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#### \* No conflict of interest

The pioneer : Charles Nicolle Book : Le destin des maladies infectieuses, 1933 Republished in 1993

- \* Follower of Pasteur
- \* **Nobel prize winner** (discovery of the mode of transmission of typhus)
- \* Director of the Pasteur institute of Tunis
- Wrote in the 1930's that chronic diseases could be linked to « Les infections inapparentes » (silent infections, occult infections...)

#### Willy Burgdorfer was a strong supporter of Charles Nicolle's theories

- \* PhD thesis on relapsing fever in East Africa due to Borrelia duttonii
- \* Published in the 1950's about « Occult infections »
- \* **Studied the Rocky Mountain Spotted Fever** (Rickettsia rickettsii)
- \* « Discovered » Borrelia burgdorferi
- \* Worked on Coxiella burnetii (considered as a bioweapon)
- Made Rickettsia helvetica (« l'agent suisse ») from American and Swiss genomes of Rickettsiae, present in the blood of patients during the initial Lyme disease oubreak

## The pioneers Giroud & Jadin

- Prof. Paul Giroud, from the Pasteur Institute, France, assistant of Charles Nicolle
- Prof. Jean Baptiste Jadin, from the Pince Leopold Institute of Antwerp, Belgium, field researcher in Congo Kinshasa and Rwanda. Father of Dr Cécile Jadin
- \* Loo & Menier, Annales de Médecine Physiologique 1960. Possible link between intracellular bacteria and neuropsychiatric diseases
- \* In the 1980s, they described :
  - \* associations of certain diseases and protozoans (single-cell parasites)
  - \* the possible pathogenic role of intracellular bacteria (Rickettsia spp, Rickettsia-like particles and Chlamydia spp)
  - developed semi-quantitative antibody tests (not recognized) and treated many patients with doxycycline

#### **Crypto-infections in dogs**

- \* Donatien & Lestoguard, 1935
- \* Intracellular bacteria in the blood of sick dogs
- \* Rickettsia canis, renamed Ehrlichia canis

# Multiple terms used to define crypto-infections

- Infections inapparentes (Charles Nicolle)
- \* Silent infections
- Occult infections (Willy Burgdorfer)
- \* Hidden infections
- \* Stealth infections
- \* Cold infections (Luc Montagnier)
- \* Post tick-bite syndrome
- \* Tick-associated poly-organic syndrome (TAPOS) (Christian Perronne)
- \* Poly-organic chronic syndrome
- Multiple systemic infectious disease syndrome (MSIDS) (Richard Horowitz)
- \* **Persistent polymorphic syndrome possibly due to a tick-bite** (SPPT in French) (French High Authority for Health, HAS)

## **Crypto-infections**

- \* Missed by the radar of the institutions and methods
- \* Confusing variety of disorders and symptoms
- \* Borrelia spp :
  - \* ability to change form
  - \* to modulate the biochemistry of their receptors
  - \* allowing them to remain « hidden » for months or years
  - not recognized by the immune system, inaccessible to antibiotics
- \* Borrelia antigens at the surface of host cells induce autoimmunity
- \* Reliable diagnostic tests have never been developed

Our cells are made from tubules of spirochetes (Microcosmos, Sagan & Margulis)

- \* Bronchial cells with undulating cilia
- \* Insulin secretion through microtubules
- \* Neurons
- \* Visual cells
- \* Flagella of sperms
- Cell division (mitosis): microtubules arranged in beams that allow chromosomes to split
- \* Etc.

#### Lack of research

- \* Lack of research into many chronic inflammatory, autoimmune and degenerative diseases (e.g. funding of HIV research vs funding of Lyme research !)
- \* Ignoring the multiple examples of the microbial origin of some cancers
- \* Veterinarians have often better diagnostic tests
- \* WHO : « One health » : animals and humans share the same environment

# We only find what we look for... Otherwise, it « does not exist »

- \* Muriel Vayssier Taussat et al. Emerg Infect Dis, 2016.
- \* PCRs for Bartonella spp on blood samples from « chronic Lyme » patients. They found :
- \* **Bartonella already known in humans** : Bartonella henselae
- \* But also... other Bartonellae previously unknown in humans (but already known by vets in animals):
  - \* Bartonella doshiae
  - \* Bartonella schoenbuchensis
  - \* Bartonella tribocorum

#### Lack of research on treatments

Not a single randomized study to evaluate a really prolonged (> 3 months) antibiotic treatment for post-treatment Lyme disease syndrome (PTLDS)

**Projects not funded** 

The theory of spontaneous generation : never ceases to be resurrected

- \* Diseases can occur from nothing
- \* Idiopathic diseases = diseases of « idiots » (experts, not patients)

## Organic or psychic ?

- Crypto-infections, the missing link between Pasteur and Freud
- \* Both studied unexplained diseases
- Chronic Lyme patients and patients suffering from other crypto-infections are in a no-man's land between microbiology and psychoanalysis

#### Ticks can transmit many microbes

#### \* Bacteria:

- Borrelia spp, Ehrlichia chaffeensis (US), Ehrlichia canis, Anaplasma phagocytophilum (Europe), Rickettsia spp, Francisella tularensis, Coxiella burnetii, Bartonellla spp, Neoehrlichia mikurensis
- \* **Parasites :** Babesia spp, Theileria spp.

