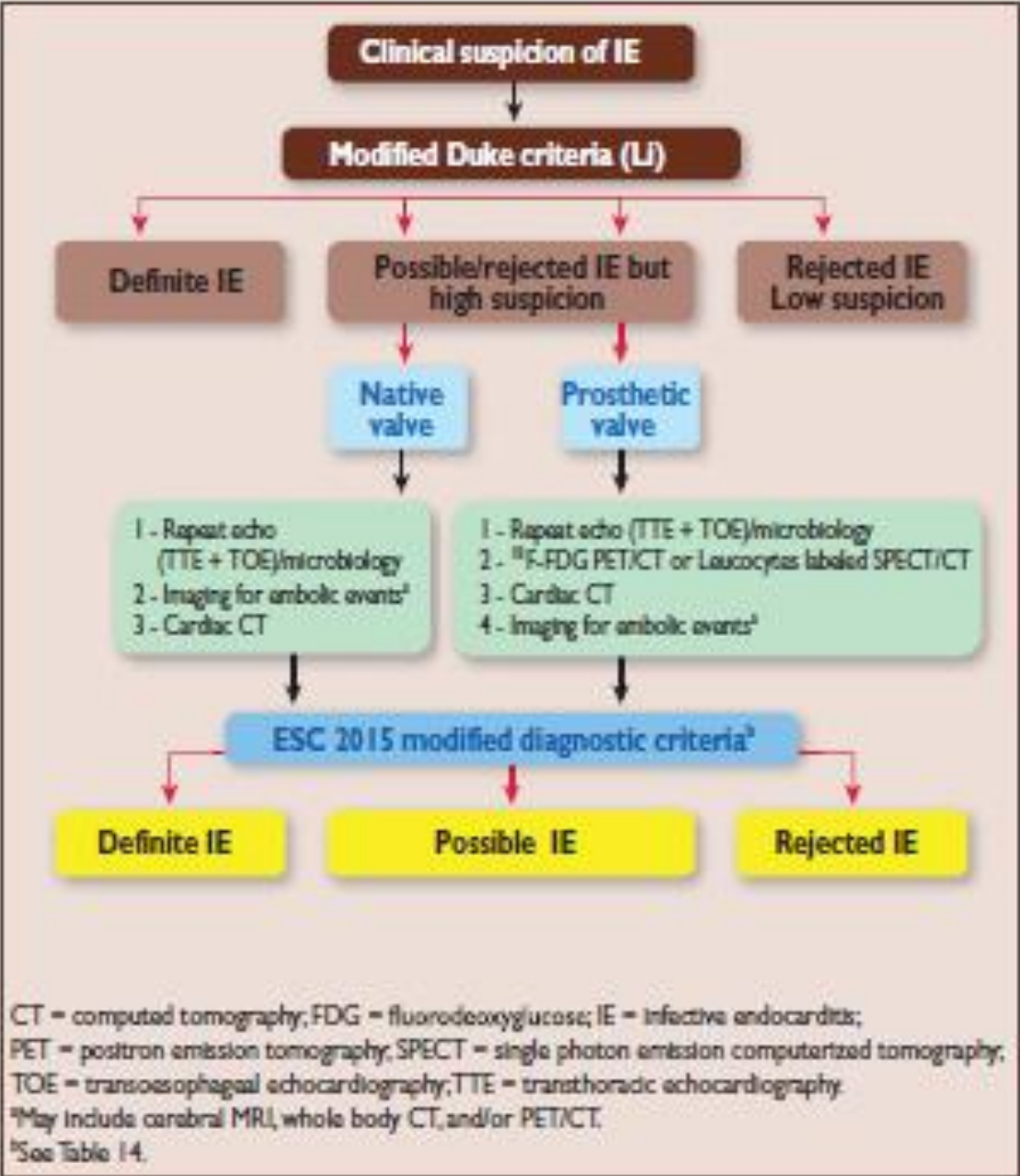
## Minor criteria

1. Predisposition such as predisposing heart condition, or injection drug use.
2. Fever defined as temperature  $>38^{\circ}\text{C}$ .
3. Vascular phenomena (including those detected by imaging only): major arterial emboli, septic pulmonary infarcts, infectious (mycotic) aneurysm, intracranial haemorrhage, conjunctival haemorrhages, and Janeway's lesions.
4. Immunological phenomena: glomerulonephritis, Osler's nodes, Roth's spots, and rheumatoid factor.
5. Microbiological evidence: positive blood culture but does not meet a major criterion as noted above or serological evidence of active infection with organism consistent with IE.

CT = computed tomography; FDG = fluorodeoxyglucose; HACEK = *Haemophilus parainfluenzae*, *H. aphrophilus*, *H. paraphrophilus*, *H. influenzae*, *Actinobacillus actinomycetemcomitans*, *Cardiobacterium hominis*, *Eikenella corrodens*, *Kingella kingae*, and *K. denitrificans*; IE = infective endocarditis; Ig = immunoglobulin; PET = positron emission tomography; SPECT = single photon emission computerized tomography. Adapted from Li et al.<sup>87</sup>

TL: Habib G et al. Eur. H. J 2015. doi/10.1093/eurheartf/ ehv 319

# European Society of Cardiology 2015 algorithm for diagnosis of infective endocarditis



TL: Habib G et al. Eur. H. J 2015. doi/10.1093/eurheartf/ ehv 319

# Chẩn đoán phân biệt

- Nhiễm trùng huyết
- Viêm phổi
- Viêm màng não
- Áp xe não
- Sốt rét
- VMNT cấp
- Viêm mạch máu
- Thấp tim
- Lao phổi
- U nhầy nhĩ trái



# Các biến chứng của VNTMNT

- Suy tim
- Thuyên tắc
- Rối loạn dẫn truyền
- Biểu hiện thần kinh
- Túi phình mycotic (mycotic aneurysm)

# Điều trị

- Nguyên tắc điều trị:
  - Kháng sinh diệt khuẩn
  - Liều cao
  - Dài ngày (4-6 tuần)
- Lựa chọn kháng sinh theo :
  - Loại vi trùng
  - Cơ địa người bệnh (có tiêm ma túy? Van nhân tạo?)
  - Cấp hay bán cấp



