

Claims

1. A method for the prognosis of overall survival, cancer recurrence or response to treatment for a patient suffering from cancer, the method comprising:

5 (a) examining a sample from the patient to determine whether the patient is human epidermal growth factor receptor 2 (HER2) positive or negative based on a predetermined level of HER2; and

(b) measuring WW domain-binding protein 2 (WBP2) levels in the patient's sample,

10 wherein a result in step (a) and a result in step (b) provides a prognosis of overall survival, cancer recurrence or response to treatment for the patient.

2. The method of claim 1, wherein the results in steps (a) and (b) are compared to a set of predetermined expression level results from a comparison
15 population.

3. The method of claim 2, wherein the comparison population is stratified into a plurality of subgroups determining the aggressiveness of a cancer.
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4. The method of claim 3, wherein each subgroup is referenced from a reference group of HER2 negative patients comprising WBP2 expression below a predetermined level.

25 5. The method of claim 4, wherein a sample with WBP2 expression levels above the predetermined level and a HER2 positive patient provides the prognosis that the patient has an approximate 4 to 5 times lower chance of overall survival compared to the reference group.

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6. The method of claim 4, wherein a sample with WBP2 expression levels below the predetermined level and a HER2 positive patient provides the prognosis that the patient has an approximate 1 to 2 times lower chance of overall survival compared to the reference group.

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7. The method of claim 4, wherein a sample with WBP2 expression levels above the predetermined level and a HER2 negative patient provides the prognosis that the patient has an approximate 2 to 3 times lower chance of overall survival compared to the reference group.

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8. The method of claim 4, wherein a sample with WBP2 expression levels above the predetermined level and a HER2 positive patient provides the prognosis that the patient has an approximate 2 to 3 times higher chance of cancer recurrence compared to the reference group.

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9. The method of claim 4, wherein a sample with WBP2 expression levels below the predetermined level and a HER2 positive patient provides the prognosis that the patient has an approximate 1 times higher chance of cancer recurrence compared to the reference group.

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10. The method of claim 4, wherein a sample with WBP2 expression levels above the predetermined level and a HER2 negative patient provides the prognosis that the patient has an approximate 1 to 2 times higher chance of cancer recurrence compared to the reference group.

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11. The method of any one of claims 1 to 5, wherein a sample with WBP2 expression levels above the predetermined level and a HER2 positive result predicts that the patient is likely to respond to treatment.

12. The method of any one of claims 1 to 5, wherein a sample with WBP2 expression levels below the predetermined level and a HER2 positive result predicts that the patient is less likely to respond to treatment.

5 13. The method of claim 11 or 12, wherein the treatment comprises a HER2 antagonist.

14. The method of claim 13, wherein the HER2 antagonist comprises Herceptin (Trastuzumab), Pertuzumab, Lapatinib, Lapatinib in combination with
10 capecitabine, Trastuzumab emtansin, Ado-trastuzumab, Neratinib, Amrubicin or Dasatinib.

15. The method of any one of the preceding claims, wherein the WBP2 levels in the patient's sample are measured with at least one probe
15 adapted to target a WBP2 protein.

16. The method of claim 15, wherein the probe is an antibody.

17. The method of claim 15, wherein the probe is an aptamer.
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18. The method of claim 15, wherein the probe is a peptide.

19. The method of any one of the preceding claims, wherein the cancer is breast cancer.
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20. The method of any one of the preceding claims, wherein the method is an *in vitro* method.

21. The method of any one of claims 1 to 19, wherein the method is
30 an *in vivo* method.

22. A kit for identifying in a sample the amount of human epidermal growth factor receptor 2 (HER2) and the amount of WW domain-binding protein 2 (WBP2), the kit comprising:

- (a) at least one first probe adapted to detect and measure a human
5 epidermal growth factor receptor 2 (HER2) level in the sample to determine whether the sample is HER2 positive or HER2 negative; and
- (b) at least one second probe adapted to detect and measure WW domain-binding protein 2 (WBP2) levels in the sample.

10 23. The kit of claim 22, wherein the first probe is adapted to target a HER2 gene.

24. The kit of claim 22, wherein the second probe is adapted to target
15 a WBP2 gene.

25. The kit of claim 22, wherein the first probe is adapted target a
HER2 protein.

26. The kit of claim 22, wherein the second probe is adapted to target
20 a WBP2 protein.

27. The kit of claim 25 or 26, wherein the first and second probe is an
antibody.

25 28. The kit of claim 25 or 26, wherein the first and second probe is an
aptamer.

29. The kit of claim 25 or 26, wherein the first and second probe is a
peptide.

30. The kit of any one of claims 22 to 29, the kit further comprises written instructions for examining a sample to determine a prognosis of overall survival, cancer recurrence or response to treatment for the patient.

5 31. An *in vitro* method for determining the prognosis of overall survival, cancer recurrence or response to treatment, the method comprising:

(a) measuring the level of human epidermal growth factor receptor 2 (HER2) in a sample;

10 (b) measuring the level of WW domain-binding protein 2 (WBP2) in the sample; and

(c) determining whether the level of HER2 and WBP2 are above or below a predetermined level wherein a result in step (c) provides a prognosis of overall cancer survival, cancer recurrence or response to cancer treatment.

15 32 A method for the prognosis of survival or response to a treatment that targets EGFR or HER2 in a patient suffering from cancer comprising the steps of measuring WBP2 levels in a patient sample.