



اللهم وفقني في دراستي، ونور بالكتاب بصري، وأشرح به
صدري، واستعمل به بدني، وأطلق به لساني، وقوّ به عزمي
بجولك وقوتك، فإنه لا حول ولا قوة إلا بك يا أرحم الراحمين

كل ما في السلايدات مطلوب حفظه و الملخصات
المذكورة في الملف هي وسيلة لتسهيل ربط المعلومات و
تضمن ملاحظات حكمتها الدكتور

وفقكم الله...

Biofilms

Pharmaceutical Microbiology

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Thin & Slimy Colonies or Communities of bacteria Biofilms
in which adheres to a surface are capable to produce their own extracellular matrix

Definition

ECM Provided certain benefits to the bacteria

- Biofilms are communities of surface-associated microorganisms encased in a self-produced extracellular matrix. ECM

embedded in ECM Substratum قاعدة بعض certain surface على بكون

- Biofilm formation is a nearly universal bacterial trait, and biofilms are found on almost all natural and artificial surfaces.

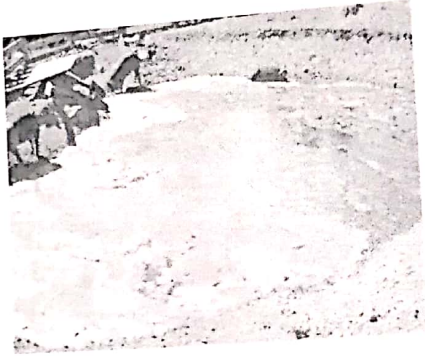
التجمع البكتيري مع بعض Colonies شكل لكن طبيعياً كـ مجموع من خلية جانب خلية

- Some examples of biofilms are the plaque that forms on teeth and ① the slime that forms on surfaces in watery areas. ②

Biofilms → Beneficial to the bacteria
→ Harmful to Humans (cause disease)

Plaque هو تراكم Secretion على الأسنان
Bacterial colony هو وسيلة ولازم سهم تتلفها دوماً

Examples of biofilms



Old Science

- Biofilms science and technology has been an active field of study since the late seventies when the first definition of biofilms was brought to public attention by Bill Costerton and coworkers in 1978

- **Significance: Protect and allow bacteria to survive and thrive in hostile environments.** Withstand host immune responses, much less susceptible to antibiotics and disinfectants.

→ They can for example exchange their genomes, resist antibiotic,

Survive in high threatening condition

low nutrition, change in temperature & pH

Yellow is the *Staphylococcus aureus*



High-resolution scanning electron microscopy image of a community associated methicillin-resistant *Staphylococcus aureus* biofilm embedded in extracellular matrix, formed on implanted catheter tubing during experimental biofilm infection

methicillin-resistant

Staphylococcus aureus

Catheter tube وبكون موجود بـ

Biofilm وهي embedded inside matrix

Human Infections linked to biofilms

هذه هي
الأمراض

Table 1 Partial list of human infections involving biofilms

Infection or disease	Common biofilm bacterial species
Dental caries	Acidogenic Gram-positive cocci (e.g., <i>Streptococcus</i>)
Periodontitis	Gram-negative anaerobic oral bacteria
Otitis media	Nontypable strains of <i>Haemophilus influenzae</i>
Musculoskeletal infections	Gram-positive cocci (e.g., <i>Staphylococcus</i>)
Necrotizing fasciitis	Group A <i>Streptococcus</i>
Biliary tract infection	Enteric bacteria (e.g., <i>Escherichia coli</i>)
Osteomyelitis	Various bacterial and fungal species—often mixed
Bacterial prostatitis	<i>E. coli</i> and other Gram-negative bacteria
Native valve endocarditis	Viridans group streptococci
Cystic fibrosis pneumonia	<i>P. aeruginosa</i> and <i>Burkholderia cepacia</i>
Melioidosis	<i>Pseudomonas pseudomallei</i>
Neurological infections	
LUL pneumonia	Gram-negative rods
Sutures	<i>Staphylococcus epidermidis</i> and <i>S. aureus</i>
Exit sites	<i>S. epidermidis</i> and <i>S. aureus</i>
Arteriovenous shunts	<i>S. epidermidis</i> and <i>S. aureus</i>
Schleral buckles	Gram-positive cocci
Contact lens	<i>P. aeruginosa</i> and Gram-positive cocci
Urinary catheter cystitis	<i>E. coli</i> and other Gram-negative rods
Peritoneal dialysis (CAPD) peritonitis	A variety of bacteria and fungi
IUDs	<i>Actinomyces israelii</i> and many others
Endotracheal tubes	A variety of bacteria and fungi
Hickman catheters	<i>S. epidermidis</i> and <i>C. albicans</i>
Central venous catheters	<i>S. epidermidis</i> and others
Mechanical heart valves	<i>S. aureus</i> and <i>S. epidermidis</i>
Vascular grafts	Gram-positive cocci
Biliary stent blockage	A variety of enteric bacteria and fungi
Orthopedic devices	<i>S. aureus</i> and <i>S. epidermidis</i>
Penile prostheses	<i>S. aureus</i> and <i>S. epidermidis</i>

