

Cystic Fibrosis Research News

Title:

IL-22 exacerbates weight loss in a murine model of chronic pulmonary *Pseudomonas aeruginosa* infection

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What was your research question?

We asked what role interleukin-22 (IL-22), a type of signal used by the immune system, has in people with cystic fibrosis (CF) infected with the bacteria *Pseudomonas aeruginosa*.

Why is this important?

Pseudomonas aeruginosa infection is common in individuals with CF and often results in chronic (long-term) lung infections. There are no current treatments that reliably prevent or clear this infection. The immune response acting against *Pseudomonas* is incompletely understood but may help prevent or clear the bacteria. IL-22 protects against a number of lung infections, but its role against *Pseudomonas* respiratory infection was unknown. IL-22 also helps repair lung tissue damage. Therefore, IL-22 may be important in preventing damaging lung infection with *Pseudomonas* in people with CF.

What did you do?

We examined levels of IL-22 in lung tissue from people with CF with a *Pseudomonas* infection, who were undergoing lung transplantation. We used a mouse to model chronic *Pseudomonas* lung infection and looked to see if IL-22 was produced and what the effect of IL-22 was against the bacteria. Mice which could not produce IL-22 were compared to mice

Cystic Fibrosis Research News

with normal IL-22 levels to discover the effect of IL-22 in preventing and controlling ongoing *Pseudomonas* infection. In addition we examined the impact on survival, lung damage and weight-loss related to infection.

What did you find?

We showed that patients with CF and *Pseudomonas* infection have IL-22 in the airways and along the inner airway lining. In mice infected with *Pseudomonas*, we showed that immune responses able to produce IL-22 developed against the bacteria. However, while we found that the absence of IL-22 in the mouse infection model did not affect the rates of ongoing *Pseudomonas* infection, level of bacteria in the lung, amount of tissue damage and animal survival. Interestingly, we found that those animals with no IL-22 lost less weight than those with it. IL-22 appears to be linked to be exacerbating weight loss in *Pseudomonas* infection.

What does this mean and reasons for caution?

Our findings show that IL-22 is produced in the immune response against *Pseudomonas* infection both in people with CF and a mouse model. However, in the mouse model, IL-22 is unable to prevent or reduce persistent lung infection. Surprisingly we found that IL-22 may play a role in the weight-loss associated with lung *Pseudomonas* infection. This is important for patients with CF as weight-loss is a significant problem that affects long-term health and survival. We did not investigate in this study the way which IL-22 affects weight-loss. In addition, the effects of IL-22 were examined in the mouse model, whether a similar effect exists in people with CF needs to be confirmed.

What's next?

The effect of IL-22 in exacerbating weight-loss in response to lung infection with *Pseudomonas* infection is important so further work is needed to investigate the mechanism(s) by which IL-22 cause weight-loss.

Original manuscript citation in PubMed

<http://www.ncbi.nlm.nih.gov/pubmed/?term=IL-22+exacerbates+weight+loss+in+a+murine+model+of+chronic+pulmonary+Pseudomonas+infection>