# The bacteria Borrelia burgdorferi s.l. and the clinical symptoms of Lyme Borreliosis



#### Lecture

#### for the Yuncheng district, Shanxi

November 2016

### What is Lyme Disease or Lyme Borreliosis

Lyme disease (LD) is a multisystemic disease **caused by Borrelia burgdorferi s.l**., a bacteria of the spirochete species.

It is called after the Swiss entomologist Dr. Burgdorfer, who detected it **1981** in the midgut of the tick **Ixodes scapularis** Since then, at least 19 species of Borr.burgd.s.l. have been described worldwide, 3 of them have been found so far in China: B.garinii, B.afzelii and B. valaisiana.

Qin Hao et al.: Distribution of Borr.burgd.s.l. in China in: J Clin Microbiol 2011 Feb;49(2) 647-50

## First Case of Lyme disease in Shanxi province 2014

- Chinese Journal of Vector Biology and Control Nov. 4, 2016 Oct. 23, 2016
- <u>Chinese Journal of Vector Biology and</u>
  <u>Control</u> 2014, <u>Vol. 25</u> Issue (4): 318-319:The first case of Lyme disease in Shanxi province, China: a case study by GENG Zhen<sup>1</sup>, Li Guohua<sup>2</sup>, HOU Xue-xia<sup>1</sup>, ZHANG Lin<sup>1</sup>, HAO

## **Content of this article**

Abstract Objective: To provide a scientific basis for confirmation of typical Lyme disease by serological analysis of the first suspected case of Lyme disease in Shanxi province, China and to improve the clinicians'recognition of Lyme disease for prompt diagnosis and treatment.

**Conclusion:** Based on the clinical history and progression of disease, as well as the serological results of this research, the diagnosis of Lyme disease can be confirmed, indicating the **existence of typical Lyme disease in Shanxi province**.

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### Does Lyme Disease or Lyme Borreliosis exist in other Chinese provinces too?

- The first case of Lyme disease (LD) in China was reported in Hailin county, Heilongjiang province **1987**.
- Thereafter, LD has been documented in more than 20 provinces and autonomous regions.
- Genotyping studies for Borr.burgdorferi s.l. have since then been conducted.
- Serological studies for Immunoglobulin G (IgG) antibodies against Borr. burgd. have been done like in Zhejiang province and Zhuolu county.

### Tick-borne diseases (TBD) are on the rise!

- In 30,000 randomly sampled people from 20 different P/A/M of China, the serological positivity of LD was 1,06 % -12,8%. The mean positivity rate was overall 5,06 % and was 5.33 % in forest areas. In Shaanxi province even 7,7%-8,8%.
- The mean <u>morbidity</u> was 2,84 % in the forests of Northeastern China.
- In Shanxi province the seroprevalence of B. burgd. was 5.54 % (from a total of 397 samples)
- In recent years, an increasing incidence of LD has been reported all over the world, which causes significant harm to livestock and human health and lifes(in USA 300 000 cases/yr)

Xian-Bo Wu et al: Distribution of tick-borne diseases in China, Parasites & Vectors, 2013,6:119

## Where have Borr.burgd. s.l. been found in China so far?

- Human cases of Lyme Borreliosis have been confirmed in 29 provinces/municipalities (P/M). Natural foci are present in at least 19 P/M in China. 3 new natural epidemic focuses exist in Shaanxi Province (Li Zhi-qing et al.:Chin.J.Hyg.Insect.Equipm. 5/2010)
- In Heilongjiang, Liaoning, Inner Mongolia over 3 million people get tick bites annually. Ca. 30, 000 of those become infected. Ca.10 % of the new cases may **turn into chronic infections** over the next 2-17 yrs without treatment.
- Co-infections with other tick borne pathogens like Anaplasma phagocytophilum (HGA) are present. In Shanxi province 1,26% of ticks carry B.b. **and** HGA.