#### \* Viruses

\* **Symbiotes :** *Midichloria mitochondrii* (live in mitochondria and able to destroy them). Similarities between symptoms of « chronic Lyme » and mitochondrial diseases

Real time micro-organisms PCR in 104 patients with polymorphic signs and symptoms that may be related to tickbite. Lacout A. ... Perronne C. et al. Eur J Microbiol Immunol 2021

- \* Real time PCR from saliva, urine, venous blood, capillary blood
- PCR for : Borrelia burgdorferi sensu lato, Borrelia miyamotoi, Borrelia hermsii, Bartonella spp., Bartonella quintana, Bartonnella henselae, Ehrlichia spp., Anaplasma spp., Rickettsia spp., Coxiella burnetii, Brucella spp., Francisella tularensis, Mycoplasma spp., Chlamydia spp., Babesia spp., Theileria spp.
- \* **48% of patients: polyinfected,** 25% harboured at least 3 different micro-organisms
- \* **Borreliae were not frequently isolated**, far behind Mycoplasma, Rickettsia, and Ehrlichia
- \* **Piroplams** (Babesia and especially Theileria) were frequently isolated
- \* The most sensitive matrices were : saliva, then urine and capillary blood. Venous blood was the less sensitive.

Borrelia miyamotoi : 43 cases diagnosed in France by real time PCR in patients with persistent polymorphic signs and symptoms. Franck M. ... Perronne C. et al. Front Med 2020.

- \* **Borrelia miyamotoi is a relapsing fever Borrelia** which can induce symptoms related to Lyme disease
- \* **Discovered in 1995** (rodents and ticks)
- \* Real time PCR from blood of 824 patients with chronic symptoms
- \* PCR positive in 43 patients (5,22%)
- \* Erythema migrans : 16%
- \* All patients : fatigue, joint pain, neuro-cognitive disorders
- \* Half of patients : chills, hot flushes, sweats
- \* **41,9% of patients :** respiratory disorders (chest tightness, lack of air)
- \* 35,5% of patients : episodes of relapsing fever
- \* Usually, no cross reaction with *B. burgdorferi* serology

Many crypto-infections : not due to tick bites

#### The infectious cause of autism ?

- Many autistic children have improved or were cured thanks to anti-infectious treatments (antibiotics <u>+ antiparasitic or</u> antifungic drugs) and low gluten diet
- \* Acute digestive or cutaneous symptoms may precede the occurrence of autistic symptoms
- \* Sutarella and Ruminococcus are found in the feces of autistic children
- \* In France, in 2012, I tried with Dr Philippe Raymond, to set a randomized clinical trial (azithromycin vs placebo) to confirm the efficacy of antibiotics (« Autibiotic » trial). Initially supported by the Minister of health, the trial was attacked and abandoned !



- \* Randomized placebo controlled studies are published
- Showing the efficacy of antibiotics (at least partial and transient)
- \* The link between *Bartonella spp.* and psychiatric diseases

# The infectious cause of Alzheimer's disease (1)

- The fight between Emil Kraepelin (Alois Alzheimer's boss) and Sigmund Freud about the organic or the psychological cause of neuropsychiatric diseases.
- \* By comparison with syphilis and its agent Treponema pallidum, Alzheimer was convinced that a bacterium was responsible for « his » disease
- Neurosyphilis, Alzheimer's disease and neuroborreliosis : brain sections show the same lesions

# The infectious cause of Alzheimer's disease (2)

- \* Alan Mc Donald, Judith Miklossy. Confirmed by Allen et al. J Neuroinfect Dis 2016.
- \* Alzheimer's disease is a neurospirochetosis
- \* Fulfill the criteria of Henle-Koch postulate
- \* Borreliae and non-syphilitic Treponemae were evidenced in the brains
- \* Link between damaged gums and spoiled teeth (Porphyromonas gingivalis) (dentist St John Crean)



- \* TNF-receptor associated periodic syndrome
- \* One patient suffering for years of episodes of high grade fever and symptoms. Cured after 4 months of antibiotic and hydroxychloroquine. Long term follow-up
- \* Galpérine et al. J Global Infect Dis 2020



- \* Biscardi et al. Mycoplasma pneumoniae and asthma in children. Clin Infect Dis 2004
- \* Role of Chlamydia

## The bacterial cause of arteriosclerosis ?

- \* IV injection of Chlamydia pneumoniae in animals : provides atheroma plaques
- \* Animals : efficacy of doxycycline on aortic aneurysms
- \* Humans : several clinical trials were inconclusive
- Dutch study, Arnoud-Meyer et al 2013. Patients with a small aortic aneurysm. Doxycycline vs placebo. Significant increase in the size of aneurysm with doxycycline (local herx with inflammatory reaction ?)
- \* Fluoroquinolones may worsen some aneurysms (herx?)

