

425, and a VH domain comprising a CDR-H1 comprising the amino acid sequence of SEQ ID NO: 426, a CDR-H2 comprising the amino acid sequence of SEQ ID NO: 427, and a CDR-H3 comprising the amino acid sequence of SEQ ID NO: 428.

In another aspect, the present disclosure relates to a pharmaceutical composition comprising the polypeptide as disclosed herein and a pharmaceutically acceptable carrier.

In another aspect, the present disclosure relates to use of the polypeptide as disclosed herein in the manufacture of a medicament for treating a DLL3 expressing tumor or DLL3 expressing cancer disease or a DLL3 expressing metastatic cancer disease.

In another aspect, the present disclosure relates to a kit comprising the polypeptide as disclosed herein and a pharmaceutically acceptable excipient.

Thus, in a first aspect, the present invention provides a bispecific antibody construct comprising a first binding domain which binds to human DLL3 on the surface of a target cell and a second binding domain which binds to human CD3 on the surface of a T cell, wherein the first binding domain binds to an epitope of DLL3 which is comprised within the region as depicted in SEQ ID NO: 260.

It must be noted that as used herein, the singular forms “a”, “an”, and “the” include plural references unless the context clearly indicates otherwise. Thus, for example, reference to “a reagent” includes one or more of such different reagents and reference to “the method” includes reference to equivalent steps and methods known to those of ordinary skill in the art that could be modified or substituted for the methods described herein.

Claims

1. A polypeptide comprising a first binding domain which binds to human delta-like 3 (DLL3) on the surface of a target cell and a second binding domain which binds to human CD3 on the surface of a T cell, wherein the first binding domain binds to an epitope of DLL3 within the amino acid sequence of SEQ ID NO: 258,

wherein the first binding domain comprises a VH region comprising CDR-H1, CDR-H2 and CDR-H3 and a VL region comprising CDR-L1, CDR-L2 and CDR-L3 selected from the group consisting of:

- a) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 31, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 32, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 33, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 34, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 35 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 36;
- b) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 41, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 42, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 43, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 44, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 45 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 46;
- c) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 51, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 52, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 53, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 54, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 55 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 56;
- d) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 61, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 62, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 63, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 64, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 65 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 66;
- e) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 71, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 72, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 73, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 74, CDR-L2

comprising the amino acid sequence of SEQ ID NO: 75 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 76;

f) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 81, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 82, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 83, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 84, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 85 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 86;

g) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 91, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 92, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 93, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 94, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 95 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 96;

h) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 101, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 102, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 103, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 104, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 105 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 106; and

i) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 111, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 112, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 113, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 114, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 115 and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 116,

and wherein the second binding domain comprises a VL region comprising CDR-L1, CDR-L2 and CDR-L3 and a VH region comprising CDR-H1, CDR-H2 and CDR-H3 selected from the group consisting of:

a) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 342, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 343, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 344; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 345, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 346, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 347;

b) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 351, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 352, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 353; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 354, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 355, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 356;

c) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 360, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 361, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 362; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 363, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 364, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 365;

d) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 369, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 370, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 371; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 372, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 373, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 374;

e) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 378, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 379, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 380; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 381, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 382, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 383;

f) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 387, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 388, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 389; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 390, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 391, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 392;

g) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 396, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 397, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 398; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 399, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 400, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 401;

h) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 405, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 406, CDR-L3 comprising the amino acid sequence

of SEQ ID NO: 407; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 408, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 409, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 410;

i) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 414, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 415, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 416; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 417, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 418, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 419; and

j) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 423, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 424, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 425; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 426, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 427, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 428.