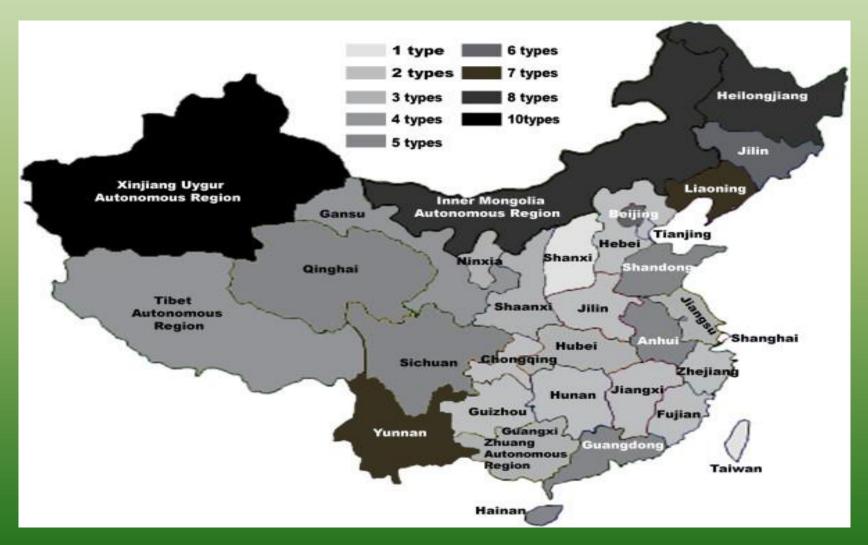
Xian-Bo Wu et al:Distribution of tick-borne diseases in China, Parasites & Vectors, 2013,6:119 ff Hao Qin et al: Seroepidemiol. Investigation of Lyme Disease...Biomed Environ Sci, 2013;26(3) 185 ff

### Distribution of Tick-Borne Diseases (TBD) across China

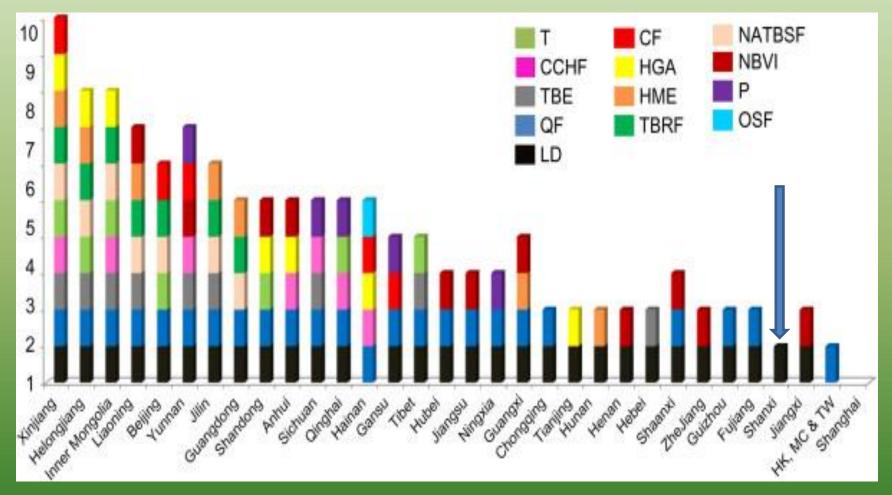


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### **The Prevalences of TB-Diseases in China**



### Types of tick-borne diseases in China`s P/A/M



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# Ixodes persulcatus, the carrier of Borrelia burgdorferi s.l. in China



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## What are the differences of Lyme disease in Europe and China

- The carrier of the spirochete Borr. burgd. s.l. in China is Ixodes persulcatus, in Europe Ixodes ricinus and in USA Ixodes scapularis.
- The following slides deal with the symptoms and the therapy of an infection with Borr. burgdorferi s.l.. The symptoms of LD are in China nearly the same as in Europe, but the carrier (Ixodes persulcatus) is different. The ticks can be carried by different animals like mice, deer or(most often) by birds from one place to the next.

