

## Supplementary Information

# 2-Aminopyridine Nucleobase Improves Triple Helical Recognition of RNA and DNA when Used Instead of Pseudoisocytosine in Peptide Nucleic Acids

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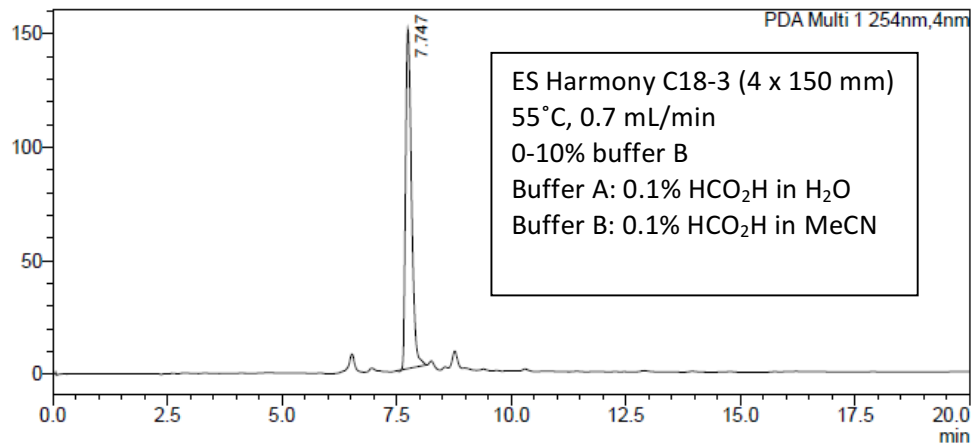
Department of Chemistry, Binghamton University, The State University of New York, Binghamton, New York 13902, United States.

**Table S1.** Deconvolution of LCMS analysis of synthesized PNAs.

	<b>Mass<sub>(calc)</sub></b>	<b>Mass (M+6, M+5...)</b>
<b>PNA1</b>	2349	471, 588, 784, 1176
<b>PNA2</b>	2381	477, 596, 795, 1191
<b>PNA3</b>	2451	491, 614, 818, 1173
<b>PNA4</b>	2366	474, 593, 790, 1184
<b>PNA5</b>	3413	684, 854, 1139, 1708
<b>PNA6</b>	3516	704, 880, 1173, 1759
<b>PNA7</b>	3833	768, 959, 1279, 1918
<b>PNA8</b>	3816	764, 955, 1273, 1909
<b>PNA9</b>	3833	768, 959, 1279, 1918
<b>PNA10</b>	5008	836, 1002, 1253, 1670
<b>PNA11</b>	6944	1131, 1319, 1583, 1979
<b>PNA12</b>	6893	1124, 1311, 1573, 1966

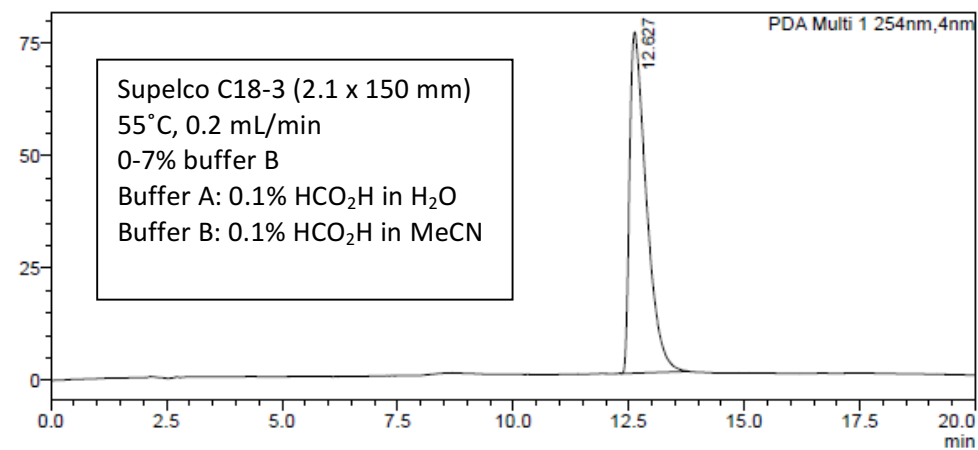
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mAU



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MS Spectrum

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Spectrum Mode:Single 12.883(624) Base Peak:470.90(2952107)  
BG Mode:None Segment 1 - Event 1

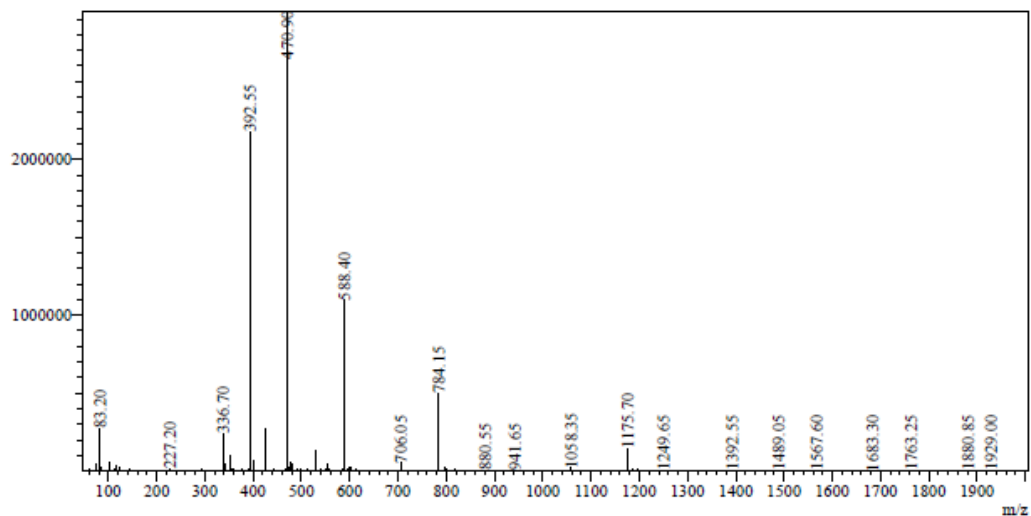
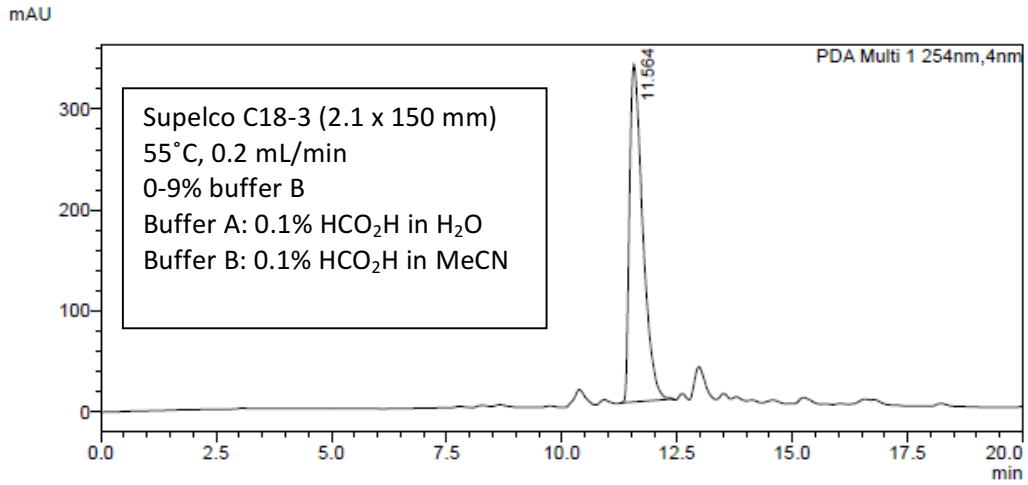
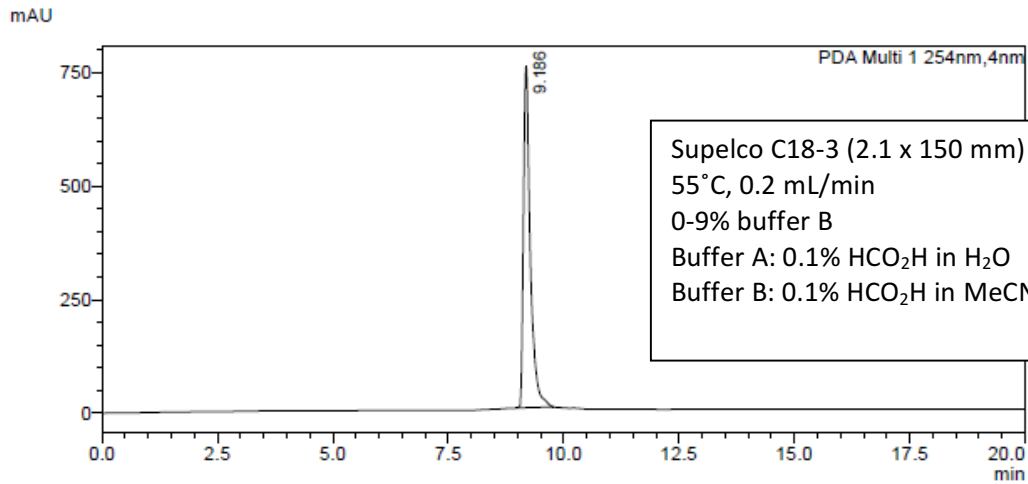


Figure S1. Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for PNA1.

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<Chromatogram>



MS Spectrum

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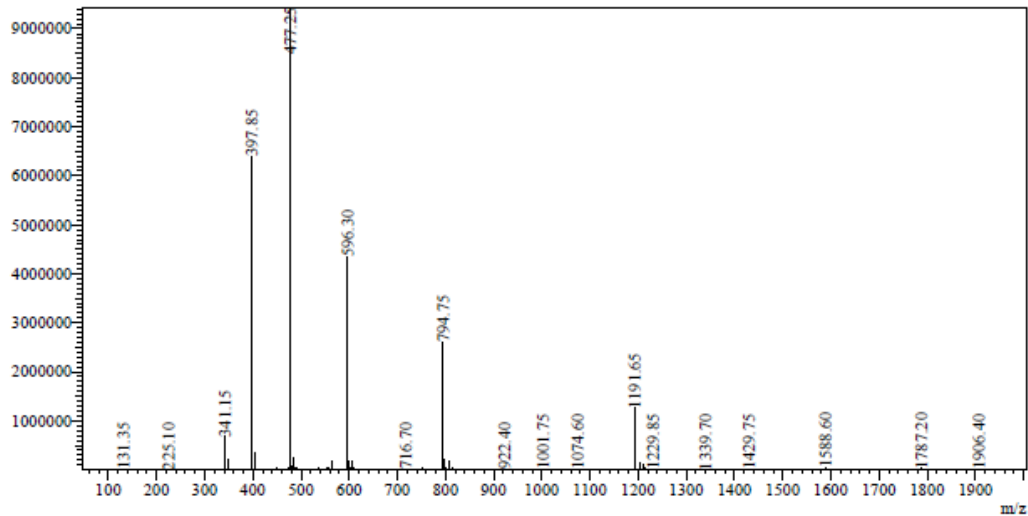
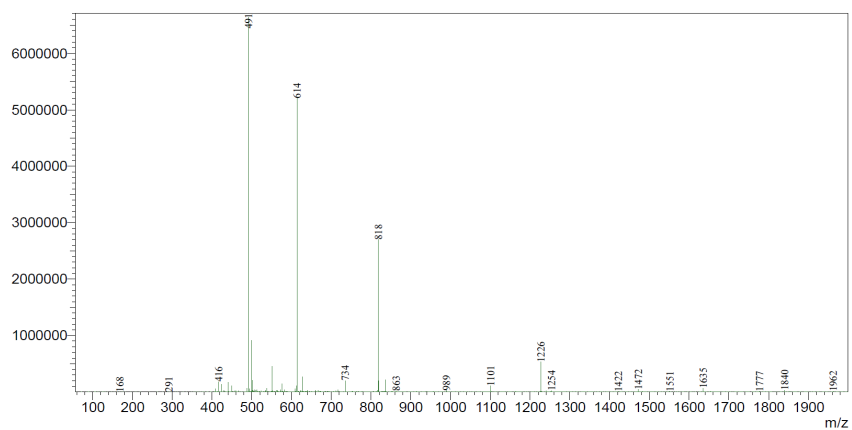
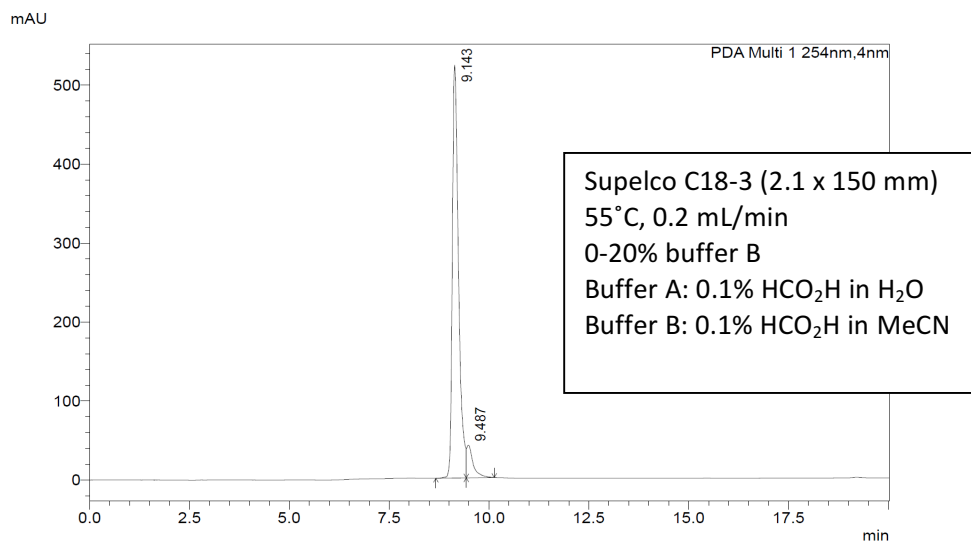
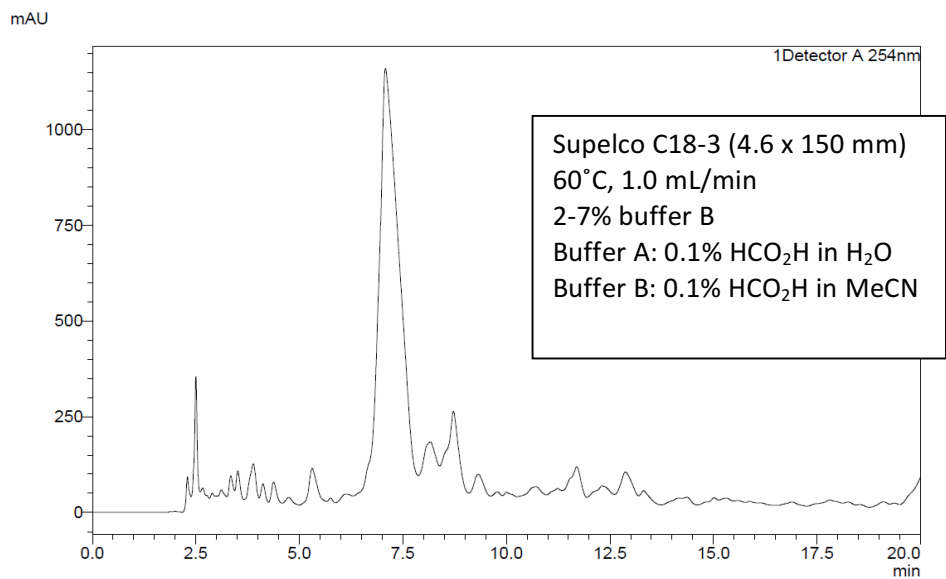
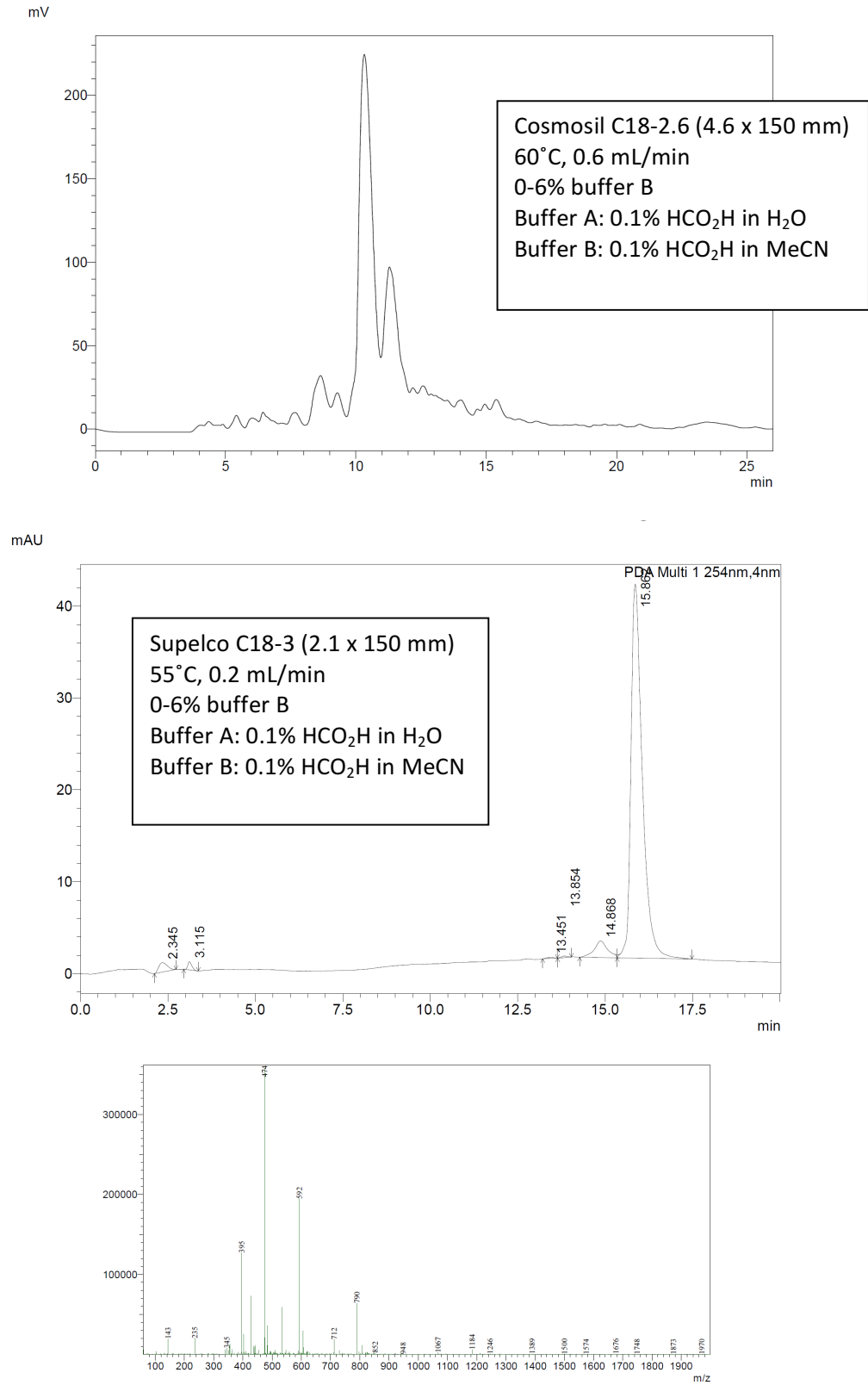


