



CAR-T cells can be used for:

- 1. Compound screening
- 2. Antibody screening
- 3. Co-stimulatory and activation domain comparison
- 4. Personalized medicine and donor variations for CAR-T screening
- 5. Checkpoint inhibitors
- 6. Safety switches and regulators of CAR-T functions
- 7. Pre-clinical in vivo models
- 8. Treg and T memory cells in CAR-T setting
- 9. CAR-T signaling, tumor microenvironment
- 10. Proof of concept studies for clinical trials

The structure of CAR from Promab's available CAR-T cells targeting EGFR antigen EGFR (epidermal growth factor receptor (EGFR; ErbB-1; HER1) is a cell-surface receptor for the epidermal growth factor family (EGF family) receptors (Her-2, Her-3, Her-4). EGFR-1 is overepxressed in a number of tumors: breast, lung, glioblastoma and other types and can be used as a marker for CAR-T immunotherapy. The EGFR-CAR-T cells are available with CD28, 4-1BB and GITR co-activation domains (Figure 1).

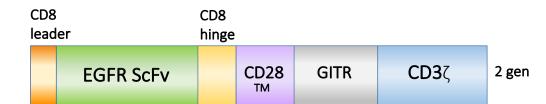


Figure 1. CAR-T cells expressing the above constructs are available from Promab targeting EGFR antigen. ScFv, single chain variable fragment. These CAR-T cells are generated with EGFR-GITR-CD3 zeta construct.

To date Promab generated 2nd or 3rd generation CAR and CAR controls (2nd generation of CAR is shown in Figure 1, CAR-T cells and CAR-Natural Killer (NK) effector cells against cancer target cells that show excellent functionality, including dose-dependent and target cell-specific cytotoxic activity.

These cells can be tested with CAR-T in cytotoxic assays and used for testing modulators of immune checkpoint inhibitors (PD-1, CTLA-4 pathways) or activators of immune response, small molecules affecting T cell or T reg activity.





Data MDA-231 cells 0.9 0.8 စို 0.7 **Target Cells alone** ੂ ਭ0.6 **Target Cells alone** 0.5 80.5 + Non-transduced T cells Ĕ 0.4 + Mock ScFv-CD28-CD3 CAR-T cells 0.3 + EGFR-GITR-CD3-CAR-T cells 0.2 0.1 23.2 21.2 33.2 35.2

Figure 2. RTCA cytotoxicity activity of effector EGFR-GITR-CD3-CAR-T cells against MDA-231 target cells. Effector:Target cells ratio=10:1.