

Doctors: Long-lasting cell treatment cured leukemia patients

February 6 2022, by Laura Ungar







This undated photo provided by the family shows Doug Olson, center foreground, of Pleasanton, Calif., on his 75th birthday. In 2010, doctors treated Olson's leukemia with an experimental gene therapy that transformed his own blood cells into cancer killers. More than a decade later, there's no sign of cancer in his body. Credit: Family Photo via AP

In 2010, doctors treated Doug Olson's leukemia with an experimental gene therapy that transformed some of his blood cells into cancer killers. More than a decade later, there's no sign of cancer in his body.

The treatment cured Olson and a second patient, according to the University of Pennsylvania doctors, who said it was the first time the therapy had been studied for so long.

"I'm doing great right now. I'm still very active. I was running half marathons until 2018," said Olson, 75, who lives in Pleasanton, California. "This is a cure. And they don't use the word lightly."

His doctors describe the two cases in a study published Wednesday in the journal *Nature*. They say the two examples show the treatment, called CAR-T cell therapy, can attack <u>cancer</u> immediately, then stay inside the body for years and evolve there to keep the disease at bay. Such so-called "living drugs" are now used by thousands around the world to treat certain blood cancers.

Based on the results, "we can now conclude that CAR-T cells can actually cure patients of leukemia," said Dr. Carl June, one of the authors of the study.

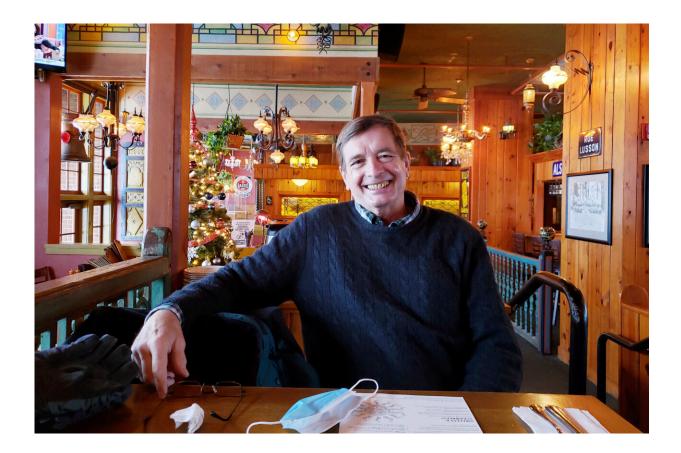
The <u>one-time treatment</u> involves collecting the patient's own T cells, white <u>blood cells</u> key to the <u>immune system</u>, and genetically changing



them in the lab so that they will find and attack cancer cells. The modified cells are given back to the patient through IV.

By the time Olson got the treatment, he'd been fighting cancer for years. When doctors diagnosed him with chronic lymphocytic leukemia in 1996, he said, "I thought I had months to live."

He eventually underwent chemotherapy and, at one point, his physician, Dr. David Porter, suggested he may need a bone marrow transplant. Porter also raised the idea of joining a CAR-T therapy study. Olson, CEO of a New Hampshire lab products company, said he was excited by the science and eager to avoid the transplant.



This 2021 photo provided by the family shows Doug Olson of Pleasanton, Calif., in Bend, Ore. In 2010, doctors treated Olson's leukemia with an experimental



gene therapy that transformed his own blood cells into cancer killers. More than a decade later, there's no sign of cancer in his body. Credit: Family Photo via AP

A couple weeks after getting the treatment, he felt sick for about a week and was hospitalized for three days.

"It was the very next week he sat me down and he said, 'We cannot find a single cancer cell in your body," Olson recalled.

The other patient, retired corrections officer Bill Ludwig, had similar results.

Over time, researchers said, the modified cells evolved, many turning into "helper" cells that work with the cancer-killing cells. Helper cells eventually became dominant in both patients.

Study author J. Joseph Melenhorst said they were able to isolate and analyze the cells using new technologies, which gave them "very good insight" into how they persisted in the patients' bodies.

Dr. Armin Ghobadi of Washington University in St. Louis, an expert in gene and cellular immunotherapy for cancer, called the findings "incredible." Though the word "cure" is rarely used in cancer, he said it appears these patients were "most likely" cured.

He was intrigued by the persistence of the CAR-T <u>cells</u> and the way the living drug evolves.

"That's just really beautiful to see," said Ghobadi, who was not involved in the study.





This 2022 photo provided by the family shows Doug Olson of Pleasanton, Calif., at Joshua Tree National Park in California. In 2010, doctors treated Olson's leukemia with an experimental gene therapy that transformed his own blood cells into cancer killers. More than a decade later, there's no sign of cancer in his body. Credit: Family Photo via AP

At this point, June said, tens of thousands of patients are being treated with CAR-T cell therapies, which have been approved for certain blood cancers by health authorities around the world, including the U.S. Food and Drug Administration. The agency first approved a CAR-T therapy



treatment in 2017 developed by Penn and the drugmaker Novartis for childhood leukemia.

The *Nature* study was paid for partly by the Novartis Institute for Biomedical Research and partly by National Institutes of Health grants.

Scientists hope to see wider use of CAR-T therapies in the future for other cancers. Last year, a CAR-T cell <u>therapy</u> was approved for multiple myeloma, the most common malignancy of the bone marrow in adults. According to the Leukemia & Lymphoma Society, leukemia, lymphoma and myeloma were expected to make up just under 10% of the 1.9 million new cancer cases in the U.S. last year.

"But the big scientific challenge—and it's a big one—is how to make this work in solid cancers," like those in the lung, colon and other places, June said.

Even in blood cancers, there are challenges. The therapies are expensive, running into the hundreds of thousands of dollars just for the drugs. And there's the risk of significant side effects, including an immune overreaction called "cytokine release syndrome" and nervous system-related problems such as brain swelling.

Both of the Penn patients did extremely well after the treatment. Ludwig traveled the country with his wife in a motor home and celebrated family milestones before dying early last year of COVID-19 complications.

Olson said he's extremely grateful for the decade of life since doctors used cutting-edge science to save him.

"What's changed is the dimension of hope. The pace of discovery takes your breath away," he said. "It's a brand new world."



More information: J. Joseph Melenhorst, Decade-long leukemia remissions with persistence of CD4+ CAR T-cells, *Nature* (2022). DOI: <u>10.1038/s41586-021-04390-6</u>. <u>www.nature.com/articles/s41586-021-04390-6</u>

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