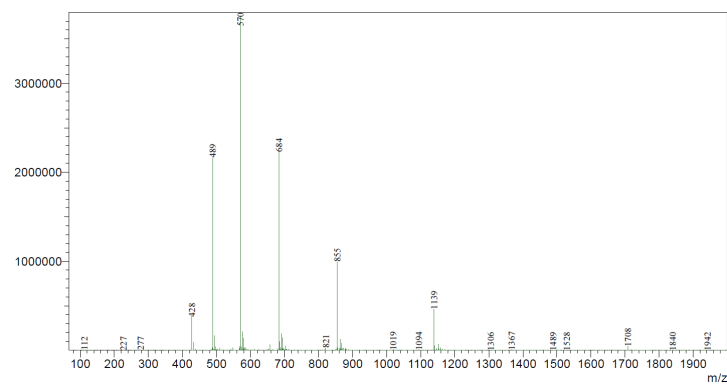
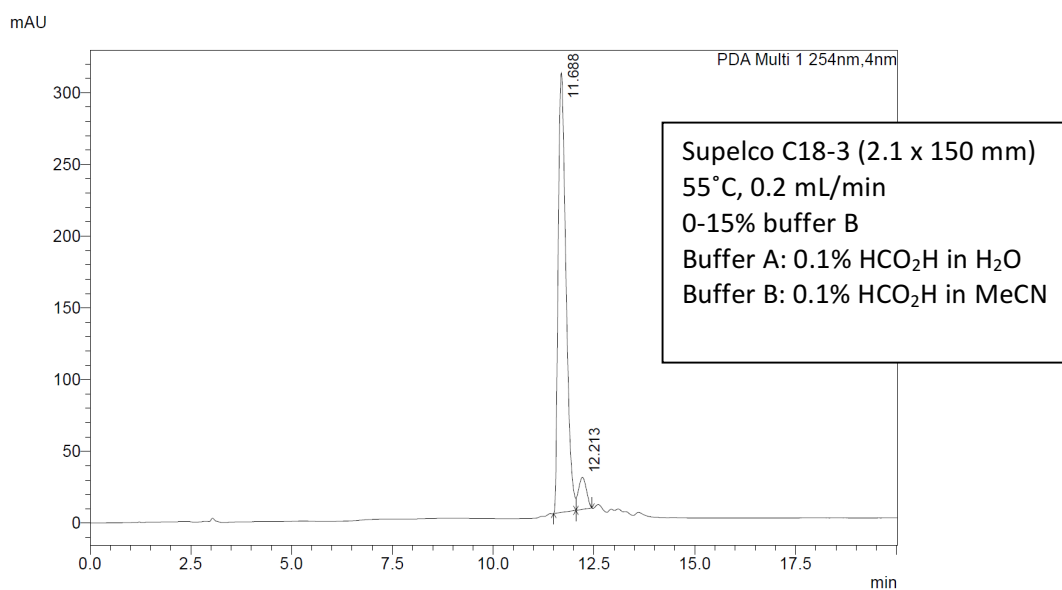
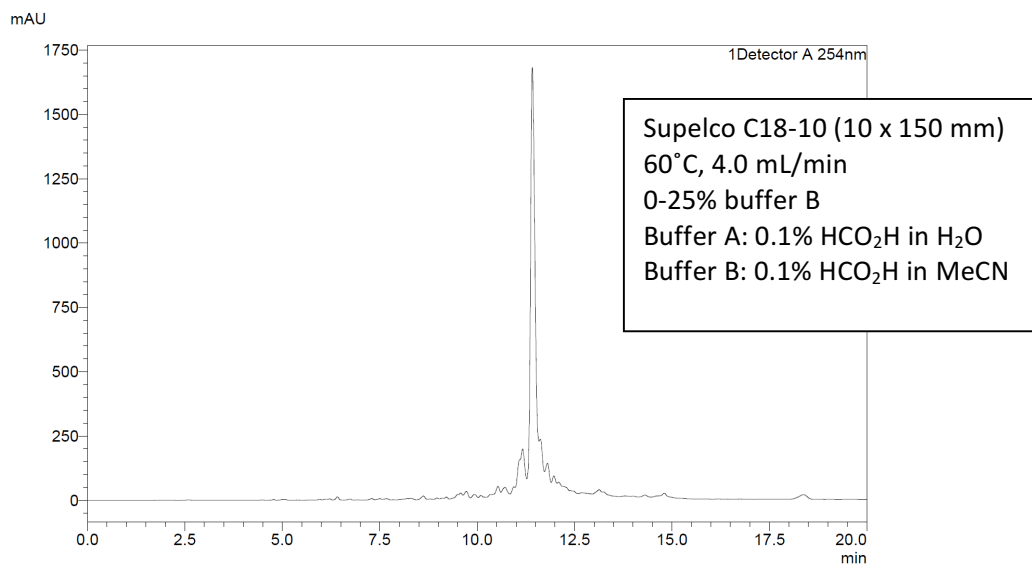
Figure S2. Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for PNA2.



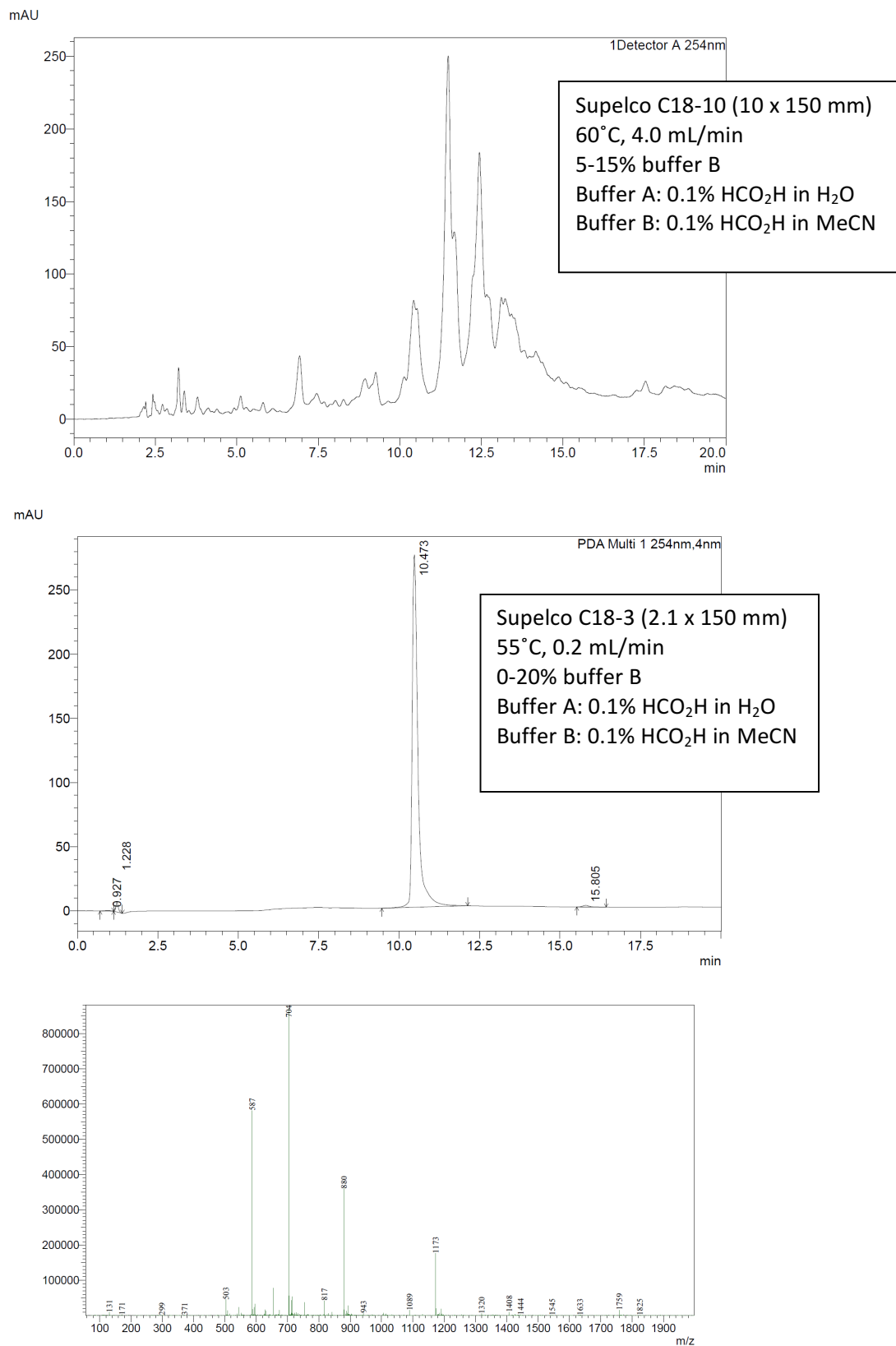
**Figure S3.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA3**.



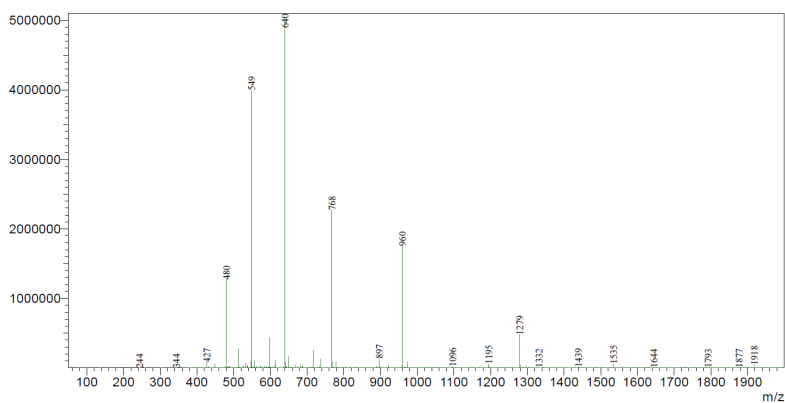
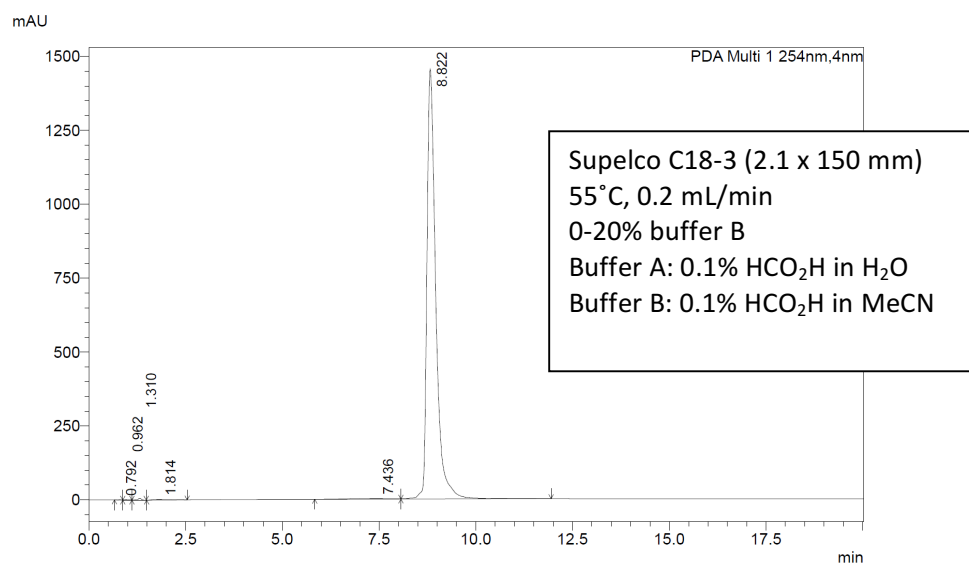
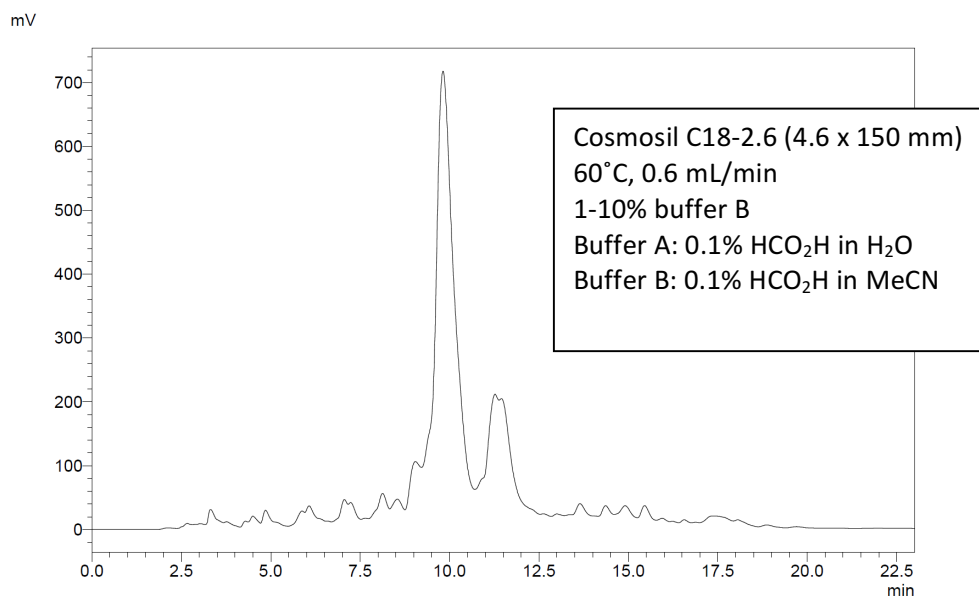
**Figure S4.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA4**.



