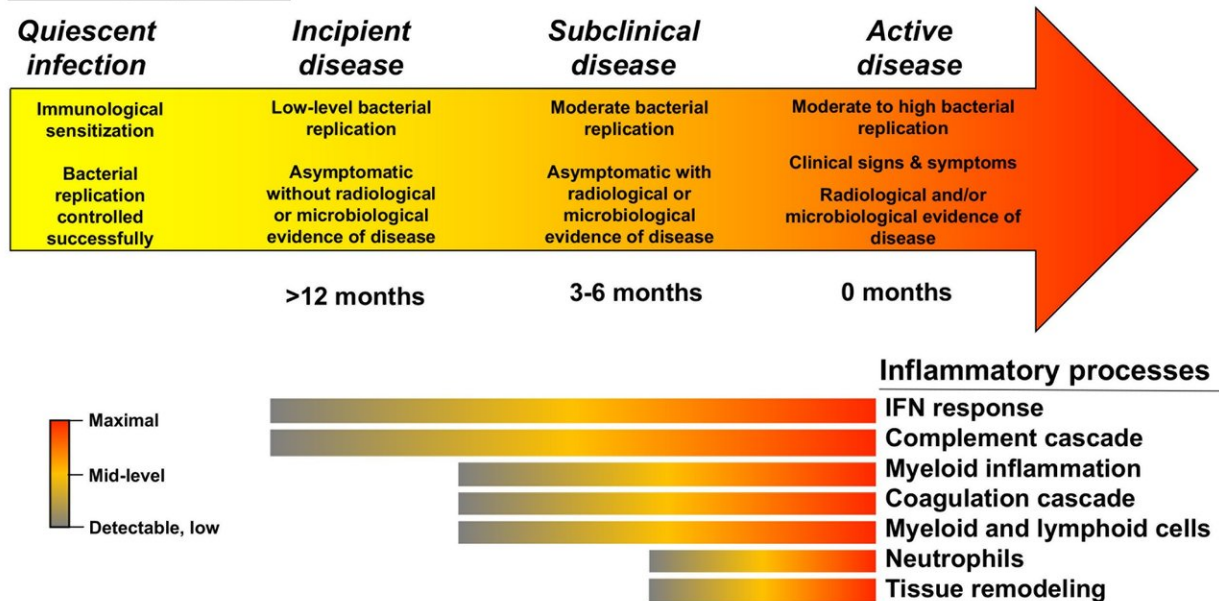


Progression from infection to pulmonary tuberculosis follows distinct timeline

November 16 2017

Disease progression



Progression from quiescent *Mycobacterium tuberculosis* infection, via incipient and subclinical disease, to active pulmonary tuberculosis disease in humans.
Credit: Scriba TJ, et al (2017)

Researchers have uncovered a sequence of biological processes that occur in humans infected with the bacterium *Mycobacterium tuberculosis* as the infection progresses to pulmonary tuberculosis, according to new research published in *PLOS Pathogens*.

M. tuberculosis infects about a quarter of all people worldwide. Some [infected people](#) remain healthy, but five to 15 percent eventually develop active tuberculosis disease, which can be deadly. The most common form of the disease occurs in the lungs and is known as pulmonary tuberculosis. Pulmonary tuberculosis can be cured, but little is known about how it develops from the initial infection.

To better understand the mechanisms that underlie development of tuberculosis, Thomas Scriba of the University of Cape Town, South Africa, and colleagues followed 150 adolescents infected with M. tuberculosis for several years. 106 of the [study participants](#) remained healthy, but 44 went on to develop pulmonary tuberculosis within a few years of initial infection.

During the study period, the researchers took regular measurements to monitor and compare immune system activity between individuals who remained healthy and those who eventually fell ill. They found that some differences were detectable as early as one to two years before diagnosis, while some were only detectable just before active disease began.

Eighteen months before diagnosis, the researchers found, individuals had elevated activity of immune system signaling molecules known as interferons, which aid in fighting infection. They also had elevated activity of the complement system, another immune system component. In the days just before diagnosis, additional immune changes occurred, including increased activity of [white blood cells](#).

The scientists also analyzed which genes were expressed in T cells (a type of white blood cell) purified from study participants' blood. They found that certain genes associated with T cells' response to infection were suppressed in those who later developed [pulmonary tuberculosis](#).

Overall, these findings lay out a clearer timeline of biological events that

occur along the path from infection to disease. With further research, this knowledge could aid development of new strategies for diagnosis, vaccination, and treatment.

More information: Scriba TJ, Penn-Nicholson A, Shankar S, Hraha T, Thompson EG, Sterling D, et al. (2017) Sequential inflammatory processes define human progression from M. tuberculosis infection to tuberculosis disease. *PLoS Pathog* 13(11): e1006687.

doi.org/10.1371/journal.ppat.1006687

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