

Efficacy evaluation of optimized CAR-T therapy against refractory epithelial ovarian cancer



ONCOLOGY	Candidate
Product Type	CAR-T cells
Indication	Refractory epithelial ovarian cancer
Target	Mesothelin (MSLN)
MoA(Mechanism of Action)	Chimeric antigen receptor (CAR) on CAR-T binds to mesothelin (MSLN) on cancer cells. The binding of CAR to MSLN then triggers the activation signals (through 41BB and CD3z domain of CAR) into CAR-T cells and allows the cytotoxic effect of CAR-T to eliminate cancer cells
Competitiveness	MSLN is the most targeted antigen and attractive target for CAR-T therapy against solid tumors. MSLN, a cell surface glycoprotein, is highly expressed in multiple solid tumors including ovarian cancer, whereas it is expressed at low levels in normal mesothelial tissues. However, all MSLN CAR-T being developed based on SS1-scFv harboring antigen binding moiety to membrane-distal epitope of MSLN have failed in clinical trials due to limited efficacy. Here, we developed a new version of scFv which has binding affinity to Region II domain (middle) of MSLN. MSLN CAR-T based on the new version of scFv showed anti-tumor activity higher than that of SS1-based CAR-T and further optimization led to the version of efficacy with complete remission in MSLN-positive tumor xenograft mice.
Development Stage	Candidate
Route of Administration	Parenteral-intravenous