هذه هي
أمراض
التي
تتسببها
العدوى
بـ

Dental caries & Otitis media & Some respiratory tract infection like Cystic fibrosis Pneumonia & Endocarditis

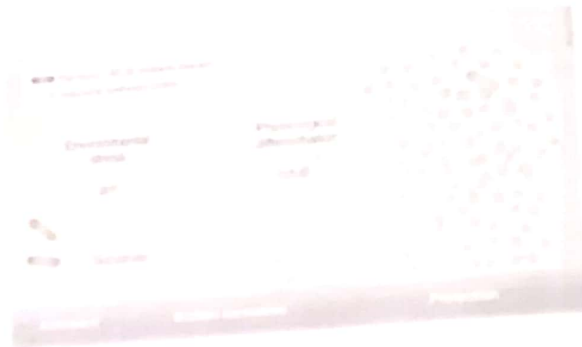
→ Harmful & slowly & Persistent infection

Biofilm formation and growth

- Basic structural unit of a biofilm is the microcolonies or cell cluster formed by the surface adherent bacterial cells.

- **IMPORTANT FACTORS IN BIOFILM FORMATION** Basic factors are:-

- ① • **Cell attachment**
- ② • **Extracellular polysaccharides (EPS)**
- ③ • **Cell-cell communication (Quorum sensing)**



cell clusters → Microcolonies → Basic structure unite all → Biofilm formation → Adhesion on surface → Biofilm

increasing of bacterial number because of further aggregation of bacteria

Protection of bacterial aggregation by ECM

Basic units of biofilm

- ① • **Microcolonies** of **homogenous** or **heterogenous** bacterial species.
- ② • **Matrix**: A **glycocalyx matrix** made up of **extracellular polymeric substance (EPS)** surrounds the microcolonies and anchors the bacterial cell to the substrate.

Functions:

- Maintains the integrity of biofilms
- Prevents desiccation.
- Resists antimicrobial agents
- Create a nutritionally rich environment

matrix can

- ① Provide full Protection
- ② Maintaining the integrity of biofilm
- ③ Prevent dehydration or desiccation
- ④ Resist antimicrobial agent
- ⑤ Create a nutritionally rich environment

Transferring of bacteria to reach surface

التي تستقر في السطح *Flagella*

Bacterial adhesion في السطح *Pili*

Quorum sensing

- ③ • is a communication mechanism between bacteria that allows specific processes to be controlled.
- Involves the regulation of expression of specific genes through the accumulation of signaling compounds that mediate intercellular communication.
Communication between bacteria so there is bacterial expression of certain gene through accumulation of signaling compound
- Dependent on cell density and mediated through signaling compounds.
- The signals are thought to allow cross-talk between species, causing them to increase their production of exo-polysaccharide and the factors that increase their virulence. ②

إشارات
Signals ①

Quorum Sensing → Expressing of certain gene are responsible for synthesis or formation of signaling compound which is important for communication between cells

Mechanisms of Biofilm formation



- Formation of a biofilm begins with the attachment of free-floating microorganisms to a surface. These first colonists adhere to the surface initially through weak, reversible van der Waals forces. If the colonists are not immediately separated from the surface, they can anchor themselves more permanently using cell adhesion structures such as pili

- in a nutrition containing fluid.
- 3 major component involve in biofilm formation:-