# Antibiotic treatment of infective endocarditis due to oral streptococci and Streptococcus bovis group<sup>a</sup> ( 1)

Antibiotic	Dosage and route	Duration (weeks)	Class <sup>b</sup>	Level <sup>c</sup>	Ref. <sup>d</sup>	Comments
<b>Strains penicillin-susceptible (MIC ≤ 0.125 mg/L) oral and digestive streptococci</b>						
<b>Standard treatment: 4-week duration</b>						
Penicillin G or Amoxicillin <sup>e</sup> or Ceftriaxone <sup>f</sup>	12–18 million U/day i.v. either in 4–6 doses or continuously  100–200 mg/kg/day i.v. in 4–6 doses  2 g/day i.v. or i.m. in 1 dose  <b>Paediatric doses:<sup>g</sup></b> Penicillin G 200,000 U/kg/day i.v. in 4–6 divided doses Amoxicillin 300 mg/kg/day i.v. in 4–6 equally divided doses Ceftriaxone 100 mg/kg/day i.v. or i.m. in 1 dose	4  4  4	I  I  I	B  B  B	6,8, 135– 139	Preferred in patients > 65 years or with impaired renal or VIII (vestibulocochlear) cranial nerve functions. 6-week therapy recommended for patients with PVE

# Antibiotic treatment of infective endocarditis due to oral streptococci and Streptococcus bovis group<sup>a</sup> ( 2)



Standard treatment: 2-week duration						
Penicillin G or Amoxicillin <sup>e</sup> or Ceftriaxone <sup>f</sup> <b>combined with</b> Gentamicin <sup>h</sup> or Netilmicin	12–18 million U/day i.v. either in 4–6 doses or continuously  100–200 mg/kg/day i.v. in 4–6 doses  2 g/day i.v. or i.m. in 1 dose  3 mg/kg/day i.v. or i.m. in 1 dose  4–5 mg/kg/day i.v. in 1 dose  <b>Paediatric doses:<sup>g</sup></b> Penicillin G, amoxicillin, and ceftriaxone as above Gentamicin 3 mg/kg/day i.v. or i.m. in 1 dose or 3 equally divided doses	2	I	B	6,8, 127, 135– 138	Only recommended in patients with non-complicated NVE with normal renal function.          Netilmicin is not available in all European countries.
<b>In beta-lactam allergic patients<sup>i</sup></b>						
Vancomycin <sup>i</sup>	30 mg/kg/day i.v. in 2 doses  <b>Paediatric doses:<sup>g</sup></b> Vancomycin 40 mg/kg/day i.v. in 2 or 3 equally divided doses	4	I	C		6-week therapy recommended for patients with PVE

# Antibiotic treatment of infective endocarditis due to oral streptococci and Streptococcus bovis group<sup>a</sup> ( 3)

Strains relatively resistant to penicillin (MIC 0.250–2 mg/L) <sup>k</sup>						
Standard treatment						
Penicillin G or Amoxicillin <sup>e</sup> or Ceftriaxone <sup>f</sup> combined with Gentamicin <sup>h</sup>	24 million U/day i.v. either in 4–6 doses or continuously	4	I	B	6,8, 135, 136	6-week therapy recommended for patients with PVE
	200 mg/kg/day i.v. in 4–6 doses	4	I	B		
	2 g/day i.v. or i.m. in 1 dose	4	I	B		
	3 mg/kg/day i.v. or i.m. in 1 dose	2	I	B		
In beta-lactam allergic patients <sup>i</sup>						
Vancomycin <sup>i</sup> with Gentamicin <sup>k</sup>  <b>Paediatric doses:<sup>e</sup></b> As above	30 mg/kg/day i.v. in 2 doses	4	I	C		6-week therapy recommended for patients with PVE
	3 mg/kg/day i.v. or i.m. in 1 dose	2	I	C		

C<sub>min</sub> = minimum concentration; IE = infective endocarditis; Im. = intramuscular; Iv. = intravenous; MIC = minimum inhibitory concentration; NVE = native valve endocarditis; PVE = prosthetic valve endocarditis; U = units.

<sup>a</sup>Refer to text for other streptococcal species; <sup>b</sup>Class of recommendation; <sup>c</sup>Level of evidence; <sup>d</sup>Reference(s) supporting recommendations; <sup>e</sup>Or ampicillin, same dosages as amoxicillin; <sup>f</sup>Preferred for outpatient therapy; <sup>g</sup>Paediatric doses should not exceed adult doses; <sup>h</sup>Renal function and serum gentamicin concentrations should be monitored once a week. When given in a single daily dose, pre-dose (trough) concentrations should be < 1 mg/L and post-dose (peak; 1 hours after injection) serum concentrations should be ~10–12 mg/L<sup>148</sup>; <sup>i</sup>Penicillin desensitization can be attempted in stable patients; <sup>j</sup>Serum vancomycin concentrations should achieve 10–15 mg/L at pre-dose (trough) level, although some experts recommend to increase the dose of vancomycin to 45–60 mg/kg/day Iv. in 2 or 3 divided doses to reach serum trough vancomycin levels (C<sub>min</sub>) of 15–20 mg/L as in staphylococcal endocarditis. However, vancomycin dose should not exceed 2 g/d unless serum levels are monitored and can be adjusted to obtain a peak plasma concentration of 30–45 µg/mL 1 hour after completion of the Iv. infusion of the antibiotic; <sup>k</sup>Patients with penicillin-resistant strains (MIC > 2 mg/L) should be treated as enterococcal endocarditis (see Table 18).



# Antibiotic treatment of infective endocarditis due to *Staphylococcus* spp (1)

Antibiotic	Dosage and route	Duration (weeks)	Class <sup>i</sup>	Level <sup>l</sup>	Ref. <sup>k</sup>	Comments
<b>Native valves</b>						
<b>Methicillin-susceptible staphylococci</b>						
(Flu)cloxacillin or oxacillin	12 g/day I.v. in 4–6 doses	4–6	I	B	6,8, 128, 135, 136, 158	Gentamicin addition is not recommended because clinical benefit has not been demonstrated and there is increased renal toxicity
	<b>Paediatric doses:</b> <sup>g</sup> 200–300 mg/kg/day I.v. in 4–6 equally divided doses					
<b>Alternative therapy*</b> Cotrimoxazole <sup>a</sup>  with Clindamycin	Sulfamethoxazole 4800 mg/day and Trimethoprim 960 mg/day (I.v. in 4–6 doses)	1 I.v. + 5 oral intake	IIb	C		*for <i>Staphylococcus aureus</i>
	1800mg/day I.v. in 3 doses	1	IIb	C		
	<b>Paediatric doses:</b> <sup>g</sup> Sulfamethoxazole 60 mg/kg/day and Trimethoprim 12 mg/kg/day (I.v. in 2 doses) Clindamycin 40 mg/kg/day (I.v. in 3 doses)					



