

What is claimed is:

1. Use of an antibody drug conjugate in the manufacture of a medicament for A
~~method of treating~~ urothelial or bladder cancer in a human subject having liver metastases,
wherein an effective amount of the antibody drug conjugate is to be administered to
~~comprising administering to the subject having liver metastases an effective amount of an~~
~~antibody drug conjugate,~~

wherein the subject has received an immune checkpoint inhibitor (CPI)
therapy, and

wherein the antibody drug conjugate comprises an antibody or antigen binding
fragment thereof that binds to 191P4D12 conjugated to one or more units of monomethyl
auristatin E (MMAE), wherein the antibody or antigen binding fragment thereof comprises a
heavy chain variable region comprising complementarity determining regions (CDRs)
comprising the amino acid sequences of the CDRs of the heavy chain variable region set
forth in SEQ ID NO:22 and a light chain variable region comprising CDRs comprising the
amino acid sequences of the CDRs of the light chain variable region set forth in SEQ ID
NO:23.

2. Use of an antibody drug conjugate in the manufacture of a medicament for A
~~method of treating~~ urothelial or bladder cancer in a human subject having a primary site of
tumor in the upper urinary tract, ~~comprising~~ wherein an effective amount of the antibody
drug conjugate is to be administeringadministered to the subject having a primary site of
tumor in the upper urinary tract ~~an effective amount of an antibody drug conjugate,~~

wherein the subject has received an immune checkpoint inhibitor (CPI)
therapy, and

wherein the antibody drug conjugate comprises an antibody or antigen binding
fragment thereof that binds to 191P4D12 conjugated to one or more units of monomethyl
auristatin E (MMAE), wherein the antibody or antigen binding fragment thereof comprises a
heavy chain variable region comprising complementarity determining regions (CDRs)
comprising the amino acid sequences of the CDRs of the heavy chain variable region set
forth in SEQ ID NO:22 and a light chain variable region comprising CDRs comprising the
amino acid sequences of the CDRs of the light chain variable region set forth in SEQ ID
NO:23.

3. Use of an antibody drug conjugate in the manufacture of a medicament for A
~~method of treating~~ urothelial or bladder cancer in a human subject, ~~comprising administering~~

~~to the subject wherein~~ an effective amount of an antibody drug conjugate is to be administered to the subject,

wherein the subject has received an immune checkpoint inhibitor (CPI) therapy,

wherein the subject had progression or recurrence of the cancer during or following the CPI therapy, and

wherein the antibody drug conjugate comprises an antibody or antigen binding fragment thereof that binds to 191P4D12 conjugated to one or more units of monomethyl auristatin E (MMAE), wherein the antibody or antigen binding fragment thereof comprises a heavy chain variable region comprising complementarity determining regions (CDRs) comprising the amino acid sequences of the CDRs of the heavy chain variable region set forth in SEQ ID NO:22 and a light chain variable region comprising CDRs comprising the amino acid sequences of the CDRs of the light chain variable region set forth in SEQ ID NO:23.

4. The ~~method~~use of any one of claims 1 to 3, wherein the subject has a duration of response of (i) at least or about 7 months following the treatment or (ii) ranging from 5 to 9 months following the treatment.

~~5. The method of any one of claims 1 to 3, wherein the subject has a duration of response ranging from 5 to 9 months following the treatment.~~

~~65.~~ The ~~method~~use of claim 1, wherein the subject has a progression free survival (i) of at least or about 4 months following the treatment or (ii) ranging from 4 to 9 months following the treatment.

~~76.~~ The ~~method~~use of claim 2 or 3, wherein the subject has a progression free survival (i) of at least or about 5 months following the treatment or (ii) ranging from 5 to 9 months following the treatment.

~~8. The method of claim 1, wherein the subject has a progression free survival ranging from 4 to 9 months following the treatment.~~

~~9. The method of claim 2 or 3, wherein the subject has a progression free survival ranging from 5 to 9 months following the treatment.~~

~~107.~~ The ~~method~~use of claim 1, wherein the subject has an overall survival (i) of at least or about 9 months following the treatment or (ii) ranging from 9 to 19 months following the treatment.

