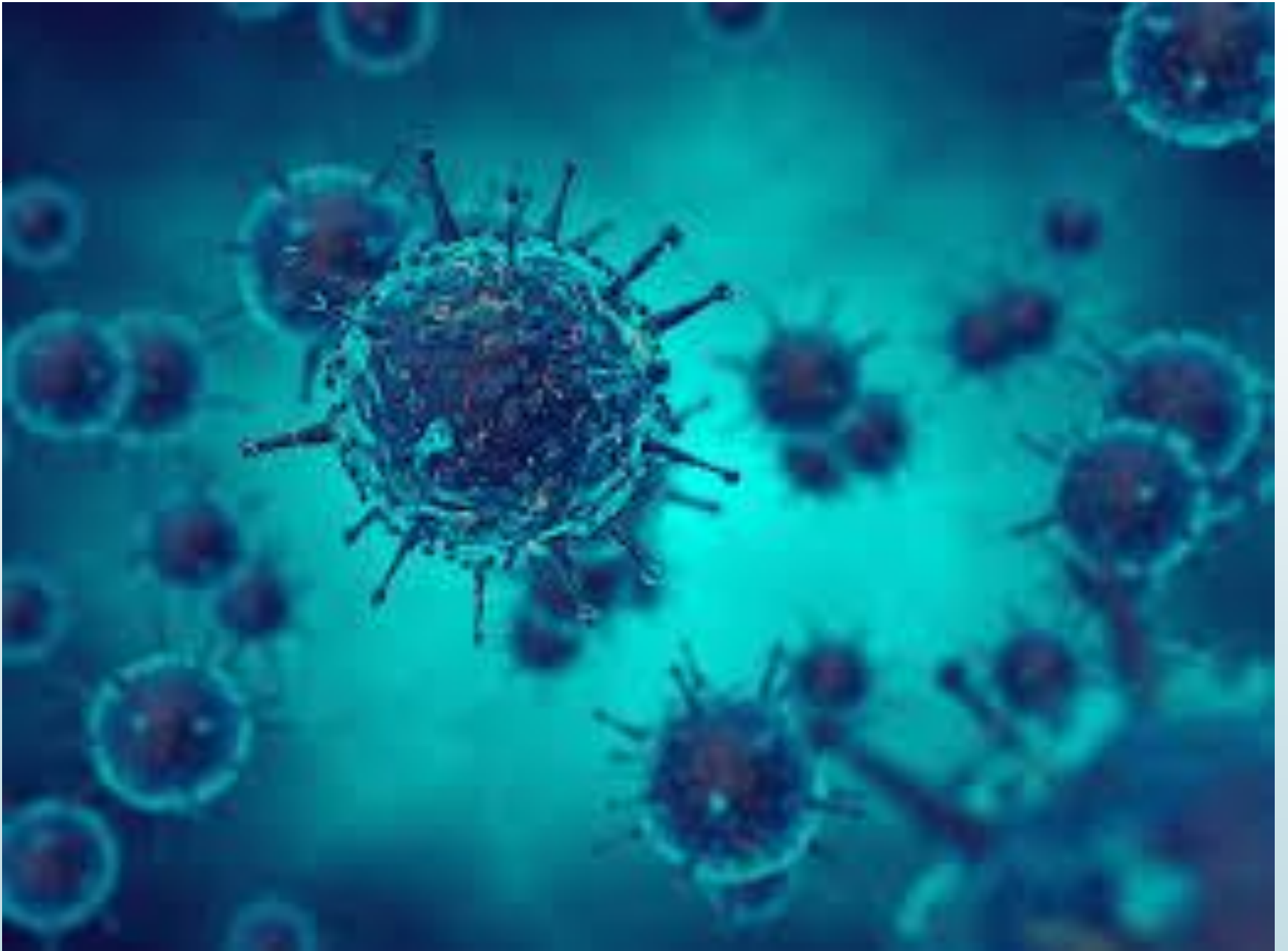
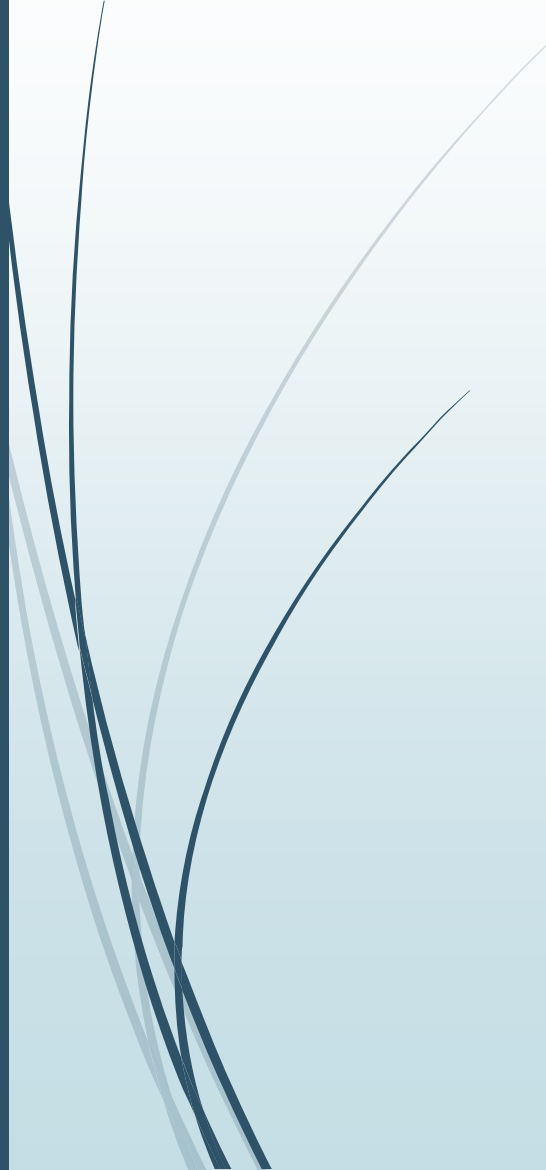