A European tick family (Ixodes ricinus) with its family members: Larva, nymph, adult female and adult male tick. All except the male may transmit Borrelia burgdorferi s.l.



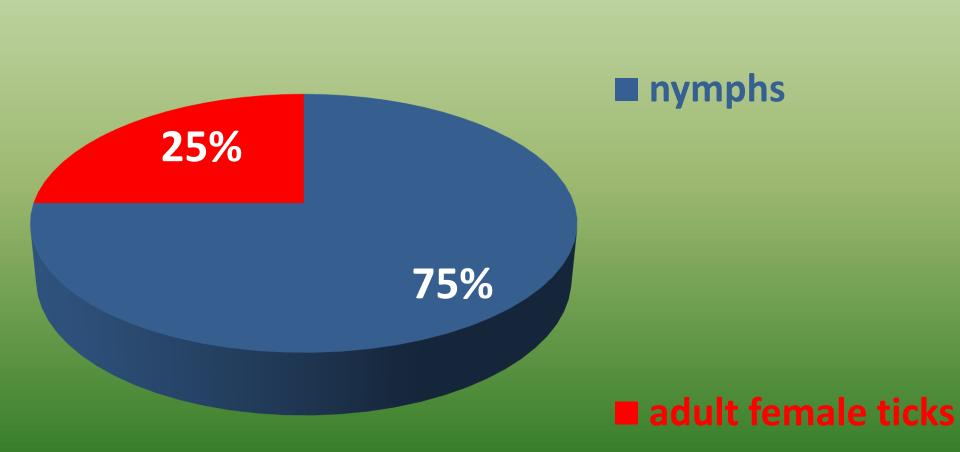
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**Foto: Polack** 

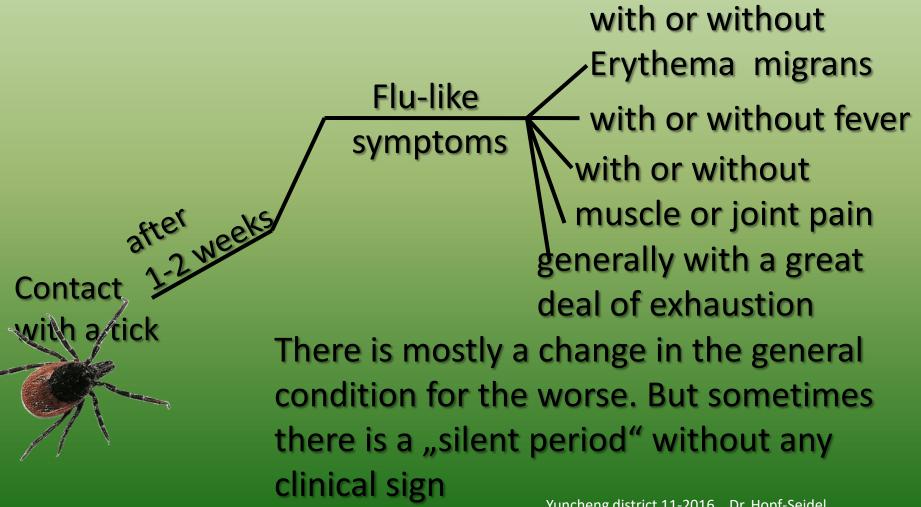
# The tiny nymph of a tick is the most dangerous transmitter of Borrelia s.l.



# Transmission rate from nymphs and adult female ticks to humans



### Symptoms in early stage of infection with **Borrelia burgdorferi s.l.**



Adult female tick firmly attached to a human host with a beginning Erythema migrans (EM)

Foto: Polack

### Typical bull's eye rash or Erythema migrans (EM)



If this early sign arises after a tick bite, **immediate** antibiotic treatment is necessary.

