

- 2 Fig. S1. The degree of miRNA functional enrichment in stages II–III.
- 3 (a). The degree of miRNA functional enrichments in stage II.

- 4 (b). The degree of miRNA functional enrichments in stage III.
- 5 (c). The degree of miRNA functional enrichments in stage IV.

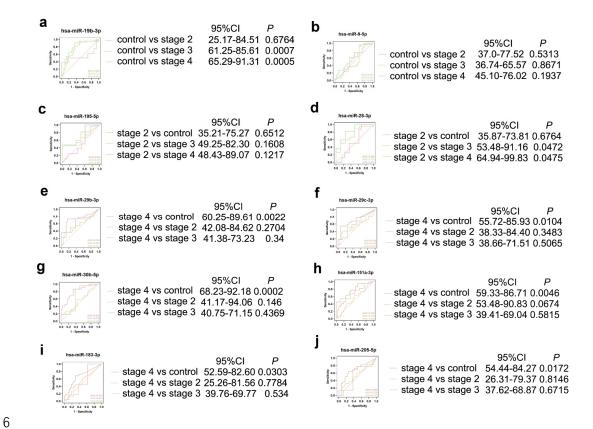


Fig. S2. ROC curve results of 10 miRNAs that did not show significant diagnostic

8 value.

- 9 (a). ROC curves of hsa-miR-19b-3p differentiated stage II, III and IV from control.
- 10 **(b)**. ROC curve of hsa-miR-9-5p differentiated stage II, III and IV from control.
- (c). ROC curve of hsa-miR-195-5p differentiated stage II from control, stage III and
- 12 stage IV.
- 13 (d). ROC curve of hsa-miR-28-3p differentiated stage II from control, stage III and
- 14 stage IV.
- 15 (e). ROC curve of hsa-miR-29b-3p differentiated stage IV from control, stage II and
- 16 stage III.
- 17 (f). ROC curve of hsa-miR-29c-3p differentiated stage IV from control, stage II and
- 18 stage III.

- 19 (g). ROC curve of hsa-miR-30b-5p differentiated stage IV from control, stage II and
- 20 stage III.
- 21 (h). ROC curve of hsa-miR-151a-3p differentiated stage IV from control, stage II and
- 22 stage III.
- 23 (i). ROC curve of hsa-miR-183-3p differentiated stage IV from control, stage II and
- 24 stage III.
- 25 (j). ROC curve of hsa-miR-205-5p differentiated stage IV from control, stage II and
- 26 stage III.

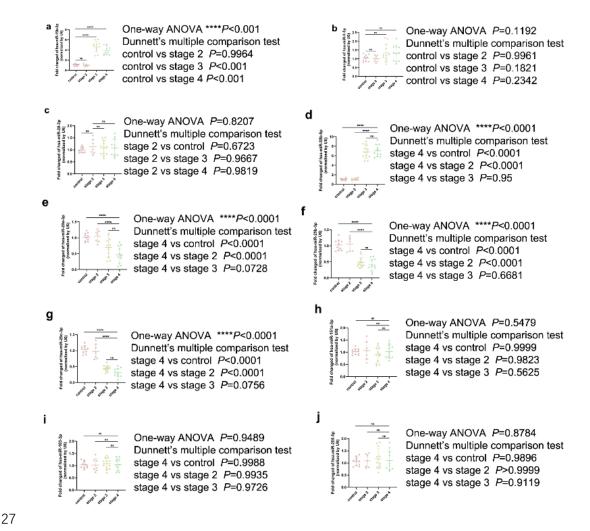


Fig. S3. qRT-PCR results of 10 miRNAs whose specific expression failed to be

29 verified by this analysis.

- 30 (a). The expressions of hsa-miR-19b-3p in stage II, stage III and stage IV.
- 31 **(b)**. The expressions of hsa-miR-9-5p in stage II, stage III and stage IV.
- 32 (c). The expressions of hsa-miR-28-3p in stage II, stage III and stage IV.
- 33 (d). The expressions of hsa-miR-30b-5p in stage II, stage III and stage IV.
- 34 (e). The expressions of hsa-miR-29a-3p in stage II, stage III and stage IV.
- 35 **(f)**. The expressions of hsa-miR-29b-3p in stage II, stage III and stage IV.
- 36 (g). The expressions of hsa-miR-29c-3p in stage II, stage III and stage IV.
- 37 (h). The expressions of hsa-miR-151a-3p in stage II, stage III and stage IV.
- 38 (i). The expressions of hsa-miR-183-3p in stage II, stage III and stage IV.
- 39 (j). The expressions of hsa-miR-205-5p in stage II, stage III and stage IV. Data were
- analyzed by one-way ANOVA with Dunnett's multiple comparisons test (mean \pm SD;
- 41 n = 40, ****P < 0.0001).

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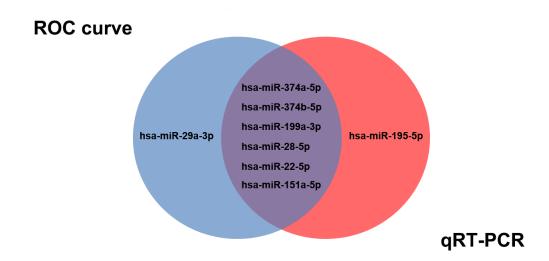


Fig. S4. Venn diagram of miRNAs validated by ROC curve and qRT-PCR.

