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COURSE AND FORMS OF INFECTION – I

The 10th lecture for the 2nd-year students, April 20th, 2015

What is the pathogenesis? – revision

- Pathogenesis explains the <u>origin and</u> <u>development of pathological symptoms</u>
- What does the pathogenesis of infection include?
- The <u>way the agent spreads</u> through the macroorganism
- Mechanisms of <u>defence against it</u>
- Actual <u>causes of symptoms</u>:
 a) either the infectious agent itself,
 b) or the reaction of macroorganism to it

Spreading by means of <u>lymph</u> – revision

- skin → regional lymphatic nodes: pyogenic cocci, F. tularensis, Y. pestis; arboviruses
- oropharynx, tonsils → <u>cervical nodes</u>: S. pyogenes, C. diphtheriae, M. tuberculosis, anaerobes (Actinomyces israeli, Prevotella), T. gondii
- lungs → <u>hilar nodes</u>: M. tbc, B. anthracis, other respiratory pathogens
- genital mucosa → inguinal nodes: Treponema pallidum, Ch. trachomatis L1-L3, H. ducreyi
- Peyer plaques → <u>mesenteric nodes</u>: Yersinia enterocolitica, enteric adenoviruses, enteroviruses

Spreading by means of <u>blood</u> – revision

- Agents of <u>all generalized infections</u>: exanthematic viruses, enteroviruses, arboviruses, *Treponema pallidum*, *Salmonella* Typhi and many others
- Agents of <u>pneumonia</u> commonly appear in blood: especially Strept. pneumoniae
- Sometimes agents of other systemic and local infections: during meningitis, pyelonephritis (urosepsis), suppurating wounds and suchlike

Spreading *per continuitatem* – revision

- From <u>cell to cell</u>: HSV, RSV, listeriae, yersiniae
- By means of secretion down the mucosa: agents of respiratory, enteric and urogenital infections
- From the site of <u>arthropod biting to its vicinity</u>: arboviruses, *Borrelia burgdorferi*
- From the wound to adjacent tissue: Streptococcus pyogenes, Clostridium perfringens
- From the middle ear to meninges: S. pneumoniae, Haemophilus influenzae type b
- From lungs to pleura: agents of pneumonia

Spreading along <u>nerves</u> – revision

Either <u>axonally</u> (within nerve fibres)

or by progressive infection of Schwann sheath

HSV, VZV, B-virus, rabies virus Mycobacterium leprae Naegleria fowleri tetanic toxin

Elimination of agent from the body – revision

From the mucosa of respiratory tract and oral cavity, intestine, urogenital tract, eye **From skin lesions** By means of urine **From blood**

Elimination from respiratory tract – revision

Sneezing:

in particular agents of common cold (rhinoviruses, coronaviruses), from bacteria e.g. Neisseria meningitidis **Coughing:** other respiratory viruses (primarily influenza virus), exanthematic viruses (VZV, morbilli virus, rubella virus), Neiss. meningitidis, Bordetella pertussis, Mycob. tuberculosis, Yersinia pestis

Elimination from alimentary tract – revision

Saliva:

- HSV, EBV, mumps virus, *Str. pyogenes* Stool:
- enteroviruses (incl. poliovirus), HAV, HEV salmonellae incl. Salm. Typhi, shigellae, EPEC, ETEC etc., V. cholerae, C. difficile Entamoeba histolytica, Giardia lamblia Ascaris lumbricoides, Taenia saginata

Elimination from urogenital tract – revision

- From diseased mucosae:
- Agents of <u>classic venereal infections</u>: in Europe Neiss. gonorrhoeae, Treponema pallidum
- Agents of <u>other</u> sexually transmitted diseases (<u>STD</u>): *Chlamydia trachomatis* serotypes D-K, papillomaviruses, HSV-2
- By means of urine:
- Salmonella Typhi
- Agents of congenital infections (rubella virus, CMV) Exotic viruses of hemorrhagic fevers (Ebola)

Elimination <u>from skin</u> lesions – revision

Staphylococcus aureus Streptococcus pyogenes Varicella-zoster virus (agent of chickenpox and shingles) **Papillomaviruses (agents of warts) Dermatophytes** (e.g. *Trichophyton rubrum*, **Microsporum canis, Epidermophyton** floccosum)

Sarcoptes scabiei (itch-mite)

Elimination from blood – revision

By means of vectors:

tick-borne encephalitis virus – ticks, yellow fever virus – mosquitoes

Rickettsia prowazekii – lice, Yersinia pestis – fleas, Borrelia recurrentis – lice Malaric plasmodia – mosquitoes

By means of small <u>cracks in mucosa</u>: HBV, HIV

Infection

Definition:

Infection = a relation between the pathogenic microbe and the macroorganism (= ecological point of view)

Infection colonization:

- **Infection** = situation when an etiological agent
- 1) penetrates into an organism and multiplies in it, or
- 2) it settles on bodily surfaces (skin or mucosae) and unfavourably affects them

Colonization = situation when

- 1) a non-pathogenic microbe settles on a bodily surface, or
- 2) a pathogen located there does not cause pathological symptoms