~~118.~~ The ~~method~~use of claim 2, wherein the subject has an overall survival (i) of at least or about 12 months following the treatment or (ii) ranging from 9 to 19 months following the treatment.

~~129.~~ The ~~method~~use of claim 3, wherein the subject has an overall survival (i) of at least or about 11 months following the treatment or (ii) ranging from 9 to 19 months following the treatment.

~~13.—The method of any one of claims 1 to 3, wherein the subject has an overall survival ranging from 9 to 19 months following the treatment.~~

~~14~~10. The ~~method~~use of any one of claims 1 to 3, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

(i) the percentage of the subjects having complete response in the treated population is at least or about 4%; or

(ii) the percentage of the subjects having partial response in the treated population is at least or about 35%.

~~15.—The method of any one of claims 1 to 3, wherein a population of the subjects is treated by the methods, and wherein the percentage of the subjects having partial response in the treated population is at least or about 35%.~~

~~16~~11. The ~~method~~use of claim 1, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein overall response rate in the treated population is at least or about 35%.

~~17~~12. The ~~method~~use of claim 2, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein overall response rate in the treated population is at least or about 43%.

~~18~~13. The ~~method~~use of claim 3, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein overall response rate in the treated population is at least or about 39%.

~~19~~14. The ~~method~~use of any one of claims 1 to 3, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

(i) the percentage of the subjects having stable disease in the treated population is at least or about 30%;

(ii) median duration of response in the treated population is at least or about 7 months; or

(iii) duration of response in the treated population ranges from 5 to 9 months.

20.—~~The method of any one of claims 1 to 3, wherein a population of the subjects is treated by the methods, and wherein median duration of response in the treated population is at least or about 7 months.~~

21.—~~The method of any one of claims 1 to 3, wherein a population of the subjects is treated by the methods, and wherein duration of response in the treated population ranges from 5 to 9 months.~~

2215. The ~~method~~use of claim 1, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

(i) median progression free survival in the treated population is at least or about 4 months; or

(ii) progression free survival in the treated population ranges from 4 to 9 months.

23.—~~The method of claim 1, wherein a population of the subjects is treated by the methods, and wherein progression free survival in the treated population ranges from 4 to 9 months.~~

2416. The ~~method~~use claim 2 or 3, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

(i) median progression free survival in the treated population is at least or about 5 months; or

(ii) progression free survival in the treated population ranges from 5 to 9 months.

25.—~~The method of claim 2 or 3, wherein a population of the subjects is treated by the methods, and wherein progression free survival in the treated population ranges from 5 to 9 months.~~

2617. The ~~method~~use of claim 1, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

(i) median overall survival in the treated population is at least or about 9 months

(ii) overall survival in the treated population ranges from 9 to 19 months;

(iii) the complete response rate is at least or about 4% for the treated population;

(iv) the partial response rate is at least or about 35% for the treated population;

(v) overall response rate is at least or about 35% for the treated population;

(vi) the median duration of response is at least or about 7 months for the treated population;

(vii) the duration of response is from 5 to 9 months for the treated population;

(viii) the median progression free survival is at least or about 4 months for the treated population;

- (ix) the progression free survival is from 4 to 9 months for the treated population;
- (x) the median overall survival is at least or about 9 months for the treated population; or
- (xi) the overall survival is from 9 to 19 months for the treated population.

2718. The ~~method~~use of claim 2, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

- (i) median overall survival in the treated population is at least or about 12 months;
 - (ii) overall survival in the treated population ranges from 9 to 19 months;
 - (iii) the complete response rate is at least or about 4% for the treated population;
 - (iv) the partial response rate is at least or about 35% for the treated population;
 - (v) overall response rate is at least or about 43% for the treated population;
 - (vi) the median duration of response is at least or about 7 months for the treated population;
 - (vii) the duration of response is from 5 to 9 months for the treated population;
 - (viii) the progression free survival is from 4 to 9 months for the treated population;
 - (ix) the median progression free survival is at least or about 5 months for the treated population;
 - (x) the progression free survival is from 5 to 9 months for the treated population;
 - (xi) the median overall survival is at least or about 12 months for the treated population;
- or
- (xii) the overall survival is from 9 to 19 months for the treated population.