# Antibiotic treatment of infective endocarditis due to Staphylococcus spp (2)

Penicillin-allergic patients <sup>b</sup> or methicillin-resistant staphylococci						
Vancomycin <sup>b, **</sup>	30–60 mg/kg/day I.v. in 2–3 doses  Paediatric doses: <sup>e</sup> 40 mg/kg/day I.v. in 2–3 equally divided doses	4–6	I	B	6,8, 135, 136	<p><b>Cephalosporins</b> (cefazolin 6 g/day or cefotaxime 6 g/day I.v. in 3 doses) are recommended for penicillin-allergic patients with non-anaphylactic reactions with methicillin-susceptible endocarditis</p> <p><b>Daptomycin</b> is superior to vancomycin for MSSA and MRSA bacteraemia with vancomycin MIC &gt; 1 mg/L</p> <p>*for <i>Staphylococcus aureus</i></p>
<b>Alternative therapy<sup>**</sup>:</b> Daptomycin <sup>c,d</sup>	10 mg/kg/day I.v. once daily  Paediatric doses: <sup>e</sup> 10 mg/kg/day I.v. once daily	4–6	IIa	C		
<b>Alternative therapy*</b> Cotrimoxazole <sup>a</sup>  with Clindamycin	Sulfamethoxazole 4800 mg/day and Trimethoprim 960 mg/day (I.v. in 4–6 doses)  1800mg/day IV in 3 doses	1 I.v. + 5 oral intake  1	IIb	C		





# Antibiotic treatment of infective endocarditis due to *Staphylococcus* spp (3)

Prosthetic valves						
Methicillin-susceptible staphylococci						
(Flu)cloxacillin or oxacillin with Rifampin <sup>g</sup> and Gentamicin <sup>f</sup>	12 g/day Iv. in 4–6 doses	≥ 6	I	B	6,8, 135, 136	Starting rifampin 3–5 days later than vancomycin and gentamicin has been suggested by some experts.  Gentamicin can be given in a single daily dose in order to reduce renal toxicity
	900–1200 mg Iv. or orally in 2 or 3 divided doses	≥ 6	I	B		
	3 mg/kg/day Iv. or Im. in 1 or 2 doses	2	I	B		
	<b>Paediatric doses:<sup>g</sup></b> Oxacillin and (flu)cloxacillin as above Rifampin 20 mg/kg/day Iv. or orally in 3 equally divided doses					







# Antibiotic treatment of infective endocarditis due to *Staphylococcus* spp (4)

Penicillin-allergic patients <sup>b</sup> and methicillin-resistant staphylococci						
Vancomycin <sup>b</sup> with Rifampin <sup>a</sup> and Gentamicin <sup>f</sup>	30–60 mg/kg/day I.v. in 2–3 doses	IV 6	I	B	6,8, 135, 136	<b>Cephalosporins</b> (cefazolin 6 g/day or cefotaxime 6 g/day I.v. in 3 doses) are recommended for penicillin-allergic patients with non-anaphylactic reactions with methicillin-susceptible endocarditis. Starting rifampin 3–5 days later than vancomycin and gentamicin has been suggested by some experts. Gentamicin can be given in a single daily dose in order to reduce renal toxicity
	900–1200 mg I.v. or orally in 2 or 3 divided doses	IV 6	I	B		
	3 mg/kg/day I.v. or I.m. in 1 or 2 doses	2	I	B		
<b>Paediatric dosing:</b> <sup>g</sup> As above						

AUC = area under the curve; C<sub>min</sub> = minimum concentration; IE = infective endocarditis; MIC = minimum inhibitory concentration; MRSA = methicillin-resistant *Staphylococcus aureus*; MSSA = methicillin-susceptible *S. aureus*; PVE = prosthetic valve endocarditis.

<sup>a</sup>Renal function, serum Cotrimoxazole concentrations should be monitored once/week (twice/week in patients with renal failure); <sup>b</sup>Serum trough vancomycin levels (C<sub>min</sub>) should be ≥ 20 mg/L. A vancomycin AUC/MIC >400 is recommended for MRSA infections; <sup>c</sup>Monitor plasma CPK levels at least once a week. Some experts recommend adding cloxacillin (2 g/4 h i.v.) or fosfomycin (2 g/6 h I.v.) to daptomycin in order to increase activity and avoid the development of daptomycin resistance; <sup>d</sup>Daptomycin and fosfomycin are not available in some European countries; <sup>e</sup>Rifampin is believed to play a special role in prosthetic device infection because it helps eradicate bacteria attached to foreign material.<sup>157</sup> The sole use of rifampin is associated with a high frequency of microbial resistance and is not recommended. Rifampin increases the hepatic metabolism of warfarin and other drugs; <sup>f</sup>Renal function and serum gentamicin concentrations should be monitored once/week (twice/week in patients with renal failure); <sup>g</sup>Paediatric doses should not exceed adult doses; <sup>h</sup>Penicillin desensitization can be attempted in stable patients; <sup>i</sup>Class of recommendation; <sup>j</sup>Level of evidence; <sup>k</sup>Reference(s) supporting recommendations.

\*\* No clinical benefit of adding rifampicin or gentamicin

# Antibiotic treatment of infective endocarditis due to *Enterococcus* spp

Antibiotic	Dosage and route	Duration, weeks	Class <sup>e</sup>	Level <sup>h</sup>	Ref. <sup>i</sup>	Comments
<b>Beta-lactam and gentamicin-susceptible strains (for resistant isolates see <sup>a,b,c</sup>)</b>						
Amoxicillin* with Gentamicin <sup>d</sup>	200 mg/kg/day i.v. in 4–6 doses	4–6	I	B	6,8, 129, 135, 136, 186	6-week therapy recommended for patients with >3 months symptoms or PVE
	3 mg/kg/day i.v. or i.m. in 1 dose	2–6**	I	B		
<b>Paediatric doses:<sup>e</sup></b> Ampicillin 300 mg/kg/day i.v. in 4–6 equally divided doses Gentamicin 3 mg/kg/day i.v. or i.m. in 3 equally divided doses						
Ampicillin with Ceftriaxone	200 mg/kg/day i.v. in 4–6 doses	6	I	B	183–185	This combination is active against <i>Enterococcus faecalis</i> strains with and without HLAR, being the combination of choice in patients with HLAR <i>E. faecalis</i> endocarditis.
	4 g/day i.v. or i.m. in 2 doses	6	I	B		
<b>Paediatric doses:<sup>e</sup></b> Amoxicillin as above Ceftriaxone 100 mg/kg/12 h i.v. or i.m.						This combination is not active against <i>E. faecium</i>
Vancomycin <sup>f</sup> with Gentamicin <sup>d</sup>	30 mg/kg/day i.v. in 2 doses	6	I	C		
	3 mg/kg/day i.v. or i.m. in 1 dose	6	I	C		
<b>Paediatric doses:<sup>e</sup></b> Vancomycin 40 mg/kg/day i.v. in 2–3 equally divided doses. Gentamicin as above						