**Figure S5.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for PNA5.

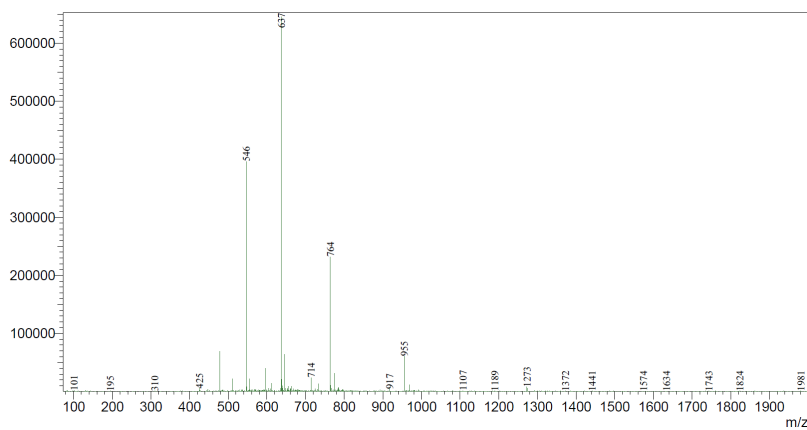
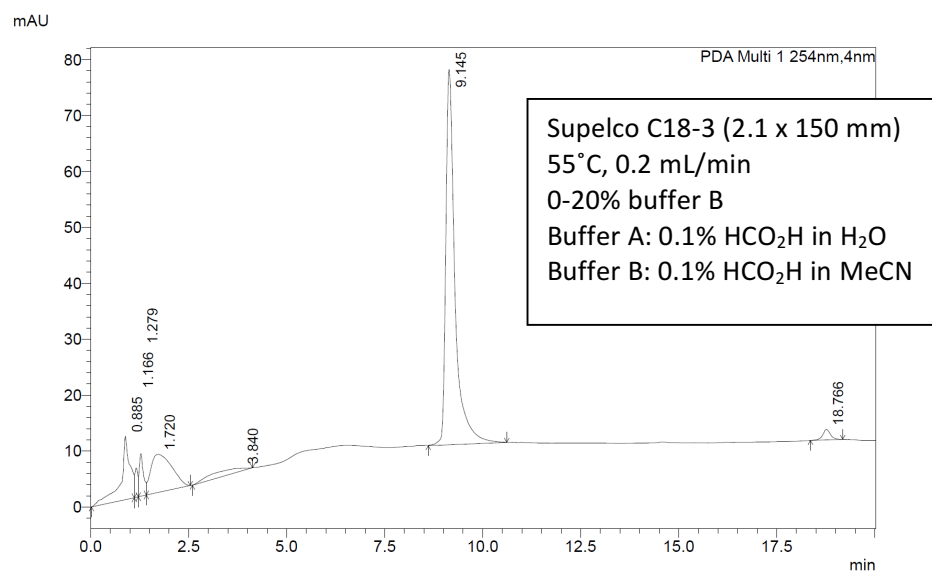
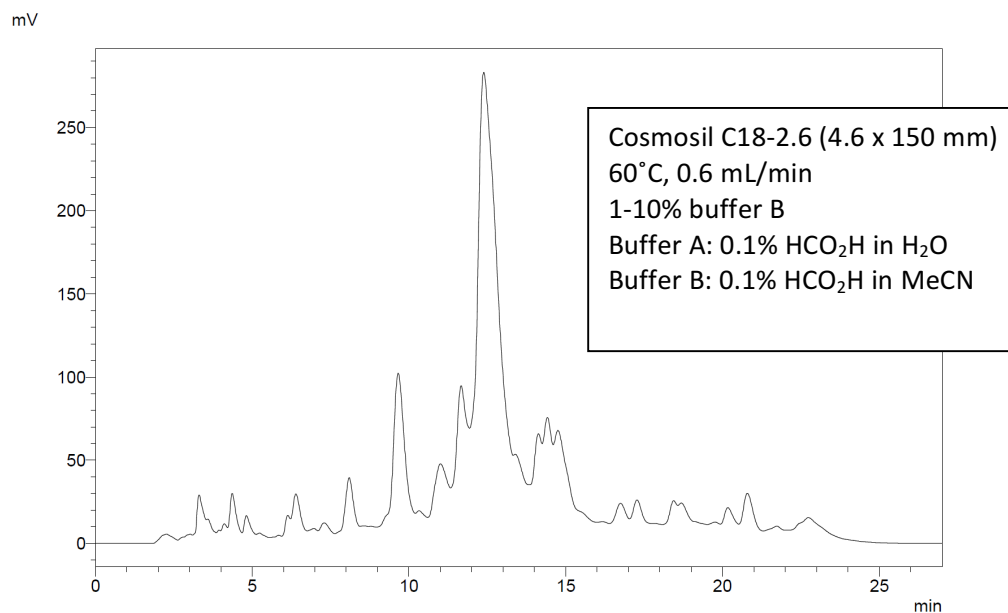


**Figure S6.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA6**.

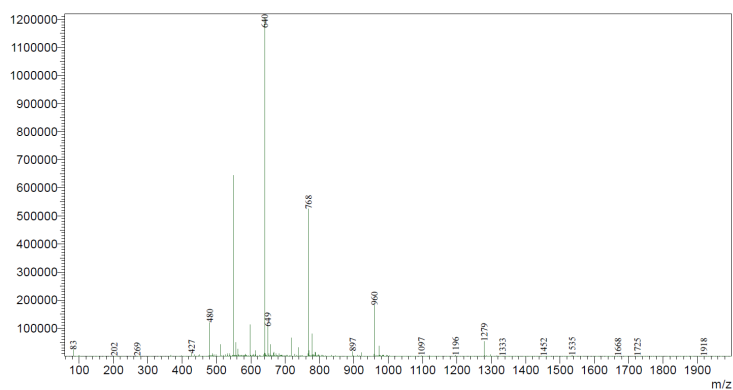
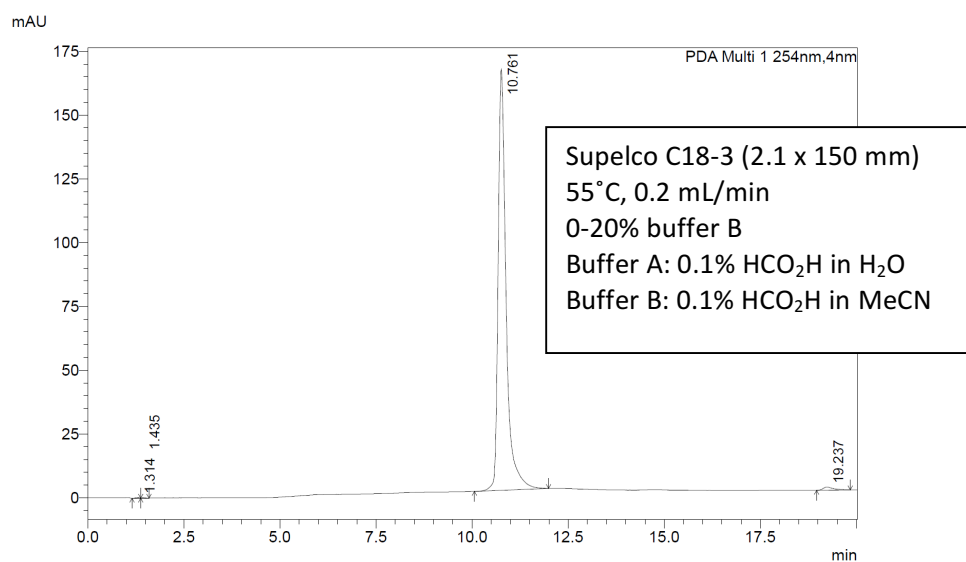
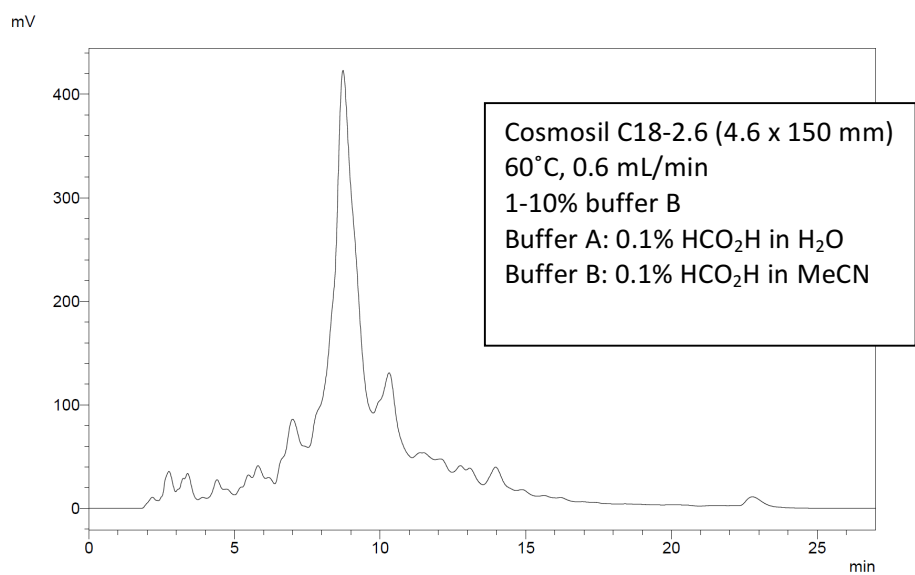


**Figure S7.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA7**.

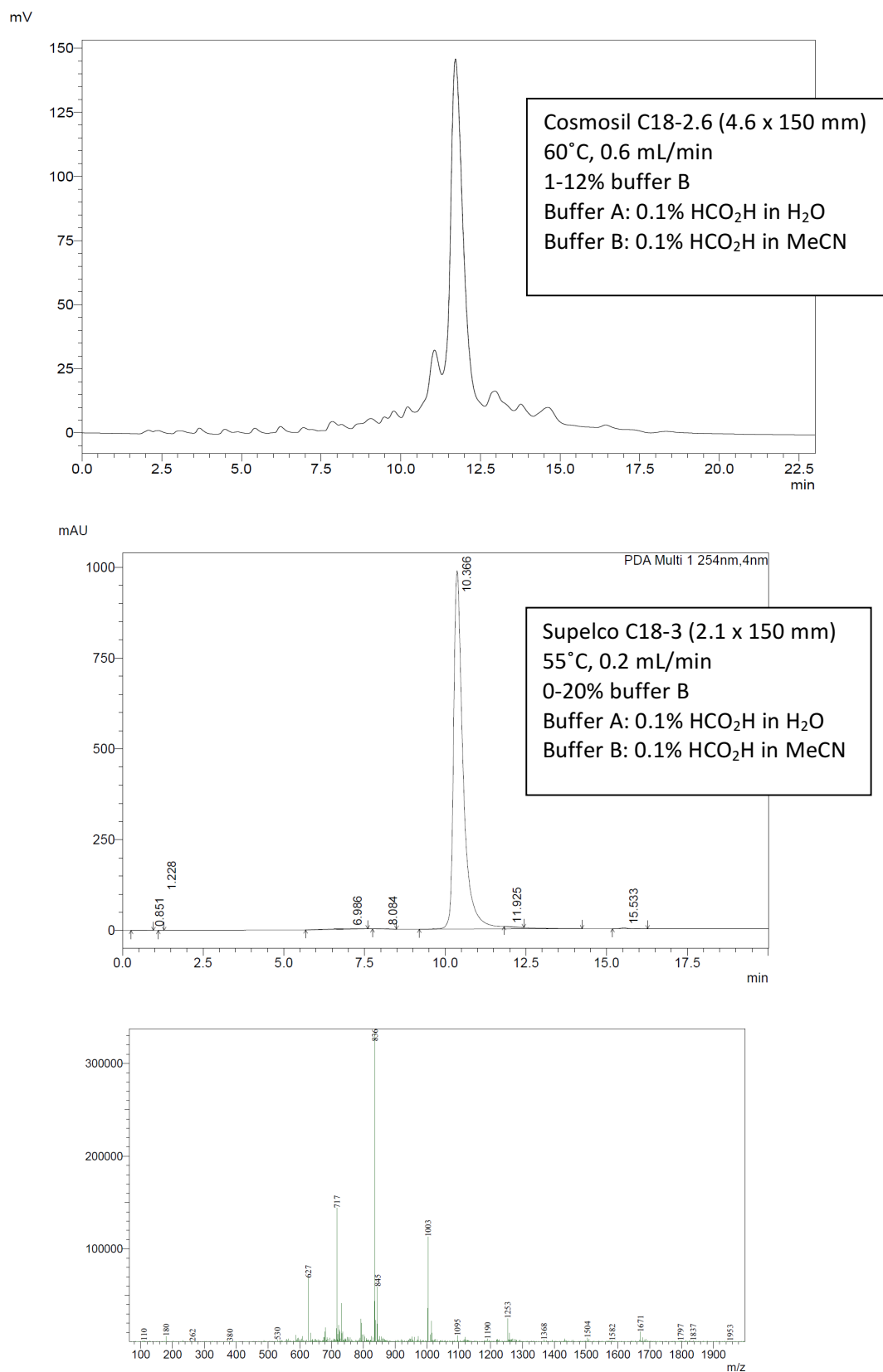




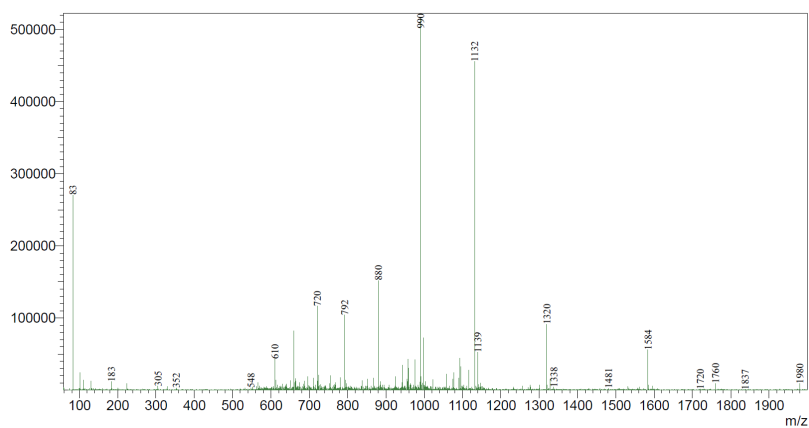
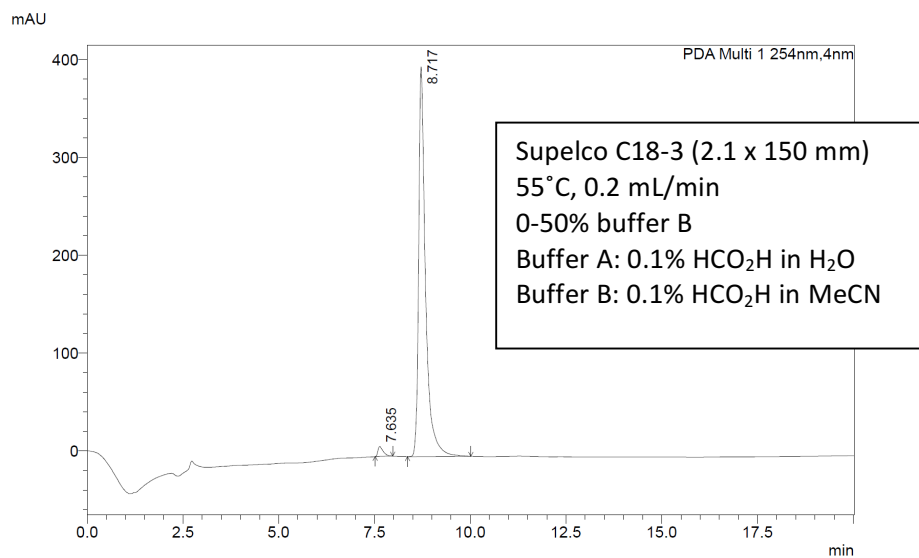
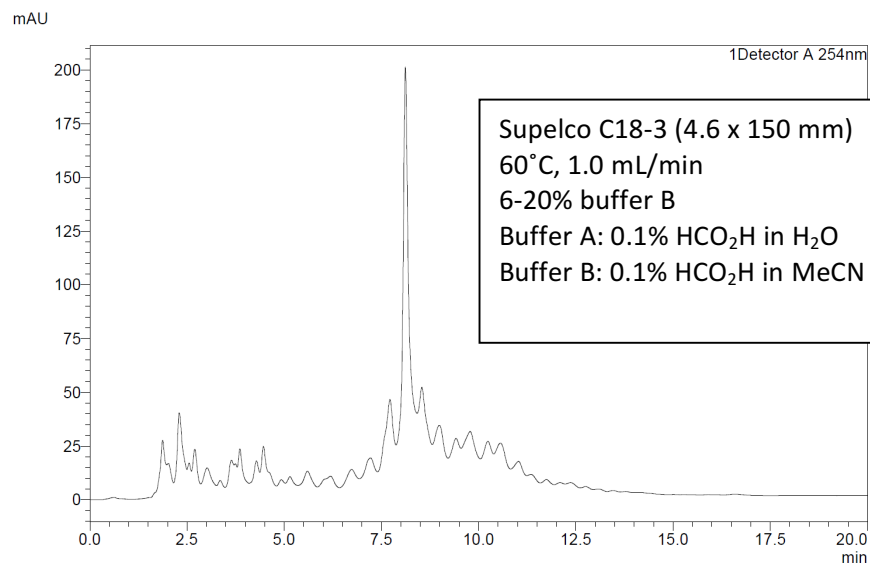
**Figure S9.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA8**.