## **Rheumatoid arthritis**

 Several randomized studies showing the efficacy of long term anti-infective treatment

- \* Minocycline > placebo (O'Dell et al. Arthritis Rheum 1997)
- \* Doxycycline : conflicting results
- \* Hydroxychloroquine alone : moderate effect
- \* Clarithromycine > placebo (Ogrendik et al. Curr Med Res Opin 2007)
- \* Only the « anti-inflammatory » effect of antibiotics and hydroxychloroquine ?

## Helicobacter pylori

- Agent of gastric and duodenal peptic ulcer
  - \* Ulcer considered as a psychosomatic disease
  - \* Gastric spirochetes already seen in the microscope in 1875!
- \* Oncogenic
- \* Link with the severity of Parkinson's disease (Tan 2015)
- \* Role of non pylori species of Helicobacter in hepatobiliary diseases ?

## **Relapsing fevers**

- \* 15 species of Borrelia transmitted by ticks or lice
- Relapsing fever reported in Madagascar in the 18th century (Drury, 1729)
- « Tick fever » reported by Dr Livingstone (explorer and missionary in East Africa)
- \* After World war 1: Hispano-african relapsing fever (Spirochaeta hispanicum renamed Borrelia hispanicum)
- \* 1950's : Studies by Willy Burgdorfer
- Sahel : high proportion of « malaria » cases are in fact borreliosis (Vial et al, Lancet 2006, Mediannikov et al, Emerg Infect Dis 2014)
- \* Diagnosis : microscopic blood smears
- \* Not anymore diagnosed in « modern » medicine

#### Whipple's disease

\* Previouly often considered as psychosomatic

\* Until the discovery of Tropheryma whipplei

#### **Mycobacterial crypto-infections**

\* The example of :

- \* Sarcoidosis
- \* Crohn' disease

# The infectious origin of sarcoidosis (1)

- \* Due to L-forms of Mycobacteria spp. No cell wall : no staining of acid fast bacilli
- Rare cases of simultaneous onset of sarcoidosis symptoms in persons exposed to non tuberculous Mycobacteria (e.g. jacuzzi)
- \* Reported cases of isolation of non-TB Mycobacteria from bronchial samples
- \* Correlation with anti-mycobacterial antibodies
- \* Link with Chlamydia infection
- \* More frequent in nursing personnel
- \* French Antilles : West Indians have no additional risk, but after living in Metropolitan France, their risk increases. Environmental factors.

# The infectious origin of sarcoidosis (2)

- I was vice-director of the National reference lab for tuberculosis and other *Mycobacteria*, Pasteur Institute, Paris
- \* I cured patients with sarcoidosis prescribing antimycobacterial combinations used for *Mycobacterium avium* in AIDS patients.
- \* 1996. Clinical trial « Mycosarc » to confirm the efficacy of clarithromycin. Promotor : Pasteur Institute. The protocol was criticized, attacked and finally abandoned

# The infectious origin of Crohn's disease (1)

- \* Due to Mycobacterium avium subspecies paratuberculosis, cause of Johne's disease in cattle (bovine diarrhoea)
- Oubreaks of human cases of Crohn's disease in villages living downstream from the fields occupied by herds contaminated by Johne's disease
- \* Human cases of transmission within couples
- \* *M. avium subsp paratuberculosis* : isolated by culture or PCR from human colon biopsies
- \* Culture only possible in liquid media, kept 4 to 6 months
- \* Causality fulfills the Henle-Koch criteria (Greenstein, Lancet Infect Dis 2003)

# The infectious origin of Crohn's disease (2)

#### \* English open label clinical study.

- \* In 52 patients, 100% efficacy of a combination of clarithromycin (or azithromycin) and rifabutin. Two years follow-up (Gui et al. J Antimicrob Chemother 1997).
- \* Study not taken into account because of the lack of placebo !
- I cured patients and designed a clinical trial which was also attacked and abandoned