Hospital Acquired Infectious Diseases

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Objectives

- Nosocomial infection (HAI)
- CDC hot list
- Common nosocomial pathogens
- Modes of transmission
- Antibiotic-resistant nosocomial pathogens
- Control of nosocomial infections

Introduction for Nosocomial Infections

- ▶ Acquired during hospital stay
- ▶ Asymptomatic and symptomatic patient can be considered infected
- ▶ Infection may be carried by:
 - ▶ Healthcare workers
 - ▶ Family members
- ▶ New antibiotic research is decreasing
- ▶ Account for 5% to 10% infection rates in USA
- ▶ > 40% in other developing countries
- ▶ Huge \$\$ cost involved in treatment





Types of nosocomial infections

- The CDC has classified 13 types of nosocomial infections based on biological and clinical data:
 - UTI
 - Surgical sites
 - Gastroenteritis
 - Meningitis
 - Respiratory infections
- Increasing rates occur because of cancer therapy, organ transplants, immunotherapy, invasive diagnostic techniques
- Not so good news! HAI have increased 17% to 30% in last 5 years

Types of nosocomial infections

- ▶ Bacteria are responsible for about 90% of infections
- ▶ Protozoans, fungi, viruses and mycobacteria are less contributing compared to bacterial infections
- ▶ Out of these- these are of “concern”
 - ▶ Enterococci
 - ▶ *P. aeruginosa*
 - ▶ *S. aureus*
 - ▶ *E. coli*





Commonly isolated nosocomial pathogens found in health care facilities

- *Acinetobacter baumannii*
- *Clostridium difficile*
- Carbapenem-Resistant *Enterobacteriaceae* (CRE)
- *Escherichia coli*
- *Klebsiella pneumoniae*
- Methicillin-Resistant *Staphylococcus aureus*
- *Pseudomonas aeruginosa*
- *Staphylococcus aureus*
- Vancomycin-Resistant *Enterococci*



Virulence factors

- ▶ Virulence factors:
 - ▶ Innate ability of bacteria to survive and thrive
 - ▶ Invade the host
 - ▶ Cause disease
- ▶ Adherence Factors
 - ▶ Many pathogenic bacteria colonize mucosal sites by using pili(fimbriae) to adhere to cells
- ▶ Toxins
 - ▶ Endotoxins
 - ▶ Part of cell wall of gram negative bacteria
 - ▶ Create fever and inflammation
 - ▶ Exotoxins
 - ▶ Secreted by bacteria causing inhibition of immune responses
 - ▶ E-coli and C-Diff
- ▶ Destructive enzymes- break down cell walls
- ▶ Immune modulators- break down the host's immunoglobulins using proteases

