CLAIMS

1. A stable aqueous pharmaceutical composition comprising a bispecific epidermal growth factor receptor (EGFR)/hepatocyte growth factor receptor (c-Met) antibody and a hyaluronidase, wherein the antibody comprises:
    a. a first heavy chain (HC1) comprising a HC1 variable region 1 (VH1) comprising the amino acid sequence of SEQ ID NO:13;
    b. a first light chain (LC1) comprising a light chain variable region 1 (VL1) comprising the amino acid sequence of SEQ ID NO:14;
    c. a second heavy chain (HC2) comprising a HC2 variable region 2 (VH2) comprising the amino acid sequence of SEQ ID NO:15;
    d. a second light chain (LC2) comprising a light chain variable region 2 (VL2) comprising the amino acid sequence of SEQ ID NO:16;

   and wherein the composition comprises about 1,050 mg to about 2,240 mg of the bispecific EGFR/c-Met antibody and about 13,000 U to about 28,000 U of the hyaluronidase.
2. The stable composition of claim 1 wherein the composition comprises about 1,050 mg of the bispecific EGFR/c-Met antibody.
3. The stable composition of claim 1 wherein the composition comprises about 1,400 mg of the bispecific EGFR/c-Met antibody.
4. The stable composition of claim 1 wherein the composition comprises about 1,575 mg of the bispecific EGFR/c-Met antibody.
5. The stable composition of claim 1 wherein the composition comprises about 1,600 mg of the bispecific EGFR/c-Met antibody.
6. The stable composition of claim 1 wherein the composition comprises about 2,100 mg of the bispecific EGFR/c-Met antibody.
7. The stable composition of claim 1 wherein the composition comprises about 2,240 mg of the bispecific EGFR/c-Met antibody.
8. A stable aqueous pharmaceutical composition comprising:
    a) about 144 mg/mL to about 176 mg/mL of a bispecific epidermal growth factor receptor (EGFR)/hepatocyte growth factor receptor (c-Met) antibody, the bispecific antibody comprising:

       a first heavy chain (HC1) comprising a HC1 variable region 1 (VH1);

a first light chain (LC1) comprising a light chain variable region 1 (VL1); a second heavy chain (HC2) comprising a HC2 variable region 2 (VH2); and a second light chain (LC2) comprising a light chain variable region 2 (VL2), wherein the VH1 comprises a heavy chain complementarity determining region 1 (HCDR1), a HCDR2 and a HCDR3 amino acid sequences of SEQ ID NOs: 1, 2, and 3, respectively; the VL1 comprises a light chain complementarity determining region 1 (LCDR1), a LCDR2 and a LCDR3 amino acid sequences of SEQ ID NOs: 4, 5 and 6, respectively, the VH2 comprises the HCDR1, the HCDR2 and the HCDR3 amino acid sequences of SEQ ID NOs: 7, 8 and 9, respectively; and the VL2 comprises the LCDR1, the LCDR2 and the LCDR3 amino acid sequences of SEQ ID NOs: 10, 11 and 12, respectively;

b) about 10 mM to about 50 mM of acetate and/or pharmaceutically acceptable acetate salt,
c) about 6.8% (w/v) to about 10.2% (w/v) of sucrose,
d) about 0.036% (w/v) to about 0.084% (w/v) of polysorbate 80 (PS80),
e) about to 0.8 mg/mL to about 1.2 mg/mL of methionine,
f) about 16 µg/mL to about 24 µg/mL of ethylenediaminetetraacetic acid (EDTA),
g) optionally, about 1,000 U/mL to about 3,000 U/mL of hyaluronidase; and
h) a pH from about 5.2 to about 6.2.

9. The stable aqueous pharmaceutical composition of claim 8, wherein the bispecific EGFR-cMet antibody comprises an HC1 variable region comprising the amino acid sequence of SEQ ID NO:13 and a LC1 variable region comprising the amino acid sequence of SEQ ID NO:14.
10. The stable aqueous pharmaceutical composition of claim 8 or claim 9, wherein the bispecific EGFR-cMet antibody comprises a HC2 variable region comprising the amino acid sequence of SEQ ID NO:15 and a LC2 variable region comprising the amino acid sequence of SEQ ID NO:16.
11. The stable aqueous pharmaceutical composition of any one of claims 8-10, wherein the HC1 comprises the amino acid sequence of SEQ ID NO:17 and the LC1 comprises the amino acid sequence of SEQ ID NO:18.
12. The stable aqueous pharmaceutical composition of any one of claims 8-11, wherein the HC2 comprises the amino acid sequence of SEQ ID NO: 19 and the LC2 comprises the amino acid sequence of SEQ ID NO:20.