2. The polypeptide according to claim 1, wherein the second binding domain comprises a VL region comprising the amino acid sequence selected from the group consisting of: SEQ ID NO: 349, SEQ ID NO: 358, SEQ ID NO: 367, SEQ ID NO: 376, SEQ ID NO: 385, SEQ ID NO: 394, SEQ ID NO: 403, SEQ ID NO: 412, SEQ ID NO: 421, and SEQ ID NO: 430.
3. The polypeptide according to claim 1, wherein the second binding domain comprises a VH region comprising the amino acid sequence selected from the group consisting of: SEQ ID NO: 348, SEQ ID NO: 357, SEQ ID NO: 366, SEQ ID NO: 375, SEQ ID NO: 384, SEQ ID NO: 393, SEQ ID NO: 402, SEQ ID NO: 411, SEQ ID NO: 420, and SEQ ID NO: 429.
4. The polypeptide according to claim 1, wherein the second binding domain comprises a VH region and a VL region comprising the pair of amino acid sequences, respectively, selected from the group consisting of: SEQ ID NOs: 348 and 349; SEQ ID NOs: 357 and 358; SEQ ID NOs: 366 and 367; SEQ ID NOs: 375 and 376; SEQ ID NOs: 384 and 385; SEQ ID NOs: 393 and 394; SEQ ID NOs: 402 and 403; SEQ ID NOs: 411 and 412; SEQ ID NOs: 420 and 421; and SEQ ID NOs: 429 and 430.
5. The polypeptide according to claim 1, wherein the second binding domain comprises an amino acid sequence selected from the group consisting of: SEQ ID NO: 350, SEQ ID NO: 359, SEQ ID NO: 368, SEQ ID NO: 377, SEQ ID NO: 386, SEQ ID NO: 395, SEQ ID NO: 404, SEQ ID NO: 413, SEQ ID NO: 422, and SEQ ID NO: 431.

6. The polypeptide according to claim 1, wherein the first binding domain comprises a VH region comprising the amino acid sequence selected from the group consisting of: SEQ ID NO: 37, SEQ ID NO: 47, SEQ ID NO: 57, SEQ ID NO: 67, SEQ ID NO: 77, SEQ ID NO: 87, SEQ ID NO: 97, SEQ ID NO: 107, SEQ ID NO: 117, SEQ ID NO: 435, and SEQ ID NO: 529.
7. The polypeptide according to claim 1, wherein the first binding domain comprises a VL region comprising the amino acid sequence selected from the group consisting of: SEQ ID NO: 38, SEQ ID NO: 48, SEQ ID NO: 58, SEQ ID NO: 68, SEQ ID NO: 78, SEQ ID NO: 88, SEQ ID NO: 98, SEQ ID NO: 108, SEQ ID NO: 118, SEQ ID NO: 436, and SEQ ID NO: 530.
8. The polypeptide according to claim 1, wherein the first binding domain comprises a VH region and a VL region comprising the pair of amino acid sequences, respectively, selected from the group consisting of: SEQ ID NOs: 37 and 38; SEQ ID NOs: 47 and 48; SEQ ID NOs: 57 and 58; SEQ ID NOs: 67 and 68; SEQ ID NOs: 77 and 78; SEQ ID NOs: 87 and 88; SEQ ID NOs: 97 and 98; SEQ ID NOs: 107 and 108; SEQ ID NOs: 117 and 118; SEQ ID NOs: 435 and 436; and SEQ ID NOs: 529 and 530.
9. The polypeptide according to claim 1, wherein the first binding domain comprises a polypeptide comprising the amino acid sequence selected from the group consisting of: SEQ ID NO: 39, SEQ ID NO: 49, SEQ ID NO: 59, SEQ ID NO: 69, SEQ ID NO: 79, SEQ ID NO: 89, SEQ ID NO: 99, SEQ ID NO: 109, SEQ ID NO: 119, SEQ ID NO: 437 and SEQ ID NO: 531.
10. A polypeptide comprising a first binding domain which binds to human DLL3 on the surface of a target cell and a second binding domain which binds to human CD3 on the surface of a T cell, wherein the first binding domain comprises a VH region comprising CDR-H1, CDR-H2 and CDR-H3 and a VL region comprising CDR-L1, CDR-L2 and CDR-L3, and the second binding domain comprises a VH region comprising CDR-H1, CDR-H2 and CDR-H3 and a VL region comprising CDR-L1, CDR-L2 and CDR-L3,

wherein the CDR-H1, CDR-H2 and CDR-H3 and CDR-L1, CDR-L2 and CDR-L3 of the first binding domain are selected from the group consisting of:

- a) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 31, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 32, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 33, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 34, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 35, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 36;

b) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 41, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 42, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 43, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 44, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 45, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 46;

c) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 51, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 52, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 53, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 54, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 55, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 56;

d) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 61, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 62, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 63, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 64, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 65, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 66;

e) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 71, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 72, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 73, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 74, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 75, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 76;

f) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 81, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 82, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 83, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 84, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 85, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 86;

g) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 91, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 92, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 93, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 94, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 95, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 96;

h) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 101, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 102, CDR-H3 comprising the amino acid

sequence of SEQ ID NO: 103, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 104, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 105, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 106; and

i) CDR-H1 comprising the amino acid sequence of SEQ ID NO: 111, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 112, CDR-H3 comprising the amino acid sequence of SEQ ID NO: 113, CDR-L1 comprising the amino acid sequence of SEQ ID NO: 114, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 115, and CDR-L3 comprising the amino acid sequence of SEQ ID NO: 116; and

wherein the CDR-H1, CDR-H2 and CDR-H3 and CDR-L1, CDR-L2 and CDR-L3 of the second binding domain are selected from the group consisting of:

a) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 342, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 343, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 344; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 345, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 346, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 347;

b) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 351, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 352, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 353; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 354, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 355, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 356;

c) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 360, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 361, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 362; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 363, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 364, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 365;

d) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 369, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 370, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 371; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 372, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 373, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 374;

- e) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 378, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 379, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 380; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 381, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 382, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 383;
 - f) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 387, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 388, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 389; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 390, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 391, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 392;
 - g) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 396, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 397, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 398; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 399, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 400, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 401;
 - h) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 405, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 406, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 407; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 408, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 409, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 410;
 - i) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 414, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 415, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 416; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 417, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 418, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 419; and
 - j) CDR-L1 comprising the amino acid sequence of SEQ ID NO: 423, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 424, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 425; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 426, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 427, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 428.
11. The polypeptide of claim 10, wherein the CDR-H1, CDR-H2 and CDR-H3 and CDR-L1, CDR-L2 and CDR-L3 of the second binding domain are CDR-L1 comprising the amino acid

- sequence of SEQ ID NO: 423, CDR-L2 comprising the amino acid sequence of SEQ ID NO: 424, CDR-L3 comprising the amino acid sequence of SEQ ID NO: 425; CDR-H1 comprising the amino acid sequence of SEQ ID NO: 426, CDR-H2 comprising the amino acid sequence of SEQ ID NO: 427, and CDR-H3 comprising the amino acid sequence of SEQ ID NO: 428.
12. The polypeptide of claim 11, wherein the first binding domain comprising a VH and a VL region comprising a pair of amino acid sequences, respectively, selected from the group consisting of: SEQ ID NOs: 37 and 38; SEQ ID NOs: 47 and 48; SEQ ID NOs: 57 and 58; SEQ ID NOs: 67 and 68; SEQ ID NOs: 77 and 78; SEQ ID NOs: 87 and 88; SEQ ID NOs: 97 and 98; SEQ ID NOs: 107 and 108; SEQ ID NOs: 117 and 118; SEQ ID NOs: 435 and 436; and SEQ ID NOs: 529 and 530.
 13. The polypeptide of claim 12, wherein the second binding domain comprises a VH comprising the amino acid sequence of SEQ ID NO: 429 and a VL region comprising the amino acid sequence of SEQ ID NO: 430.
 14. The polypeptide of claim 13, wherein the polypeptide comprises the amino acid sequence of SEQ ID NO: 40, SEQ ID NO: 50, SEQ ID NO: 60, SEQ ID NO: 70, SEQ ID NO: 80, SEQ ID NO: 90, SEQ ID NO: 100, SEQ ID NO: 110, SEQ ID NO: 120, SEQ ID NO: 211, SEQ ID NO: 212, SEQ ID NO: 213, SEQ ID NO: 214, SEQ ID NO: 215, SEQ ID NO: 216, SEQ ID NO: 217, SEQ ID NO: 438 or SEQ ID NO: 532.
 15. The polypeptide of claim 14, wherein the polypeptide comprises the amino acid sequence of SEQ ID NO: 517, SEQ ID NO: 518, SEQ ID NO: 519, SEQ ID NO: 520, SEQ ID NO: 521, SEQ ID NO: 522, SEQ ID NO: 523 or SEQ ID NO: 524.
 16. The polypeptide according to claim 11 or 15, wherein the first binding domain comprises the amino acid sequence of SEQ ID NO: 39 or SEQ ID NO: 437, and wherein the second binding domain comprises the amino acid sequence of SEQ ID NO: 431.
 17. A polypeptide comprising a first binding domain which binds to human DLL3 on the surface of a target cell and a second binding domain which binds to human CD3 on the surface of a T cell, wherein the first binding domain comprises a VH region comprising a CDR-H1 comprising the amino acid sequence of SEQ ID NO: 31, a CDR-H2 comprising the amino acid sequence of SEQ ID NO: 32, and a CDR-H3 comprising the amino acid sequence of SEQ ID NO: 33, and a VL region comprising a CDR-L1 comprising the amino acid sequence of SEQ ID NO: 34, a CDR-L2 comprising the amino acid sequence of SEQ ID NO: 35, and a CDR-L3 comprising the amino acid sequence of SEQ ID NO: 36, and wherein the second binding

