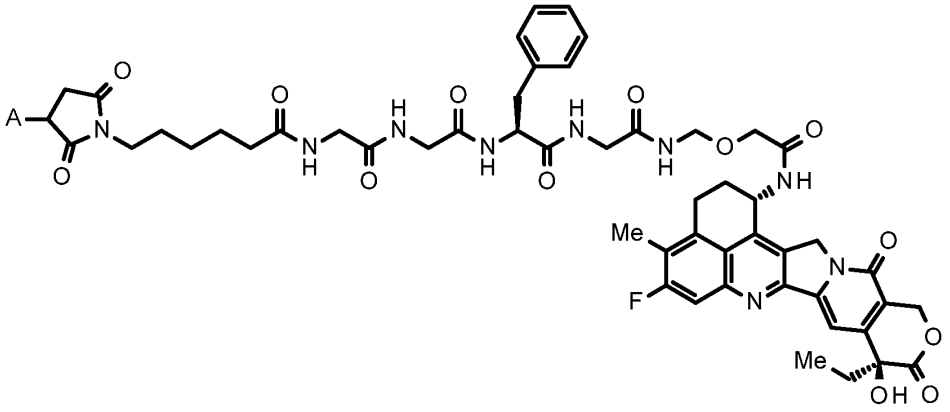
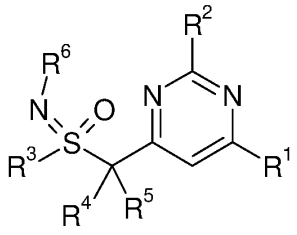
CLAIMS

1. A pharmaceutical product comprising an anti-HER2 antibody-drug conjugate and an ATR inhibitor for administration in combination, wherein the anti-HER2 antibody-drug conjugate is an antibody-drug conjugate in which a drug-linker represented by the following formula:



wherein A represents the connecting position to an antibody, is conjugated to an anti-HER2 antibody via a thioether bond.

2. The pharmaceutical product according to claim 1, wherein the ATR inhibitor is a compound represented by the following formula (I):

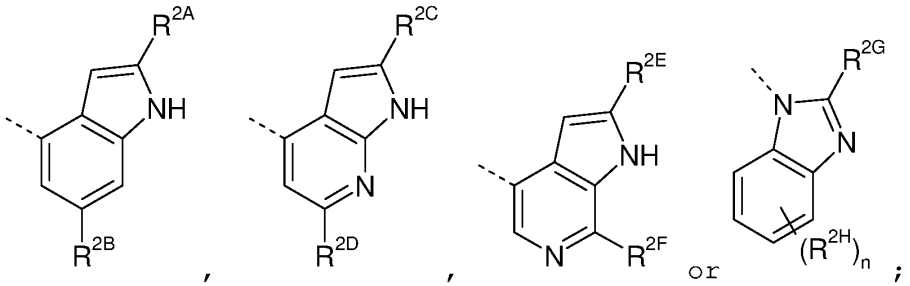


(I)

wherein:

R^{1} is selected from morpholin-4-yl and 3-methylmorpholin-4-yl;

R^{2} is



, , or ;

n is 0 or 1;

R^{2A}, **R**^{2C}, **R**^{2E} and **R**^{2F} each independently are hydrogen or methyl;

R^{2B} and **R**^{2D} each independently are hydrogen or methyl;

R^{2G} is selected from $-NHR^{7}$ and $-NHCOR^{8}$;

R^{2H} is fluoro;

R^{3} is methyl;

R^{4} and **R**^{5} are each independently hydrogen or methyl, or **R**^{4} and **R**^{5} together with the atom to which they are attached form Ring A;

Ring A is a C_{3-6}cycloalkyl or a saturated 4-6 membered heterocyclic ring containing one heteroatom selected from O and N;

R^{6} is hydrogen;

R^{7} is hydrogen or methyl;

R^{8} is methyl,

or a pharmaceutically acceptable salt thereof.

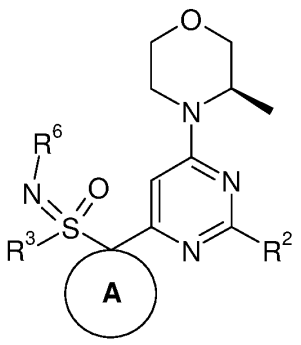
3. The pharmaceutical product according to claim 2 wherein, in formula (I), $\mathbf{R^4}$ and $\mathbf{R^5}$ together with the atom to which they are attached form Ring A, and Ring A is a C_{3-6}cycloalkyl or a saturated 4–6 heterocyclic ring containing one heteroatom selected from O and N.

