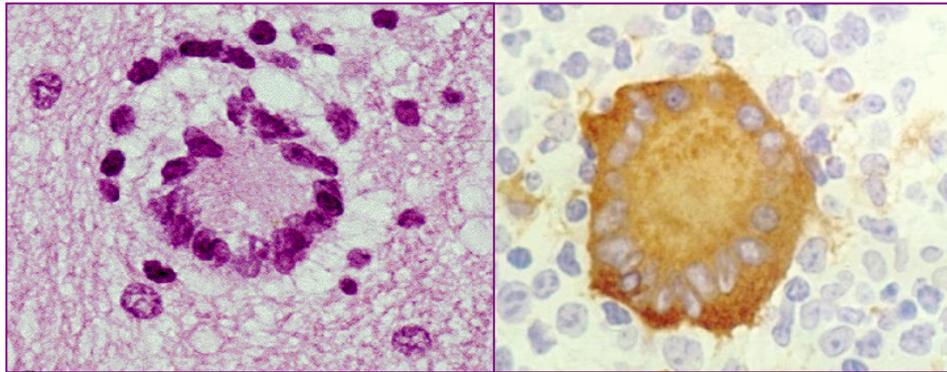


Carrefour Pathologie 2017

Quelle place et quel avenir pour la pathologie infectieuse en ACP?



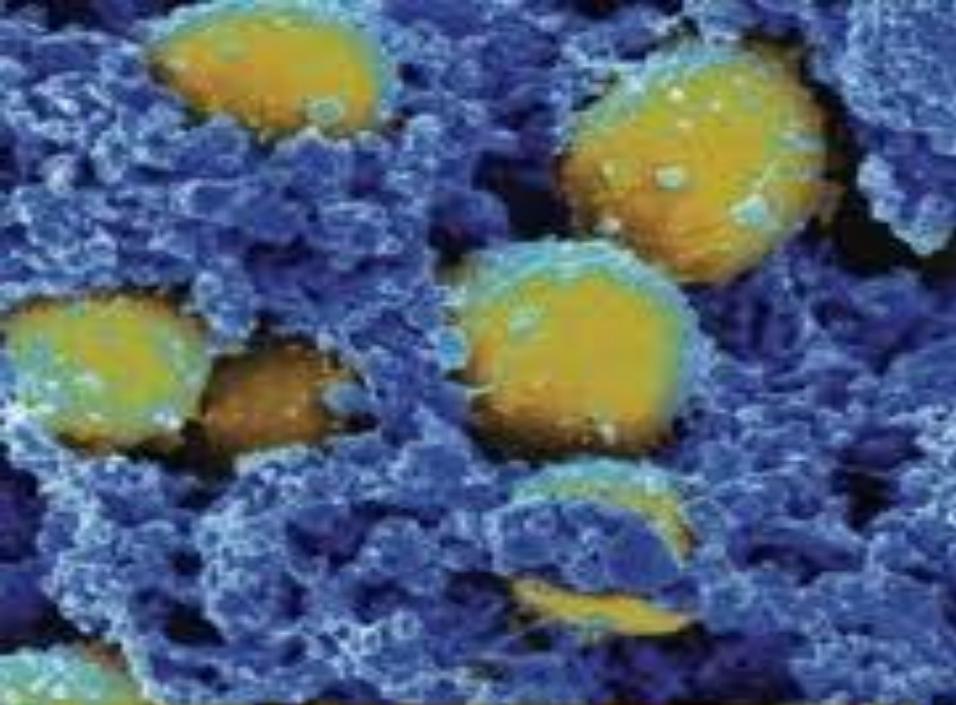
Paul Hofman

Laboratory of Clinical and Experimental Pathology, Louis Pasteur Hospital,
University Côte d'Azur, Nice

« Infectious diseases are our faithful companions throughout life »



Charles Nicolle (Nobel prize in Medicine, 1928), Pasteur Institut, Paris



Charles Nicolle

Destin des maladies infectieuses

essai



« Infectious diseases are our faithful companions throughout life »



**Ebola river
(Democratic Republic of Congo)**



Ebola virus



Kindly from Pr N Pasaka, Kinsasha, RDC



The Nobel Prize in Physiology or Medicine 2015



Ill. N. Elmehed. © Nobel
Media AB 2015.

William C. Campbell

Prize share: 1/4



Ill. N. Elmehed. © Nobel
Media AB 2015.

Satoshi Ōmura

Prize share: 1/4

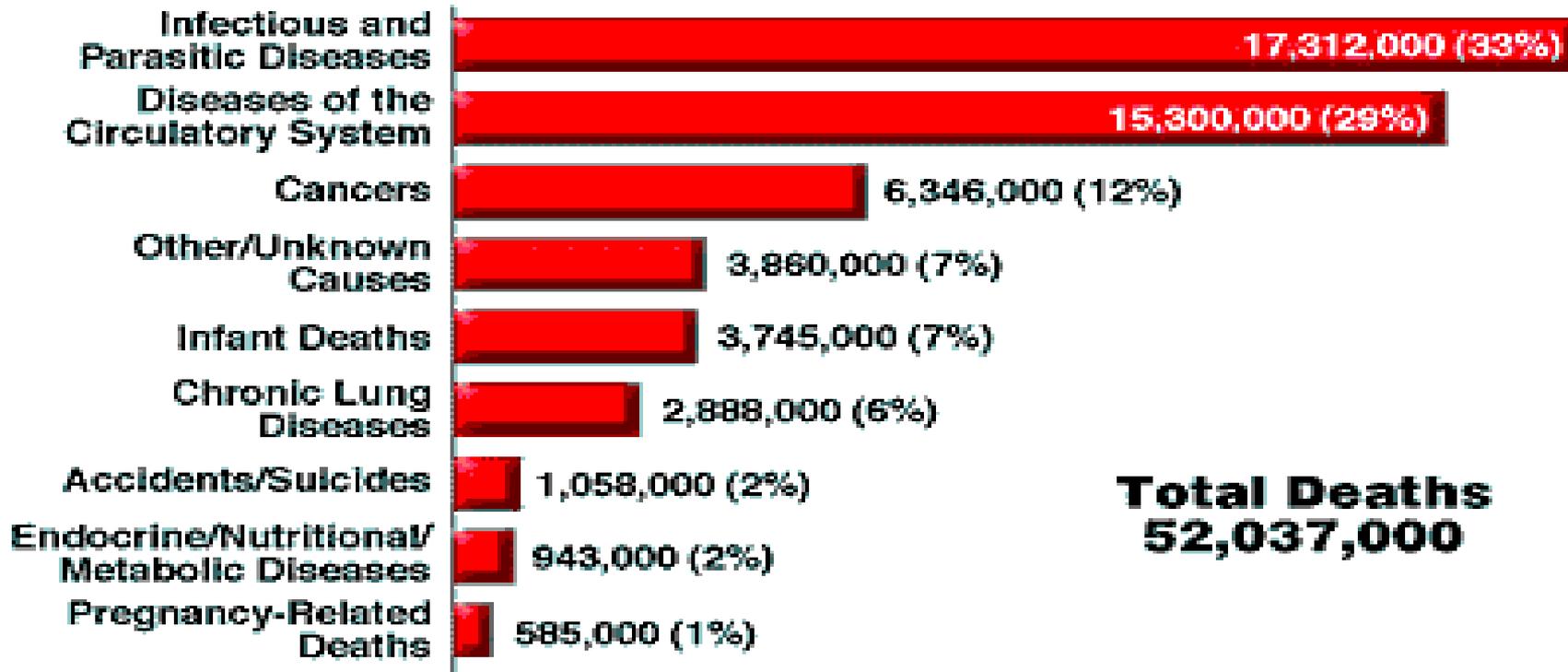


Ill. N. Elmehed. © Nobel
Media AB 2015.

Youyou Tu

Prize share: 1/2

Worldwide Causes of Death



Which input of infectious diseases
in a daily practice for a pathology laboratory?

**Different scenarii for different challenges for a
surgical pathologist!**

The first scenario.....

