

Supporting Information

for

Towards universal stimuli-responsive Drug Delivery Systems based on the tetrazole-containing polymers: synthesis of pillar[5]arenes and their self-assembly into nanocontainers

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Table of contents

1. NMR, MALDI TOF MS, IR spectra of the compounds 6-12	3
2. Crystal data	20
3. Figure S24. UV spectra and Bindfit (Fit data to 1:1, 1:2 and 2:1 Host-Guest equilibria).....	21
4. Table S3. Dynamic light scattering.	29
5. Scanning electron microscopy.....	34
6. Fluorescence spectra.....	39

1. NMR, MALDI TOF MS, IR spectra of the compounds 6-12

Figure S1. ^1H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(4-methylbenzylsulfonate-1-ethoxy)-pillar[5]arene (**6**), CDCl_3 , 298 K, 400 MHz.

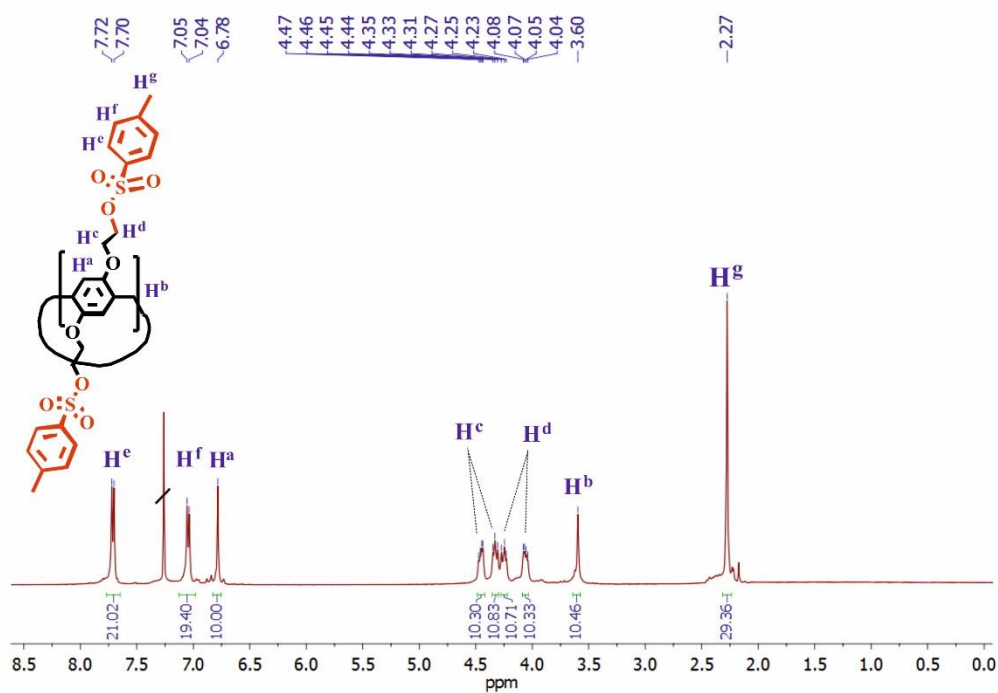


Figure S2. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(4-methylbenzylsulfonate-1-ethoxy)-pillar[5]arene (**6**), CDCl_3 , 298 K, 100 MHz.