domain comprises a VL domain comprising a CDR-L1 comprising the amino acid sequence of SEQ ID NO: 423, a CDR-L2 comprising the amino acid sequence of SEQ ID NO: 424, a CDR-L3 comprising the amino acid sequence of SEQ ID NO: 425, and a VH domain comprising a CDR-H1 comprising the amino acid sequence of SEQ ID NO: 426, a CDR-H2 comprising the amino acid sequence of SEQ ID NO: 427, and a CDR-H3 comprising the amino acid sequence of SEQ ID NO: 428.

18. The polypeptide according to claim 17, wherein the first binding domain comprises a VH region comprising the amino acid sequence of SEQ ID NO: 435 and a VL region comprising the amino acid sequence of SEQ ID NO: 436, and the second binding domain comprises a VH region comprising the amino acid sequence of SEQ ID NO: 429 and a VL region comprising the amino acid sequence of SEQ ID NO: 430, or

wherein the first binding domain comprises a VH region comprising the amino acid sequence of SEQ ID NO: 37 and a VL region comprising the amino acid sequence of SEQ ID NO: 38, and the second binding domain comprises a VH region comprising the amino acid sequence of SEQ ID NO: 429 and a VL region comprising the amino acid sequence of SEQ ID NO: 430.

19. The polypeptide of claim 17 or 18, wherein the polypeptide comprises the amino acid sequence of SEQ ID NO: 520.
20. The polypeptide of claim 1, 10, or 17, wherein the polypeptide is in a format of two single-chain variable fragments ((scFv)₂), or a diabody or an oligomer of any of the foregoing formats.
21. A pharmaceutical composition comprising the polypeptide of claim 1, 10, 15, 17, or 19 and a pharmaceutically acceptable carrier.
22. Use of the polypeptide according to claims 1, 10, 15, 17, or 19 in the manufacture of a medicament for treating a DLL3 expressing tumor or DLL3 expressing cancer disease or a DLL3 expressing metastatic cancer disease.
23. The use according to claim 22, wherein the DLL3 expressing tumor or DLL3 expressing cancer disease is small cell lung cancer (SCLC), melanoma or neuroendocrine prostate cancer.
24. A kit comprising the polypeptide according to claims 1, 10, 15, 17, or 19 and a pharmaceutically acceptable excipient.
25. The kit according to claim 24 further comprising directions for use.

26. The polypeptide of claims 1, 10, or 17, wherein the polypeptide comprises two single-chain variable fragments ((scFv)₂).
27. The polypeptide of claims 1, 10, 15, 17, or 19, or the pharmaceutical composition of claim 21 for use in therapy.