S. aureus

- *Staphylococcus* genus is considered one of the most important pathogens
- Gram-positive cocci, non-spore forming, catalase- and coagulase-positive, immotile, facultatively anaerobe
- Mainly colonizes in nasal passages and skin





S. aureus

- 20% individuals have persistent colonization of *S. aureus*
- Decreased immunity and immunocompetent patients
- *S. aureus* infects not only the superficial but also the deep tissues
 - Toxin-mediated diseases of *S. aureus* include:
 - Food poisoning
 - Toxic shock syndrome
 - Scalded skin syndrome
- Virulence mechanisms of *S. aureus* include:
 - Toxins
 - Enzymes
 - Immune modulators



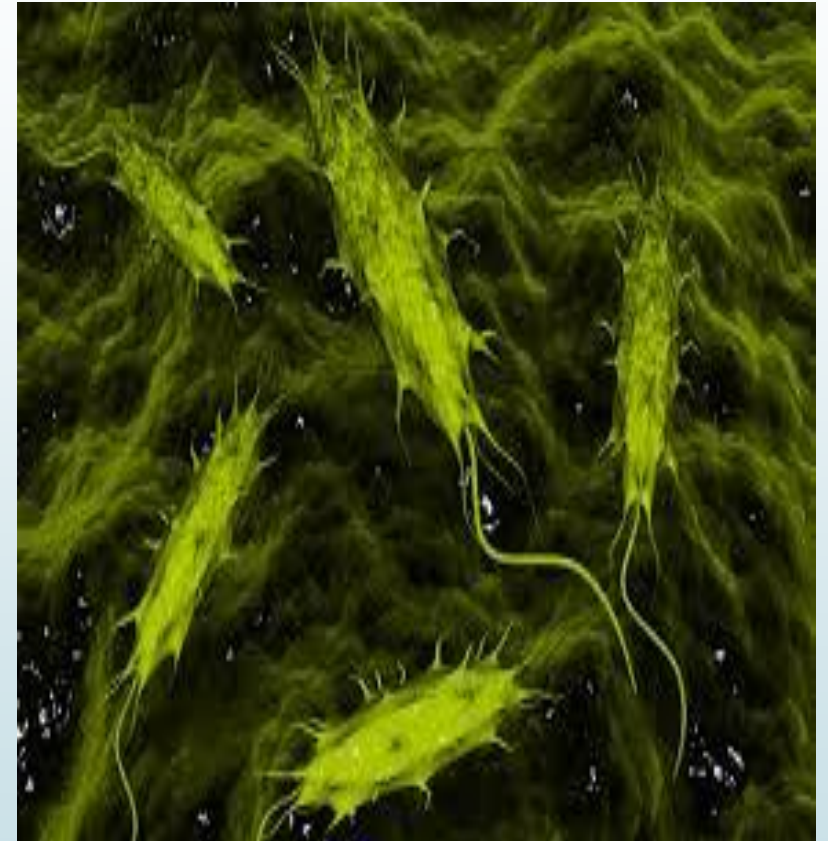


S. aureus

- ▶ In hospitals these infections can be fatal:
 - ▶ Bacteremia or sepsis when bacteria spread to the bloodstream
 - ▶ Pneumonia, which predominantly affects people with underlying lung disease including those on mechanical ventilators
 - ▶ Endocarditis (infection of the heart valves), which can lead to heart failure or stroke
 - ▶ Osteomyelitis (bone infection), which can be caused by staph bacteria traveling in the bloodstream

E-coli

- *E. coli* is an emerging nosocomial pathogen
- *E. coli* is Gram-negative and oxidase-negative facultative anaerobe bacteria
- Colonize in GI tract of human beings and other animals