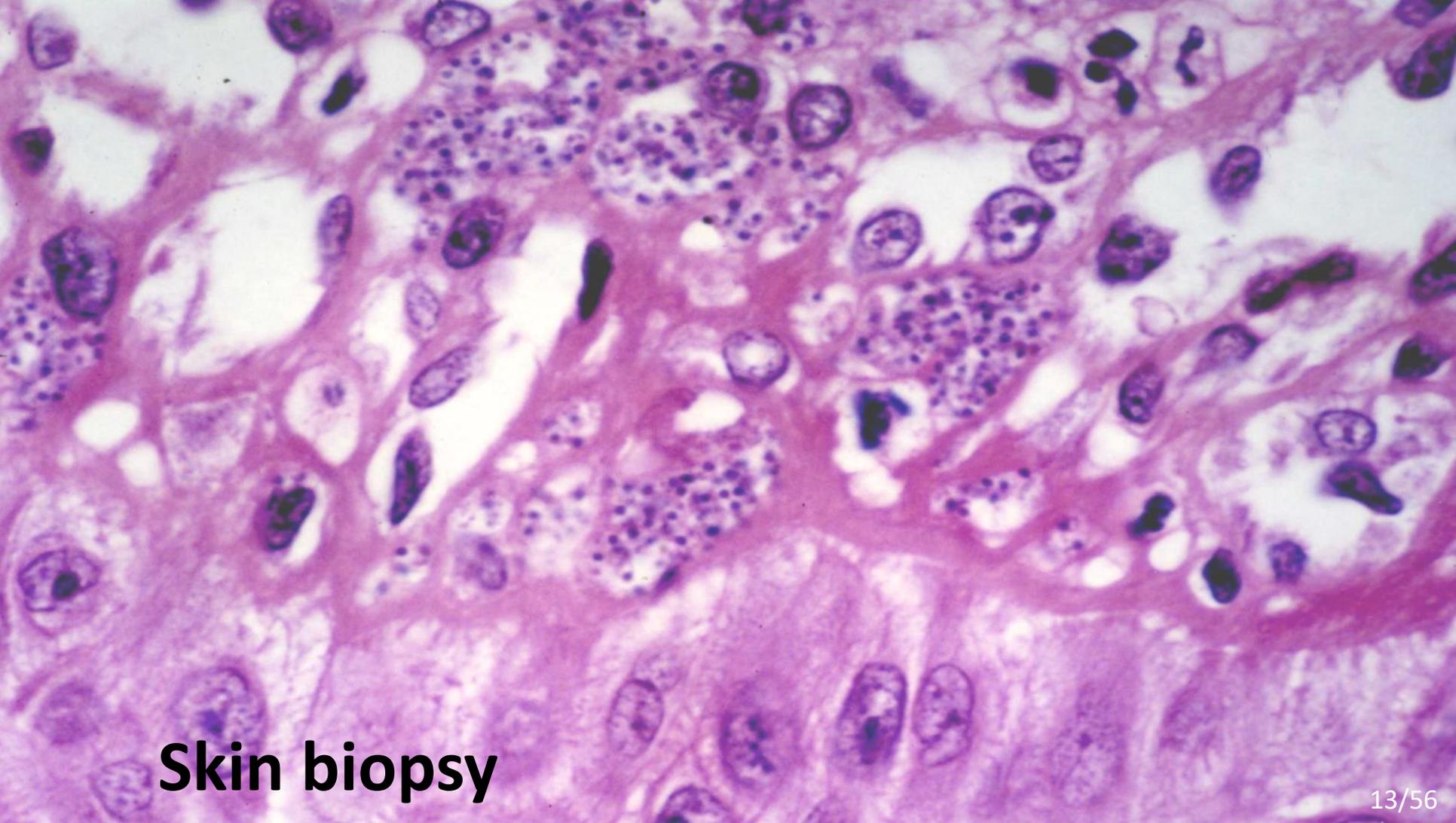
An easy diagnosis for a surgical (clinical) pathologist working in any countries....





Ten years old girl
Living in Nice
Ulcerated skin lesion of the cheek
Three months duration





Skin biopsy

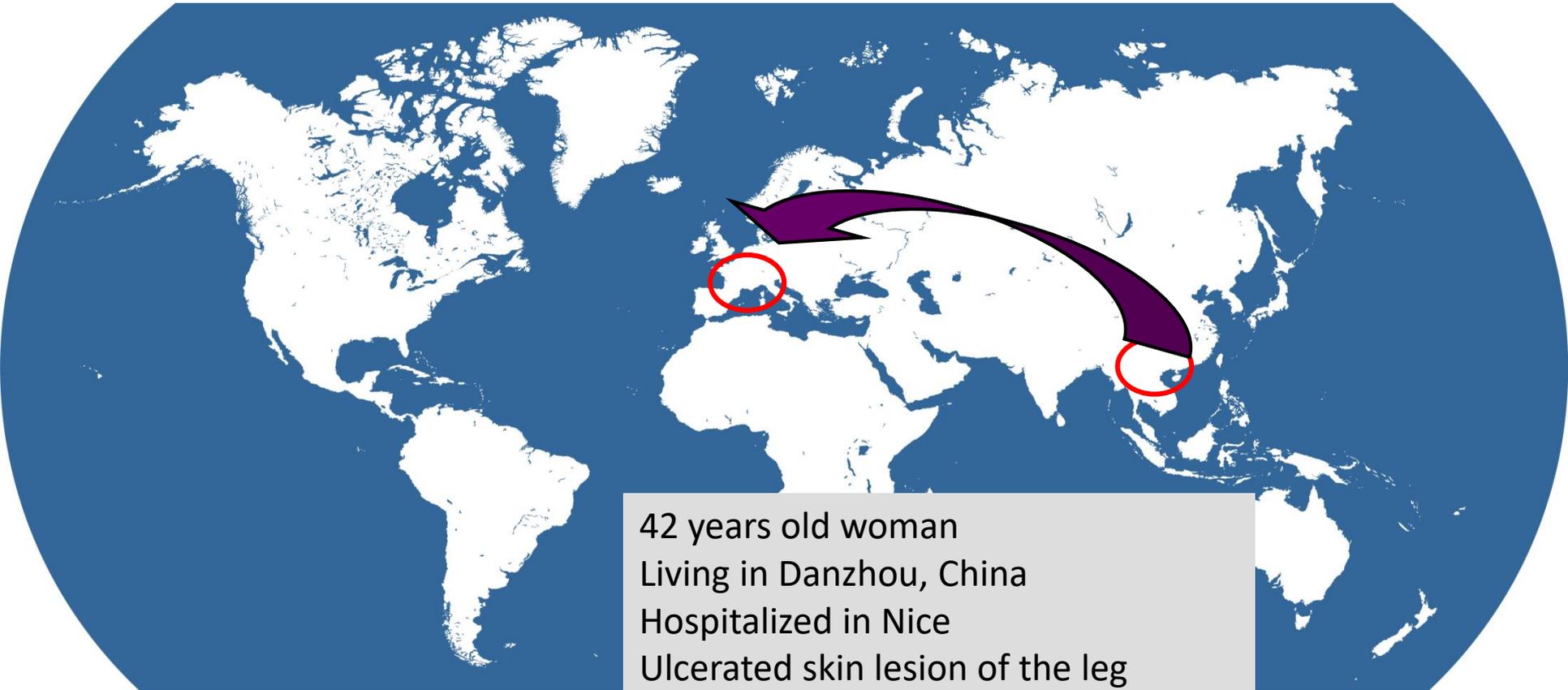
What is your diagnosis ?

Cutaneous leishmaniasis
(caused by *Leishmania infantum*)

Another scenario.....

A more complex diagnosis for a surgical (clinical) pathologist working in Europe....





42 years old woman
Living in Danzhou, China
Hospitalized in Nice
Ulcerated skin lesion of the leg
12 months duration



Skin biopsy



What is your diagnosis ?

Cutaneous sparganosis
(caused by *Sparganum proliferum*)

Asiatic food market



A very rare parasitic disease (cestode) of the skin 22/56

Another scenario.....