# Antibiotic treatment of blood culture – negative infective endocarditis (adapted from Brouqui et al <sup>193</sup>)

Pathogens	Proposed therapy <sup>a</sup>	Treatment outcome
<i>Brucella</i> spp.	Doxycycline (200 mg/24 h) plus cotrimoxazole (960 mg/12 h) plus rifampin (300–600/24 h) for ≥3–6 months <sup>b</sup> orally	Treatment success defined as an antibody titre <1:60. Some authors recommend adding gentamicin for the first 3 weeks.
<i>C. burnetii</i> (agent of Q fever)	Doxycycline (200 mg/24 h) plus hydroxychloroquine (200–600 mg/24 h) <sup>c</sup> orally (>18 months of treatment)	Treatment success defined as anti-phase I IgG titre <1:200, and IgA and IgM titres <1:50.
<i>Bartonella</i> spp. <sup>d</sup>	Doxycycline 100 mg/12 h orally for 4 weeks plus gentamicin (3 mg/24 h) i.v. for 2 weeks	Treatment success expected in ≥90%.
<i>Legionella</i> spp.	Levofloxacin (500 mg/12 h) i.v. or orally for ≥6 weeks or clarithromycin (500 mg/12 h) i.v. for 2 weeks, then orally for 4 weeks plus rifampin (300–1200 mg/24 h)	Optimal treatment unknown.
<i>Mycoplasma</i> spp.	Levofloxacin (500 mg/12 h) i.v. or orally for ≥6 months <sup>e</sup>	Optimal treatment unknown.
<i>T. whipplei</i> (agent of Whipple's disease) <sup>f</sup>	Doxycycline (200 mg/24 h) plus hydroxychloroquine (200–600 mg/24 h) <sup>c</sup> orally for ≥18 months	Long-term treatment, optimal duration unknown.

ID = infectious disease; IE = infective endocarditis; Ig = immunoglobulin; I.v. = intravenous; U = units.

<sup>a</sup>Owing to the lack of large series, the optimal duration of treatment of IE due to these pathogens is unknown. The presented durations are based on selected case reports. Consultation with an ID specialist is recommended.

<sup>b</sup>Addition of streptomycin (15 mg/kg/24 h in 2 doses) for the first few weeks is optional.

<sup>c</sup>Doxycycline plus hydroxychloroquine (with monitoring of serum hydroxychloroquine levels) is significantly superior to doxycycline.<sup>194</sup>

<sup>d</sup>Several therapeutic regimens have been reported, including aminopenicillins (ampicillin or amoxicillin, 12 g/24 h I.v.) or cephalosporins (ceftriaxone, 2 g/24 h I.v.) combined with aminoglycosides (gentamicin or netilmicin).<sup>195</sup> Dosages are as for streptococcal and enterococcal IE (Tables 16 and 18).<sup>196,197</sup>

<sup>e</sup>Newer fluoroquinolones (levofloxacin, moxifloxacin) are more potent than ciprofloxacin against intracellular pathogens such as *Mycoplasma* spp., *Legionella* spp., and *Chlamydia* spp.

<sup>f</sup>Treatment of Whipple's IE remains highly empirical. In the case of central nervous system involvement, sulfadiazine 1.5 g/6 h orally must be added to doxycycline. An alternative therapy is ceftriaxone (2 g/24 h I.v.) for 2–4 weeks or penicillin G (2 million U/4 h) and streptomycin (1 g/24 h) I.v. for 2–4 weeks followed by cotrimoxazole (800 mg/12 h) orally. Trimethoprim is not active against *T. whipplei*. Successes have been reported with long-term therapy (>1 year).

# Proposed antibiotic regimens for initial empirical treatment of infective endocarditis in acute severely ill patients (before pathogen identification)<sup>a</sup> (1)



Antibiotic	Dosage and route	Class <sup>b</sup>	Level <sup>c</sup>	Comments
<b>Community-acquired native valves or late prosthetic valves (≥ 12 months post surgery) endocarditis</b>				
Ampicillin with (Flu)cloxacillin or oxadillin with Gentamicin <sup>d</sup>	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	IIa	C	Patients with BCNIE should be treated in consultation with an ID specialist.
Vancomycin <sup>d</sup> with Gentamicin <sup>d</sup>	30–60 mg/kg/day i.v. in 2–3 doses  3 mg/kg/day i.v. or i.m. in 1 dose			

# Proposed antibiotic regimens for initial empirical treatment of infective endocarditis in acute severely ill patients (before pathogen identification) <sup>a</sup> (2)



Early PVE (<12 months post surgery) or nosocomial and non-nosocomial healthcare associated endocarditis				
Vancomycin <sup>d</sup> with Gentamicin <sup>d</sup> 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	<b>IIb</b>	<b>C</b>	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 <i>S. aureus</i> identification

BCNIE = blood culture-negative infective endocarditis; ID = infectious disease; Im. = intramuscular; Iv. = intravenous; PVE = prosthetic valve endocarditis.  
<sup>a</sup>If initial blood cultures are negative and there is no clinical response, consider BCNIE aetiology (see Section 7.10) and maybe surgery for molecular diagnosis and treatment, and extension of the antibiotic spectrum to blood culture-negative pathogens (doxycycline, quinolones) must be considered.  
<sup>b</sup>Class of recommendation.  
<sup>c</sup>Level of evidence.  
<sup>d</sup>Monitoring of gentamicin or vancomycin dosages is as described in Tables 16 and 17.

# Điều trị VNTMNT do nhóm HACEK

Antibiotic	Dosage and Route <sup>†</sup>	Duration (wk)
Ceftriaxone <sup>‡</sup>	2 gm once daily IV or IM	4
Ampicillin/Sulbactam	12 gm/24 hr IV given every 4 hr in six equally divided doses	4

- TL: Karchmer AW. Infective Endocarditis In Braunwald's Heart Disease, ed by Libby, Bonow, Mann, Zipes. Saunders Elsevier 2008, 8th ed, p 1713-1733

# Các dấu hiệu siêu âm tim, gợi ý b/n VNTMNT cần phẫu thuật

Vegetation
Persistent vegetation after systemic embolization
Anterior mitral valve leaflet vegetation, particularly if it is highly mobile with size > 10 mm*
One or more embolic events during the first 2 weeks of antimicrobial therapy*
Increase in 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 therapy†

See text for more complete discussion of indications for surgery based on vegetation characterizations.

