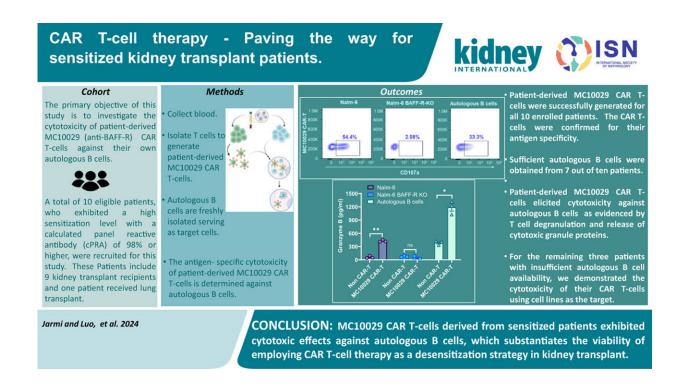


## Could CAR-T cell therapy improve kidney transplants?

April 9 2024, by Susan Buckles, Mayo Clinic News Network



Credit: Kidney International (2024). DOI: 10.1016/j.kint.2024.02.008

Chimeric antigen receptor-T cell therapy (CAR-T cell therapy) could provide a revolutionary approach to organ transplantation for patients who are hard to match and susceptible to rejection, Mayo Clinic researchers discovered.

Their pioneering research focuses on using CAR-T cells derived from



the patient's own immune system to prevent rejection of donated organs. Sensitized <u>patients</u> are those who have high levels of antibodies that cause their immune systems to react negatively to potential donor organs. These patients often face extended waiting periods for a transplant.

Research from this proof-of-concept <u>study</u> is published in *Kidney International*.

"This study was designed to evaluate the feasibility of using an immunotherapy that would remove B cells that cause sensitization in patients waiting for a kidney transplant," says Tambi Jarmi, M.D., first author on the study and a transplant nephrologist at Mayo Clinic in Florida.

"This research is one of the first steps toward applying CAR-T cells in the field of transplantation to try to make more donor organs available for transplant and reduce the wait for patients who need a new kidney."

## Research on CAR-T cell therapy and transplantation

Researchers in the lab of Hong Qin, M.D., Ph.D., developed a new type of CAR-T cell therapy that targets a protein known as B cell activating factor receptor (BAFF-R), which is present on B cells. B cells play a key role in regulating the immune system and generating antibodies. They investigated whether the BAFF-R CAR-T cells could target and remove B cells that cause sensitization.

For this study, researchers generated BAFF-R CAR-T cells from 10 patients considered sensitized to <u>organ transplantation</u>. Nine of them had undergone kidney transplants and one had received a lung transplant.

"We isolated T cells from a small quantity of the patient's blood. These CAR-T cells were then engineered to identify and neutralize B cells



carrying a specific antigen, effectively targeting and eliminating the sensitizing cells," says Dr. Qin, lead author on the study, who is an immunologist and CAR-T cell researcher.

Then, the research team compared the activity of BAFF-R CAR-T cells to T cells after they were treated with B cells from healthy donors. Researchers saw a reduction in B cells from the sensitized patient group that had been treated with patient-derived CAR-T cells when compared to the non-CAR-T control group.

"Our study demonstrated that BAFF-R CAR-T cells derived from sensitized patients exhibited killing effects against B cells, which substantiates the viability of using BAFF-R CAR-T cell therapy as a desensitization strategy in kidney transplant," says Dr. Jarmi.

Mayo Clinic now is preparing to biomanufacture the BAFF CAR-T cells at its commercial-grade facilities in Florida for testing in early-stage clinical trials.

The use of BAFF-R CAR-T cell therapy to desensitize <u>kidney transplant</u> patients is considered experimental. It is among the first investigations into the use of this immunotherapy for diseases beyond cancer. This research is a collaboration between the Mayo Clinic Regenerative Immunotherapy and CAR-T Translational Research Program, Mayo Clinic Department of Transplant Florida and Mayo Clinic Division of Hematology and Medical Oncology.

Additional research is needed to validate the findings of this study.

**More information:** Tambi Jarmi et al, CAR T-cell therapy—paving the way for sensitized kidney transplant patients, *Kidney International* 



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