

Immune cell model paves way for new treatments targeting common infection among immunocompromised children

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Researchers have successfully engineered human immune cells to model an infection common among immunocompromised people in a discovery that paves the way for new drug testing and treatments.



The research, led by the Murdoch Children's Research Institute and published in *Stem Cell Reports*, used cutting edge stem cell technology to better understand how the <u>infection</u> invades <u>immune cells</u> and causes health complications, such as lung and skin and soft tissue infections, in immunocompromised people, particularly those with <u>cystic fibrosis</u>.

Murdoch Children's researcher Dr. Shicheng Jacky Sun said the immune cell type the team created in the lab, known as a macrophage, played an important role in infection, inflammation and regeneration. But due to this function was also a natural host for germs.

"We were also able to use our stem cell model to rapidly test and screen different types of antibiotics against mycobacterium."

Murdoch Children's researcher Dr. Sohinee Sarkar said the search for effective treatments had been hampered until now by the lack of infection models to test new drugs.

"Mycobacteria opportunistically infect people with lung diseases, such as cystic fibrosis, and also causes skin and soft tissue infections in those with immunodeficiencies," she said.

"Current treatments take months and involve giving cocktails of different antibiotics with wide-ranging toxicities. Treatments often fail as the infection is highly resistant to antibiotics, leaving infected people with few other options. Patients with mycobacteria are also excluded from receiving life-saving lung transplants."



Dr. Sarkar said due to high treatment failure rates, repeated cycles of infection could greatly damage the lung tissue and accelerate progression of lung failure in those with cystic fibrosis.

"Improved treatments could mean less frequent hospital visits, shorter stays and minimal exposure to toxic antibiotics, which is particularly important for children with cystic fibrosis," she said.

Data shows 11 percent of children with cystic fibrosis test positive for mycobacteria.

Dr. Sarkar said the infection model could also be used for drug screening for other superbugs with limited treatment options.

"Some bacteria have evolved to escape our <u>immune system</u> by hiding themselves within host cells, making it difficult to treat these infections with traditional antibiotics," she said. "Our stem cell-based infection model can be readily scaled up to screen large number of drugs against such bacteria to identify new treatments."

Murdoch Children's Professor Ed Stanley, also a member of the new global stem cell medicine consortium, reNEW, added that the team was looking to extend and advance its stem cell research capability to contribute to the global efforts to find treatments and therapies for a range of currently incurable conditions.

More information: Shicheng Sun et al, Human pluripotent stem cell-derived macrophages host Mycobacterium abscessus infection, *Stem Cell Reports* (2022). DOI: 10.1016/j.stemcr.2022.07.013

Provided by Murdoch Children's Research Institute



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