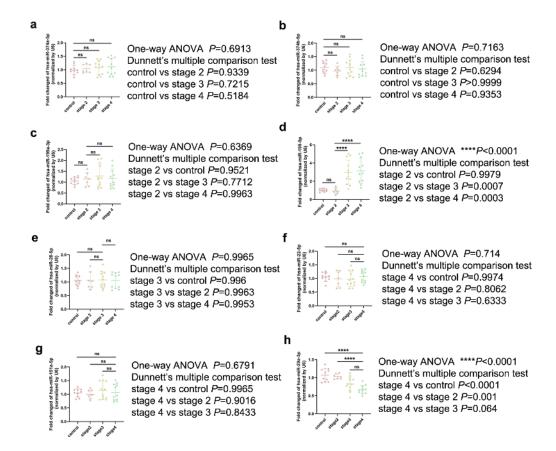


Fig. S5. The expression of miRNAs validated by ROC curves and qRT-PCR assay, in

48 the sera of PD participants.

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- 49 (a). The expressions of hsa-miR-374a-5p in stage II, stage III and stage IV.
- 50 **(b)**. The expressions of hsa-miR-374b-5p in stage II, stage III and stage IV.
- 51 (c). The expressions of hsa-miR-199a-3p in stage II, stage III and stage IV.
- 52 (d). The expressions of hsa-miR-195-5p in stage II, stage III and stage IV.
- (e). The expressions of hsa-miR-28-5p in stage II, stage III and stage IV.
- 54 (f). The expressions of hsa-miR-22-5p in stage II, stage III and stage IV.
- 55 (g). The expressions of hsa-miR-151a-5p in stage II, stage III and stage IV.
- 56 (h). The expressions of hsa-miR-29a-3p in stage II, stage III and stage IV. Data were
- analyzed by one-way ANOVA with Dunnett's multiple comparisons test (mean \pm SD;
- 58 n = 40; **** P < 0.0001).

Table S1 The primer sequences used by qRT-PCR

miRNA	Primer sequences
U6	5'-GCTCGCTTCGGCAGCA-3' (forward)
U6	5'-AACGCTTCACGAATTTGCGT-3' (reverse)
hsa-miR-374a-5p	5'-TTATAATACAACCTGATAAGTG-3' (forward)
hsa-miR-374b-5p	5'-ATATAATACAACCTGCTAAGTG-3' (forward)
hsa-miR-19-3p	5'-TGTGCAAATCCATGCAAAACTGA-3' (forward)
hsa-miR-30c-5p	5'-TGTAAACATCCTACACTCTCAGC-3' (forward)
hsa-miR-199a-3p	5'-ACAGTAGTCTGCACATTGGTTA-3' (forward)
hsa-miR-195-5p	5'-TAGCAGCACAGAAATATTGGC-3' (forward)
hsa-miR-5100	5'-TTCAGATCCCAGCGGTGCCTCT-3' (forward)
hsa-miR-28-5p	5'-AAGGAGCTCACAGTCTATTGAG-3' (forward)
hsa-miR-378a-5p	5'-CTCCTGACTCCAGGTCCTGTGT-3' (forward)
hsa-miR-22-5p	5'-AGTTCTTCAGTGGCAAGCTTTA-3' (forward)
hsa-miR-151a-5p	5'-TCGAGGAGCTCACAGTCTAGT-3' (forward)
hsa-miR-4443	5'-TTGGAGGCGTGGGTTTT-3' (forward)
hsa-miR-30b-5p	5'-TGTAAACATCCTACACTCAGCT-3' (forward)
hsa-miR-29a-3p	5'-TAGCACCATCTGAAATCGGTTA-3' (forward)
hsa-miR-29b-3p	5'-TAGCACCATTTGAAATCAGTGTT-3' (forward)
hsa-miR-29c-3p	5'-TAGCACCATTTGAAATCGGTTA-3' (forward)
hsa-miR-205-5p	5'-TCCTTCATTCCACCGGAGTCTG-3' (forward)
hsa-miR-183-3p	5'-GTGAATTACCGAAGGGCCATAA-3' (forward)
hsa-miR-21-5p	5'-TAGCTTATCAGACTGATGTTGA-3' (forward)
hsa-miR-25-5p	5'-AGGCGGAGACTTGGGCAATTG-3' (forward)
hsa-miR-1260b	5'-ATCCCACCACTGCCACCAT-3' (forward)
hsa-miR-151a-3p	5'-CTAGACTGAAGCTCCTTGAGG-3' (forward)
hsa-miR-9-5p	5'-TCTTTGGTTATCTAGCTGTATGA-3' (forward)
hsa-miR-28-3p	5'-CACTAGATTGTGAGCTCCTGGA-3' (forward)
hsa-miR-4492	5'-GAGGCTGAGCGCTCGTCA-3' (forward)

The downstream primers of miRNAs come from universal primers in the kit (miScript II RT kit, QIAGEN) except U6.