

# **Gram Negatív Pyogen Coccusok és Coccobacillusok**

**Dr. Berek Zsuzsa  
2008 Október 29**

# **Neisseria, Haemophilus, Bordetella**

## **1. Neisseria**

# Pyogen Coccusok GRAM -

**Aerob:** Oxidase +

**Neisseria**

**N. gonorrhoeae**

P

**N. meningitidis**

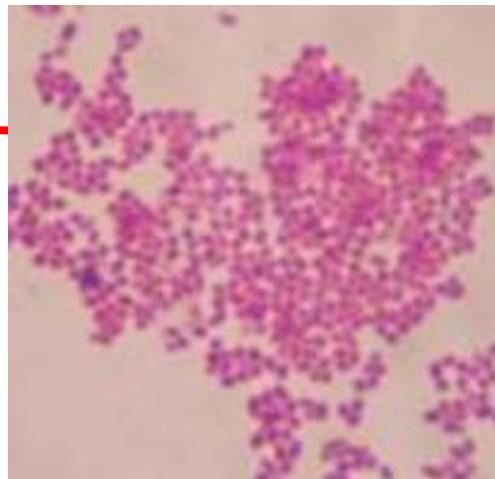
P

egyebek (N. sicca, N. subflava, N. flavesiens és  
apathogen fajok)

**Moraxella**

**M. catarrhalis**

**Anaerob:**



**Veillonella spp.**

**Veillonellae**

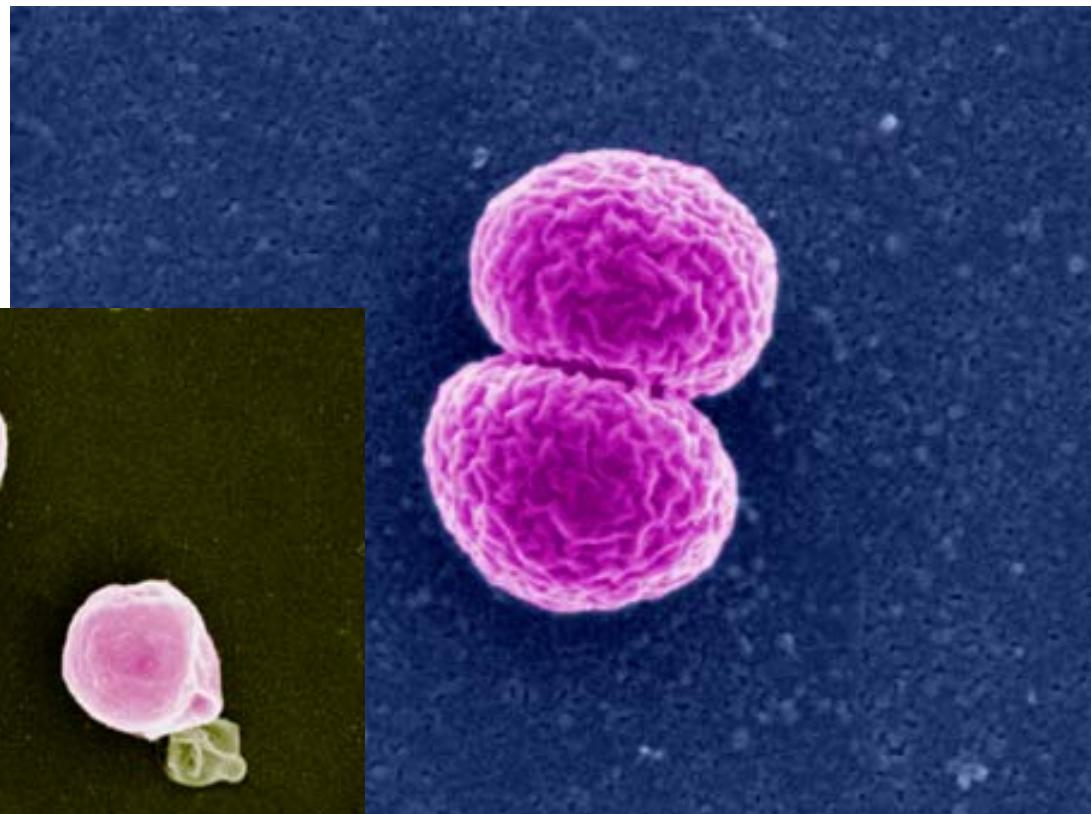
[vietsciences.free.fr](http://vietsciences.free.fr)

# N. gonorrhoeae és N. meningitidis

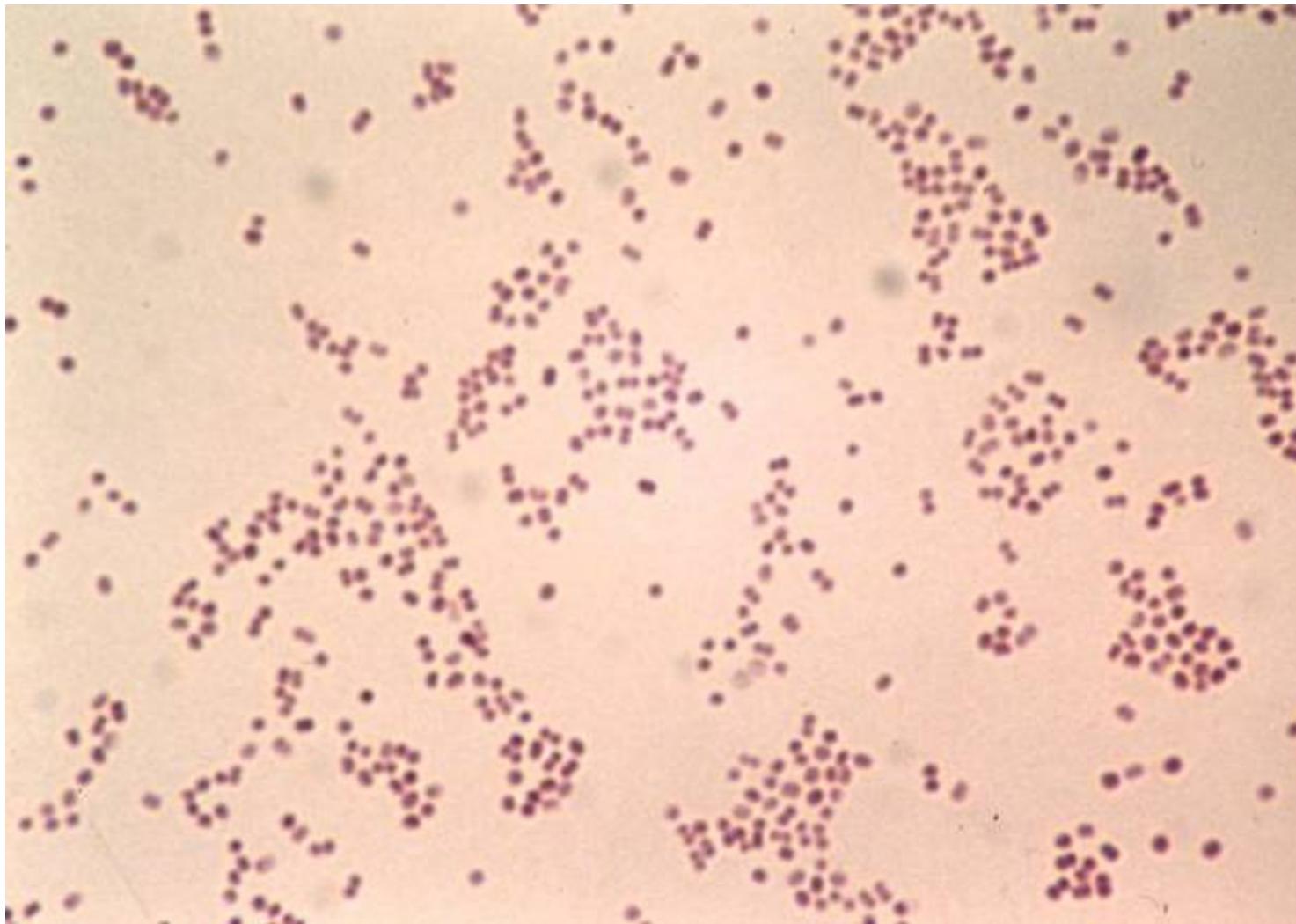
## Morphologia

Gram negatív

Diplococcus



# Gramnegativ Diplococcusok



# N. gonorrhoeae és N. meningitidis

## Tenyésztés:

Speciális, (Csokoládé agar,  
5-10% CO<sub>2</sub>)

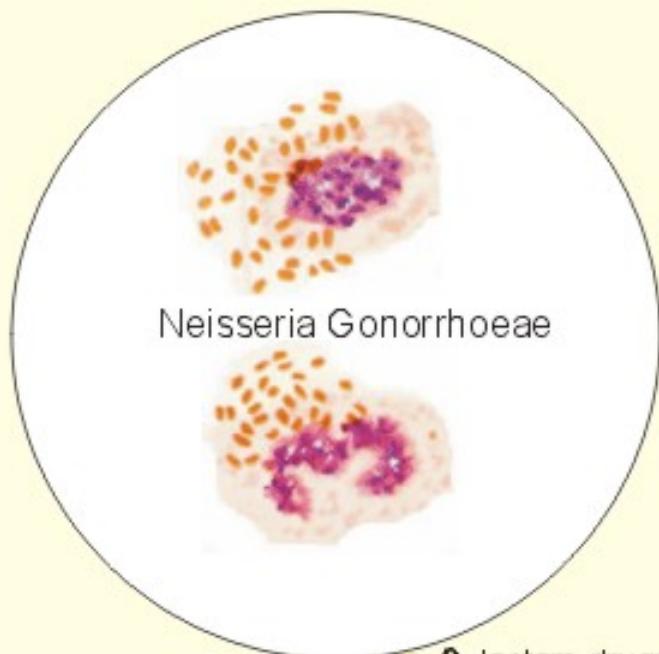
## Rezisztencia:

**Érzékenyek**, kiszáradásra,  
hőre, dezinficiensekre,  
antibiotikumokra

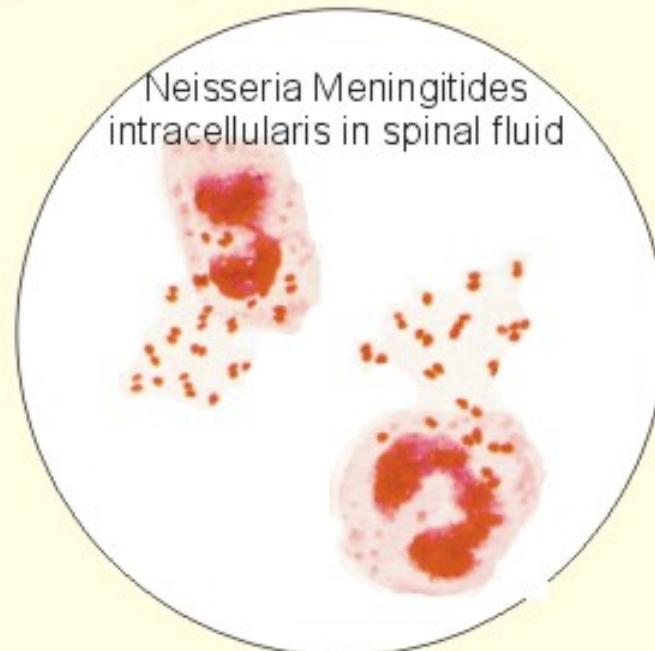
Oxidase +



## Microscopic Pictures Of *Neisseria* (Gram-negative Diplococci)



*Neisseria Gonorrhoeae*



*Neisseria Meningitidis*  
intracellularis in spinal fluid

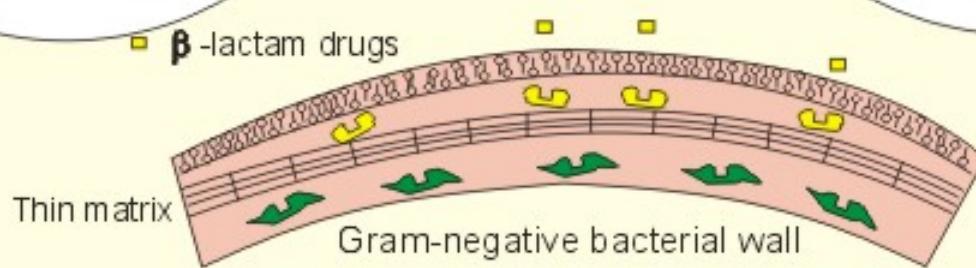


Fig. 33-3

KMc

# **N. gonorrhoeae = Gonococcus**

## **Antigének és Virulenciafaktorok:**

**Pili/Fimbriae** (Antigenvariációk!)

**IgA-Protease-ok!**

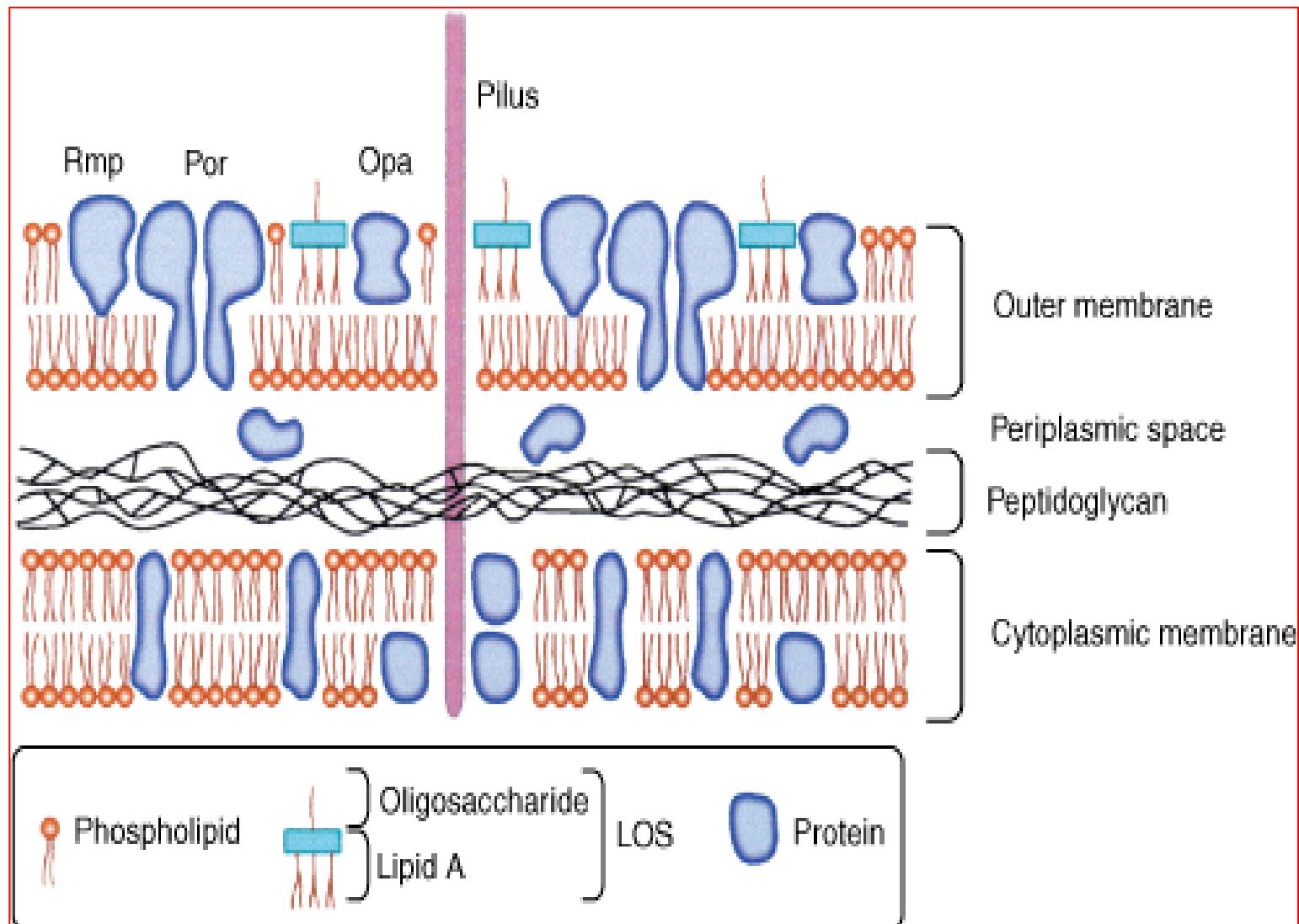
**outer membrane proteins (OMP)**

(Antigenvariációk!)