**A more complex diagnosis for a surgical pathologist working in Europe....
Is it a difficult diagnosis... or ...not ?**

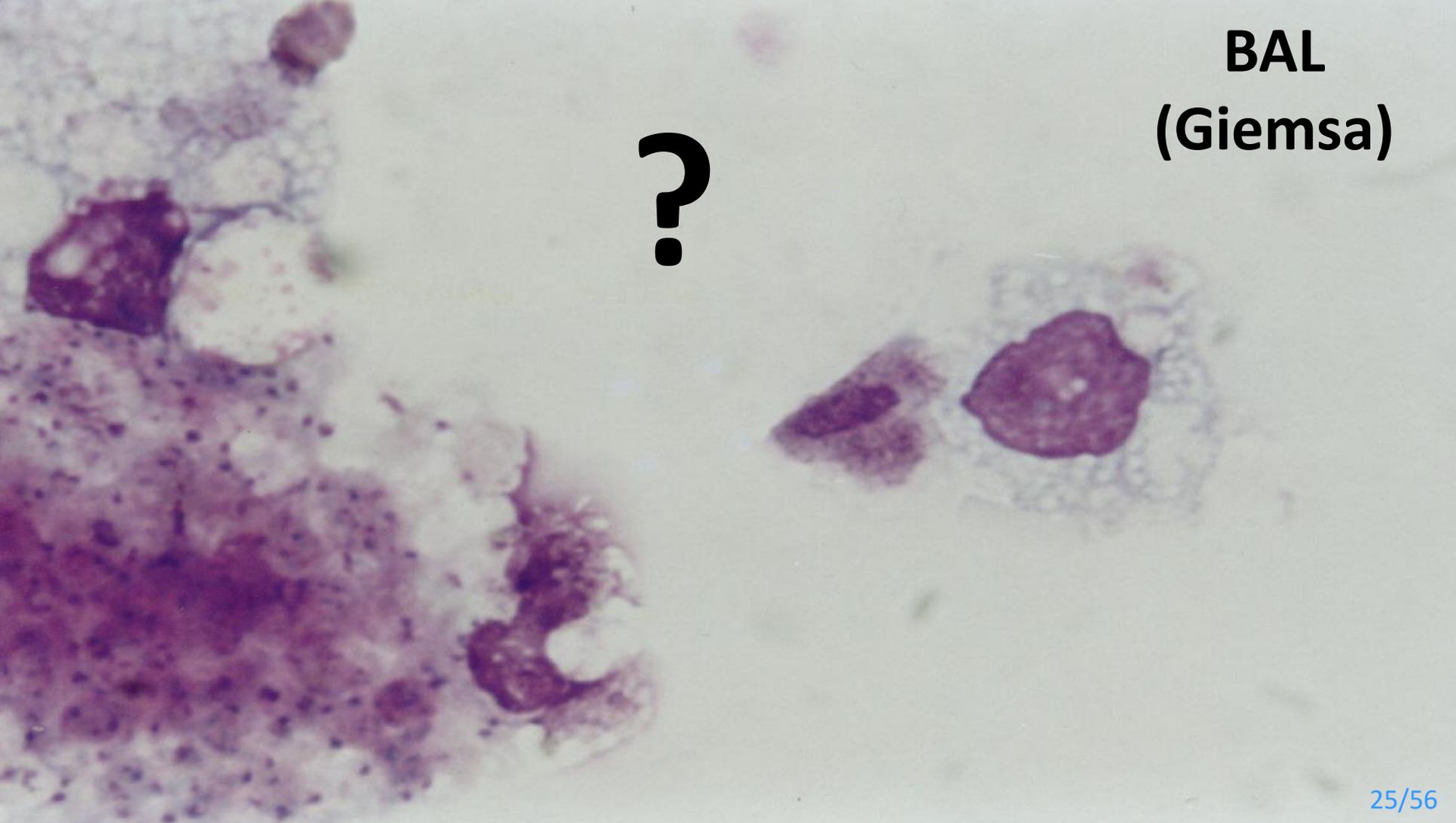




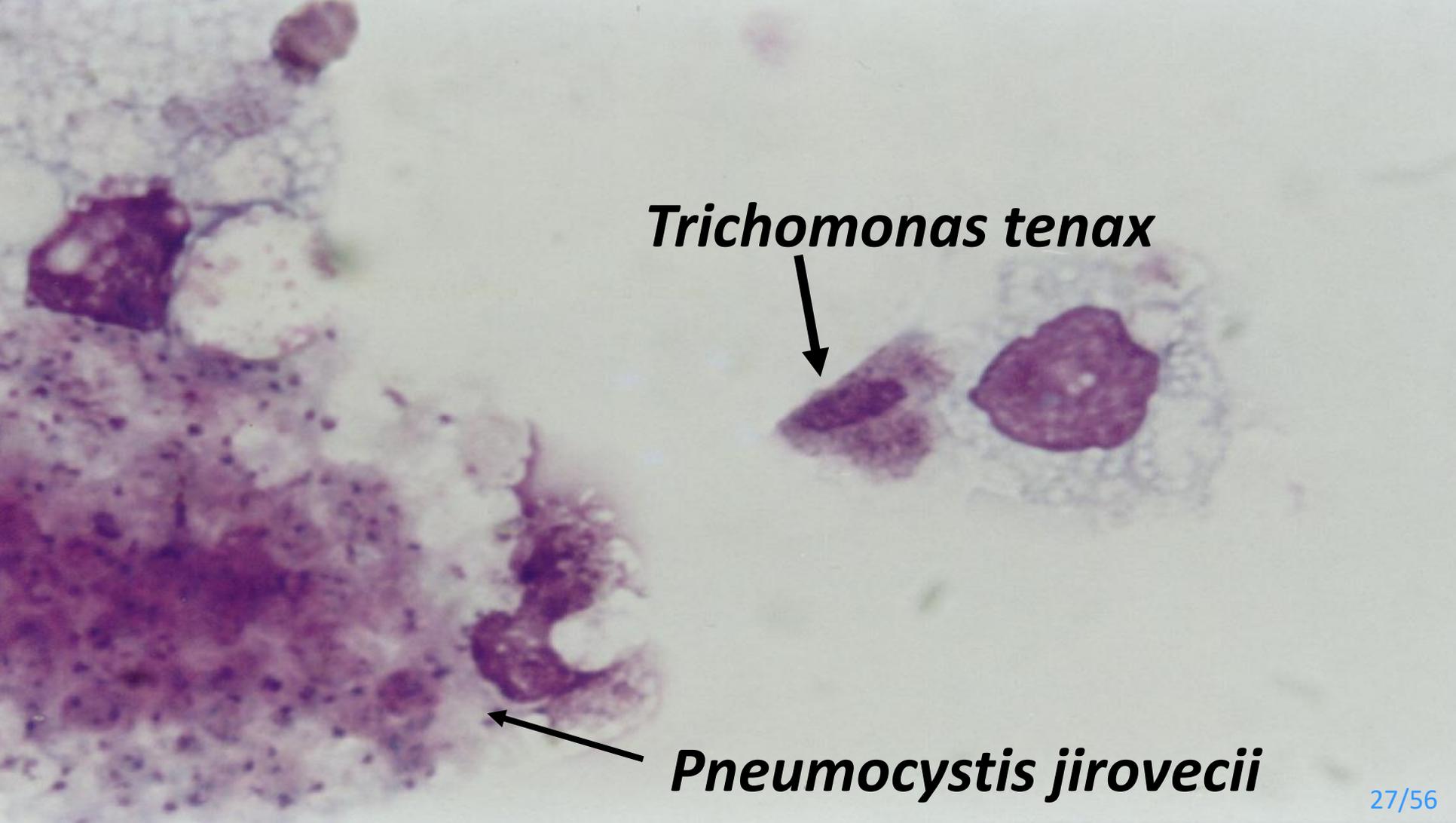
36 years old man, HIV +
Living in Nice,
Fever, polypnea

BAL
(Giemsa)

?



What is your diagnosis ?



Trichomonas tenax

Pneumocystis jirovecii

Opportunities in infectious diseases pathology practice



Increase in **opportunistic infections** following the increasing transplantation, chemotherapy and targeted therapy rates

Increase in **imported parasitic and tropical diseases** in travellers and in immigrants

Emerging complications of HIV associated diseases caused by HAART

Epidemics of new and emerging or re **emerging infectious diseases**

Optimisation of **new ancillary methods** to detect pathogens in fixed samples

Opportunity to observe **tissue lesions associated with the agent** synonymous to pathogenicity and infection!



Total: 2.5 – 3.5 million

AIDS Deaths

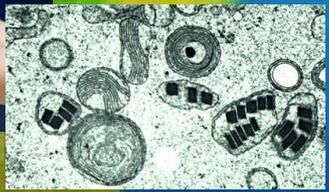


HAART and HIV

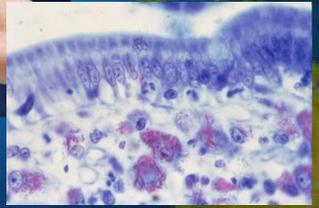
A large panel of (new) pathological induced lesions



