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computer graphics software  
for Linux Tuberculosis (TB)  
is a persistent disease and  
the causative agent,  
Mycobacterium tuberculosis  
(MTB), is acquired early in  
life and usually remains  
latent in the host throughout  
the lifetime. Despite  
effective TB control  
measures, the disease is the  
leading cause of death from

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an infectious disease worldwide. The mechanisms that allow MTB to remain latent within its host and be reactivated at times of immunosuppression are poorly understood.

Accordingly, the development of therapeutic strategies against latent MTB infection has been limited. In our recent work,



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we discovered that heparan sulfate (HS), a cell surface glycoconjugate that is associated with the host response to *M. tuberculosis*, is required for dormancy-like survival of MTB within human macrophages. We further demonstrated that the HS biosynthetic enzyme, heparan sulfate glucosaminyl 3-O-

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sulfotransferase-1  
(HS3ST1), is expressed at  
high levels during latent  
infection, suggesting that it  
plays a role in the  
acquisition of latency. In  
this project, we will further  
characterize the role of  
HS3ST1 in the  
establishment and  
maintenance of latent  
infection and the

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mechanisms of its biosynthesis and regulation. In Aim 1, we will determine the role of HS3ST1 in the acquisition of latency in mice that are deficient in specific immune cells using aerosol-based models of infection. In Aim 2, we will characterize the transcriptional regulation of HS3ST1 and the interaction

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of its encoded protein with other bacterial factors involved in the establishment and maintenance of latent infection. These studies will provide new insights into the mechanisms of latency and the interaction of the bacteria with host tissues, and will identify new targets for the development of

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treatments against latent TB. The pyruvate dehydrogenase complex is the pivotal enzyme that catalyzes the first committed step in carbohydrate metabolism by which NADH is oxidized to acetyl-CoA. It is composed of four non-identical protein components, E1, E2, E3 and the dihydrolipoamide

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acetyltransferase. The E1 component is the catalytic component of the dehydrogenase complex and consists of two tightly associated, non-covalently bound, tightly bound, non-coval

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