**LOS** (Mimikri!)

**Sejtfal Peptidoglycan** (Toxikus hatás)

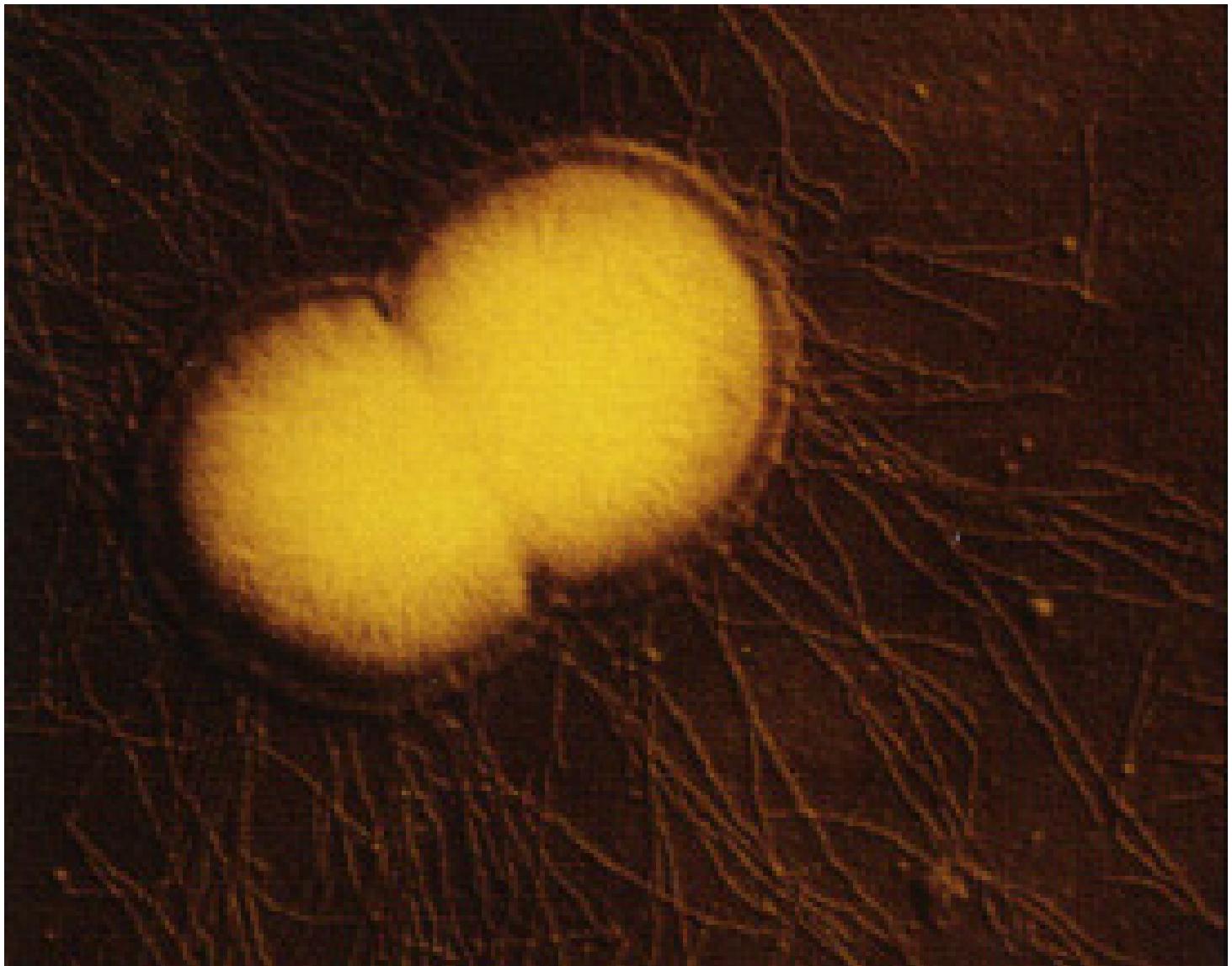
# N. gonorrhoeae = Gonococcus



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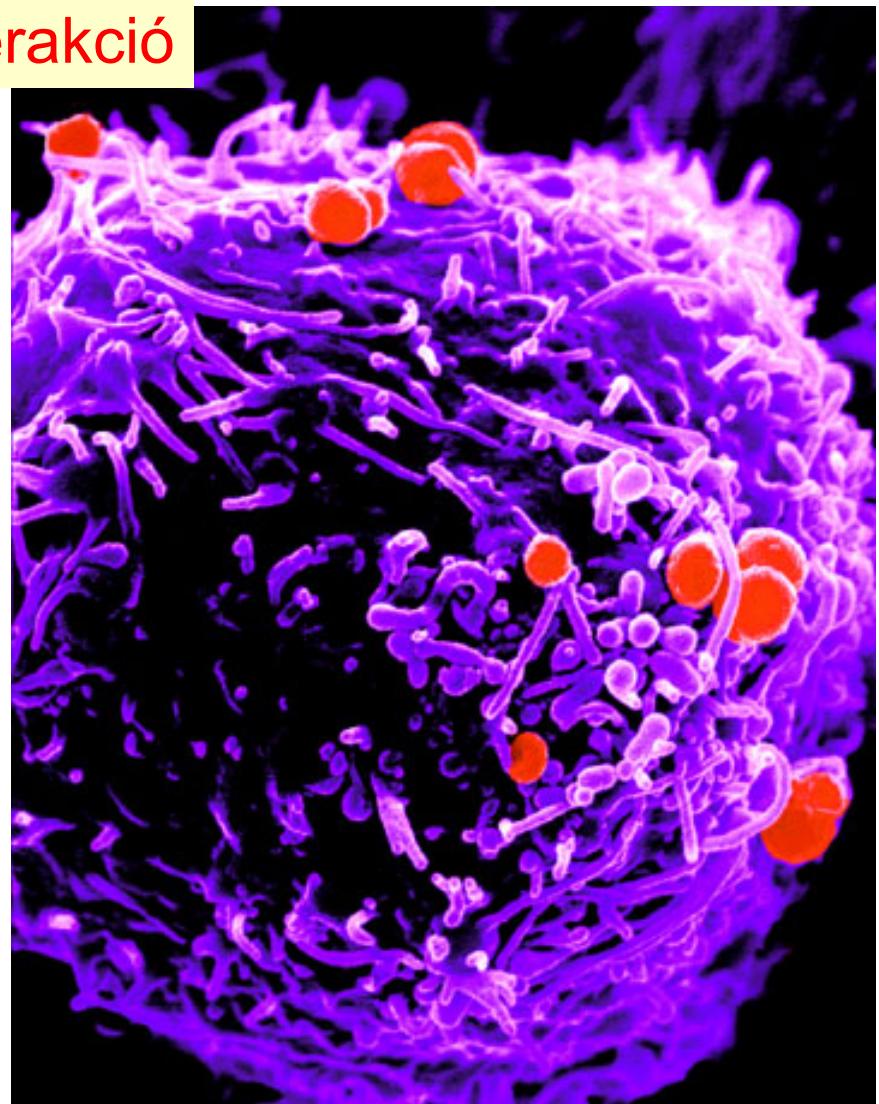
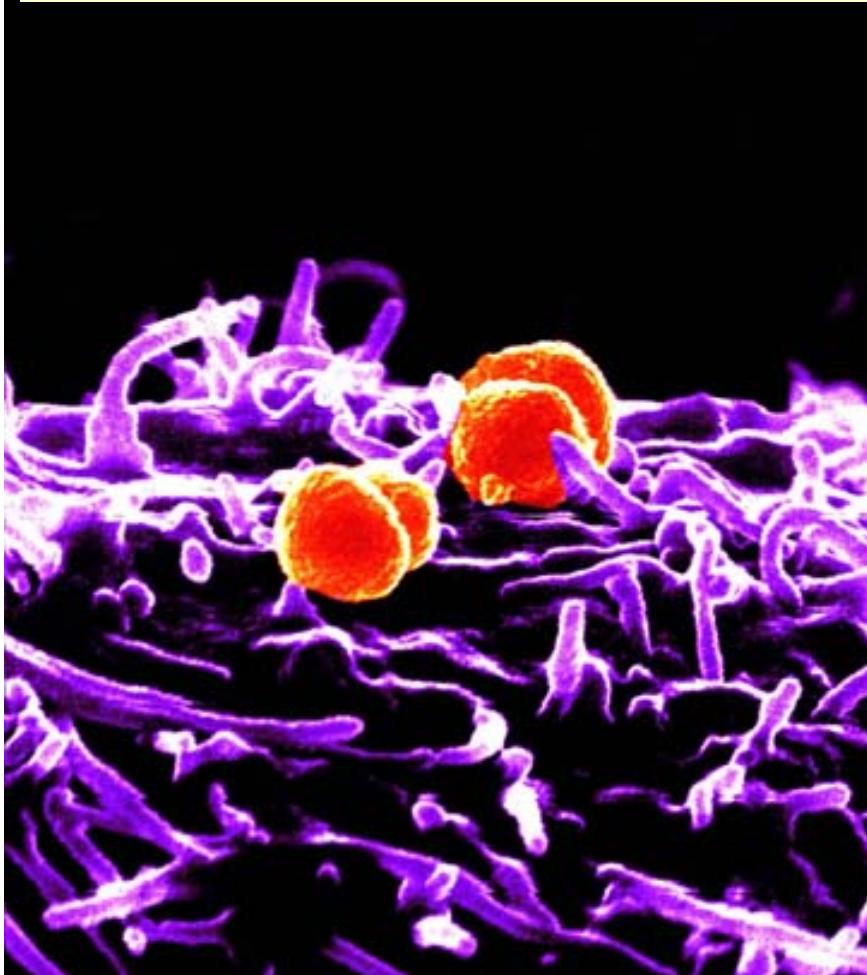
# **N. gonorrhoeae = Gonococcus**

**Pili**



# N. gonorrhoeae = Gonococcus

Gonococcus-Lymphocyta Interakció



# **N. gonorrhoeae = Gonococcus**

**Fertőzés forrása**  
beteg emberek

## **Átvitel**

- Direkt (szexuális) Kontaktus

## **Kórképek**

Gonorrhea = Kankó = Tripper

Ophthalmoblenorrhea neonatorum

Generalisatio 1%

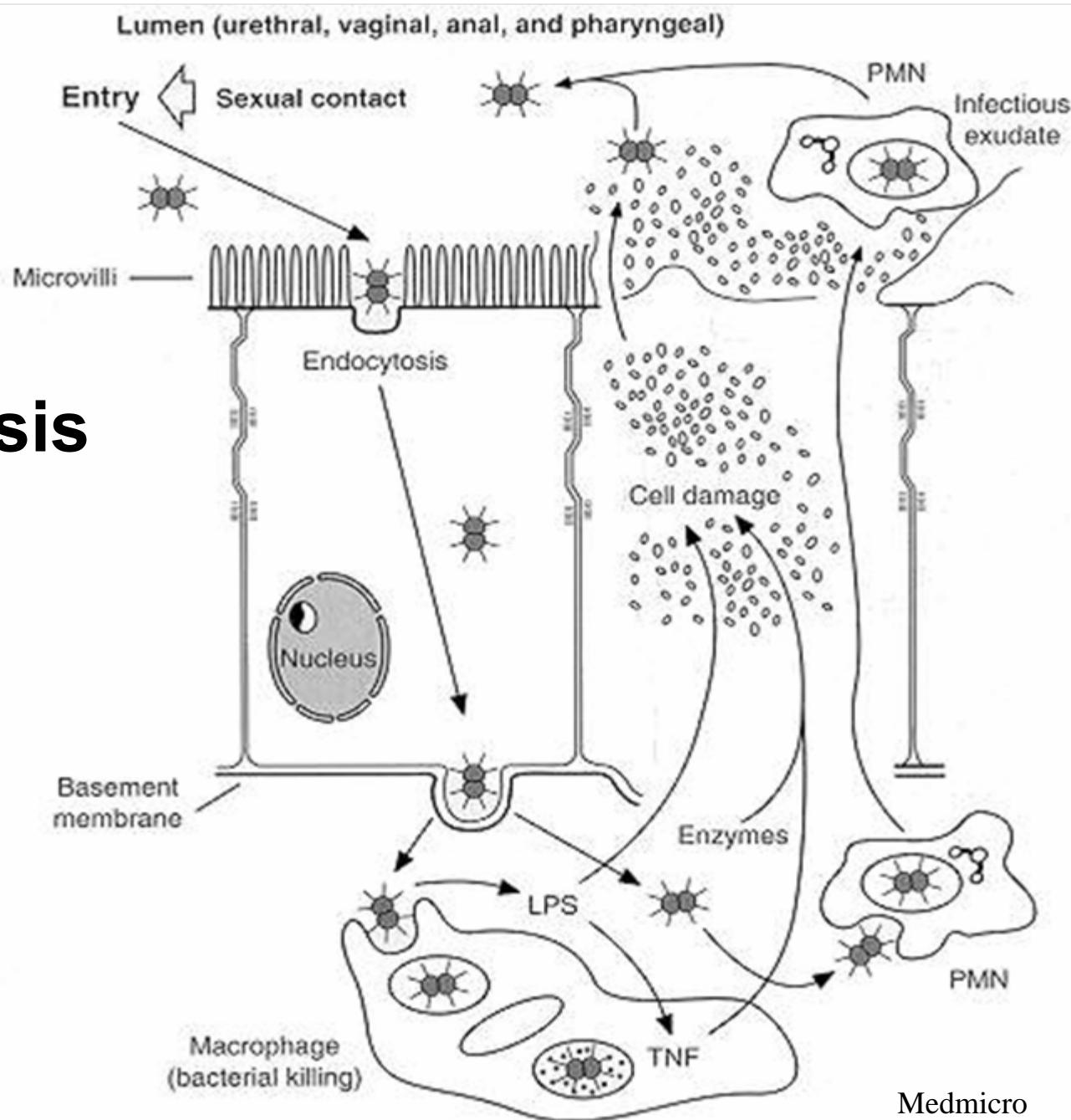
**NINCS IMMUNITÁS!**  
**(Antigenvariációk!)**



Die Erreger des Trippers (*Neisseria gonorrhoeae*, hier blau) werden von fingerförmigen Fortsätzen auf der Zelloberfläche (grün) umschlossen. Im weiteren Verlauf der Infektion dringen die Bakterien dann in die Zelle ein.

*Max-Planck-Institut für Infektionsbiologie, Volker Brinkmann*

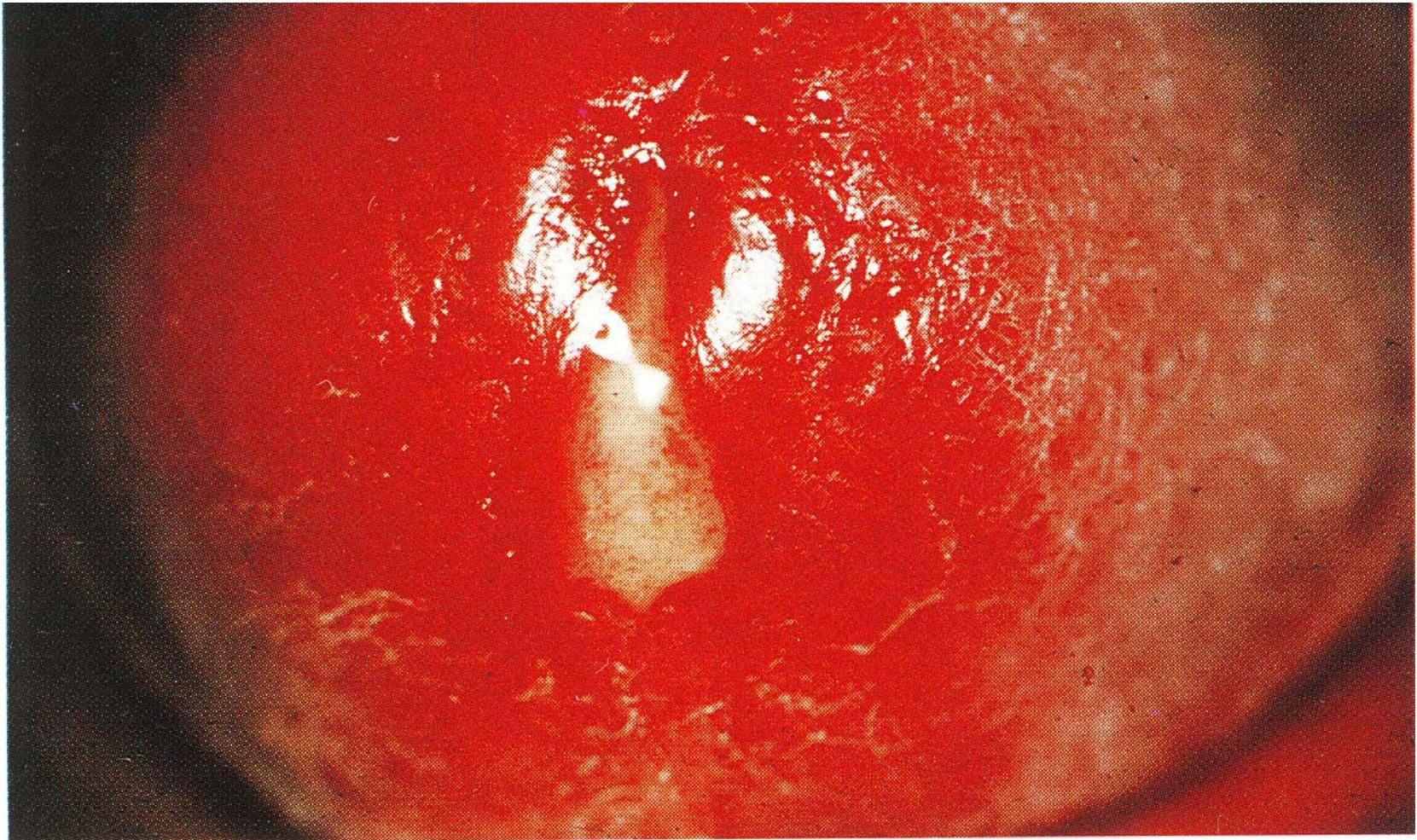
# Pathogenesis



# Gonorrhea – akut Urethritis

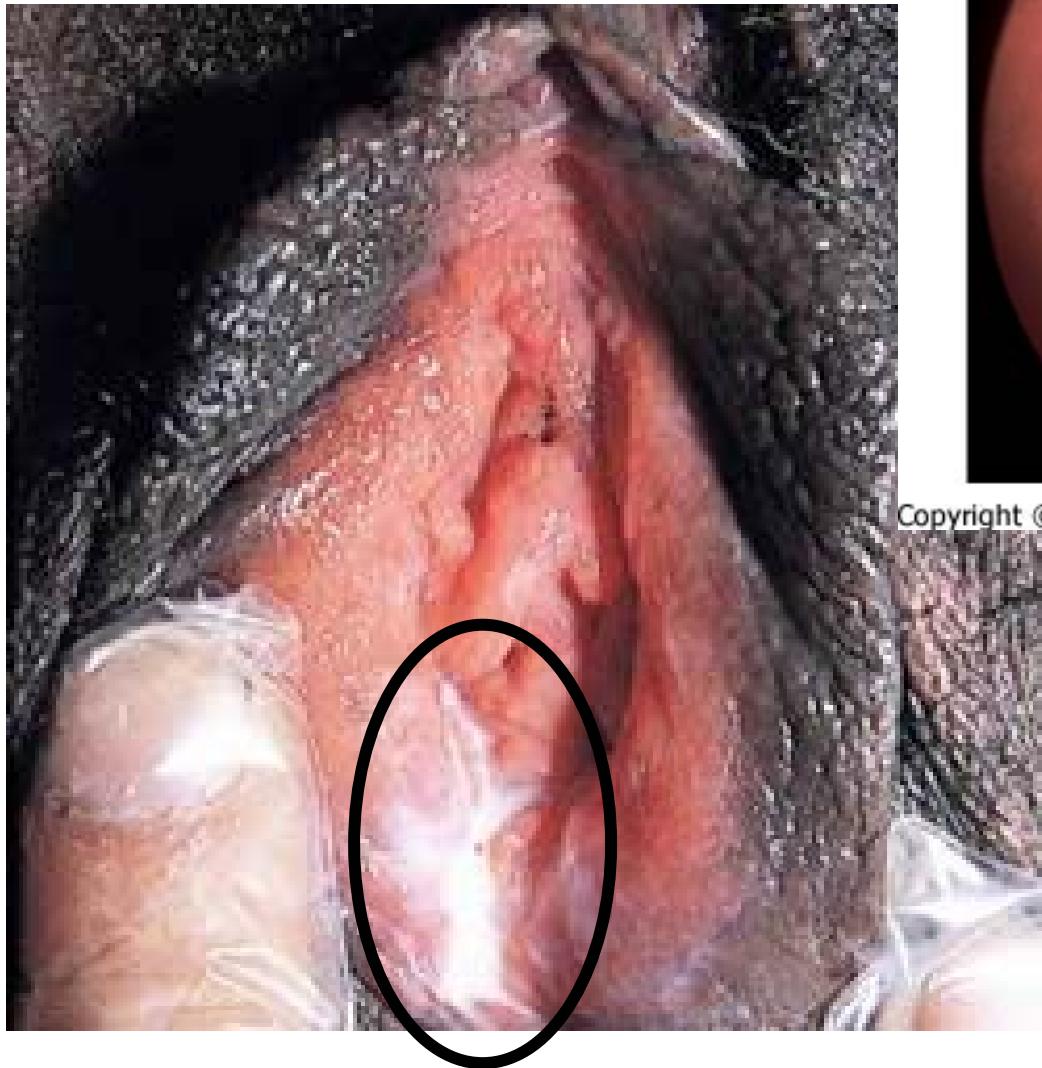


# Gonorrhea – akut Urethritis



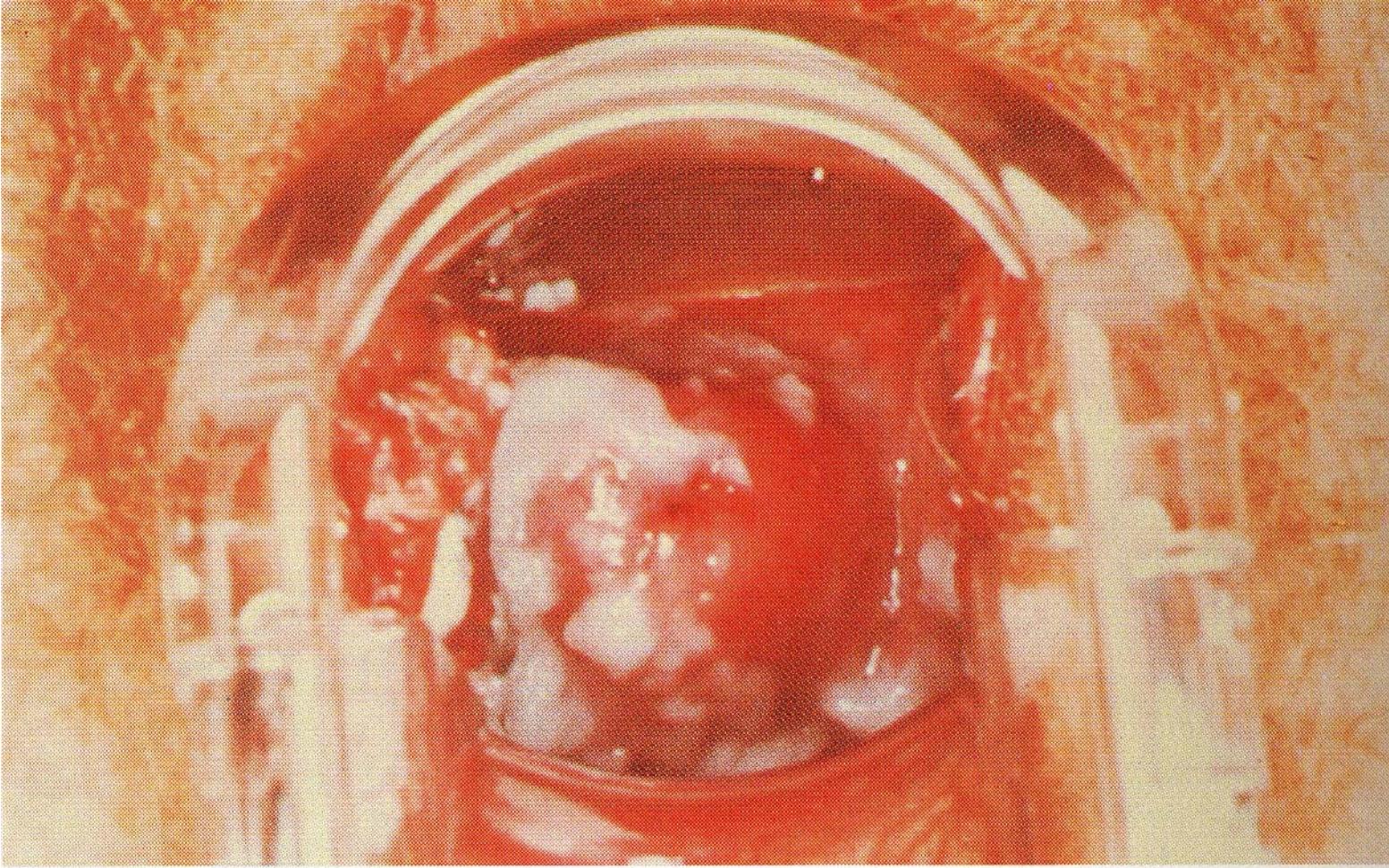
**Fig. 78** Gonococcal urethritis. Typical purulent meatal discharge with inflammation of the glans. Symptomatic gonorrhoea in males is characterized by a spontaneous purulent discharge and dysuria. Courtesy of Dr J. Clay.