# Indications and timing of surgery in left sided valve infective endocarditis (native valve endocarditis and prosthetic valve endocarditis) (1)

Indications for surgery	Timing <sup>a</sup>	Class <sup>b</sup>	Level <sup>c</sup>	Ref. <sup>d</sup>
<b>1. Heart failure</b>				
Aortic or mitral NVE or PVE with severe acute regurgitation, obstruction or fistula causing refractory pulmonary oedema or cardiogenic shock	Emergency	I	B	111,115, 213,216
Aortic or mitral NVE or PVE with severe regurgitation or obstruction causing symptoms of HF or echocardiographic signs of poor haemodynamic tolerance	Urgent	I	B	37,115, 209,216, 220,221



# Indications and timing of surgery in left sided valve infective endocarditis (native valve endocarditis and prosthetic valve endocarditis) (2)



<b>2. Uncontrolled infection</b>				
Locally uncontrolled infection (abscess, false aneurysm, fistula, enlarging vegetation)	Urgent	I	B	37,209, 216
Infection caused by fungi or multiresistant organisms	Urgent/ elective	I	C	
Persisting positive blood cultures despite appropriate antibiotic therapy and adequate control of septic metastatic foci	Urgent	IIa	B	123
PVE caused by staphylococci or non-HACEK gram-negative bacteria	Urgent/ elective	IIa	C	

# Indications and timing of surgery in left sided valve infective endocarditis (native valve endocarditis and prosthetic valve endocarditis) (3)

3. Prevention of embolism				
Aortic or mitral NVE or PVE with persistent vegetation <b>&gt;10 mm after one or more embolic episode despite appropriate antibiotic therapy</b>	Urgent	I	B	9,58,72, 113,222
Aortic or mitral NVE with vegetations >10 mm, associated with severe valve stenosis or regurgitation, and low operative risk	Urgent	IIa	B	9
Aortic or mitral NVE or PVE with isolated very large vegetations ( <b>&gt;30 mm</b> )	Urgent	IIa	B	113
Aortic or mitral NVE or PVE with isolated large vegetations (>15 mm) and no other indication for surgery <sup>a</sup>	Urgent	IIb	C	

HACEK = *Haemophilus parainfluenzae*, *Haemophilus aphrophilus*, *Haemophilus paraphrophilus*, *Haemophilus influenzae*, *Actinobacillus actinomycetemcomitans*, *Cardiobacterium hominis*, *Eikenella corrodens*, *Kingella kingae* and *Kingella denitrificans*; HF = heart failure; IE = infective endocarditis; NVE = native valve endocarditis; PVE = prosthetic valve endocarditis.

<sup>a</sup>Emergency surgery: surgery performed within 24 h; urgent surgery: within a few days; elective surgery: after at least 1–2 weeks of antibiotic therapy.

<sup>b</sup>Class of recommendation.

<sup>c</sup>Level of evidence.

<sup>d</sup>Reference(s) supporting recommendations.

<sup>e</sup>Surgery may be preferred if a procedure preserving the native valve is feasible.

# Management of neurological complications of infective endocarditis (1)

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
After a <b>silent embolism</b> or transient ischaemic attack, <b>cardiac surgery</b> , if indicated, is recommended without delay	I	B	105, 263
Neurosurgery or endovascular therapy is recommended for very <b>large, enlarging</b> or ruptured intracranial infectious aneurysms	I	C	
Following intracranial haemorrhage, surgery should generally be <b>postponed</b> for $\geq 1$ month	IIa	B	264–266

# Management of neurological complications of infective endocarditis (2)

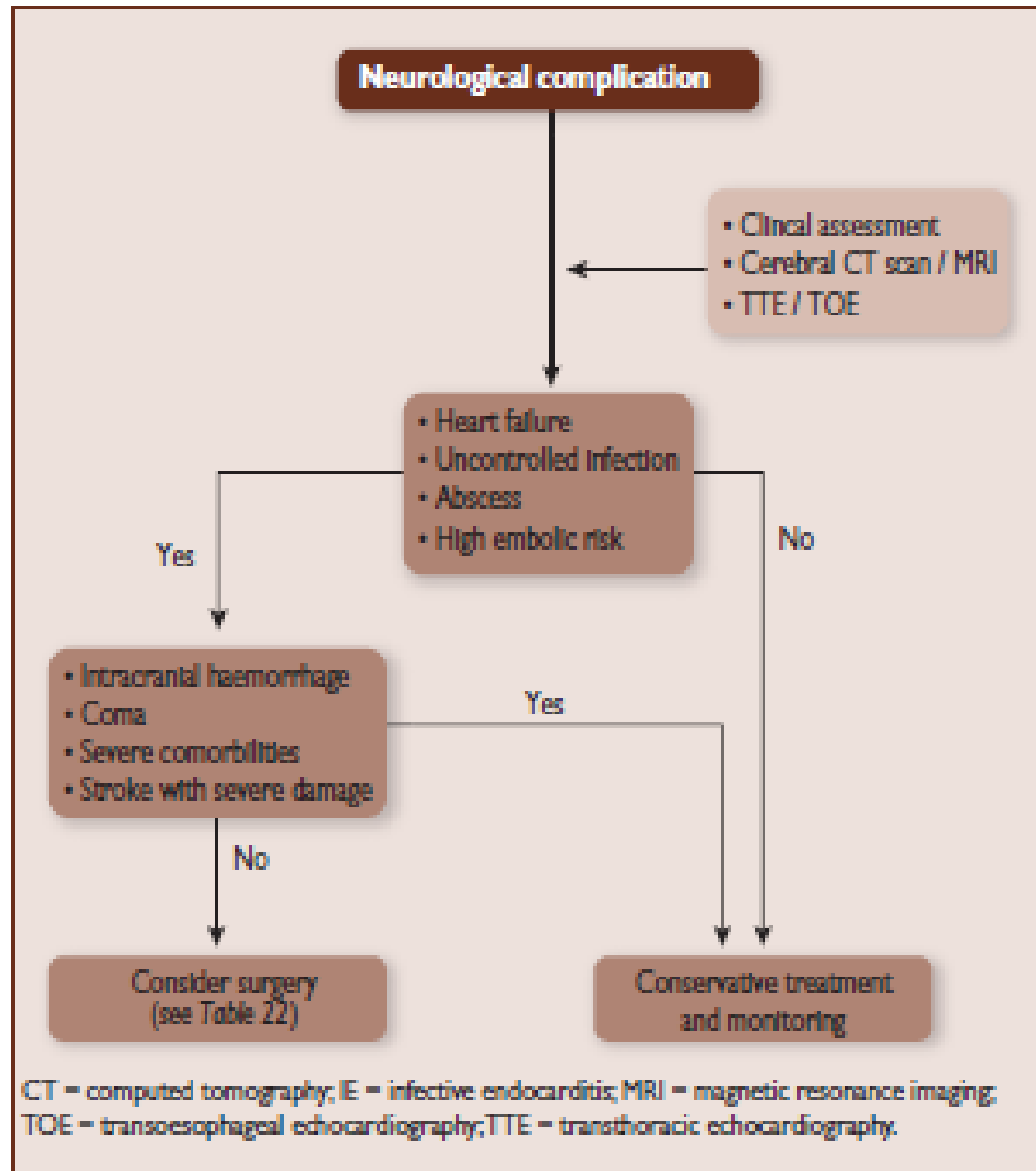


After a stroke, surgery indicated for HF, uncontrolled infection, abscess, or persistent high embolic risk should be considered without any delay as long as coma is absent and the presence of cerebral haemorrhage has been excluded by cranial CT or MRI	IIa	B	9263
Intracranial infectious aneurysms should 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	IIa	B	267, 268

CT = computed tomography; HF = heart failure; IE = infective endocarditis; MR = magnetic resonance; MRI = magnetic resonance imaging.  
<sup>a</sup>Class of recommendation.  
<sup>b</sup>Level of evidence.  
<sup>c</sup>Reference(s) supporting recommendations.