13. The stable aqueous pharmaceutical composition of any one of claims 8-12, wherein the bispecific EGFR-cMet antibody is amivantamab or a biosimilar thereof.
14. The stable aqueous pharmaceutical composition of any one of claims 8-13, wherein the bispecific EGFR-cMet antibody has a concentration of about 160 mg/mL.
15. The stable aqueous pharmaceutical composition of any one of claims 8-14, wherein the acetate and/or pharmaceutically acceptable acetate salt has a concentration of about 30 mM.
16. The stable aqueous pharmaceutical composition of any one of claims 8-15, wherein the acetate and/or pharmaceutically acceptable acetate salt comprises glacial acetic acid and/or sodium acetate trihydrate.
17. The stable aqueous pharmaceutical composition of any one of claims 8-16, comprising about 8.5% (w/v) sucrose.
18. The stable aqueous pharmaceutical composition of any one of claims 8-17, comprising about 0.06% (w/v) PS80.
19. The stable aqueous pharmaceutical composition of any one of claims 8-18, wherein the methionine comprises L-methionine and has a concentration of about 1 mg/mL.
20. The stable aqueous pharmaceutical composition of any one of claims 8-19, wherein the EDTA has a concentration of about 20 µg/mL.
21. The stable aqueous pharmaceutical composition of any one of claims 8-20, wherein the pH is about 5.7.
22. The stable aqueous pharmaceutical composition of any one of claims 8-21, wherein the hyaluronidase is a human hyaluronidase, optionally a soluble human PH20 comprising the amino acid sequence of SEQ ID NO: 25.
23. The stable aqueous pharmaceutical composition of any one of claims 8-22, wherein the concentration rHuPH20 is about 1,000 U/mL to about 3,000 U/mL.
24. The stable aqueous pharmaceutical composition of any one of claims 8-23, wherein the concentration rHuPH20 is about 2,000 U/mL.
25. The stable aqueous pharmaceutical composition of any one of claims 8-24, wherein stability is defined based on color of solution, pH, turbidity, number of subvisible particles, percentage of aglycosylated heavy chain (AGHC), percentage of new peak(s), percentage of high molecular weight species (HMWS), percentage of low molecular weight species (LMWS), percentage of sum of acidic peaks, percentage of sum of basic peaks, protein concentration, percentage of EGFR binding activity,

percentage of cMet binding activity, percentage of PS80, optionally, percentage of rHuPH20 activity, or any combination thereof.

26. The stable aqueous pharmaceutical composition of any one of claims 8-25, wherein the total volume of the composition ranges from about 6mL to about 9mL.
27. The stable aqueous pharmaceutical composition of claim 24, wherein the total volume of the composition is about 7.1mL.
28. The stable aqueous pharmaceutical composition of claim 24, wherein the total volume of the composition is about 6.6mL.
29. The stable aqueous pharmaceutical composition of claim 24, wherein the total volume of the composition is about 8.75mL.
30. The stable aqueous pharmaceutical composition of any one of claims 1-29, comprising about 160 mg/mL of the bispecific EGFR-cMet antibody, about 30 mM acetate and/or pharmaceutically acceptable acetate salt, about 8.5% sucrose, and about 1 mg/mL L-methionine with polysorbate 80 to a final concentration of about 0.06% (w/v) and EDTA to a final concentration of about 20 µg/mL, wherein the stable aqueous pharmaceutical composition has pH about 5.7,
and wherein the bispecific EGFR-cMet antibody comprises a heavy chain 1 (HC1) comprising the amino acid sequence of SEQ ID NO:17, HC2 comprising the amino acid sequence of SEQ ID NO:19, a light chain 1 (LC1) comprising the amino acid sequence of SEQ ID NO:18, and a LC2 comprising the amino acid sequence of SEQ ID NO:20.
31. The stable aqueous pharmaceutical composition of any one of claims 1-29, comprising about 160 mg/mL of the bispecific EGFR-cMet antibody, about 30 mM acetate and/or pharmaceutically acceptable acetate salt, about 8.5% sucrose, about 1 mg/mL L-methionine with polysorbate 80 to a final concentration of about 0.06% (w/v) and EDTA to a final concentration of about 20 µg/mL, and rHuPH20 to a final concentration of about 2,000 U/mL, wherein the stable aqueous pharmaceutical composition has pH about 5.7,
and wherein the bispecific EGFR-cMet antibody comprises a heavy chain 1 (HC1) comprising the amino acid sequence of SEQ ID NO:17, HC2 comprising the amino acid sequence of SEQ ID NO:19, a light chain 1 (LC1) comprising the amino acid sequence of SEQ ID NO:18, and a LC2 comprising the amino acid sequence of SEQ ID NO:20.