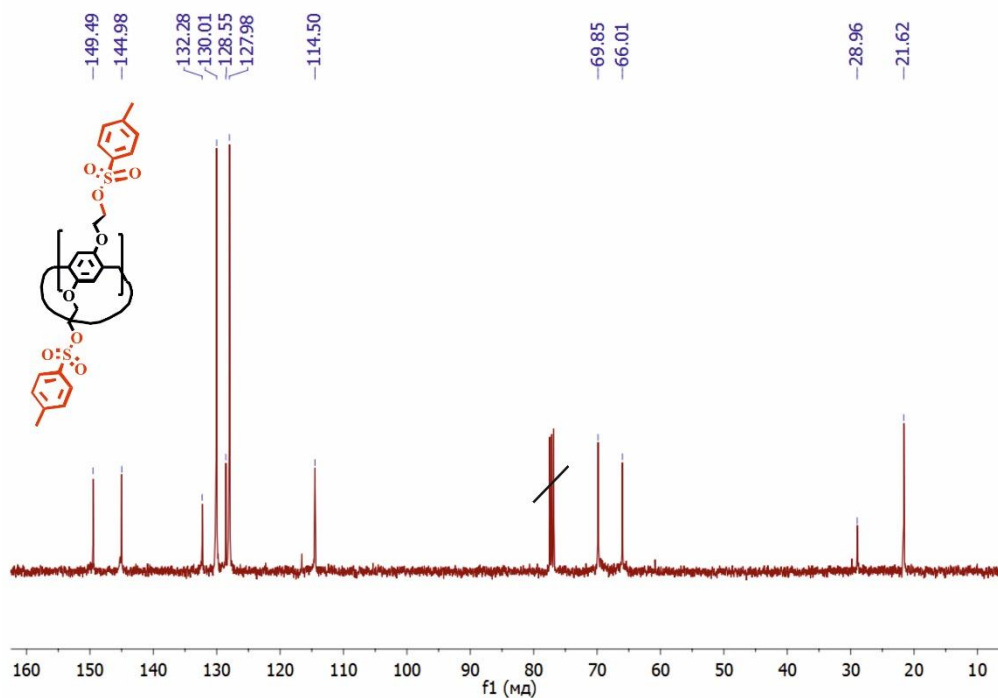


Figure S3. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-(4-methylbenzylsulfonate-1-ethoxy)-pillar[5]arene (**6**).

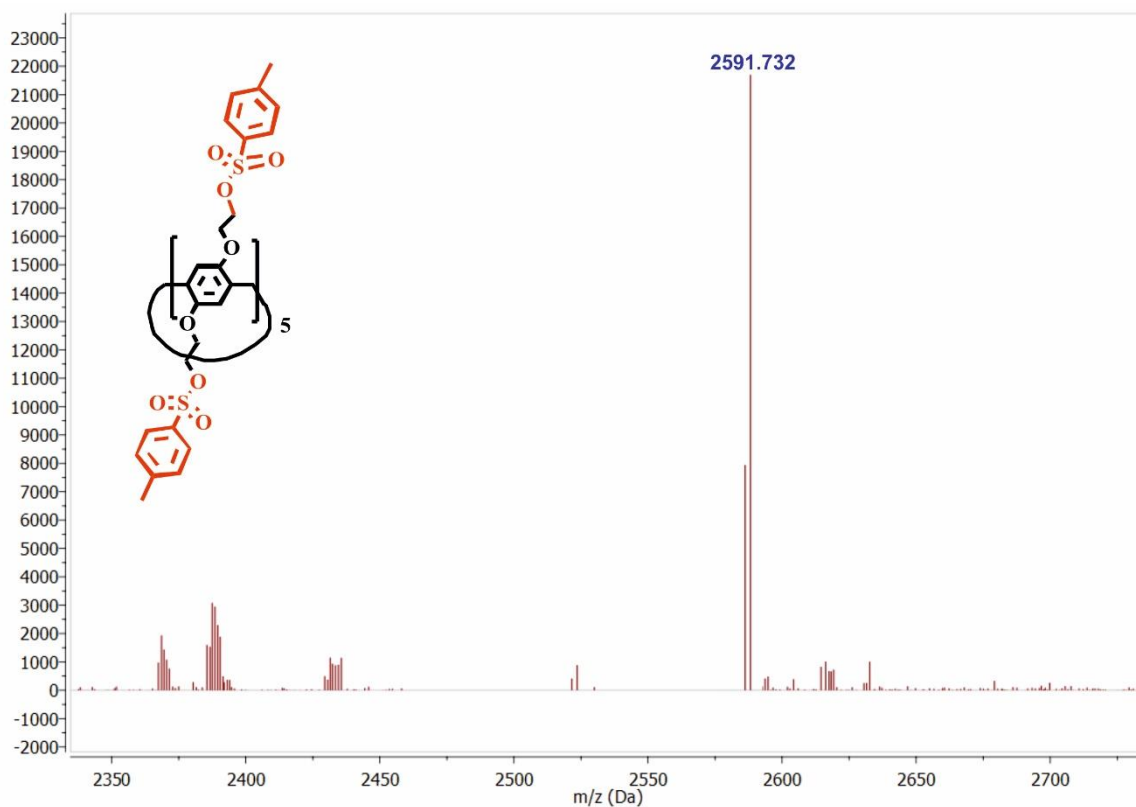


Figure S4. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(4-methylbenzylsulfonate-1-ethoxy)-pillar[5]arene (**6**).