# Gonorrhea – akut Cervicitis



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# Gonorrhea – akut Cervicitis



**Fig. 79** Gonococcal endocervicitis. View through vaginal speculum showing reddened external os through which mucopurulent secretion is exuding. The most common manifestation of gonorrhoea in females is cervicitis, which is often asymptomatic. Courtesy of Dr S. E. Thompson.

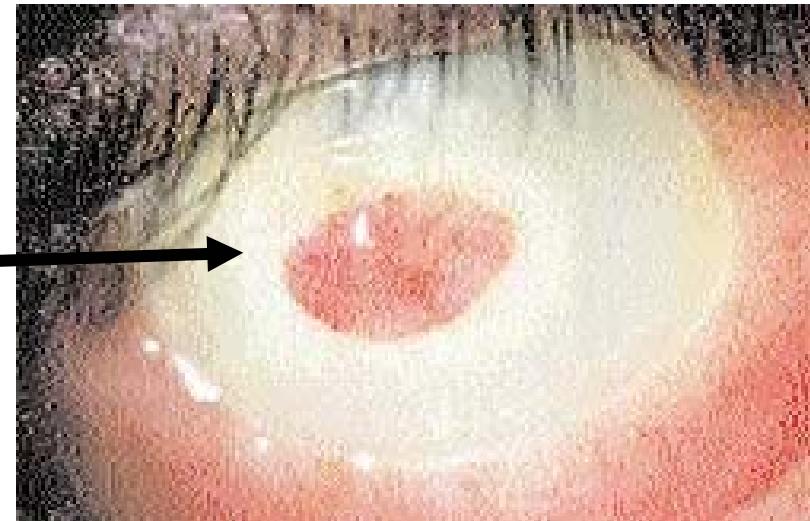
# Gonorrhea – akut Conjunktivitis Blenorrhea neonatorum



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[www.mc3.edu](http://www.mc3.edu)

**Corneal ulcers due to gonococcus are very destructive and have a tendency to perforate the cornea.**



[www.slackbooks.com](http://www.slackbooks.com)

# Gonorrhoe – Krónikus és disseminált forma

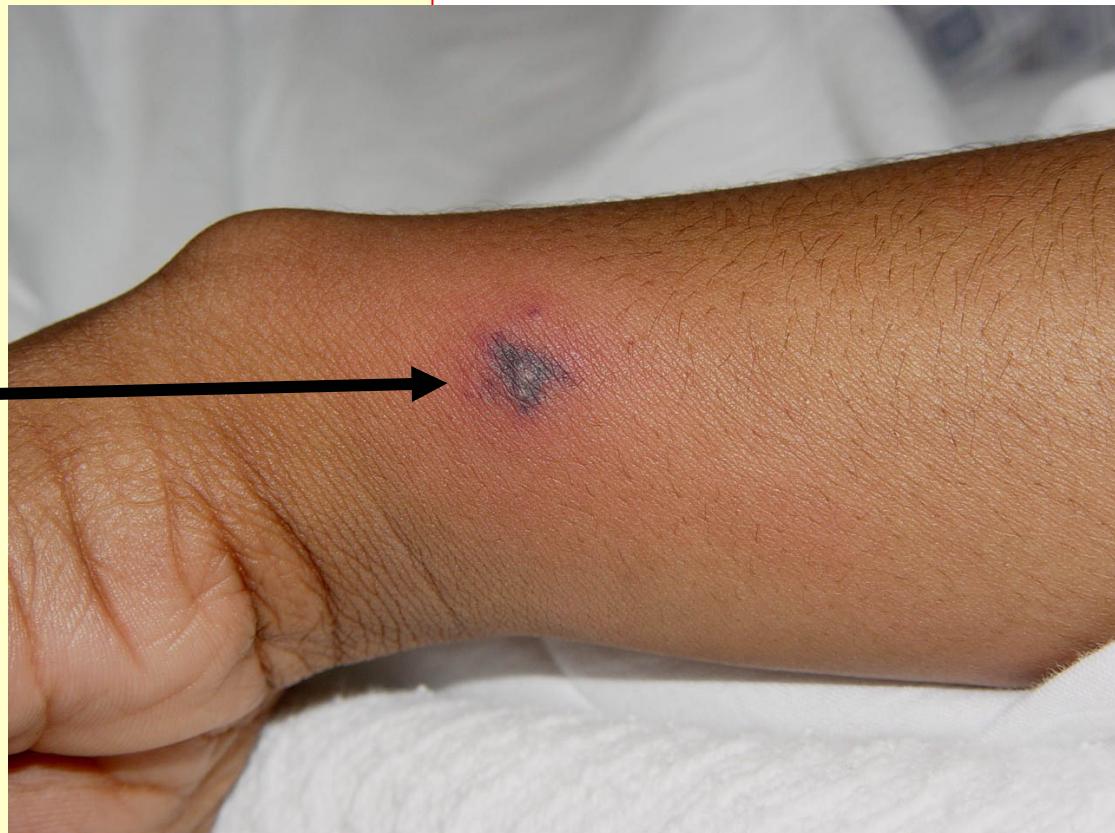
Endometritis,

Salpingitis,

Prostatitis

**purulent Arthritis,**

Vasculitis



**FONTOS!**

anorectalis Go és Pharyngitis

(„alternatív Genitáliák”)



Fig. 8.33 Gonococcal septic arthritis. Arthritis due to *N. gonorrhoeae* in a 24-year-old woman, showing marked erythema and swelling of the right ankle and leg. By courtesy of Dr. T.F. Sellers Jr.

Fig. 8.33 Gonococcal arthritis. Dactylitis secondary to gonococcal bacteraemia. By courtesy of Dr. S.E. Thompson



# Gonorrhea – Diagnosis – csak akut esetben!

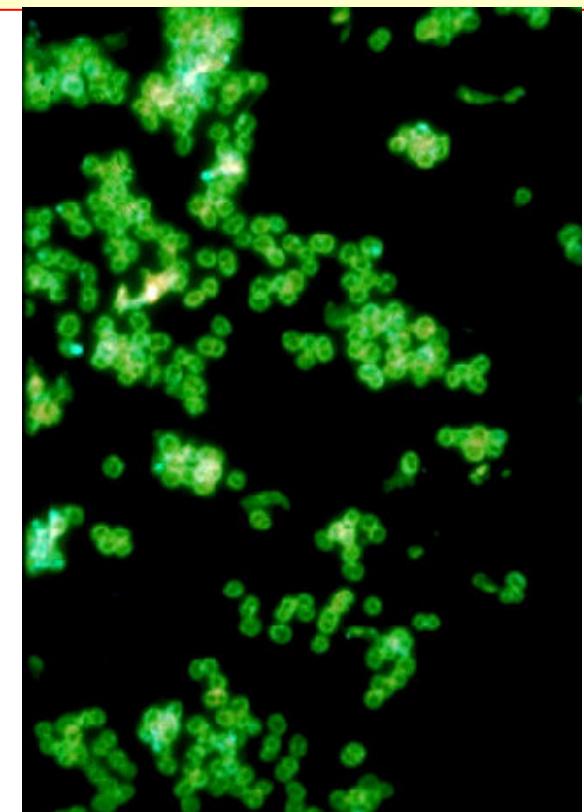
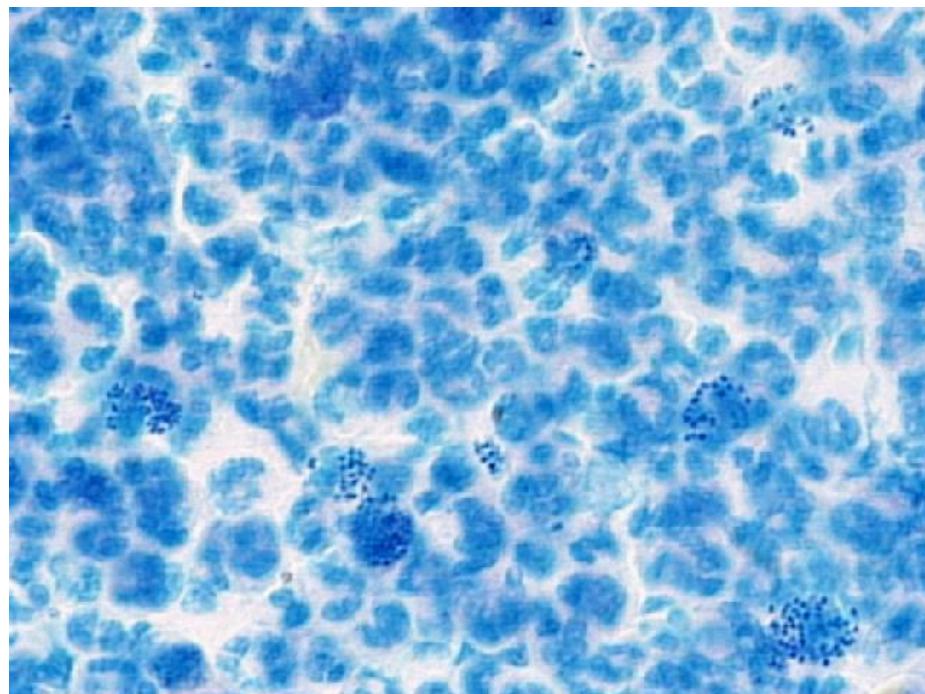
**Mikroszkópos vizsgálat**

Kórokozó kimutatása

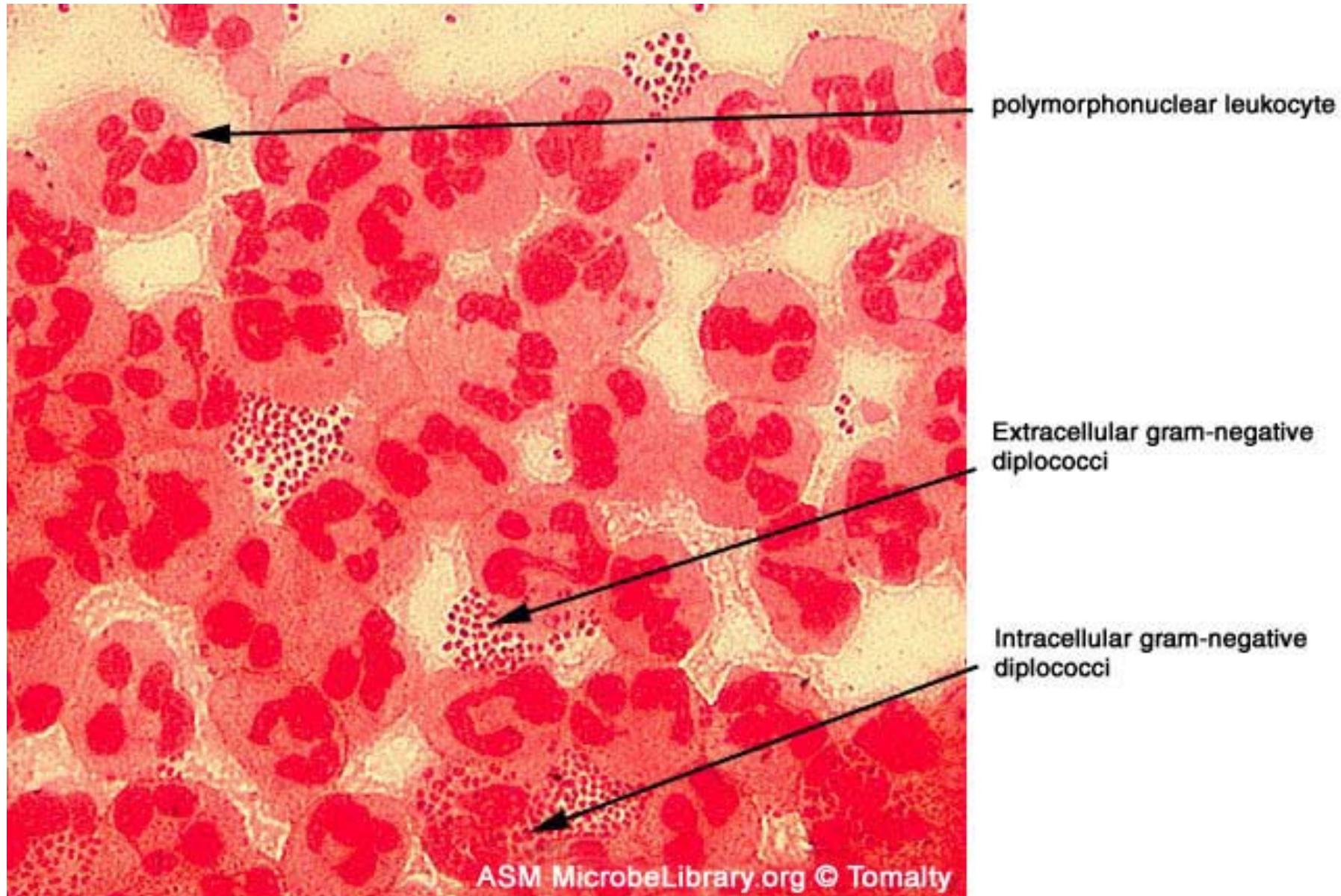
**Gram festés**

Methylenkék festés, Direkt Immunofluoreszcens (DIF)

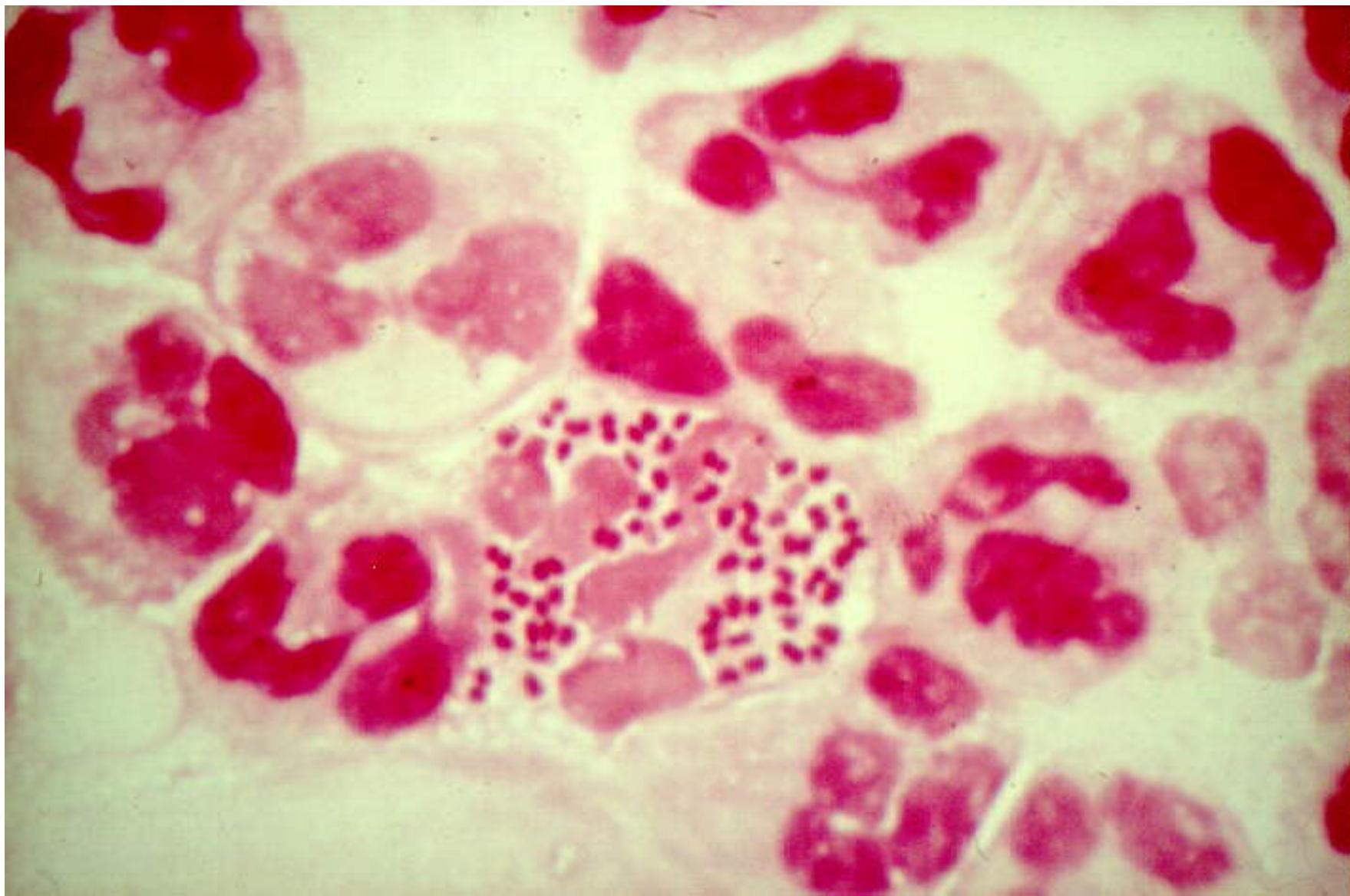
[www2.mf.uni-lj.si](http://www2.mf.uni-lj.si),  
[www.uni-ulm.de](http://www.uni-ulm.de),  
pathmicro.med.sc.edu



# GO – Gram festés – előzetes eredmény!



# GO – Gram festés – előzetes eredmény!



# Gonorrhea – Diagnosis

Tenyésztés:

„bedside” Thayer-Martin táptalaj  
és Csokoládéagar, 5% CO<sub>2</sub>

Identifikálás: ox+, glu+, mal-

Antigénkimutatás:  
Latex-agglutinatio



## Therápia:

3. Generációs Cephalosporin (Ceftriaxone)  
vagy Spectinomycin (Aminoglycoside)



## Prophylaxis:

### GO

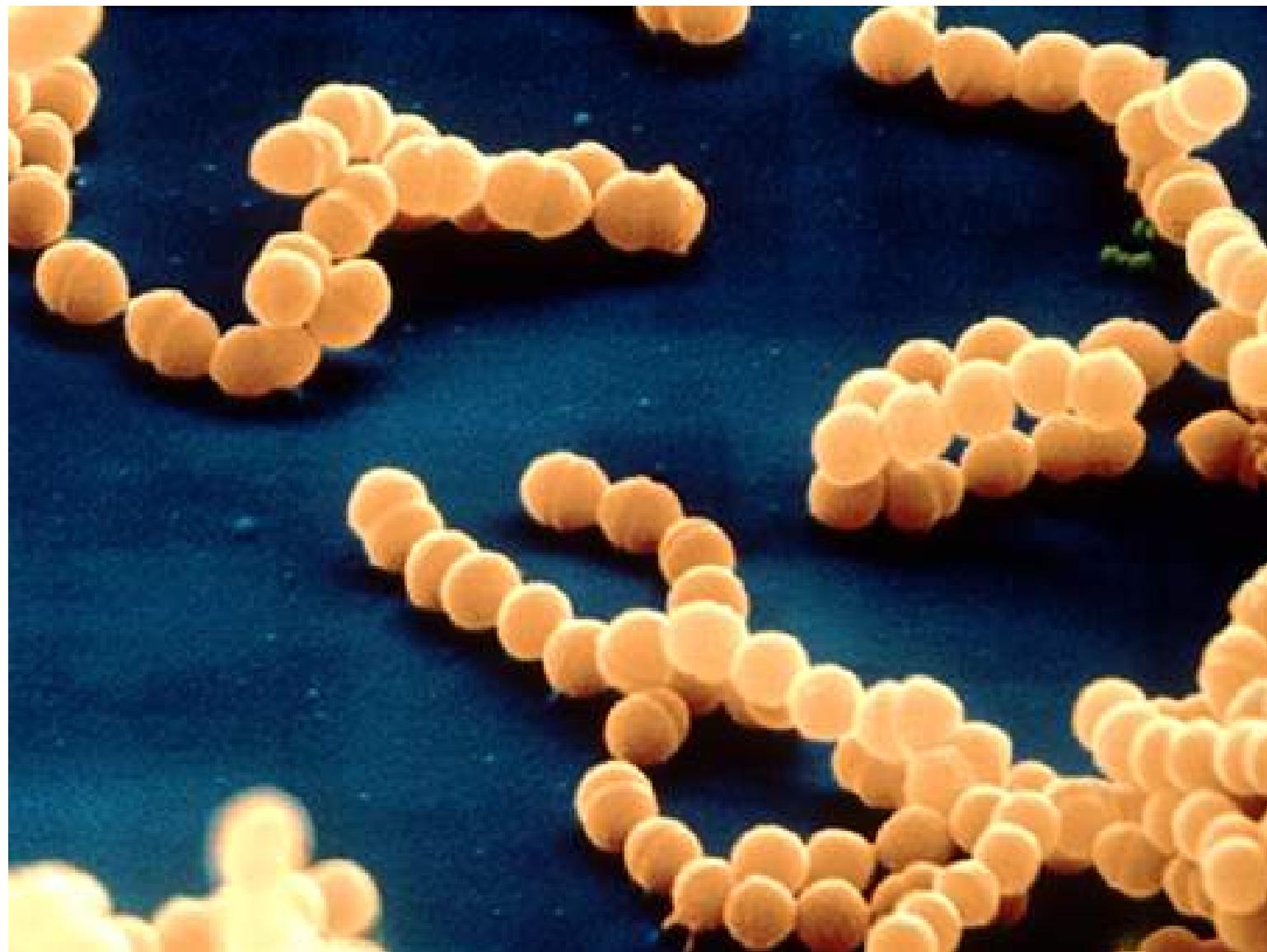
- Nem megkapni! (safe sex)
- Lehetséges fertőző források, kontaktok felderítése
- Korai Diagnosis és kezelés