# Therapeutic strategies of patients with infective endocarditis and neurological complications



# Factors associated with an increased rate of relapse

- Inadequate antibiotic treatment (agent, dose, duration)
- Resistant microorganisms, i.e. *Brucella* spp., *Legionella* spp., *Chlamydia* spp., *Mycoplasma* spp., *Mycobacterium* spp., *Bartonella* spp., *Coxiella Burnetii*, fungi
- Polymicrobial infection in an IVDA
- Empirical antimicrobial therapy for BCNIE
- Periannular extension
- Prosthetic valve IE
- Persistent metastatic foci of infection (abscesses)
- Resistance to conventional antibiotic regimens
- Positive valve culture
- Persistence of fever at the seventh postoperative day
- Chronic dialysis

BCNIE = blood culture-negative infective endocarditis; IE = infective endocarditis; IVDA = intravenous drug abuser.

# Cardiac device related infective endocarditis: diagnosis, treatment and prevention (1)

CIED: cardiovascular Implantable Electronic Device

CDRIE: Cardiac device related infective endocarditis



Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
<b>A. Diagnosis</b>			
1. Three or more sets of blood cultures are recommended before prompt initiation of antimicrobial therapy for CIED infection	I	C	
2. Lead-tip culture is indicated when the CIED is explanted	I	C	
3. TOE is recommended in patients with suspected CDRIE with positive or negative blood cultures, independent of the results of TTE, to evaluate lead-related endocarditis and heart valve infection	I	C	
4. Intracardiac echocardiography may be considered in patients with suspected CDRIE, positive blood cultures and negative TTE and TOE results	IIb	C	
5. Radiolabelled leucocyte scintigraphy and <sup>18</sup> F-FDG PET/CT scanning may be considered additive tools in patients with suspected CDRIE, positive blood cultures and negative echocardiography	IIb	C	

# Cardiac device related infective endocarditis: diagnosis, treatment and prevention (2)

CIED: cardiovascular Implantable Electronic Device  
 CDRIE: Cardiac device related infective endocarditis

<b>B. Principles of treatment</b>			
1. Prolonged (i.e. before and after extraction) antibiotic therapy and complete hardware (device and leads) <b>removal</b> are recommended in definite CDRIE, as well as in presumably isolated pocket infection	I	C	
2. Complete hardware removal should be considered on the basis of <b>occult infection</b> without another apparent source of infection	IIa	C	
3. In patients with NVE or PVE and an intracardiac device with no evidence of associated device infection, complete hardware extraction may be considered	IIb	C	
<b>C. Mode of device removal</b>			
1. Percutaneous extraction is recommended in most patients with CDRIE, <b>even those with vegetations &gt; 10 mm</b>	I	B	382, 391, 405

Continued



# Cardiac device related infective endocarditis: diagnosis, treatment and prevention (3)

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
2. Surgical extraction should be considered if percutaneous extraction is incomplete or impossible or when there is associated severe destructive tricuspid IE	IIa	C	
3. Surgical extraction may be considered in patients with large vegetations (>20 mm)	IIb	C	
<b>D. Reimplantation</b>			
1. After device extraction, reassessment of the need for reimplantation is recommended	I	C	
2. When indicated, definite reimplantation should be postponed if possible, to allow a few days or weeks of antibiotic therapy	IIa	C	
3. A 'temporary' ipsilateral active fixation strategy may be considered in pacemaker-dependent patients requiring appropriate antibiotic treatment before reimplantation	IIb	C	
4. Temporary pacing is not routinely recommended	III	C	

TL: Habib G et al. Eur. H. J 2015.  
doi/10.1093/eurheartf/ ehv 319

# Cardiac device related infective endocarditis: diagnosis, treatment and prevention (4)

E. Prophylaxis			
1. Routine antibiotic prophylaxis is recommended before device implantation	I	B	367, 368, 373
2. Potential sources of sepsis should be eliminated $\geq 2$ weeks before implantation of an intravascular/ cardiac foreign material, except in urgent procedures	IIa	C	

CDRIE = cardiac device-related infective endocarditis; CIED = cardiac implantable electronic device; FDG = fluorodeoxyglucose; IE = infective endocarditis; NVE = native valve endocarditis; PET = positron emission tomography; PVE = prosthetic valve endocarditis; TOE = transoesophageal echocardiography; TTE = transthoracic echocardiography.

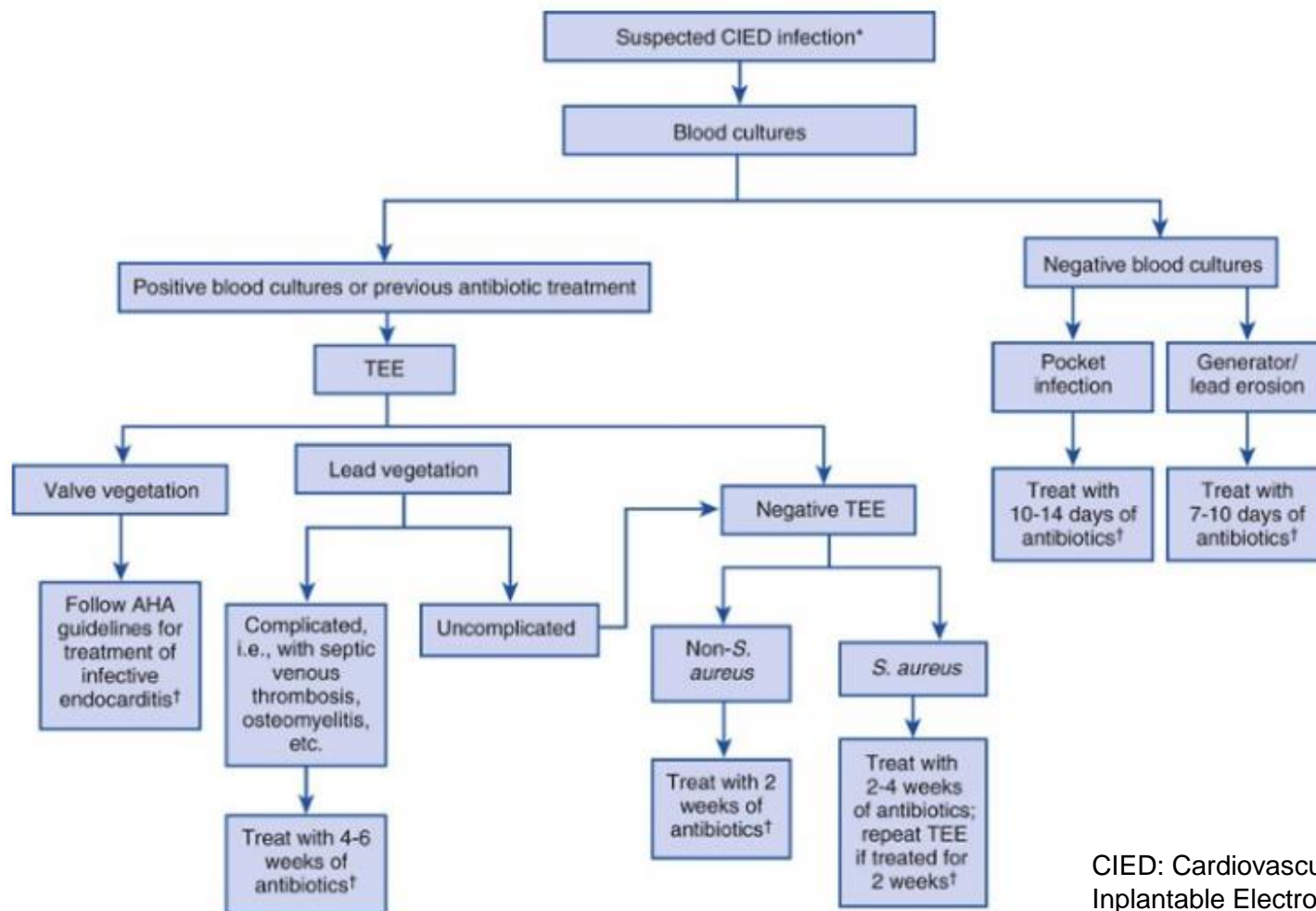
<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