**Figure S8.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA9**.



**Figure S10.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA10**.



**Figure S11.** Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for **PNA11**.

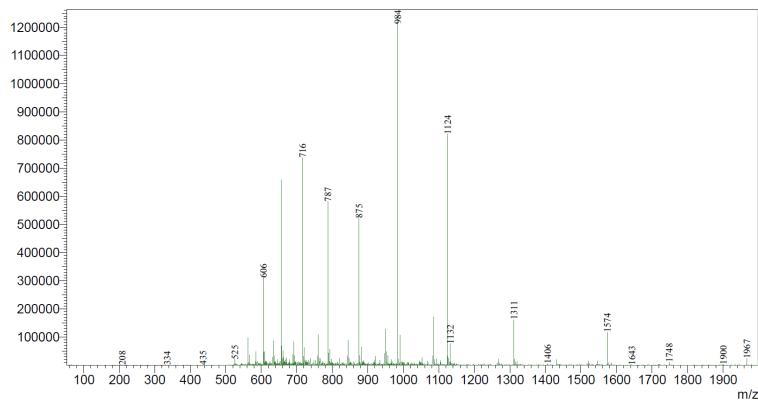
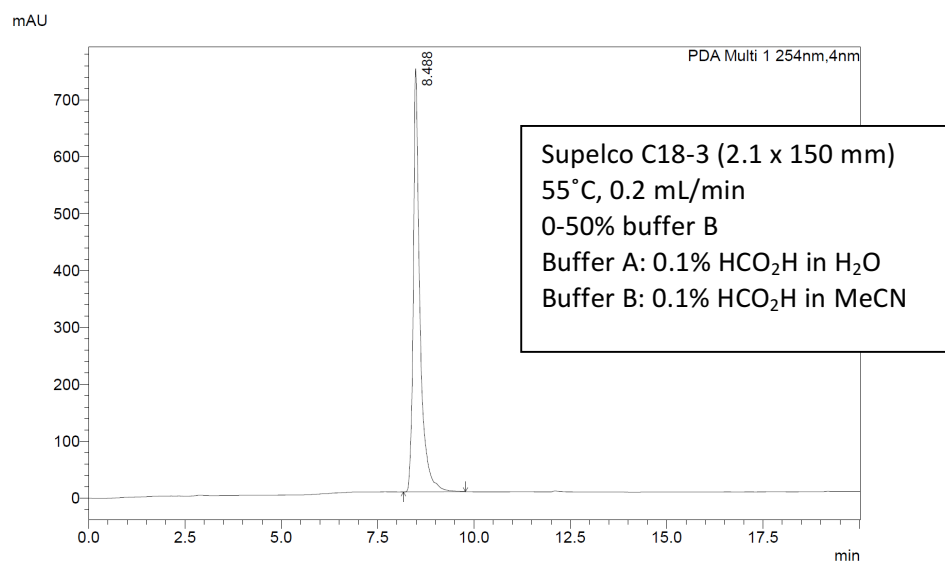
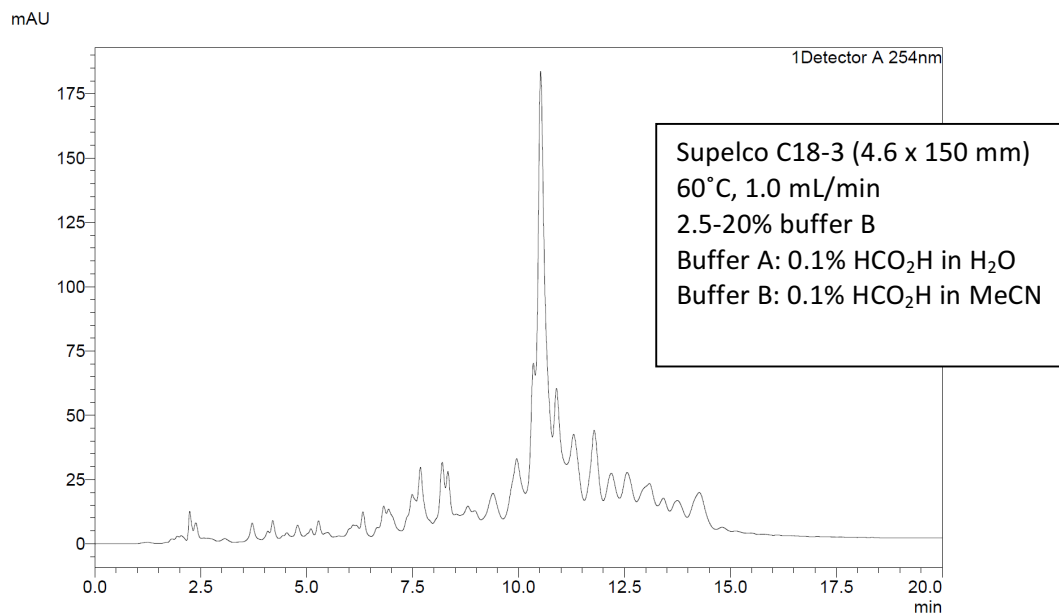
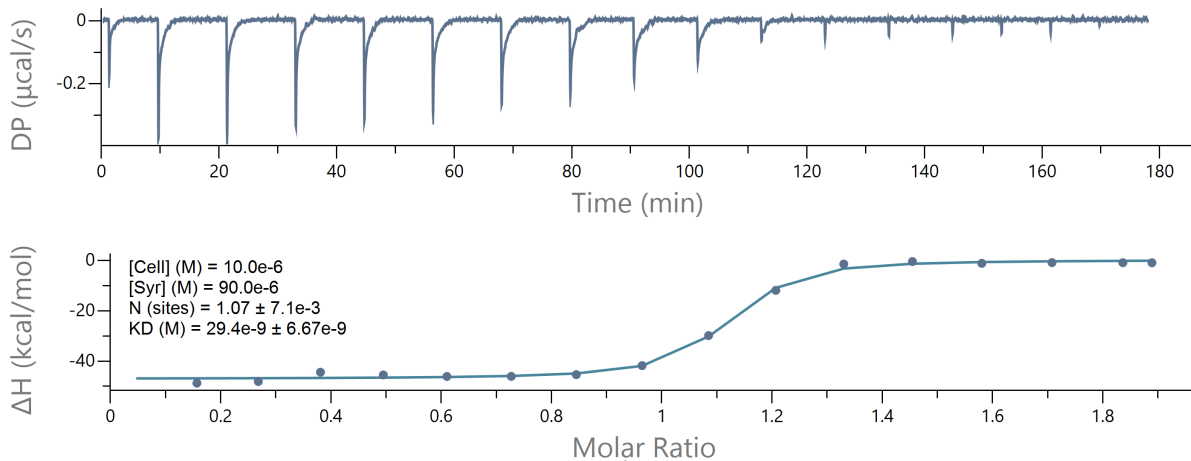
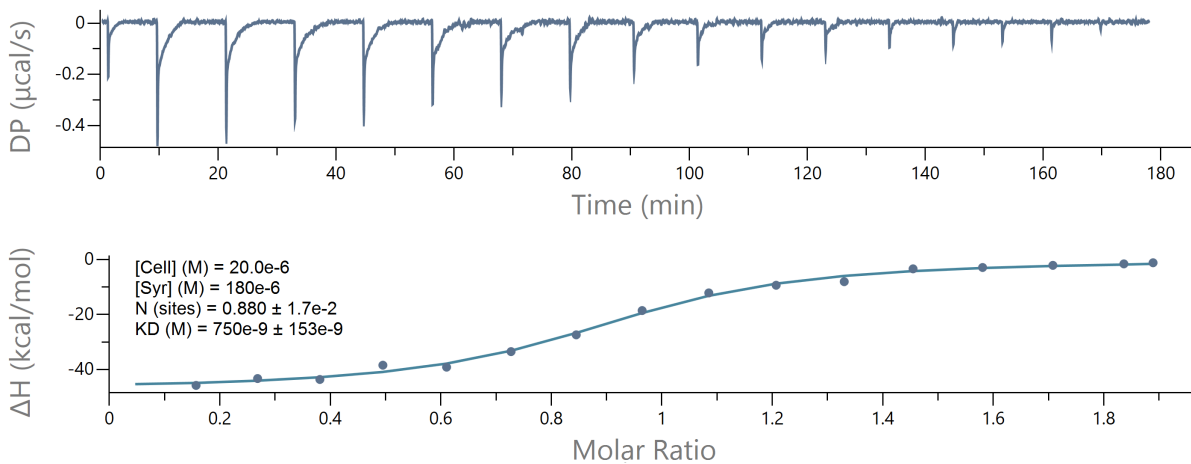


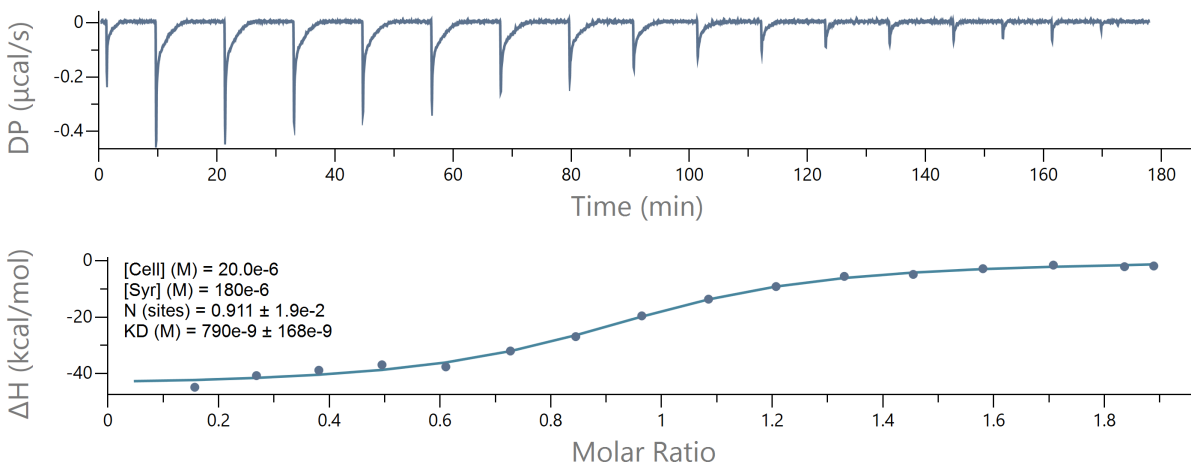
Figure S12. Crude (top) and purified (middle) chromatograms and mass spectrum (bottom) for PNA12.



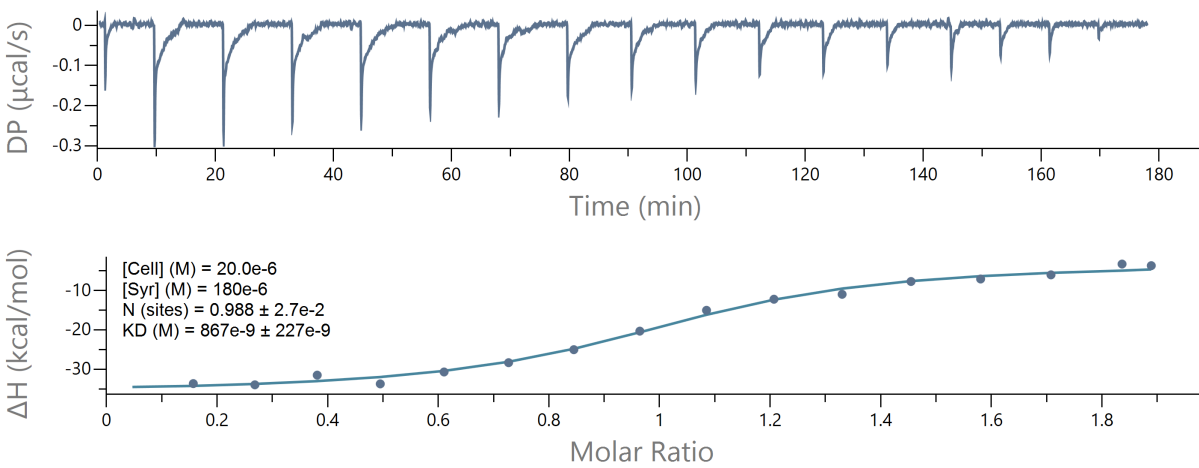
**Figure S13.** Representative ITC result of PNA1 against rHRP1.



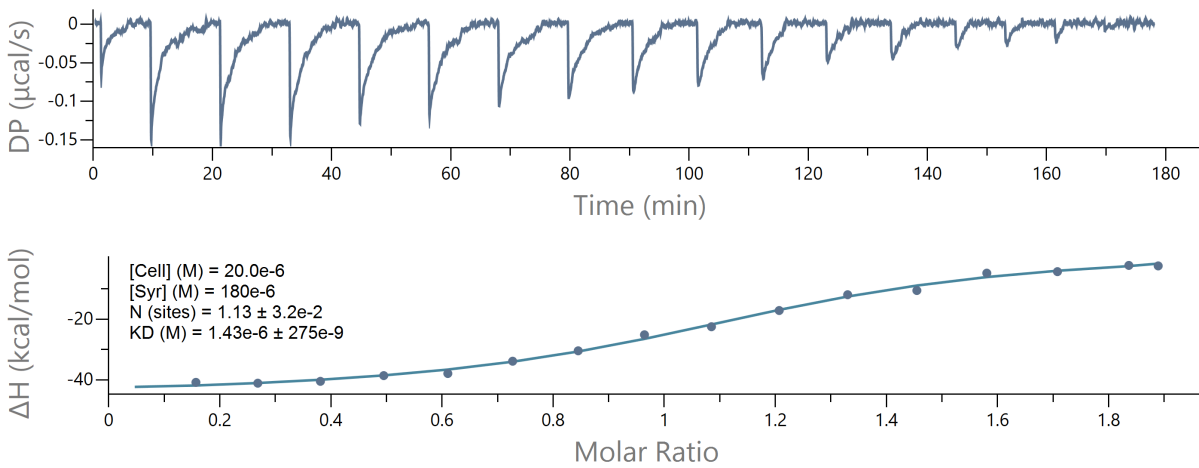
**Figure S14.** Representative ITC result of PNA1 against rHRP2.



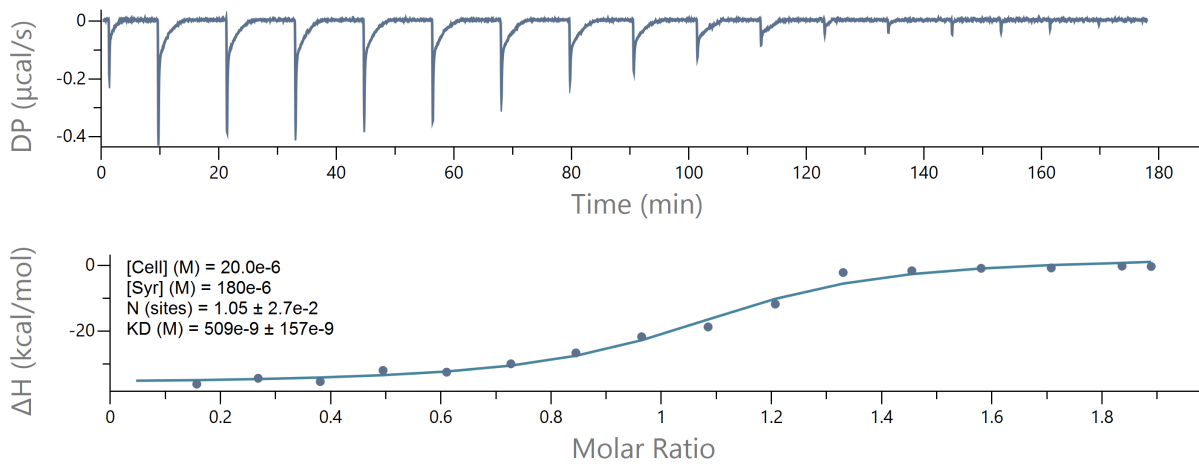
**Figure S15.** Representative ITC result of PNA1 against rHRP3.