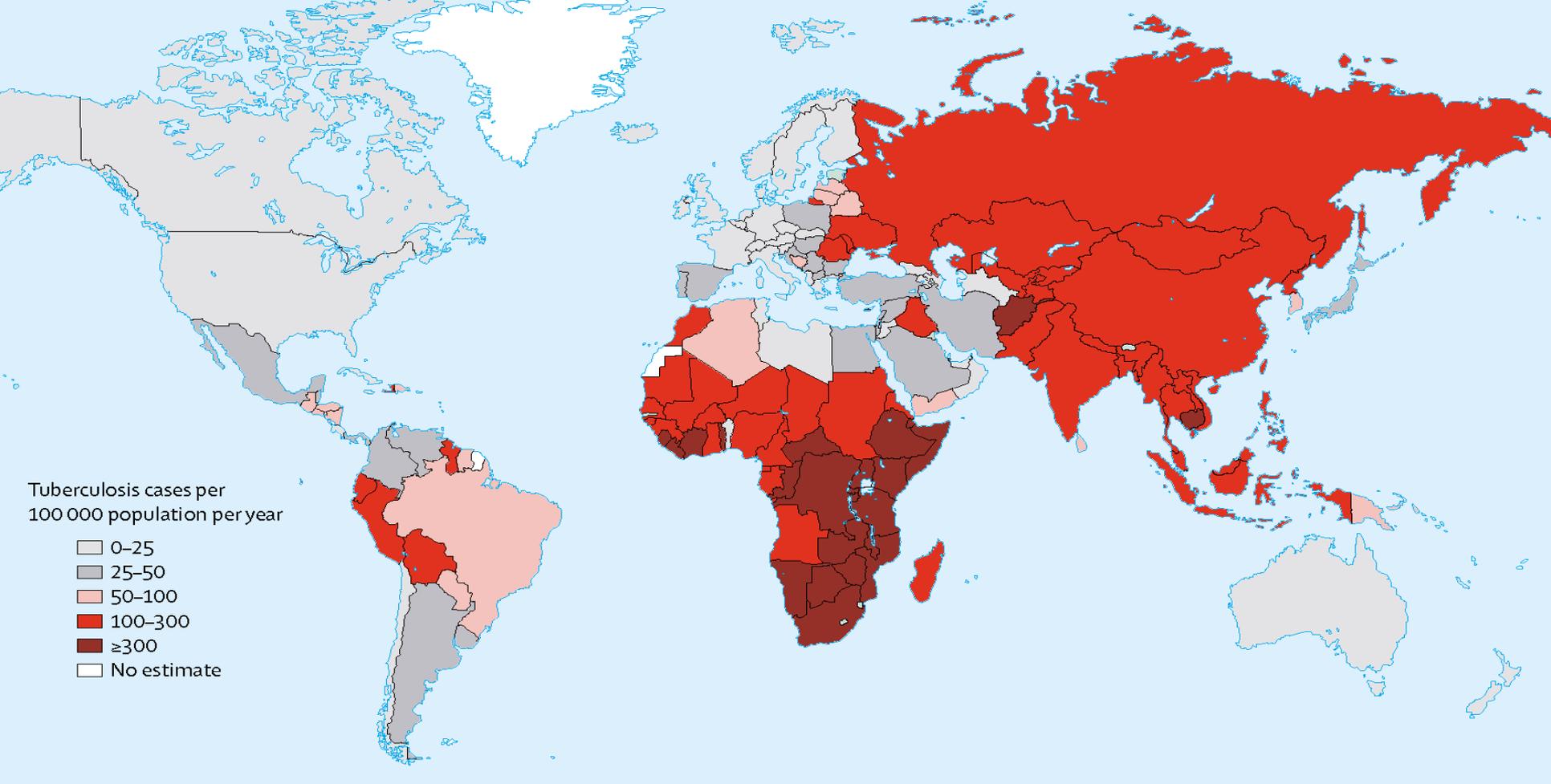
Dystrophies



Toxicities



IRIS



Tuberculosis: 1, 500.000 deaths per year

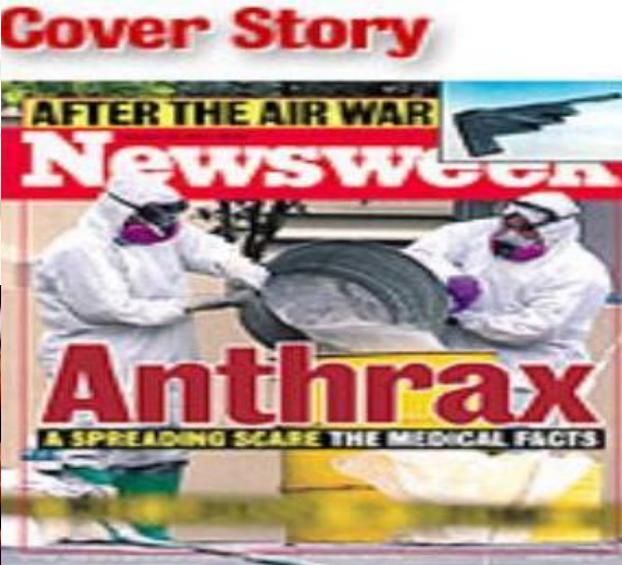
« Infectious diseases are our faithful companions throughout life »

An aerial photograph of a large iceberg in the ocean. The visible tip of the iceberg is small and white, while the vast, submerged portion is a deep blue, illustrating the concept that infectious diseases are often hidden or underestimated. A blue oval with a black border is superimposed over the submerged part of the iceberg, containing the text.

Emerging and re emerging infectious diseases

Emerging infectious diseases (1991-1993)



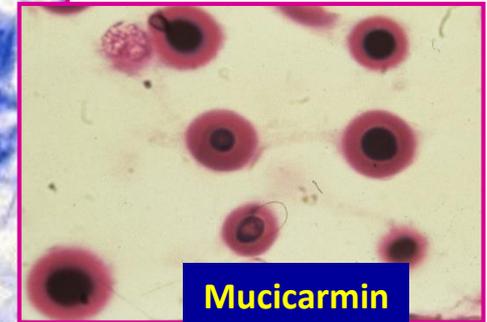
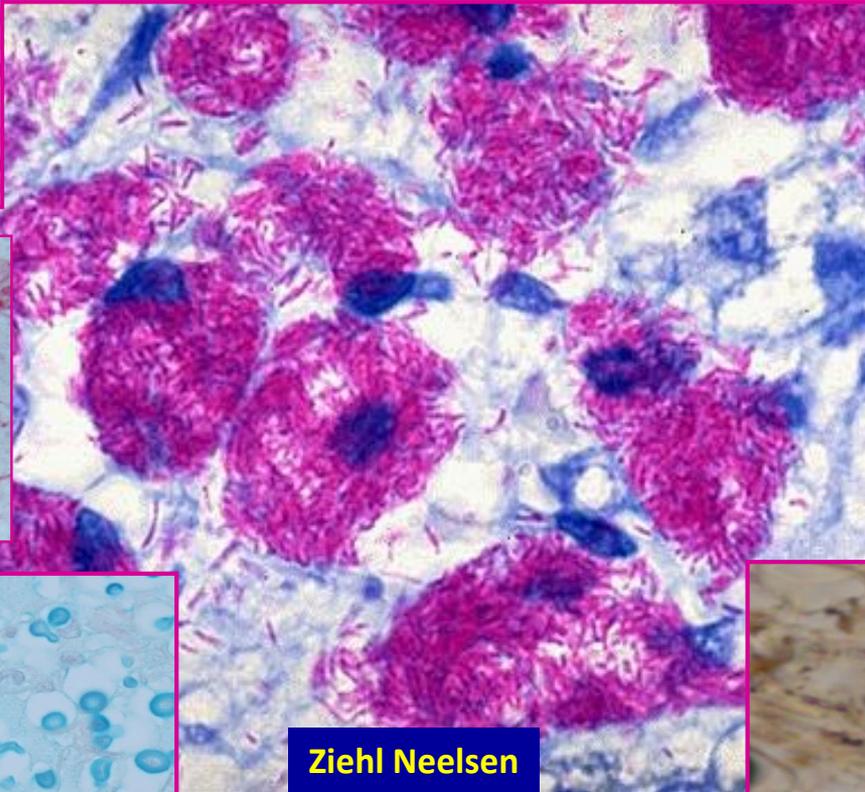
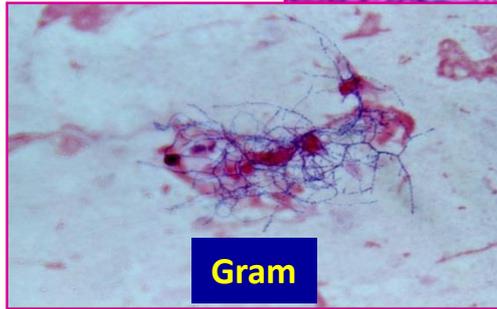
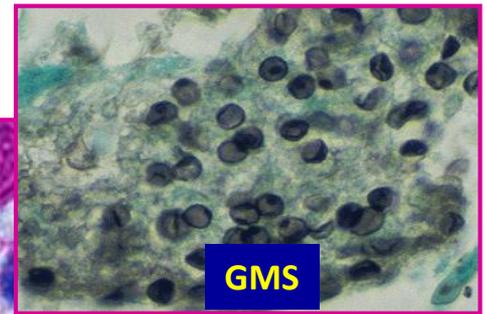
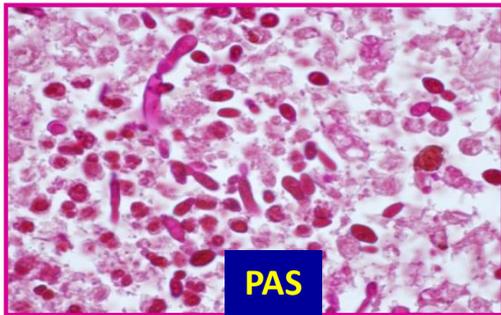