E-coli

- *E. coli* is responsible for a number of diseases:
 - UTI
 - Septicemia
 - Pneumonia
 - Neonatal meningitis (Group B Streptococci)
 - Peritonitis
 - Gastroenteritis
- Virulence factors are endotoxins, capsule, adhesions and type 3 secretion systems
- Specialized virulence factors are seen in cases of UTI and gastroenteritis

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Vancomycin-Resistant Enterococci (VRE)

- Vancomycin-Resistant Enterococci
- 17 different species of Enterococci
- Resistant to Vancomycin
 - Used to treat infections caused by enterococci
- Normally present in the human intestines and in the female genital tract
- Can sometimes cause infections
- Most Vancomycin-Resistant Enterococci infections occur in hospitals

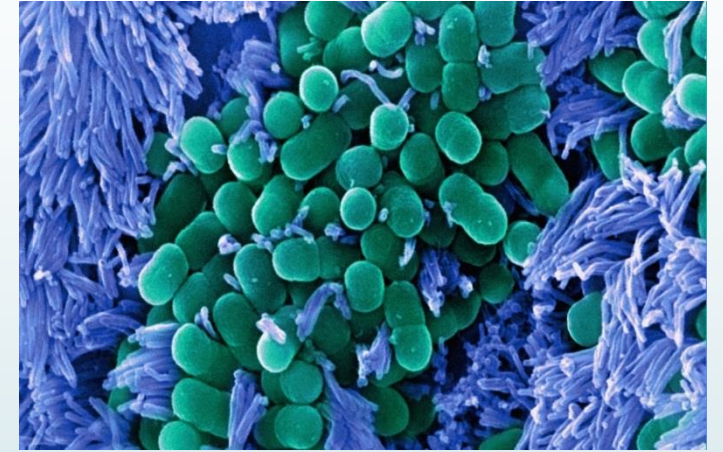
Enterococci

- ▶ 2nd leading cause of hospital acquired infections worldwide
 - ▶ Nursing home admissions?
- ▶ 20%–30% of all infections in the USA
- ▶ Facultative anaerobic Gram-positive enteric microbes
- ▶ Part of normal microbiota in female genital tract and gastrointestinal tract as well
- ▶ Involved in the blood-borne infections; UTI, wound infections, and surgical procedures infections
- ▶ Virulence factors include extracellular surface proteins, cytolysin, adhesions, hemolysins, gelatinase, extracellular superoxide and aggregation substances



Clostridium difficile (C. difficile)

- Nosocomial pathogen which mainly causes diarrhea worldwide
- Gram-positive bacillus
- Anaerobic and spore-forming bacteria
- Colonizes in intestinal tract and serves as part of normal microbiota
- Diseases caused by toxins produced by *C. difficile*:
 - Colitis 15%–25% cases
 - Diarrhea
- Major virulence factors for *C. difficile* are toxins, fimbriae, capsule and hydrolytic enzymes





Modes of transmission

- ▶ *S. aureus*
 - ▶ Infected individuals' skin or contact via shared items and surfaces like door handles, benches, towels and taps
- ▶ *E. coli*
 - ▶ Transmitted through person to person, environment or contaminated water and food
- ▶ Vancomycin-resistant enterococci
 - ▶ Patients with diarrhea
 - ▶ Room items such as surfaces and equipment's act as reservoirs
 - ▶ Can survive on these surfaces for days or weeks



Modes of transmission

- Klebsiella pneumonia
 - Transmitted by person-to-person contact
 - Healthcare professionals do not wash or clean hands after checking a contaminated patient
 - Ventilators, catheters or exposed wounds can be the source of its transmission
 - Transmitted through stool , patients' hands and pharynx
- P. aeruginosa
 - Common reservoirs include breast pumps, incubators, sinks and hands of hospital staff and hand soaps

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Modes of transmission

- *C. difficile*
 - Spores of *C. difficile* can hold for months
 - Infected intestinal patients are major sites acted as reservoirs
 - Inanimate objects such as door knobs, stethoscopes, ventilators, etc
 - Hospital staff along with hospital settings are also part of the problem



Antibiotic-resistant nosocomial pathogens

- Multi-drug-resistant nosocomial organisms include
 - MRSA
 - Vancomycin-Resistant Enterococci
 - *P. aeruginosa*
 - *K. pneumonia*
 - *C. difficile* shows natural resistance
- 50% to 60% of hospital-acquired infections are caused by resistant pathogens
- Improper use of antibiotics is the major cause of this drug resistance