## The infectious causes of cancers

- \* Not only viruses (HTLV-1, EBV, HBV, HCV, HPV, HHV-8, etc.)
- \* Bacteria:
  - \* Helicobacter pylori
    - \* Gastric adenocarcinoma, gastric lymphoma (MALT)
  - \* Campylobacter jejuni
    - \* Immunoproliferative disease of the small intestine
  - \* Neoehrlichia mikurensis
    - \* Malignant lymphomas
  - \* Porphyromonas gingivalis
    - \* Oesophageal cancer
  - \* Bartonella spp.
    - \* Malignant melanoma

#### \* Parasites :

- \* Theileria (piroplasms)
  - \* Malignant lymphomas
  - \* Able to transform healthy human cells into cancerous cells

#### **Crypto-infections : Code of silence**

- Hidden infections, but also hidden medical and scientific informations about these diseases
- Many published data not included in « official » recommendations
- \* The reign of censorship, particularly in the medical journals
- Major medical journals at the orders to publish misleading editorials

#### Silent epidemics are denied

- \* Lack of reliable diagnostic tests
- \* Lack of good statistics
- \* Difficult to convince politicians of the reality of a huge global problem, which should require fast-track research

# The ridiculous planned disappearance of infectious diseases

- \* After World war II, microbiology is denigrated
  - Advances in hygiene, nutrition, vaccines and anti-infective drugs, especially antibiotics
  - Would quickly sweep away microbes, intruders worthy only of the Middle Ages
- \* But... Life is either infectious or it does not exist at all
- Promotion of « more noble » specialties : genetics, immunology, statistics...

# Cost of chronic Lyme disease and other crypto-infections

- \* Wandering from many doctors to many medical departments
- \* Multiple examinations
- \* Multiple inefficient treatments
- \* Multiple alternative treatments (e.g. phytotherapy), or alternative tests, not reimbursed
- \* Multiple hospitalizations, including in psychiatry wards
- \* Multiple sick leaves
- \* Chronic handicap

# Cost of chronic Lyme disease, Belgium. University of Gent

- Cost of illness in patients with post-treatment Lyme disease syndrome in Belgium. Willems R, Verhaeghe N, Perronne C, Borgemans L, Annemans L. Eur J Public Health. 2023
- \* Mean annual direct costs per patient = 4.618 euros
  - \* Out of pocket expenditures : 49,5%
- \* At population level :
  - \* Direct costs = 19,4 millions euros
  - \* Indirect costs = 151,5 millions euros

# Chronic pathologies : multiple causes of persistence

- \* Microbes are not the only cause
- Anti-infective agents cannot achieve cure in all patients
- \* Genetic factors
- \* Auto-immune processes
- \* Stress
- \* Heavy metals
- \* Psychological traumas
- \* Domestic violence
- Sexual violence during chilhood
- \* Food habits
- \* Physical activities

# Crypto-infections: Triggering factors of chronic syndromes or relapses

\* Stress

\* Some acute infections able to transitorily decrease the immune defenses

- \* Infectious mononucleosis (EBV virus)
- \* Cytomegalovirus (CMV)
- \* Malaria
- \* Dengue
- \* Chikungunya
- \* Covid-19 (Long Covid)

# Military affairs : lab modifications of ticks and Borreliae

- \* Modified by man (bioweapons)
  - **Book « Bitten »**, Kris Newby, 2019, Harper (archives from Willy Burgdorfer's lab)
  - Motion voted by the US Chamber of Representatives, July 2019. Establishment of an Investigative Committee on the Pentagone activities

## « Bitten », Kris Newby, Harper The secret archives of Willy Burgdorfer



The Secret History of

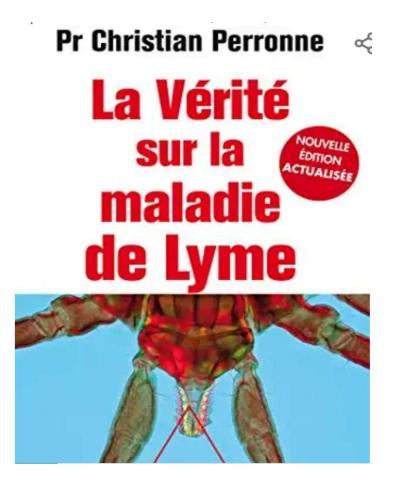
Lyme Disease and

Biological

Weapons

KRIS NEWBY

Books about Lyme disease and other crypto-infections (Odile Jacob, Hammersmith)





Denial, censorship and suppression – the truth about what lies behind chronic disease

#### DR CHRISTIAN PERRONNE

"There are many occult infections triggering a cascade of infection, inflammation and autoimmunity, of which Borrelia, the bacterium causing Lyme disease, is the prototype" Dr Jack Lambert, Consultant in Infectious Diseases, Dublin, Ireland