## Ophthalmia neonatorum:

Credé szemcsepp 1% ezüst-acetát (-nitrát)

**Védőoltás NINCS! (Antigenvariációk!)**

# **N. meningitidis = Meningococcus**



scanning EM

# **N. meningitidis = Meningococcus**

**Antigének és Virulenciafaktorok:**

**TOK** – Polysaccharide

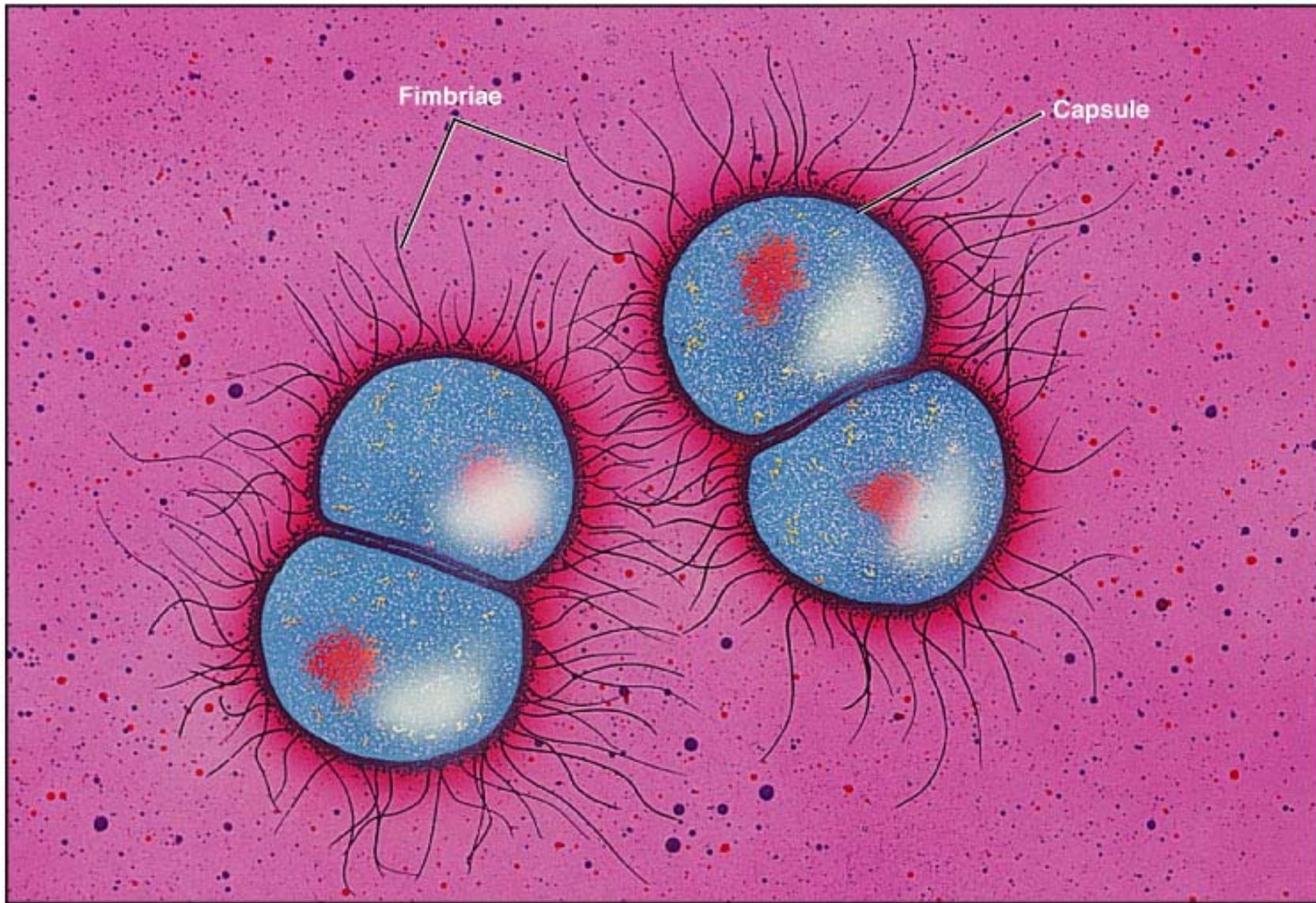
12 Serotypus (A, B, C, W135, Y!)

**Pili/Fimbriae**

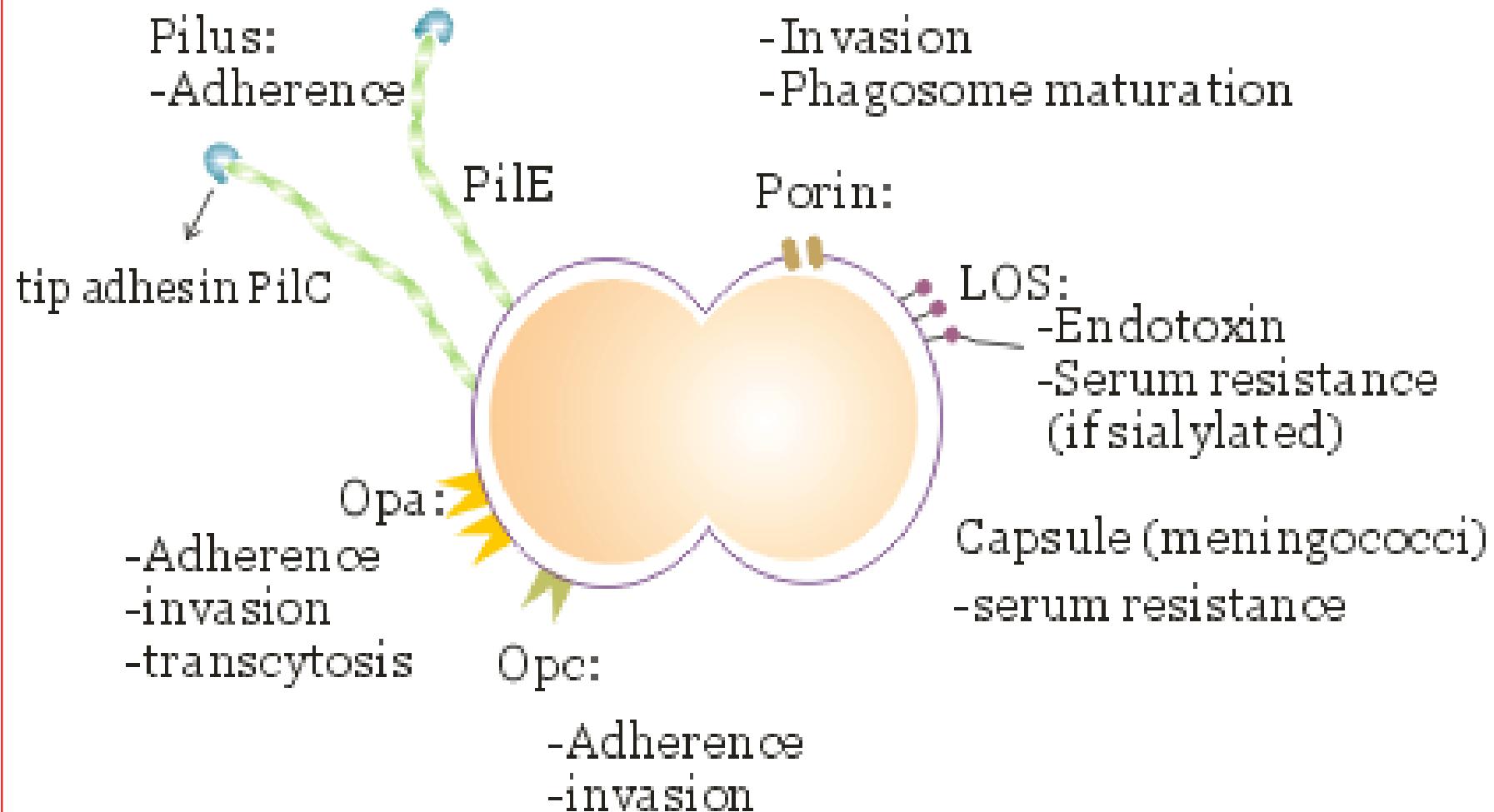
**IgA-Protease!**

**Outer Membrane Protein-ek (OMP)**

**LOS** (Mimikri, szializálva Serum-rezisztens!)



# Meningococcus



# **N. meningitidis = Meningococcus**

## **Fertőzés forrása**

emberek – hordozók (betegek, egészségesek)

## **Átvitel, behatolás**

- Direkt, cseppfertőzéssel
- Orrüreg, Torok

## **Kórképek**

Pharyngitis

**Meningitis cerebrospinalis epidemica**

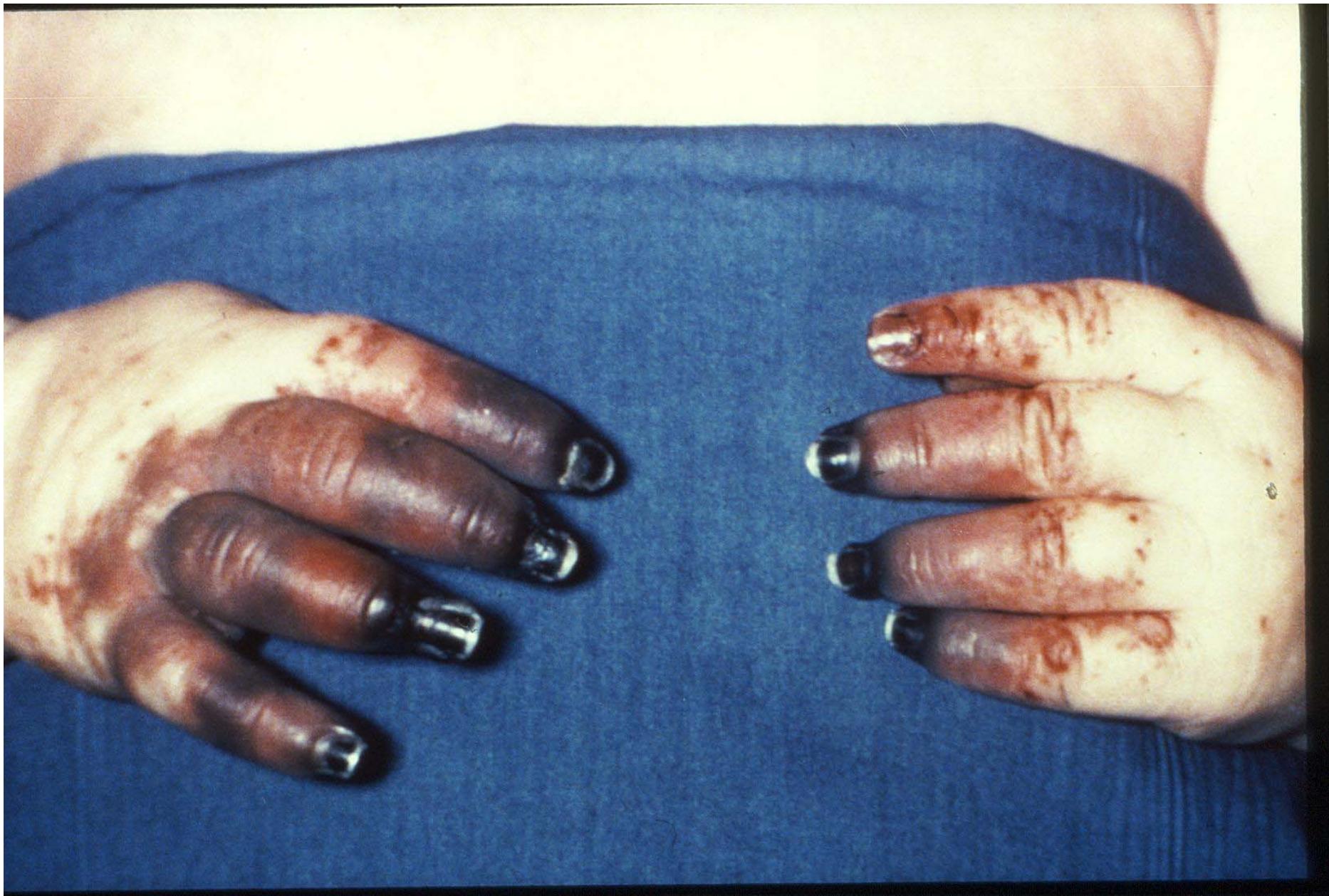
Sepsis = **Waterhouse-Friderichsen Syndroma**



**Fig. 10.56 Acute meningococcaemia. Note the variable size of the lesions and their peripheral distribution. Some of the lesions are obviously purpuric, others macular or papular.**



**Fig. 10.60 Acute meningococcaemia. Petechia on bulbar conjunctiva.**



**Fig. 10.62 Acute meningococcaemia. Gangrene of the extremities following a near-fatal illness with hypotension.**



**Fig. 10.63 Acute meningococcaemia. Gangrene of both legs in a black man with acute meningococcal infection. Bilateral below knee amputations were later required.**



The characteristic skin rash of meningococcal septicaemia, caused by *Neisseria meningitidis*. (Courtesy of Wellcome Trust Photographic Library)  
[srs.dl.ac.uk](http://srs.dl.ac.uk)

# Waterhouse – Friderichsen Syndroma - diagnosis

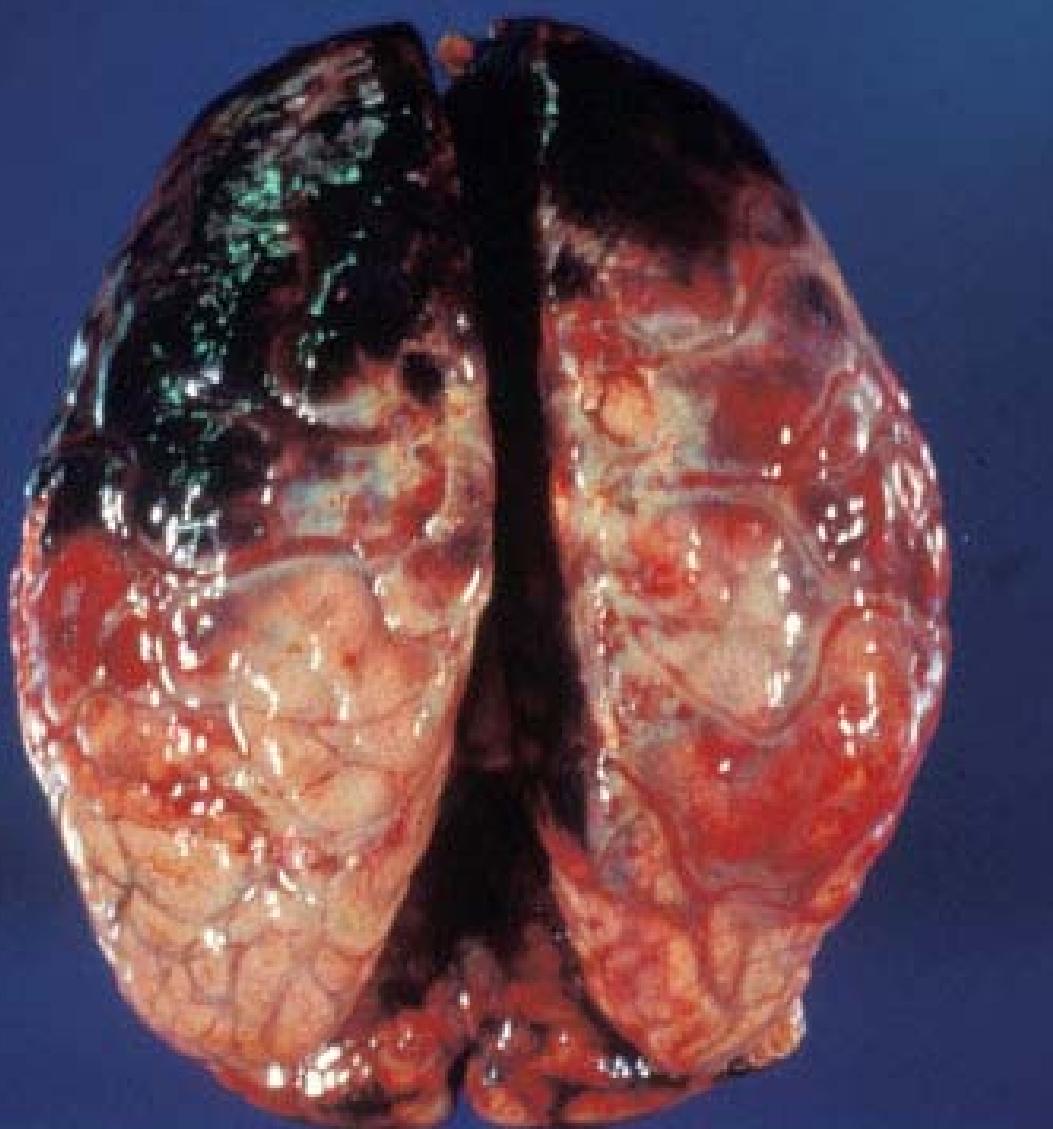




**Waterhouse- Friderichsen Syndrome:** schwere nekrotisierende Hautläsionen bei Meningokokkensepsis mit Verbrauchskoagulopathie  
(R. E. Rieger, Univ.-Kinderklinik Marburg).



**The patient with Waterhouse-Friderichsen syndrome has sepsis with DIC and marked purpura.**



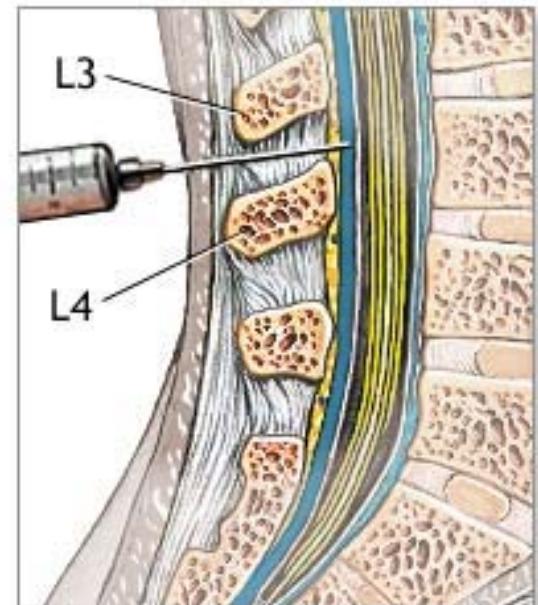
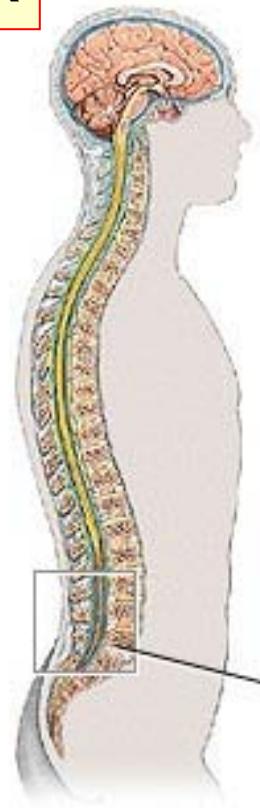
Purulent meningitis with hemorrhage in the frontal lobe (gross findings).