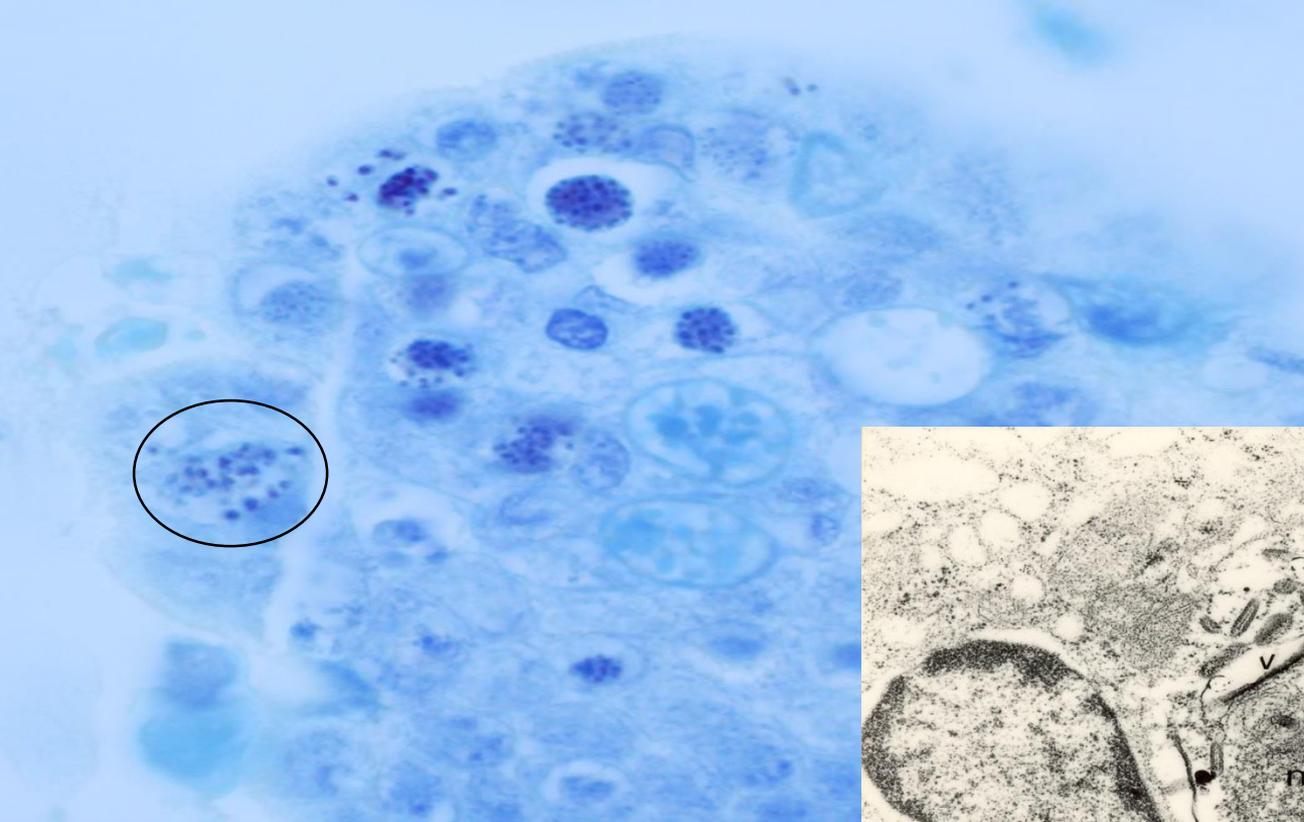
18 Feb 2017 – “Bioterrorists could one day kill hundreds of millions of people in an attack more deadly than nuclear war”



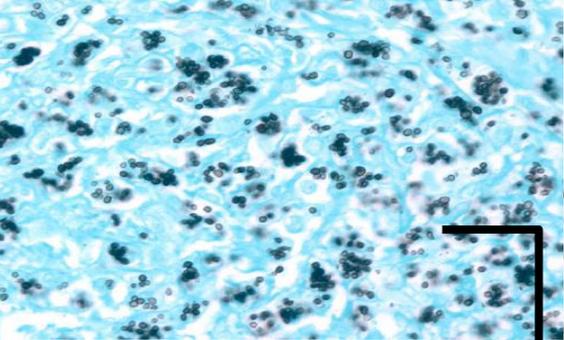
Bill Gates will warn world leaders !

Which input for the ancillary methods (old and current) in infectious diseases pathology ?

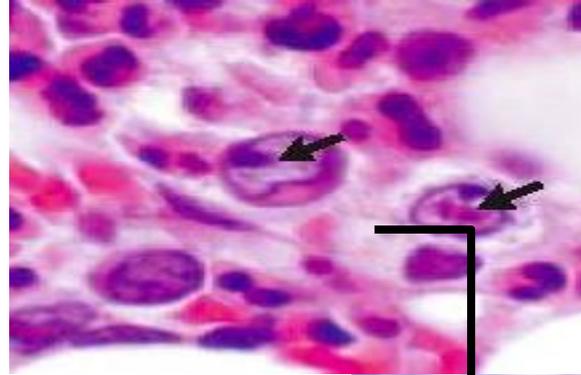
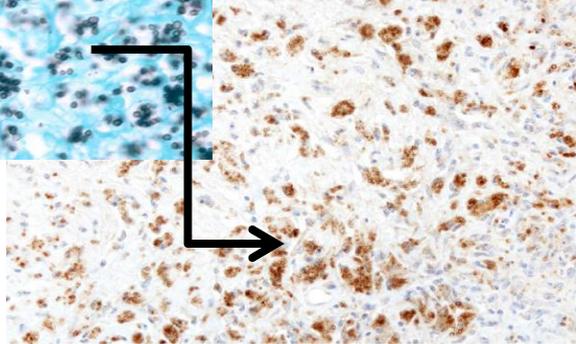




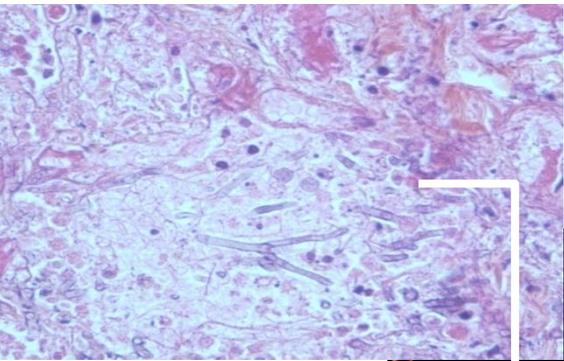
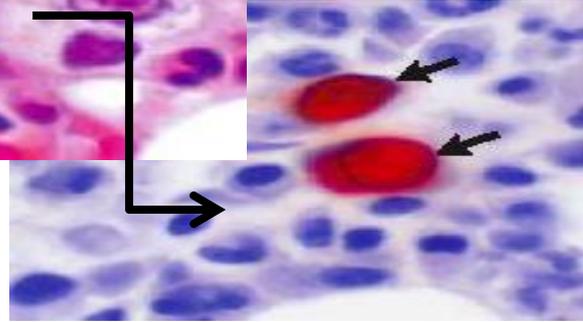
Microsporidiosis
(*Enterocytozoon bieneusii*)



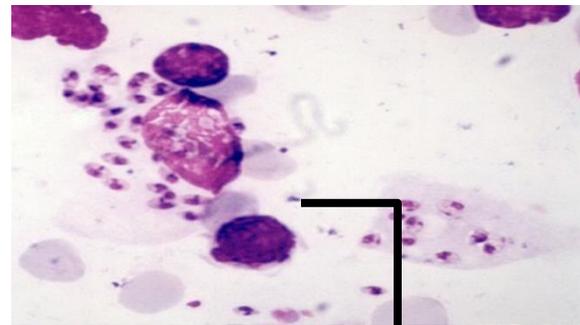
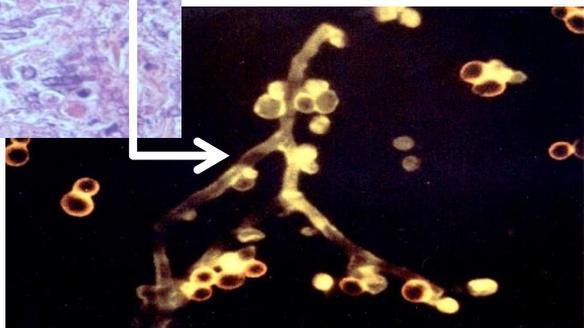
AC-Anti-Histoplasma



