

## Infective Endocarditis (IE)

**Definition:** Infective endocarditis (IE) is an inflammation of the endocardium, the innermost layer of the heart and the lining of heart valves.

**Primary Cause:** Usually microbial infection, but there are also non-infectious types, such as NBTE (Non-Bacterial Thrombotic Endocarditis).

**Historical Context:** Once termed "bacterial endocarditis" until it was found fungi and non-bacterial agents could cause similar conditions.

Three Independent Predisposing Factors for IE:

### 1. Transient Bacteremia:

Can be nosocomial (hospital-acquired) or induced by invasive medical procedures or inappropriate antibiotic use.

Preventive antibiotics are often prescribed when IE risk is identified.

### 2. Structural Heart Defects:

Defects can allow bacteria to adhere to and colonize the heart.

### 3. Vegetation Formation:

Infected areas become encapsulated by a fibrin clot, cutting off blood supply, making it a metabolically inactive infection resistant to immune clearance.

IE as a Rare, Life-threatening Disease:

It has long-term effects on survivors and is deadly without treatment.

Can involve multiple organ systems and become fatal if untreated.

**Dental Prophylaxis:** Antibiotics are commonly prescribed before dental procedures to prevent oral flora from entering the bloodstream.

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## Epidemiology

**Incidence:** Roughly 1-10 cases per 100,000 person-years.

**Risk Factors by Region:**

Low-income countries: Rheumatic heart disease remains a primary risk factor (up to two-thirds of cases).

High-income countries: Degenerative valve disease, diabetes, cancer, intravenous drug use, and congenital heart disease are now leading risk factors.

**Age Trends:**

Mean patient age has increased (now >50 years), likely due to improved health and longevity.

## Mortality:

High mortality if untreated; even with treatment, IE has about a 25% mortality rate.

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## Causative Agents of IE

### Predominant Organisms:

Gram-positive cocci account for over 90% of cases worldwide.

Developed countries: *Staphylococcus aureus* is most common.

Developing countries: *Streptococcus viridans* is more frequent.

Enterococci (e.g., *E. faecalis*, *E. faecium*) and *Streptococcus bovis* account for 5-10% of cases.

### Prosthetic Valves:

Common infection site; infections often by *S. aureus*.

IV Drug Use: Right-sided (tricuspid) valve IE is common in IV drug users.

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## Predisposing Factors for Endocarditis

Historical Factor: Rheumatic disease, caused by Group A *Streptococcus pyogenes*, commonly seen in children after strep throat.

### Modern Factors:

Congenital heart disease, prosthetic heart valves, pacemakers, post-pneumonia/meningitis.

Periodontal disease or dental procedures.

Medical procedures (e.g., dental extractions, implants, hemodialysis, cystoscopy, colonoscopy).

Infections in IV Drug Users: Infections from skin or mucosal flora introduced to the bloodstream.

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## Microbiology Overview

### Key Bacterial Causes:

*Staphylococcus* and *Streptococcus* species are responsible for ~80% of IE cases.

### *Staphylococcus aureus*:

Most common cause in high-income countries and common in IV drug users.

### Other Bacterial Groups:

HACEK group (*Haemophilus*, *Aggregatibacter*, *Cardiobacterium*, *Eikenella*, *Kingella*): Rare, healthcare-associated, Gram-negative organisms.

Fungal Causes (e.g., *Candida albicans*, *Aspergillus*): Rare but severe, typically in hospitalized or immunosuppressed patients.

Microbiological Differentiation:

Staphylococcus vs. Streptococcus:

Catalase test: Staphylococcus (+), Streptococcus (-).

Staphylococcus species are further classified by coagulase (e.g., *S. aureus* is coagulase-positive).

Types of IE in Prosthetic Valve Patients:

1. Early (within 1 month post-surgery): Commonly due to coagulase-negative Staphylococcus species, like *S. epidermidis*.

2. Late (after 1 month): Primarily due to *S. aureus*.

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Microbial Causes Summary

Gram-positive Cocci: Common pathogens include Staphylococcus, Streptococcus, Enterococcus.

Viridans streptococci (oral flora) frequently cause IE in developing regions.

Fungal Endocarditis:

Often related to *Candida*, especially in catheterized or immunosuppressed patients.

Diagnosed through culturing on specific media (e.g., Sabouraud dextrose agar).

Yeast and Pseudohyphae:

*Candida* can exist in both yeast and pseudohyphal forms, depending on the environment.

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Pathophysiology of IE

Mechanism:

Bacteria enter the bloodstream, adhere to altered heart valves, and form vegetations (biofilms).

Vegetations can cause septic emboli, ischemia, or infarction in various organs.

Clinical Significance:

Biofilms protect bacteria from the immune system and require prolonged IV antibiotic therapy for effective treatment.

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## Clinical Features

### Diverse and Non-specific Presentation:

IE diagnosis is often delayed due to its non-specific symptoms.

Acute IE: Rapidly damages cardiac structures and is often fatal within weeks if untreated.

Subacute IE: Progresses slowly unless complicated by major embolic event or ruptured mycotic aneurysm, rarely metastasizes, and can take months to develop complications.

### Key Indicators:

Fever of unknown origin with a new or changing heart murmur suggests IE until proven otherwise.

### Manifestations:

#### Cardiac:

Heart murmurs, valvular damage, congestive heart failure in 30-40% of cases.

#### Noncardiac:

Janeway Lesions: Painless, due to septic emboli.

Osler's Nodes: Painful, immune-mediated.

Subungual Hemorrhage and conjunctival Hemorrhage: Signs of vascular involvement.

Roth's Spots: retinal hemorrhages with clear centers, an immunologic phenomenon.

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## Diagnosis

### Core Diagnostic Tools:

Blood Culture: Essential first step; IE typically causes continuous bacteremia.

Echocardiography: Should be done for all suspected IE cases.

### Modified Duke Criteria:

Combines pathologic, major, and minor criteria for diagnosis:

Pathologic: Microorganisms identified in vegetations.

Major: Positive blood cultures and endocardial involvement on imaging.

Minor: Fever, predisposing heart condition, vascular phenomena, immunologic phenomena, microbiologic evidence.

### Diagnosis Confirmation:

Requires combinations of criteria: 1 pathologic or 2 major, or 1 major + 3 minor, or all 5 minor criteria.

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## Management

### Antimicrobial Therapy:

Vancomycin and Gentamicin should be initiated immediately after blood culture. Prolonged IV therapy often required due to biofilm complications.

### Surgical Treatment:

Corrective surgery may be necessary for cardiac defects to prevent future IE episodes.

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## Prevention

### Antibiotic Prophylaxis:

Previously recommended for many procedures; now typically reserved for high-risk patients. Policies vary by country (e.g., the UK restricts prophylaxis to high-risk patients).