4. The pharmaceutical product according to claim 2 or claim 3 wherein, in formula (I), Ring A is a cyclopropyl, tetrahydropyranyl or piperidinyl ring.

5. The pharmaceutical product according to any one of claims 2 to 4 wherein, in formula (I), $\mathbf{R^{2A}}$ is hydrogen; $\mathbf{R^{2B}}$ is hydrogen; $\mathbf{R^{2C}}$ is hydrogen; $\mathbf{R^{2D}}$ is hydrogen; $\mathbf{R^{2E}}$ is hydrogen; and $\mathbf{R^{2F}}$ is hydrogen.

6. The pharmaceutical product according to any one of claims 2 to 5 wherein, in formula (I), R^1 is 3-methylmorpholin-4-yl.

7. The pharmaceutical product according to any one of claims 2 to 6 wherein the compound of formula (I) is a compound of formula (Ia):



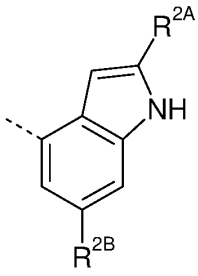
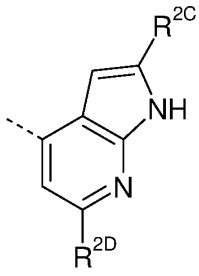
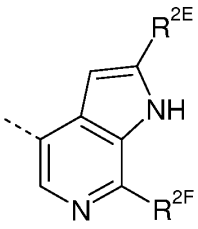
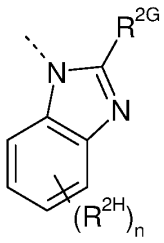
(Ia)

or a pharmaceutically acceptable salt thereof.

8. The pharmaceutical product according to claim 7 wherein, in formula (Ia):

Ring A is cyclopropyl ring;

R^2 is

;

n is 0 or 1;

R^{2A} is hydrogen;

R^{2B} is hydrogen;

R^{2C} is hydrogen;

R^{2D} is hydrogen;

R^{2E} is hydrogen;

R^{2F} is hydrogen;

R^{2G} is $-NHR^7$;

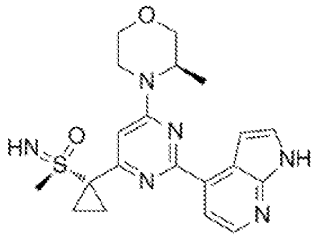
R^{2H} is fluoro;

R^{3} is a methyl group;

R^{6} is hydrogen; and

R^{7} is hydrogen or methyl.

9. The pharmaceutical product according to claim 2, wherein the ATR inhibitor is AZD6738 represented by the following formula:



or a pharmaceutically acceptable salt thereof.

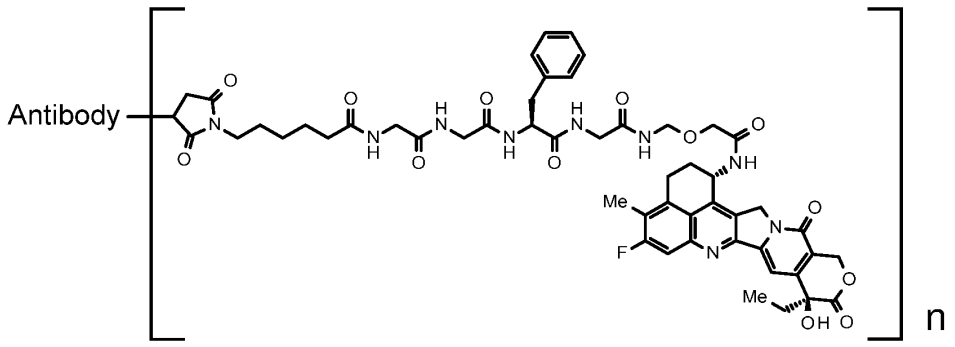
10. The pharmaceutical product according to any one of claims 1 to 9, wherein the anti-HER2 antibody is an antibody comprising a heavy chain comprising CDRH1 consisting of an amino acid sequence represented by SEQ ID NO: 3, CDRH2 consisting of an amino acid sequence represented by SEQ ID NO: 4 and CDRH3 consisting of an amino acid sequence represented by SEQ ID NO: 5, and a light chain comprising CDRL1 consisting of an amino acid sequence represented by SEQ ID NO: 6, CDRL2 consisting of an amino acid sequence consisting of amino acid residues 1 to 3 of SEQ ID NO: 7 and CDRL3 consisting of an amino acid sequence represented by SEQ ID NO: 8.