**Acute hemorrhage in bilateral adrenals caused acute adrenal insufficiency (Waterhouse-Friderichsen syndrome).**

# Meningitis Diagnosis

Vizsgálati anyagok:  
**Liquor!** - Lumbalpunctio  
**Vér**  
Hordozóktól: Torokváladék



Columna  
vertebral

# Meningitis Diagnosis

Kórokozó kimutatás

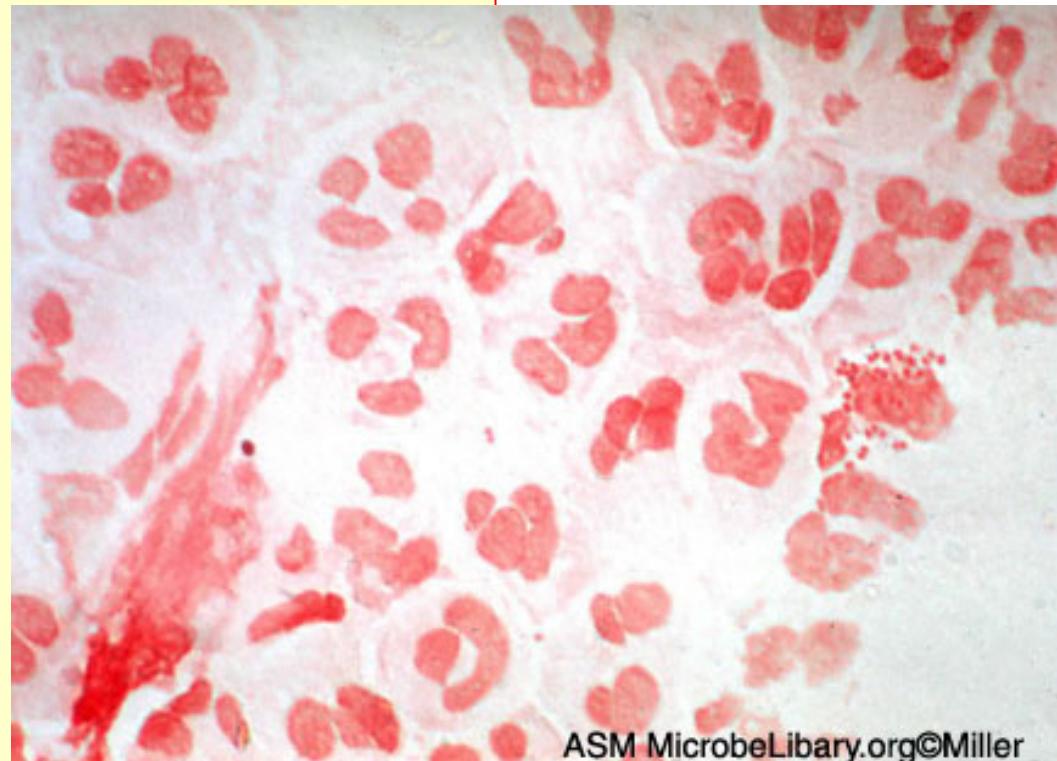
**Mikroszkópos Vizsgálat**  
(Liquor, Haemokultura)

Tenyésztés

Liquor, Vér, Torok

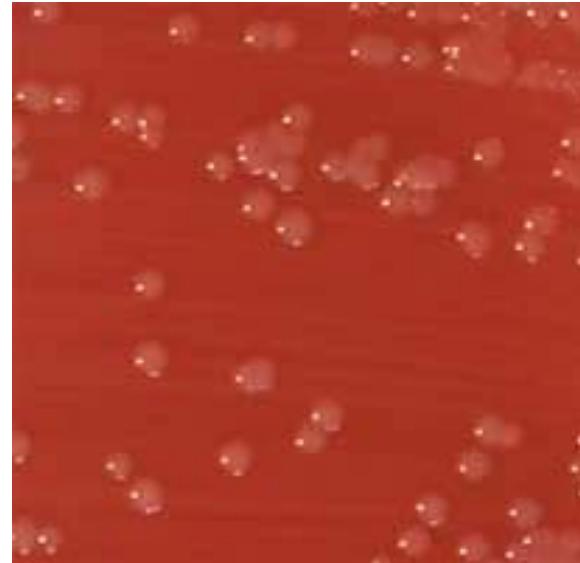
Direkt Ag Kimutatás

(Liquor) – Latex-agglutinatio

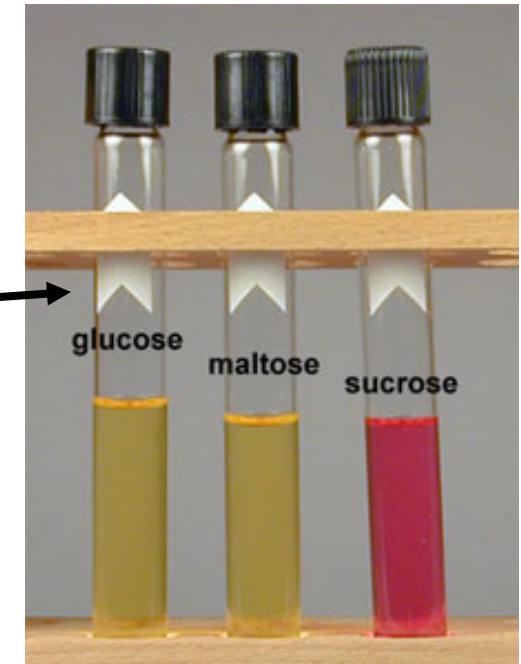


# Diagnosis *N. meningitidis*

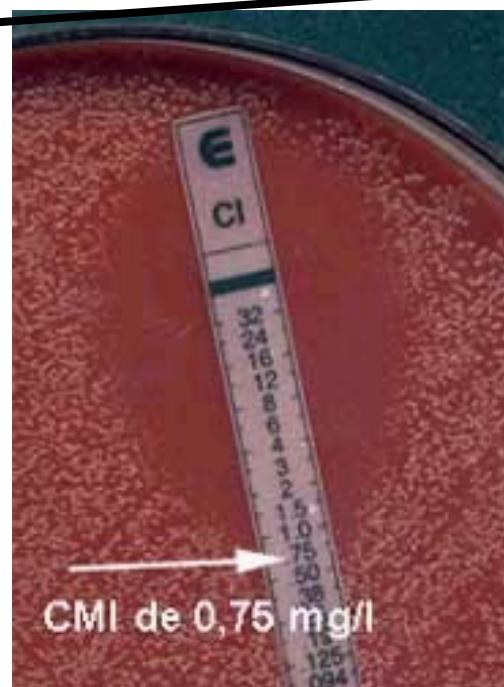
Tenyésztés:  
Véresagar,  
Csokoládéagar



Identifikálás:  
glu+, mal+



MIC (E-test)



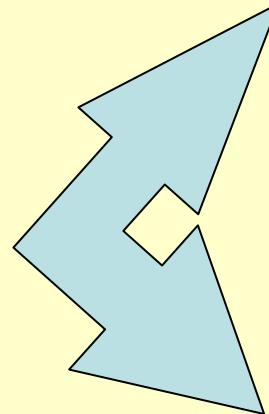
# Meningococcus meningitis

## Therapia:

Penicillinek és/vagy

Cephalosporin (III. Gen.)

Nincs Beta-lactamase termelés



## Prophylaxis:

Aktív Immunizálás

Védőoltás:

- Rizikócsoportok
  - Utazás!
- (Meningitis övezet!)

## Chemoprophylaxis:

Rifampicin (Kontaktok)





# Meningitis övezet

# **Neisseria meningitidis - B**

**Európa!**

**NINCS Védőoltás!**

**Rifampicin**



# ***Neisseria, Haemophilus, Bordetella***

**2. *Haemophilus***

# Kicsi, Gram negatív pálcák/coccobacillusok

Genus

**Haemophilus**

Faj

**H. influenzae P**

**H. parainfluenzae**

**H. aegyptius P**

**H. ducreyi P**

Bordetella

**B. pertussis P**

**B. parapertussis**

P: Pathogen

# *Haemophilus influenzae*

## Morphologia:

Gram - Coccobacillus,  
ca. 1 µm

## Tenyésztés:

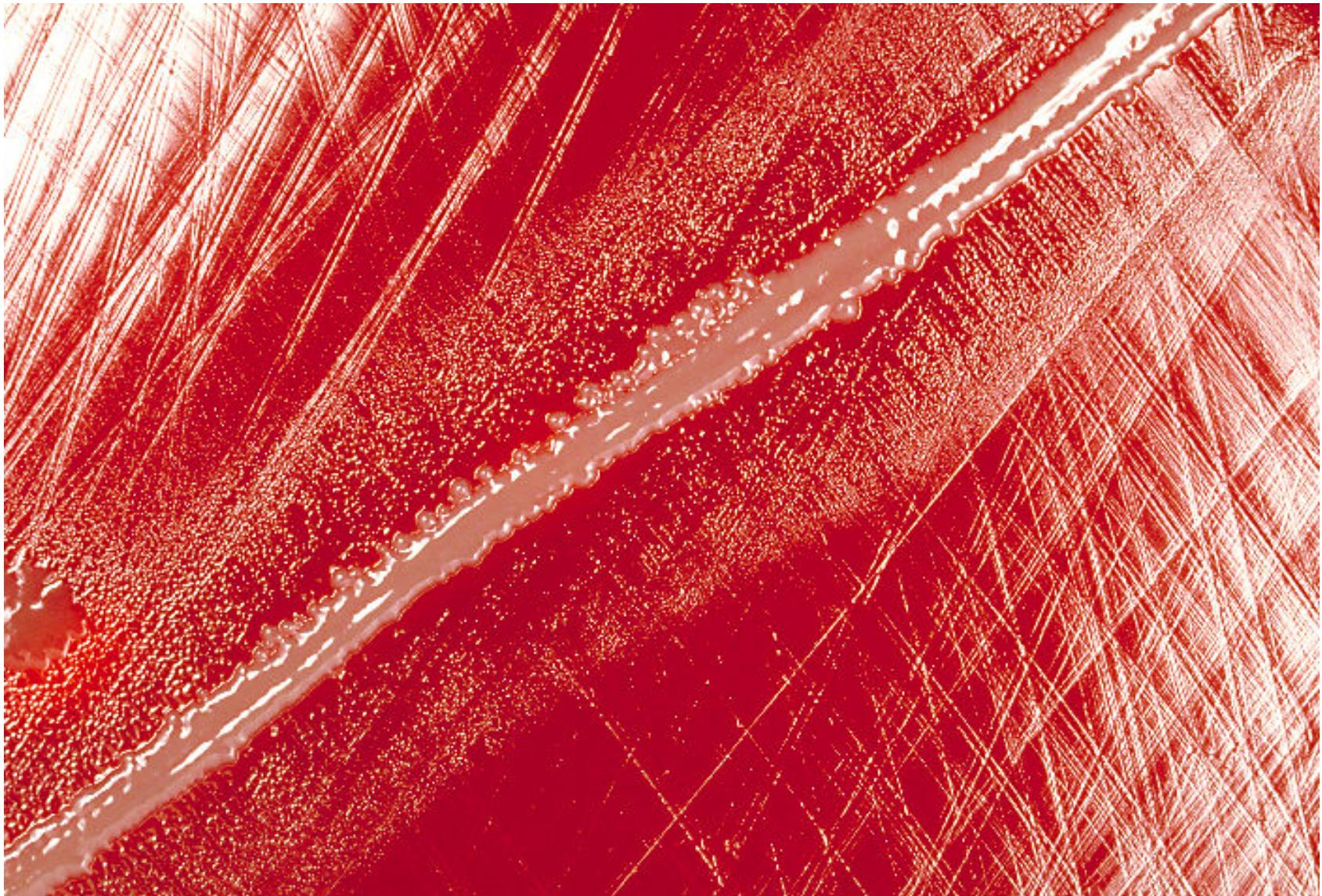
Növekedési faktorok !

(Csokoládé agar,  
X= Haem, V= NAD,

Szatellitizmus,

„Dajkajelenség”)





# *Haemophilus influenzae*

Antigének és Virulenciafaktorok:

TOK – Polysaccharid

Typen: a, b, c, d, e, f (HiB!)

IgA-Protease!

Felszínen:

Outer Membrane Proteine (OMP)

LPS



*Haemophilus  
influenzae Typ b  
(Hib)*

[www.soundmedicine.iu.edu](http://www.soundmedicine.iu.edu)

# *Haemophilus influenzae*

Kórképek:

Meningitis, Sepsis

Cellulitis

Felső légutak:

Epiglottitis!, Nasopharyngitis, Sinusitis, Otitis media

Alsó légutak:

Bronchitis, Pneumonia

# *Haemophilus influenzae*



## Sepsis

An infant with severe vasculitis with disseminated intravascular coagulation (DIC) with gangrene of the hand secondary to *Haemophilus influenzae* type b septicemia - prior to the availability of the Hib vaccine.

-Image provided by: Visual Red Book on CD-ROM-

[www.ecbt.org](http://www.ecbt.org)

-(2000 Red Book: 25th Edition, Report of the Committee on Infectious Diseases)

# *Haemophilus influenzae*



**Periorbital cellulitis.**

© Neal Halsy, MD [www.cispimmunize.org](http://www.cispimmunize.org)

# *Haemophilus influenzae*

Kórképek:

**Meningitis, Sepsis**

Cellulitis

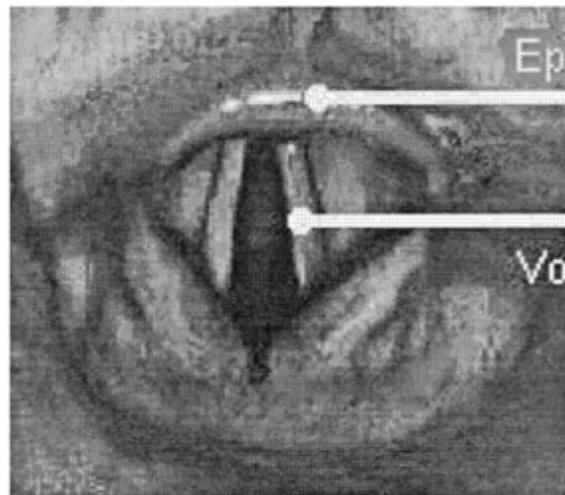
Felső légutak:

**Epiglottitis!, Nasopharyngitis, Sinusitis, Otitis media**

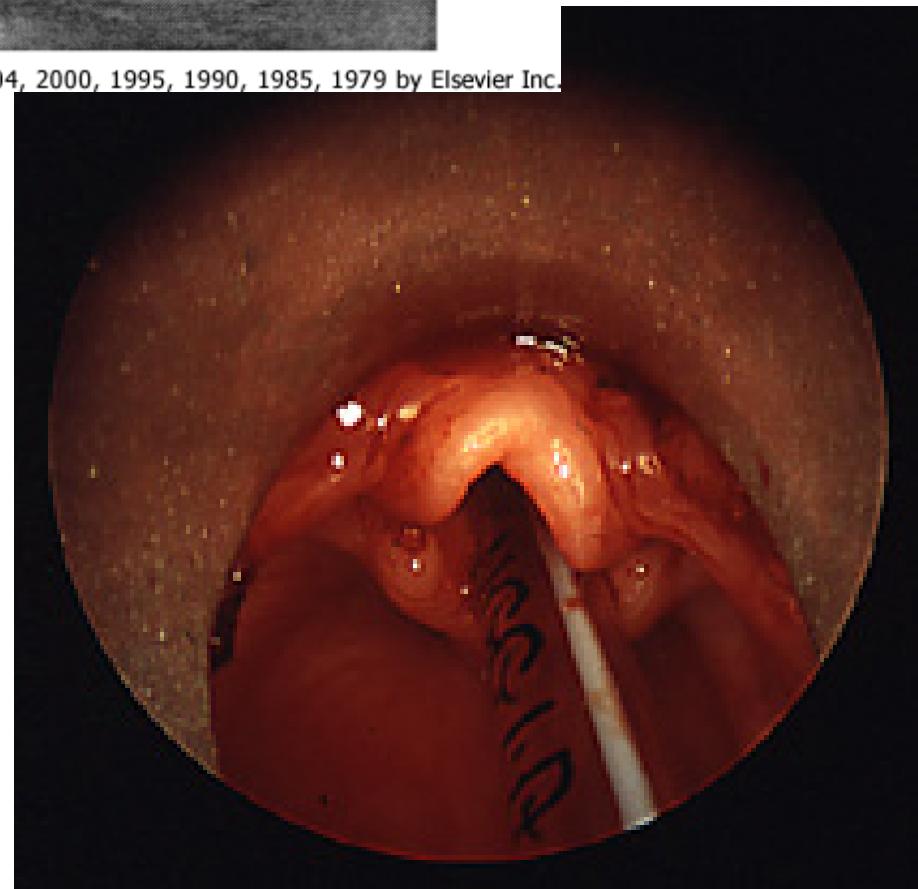
Alsó légutak:

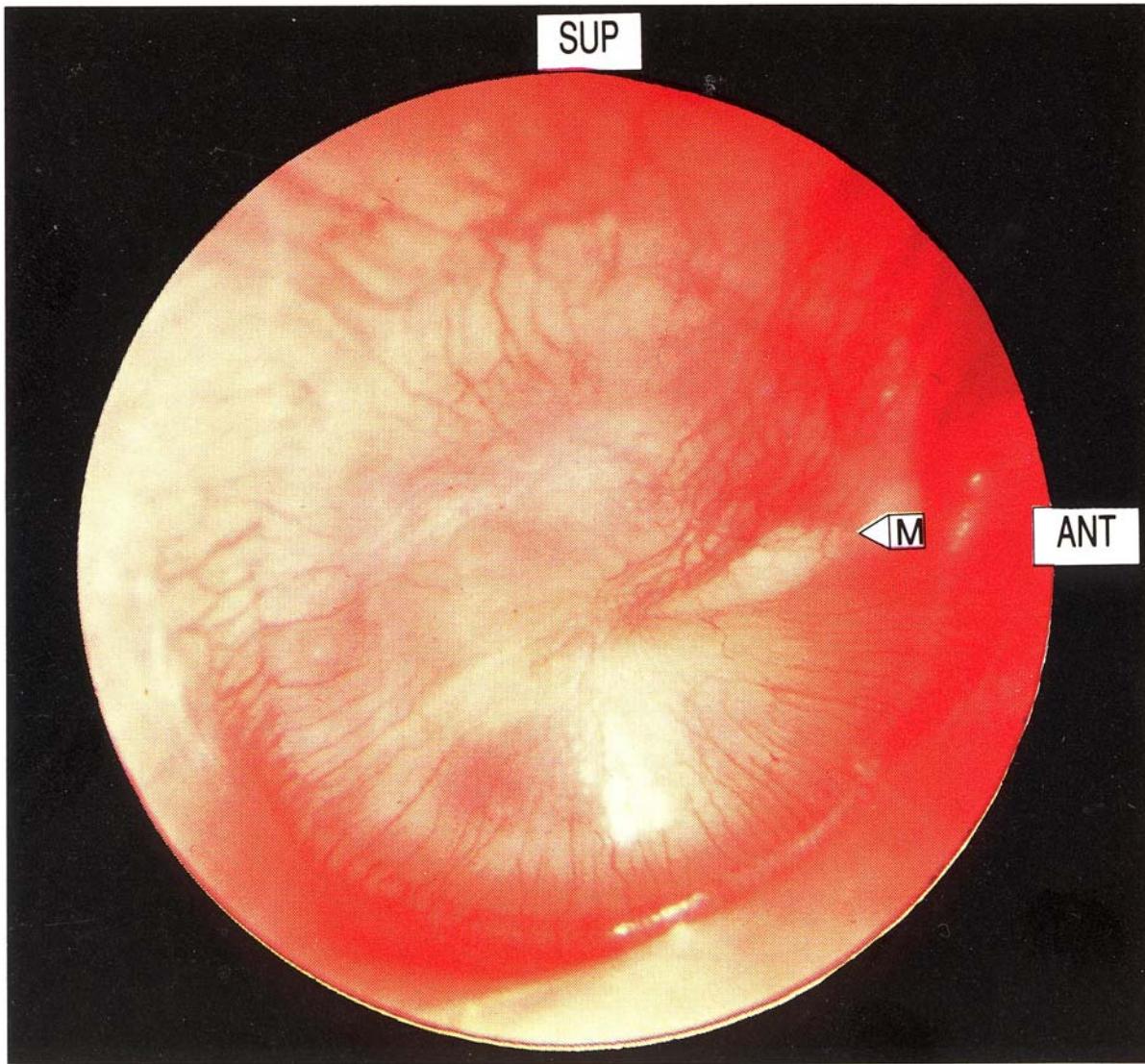
Bronchitis, Pneumonia

# HIB-epiglottitis

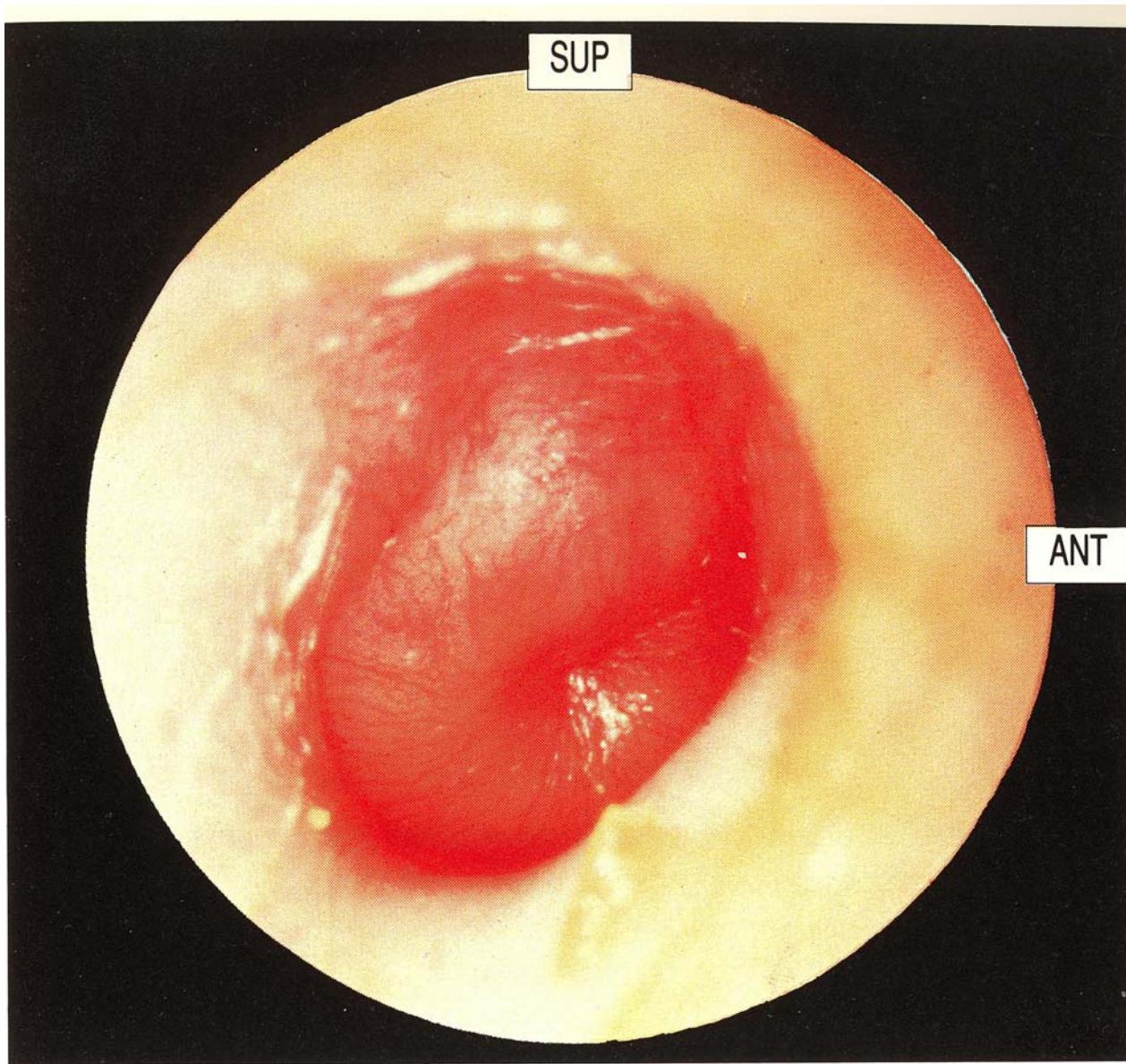


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**Fig. 10** Acute otitis media. Early stage showing mild injection of the drum, especially in the region of the malleus (M). The most common bacterial causes are *Streptococcus pneumoniae* and *Haemophilus influenzae*, with a smaller proportion of cases caused by *Streptococcus pyogenes* or *Branhamella catarrhalis*. Courtesy of Dr M. Chaput de Saintonge.



**Fig. 11** Acute otitis media. Advanced stage showing bulging of the drum on both sides of the malleus, which is obscured. These features are seen just before the drum perforates. By courtesy of Dr M. Chaput de Saintonge.

# *Haemophilus influenzae*

## Diagnosis:

Vizsgálati anyagok

- LIQUOR! (CSF)
- Minták a fertőzés helyéről (orr, torok, köpet stb.)

## Kórokozó kimutatás:

Mikroszkópos vizsgálat, Tenyésztés,  
Tok Ag kimutatás (Latex-agglutinatio)

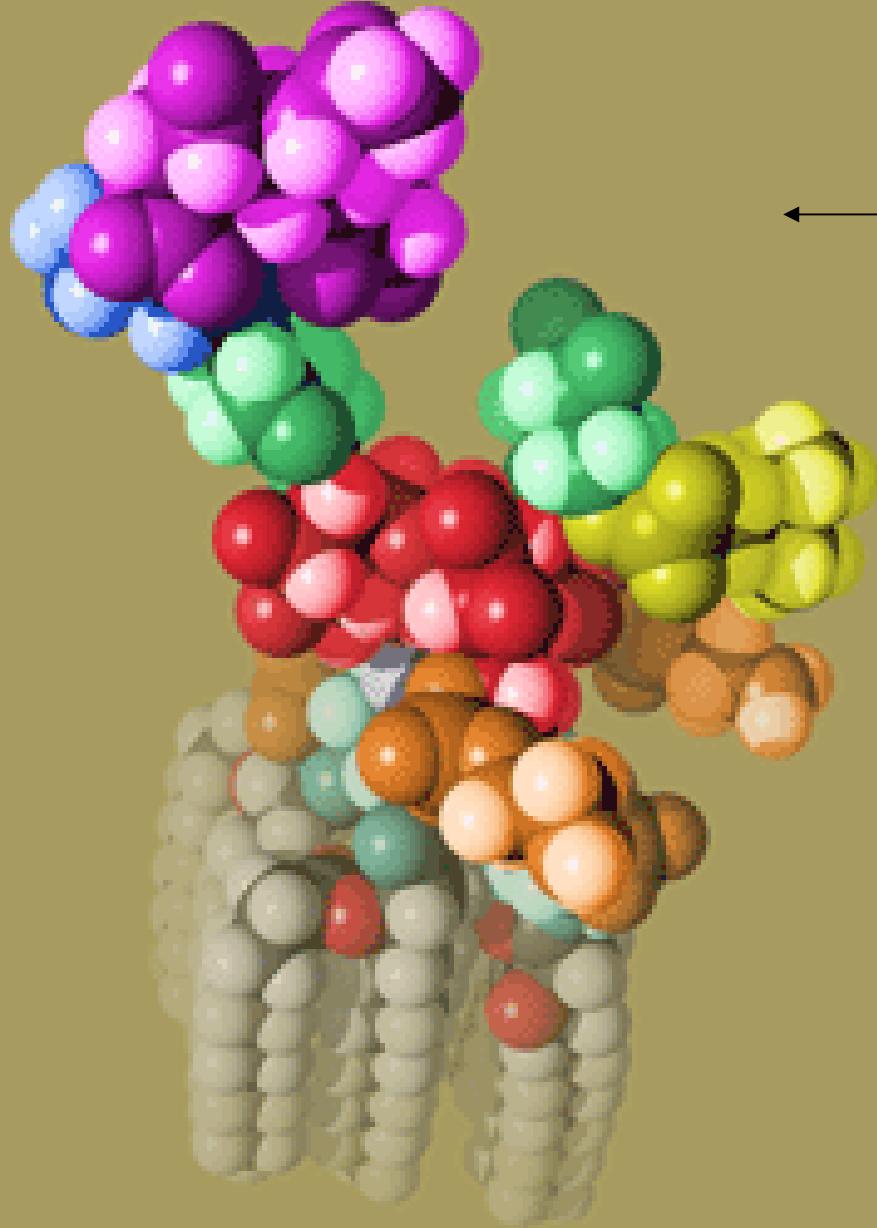
## Therapia:

1. Ampicillin + III. gen. Cephalosporin
2. Ampicillin + Aminoglycoside

## Prophylaxis:

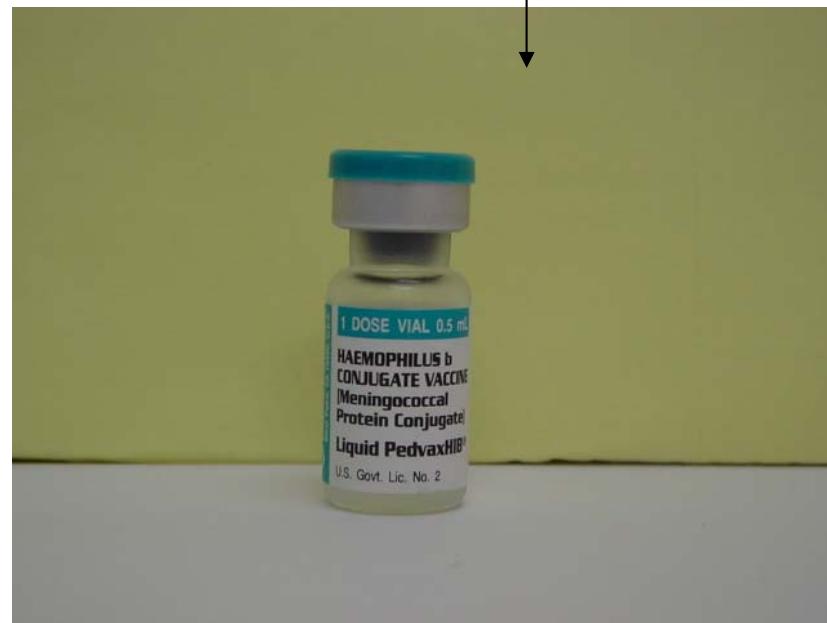
Aktív Immunizálás - **HiB Konjugált-Védőoltás**

(Polysaccharid + Protein)



## Lipopolsaccharid Extrakt - Vakcina

[ibs-isb.nrc-cnrc.gc.ca](http://ibs-isb.nrc-cnrc.gc.ca)



# *Haemophilus ducreyi*

Kórkép:

**Ulcus molle = Chancroid = lágy Schanker/fekély**

# *Haemophilus aegyptius*

Kórkép: Brazíliai purpurás láz

# *Haemophilus parainfluenzae*

Pharyngitis, Endocarditis, Conjunctivitis

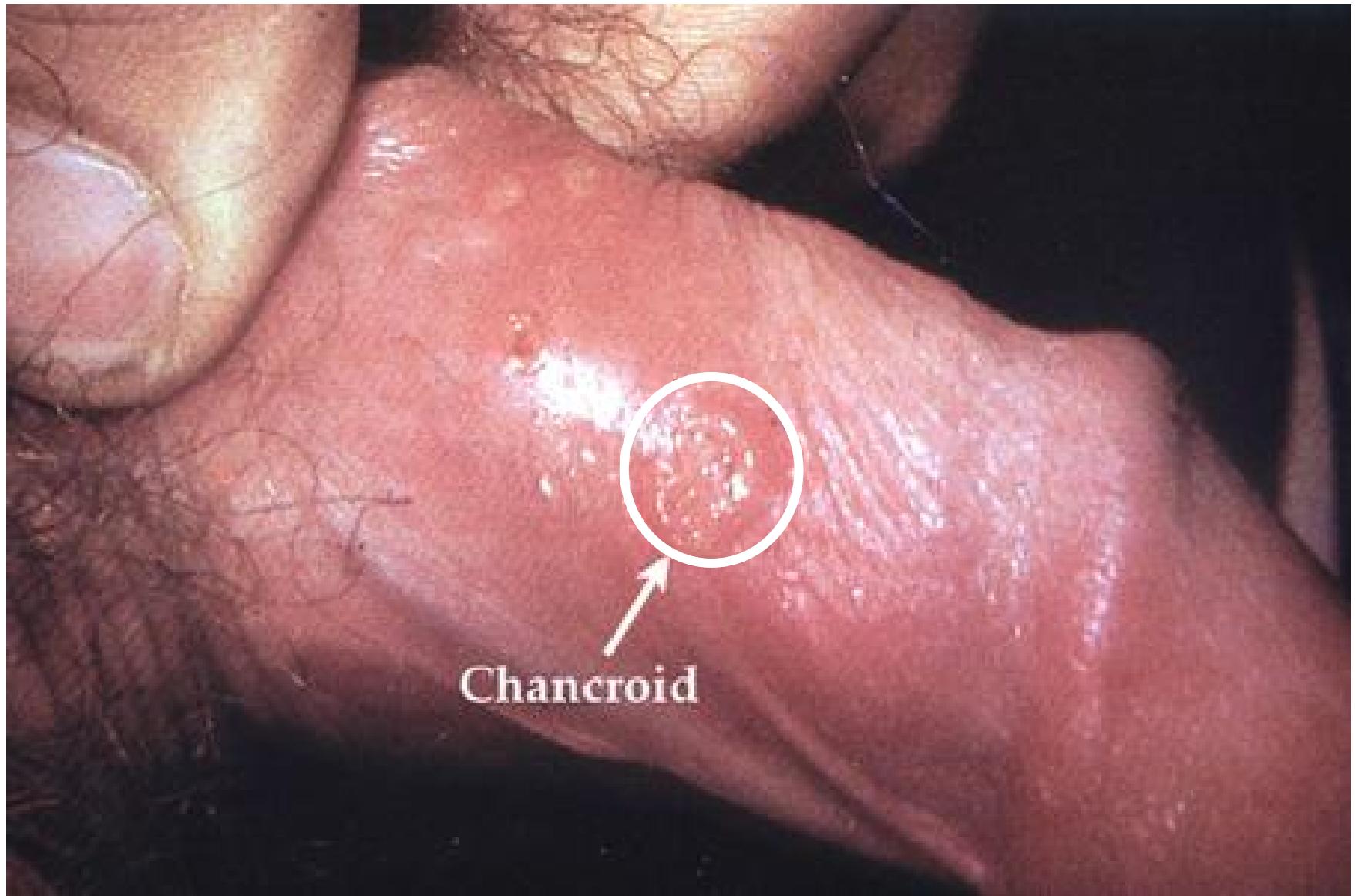
## Ulcus molle



# Ulcus molle



<http://dermis.net>



Chancroid



**Chancroid in female**

# **Neisseria, Haemophilus, Bordetella**

**3. Bordetella**

# Kicsi, Gram negatív pálcák/coccobacillusok

Genus

**Haemophilus**

Faj

**H. influenzae P**

**H. parainfluenzae**

**H. aegyptius P**

**H. ducreyi P**

**Bordetella**

**B. pertussis P**

**B. parapertussis**

P: Pathogen

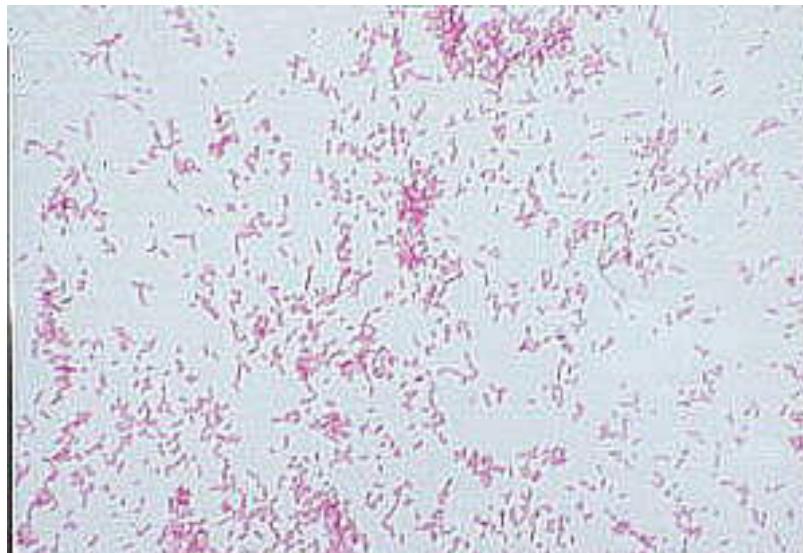
# *Bordetella pertussis*

## Morphologia:

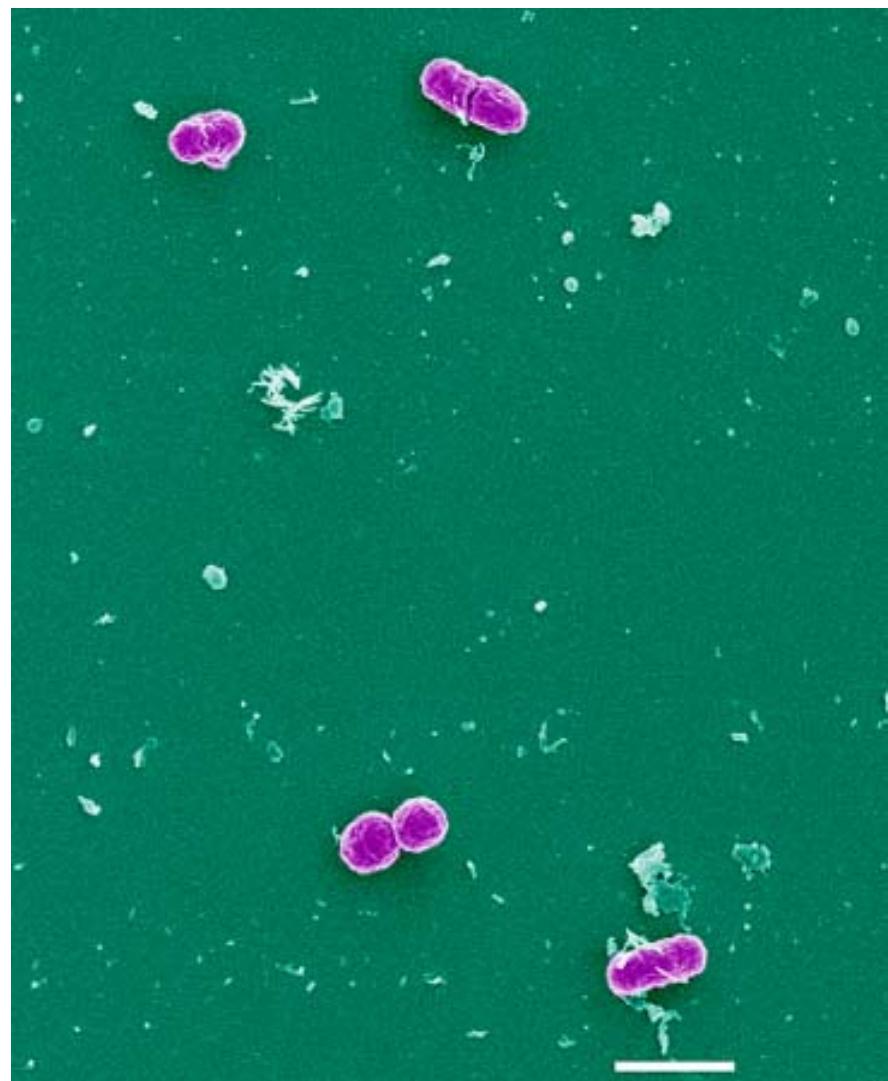
Gram negativ

Coccobacillus,

ca. 1  $\mu\text{m}$



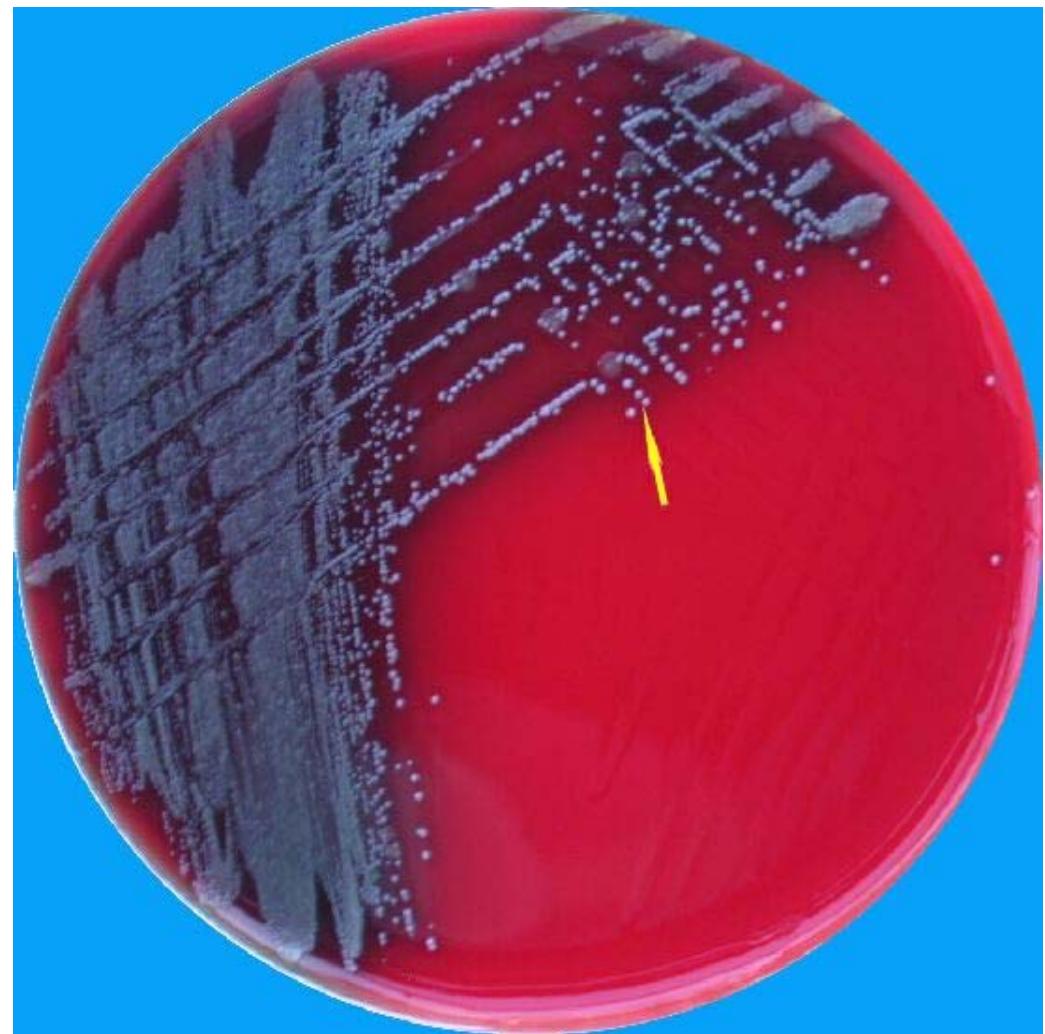
Gram stain of *B. pertussis*



# *Bordetella pertussis*

Tenyésztés:

Speciális  
Bordet – Gengou



# *Bordetella pertussis*

## Antigének és Virulenciafaktorok:

Tok

Fimbriák, filamentózus Haemagglutinin  
Outer Membrane Proteine (OMP)

LPS

Pertactin

## Extracelluláris Toxinok:

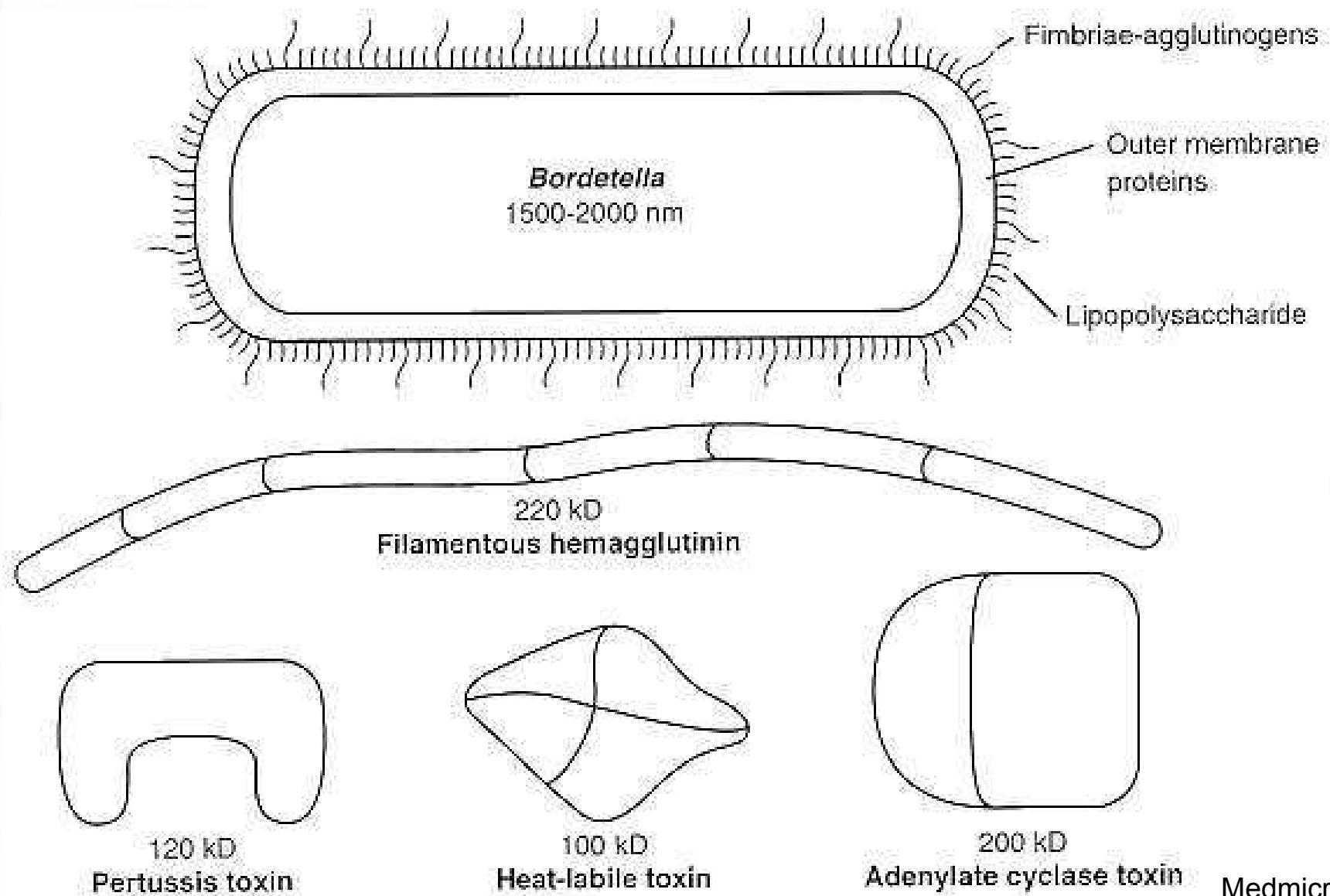
Pertussis Toxin (AB toxin)

Adenylatcyclase Toxin

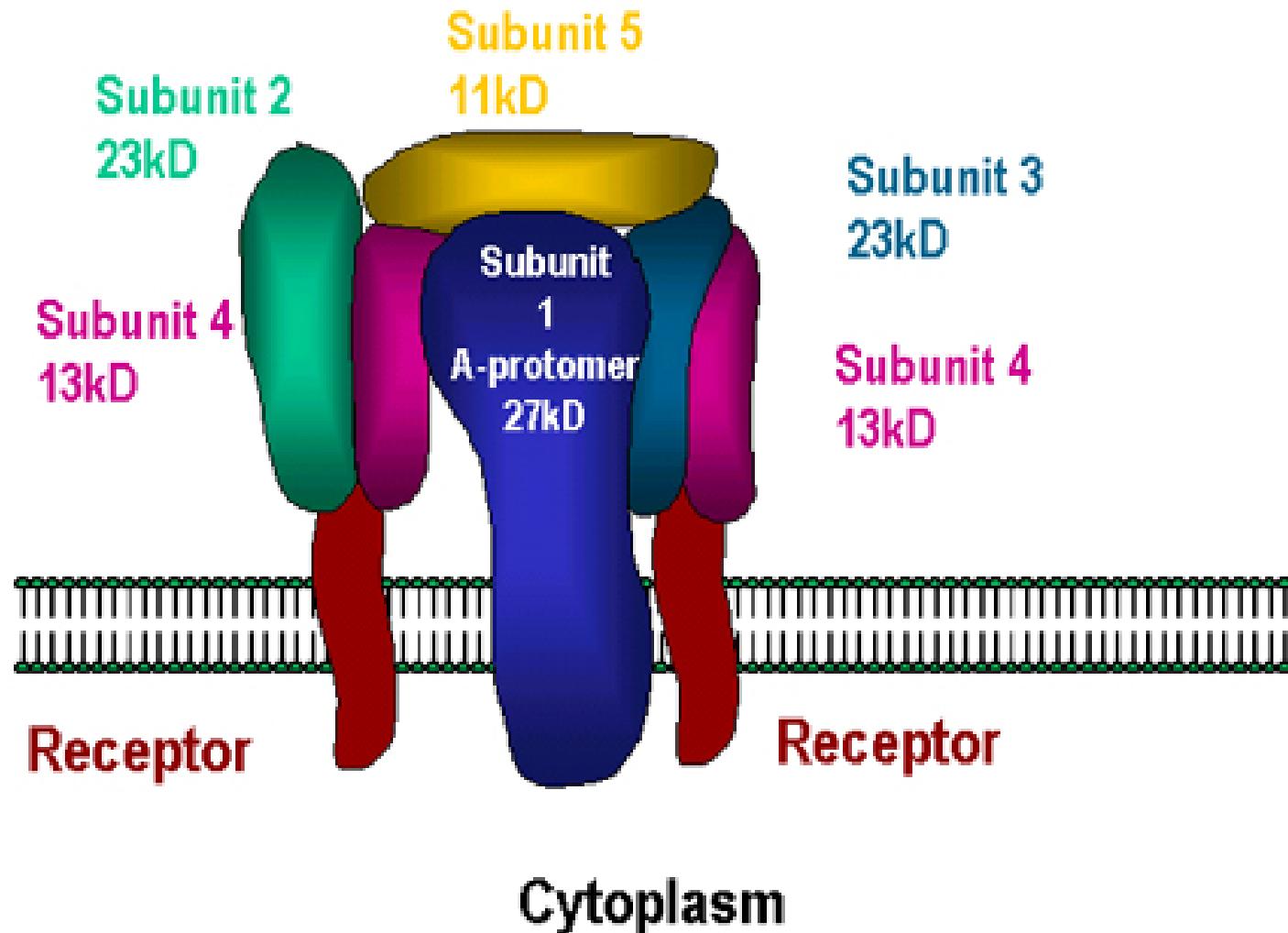
Trachealis cytotoxin

Dermatonecrotikus Toxin

## FIGURE 31-2 Virulence factors of *B pertussis*.



# Pertussis toxin



# *Bordetella pertussis*

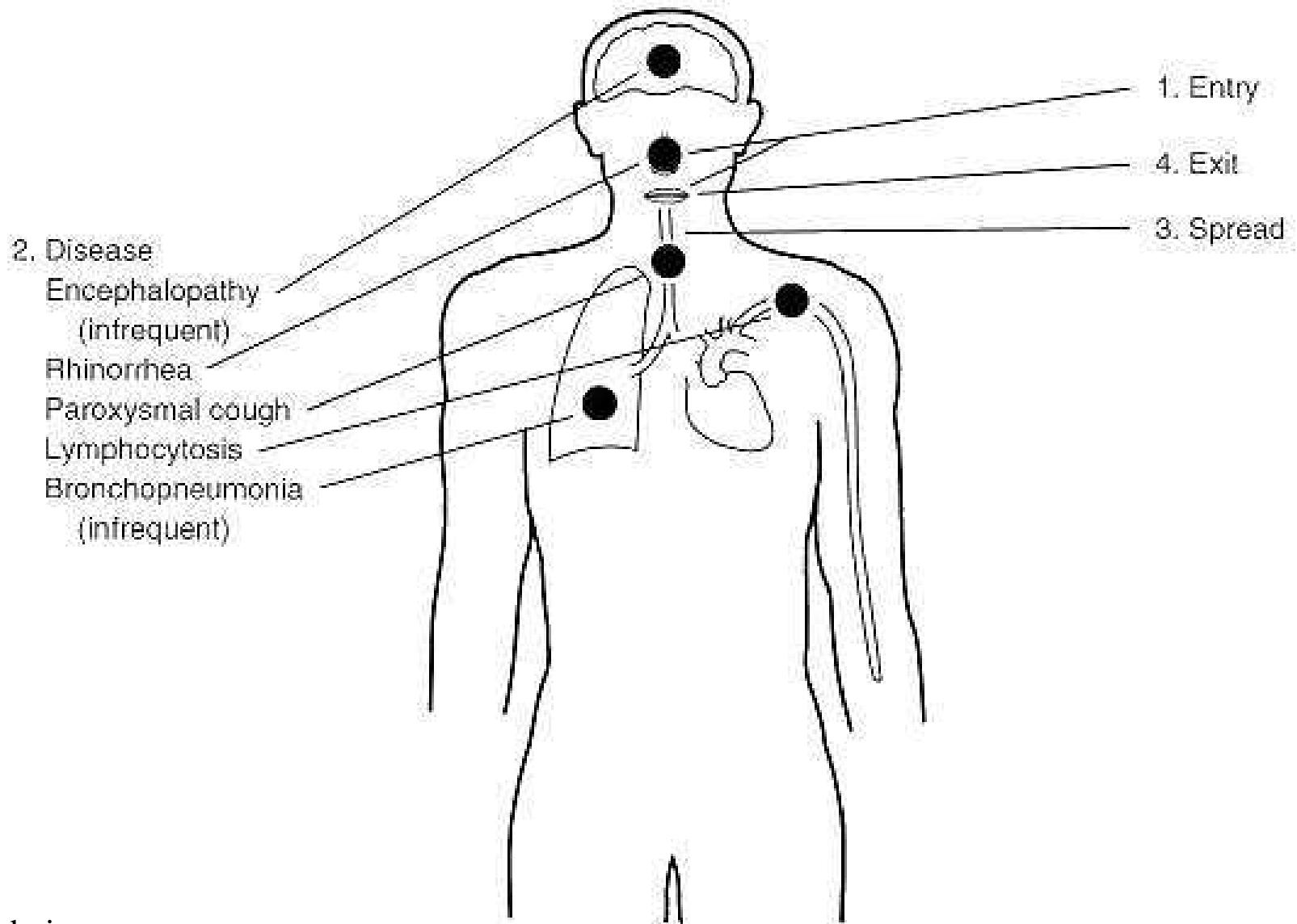
## Pathogenesis, Fertőzés:

Fertőzés forrása: Betegek – prodromalis és katarrhalis Stadiumban

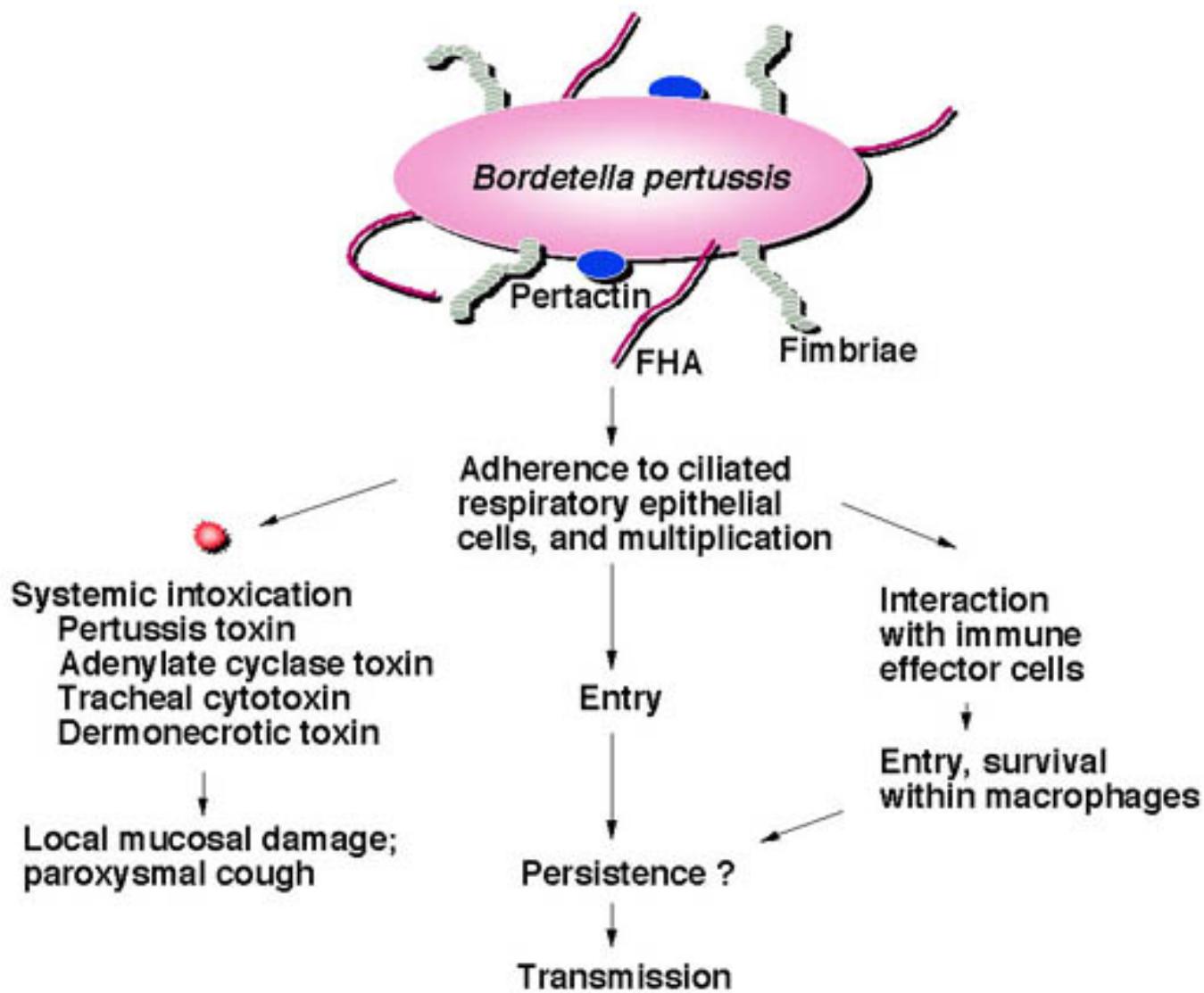
Behatolási kapu: Légzőtraktus

Átvitel: Cseppfertőzés → ÉRZÉKENY!  
55°C; 30'

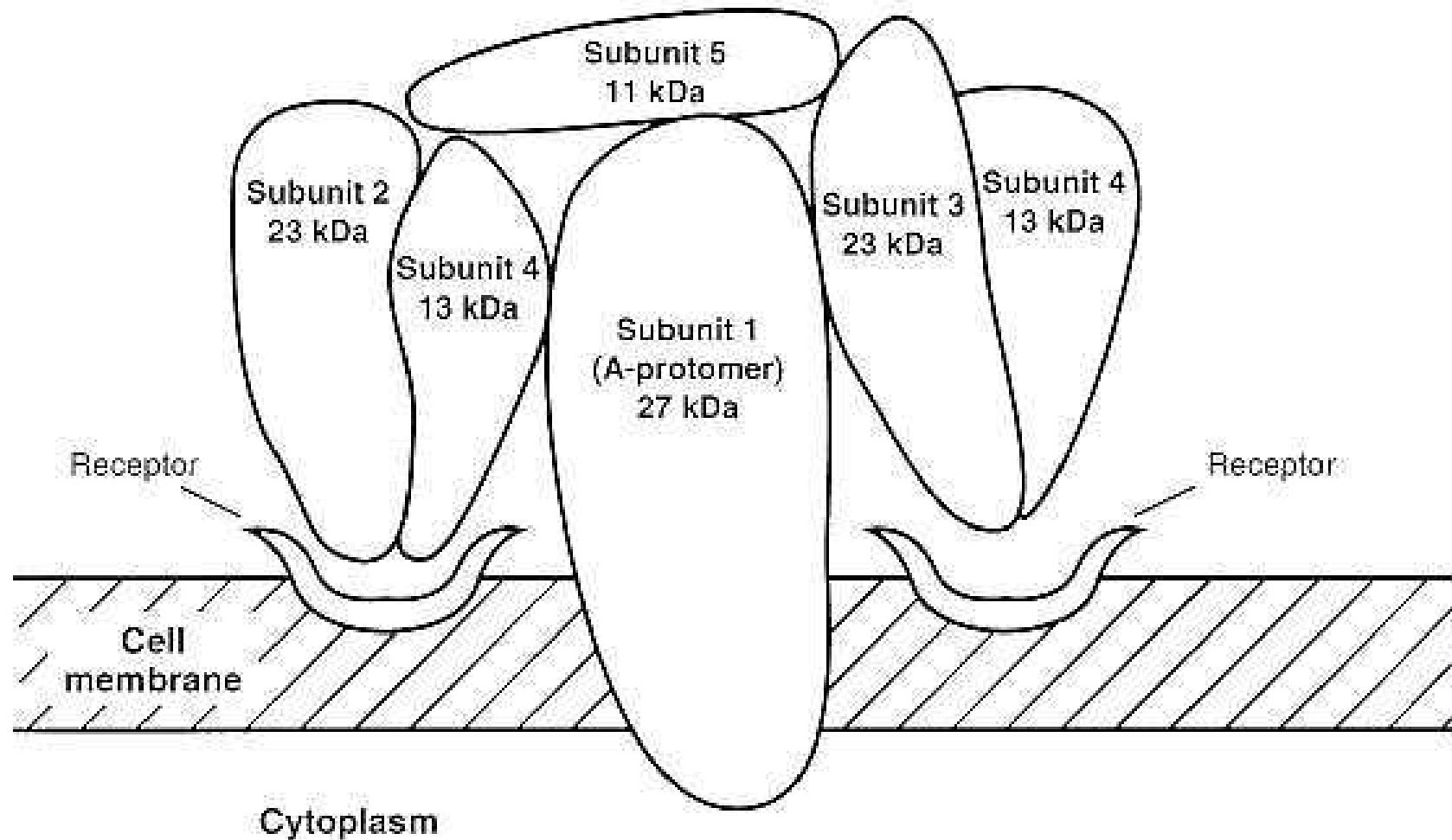
# FIGURE 31-1 Pathogenesis of whooping cough.