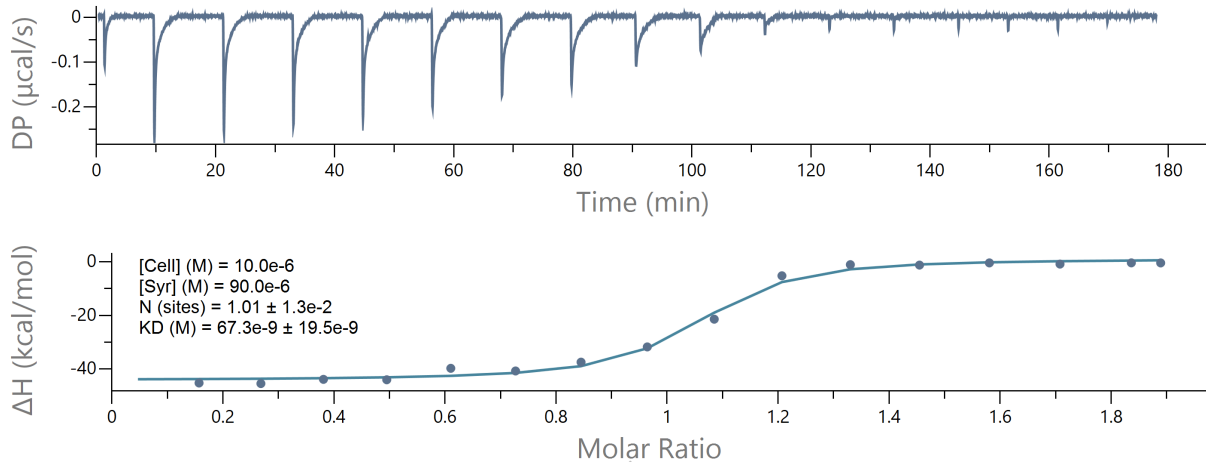
**Figure S16.** Representative ITC result of PNA1 against rHRP4.



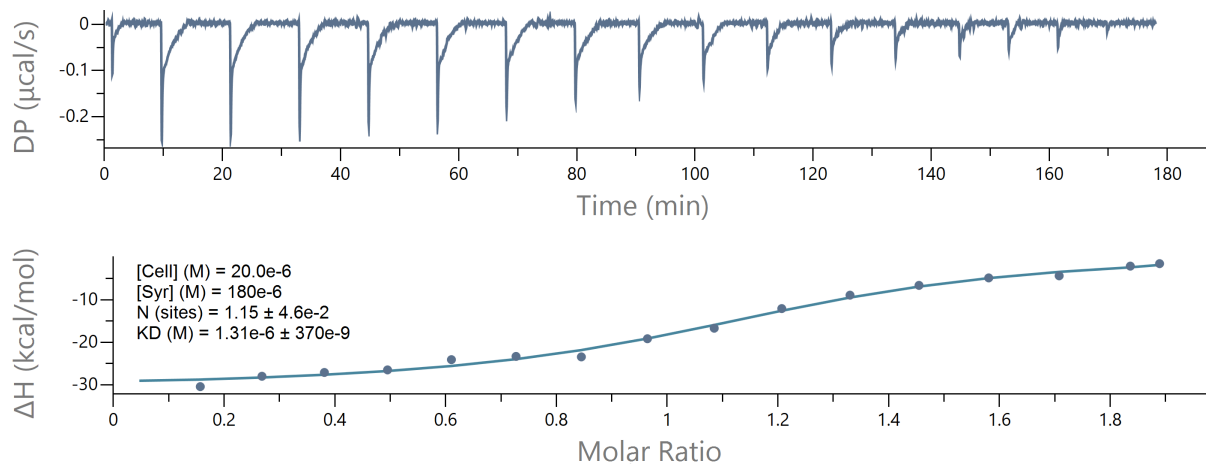
**Figure S17.** Representative ITC result of PNA1 against dHRP1.



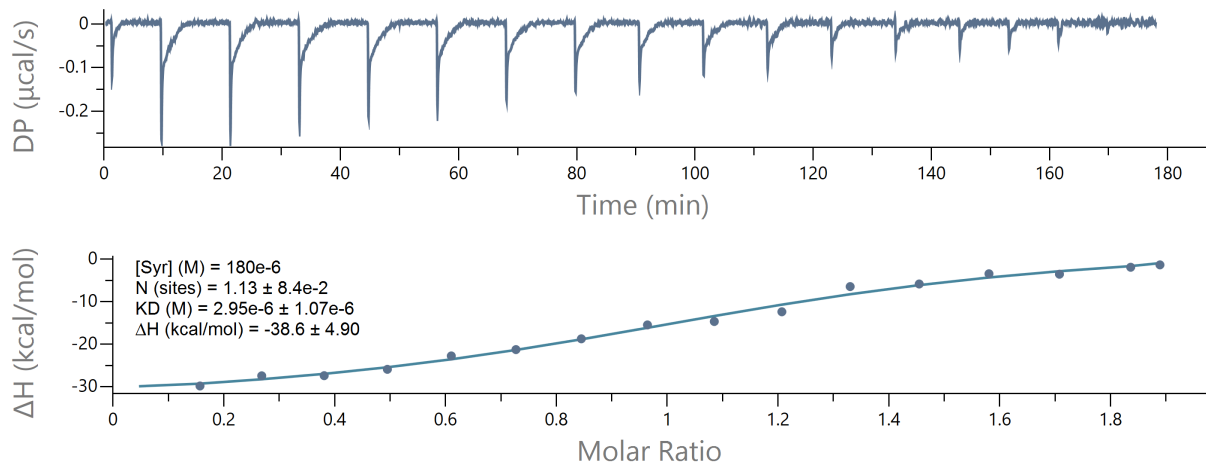
**Figure S18.** Representative ITC result of PNA2 against rHRP



**Figure S19.** Representative ITC result of PNA2 against rHRP2.

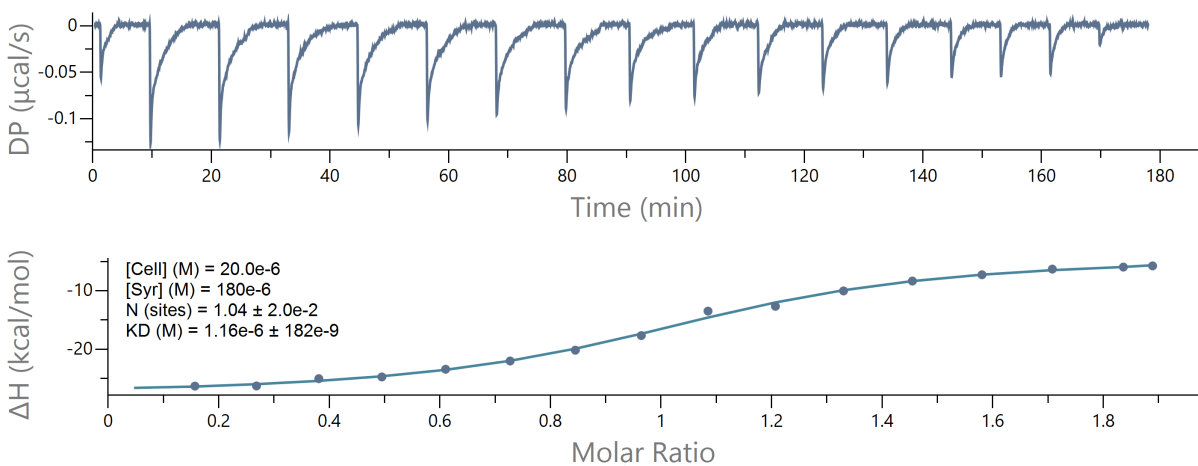


**Figure S20.** Representative ITC result of PNA2 against rHRP3.

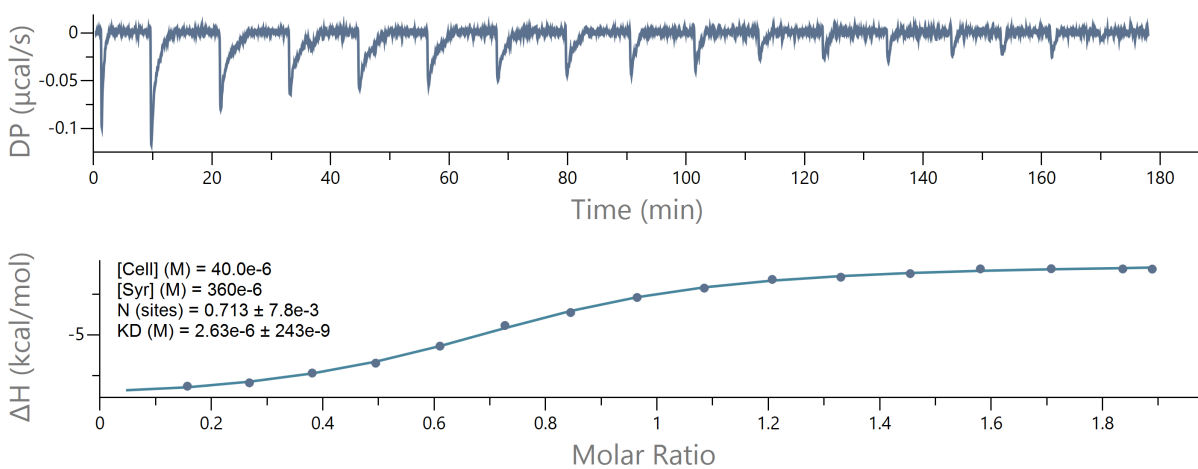


**Figure S21.** Representative ITC result of PNA2 against rHRP4.

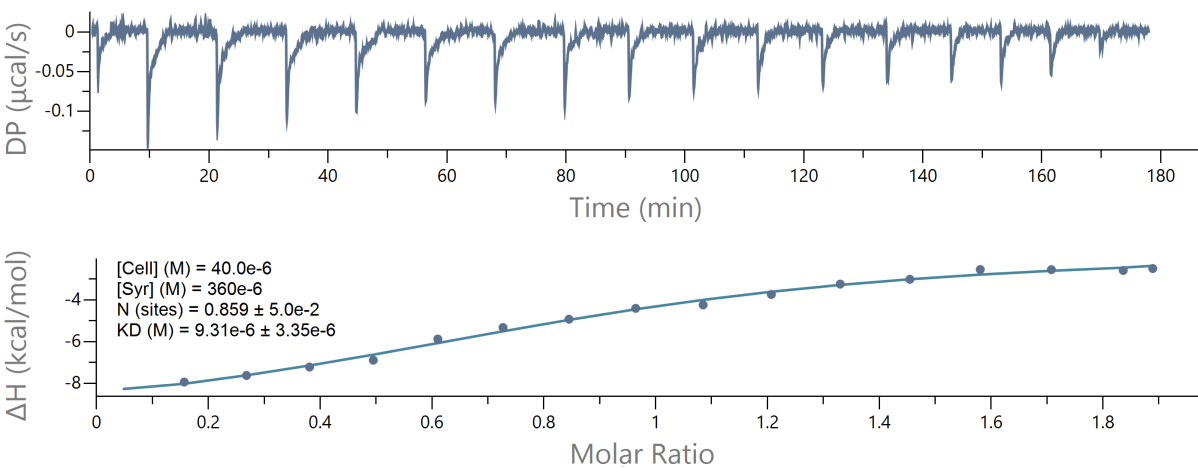




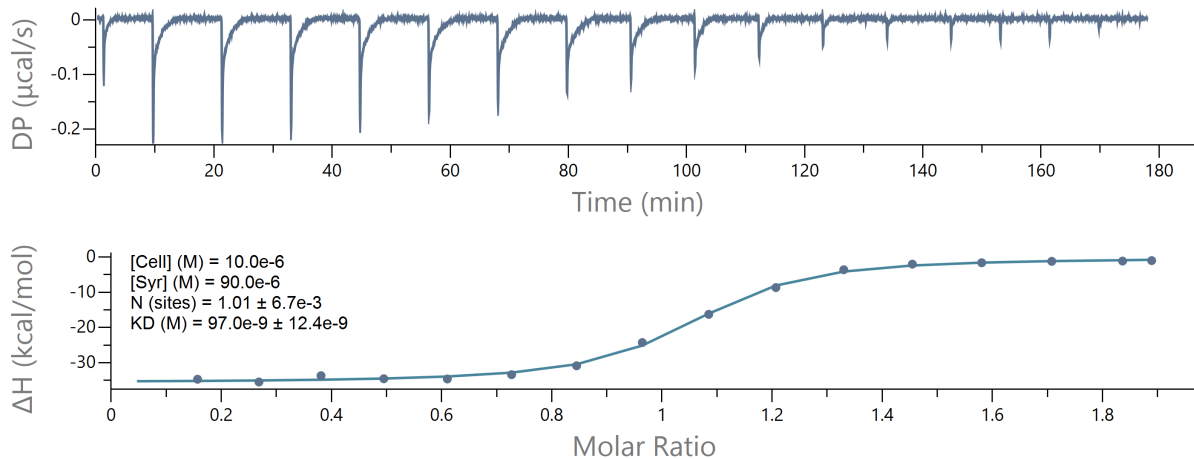
**Figure S22.** Representative ITC result of PNA2 against dHRP2.



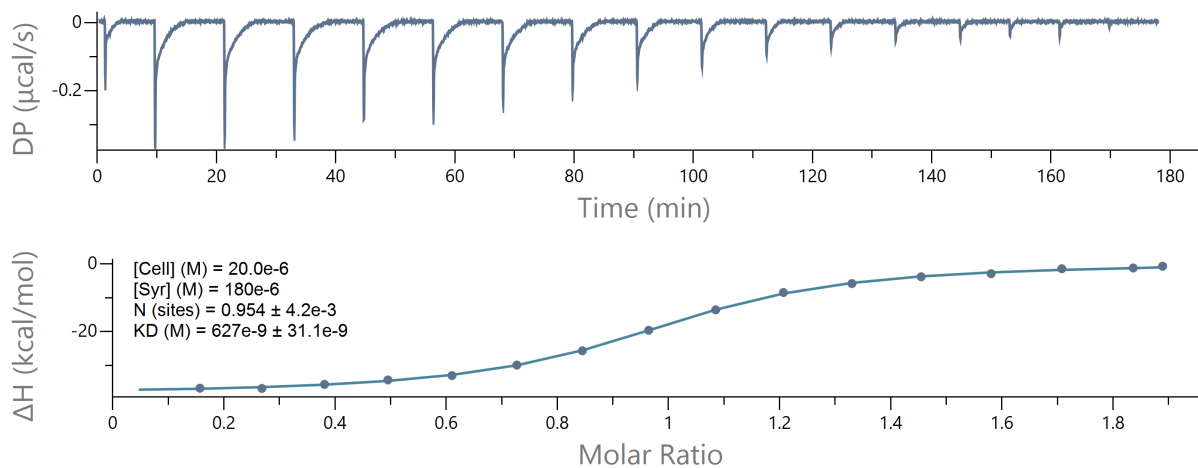
**Figure S23.** Representative ITC result of PNA3 against rHRP1.



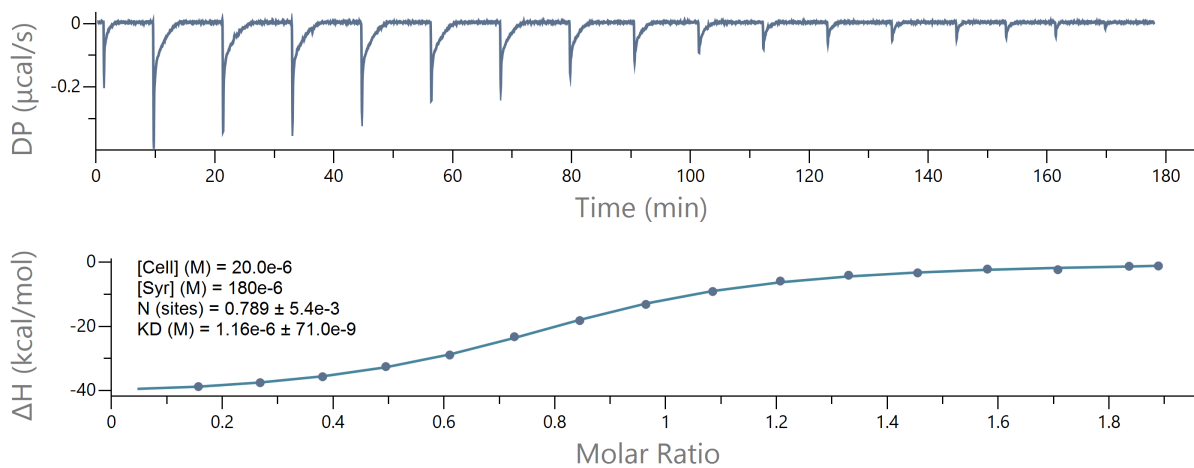
**Figure S24.** Representative ITC result of PNA3 against dHRP1.