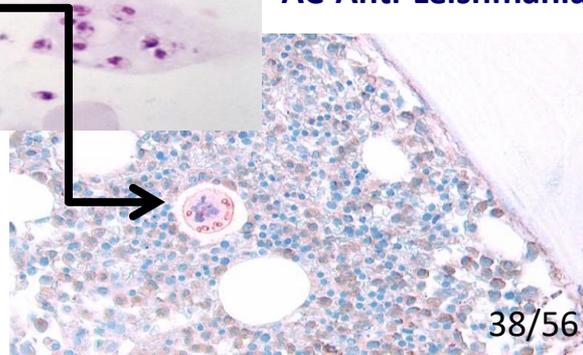
AC-Anti-Parvovirus



AC-Anti-Aspergillus



AC-Anti-Leishmania

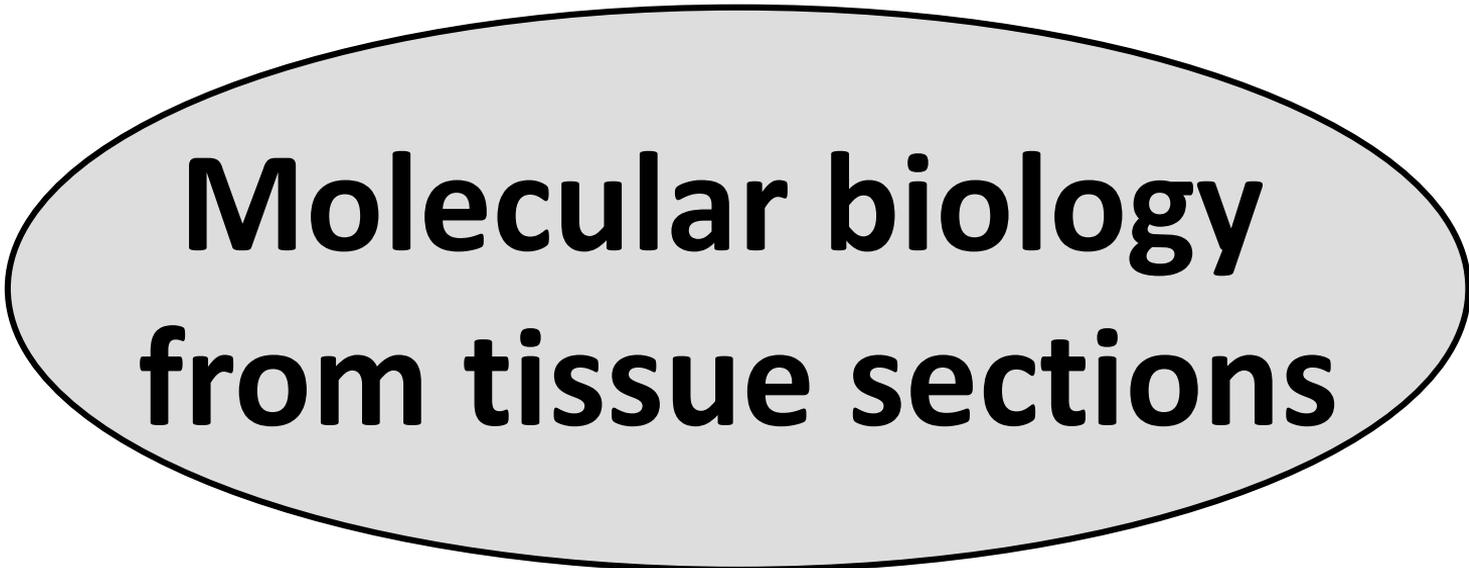


Paraffin sections

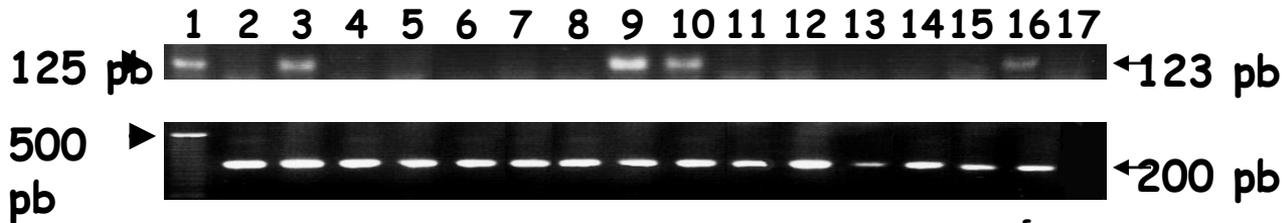
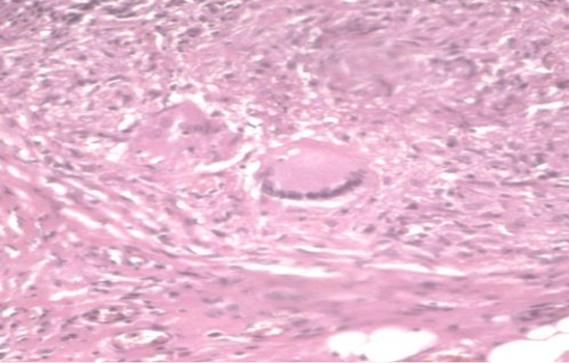
Escherichia coli

Streptococcus aureus

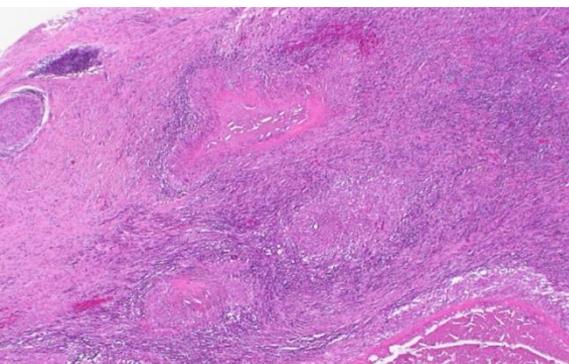
Salmonella typhimurium



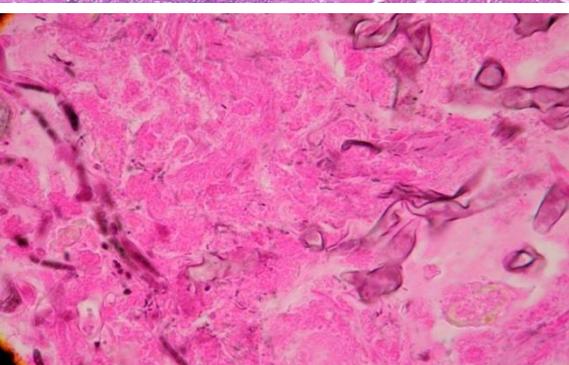
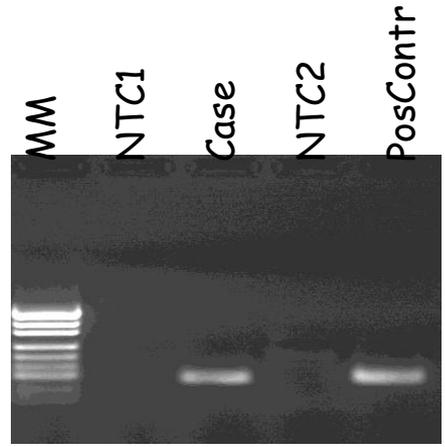
**Molecular biology
from tissue sections**



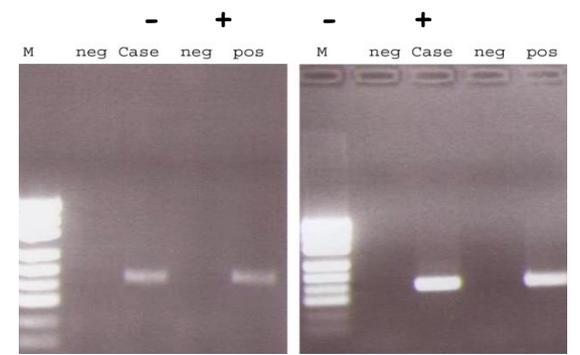
Mycobacterium tuberculosis



Brucella melitensis



Aspergillus sp. & Mucorales sp.



Opportunities in infectious diseases pathology practice

Increased opportunistic infections following the increasing transplantation, chemotherapy and targeted therapy rates

Increase in imported infectious and tropical diseases in returning travellers and in immigrants

Emerging complications of HIV associated diseases caused by HAART

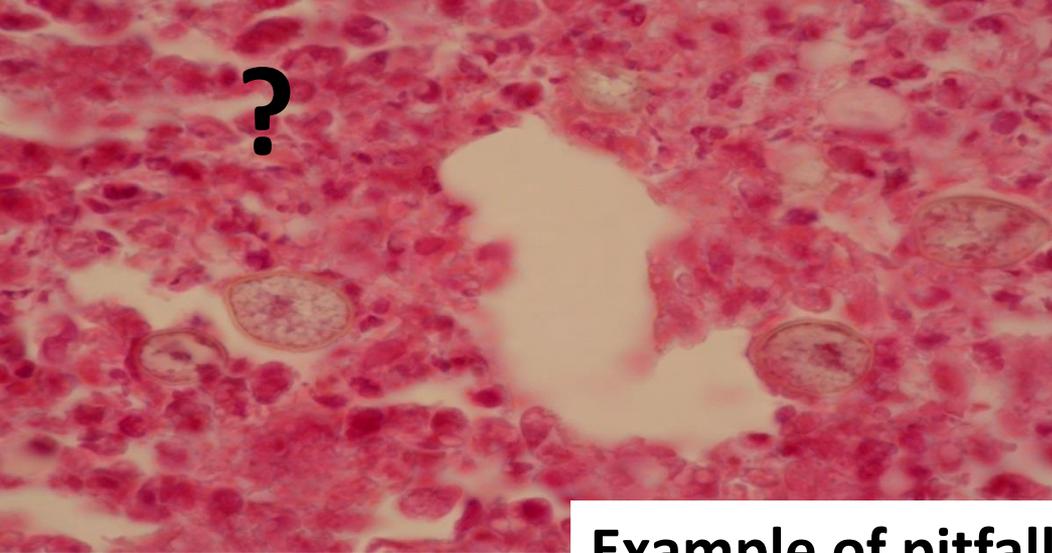
Epidemics of new and emerging or re emerging infectious diseases

Optimisation and development of new ancillary methods to detect pathogens in fixed samples

Opportunity to observe tissue lesions associated with the agent which is synonymous to pathogenicity and infection



Differential diagnosis: non pathogenic lesions mimicking an infection



Nasal mucosa

Example of pitfall in ID Pathology field

Nasal mucosa



Paracoccidioidomycosis



Nasal mucosa

Myospherulosis



Nasal mucosa

6 constraints in infectious diseases pathology activity



Weak ability to get a **full time job** for doing pathology infectious diseases diagnostic

Weak possibility to have **training in tropical diseases** pathology in tropical countries

Difficulty to maintain a **broad number of antibodies** targeted infectious agents

Low budget for the development of a **molecular pathology infectious** disease unit

Set up weekly **meetings with physicians** concerned by infectious diseases pathology

Sustainability of **autopsy activity** in institutions



3 potential orientation for tomorrow (?)