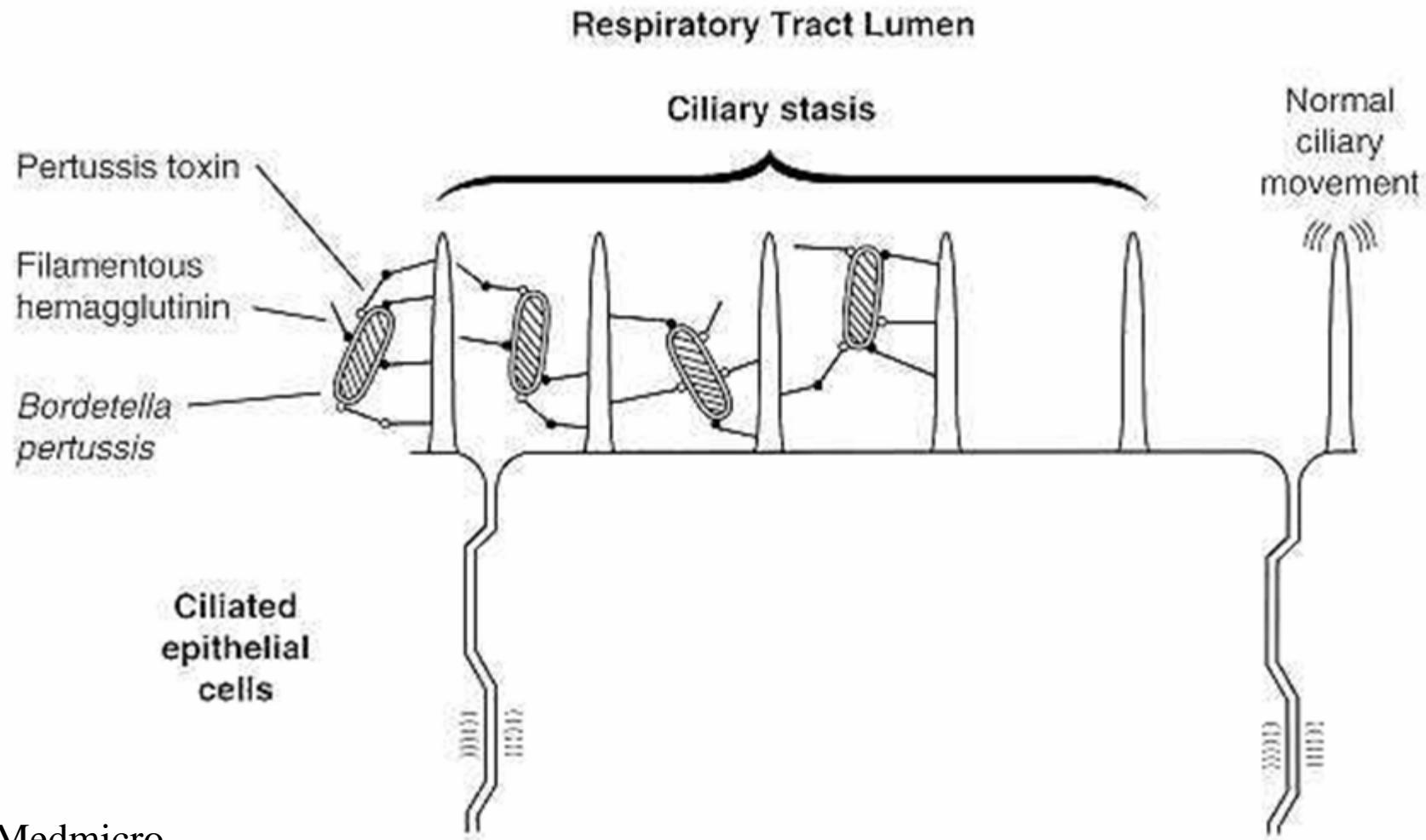
## Pathogenesis of *Bordetella pertussis*



# FIGURE 31-3 Binding of pertussis toxin to cell membranes.



## FIGURE 31-4 Synergy between pertussis toxin and the filamentous hemagglutinin in binding to ciliated respiratory epithelial cells.



# *Bordetella pertussis*

Kórkép:

**Szamárköhögés / Pertussis**

(Peribronchialis gyulladás, Intersticiális Pneumonia)

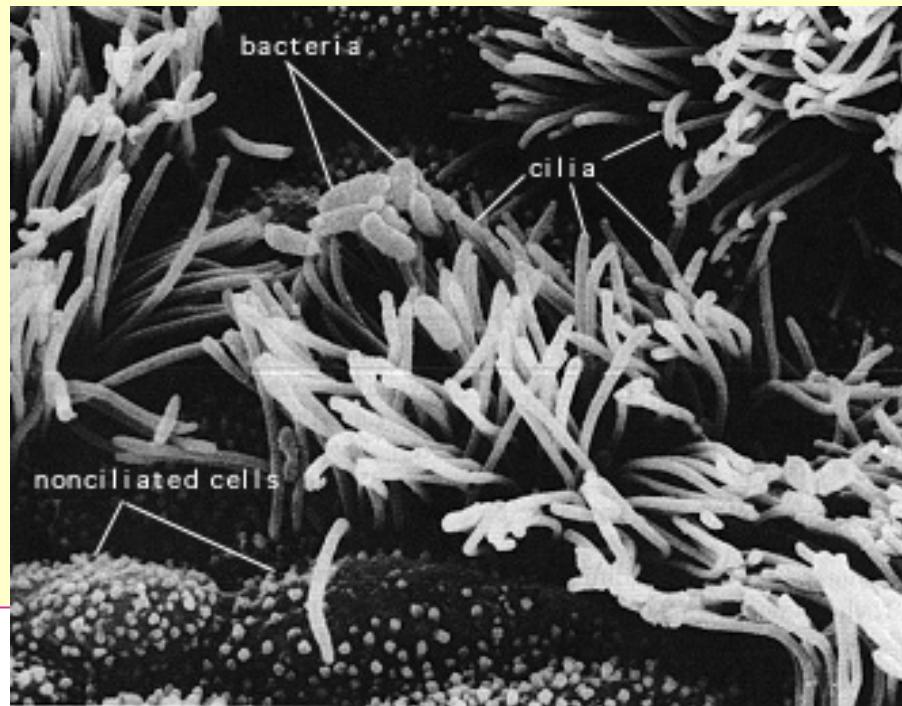
**4-Phasis:**

Prodroma,

Katarrhalis,

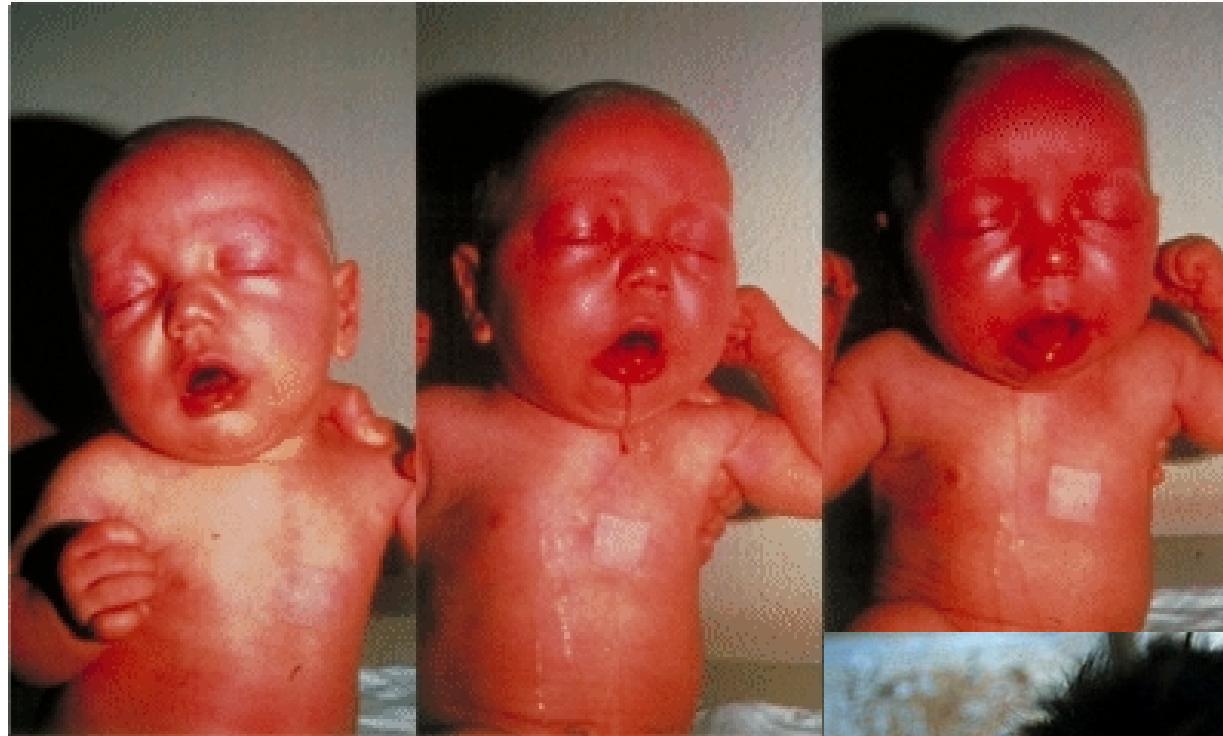
Paroxysmalis,

Rekonvalescens



Colonization of tracheal epithelial cells by *B. pertussis*

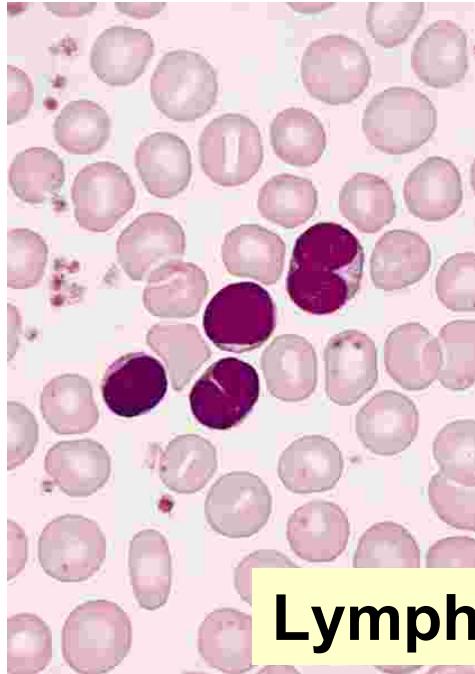
[web.umr.edu/~microbio](http://web.umr.edu/~microbio)



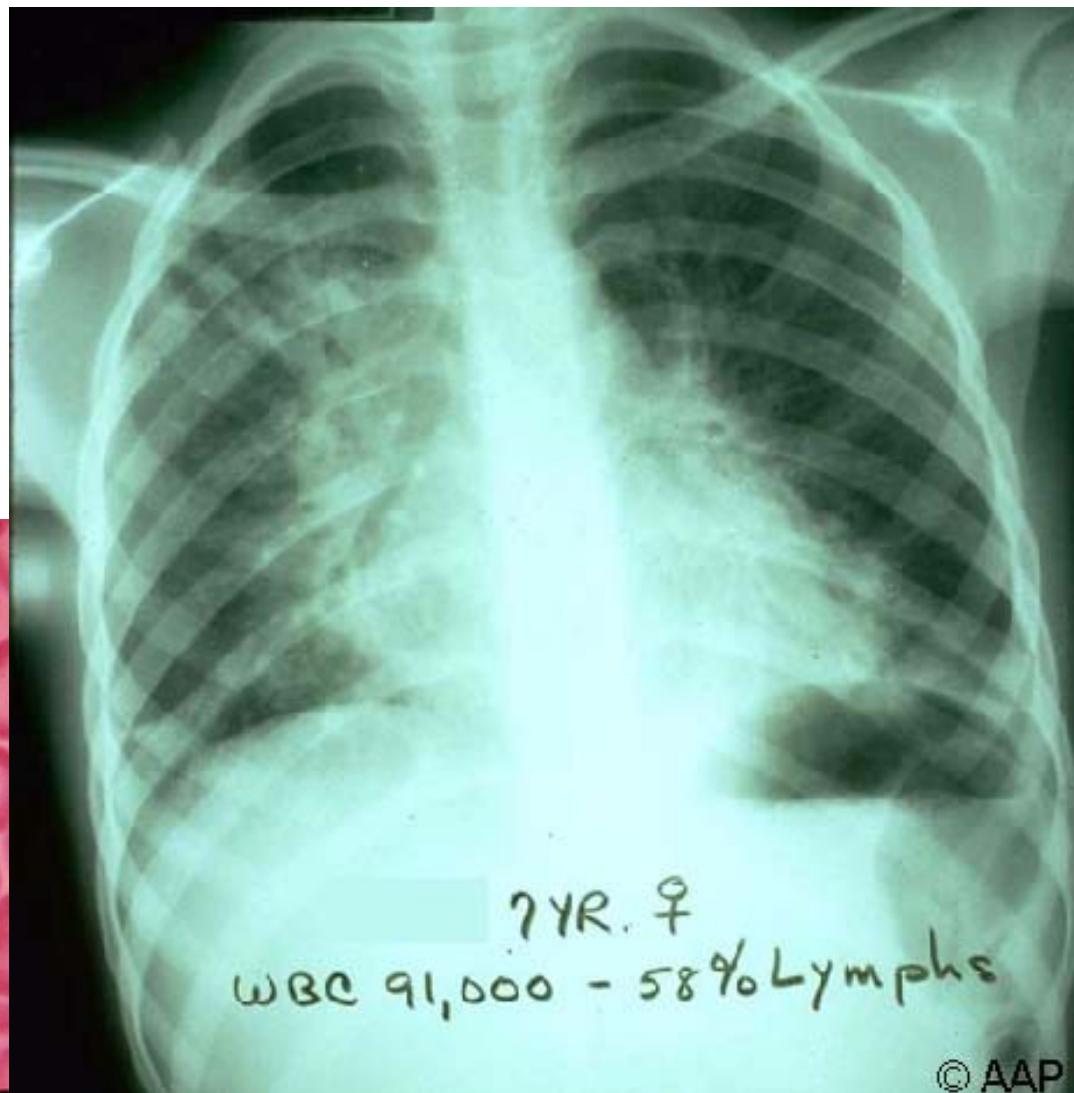
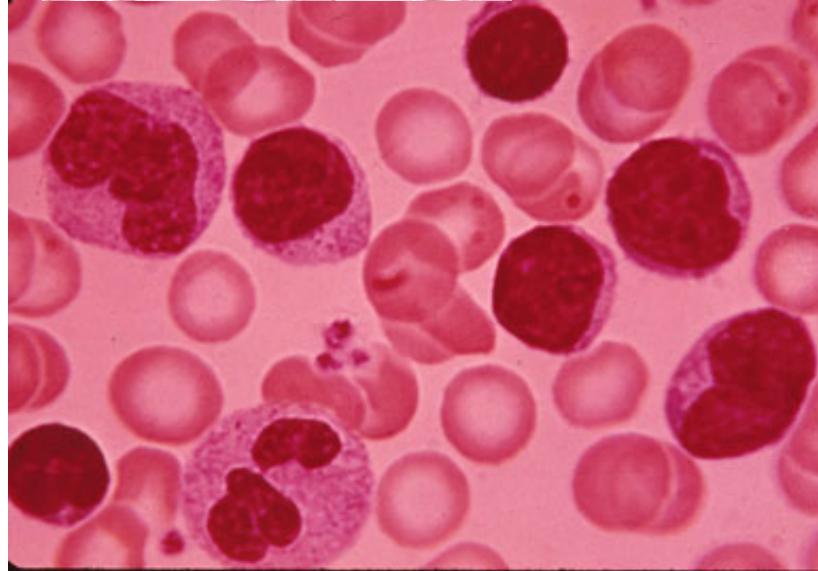
## Pertussis – paroxysmalis Phasis



# Pertussis - Diagnosis



Lymphocytosis



© AAP

# *Bordetella pertussis*

## Diagnosis

### Tenyésztés:

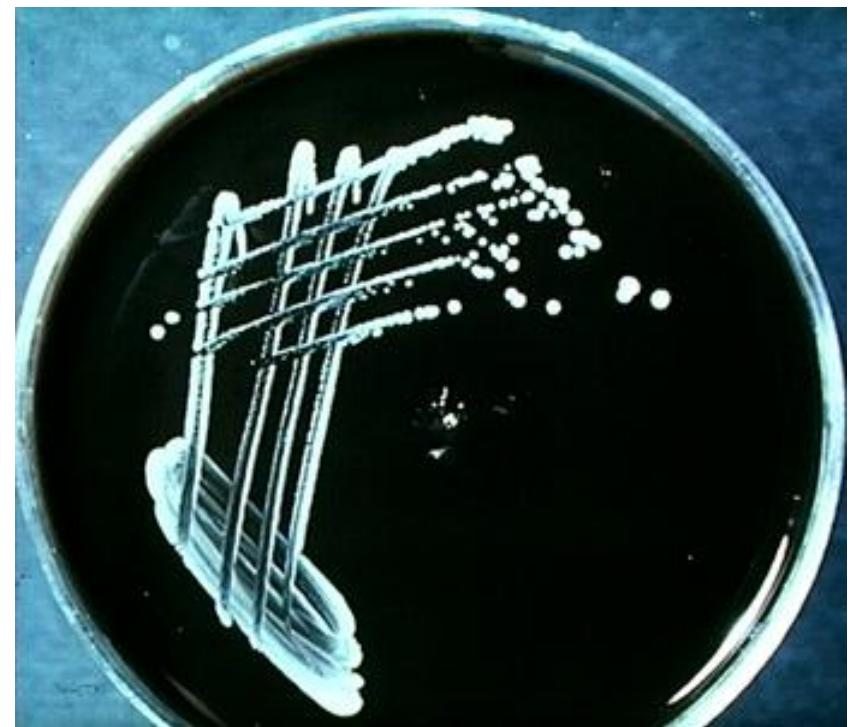
Bordet – Gengou  
Direkt ráköhögni!

Charcoal Medium →

### Szerológia:

IgM, IgA, IgG kimutatás

PCR



# ***Bordetella pertussis***

**Therapia:**

Makrolid

**Prophylaxis:**

Aktív Immunizálás – acelluláris oltóanyag **DaPT**,  
benne:

Toxoid

FH/Pilus

Pertactin

**DiPerTe** – előlt B. pertussis baktérium



Korfu, 2006

VÉGE