<sup>c</sup>Reference(s) supporting recommendations.

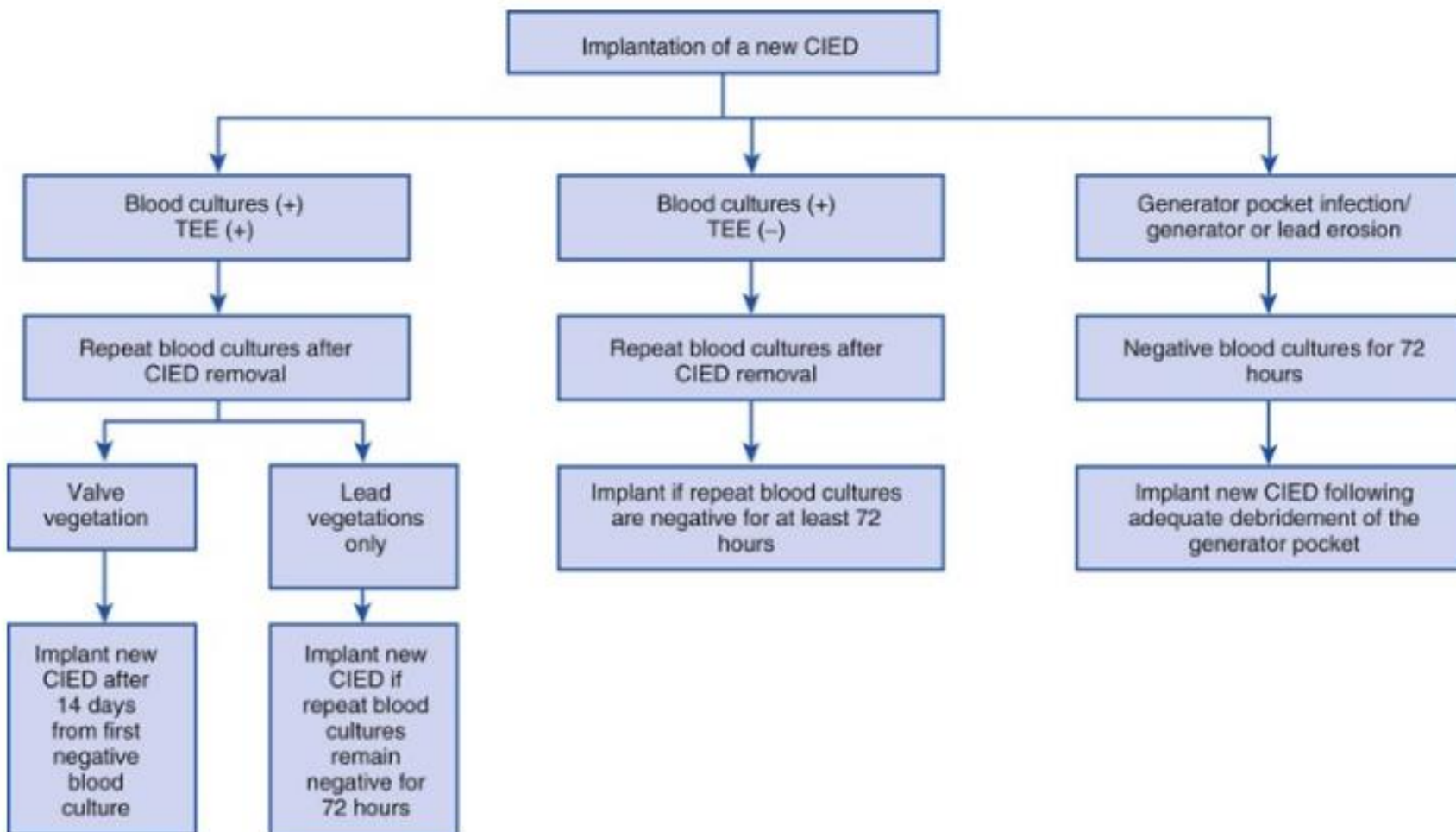
TL: Habib G et al. Eur. H. J 2015.  
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# Tiếp cận điều trị bệnh nhân người lớn bị nhiễm CIED



CIED: Cardiovascular Implantable Electronic Device

# Tiếp cận cấy dụng cụ mới sau gỡ bỏ CIED nhiễm trùng



# Indications for surgical treatment of right sided infective endocarditis



Recommendation	Class <sup>a</sup>	Level <sup>b</sup>
Surgical treatment should be considered in the following scenarios: <ul style="list-style-type: none"><li>• Microorganisms difficult to eradicate (e.g. persistent fung) or bacteraemia for &gt; 7 days (e.g. <i>S. aureus</i>, <i>P. aeruginosa</i>) despite adequate antimicrobial therapy or</li><li>• Persistent tricuspid valve vegetations &gt; 20 mm after recurrent pulmonary emboli with or without concomitant right heart failure or</li><li>• Right HF secondary to severe tricuspid regurgitation with poor response to diuretic therapy</li></ul>	IIa	C

HF = heart failure.

