

**Table S1.** Molar composition of the prepolymerization mixtures used in the experimental design and resulting binding affinities of the corresponding nanoMIPs.

Polymer	AA %mol	tBAM %mol	NIPAM %mol	BIS %mol	$K_{eq} \pm 1$ s.e. ( $10^6$ M $^{-1}$ )
P1	10	57	30	3	$2.05 \pm 1.12$
P2	20	46	30	4	$2.44 \pm 0.68$
P3	20	48	30	2	$4.01 \pm 0.62$
P4	30	35	30	5	$3.99 \pm 0.57$
P5	30	37	30	3	$1.39 \pm 0.27$
P6	30	39	30	1	$3.00 \pm 0.75$
P7	40	26	30	4	$3.08 \pm 0.83$
P8	40	28	30	2	$0.63 \pm 0.76$
P9	50	17	30	3	$0.37 \pm 0.73$

**Table S2.** Non linear fitting of langmuirian binding models for ciprofloxacin-imprinted nanoMIPs.

Polymer	$K_{eq} \pm 1$ s.e. ( $10^6$ M $^{-1}$ )	$B_{max} \pm 1$ s.e. (nmol g $^{-1}$ )	r $^2$	Fit std. error	F-value
P1	$2.05 \pm 1.12$	$8.3 \pm 2.1$	0.905	0.417	76.0
P2	$2.44 \pm 0.68$	$7.6 \pm 1.1$	0.950	0.294	153.5
P3	$4.01 \pm 0.62$	$8.8 \pm 0.7$	0.981	0.245	409.5
P4	$3.99 \pm 0.57$	$5.4 \pm 0.3$	0.973	0.160	283.5
P5	$1.39 \pm 0.27$	$10.7 \pm 1.4$	0.990	0.150	806.2
P6	$3.00 \pm 0.75$	$18.8 \pm 2.9$	0.979	0.442	381.9
P7	$3.08 \pm 0.83$	$9.5 \pm 1.4$	0.951	0.397	153.9
P8	$0.63 \pm 0.76$	$19.8 \pm 20.8$	0.952	0.236	153.5
P9	$0.37 \pm 0.73$	$43.8 \pm 77.8$	0.920	0.583	91.48

**Table S3.** Non linear fitting of 6-parameters polynomial model for the two-factor central composite (d=2, n=9) experimental design.

Model:  $z = a_0 + a_1x + a_2y + a_3x^2 + a_4y^2 + a_5xy$  ( $r^2 = 0.921$ , fit std. error = 0.607, F-value = 6.980)

Parameters	value $\pm 1$ s.e.	t-value	P
a <sub>0</sub>	$15.686 \pm 3.697$	4.243	0.0240
a <sub>1</sub>	$-5.792 \pm 1.351$	-4.287	0.0233
a <sub>2</sub>	$-0.322 \pm 0.140$	-2.304	0.1046
a <sub>3</sub>	$0.496 \pm 0.150$	3.306	0.0455
a <sub>4</sub>	$-0.001 \pm 0.001$	-0.381	0.7287
a <sub>5</sub>	$0.101 \pm 0.033$	3.073	0.0544

Polymer	z value ( $K_{eq}$ )	z predicted	Residual	Residual %
P1	2.05	2.54	-0.49	23.8
P2	2.44	1.89	0.55	22.4
P3	4.01	3.47	0.54	13.5
P4	3.99	4.14	-0.15	-3.7
P5	1.39	1.71	-0.32	-22.7
P6	3.00	3.24	-0.24	-8.00
P7	3.08	2.84	0.24	7.70
P8	0.63	0.36	0.26	41.7
P9	0.37	0.39	-0.02	-5.31

**Table S4.** Recovery of ciprofloxacin in human urine after dilution 9 + 1 v/v with 50 mmol L<sup>-1</sup> MES buffer, pH 4.5 and MISPE. t-values for pairwise comparison of samples at different concentration.

<b>μg mL<sup>-1</sup></b>	<b>0.2</b>	<b>0.5</b>	<b>1.0</b>	<b>1.5</b>	<b>2.0</b>
<b>0.2</b>	-	0.432	0.821	0.477	0.910
<b>0.5</b>	0.432	-	0.316	0.938	0.290
<b>1.0</b>	0.821	0.316	-	0.352	0.880
<b>1.5</b>	0.477	0.938	0.352	-	0.334
<b>2.0</b>	0.910	0.290	0.880	0.334	-