2819. The ~~method~~use of claim 3, wherein a population of the subjects is to be treated by the ~~methods~~medicament, and wherein:

- (i) median overall survival in the treated population is at least or about 11 months
- (ii) overall survival in the treated population ranges from 9 to 19 months;
- (iii) the complete response rate is at least or about 4% for the treated population;
- (iv) the partial response rate is at least or about 35% for the treated population
- (v) overall response rate is at least or about 39% for the treated population;
- (vi) the median duration of response is at least or about 7 months for the treated population;
- (vii) the duration of response is from 5 to 9 months for the treated population;
- (viii) the progression free survival is from 4 to 9 months for the treated population;
- (ix) the median progression free survival is at least or about 5 months for the treated population;
- (x) the progression free survival is from 5 to 9 months for the treated population;
- (xi) the median overall survival is at least or about 11 months for the treated population;

or

(xii) the overall survival is from 9 to 19 months for the treated population.

29. — ~~The method of any one of claims 1 to 3, wherein a population of the subjects is treated by the methods, and wherein overall survival in the treated population ranges from 9 to 19 months.~~

30. — ~~The method of any one of claims 1 to 3, wherein the complete response rate is at least or about 4% for a population of subjects treated with the method.~~

31. — ~~The method of any one of claims 1 to 3, wherein the partial response rate is at least or about 35% for a population of subjects treated with the method.~~

32. — ~~The method of claim 1, wherein overall response rate is at least or about 35% for a population of subjects treated with the method.~~

33. — ~~The method of claim 2, wherein overall response rate is at least or about 43% for a population of subjects treated with the method.~~

34. — ~~The method of claim 3, wherein overall response rate is at least or about 39% for a population of subjects treated with the method.~~

36. — ~~The method of any one of claims 1 to 3, wherein the duration of response is from 5 to 9 months for a population of subjects treated with the method.~~

37. — ~~The method of claim 1, wherein the median progression free survival is at least or about 4 months for a population of subjects treated with the method.~~

38. — ~~The method of any one of claims 1 to 3, wherein the progression free survival is from 4 to 9 months for a population of subjects treated with the method.~~

39. — ~~The method of claim 2 or 3, wherein the median progression free survival is at least or about 5 months for a population of subjects treated with the method.~~

40. — ~~The method of claim 2 or 3, wherein the progression free survival is from 5 to 9 months for a population of subjects treated with the method.~~

41. — ~~The method of claim 1, wherein the median overall survival is at least or about 9 months for a population of subjects treated with the method.~~

42. — ~~The method of claim 2, wherein the median overall survival is at least or about 12 months for a population of subjects treated with the method.~~

43. — ~~The method of claim 3, wherein the median overall survival is at least or about 11 months for a population of subjects treated with the method.~~

44. — ~~The method of any one of claims 1 to 3, wherein the overall survival is from 9 to 19 months for a population of subjects treated with the method.~~

~~4520.~~ The ~~method~~use of any one of claims 1 to ~~4419~~, wherein the subject is a subject that received platinum-based chemotherapy.

~~4621.~~ The ~~method~~use of any one of claims 1 to ~~4520~~, wherein the cancer is urothelial cancer, and wherein the human subject has locally advanced or metastatic urothelial carcinoma.

~~4722.~~ The ~~method~~use of any one of claims 1 to ~~4621~~, wherein the subject has one or more of the conditions selected from the group consisting of:

- (i) absolute neutrophil count (ANC) no less than $1500/\text{mm}^3$;
- (ii) platelet count no less than $100 \times 10^9/\text{L}$;
- (iii) hemoglobin no less than 9 g/dL;
- (iv) serum bilirubin no more than either of 1.5 times of upper limit of normal (ULN) or 3 times ULN for patients with Gilbert's disease;
- (v) CrCl no less than 30 mL/min, and
- (vi) alanine aminotransferase and aspartate aminotransferase no more than 3 fold of ULN;

optionally wherein the subject has all of conditions (i) to (vi); further optionally wherein the CrCl is measured by 24 hour urine collection or estimated by the Cockcroft-Gault criteria.