11. The pharmaceutical product according to any one of claims 1 to 9, wherein the anti-HER2 antibody is an antibody comprising a heavy chain comprising a heavy chain variable region consisting of an amino acid sequence represented by SEQ ID NO: 9 and a light chain comprising a light chain variable region consisting of an amino acid sequence represented by SEQ ID NO: 10.

12. The pharmaceutical product according to any one of claims 1 to 9, wherein the anti-HER2 antibody is an antibody comprising a heavy chain consisting of an amino acid sequence represented by SEQ ID NO: 1 and a light chain consisting of an amino acid sequence represented by SEQ ID NO: 2.

13. The pharmaceutical product according to any one of claims 1 to 9, wherein the anti-HER2 antibody is an antibody comprising a heavy chain consisting of an amino acid sequence represented by SEQ ID NO: 11 and a light chain consisting of an amino acid sequence represented by SEQ ID NO: 2.

14. The pharmaceutical product according to any one of claims 1 to 13, wherein the anti-HER2 antibody-drug conjugate is represented by the following formula:



wherein 'Antibody' indicates the anti-HER2 antibody conjugated to the drug-linker via a thioether bond, and n indicates an average number of units of the drug-linker conjugated per antibody molecule in the antibody-drug conjugate, wherein n is in the range of from 7 to 8.

15. The pharmaceutical product according to any one of claims 1 to 14, wherein the anti-HER2 antibody-drug conjugate is trastuzumab deruxtecan (DS-8201).

16. The pharmaceutical product according to any one of claims 1 to 15, wherein the product is a composition comprising the anti-HER2 antibody-drug conjugate and the ATR inhibitor, for simultaneous administration.

17. The pharmaceutical product according to any one of claims 1 to 15, wherein the product is a combined preparation comprising the anti-HER2 antibody-drug conjugate and the ATR inhibitor, for sequential or simultaneous administration.

18. The pharmaceutical product according to any one of claims 1 to 17, wherein the product is for treating cancer.

19. The pharmaceutical product according to claim 18, wherein the cancer is at least one selected from the group consisting of breast cancer, gastric cancer, colorectal cancer, lung cancer, esophageal cancer, head-and-neck cancer, esophagogastric junction adenocarcinoma, biliary tract cancer, Paget's disease, pancreatic cancer, ovarian cancer, uterine carcinosarcoma, urothelial cancer, prostate cancer, bladder cancer, gastrointestinal stromal tumor, digestive tract stromal tumor, uterine cervix cancer, squamous cell carcinoma, peritoneal cancer, liver cancer, hepatocellular cancer, corpus uteri carcinoma, kidney cancer, vulval cancer, thyroid cancer, penis cancer, leukemia, malignant lymphoma, plasmacytoma, myeloma, glioma, glioblastoma multiforme, osteosarcoma, sarcoma, and melanoma.

20. The pharmaceutical product according to claim 18, wherein the cancer is breast cancer.

21. The pharmaceutical product according to claim 20, wherein the breast cancer has a HER2 status score of IHC 3+.

22. The pharmaceutical product according to claim 20, wherein the breast cancer is HER2 low-expressing breast cancer.

23. The pharmaceutical product according to claim 20, wherein the breast cancer has a HER2 status score of IHC 2+.

24. The pharmaceutical product according to claim 20, wherein the breast cancer has a HER2 status score of IHC 1+.

25. The pharmaceutical product according to claim 20, wherein the breast cancer has a HER2 status score of IHC >0 and <1+.

26. The pharmaceutical product according to claim 20, wherein the breast cancer is triple-negative breast cancer.

27. The pharmaceutical product according to claim 18, wherein the cancer is gastric cancer.

28. The pharmaceutical product according to claim 18, wherein the cancer is colorectal cancer.

29. The pharmaceutical product according to claim 18, wherein the cancer is lung cancer.

30. The pharmaceutical product according to claim 29, wherein the lung cancer is non-small cell lung cancer.

31. The pharmaceutical product according to claim 18, wherein the cancer is pancreatic cancer.

32. The pharmaceutical product according to claim 18, wherein the cancer is ovarian cancer.

33. The pharmaceutical product according to claim 18, wherein the cancer is prostate cancer.

34. The pharmaceutical product according to claim 18, wherein the cancer is kidney cancer.

35. The pharmaceutical product according to claim 18, wherein cancer cells of the cancer are SLFN11-deficient.

36. The pharmaceutical product according to claim 18, wherein SLFN11 expression is lower in the cancer cells of a patient relative to the patient's SLFN11-expressing non-cancer cells.