Relationship between the microbe and the host – revision The relationship is a dynamic one and influenced by the environment: microbe → host

Illness is not a rule – peaceful coexistence is usually better for the parasite In spite of that the host tries to get rid of the parasite – to destroy, remove or at least to keep it in one spot

environment

Course of infection – I

Four components can be distinguished during the course of infection:

- Incubation time
- Prodromes
- Typical syndrome (= complex of symptoms) of the infectious disease
- Convalescence

Course of infection – II

Incubation time

salmonellosis ½–1 day, influenza 1–2 days, tbc 2–8 weeks, hepatitis B 90–100 days

<u>Prodrom</u>es

- not always; nonspecific (†T, headache, feeling ill etc.), several hours to days
- <u>Typical syndrome</u> of infectious disease as described in textbooks

Convalescence

from subsiding troubles till normalization of laboratory results (except antibodies!)

Course of infection – III

Relapse

the same agent, infection comes on again during the convalescence

<u>Recurrence</u>

the agent remains in the body, infection comes on again only after recovery (Brill-Zinsser disease = recurrence of epidemic typhus)

Reinfection

new infection by the same agent from outside

Superinfection

infection by another agent before recovery from the first infection

Forms of infection

Inapparent infection (without symptoms) sole consequence: development of immunity (usually by means of antibodies) <u>Manifest infection (with symptoms)</u> subclinical: non-characteristic signs only abortive: only some symptoms or slightly manifested ones clinical: typical signs as in textbooks foudroyant, fulminant: very abrupt, with dramatic symptoms

Duration of infection

<u>Acute</u>: days (common cold, salmonellosis) to weeks (majority of infections)

<u>Subacute</u>: months – either as a complication of any infection, or as the rule (some kinds of hepatitis, warts, sepsis lenta)

<u>Chronic</u>: years (tbc, lepra, dermatomycoses, parasitic infections)

Fulminant, foudroyant: very rapid course – hours (meningococcal sepsis)

Extent of infection

<u>Local</u>: portal of entry & regional nodes, or a specific organ (common cold, ringworm, warts, uncomplicated gonorrhoea, abscessus in an organ)

Systemic: whole organ system (influenza, lung tbc, meningitis, extensive pyodermia, pyelonephritis, pelvic inflammatory disease)

<u>Generalized</u>: regularly (exanthematic viroses, typhoid fever, exanthematic typhus), or as a complication (sepsis after injury, during cystitis or cholecystitis, salmonellosis in a newborn)

Focal infection – I

Focal infection theory:

chronic infection limited to a certain focus can result in a systemic illness with symptoms in quite a different site

Concept of focal infection used to be very fashionable formerly in diverse medical branches

In the name of so-called sanation of focuses thousands of patients were bona fide subjected to tooth extractions, tonsillectomies, cholecystectomies and other surgical interventions without proving the usefulness of these procedures by controlled studies

Focal infection – II

- The <u>connection between systemic disease and a</u> <u>local infection has been proved only in</u>
- <u>rheumatic fever</u> inflammation of heart, kidneys and joints after tonsillar infection by Streptococcus pyogenes
- <u>Reiter's syndrome</u> reactive arthritis after

 sexually transmitted urogenital infection by
 Chlamydia trachomatis serotypes D-K,
 intestinal infection caused by pathogens from
 genus Salmonella, Shigella, Yersinia or
 Campylobacter
- <u>hemolytic-uremic syndrome</u> after intestinal infection by <u>Escherichia coli</u> serotype O157:H7
- <u>sterile mykids</u> e.g. on palms during tinea pedis

Special types of <u>chronic</u> infections

Inapparent chronic infections can be clasified as

- 1. <u>latent</u>: agent hides in a non-infectious form, or it escapes from the infected cell after an activation of infection only
 - **HSV and VZV:** nerve ganglia cells, <u>CMV</u>: kidney and salivary glands cells, <u>EBV</u>: lymphocytes
- 2. <u>persistent</u>: agent can be detected by routine methods, because it is present mostly in an infectious form
- Both types are markers of failing immunity Both types can be activated

Examples of persistent infections

<u>Bacterial</u>: *Rickettsia prowazekii* (activation of exanthematic typhus = m. Brill-Zinsser), *Salmonella* Typhi (carriers), *Mycob. tbc* (lymphatic nodes)

<u>Viral</u>: HBV (hepatocytes), adenoviruses (adenoids), JCV and BKV (kidneys), congenital infections by CMV and rubella virus

<u>Parasitary</u>: hypnozoites of *Plasmodium ovale* and *P. vivax* (liver), *Toxoplasma gondii* bradyzoites (nodes, muscles, brain)

Recommended reading material

Paul de Kruif: Microbe Hunters Paul de Kruif: Men against Death Axel Munthe: The Story of San Michele Sinclair Lewis: Arrowsmith André Maurois: La vie de Sir Alexander Fleming Hans Zinsser: Rats, Lice, and History Michael Crichton: Andromeda Strain Albert Camus: Peste Victor Heisser: An American Doctor Odyssey Richard Preston: The Hot Zone

> Please mail me other suggestions at: <u>mvotava@med.muni.cz</u> Thank you for your attention