~~48.—The method of claim 47, wherein the subject has all of conditions (i) to (vi) of claim 47.~~

~~49.—The method of claim 47 or 48, wherein the CrCl is measured by 24 hour urine collection or estimated by the Cockcroft-Gault criteria.~~

~~5023.~~ The ~~method~~use of any one of claims 1 to ~~4922~~, wherein:

- (i) the subject has no more than Grade 2 sensory or motor neuropathy; and/or
- (ii) the subject has no active central nervous system metastases; and/or
- (iii) the subject has no uncontrolled diabetes; optionally wherein the uncontrolled diabetes is determined by hemoglobin A1c (HbA1c) no less than 8% or HbA1c between 7 and 8% with associated diabetes symptoms that are not otherwise explained; further optionally wherein the associated diabetes symptoms comprise or consist of polyuria, polydipsia, or both polyuria and polydipsia.

~~51.—The method of any one of claims 1 to 50, wherein the subject has no active central nervous system metastases.~~

~~52.—The method of any one of claims 1 to 51, wherein the subject has no uncontrolled diabetes.~~

53. — ~~The method of claim 52, wherein the uncontrolled diabetes is determined by hemoglobin A1c (HbA1c) no less than 8% or HbA1c between 7 and 8% with associated diabetes symptoms that are not otherwise explained.~~

54. — ~~The method of claim 53, wherein the associated diabetes symptoms comprise or consist of polyuria, polydipsia, or both polyuria and polydipsia.~~

5524. The ~~method~~use of any one of claims 1 to 5423, wherein:

(i) the CPI therapy is a therapy of programmed death receptor-1 (PD-1) inhibitor; optionally wherein the PD-1 inhibitor is nivolumab or pembrolizumab; or (ii) the CPI therapy is a therapy of programmed death-ligand 1 (PD-L1) inhibitor; optionally wherein the PD-L1 inhibitor is selected from a group consisting of atezolizumab, avelumab, and durvalumab.

56. — ~~The method of any one of claims 1 to 54, wherein the CPI therapy is a therapy of programmed death-ligand 1 (PD-L1) inhibitor.~~

57. — ~~The method of claim 55, wherein PD-1 inhibitor is nivolumab or pembrolizumab.~~

58. — ~~The method of claim 56, wherein PD-L1 inhibitor is selected from a group consisting of atezolizumab, avelumab, and durvalumab.~~

5925. The ~~method~~use of any one of claims 1 to 5824, wherein the antibody or antigen binding fragment thereof comprises CDR-H1 comprising the amino acid sequence of SEQ ID NO:9, CDR-H2 comprising the amino acid sequence of SEQ ID NO:10, CDR-H3 comprising the amino acid sequence of SEQ ID NO:11; CDR-L1 comprising the amino acid sequence of SEQ ID NO:12, CDR-L2 comprising the amino acid sequence of SEQ ID NO:13, and CDR-L3 comprising the amino acid sequence of SEQ ID NO:14, or

wherein the antibody or antigen binding fragment thereof comprises CDR-H1 comprising the amino acid sequence of SEQ ID NO:16, CDR-H2 comprising the amino acid sequence of SEQ ID NO:17, CDR-H3 comprising the amino acid sequence of SEQ ID NO:18; CDR-L1 comprising the amino acid sequence of SEQ ID NO:19, CDR-L2 comprising the amino acid sequence of SEQ ID NO:20, and CDR-L3 comprising the amino acid sequence of SEQ ID NO:21.

6026. The ~~method~~use of any one of claims 1 to 5824, wherein the antibody or antigen binding fragment thereof comprises CDR-H1 consisting of the amino acid sequence of SEQ ID NO:9, CDR-H2 consisting of the amino acid sequence of SEQ ID NO:10, CDR-H3 consisting of the amino acid sequence of SEQ ID NO:11; CDR-L1 consisting of the amino acid sequence of SEQ ID NO:12, CDR-L2 consisting of the amino acid sequence of SEQ ID NO:13, and CDR-L3 consisting of the amino acid sequence of SEQ ID NO:14, or

wherein the antibody or antigen binding fragment thereof comprises CDR-H1 consisting of the amino acid sequence of SEQ ID NO:16, CDR-H2 consisting of the amino acid sequence of SEQ ID NO:17, CDR-H3 consisting of the amino acid sequence of SEQ ID NO:18; CDR-L1 consisting of the amino acid sequence of SEQ ID NO:19, CDR-L2 consisting of the amino acid sequence of SEQ ID NO:20, and CDR-L3 consisting of the amino acid sequence of SEQ ID NO:21.