37. A pharmaceutical product as defined in any one of claims 1 to 17, for use in treating cancer.

38. The pharmaceutical product for the use according to claim 37, wherein the cancer is at least one selected from the group consisting of breast cancer, gastric cancer, colorectal cancer, lung cancer, esophageal cancer, head-and-neck cancer, esophagogastric junction adenocarcinoma, biliary tract cancer, Paget's disease, pancreatic cancer, ovarian cancer, uterine carcinosarcoma, urothelial cancer, prostate cancer, bladder cancer, gastrointestinal stromal tumor, digestive tract stromal tumor, uterine cervix cancer, squamous cell carcinoma, peritoneal cancer, liver cancer, hepatocellular cancer, corpus uteri carcinoma, kidney cancer, vulval cancer, thyroid cancer, penis cancer, leukemia, malignant lymphoma, plasmacytoma, myeloma, glioma, glioblastoma multiforme, osteosarcoma, sarcoma, and melanoma.

39. The pharmaceutical product for the use according to claim 37, wherein the cancer is breast cancer.

40. The pharmaceutical product for the use according to claim 39, wherein the breast cancer has a HER2 status score of IHC 3+.

41. The pharmaceutical product for the use according to claim 39, wherein the breast cancer is HER2 low-expressing breast cancer.

42. The pharmaceutical product for the use according to claim 39, wherein the breast cancer has a HER2 status score of IHC 2+.

43. The pharmaceutical product for the use according to claim 39, wherein the breast cancer has a HER2 status score of IHC 1+.

44. The pharmaceutical product for the use according to claim 39, wherein the breast cancer has a HER2 status score of IHC >0 and <1+.

45. The pharmaceutical product for the use according to claim 39, wherein the breast cancer is triple-negative breast cancer.

46. The pharmaceutical product for the use according to claim 37, wherein the cancer is gastric cancer.

47. The pharmaceutical product for the use according to claim 37, wherein the cancer is colorectal cancer.

48. The pharmaceutical product for the use according to claim 37, wherein the cancer is lung cancer.

49. The pharmaceutical product for the use according to claim 48, wherein the lung cancer is non-small cell lung cancer.

50. The pharmaceutical product for the use according to claim 37, wherein the cancer is pancreatic cancer.

51. The pharmaceutical product for the use according to claim 37, wherein the cancer is ovarian cancer.

52. The pharmaceutical product for the use according to claim 37, wherein the cancer is prostate cancer.

53. The pharmaceutical product for the use according to claim 37, wherein the cancer is kidney cancer.

54. The pharmaceutical product for the use according to claim 37, wherein cancer cells of the cancer are SLFN11-deficient.

55. The pharmaceutical product for the use according to claim 37, wherein SLFN11 expression is lower in the cancer cells of a patient relative to the patient's SLFN11-expressing non-cancer cells.

56. Use of an anti-HER2 antibody-drug conjugate or an ATR inhibitor in the manufacture of a medicament for administration of the anti-HER2 antibody-drug conjugate

and the ATR inhibitor in combination, wherein the anti-HER2 antibody-drug conjugate and the ATR inhibitor are as defined in any one of claims 1 to 15, for treating cancer.

57. The use according to claim 56, wherein the cancer is at least one selected from the group consisting of breast cancer, gastric cancer, colorectal cancer, lung cancer, esophageal cancer, head-and-neck cancer, esophagogastric junction adenocarcinoma, biliary tract cancer, Paget's disease, pancreatic cancer, ovarian cancer, uterine carcinosarcoma, urothelial cancer, prostate cancer, bladder cancer, gastrointestinal stromal tumor, digestive tract stromal tumor, uterine cervix cancer, squamous cell carcinoma, peritoneal cancer, liver cancer, hepatocellular cancer, corpus uteri carcinoma, kidney cancer, vulval cancer, thyroid cancer, penis cancer, leukemia, malignant lymphoma, plasmacytoma, myeloma, glioma, glioblastoma multiforme, osteosarcoma, sarcoma, and melanoma.