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

# Recommendations for the use of antithrombotic therapy (1)

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref. <sup>c</sup>
Interruption of antiplatelet therapy is recommended in the presence of major bleeding	I	B	257
In intracranial haemorrhage, interruption of all anticoagulation is recommended	I	C	
In ischaemic stroke without haemorrhage, replacement of oral anticoagulant (anti-vitamin K) therapy by unfractionated or low molecular weight heparin for 1–2 weeks should be considered under close monitoring <sup>d</sup>	IIa	C	

## Recommendations for the use of antithrombotic therapy (2)

In patients with intracranial haemorrhage and a mechanical valve, unfractionated or low molecular weight heparin should be reinitiated <u>as soon as possible following multidisciplinary discussion</u>	<b>IIa</b>	<b>C</b>	
In the <u>absence of stroke</u> , replacement of oral anticoagulant therapy by unfractionated or low molecular weight heparin for 1–2 weeks should be considered in the case of <i>Staphylococcus aureus</i> IE under close monitoring	<b>IIa</b>	<b>C</b>	
Thrombolytic therapy is not recommended in patients with IE	<b>III</b>	<b>C</b>	

IE = infective endocarditis.

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

<sup>c</sup>Reference(s) supporting recommendations.

<sup>d</sup>There is very limited experience with new oral anticoagulant treatment in the field of IE.

## Recommendations for the use of antithrombotic therapy (3)

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>
<b>1. Prophylaxis/prevention</b>		
<p>Antibiotic prophylaxis should be considered for patients at highest risk for IE</p> <ul style="list-style-type: none"> <li>a. Patients with any prosthetic valve, including transcatheter valve, or those in whom any prosthetic material was used for cardiac valve repair</li> <li>b. Patients with a previous episode of IE</li> <li>c. Patients with congenital heart disease (i.e. any type of cyanotic congenital heart disease or any type of congenital heart disease repaired with a prosthetic material)</li> </ul>	<b>IIa</b>	<b>C</b>
Antibiotic prophylaxis is not recommended in other forms of valvular or congenital heart disease	<b>III</b>	<b>C</b>



# Recommendations for the use of antithrombotic therapy (4)

Dental procedures		
Antibiotic prophylaxis should only be considered for dental procedures requiring manipulation of the gingival or periapical region of the teeth or perforation of the oral mucosa	IIa	C
Antibiotic prophylaxis is not recommended for local anaesthetic injections in non-infected tissues, treatment of superficial caries, removal of sutures, dental X-rays, placement or adjustment of removable prosthodontic or orthodontic appliances or braces, or following the shedding of deciduous teeth or trauma to the lips and oral mucosa	III	C
Other procedures		
Antibiotic prophylaxis is not recommended for respiratory tract procedures, including bronchoscopy or laryngoscopy, transnasal or endotracheal intubation, gastroscopy, colonoscopy, cystoscopy, vaginal or caesarean delivery, TOE or skin and soft tissue procedures	III	C

TL: Habib G et al. Eur. H. J 2015. doi/10.1093/eurheartf/ ehv 319

# Cheá ñoã phoøng ngöøa VNTMNT treân beänh nhaân coù laøm thuû thuaät ôu raêng, mieäng vaø ñöông hoâ haáp treân



Setting	Regimen Administered 30-60 Min before Procedure*
Standard regimen <sup>†</sup>	Amoxicillin 2.0 gm PO
Amoxicillin/penicillin-allergic patients	Cephalexin 2 gm PO <sup>†</sup> <i>or</i> Azithromycin or clarithromycin 500 mg PO <i>or</i> Clindamycin 600 mg PO
Patients unable to take oral medications	Ampicillin 2.0 gm IM or IV <i>or</i> Cefazolin or ceftriaxone 1 gm IV <sup>†</sup>
Ampicillin/amoxicillin/penicillin-allergic patients unable to take oral medications	Clindamycin 300 mg IV 30 min before procedure, then 150 mg 6 hr after initial dose

# Các trường hợp bệnh có nguy cơ cao VNTMNT cần phòng ngừa bệnh khi làm thủ thuật về răng/ khuyến cáo AHA 2007

- Prosthetic cardiac valve
- Previous infective endocarditis
- Congenital heart disease
- Unrepaired cyanotic congenital heart disease, including those with paliative shunts and conduits
- Completely repaired congenital heart disease with prosthetic material or device either by surgery or catheter intervention during the first 6 months after the procedure
- Repaired congenital heart disease 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

# Tiên lượng



- Yếu tố tiên lượng xấu:
  - Tác nhân gây bệnh không phải là Streptococci
  - Xuất hiện suy tim
  - Tổn thương van ĐMC
  - Van nhân tạo
  - Tuổi già
  - Có áp xe cơ tim hay vòng van
- VNTMNT do Staphylococcus aureus trên người không nhiễm ma túy # 40% tử vong
- Mảnh sùi lớn: tiên lượng xấu
- VNTMNT sớm/van nhân tạo xấu hơn
- VNTMNT chậm/van nhân tạo

# Predictors of poor outcome in patients with infective endocarditis (1)

## Patient characteristics

- Older age
- Prosthetic valve IE
- Diabetes mellitus
- Comorbidity (e.g., frailty, immunosuppression, renal or pulmonary disease)

## Clinical complications of IE

- Heart failure
- Renal failure
- >Moderate area of ischaemic stroke
- Brain haemorrhage
- Septic shock



# Predictors of poor outcome in patients with infective endocarditis (2)

## Microorganism

- *Staphylococcus aureus*
- Fungi
- Non-HACEK Gram-negative bacilli

## Echocardiographic findings

- Periannular complications
- Severe left-sided valve regurgitation
- Low left ventricular ejection fraction
- Pulmonary hypertension
- Large vegetations
- Severe prosthetic valve dysfunction
- Premature mitral valve closure and other signs of elevated diastolic pressures

HACEK = *Haemophilus parainfluenzae*, *H. aphrophilus*, *H. paraphrophilus*, *H. influenzae*, *Actinobacillus actinomycetemcomitans*, *Cardiobacterium hominis*, *Eikenella corrodens*, *Kingella kingae*, and *K. denitrificans*; IE = infective endocarditis.

# Đặc điểm về chẩn đoán và điều trị VNTMNT/người cao tuổi

- Triệu chứng cơ năng và thực thể không rầm rộ, khó phát hiện do có bệnh nội khoa kèm theo (TD: Phế khí thũng, bệnh phổi mạn tắc nghẽn...)
- Cây máu sớm và SAT qua thành ngực, nhất là SATQTQ rất cần thiết
- Trắc nghiệm PCR trên DNA của vi trùng cần thiết/ một số trường hợp
- Sử dụng thuốc theo phác đồ không khác người trẻ: chỉnh liều theo độ lọc cầu thận và chức năng gan
- Theo dõi sát các biến chứng bằng lâm sàng và siêu âm tim