~~6127.~~ The ~~method~~use of any one of claims 1 to ~~6026~~6127, wherein the antibody or antigen binding fragment thereof comprises a heavy chain variable region comprising the amino acid sequence of SEQ ID NO:22 and a light chain variable region comprising the amino acid sequence of SEQ ID NO:23.

~~6228.~~ The ~~method~~use of any one of claims 1 to ~~6127~~6228, wherein the antibody comprises a heavy chain comprising the amino acid sequence ranging from the 20th amino acid (glutamic acid) to the 466th amino acid (lysine) of SEQ ID NO:7 and a light chain comprising the amino acid sequence ranging from the 23rd amino acid (aspartic acid) to the 236th amino acid (cysteine) of SEQ ID NO:8.

~~6329.~~ The ~~method~~use of any one of claims 1 to ~~6127~~6329, wherein:

- (i) the antigen binding fragment is an Fab, F(ab')₂, Fv or scFv; and/or
- (ii) the antigen binding fragment is recombinantly produced.

~~6430.~~ The ~~method~~use of any one of claims 1 to ~~6228~~6430, wherein:

- (i) the antibody is a fully human antibody;
- (ii) the antibody is an IgG1 and light chain is a kappa light chain; and/or
- (iii) the antibody is recombinantly produced.

~~65.—The method of any one of claims 1 to 62 and 64, wherein the antibody is an IgG1 and light chain is a kappa light chain~~

~~66.—The method of any one of claims 1 to 65, wherein the antibody or antigen binding fragment thereof is recombinantly produced.~~

~~6731.~~ The ~~method~~use of any one of claims 1 to ~~6630~~6731, wherein the antibody or antigen binding fragment is conjugated to each unit of MMAE via a linker.

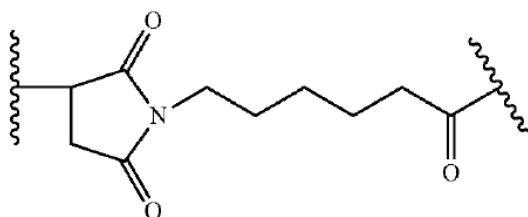
~~6832.~~ The ~~method~~use of claim ~~6731~~6832, wherein:

(i) the linker is an enzyme-cleavable linker, and wherein the linker forms a bond with a sulfur atom of the antibody or antigen binding fragment thereof; and/or

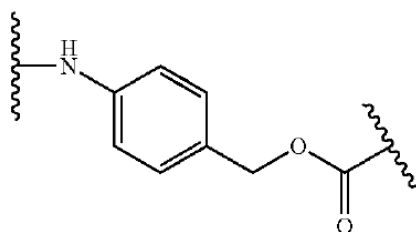
(ii) the linker has a formula of: –Aa–Ww–Yy–; wherein –A– is a stretcher unit, a is 0 or 1; –W– is an amino acid unit, w is an integer ranging from 0 to 12; and –Y– is a spacer unit, y is 0, 1, or 2.

~~69. The method of claim 67 or 68, wherein the linker has a formula of: Aa-Ww-Yy ; wherein A is a stretcher unit, a is 0 or 1; W is an amino acid unit, w is an integer ranging from 0 to 12; and Y is a spacer unit, y is 0, 1, or 2.~~

7033. The ~~method~~use of claim ~~69~~32, wherein the stretcher unit has the structure of Formula (1) below; the amino acid unit is valine-citrulline; and the spacer unit is a PAB group comprising the structure of Formula (2) below:



Formula (1)



Formula (2).

7134. The ~~method~~use of claim ~~69-32~~ or ~~7033~~, wherein the stretcher unit forms a bond with a sulfur atom of the antibody or antigen binding fragment thereof; and wherein the spacer unit is linked to MMAE via a carbamate group.