But keep in mind: Only 40-50 % of all infected persons develop an EM

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### Typical bull's eye rash (Erythema migrans)



## ...but 13 days after the tick bite, the same rash can easily be overlooked



### A rash (EM) 10 days after a tick bite`s infection with Borrelia s.l.



# An Erythema migrans can be extremely expanded, is painfree and therefore, can easily be overseen



©Foto : Dr. Hopf-Seidel

#### Erythema migrans (EM), vaguely edged out



### A lymphocytoma, the typical early sign of LD mostly in children



It can develop at all areas with soft tissue like at the ear lobes, the nipples, the cheek and the scrotum

# Lymphocytoma at the cheek, a sure sign of infection with Borrelia burgdorferi s.l.



### **Clinical symptoms of Lyme Disease/Borreliosis**

**Exhaustive tiredness** 

Mental and/or cognitive disturbances

Intense fatigue

Multilocular migrating joint aches and joint swellings

Frequent infections with viruses and bacteria

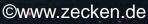
#### Sleep disturbances

#### **Frequent symptoms of (chronic) Lyme Disease**

- Headaches, shoulder-/neck-pain, jumping aches and swelling of joints, tendinitis and rupture of tendons without adequate trauma, myalgia
- Blurred vision, tinnitus, hearing loss
- Brain "fog", short memory loss, impairment of concentration and learning, mood swings
- Hormonal changes for thyroid and sexual organs
- Sleep disturbances and exhaustion, fatigue

# What happens after an infection with Borrelia s.l.

Borrelia are able to build persistent forms, the so-called round bodies (blebs, cysts, L-forms) and biofilms. They stimulate Th 1cytokines like TNF-alpha, IFN gamma, IL-1 beta with the consequence of a chronic persistent infection. <sup>11-2016</sup> Dr. Hopf-Seidel The spirochetes divide themselves every 12-24 hrs starting immediately after the host`s infection.They rebuild their cell wall continuosly. The lipopolysaccharides of the cell wall have an antigen effect and therefore the host`s immunsystem forms antibodies (IgM, IgG).





# The formation of cystes of Borrelia burgdorferi s.l. in detail



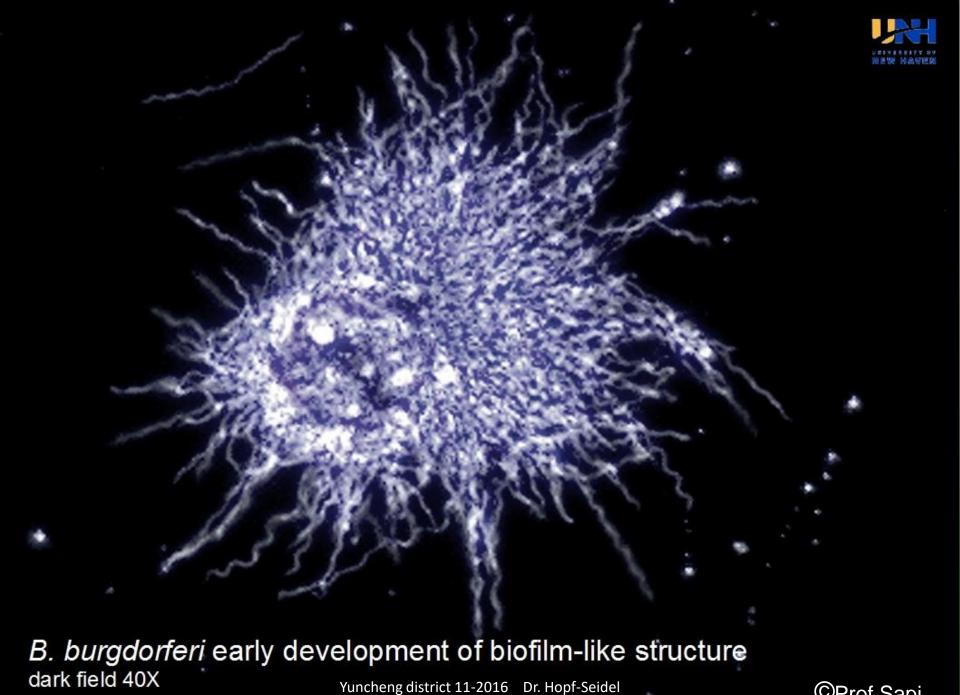
# The different stages of viable Borrelia s.l. demand adequate treatment regimes