Increasing the number of supports and of **grants** in infectious diseases pathology (be attractive to the NGP*)

Developing **telepathology** approach for optimisation of the turn around time for **consultation diagnosis** made by experts in infectious and tropical diseases pathology

Set up or **sustaining expert centres** in infectious and tropical diseases pathology

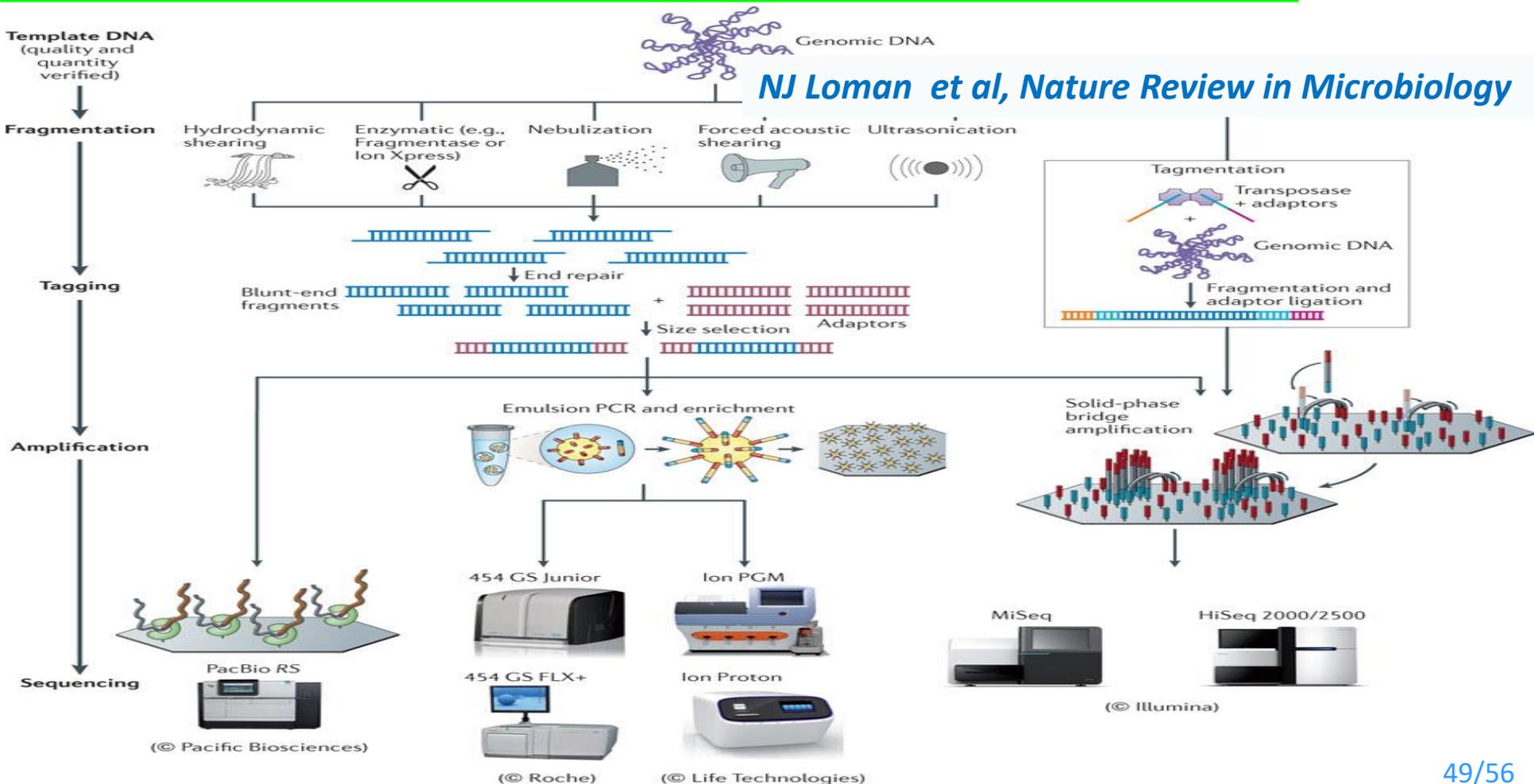
*Next Generation of Pathologists

Innovative technologies for the improvement of infectious diseases diagnostic from fixed tissue sections ?



MAIN HIGH-THROUGHPUT SEQUENCING PLATFORMS

NJ Loman et al, Nature Review in Microbiology



Spatial dynamics of HIV-1 spread From 1920 to 1960....!

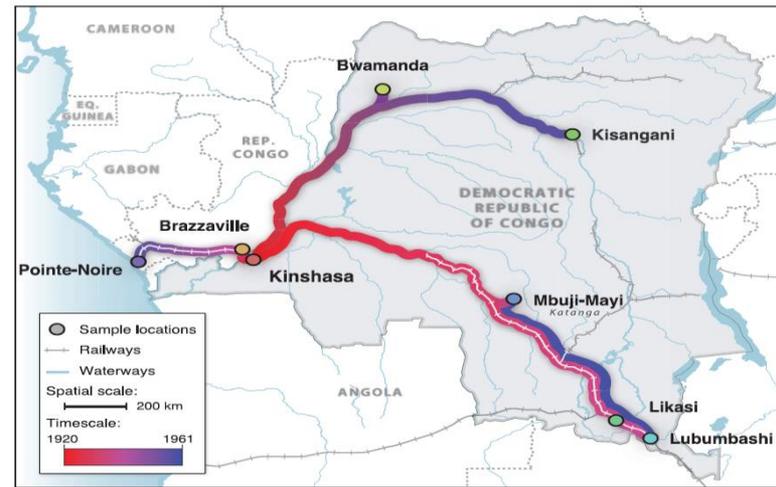
RESEARCH ARTICLE

HIV EPIDEMIOLOGY

The early spread and epidemic ignition of HIV-1 in human populations

Nuno R. Faria,^{1,2} Andrew Rambaut,^{3,4,5} Marc A. Suchard,^{6,7} Guy Baele,² Trevor Bedford,⁸ Melissa J. Ward,³ Andrew J. Tatem,^{4,9} João D. Sousa,^{2,10} Nimalan Arinaminpathy,¹ Jacques Pépin,¹¹ David Posada,¹² Martine Peeters,¹³ Oliver G. Pybus,^{1*†} Philippe Lemey^{2*†}

Thirty years after the discovery of HIV-1, the early transmission, dissemination, and establishment of the virus in human populations remain unclear. Using statistical approaches applied to HIV-1 sequence data from central Africa, we show that from the 1920s Kinshasa (in what is now the Democratic Republic of Congo) was the focus of early transmission and the source of pre-1960 pandemic viruses elsewhere. Location and dating estimates were validated using the earliest HIV-1 archival sample, also from Kinshasa. The epidemic histories of HIV-1 group M and nonpandemic group O were similar until ~1960, after which group M underwent an epidemiological transition and outpaced regional population growth. Our results reconstruct the early dynamics of HIV-1 and emphasize the role of social changes and transport networks in the establishment of this virus in human populations.



A careful analysis of the morphological features is still pivotal for identifying most pathogens in tissue sections

Asking for ancillary methods **only** if necessary

However.....