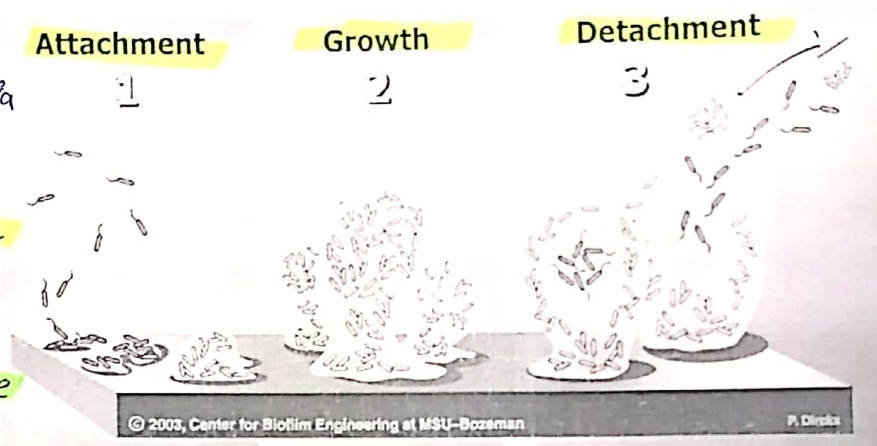
- ## Stages of Biofilm Formation

تَبُّ السُّيِّئَاتِ
Floating bacteria

that become
to be a sessile

State

State
↓
Law
↓
Community



STAGE 1

STAGE 1 Attachment

- Adsorption of inorganic and organic molecules to the solid surface, forming a thin layer termed as conditioning layer.

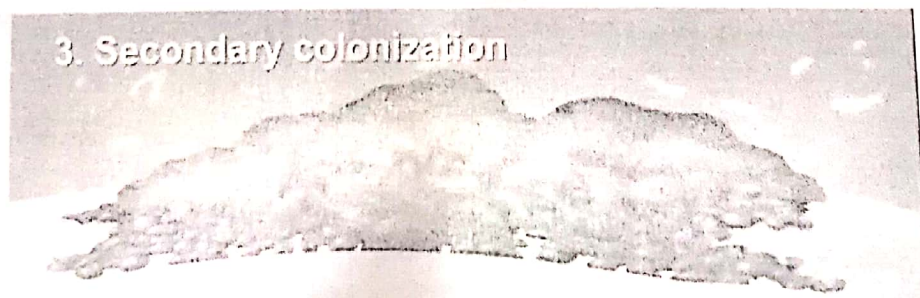
STAGE 2

- Adhesion of microbial cells to this layer.
 - Phase 1 - Transport of microbes to surface.
 - Phase 2 - Initial non specific microbial-substrate adherence phase.
 - Phase 3 - Specific microbial-substrate adherence phase.



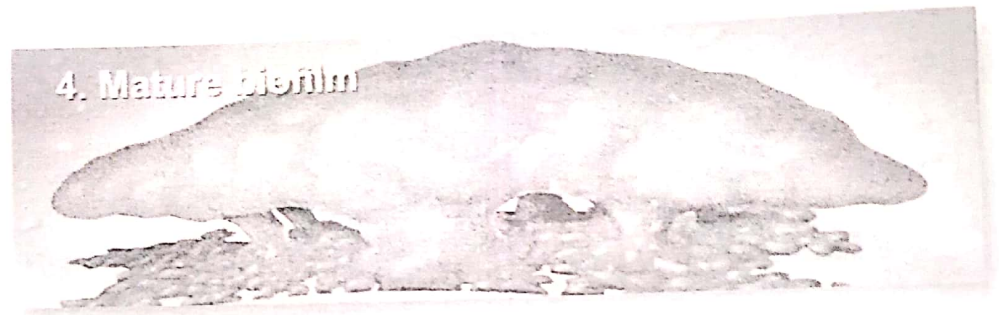
STAGE 3

- Bacterial growth and expansion.
- Monolayer of microbes attracts secondary colonizers forming microcolony, and the collection of microcolonies gives rise to the final structure of biofilm.