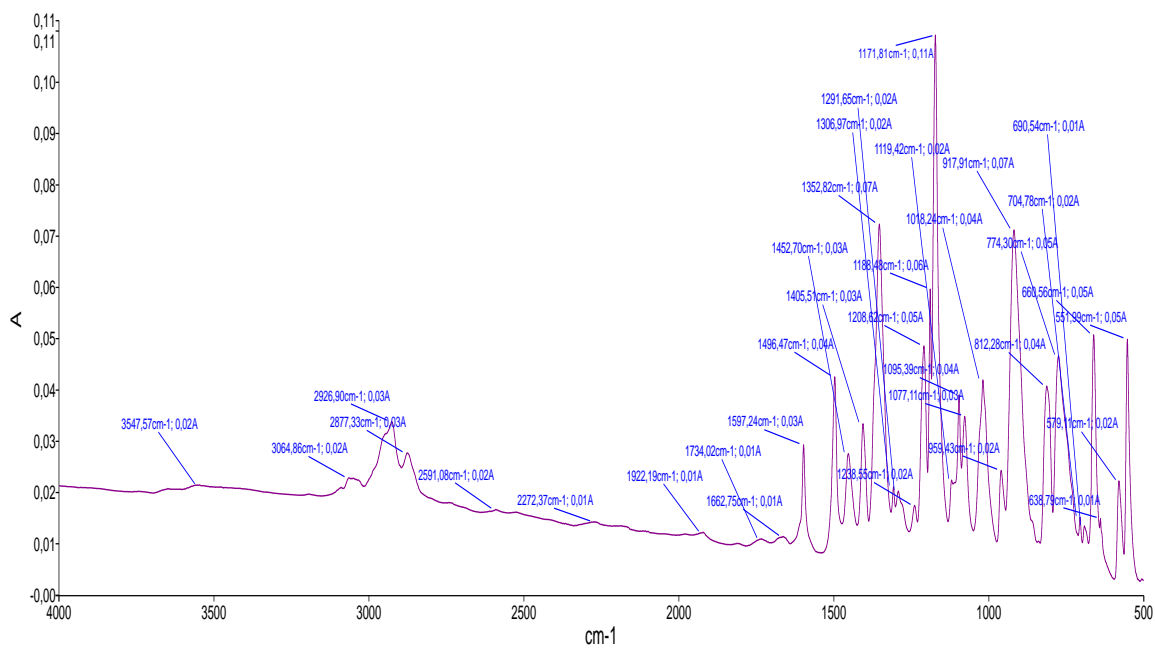


Figure S5. ^1H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[(isoindoline-1,3-dione)propoxy]-pillar[5]arene (**11**), CDCl_3 , 298 K, 400 MHz.

