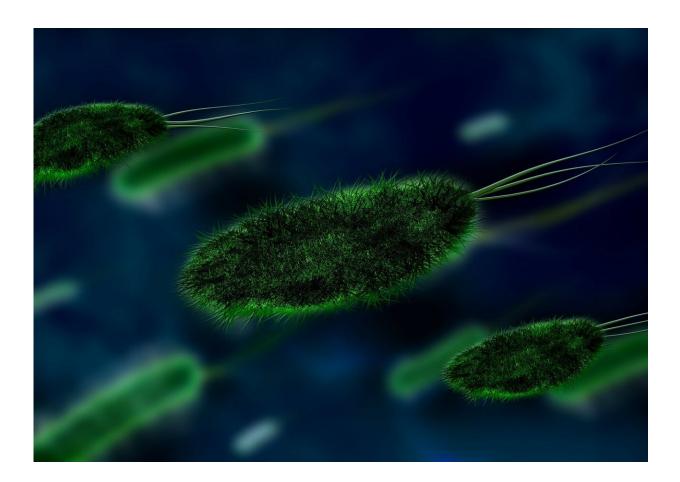


Klebsiella pneumoniae: An opportunistic pathogen harmless to some, but causes severe disease in others

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Klebsiella pneumoniae is a common species of bacteria found in our



bodies—and may even be lurking in your gut, mouth or nose right now. But it's also a notoriously harmful bacteria that can make us very ill.

It's the most common cause of <u>hospital-acquired pneumonia</u> in the US and the second most frequent cause of <u>urinary tract infections</u> worldwide, after Escherichia coli (E coli). If it infects wounds or enters the bloodstream, K pneumoniae can cause <u>bloodstream infections and</u> <u>sepsis</u>.

So how can K pneumoniae live harmlessly among the rest of the microbiome in some of us, yet cause disease in others? Understanding this may hold the key to preventing infections.

Scientists aren't entirely sure what proportion of the population carries K pneumoniae as part of their normal gut microbiome. Past attempts have had highly variable results.

For example, one survey of healthy people detected K pneumoniae in almost 4% of stool samples. Yet other studies show K pneumoniae is noticeably more common among certain groups—including <u>hospital</u> patients, people living in <u>lower income countries</u> and, in particular, among people who had traveled to <u>Asia</u>.

K pneumoniae is what's known as an opportunistic pathogen. This means that when carried in the gut, nose or mouth as part of the normal microbiota, K pneumoniae should not cause any health problems unless a person's immune system becomes compromised due to an <u>infection</u> or disease. So our microbiome can act as a reservoir of K pneumoniae, from which it can spread to other parts of the body and cause infection.

<u>One study</u> of 498 <u>intensive care patients</u> at a hospital in Australia found that half of K pneumoniae infections were caused by the patient's own K pneumoniae strain that had already been living in their gut or throat.



It's thought that K pneumoniae can spread from the gut to other parts of the body via medical devices such as <u>ventilators</u>. This type of gut-to-lung translocation has recently been observed in other pneumonia-causing species of <u>bacteria</u>, such as <u>Pseudomonas aeruginosa</u>. Surgeries can also make possible the spread of K pneumoniae to sites where it can cause infection.

Stopping the spread

Unfortunately, some K pneumoniae strains have developed <u>high levels of</u> <u>drug resistance</u>. This means that some drugs once used to treat K pneumoniae infections now no longer work.

It's particularly concerning that some strains of K pneumoniae are developing resistance to the group of antibiotics called carbapenems, which are generally only used as a <u>last resort</u> treatment when other antibiotics haven't worked. And, this resistance is becoming more widespread among the population.

There's an <u>urgent need</u> to develop alternatives to antibiotics so that cases of drug-resistant K pneumoniae can be prevented or treated. Our laboratory's research focuses on harnessing the <u>gut microbiome</u> as a potential solution.

Since carrying K pneumoniae in the gut is a known <u>risk factor</u> for subsequent infection, one route to avoiding this could be to manipulate the microbiome. This could be done by using probiotics containing beneficial species of bacteria to limit K pneumoniae in the gut. Such a solution could be especially important for people in hospitals or care homes, where K pneumoniae is more <u>prevalent</u> and infection risk is highest.

The microbiome has long been known to provide a host with a degree of



natural protection against infection via a property known as <u>colonization</u> <u>resistance</u>. This is when resident gut bacteria outcompete incoming species, including potential pathogens, and prevent them from establishing in the gut.

But microbiomes <u>vary greatly</u>—and some people carry more protective microbial communities than others.

My colleagues and I wanted to understand why some gut communities can resist the growth of harmful bacteria while others cannot. In the lab, we combined human gut bacteria into communities containing different diversities and compositions of bacterial species. We then challenged these communities with K pneumoniae (as well as other harmful bacteria, such as Salmonella).

We found that <u>diverse gut microbiomes</u> were more protective against K pneumoniae colonization. We showed that this protection was due to the resident gut bacteria using up the nutrients needed in order for invading microbes to grow. This led us to develop a way of predicting combinations of gut bacteria that can resist growth of unwanted species of bacteria such as K pneumoniae.

We are still only just beginning to understand the role that microbes play when it comes to our health. Some of these microbes, such as K pneumoniae, can even be harmful and harmless at the same time. Studying the interactions between the members of the gut microbiota is a critical area of research for microbiome scientists because it could lead to new ways of preventing or treating infections.

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