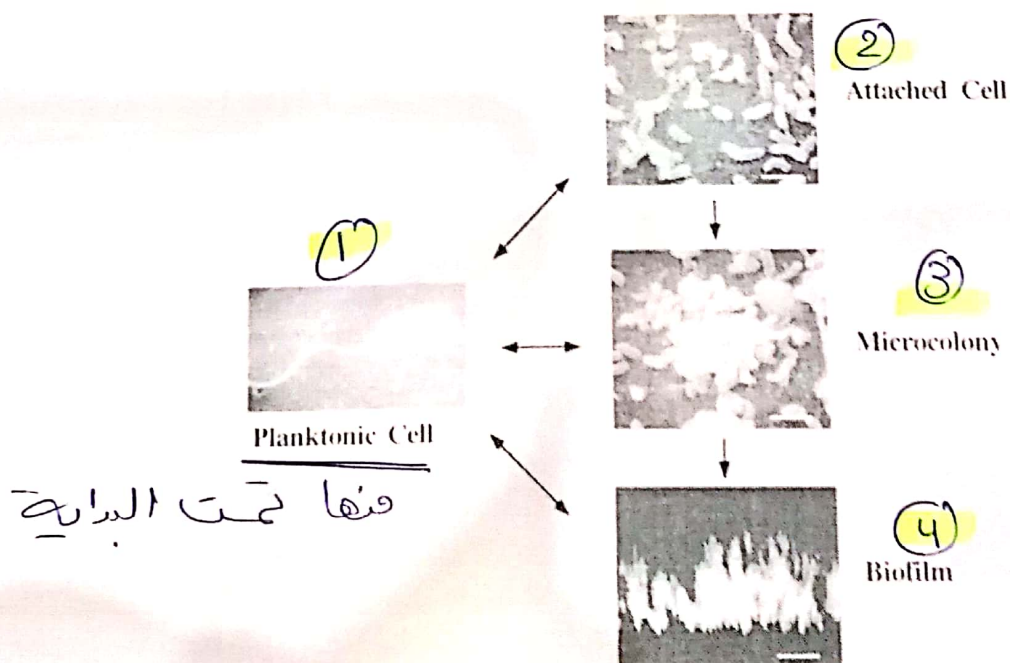


STAGE 3

- A mature biofilm will be a metabolically active community of microorganisms where individuals share duties and benefits.
- Some microorganisms help in adhering to the solid support, while some others create bridges between different species.