7235. The ~~method~~use of any one of claims 1 to ~~7134~~, wherein:

(i) the ADC comprises from 1 to 20 units of MMAE per antibody or antigen binding fragment thereof;

(ii) the ADC comprises from 1 to 10 units of MMAE per antibody or antigen binding fragment thereof;

(iii) the ADC comprises from 2 to 8 units of MMAE per antibody or antigen binding fragment thereof; or

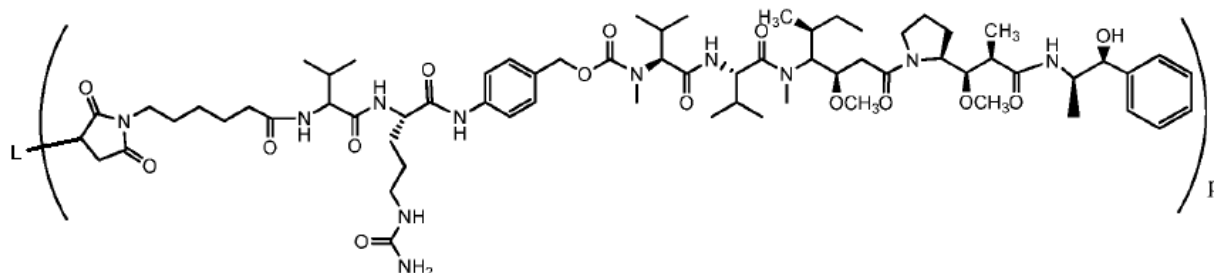
(iv) the ADC comprises from 3 to 5 units of MMAE per antibody or antigen binding fragment thereof.

~~73. The method of any one of claims 1 to 72, wherein the ADC comprises from 1 to 10 units of MMAE per antibody or antigen binding fragment thereof.~~

~~74. The method of any one of claims 1 to 73, wherein the ADC comprises from 2 to 8 units of MMAE per antibody or antigen binding fragment thereof.~~

~~75. The method of any one of claims 1 to 74, wherein the ADC comprises from 3 to 5 units of MMAE per antibody or antigen binding fragment thereof.~~

~~7636. The method of any one of claims 1 to 7335, wherein the ADC has the following structure:~~



wherein L- represents the antibody or antigen binding fragment thereof and p is from 1 to 10.

~~7737. The method of claim 7636, wherein:~~

~~(i) p is from 2 to 8~~

~~(ii) p is from 3 to 5;~~

~~(iii) p is from 3 to 4;~~

~~(iv) p is about 4; or~~

~~(v) the average p value of the effective amount of the antibody drug conjugates is about~~

~~3.8.~~

~~78. The method of claim 76 or 77, wherein p is from 3 to 5.~~

~~79. The method of any one of claims 76 to 78, wherein p is from 3 to 4.~~

~~80. The method of any one of claims 77 to 79, wherein p is about 4.~~

~~81. The method of any one of claims 76 to 79, wherein the average p value of the effective amount of the antibody drug conjugates is about 3.8.~~

~~8238. The method of any one of claims 1 to 8137, wherein:~~

~~(i) the ADC is to be administered at a dose of about 1 to about 10 mg/kg of the subject's body weight, about 1 to about 5 mg/kg of the subject's body weight, about 1 to about 2.5 mg/kg of the subject's body weight, or about 1 to about 1.25 mg/kg of the subject's body weight;~~

~~(ii) the ADC is to be administered at a dose of about 0.25 mg/kg, about 0.5 mg/kg, about 0.75 mg/kg, about 1.0 mg/kg, about 1.25 mg/kg, about 1.5 mg/kg, about 1.75 mg/kg, about 2.0 mg/kg, about 2.25 mg/kg, or about 2.5 mg/kg of the subject's body weight;~~

(iii) the ADC is to be administered at a dose of about 1 mg/kg of the subject's body weight; or

(iv) the ADC is to be administered at a dose of about 1.25 mg/kg of the subject's body weight.