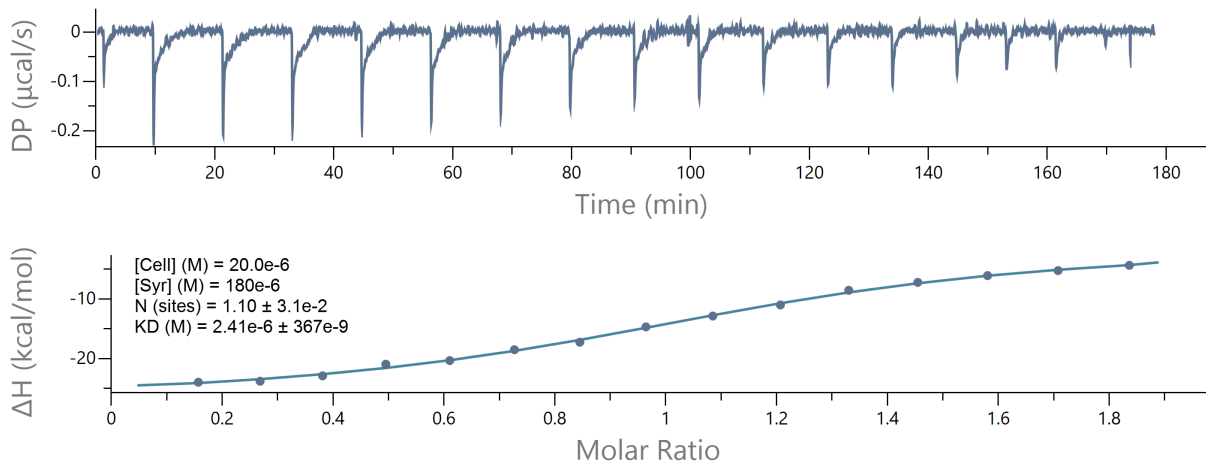
**Figure S25.** Representative ITC result of PNA4 against rHRP1.



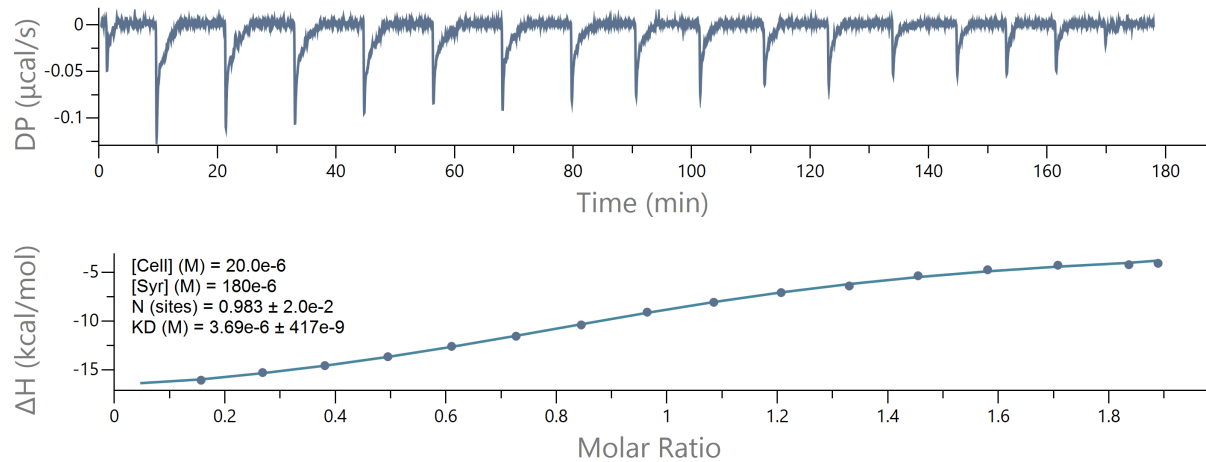
**Figure S26.** Representative ITC result of PNA4 against rHRP2.



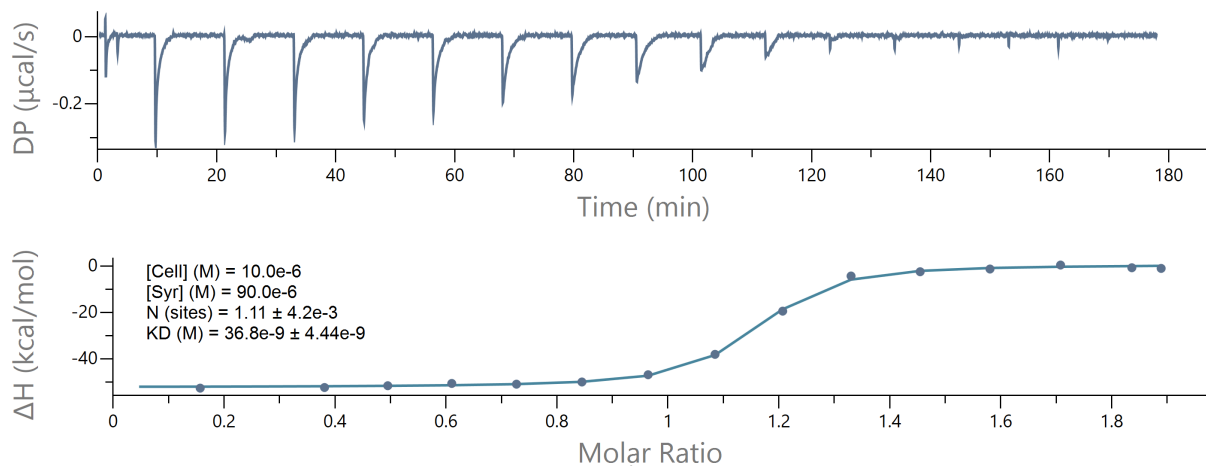
**Figure S27.** Representative ITC result of PNA4 against rHRP3.



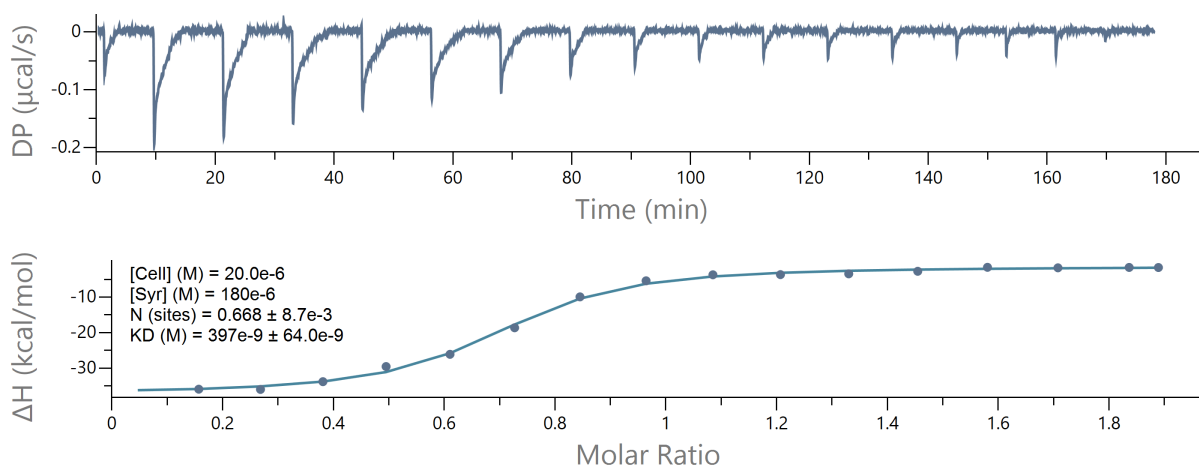
**Figure S28.** Representative ITC result of PNA4 against rHRP4.



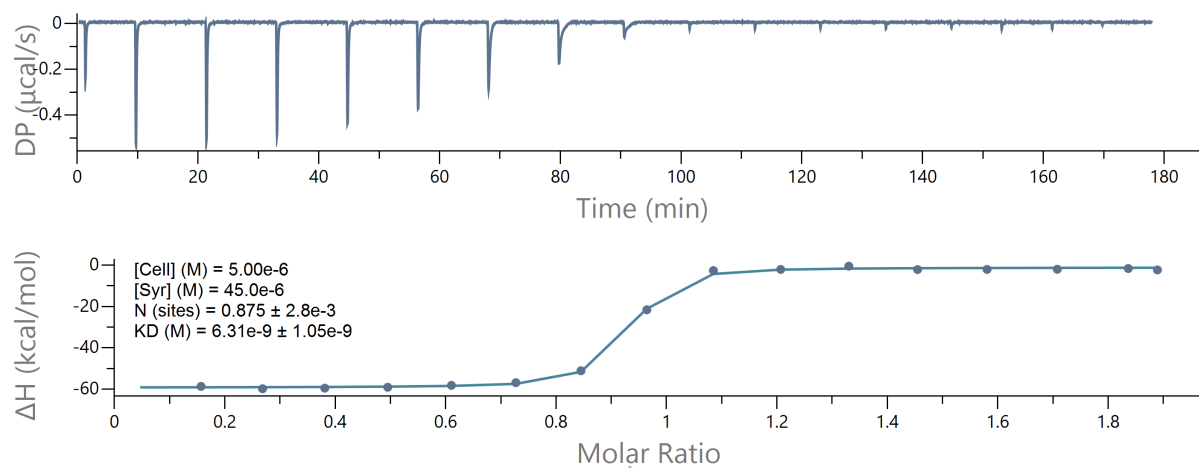
**Figure S29.** Representative ITC result of PNA4 against dHRP1.



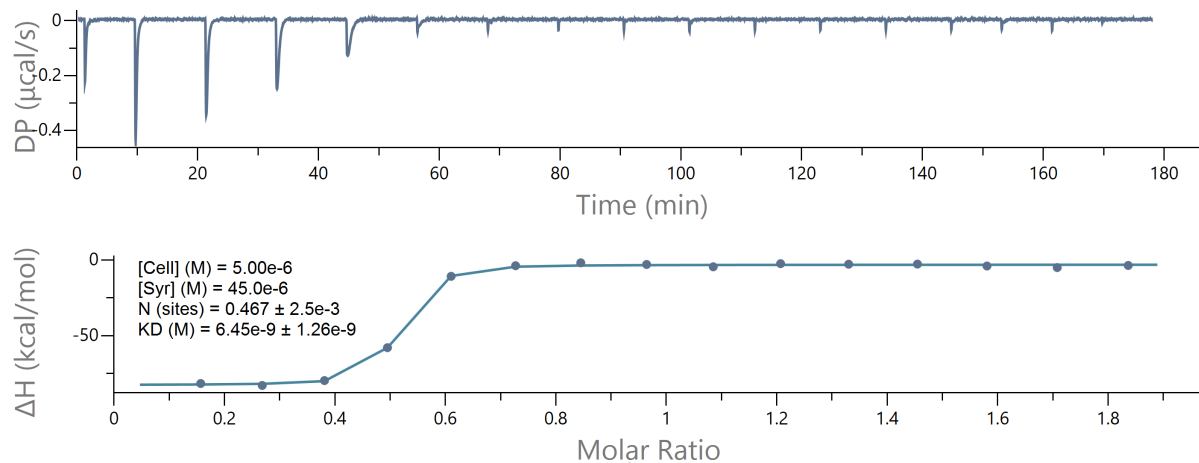
**Figure S30.** Representative ITC result of PNA5 against HRP5r.



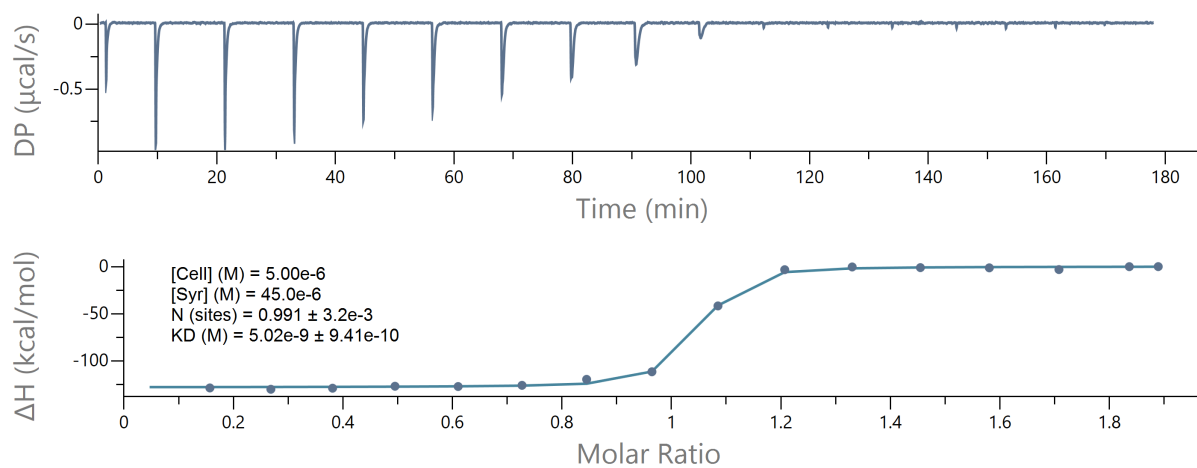
**Figure S31.** Representative ITC result of PNA5 against HRP5d.



**Figure S32.** Representative ITC result of PNA10 against ssDNA<sub>5</sub>.



**Figure S33.** Representative ITC result of PNA11 against ssDNA<sub>5</sub>.



**Figure S34.** Representative ITC result of **PNA12** against **ssDNA<sub>5</sub>**.

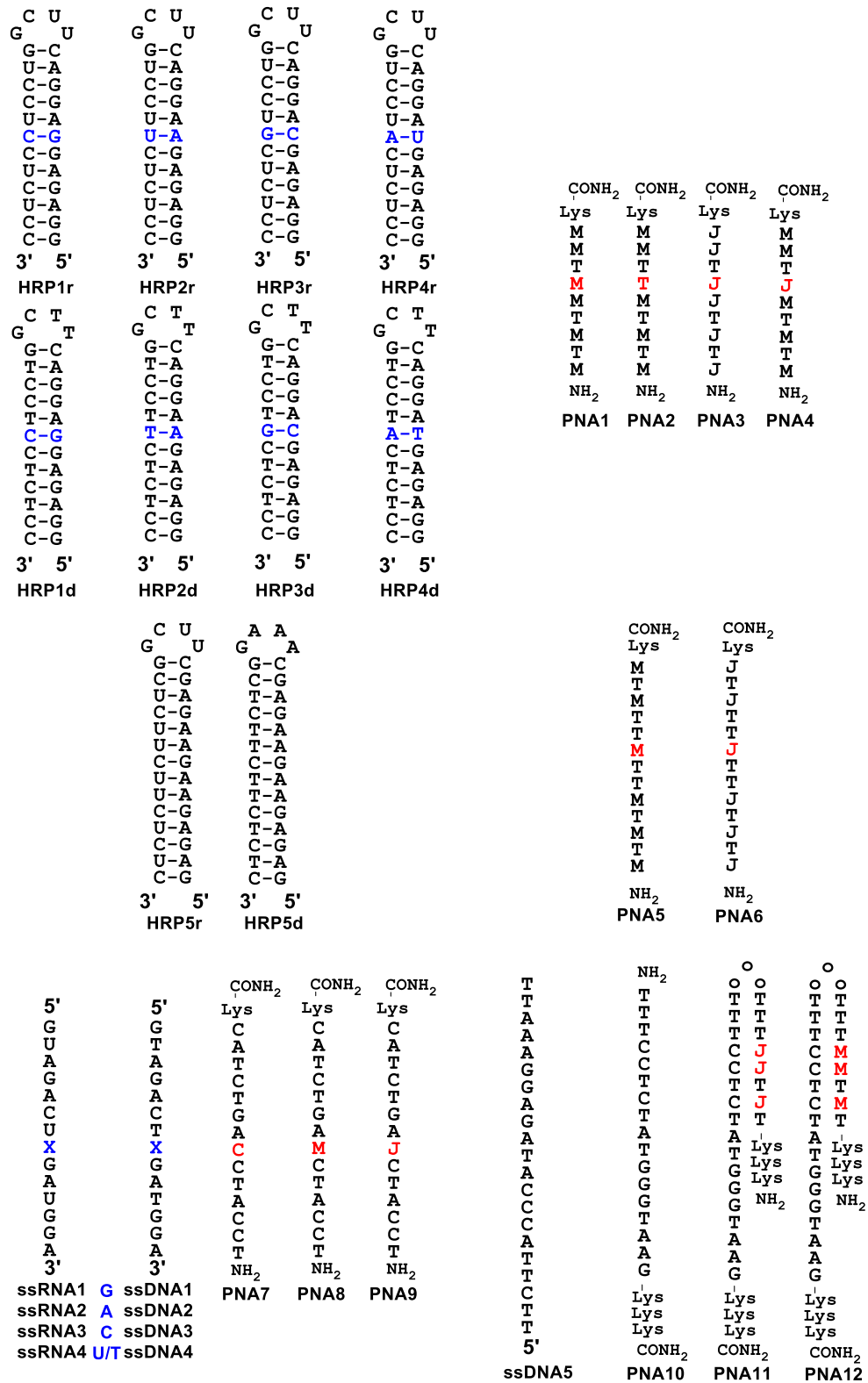


Figure S35. PNA and Oligonucleotide sequences.