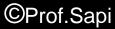
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All the <u>intra</u>cellular persister forms can be treated with Tetracyclines (best Minocyclin), or with Makrolides (Clarithromycin, Azithromycin), best in combination with Tinidazol (Fasigyn<sup>®</sup>) or Hydroxychloroquin

As long as the spirochetes divide themselves, they can be treated with cell wall synthese inhibitors like **Cephalosporines** (Amoxicillin<sup>®</sup>, Cefuroxim<sup>®</sup>) or Betalactames (Ceftriaxon<sup>®</sup>, Cefotaxim<sup>®</sup>) or Tetracyclines (Minocyclin or Doxycyclin)







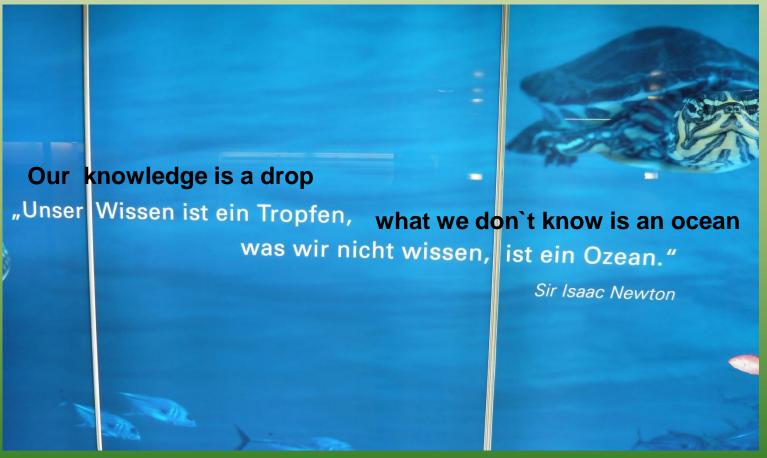
### **Treatment of early stage of Lyme Disease**

- **Amoxicillin** 3 x 1000 mg for 30 days (especially for pregnant women, for children according to weight)
- Clarithromycin 2 x 500 mg (starting with half the dose over a period of 4 days ) for 30 days for adults, weight adapted for children (7,5 mg/kg body weight bid).
- Minocyclin with 2 x 100 mg for patients of 50 -70 kg bodyweight (always start with only 50 mg/day and then increase slowly 50mg more every 3 days to prevent the possible side effects headaches and vertigo)

### **Treatment of late stage of Lyme Disease**

- Azithromycin 500 mg- 600 mg/day for 4 days, then 3 days off because of the intracellular accumulation of the drug (generally given for 6 cycles)
- Doxycyclin/Minocyclin should always be given in combination with Hydroxychloroquine (Quensyl<sup>®</sup>, Plaquenil<sup>®</sup>), **Tinidazol** (Fasigyn<sup>®</sup>) or **Rifampicin** (Eremfat<sup>®</sup>, Rimpacin<sup>®</sup>) to prevent the formation of round bodies and **biofilms**. This combined therapy helps to reduce the number of Borrelia s.l. in the body to prevent further clinical relapses.
- Biofilms are the most difficult forms of Borr. burgd. to be treated

# This applies to our present knowledge of Borreliosis as well.....



## More information you may find online: <u>www.dr-hopf-seidel.de</u> or in my book





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## Thank you for your attention