~~83. The method of any one of claims 1 to 82, wherein the ADC is administered at a dose of about 0.25 mg/kg, about 0.5 mg/kg, about 0.75 mg/kg, about 1.0 mg/kg, about 1.25 mg/kg, about 1.5 mg/kg, about 1.75 mg/kg, about 2.0 mg/kg, about 2.25 mg/kg, or about 2.5 mg/kg of the subject's body weight.~~

~~84. The method of any one of claims 1 to 83, wherein the ADC is administered at a dose of about 1 mg/kg of the subject's body weight.~~

~~85. The method of any one of claims 1 to 83, wherein the ADC is administered at a dose of about 1.25 mg/kg of the subject's body weight.~~

~~86~~39. The ~~method~~use of any one of claims 1 to ~~85~~38, wherein the ADC is to be administered by:

(i) an intravenous (IV) injection or infusion;

(ii) an IV injection or infusion three times every four-week cycle;

(iii) an IV injection or infusion on Days 1, 8 and 15 of every four-week cycle;

(iv) an IV injection or infusion over about 30 minutes three times every four-week cycle;

or

(v) an IV injection or infusion over about 30 minutes on Days 1, 8 and 15 of every four-week cycle.

~~87. The method of any one of claims 1 to 86, wherein the ADC is administered by an IV injection or infusion three times every four-week cycle.~~

~~88. The method of any one of claims 1 to 87, wherein the ADC is administered by an IV injection or infusion on Days 1, 8 and 15 of every four-week cycle.~~

~~89. The method of any one of claims 1 to 88, wherein the ADC is administered by an IV injection or infusion over about 30 minutes three times every four-week cycle.~~

~~90. The method of any one of claims 1 to 89, wherein the ADC is administered by an IV injection or infusion over about 30 minutes on Days 1, 8 and 15 of every four-week cycle.~~

~~91~~40. The ~~method~~use of any one of claims 1 to ~~90~~39, wherein:

(i) the ADC is formulated in a pharmaceutical composition comprising L-histidine, polysorbate-20 (TWEEN-20), and trehalose dehydrate;

(ii) the ADC is formulated in a pharmaceutical composition comprising about 20 mM L-histidine, about 0.02% (w/v) TWEEN-20, about 5.5% (w/v) trehalose dihydrate, and hydrochloride, and wherein the pH of the pharmaceutical composition is about 6.0 at 25°C; or

(iii) the ADC is formulated in a pharmaceutical composition comprising about 9 mM histidine, about 11 mM histidine hydrochloride monohydrate, about 0.02% (w/v) TWEEN-20, and about 5.5% (w/v) trehalose dihydrate, and wherein the pH of the pharmaceutical composition is about 6.0 at 25°C.

92. — ~~The method of any one of claims 1 to 91, wherein the ADC is formulated in a pharmaceutical composition comprising about 20 mM L-histidine, about 0.02% (w/v) TWEEN-20, about 5.5% (w/v) trehalose dihydrate, and hydrochloride, and wherein the pH of the pharmaceutical composition is about 6.0 at 25°C.~~

93. — ~~The method of any one of claims 1 to 91, wherein the ADC is formulated in a pharmaceutical composition comprising about 9 mM histidine, about 11 mM histidine hydrochloride monohydrate, about 0.02% (w/v) TWEEN-20, and about 5.5% (w/v) trehalose dihydrate, and wherein the pH of the pharmaceutical composition is about 6.0 at 25°C.~~

9441. ~~The method~~use of any one of claims 1 to 9340, wherein the ADC is enfortumab vedotin (EV) or a biosimilar thereof, wherein the EV is to be administered at a dose of about 1.25 mg/kg of the subject's body weight, and wherein the dose is to be administered by an IV injection or infusion over about 30 minutes on Days 1, 8 and 15 of every four-week cycle.

9542. ~~The method~~use of any one of claims 1 to 9441, ~~whereby~~ wherein:

(i) a population of the subjects have a complete response following the treatment;

(ii) a population of the subjects have a partial response following the treatment;

(iii) a population of the subjects have a complete response or a partial response following the treatment; or

(iv) a population of the subjects have a stable disease following the treatment.

96. — ~~The method of any one of claims 1 to 94, wherein a population of the subjects have a partial response following the treatment.~~

97. — ~~The method of any one of claims 1 to 94, wherein a population of the subjects have a complete response or a partial response following the treatment.~~

98. — ~~The method of any one of claims 1 to 94, wherein a population of the subjects have a stable disease following the treatment.~~