32. Use of the pharmaceutical composition of any one of claims 1-31 in the manufacture of a medicament for treating cancer in a subject in need thereof, wherein the pharmaceutical composition is to be administered to the subject.
33. The use of claim 32, wherein the administration is subcutaneous.
34. The use of claims 30-31, wherein the cancer comprises lung cancer, squamous cell carcinoma of the head and neck (SCCHN), hepatocellular cancer (HCC), colorectal cancer (CRC), renal cell cancer (RCC), medullary thyroid cancer (MTC), gastroesophageal cancer (GEC), mesothelioma, breast cancer (BC) or ovarian cancer (OC).
35. The use of claims 32-33, wherein the cancer comprises non-small cell lung cancer (NSCLC).
36. A method for preparing a stable aqueous pharmaceutical composition of a bispecific antibody targeting EGFR and cMet, the bispecific antibody targeting EGFR and cMet comprising a first heavy chain (HC1) comprising a HC1 variable region 1 (VH1); a first light chain (LC1) comprising a light chain variable region 1 (VL1);a second heavy chain (HC2) comprising a HC2 variable region 2 (VH2); and a second light chain (LC2) comprising a light chain variable region 2 (VL2), wherein the VH1 comprises a heavy chain complementarity determining region 1 (HCDR1), a HCDR2 and a HCDR3 comprising amino acid sequences of SEQ ID NOs: 1, 2, and 3, respectively; the VL1 comprises a light chain complementarity determining region 1 (LCDR1), a LCDR2 and a LCDR3 comprising amino acid sequences of SEQ ID NOs: 4, 5 and 6, respectively; the VH2 comprises HCDR1, HCDR2 and HCDR3 amino acid sequences of SEQ ID NOs: 7, 8 and 9, respectively; and the VL2 comprises LCDR1, LCDR2 and LCDR3 amino acid sequences of SEQ ID NOs: 10, 11 and 12, respectively; the method comprising:
    combining a composition comprising about 160 mg/mL of the bispecific antibody, about 30 mM acetate and/or pharmaceutically acceptable acetate salt, about 8.5% sucrose, and about 1 mg/mL L-methionine with polysorbate 80 to a final concentration of about 0.06% (w/v) and EDTA to a final concentration of about 20 µg/mL, optionally rHuPH20 to a final concentration of about 2,000 U/mL, wherein the stable aqueous pharmaceutical composition has about pH 5.7.
37. The method of claim 36, wherein the bispecific EGFR-cMet antibody comprises an HC1 variable region comprising the amino acid sequence of SEQ ID NO:13 and a LC1 variable region comprising the amino acid sequence of SEQ ID NO:14.

38. The method of any one of claims 36-37, wherein the bispecific EGFR-cMet antibody comprises a HC2 variable region comprising the amino acid sequence of SEQ ID NO:15 and a LC2 variable region comprising the amino acid sequence of SEQ ID NO:16.
39. The method of any one of claims 36-38, wherein the antibody comprises a heavy chain 1 (HC1) comprising the amino acid sequence of SEQ ID NO:17 and a light chain 1 (LC1) comprising the amino acid sequence of SEQ ID NO:18.
40. The method of any one of claims 36-39, wherein the antibody comprises a HC2 comprising the amino acid sequence of SEQ ID NO:19 and a LC2 comprising the amino acid sequence of SEQ ID NO:20.
41. The method of any one of claims 36-40, wherein the antibody is amivantamab or a biosimilar thereof.
42. A kit comprising the stable aqueous pharmaceutical composition of any one of claims 1-31 and instructions for use thereof.
43. An article of manufacture comprising a container holding a stable aqueous pharmaceutical composition in accordance with any one of claims 1-31.
44. The article of manufacture according to claim 43, wherein the container is a vial with a stopper pierceable by a syringe.
45. The article of manufacture according to claim 44, wherein the vial is a single-use vial.
46. A pharmaceutical composition of any one of claims 1-31 for use in the treatment of cancer.
47. Use of the stable aqueous pharmaceutical composition of claim 1 in the manufacture of a medicament for reducing infusion-related reactions in a subject treated with amivantamab, wherein the pharmaceutical formulation is to be subcutaneously administered to the subject.