Table S2. PNA1 binding affinity and thermodynamic data obtained by ITC.

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
PNA1 vs rHRP1 01	3.18E-08	3.14E+07	-52.0	41.8	1.1
PNA1 vs rHRP1 02	2.94E-08	3.40E+07	-47.1	36.8	1.1
PNA1 vs rHRP1 03	2.89E-08	3.46E+07	-49.1	38.8	1.1
Average	3.00E-08	3.34E+07	-49.4	39.1	1.1
St. Dev.	1.27E-09	1.37E+06	2.0	2.1	0.00
PNA1 vs rHRP2 01	7.78E-07	1.29E+06	-39.2	30.8	1.2
PNA1 vs rHRP2 02	7.50E-07	1.33E+06	-47.2	38.8	0.9
PNA1 vs rHRP2 03	7.96E-07	1.26E+06	-47.9	39.6	0.9
Average	7.75E-07	1.29E+06	-44.8	36.4	1.0
St. Dev.	1.89E-08	3.18E+04	3.9	4.0	0.17
PNA1 vs rHRP3 01	7.46E-07	1.34E+06	-40.2	31.8	1.0
PNA1 vs rHRP3 02	7.90E-07	1.27E+06	-45.0	36.6	0.9
PNA1 vs rHRP3 03	8.15E-07	1.23E+06	-45.2	36.9	1.0
Average	7.84E-07	1.28E+06	-43.5	35.1	1.0
St. Dev.	2.85E-08	4.71E+04	2.3	2.3	0.06
PNA1 vs rHRP4 01	8.67E-07	1.15E+06	-32.7	24.5	1.0
PNA1 vs rHRP4 02	7.39E-07	1.35E+06	-37.1	28.7	0.9
PNA1 vs rHRP4 03	8.94E-07	1.12E+06	-33.7	25.4	0.9
Average	8.33E-07	1.21E+06	-34.5	26.2	0.9
St. Dev.	6.76E-08	1.03E+05	1.9	1.8	0.06
PNA1 vs dHRP1 01	1.43E-06	6.99E+05	-48.0	40.0	1.1
PNA1 vs dHRP1 02	1.37E-06	7.30E+05	-39.3	31.3	1.2
PNA1 vs dHRP1 03	1.21E-06	8.26E+05	-52.0	43.9	1.0
Average	1.34E-06	7.52E+05	-46.4	38.4	1.1
St. Dev.	1.14E-07	6.64E+04	6.5	6.5	0.12

**Table S3. PNA2 binding affinity and thermodynamic data obtained by ITC.**

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
PNA2 vs rHRP1 01	5.92E-07	1.69E+06	-38.8	30.3	1.0
PNA2 vs rHRP1 02	5.09E-07	1.96E+06	-38.5	29.9	1.1
PNA2 vs rHRP1 03	5.11E-07	1.96E+06	-40.5	31.9	1.0
<b>Average</b>	5.37E-07	1.87E+06	-39.3	30.7	1.0
<b>St. Dev.</b>	3.87E-08	1.28E+05	0.9	0.9	0.06
PNA2 vs rHRP2 01	7.98E-08	1.25E+07	-42.4	32.7	1.0
PNA2 vs rHRP2 02	8.34E-08	1.20E+07	-41.4	31.7	1.1
PNA2 vs rHRP2 03	8.44E-08	1.18E+07	-40.4	30.7	1.1
<b>Average</b>	8.25E-08	1.21E+07	-41.4	31.7	1.1
<b>St. Dev.</b>	1.98E-09	2.94E+05	0.8	0.8	0.06
PNA2 vs rHRP3 01	9.64E-07	1.04E+06	-33.9	25.7	1.1
PNA2 vs rHRP3 02	1.04E-06	9.62E+05	-33.9	25.7	1.1
PNA2 vs rHRP3 03	1.31E-06	7.63E+05	-32.0	24.0	1.2
<b>Average</b>	1.10E-06	9.21E+05	-33.3	25.1	1.1
<b>St. Dev.</b>	1.48E-07	1.16E+05	0.9	0.8	0.06
PNA2 vs rHRP4 01	2.40E-06	4.17E+05	-39.2	31.6	1.1
PNA2 vs rHRP4 02	2.19E-06	4.57E+05	-37.1	29.4	1.1
PNA2 vs rHRP4 03	2.95E-06	3.39E+05	-38.6	31.0	1.1
<b>Average</b>	2.51E-06	4.04E+05	-38.3	30.7	1.1
<b>St. Dev.</b>	3.20E-07	4.88E+04	0.9	0.9	0.00
PNA2 vs dHRP2 01	1.03E-06	9.71E+05	-23.0	14.9	1.0
PNA2 vs dHRP2 02	9.98E-07	1.00E+06	-25.5	17.3	1.1
PNA2 vs dHRP2 03	1.16E-06	8.62E+05	-23.9	15.7	1.0
<b>Average</b>	1.06E-06	9.45E+05	-24.1	16.0	1.1
<b>St. Dev.</b>	8.58E-08	7.35E+04	1.3	1.2	0.03



Table S4. PNA3 binding affinity and thermodynamic data obtained by ITC.

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
PNA3 vs rHRP5 01	1.87E-06	5.35E+05	-7.82	-1.6	0.7
PNA3 vs rHRP5 02	2.63E-06	3.80E+05	-8.55	0.9	0.7
PNA3 vs rHRP5 03	2.50E-06	4.00E+05	-10.1	2.4	0.7
Average	2.33E-06	4.38E+05	-8.8	0.6	0.7
St. Dev.	4.07E-07	8.41E+04	1.2	2.0	0.02
PNA3 vs dHRP5 01	1.19E-05	8.40E+04	-14.7	8.0	0.8
PNA3 vs dHRP5 02	9.31E-06	1.07E+05	-8.56	1.7	0.9
Average	1.06E-05	9.57E+04	-11.6	4.9	0.8
St. Dev.	1.83E-06	1.65E+04	4.3	4.5	0.08

Table S5. PNA4 binding affinity and thermodynamic data obtained by ITC.

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
PNA4 vs rHRP1 01	9.70E-08	1.03E+07	-35.3	25.7	1.0
PNA4 vs rHRP1 02	9.54E-08	1.05E+07	-34.7	25.1	1.0
PNA4 vs rHRP1 03	9.07E-08	1.10E+07	-36.4	27.1	1.0
Average	9.44E-08	1.06E+07	-35.5	26.0	1.0
St. Dev.	3.27E-09	3.74E+05	0.9	1.0	0.01
PNA4 vs rHRP2 01	8.31E-07	1.20E+06	-30.3	21.7	0.9
PNA4 vs rHRP2 02	5.24E-07	1.91E+06	-43.6	35.1	0.5
PNA4 vs rHRP2 03	6.27E-07	1.60E+06	-38.7	30.2	1.0
Average	6.61E-07	1.57E+06	-37.5	29.0	0.8
St. Dev.	1.56E-07	3.53E+05	6.7	6.8	0.27
PNA4 vs rHRP3 01	1.16E-06	8.62E+05	-42.9	34.8	0.8
PNA4 vs rHRP3 02	1.06E-06	9.43E+05	-51.5	43.4	0.9
PNA4 vs rHRP3r 03	1.29E-06	7.75E+05	-40.7	32.4	0.8
Average	1.17E-06	8.60E+05	-45.0	36.9	0.8
St. Dev.	1.15E-07	8.41E+04	5.8	5.8	0.09
PNA4 vs rHRP4 01	2.89E-06	3.45E+05	-28.9	21.3	1.2
PNA4 vs rHRP4 02	2.68E-06	3.73E+05	-19.5	11.9	0.7
PNA4 vsr HRP4 03	2.41E-06	4.15E+05	-26.3	18.7	1.1
Average	2.66E-06	3.80E+05	-24.9	17.3	1.0
St. Dev.	2.41E-07	4.87E+04	4.9	4.9	0.24
PNA4 vs dHRP1 01	4.92E-06	2.03E+05	-27.0	19.7	1.1
PNA4 vs dHRP1 02	3.69E-06	2.71E+05	-17.2	9.8	1.0
PNA4 vs dHRP1 03	5.10E-06	1.96E+05	-22.2	15.0	1.1
Average	4.57E-06	2.23E+05	-22.1	14.8	1.1
St. Dev.	7.67E-07	4.13E+04	4.9	5.0	0.06

**Table S6.** UV-melting of **PNA1** vs matched and mismatched dsRNA and dsDNA.

Name	PNA1 vs rHRP1	PNA1 vs rHRP2	PNA1 vs rHRP3	PNA1 vs rHRP4
<b>Melting</b> <b>temp. (°C)</b>	67.1	36.2	36.6	32.9
	67.3	36.2	37.3	32.8
	66.2	35.8	36.2	32.8
	66.5	36.2	37.2	31.8
	65.6	36.9	36.9	32.6
<b>Average</b>	66.5	36.3	36.8	32.6
<b>St. Dev.</b>	0.7	0.4	0.4	0.4
Name	PNA1 vs dHRP1	PNA1 vs dHRP2	PNA1 vs dHRP3	PNA1 vs dHRP4
<b>Melting</b> <b>temp. (°C)</b>	35.2	<20	<20	<20
	34.5	<20	<20	<20
	35.5	<20	<20	<20
	35.0	<20	<20	<20
	34.8	<20	<20	<20
<b>Average</b>	35.0	-	-	-
<b>St. Dev.</b>	0.3	-	-	-

**Table S7.** UV-melting of **PNA2** vs matched and mismatched dsRNA and dsDNA.

Name	PNA2 vs rHRP1	PNA2 vs rHRP2	PNA2 vs rHRP3	PNA2 vs rHRP4
<b>Melting</b> <b>temp. (°C)</b>	46.4	70.1	35.4	34.7
	46.8	70.1	35.2	34.5
	46.5	70.4	36.0	34.3
	45.6	68.7	35.6	34.8
	46.9	68.6	34.7	34.7
<b>Average</b>	46.4	69.6	35.4	34.6
<b>St. Dev.</b>	0.5	0.8	0.4	0.2
Name	PNA2 vs dHRP1	PNA2 vs dHRP2	PNA2 vs dHRP3	PNA2 vs dHRP4
<b>Melting</b> <b>temp. (°C)</b>	<20	29.7	<20	<20
	<20	29.9	<20	<20
	<20	29.0	<20	<20
	<20	28.9	<20	<20
	<20	29.6	<20	<20
<b>Average</b>	-	29.4	-	-
<b>St. Dev.</b>	-	0.4	-	-

**Table S8.** UV-melting of **PNA3** vs matched dsRNA and dsDNA.

Name	PNA3 vs rHRP1
<b>Melting temp. (°C)</b>	37.6
	36.3
	36.9
	36.4
	37.0
<b>Average St. Dev.</b>	36.8 0.5
Name	PNA3 vs dHRP1
<b>Melting temp. (°C)</b>	<20
	<20
	<20
	<20
	<20
<b>Average St. Dev.</b>	- -

**Table S9.** UV-melting of **PNA4** vs matched and mismatched dsRNA and dsDNA.

Name	PNA4 vs rHRP1	PNA4 vs rHRP2	PNA4 vs rHRP3	PNA4 vs rHRP4
<b>Melting temp. (°C)</b>	60.0	43.8	39.4	35.1
	60.8	44.1	39.3	35.4
	61.4	43.5	38.9	36.4
	60.3	44.6	38.7	35.8
	61.1	43.1	39.3	35.8
<b>Average St. Dev.</b>	60.7 0.6	43.8 0.6	39.1 0.3	35.7 0.5
Name	PNA4 vs dHRP1	PNA4 vs dHRP2	PNA4 vs dHRP3	PNA4 vs dHRP4
<b>Melting temp. (°C)</b>	29.4	<20	<20	<20
	30.2	<20	<20	<20
	29.7	<20	<20	<20
	29.8	<20	<20	<20
	29.5	<20	<20	<20
<b>Average St. Dev.</b>	29.7 0.3	- -	- -	- -

**Table S10.** PNA5 binding affinity and thermodynamic data obtained by ITC.

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
PNA5 vs HRP5r 01	3.68E-06	2.72E+07	-53.4	42.7	0.8
PNA5 vs HRP5r 02	3.55E-06	2.82E+07	-57.2	43.2	0.7
PNA5 vs HRP5r 03	3.66E-06	2.73E+07	-54.5	47.0	0.8
<b>Average</b>	3.84E-06	2.76E+07	-54.5	44.3	0.7
<b>St. Dev.</b>	7.73E-07	5.37E+05	2.4	2.4	0.06
PNA5 vs HRP5d 01	4.54E-07	2.20E+06	-34.6	26.0	1.1
PNA5 vs HRP5d 02	3.97E-07	2.52E+06	-35.9	27.2	1.1
PNA5 vs HRP5d 03	3.01E-07	3.32E+06	-35.3	26.4	1.1
<b>Average</b>	3.63E-07	2.68E+06	-35.3	26.5	1.1
<b>St. Dev.</b>	7.00E-08	5.77E+05	0.7	0.6	0.01

**Table S11.** UV-melting of PNA5 vs matched dsRNA and dsDNA

Name	PNA5 vs HRP5r	PNA5 vs HRP5d
<b>Melting temp. (°C)</b>	90.7	85.9
	90.2	87.0
	89.8	86.5
	89.9	87.5
	89.0	87.1
<b>Average</b>	89.9	86.8
<b>St. Dev.</b>	0.6	0.6

**Table S12.** UV-melting of **PNA7** vs matched and mismatched ssRNA and ssDNA.

Name	PNA7 vs ssRNA1	PNA7 vs ssRNA2	PNA7 vs ssRNA3	PNA7 vs ssRNA4
<b>Melting</b> <b>temp. (°C)</b>	73.3	58.2	56.9	57.2
	72.8	58.1	56.7	56.9
	73.1	57.5	56.3	56.4
	73.8	57.8	56.0	56.7
	72.4	57.3	55.9	56.9
<b>Average</b>	73.2	57.8	56.4	56.8
<b>St. Dev.</b>	0.5	0.4	0.4	0.3
Name	PNA7 vs ssDNA1	PNA7 vs ssDNA2	PNA7 vs ssDNA3	PNA7 vs ssDNA4
<b>Melting</b> <b>temp. (°C)</b>	66.9	46.1	45.0	47.8
	65.5	46.2	44.2	47.5
	66.6	45.7	44.4	47.1
	65.2	45.2	44.4	47.3
	66.0	44.9	44.7	47.3
<b>Average</b>	66.0	45.6	44.5	47.4
<b>St. Dev.</b>	0.7	0.5	0.3	0.3

**Table S13.** UV-melting of **PNA8** vs matched and mismatched ssRNA and ssDNA.

Name	PNA8 vs ssRNA1	PNA8 vs ssRNA2	PNA8 vs ssRNA3	PNA8 vs ssRNA4
<b>Melting</b> <b>temp. (°C)</b>	59.9	55.7	57.7	56.1
	58.4	55.7	57.0	56.5
	59.5	55.4	56.4	55.1
	58.3	55.4	56.6	56.0
	59.5	54.9	56.0	55.2
<b>Average</b>	59.1	55.4	56.8	55.8
<b>St. Dev.</b>	0.7	0.3	0.6	0.6
Name	PNA8 vs ssDNA1	PNA8 vs ssDNA2	PNA8 vs ssDNA3	PNA8 vs ssDNA4
<b>Melting</b> <b>temp. (°C)</b>	45.0	42.4	46.8	46.0
	45.6	41.2	47.0	45.9
	45.4	41.9	46.2	45.2
	45.2	41.1	46.3	45.2
	45.2	41.1	46.0	45.4
<b>Average</b>	45.3	41.5	46.5	45.6
<b>St. Dev.</b>	0.2	0.6	0.4	0.4