58. The use according to claim 56, wherein the cancer is breast cancer.

59. The use according to claim 58, wherein the breast cancer has a HER2 status score of IHC 3+.

60. The use according to claim 58, wherein the breast cancer is HER2 low-expressing breast cancer.

61. The use according to claim 58, wherein the breast cancer has a HER2 status score of IHC 2+.

62. The use according to claim 58, wherein the breast cancer has a HER2 status score of IHC 1+.

63. The use according to claim 58, wherein the breast cancer has a HER2 status score of IHC >0 and <1+.

64. The use according to claim 58, wherein the breast cancer is triple-negative breast cancer.

65. The use according to claim 56, wherein the cancer is gastric cancer.

66. The use according to claim 56, wherein the cancer is colorectal cancer.

67. The use according to claim 56, wherein the cancer is lung cancer.

68. The use according to claim 67, wherein the lung cancer is non-small cell lung cancer.

69. The use according to claim 56, wherein the cancer is pancreatic cancer.

70. The use according to claim 56, wherein the cancer is ovarian cancer.

71. The use according to claim 56, wherein the cancer is prostate cancer.

72. The use according to claim 56, wherein the cancer is kidney cancer.

73. The use according to claim 56, wherein cancer cells of the cancer are SLFN11-deficient.

74. The use according to claim 56, wherein SLFN11 expression is lower in the cancer cells of a patient relative to the patient's SLFN11-expressing non-cancer cells.

75. The use according to any one of claims 56 to 74 wherein the medicament is a composition comprising the anti-HER2 antibody-drug conjugate and the ATR inhibitor, for simultaneous administration.

76. The use according to any one of claims 56 to 74 wherein the medicament is a combined preparation comprising the anti-HER2 antibody-drug conjugate and the

ATR inhibitor, for sequential or simultaneous administration.

77. A method of treating cancer comprising administering an anti-HER2 antibody-drug conjugate and an ATR inhibitor as defined in any one of claims 1 to 15 in combination to a subject in need thereof.

78. The method according to claim 77, wherein the cancer is at least one selected from the group consisting of breast cancer, gastric cancer, colorectal cancer, lung cancer, esophageal cancer, head-and-neck cancer, esophagogastric junction adenocarcinoma, biliary tract cancer, Paget's disease, pancreatic cancer, ovarian cancer, uterine carcinosarcoma, urothelial cancer, prostate cancer, bladder cancer, gastrointestinal stromal tumor, digestive tract stromal tumor, uterine cervix cancer, squamous cell carcinoma, peritoneal cancer, liver cancer, hepatocellular cancer, corpus uteri carcinoma, kidney cancer, vulval cancer, thyroid cancer, penis cancer, leukemia, malignant lymphoma, plasmacytoma, myeloma, glioma, glioblastoma multiforme, osteosarcoma, sarcoma, and melanoma.

79. The method according to claim 77, wherein the cancer is breast cancer.

80. The method according to claim 79, wherein the breast cancer has a HER2 status score of IHC 3+.

81. The method according to claim 79, wherein the breast cancer is HER2 low-expressing breast cancer.

82. The method according to claim 79, wherein the breast cancer has a HER2 status score of IHC 2+.

83. The method according to claim 79, wherein the breast cancer has a HER2 status score of IHC 1+.

84. The method according to claim 79, wherein the breast cancer has a HER2 status score of IHC >0 and <1+.

85. The method according to claim 79, wherein the breast cancer is triple-negative breast cancer.

86. The method according to claim 77, wherein the cancer is gastric cancer.

87. The method according to claim 77, wherein the cancer is colorectal cancer.

88. The method according to claim 77, wherein the cancer is lung cancer.

89. The method according to claim 88, wherein the lung cancer is non-small cell lung cancer.

90. The method according to claim 77, wherein the cancer is pancreatic cancer.

91. The method according to claim 77, wherein the cancer is ovarian cancer.

92. The method according to claim 77, wherein the cancer is prostate cancer.

93. The method according to claim 77, wherein the cancer is kidney cancer.

94. The method according to claim 77, wherein cancer cells of the cancer are SLFN11-deficient.

95. The method according to claim 77, wherein SLFN11 expression is lower in the cancer cells of a patient relative to the patient's SLFN11-expressing non-cancer cells.

96. The method according to any one of claims 77 to 95, wherein the method comprises administering the anti-HER2 antibody-drug conjugate and the ATR inhibitor sequentially.

97. The method according to any one of claims 77 to 95, wherein the method comprises administering the anti-HER2 antibody-drug conjugate and the ATR inhibitor simultaneously.