Antibiotic-resistant nosocomial pathogens

➤ MRSA

- β -Lactamase antibiotics including penicillin became resistant in the 1940s
- Resistance of penicillin due to its improper use
- Staphylococcal species have penicillinase enzyme that makes penicillin infective
 - Penicillinase-resistant antibiotics were developed such as cephalosporin's and methicillin's
- By the 1960s, methicillin-resistant species of *S. aureus* were first reported
- Resistance was due to the modification of penicillin-binding proteins
- β -lactam antibiotics along with their derivatives are now ineffective

Antibiotic-resistant nosocomial pathogens

- ▶ Vancomycin-Resistant Enterococci
 - ▶ Resistance in the enterococcal species due to the *vanA* and *vanB* genes
 - ▶ Part of plasmid and would spread resistance to other microbes
 - ▶ Enterococci are resistant to many different classes of antibiotics which include:
 - ▶ Penicillin
 - ▶ Ampicillin
 - ▶ Aminoglycosides
 - ▶ Tetracycline's
 - ▶ Carbapenem
 - ▶ Fluoroquinolones
 - ▶ Macrolides
 - ▶ Linezolid, daptomycin, and tigecycline current treatment options

Antibiotic-resistant nosocomial pathogens

- *P. aeruginosa* is becoming resistant due to different mechanisms
 - These mechanisms include:
 - Restricted uptake of drug
 - Modification and altered targets sites for antibiotics
- Due to this increasing resistance, complications are seen in these infections
- Ineffective drugs showing resistance:
 - Cephalosporin's
 - Trimethoprim
 - Macrolides
 - Chloramphenicol
 - Tetracycline's
 - Fluoroquinolones
- Current treatment combination of Cephalosporins and Aminoglycoside



Antibiotic-resistant nosocomial pathogens

- ▶ *Klebsiella pneumoniae*
 - ▶ Resistance to β -lactam antibiotics
 - ▶ 3rd and 4th generation Cephalosporins show resistance for *K. pneumoniae*
 - ▶ Treatment with combination therapy Aminoglycosides and Imipenem
- ▶ *C. difficile*
 - ▶ Increased use of broad-spectrum antibiotics is associated with resistance
 - ▶ Improper antibiotic use was the cause of increasing infections of *C. difficile*
 - ▶ Treatment with Vancomycin

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Control of nosocomial infections

- Need statistics on antimicrobial susceptibility
- Resistant organisms make it extremely difficult to devise a proper plan for control
- Each hospital has variant strains making it difficult to compare and share information
- Different healthcare settings at some hospitals don't share data

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Need to know the types of microorganism involved in infection

- ▶ Infectious organisms are heterogeneous in nature which makes them different from one another
- ▶ Compare the infection rates
 - ▶ Type of healthcare settings is important and services they provide
 - ▶ Admissions from nursing homes
- ▶ Management of infections differs greatly between institutions



Development of infection control programs

- Guidelines for the sterilization and disinfection
- Guidelines for the prevention of catheter-associated UTI
- Create protocols that reduce the transmission of nosocomial infections
- Surveillance of nosocomial infections along with investigation of major outbreaks
 - CDC Helps with this
- Training of healthcare professionals is extremely important for the control and prevention of infection

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Surveillance of nosocomial infections

- Ongoing, systematic collection, analysis, and interpretation of health data
- Control the infections by evaluating the efficacy of treatment and clusters of breakout infections
- Surveillance can lead to better infection control practices
- People practice better infection control practices if they are being ‘watched’



Summary

- Even with the best antibiotics, nosocomial infections are still uncontrollable
- Control of organisms responsible for nosocomial infections is much needed as they cause great economic as well as production loss
- Transmission of these infections in the hospital settings through healthcare workers can be avoided by the use of infection control practices
- Improper and frequent use of antibiotics is an important cause of drug-resistant organisms that are difficult to treat
- Hospitals should devise the infection control programs through which infection rates can be compared and controlled
- A well-managed surveillance methodology is required
- Best practice should be shared among hospitals to stop the spread of nosocomial infections.



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