**Table S14.** UV-melting of **PNA9** vs matched and mismatched ssRNA and ssDNA.

<b>Name</b>	<b>PNA9 vs ssRNA1</b>	<b>PNA9 vs ssRNA2</b>	<b>PNA9 vs ssRNA3</b>	<b>PNA9 vs ssRNA4</b>
<b>Melting temp. (°C)</b>	73.0	58.1	58.4	58.9
	72.0	57.7	58.1	59.0
	71.8	57.3	58.0	58.2
	72.7	56.9	57.8	58.6
	71.4	56.9	57.5	57.8
<b>Average</b>	72.2	57.4	58.0	58.5
<b>St. Dev.</b>	0.6	0.5	0.4	0.5
<b>Name</b>	<b>PNA9 vs ssDNA1</b>	<b>PNA9 vs ssDNA2</b>	<b>PNA9 vs ssDNA3</b>	<b>PNA9 vs ssDNA4</b>
<b>Melting temp. (°C)</b>	67.1	46.5	49.0	49.9
	65.8	45.5	49.4	49.9
	66.6	45.9	48.7	49.8
	65.9	45.6	48.7	49.7
	66.2	46.2	48.6	49.3
<b>Average</b>	66.3	45.9	48.9	49.7
<b>St. Dev.</b>	0.5	0.4	0.3	0.2

**Table S15. PNA10-PNA12 binding affinity and thermodynamic data obtained by ITC.**

Name	$K_d$ (M)	$K_a$ ( $M^{-1}$ )	$\Delta H$ (kcal/mol)	$-T\Delta S$ (kcal/mol)	N (sites)
<b>PNA10 vs ssDNA5 01</b>	6.67E-9	1.49E+8	-65.9	54.7	0.8
<b>PNA10 vs ssDNA5 02</b>	6.29E-9	1.59E+8	-65.8	54.6	0.6
<b>PNA10 vs ssDNA5 03</b>	6.31E-9	1.58E+8	-58.0	46.8	0.9
<b>Average</b>	6.42E-9	1.56E+8	-63.2	52.0	0.8
<b>St. Dev.</b>	2.14E-10	5.09E+6	4.5	4.5	0.13
<b>PNA11 vs ssDNA5 01</b>	6.93E-9	1.44E+8	-79.4	68.3	0.4
<b>PNA11 vs ssDNA5 02</b>	6.45E-9	1.55E+8	-79.5	68.3	0.5
<b>PNA11 vs ssDNA5 03</b>	6.89E-9	1.45E+8	-97.9	86.8	0.4
<b>Average</b>	6.76E-9	1.48E+8	-85	75	0.42
<b>St. Dev.</b>	2.66E-10	5.97E+6	11	11	0.06
<b>PNA12 vs ssDNA5 01</b>	5.14E-9	1.95E+8	-140	128	1.0
<b>PNA12 vs ssDNA5 02</b>	5.02E-9	1.99E+8	-128	117	1.0
<b>PNA12 vs ssDNA5 03</b>	5.33E-9	1.88E+8	-137	125	1.0
<b>Average</b>	5.16E-9	1.95E+8	-135	123	1.0
<b>St. Dev.</b>	1.56E-10	5.83E+6	6.2	5.7	0.03

**Table S16. UV-melting of PNA10-PNA12 vs matched ssDNA5.**

Name	$T_m$ ( $^{\circ}C$ )
<b>PNA10 vs ssDNA5 01</b>	76.6
<b>PNA10 vs ssDNA5 02</b>	76.8
<b>PNA10 vs ssDNA5 03</b>	76.1
<b>PNA10 vs ssDNA5 04</b>	76.3
<b>PNA10 vs ssDNA5 05</b>	76.6
<b>Average</b>	76.4
<b>St. Dev.</b>	0.3
<b>PNA11 vs ssDNA5 01</b>	91.4
<b>PNA11 vs ssDNA5 02</b>	91.5
<b>PNA11 vs ssDNA5 03</b>	90.1
<b>PNA11 vs ssDNA5 04</b>	91.2
<b>PNA11 vs ssDNA5 05</b>	90.5
<b>Average</b>	91.0
<b>St. Dev.</b>	0.6
<b>PNA12 vs ssDNA5 01</b>	93.7
<b>PNA12 vs ssDNA5 02</b>	94.1
<b>PNA12 vs ssDNA5 03</b>	94.2
<b>PNA12 vs ssDNA5 04</b>	93.9
<b>PNA12 vs ssDNA5 05</b>	94.5
<b>Average</b>	94.1
<b>St. Dev.</b>	0.3

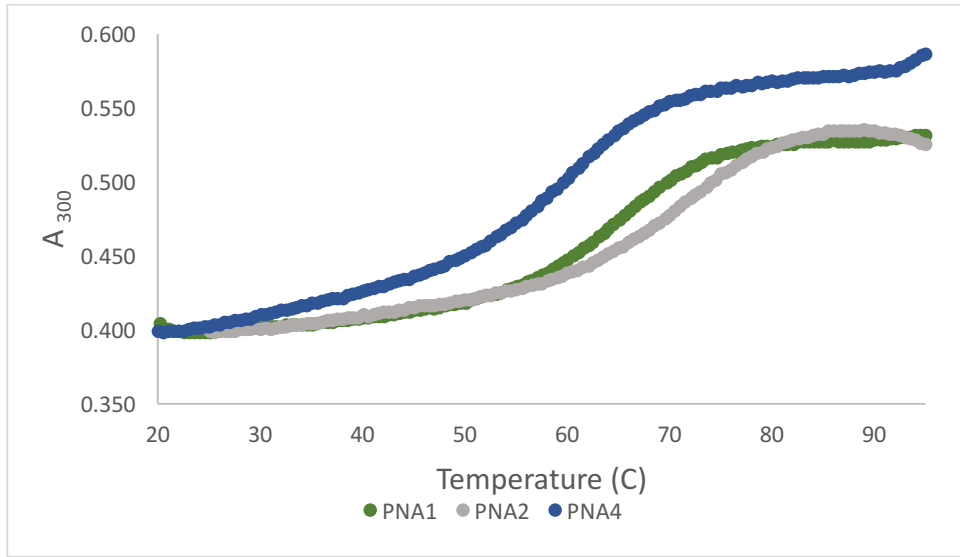


Figure S36. UV-melting at 300 nm of **PNA1** (M), **PNA2** (T), and **PNA4** (J) against matched dsRNA.

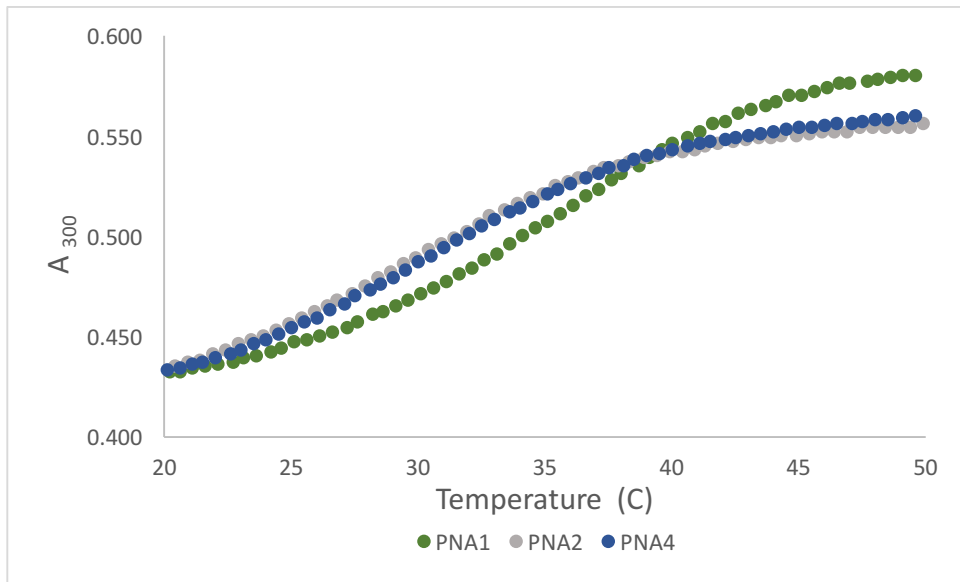
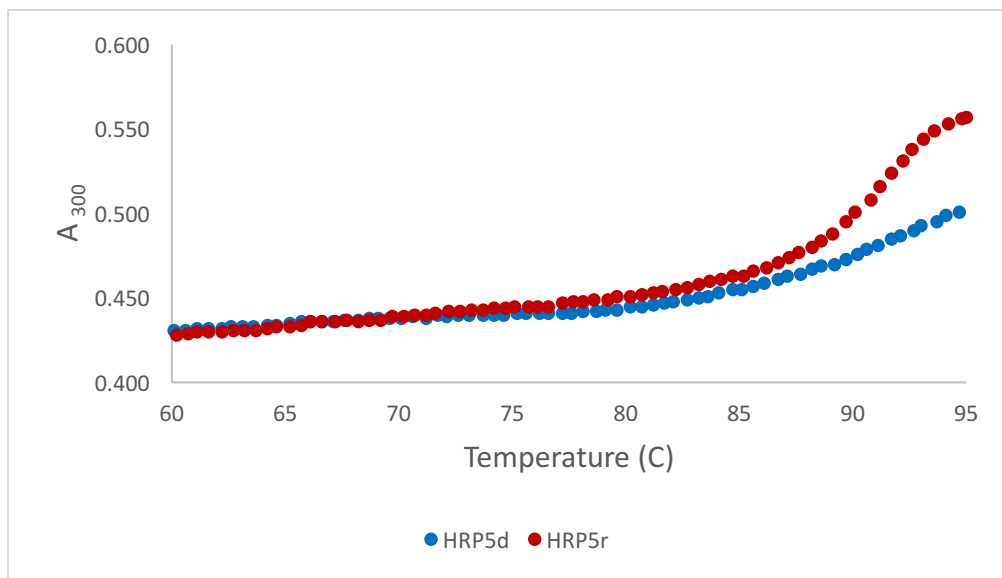
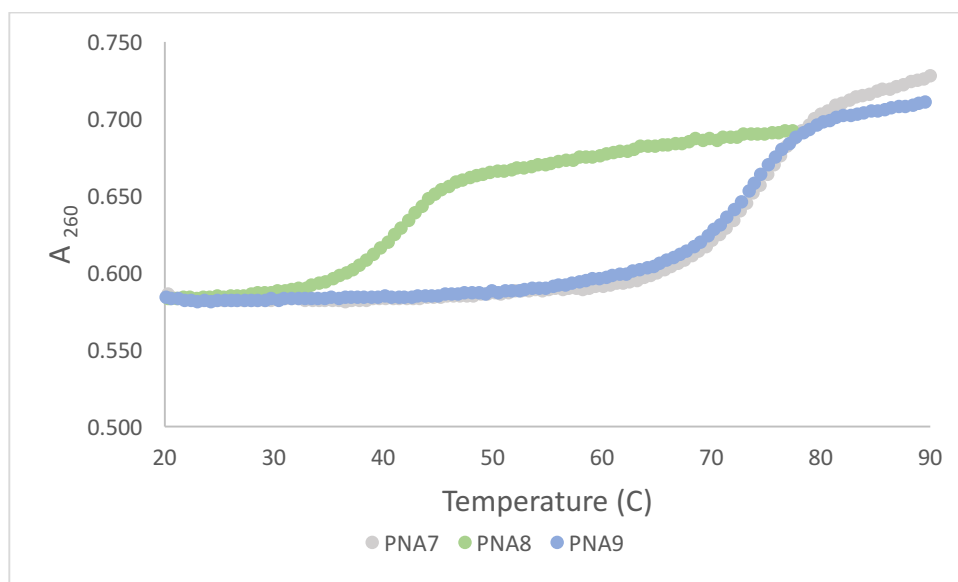


Figure S37. UV-melting at 300 nm of **PNA1** (M), **PNA2** (T), and **PNA4** (J) against matched dsDNA.

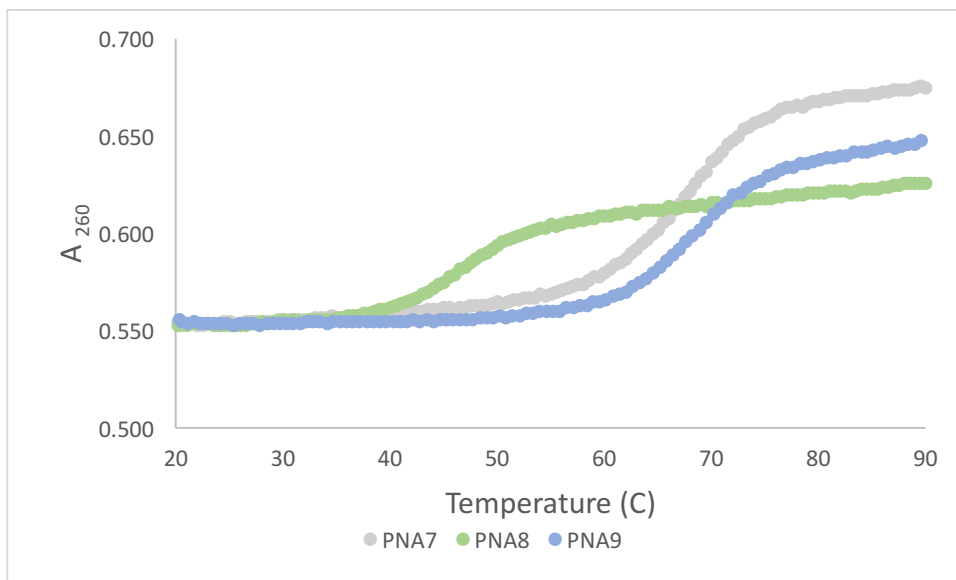




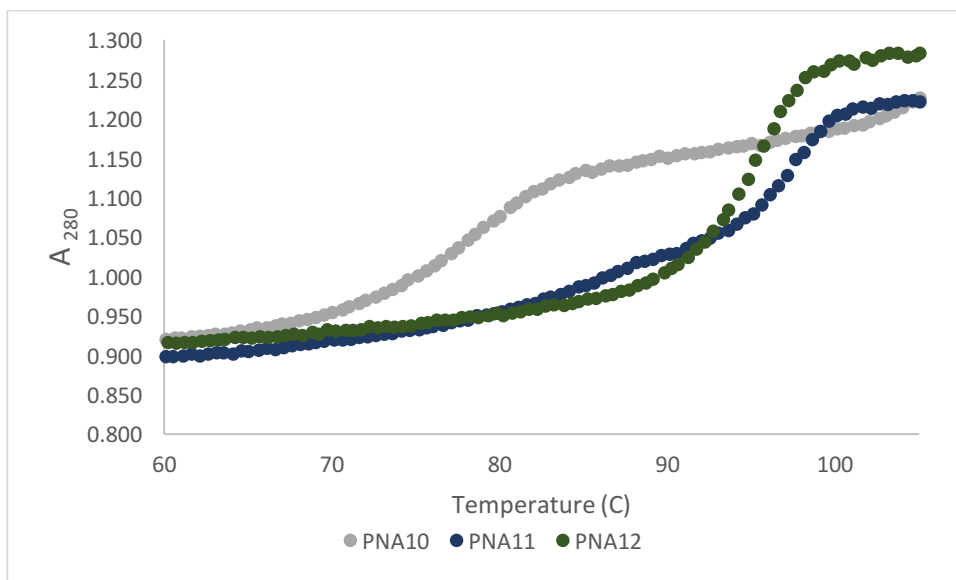
**Figure S38.** UV-melting at 300 nm of PNA5 against matched dsDNA and dsRNA.



**Figure S39.** UV-melting at 260 nm of **PNA7** (C), **PNA8** (J), and **PNA9** (J) against matched ssRNA1.



**Figure S40.** UV-melting at 260 nm of **PNA7** (C), **PNA8** (M), and **PNA9** (J) against matched ssDNA1.



**Figure S41.** UV-melting at 280nm of **PNA10**, **PNA11**, and **PNA12** against ssDNA5.