Biofilm formation in *Vibrio cholerae*



STAGE 4 : Detachment / Dispersion

خروج البكتيريا من
ومذا يساعد على
Biofilm Bacterial survival

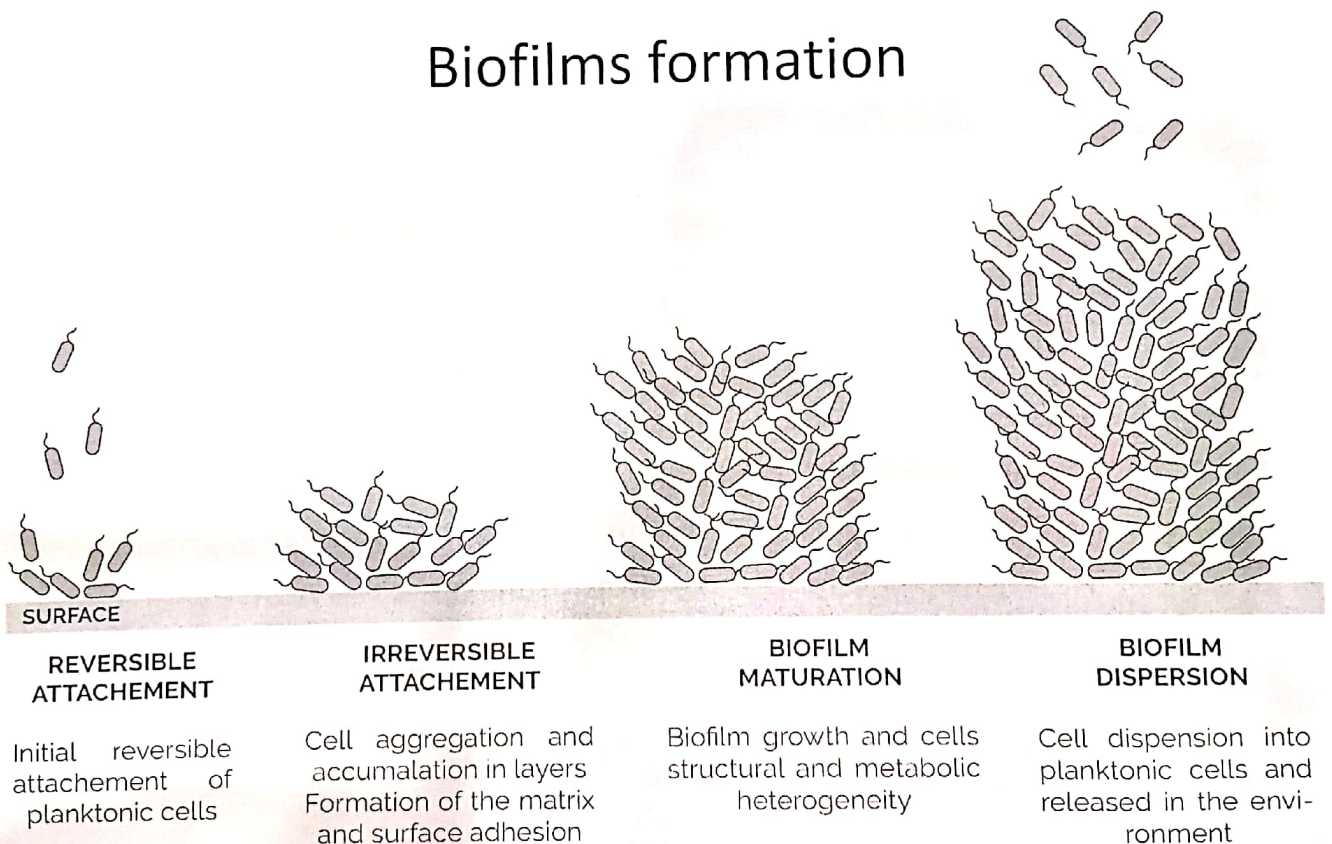
Disease spreading &

- During detachment, as a strategy of bacterial cells to leave biofilms and continue another biofilm life cycle

- Detachment plays an important role in shaping the morphological characteristics and structure of mature biofilm.



Biofilms formation



Benefits Of Biofilm To Microbes

- Helps the bacteria to survive in unfavorable environmental and nutritional conditions.
- Resistance to antimicrobial agents.
- Opportunity of genetic material exchange.
- Ability to communicate between bacterial population of same and or different species.
- Produce growth factors across species boundaries.

Role of biofilms in antibiotic resistance

How Biofilm resist antibiotics?

- Delayed penetration of the antimicrobial agent---EPS diffusional barrier ; delays rate of transport of drugs
- Bacteria intact in biofilms more resistant than single cells
- Altered growth rate of organisms in biofilms---organisms grow more slowly in biofilms---Slow uptake of antimicrobials---Actively growing and dividing cells are required for drug action

Slowly growing bacteria بكتيريا بطيئة النمو Biofilm دالة البكتيريا
 Alter growth of فز
 antimicrobial agent

Biofilms a Great threat to Implants

- A significant number of people are affected by biofilm infections which develop on medical devices implanted in the body such as catheters (tubes used to conduct fluids in or out of the body), artificial joints, and mechanical heart valves.
- When implanted material becomes colonized by microorganisms, a slow developing but persistent infection results

المسألة بالـ Biofilm <<< Infection

Endocarditis and Biofilms

↳ Infection of vascular endothelium of heart

- Microorganisms may attach and develop biofilms on components of mechanical heart valves and surrounding tissues of the heart, leading to a condition known as prosthetic valve endocarditis.
- The primary organisms responsible for this condition are S. epidermidis, S. aureus, Streptococcus spp., gram-negative bacilli, enterococci, and Candida spp.
- These organisms may originate from the skin, other indwelling devices such as central venous catheters, or dental work.

قبل ما الدكتور يحد Dental Procedure سألوا المريض
! إذا عنده Cardiac disease أو Valve replacement

Urinary catheters and Biofilms

- Urinary catheters are tubular latex or silicone devices, which when inserted may readily acquire biofilms on the inner or outer surfaces.
- The organisms commonly contaminating these devices and developing biofilms are **S. epidermidis**, **Enterococcus faecalis**, **E. coli**, **Proteus mirabilis**, **P. aeruginosa**, **K. pneumoniae**, and other **gram-negative** organisms.
- The longer the urinary catheter remains in place, the greater the tendency of these organisms to develop biofilms and result in urinary tract infections

كل ما دخلنا Solid device جسم العروق هاي
فرصة للبكتيريا لتتربى Biofilms

Longer time of replacement of catheter So high chance of development of biofilm

* نغيب البكتيريا الي ممكن تسبب Urinary tract infection
induce Biofilm formation