42 years old man, hunter in the South of France

Mediastinal lymph nodes

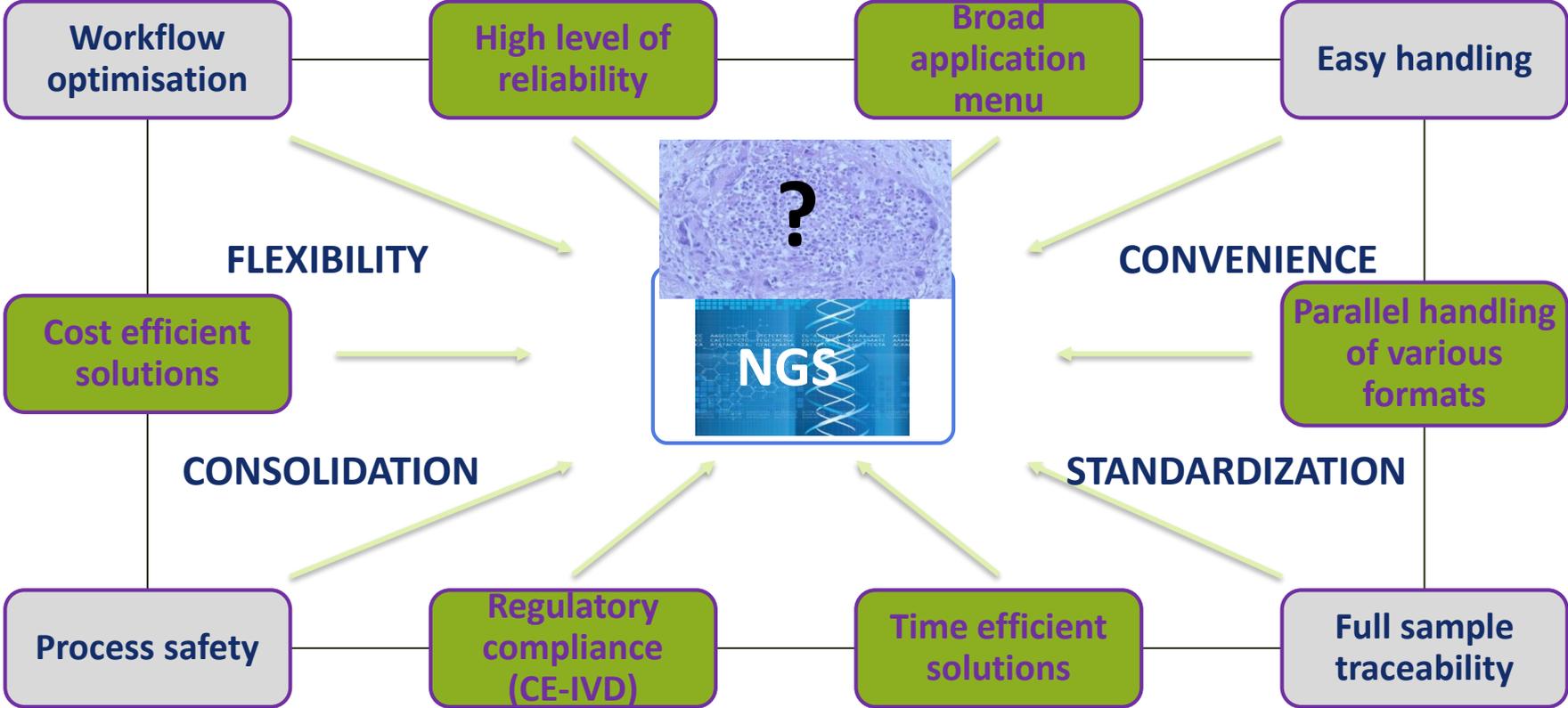
Fever

All ancillary methods negatives (including microbiology tests)

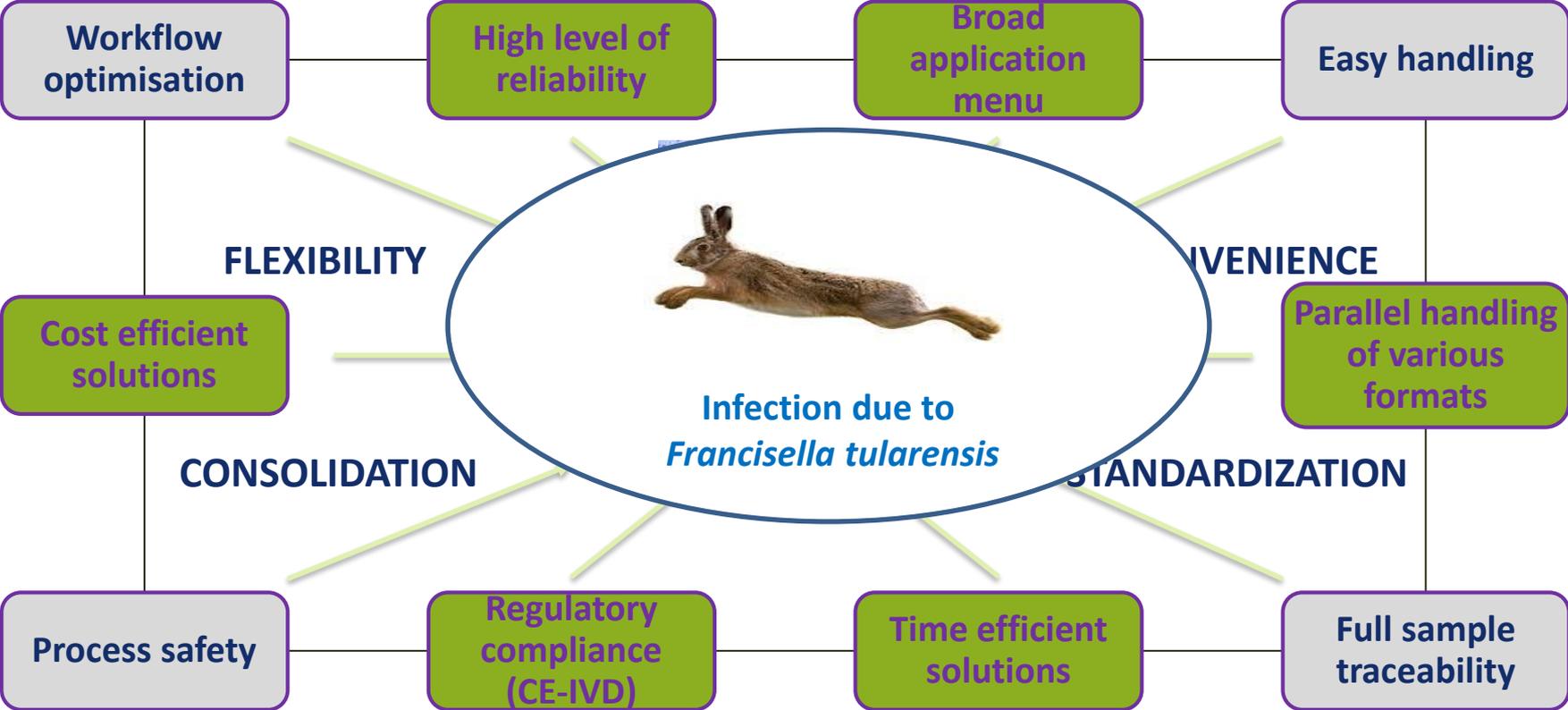
Uncertain diagnosis

?

Needs of molecular diagnosis for routine practice in Infectious diseases from fixed samples: A new challenge for the pathologists or..... for the microbiologists ?



Needs of molecular diagnosis for routine practice in Infectious diseases from fixed samples: a new challenge for the pathologists or for the microbiologists ?

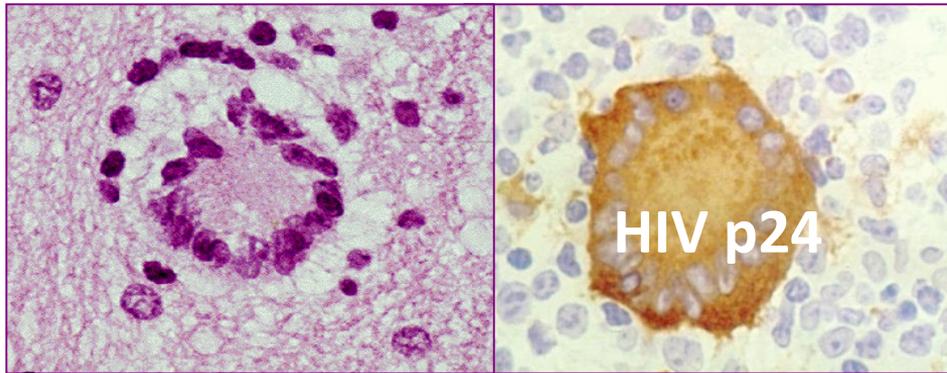


Future role for the pathologist in Infectious Diseases

- Diagnosis, prognosis and theranosis assessment of ID from tissue sections
(*similarly to the role of **the clinical pathologist** in oncology*)
- Optimisation of the molecular pathology approach of ID from tissue sections
(*similarly to the role of the **molecular pathologist** in oncology*)
- Does the molecular pathology in microbiology can be set up in a department of pathology in Europe ?
- Does challenging the microbiologist in molecular pathology still necessary ?

Carrefour Pathologie 2017

Quelle place et quel avenir de la pathologie infectieuse en ACP?



Paul Hofman

Laboratory of Clinical and Experimental Pathology, Louis Pasteur Hospital,
University Côte d'Azur, Nice