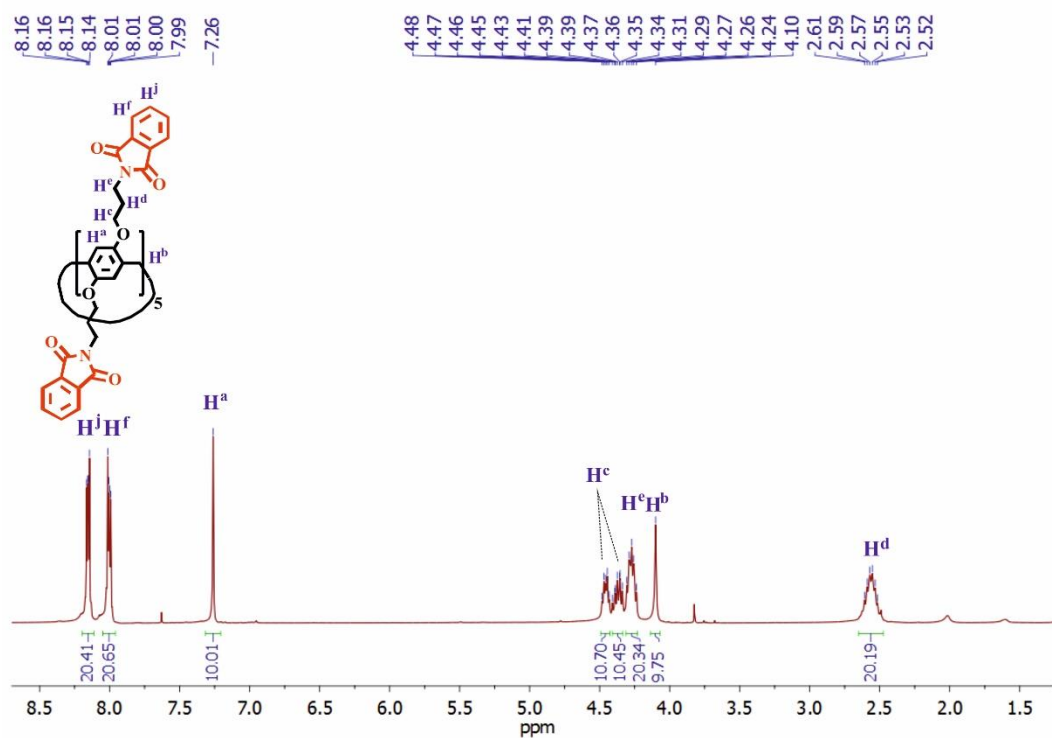


Figure S6. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[(isoindoline-1,3-dione)propoxy]-pillar[5]arene (**11**), CDCl_3 , 298 K, 100 MHz.

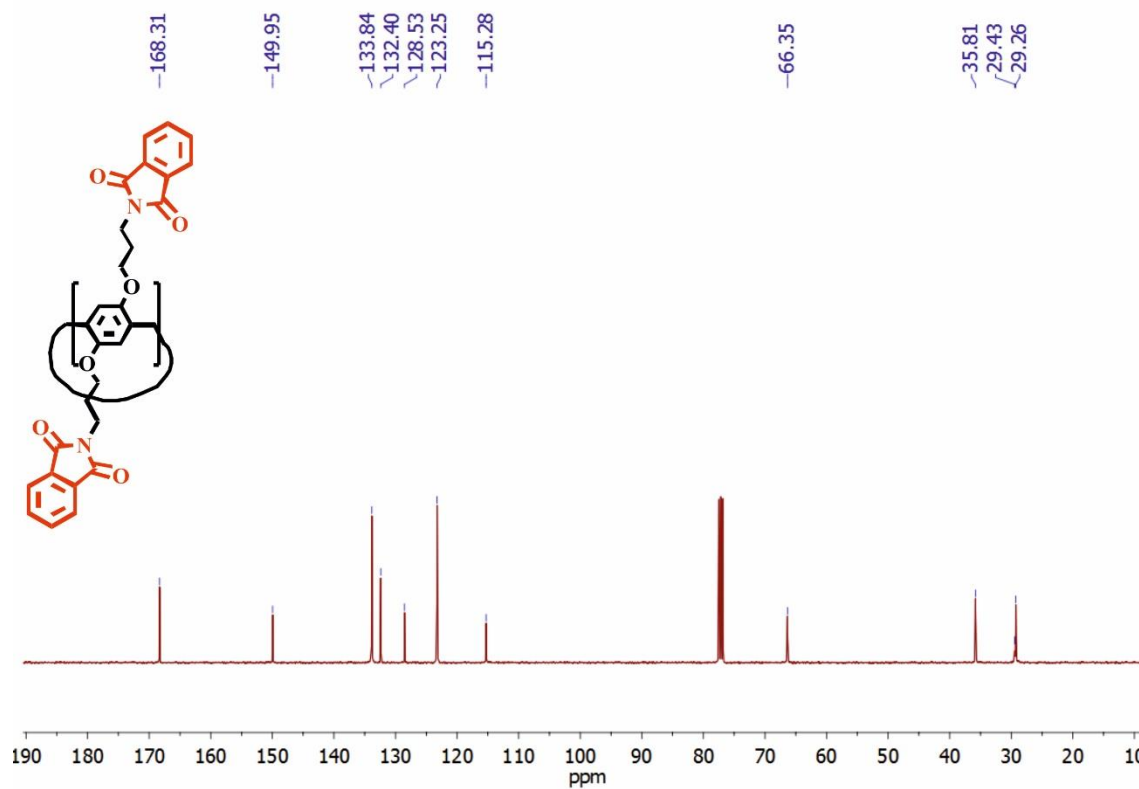


Figure S7. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[(isoindoline-1,3-dione)propoxy]-pillar[5]arene (**11**).

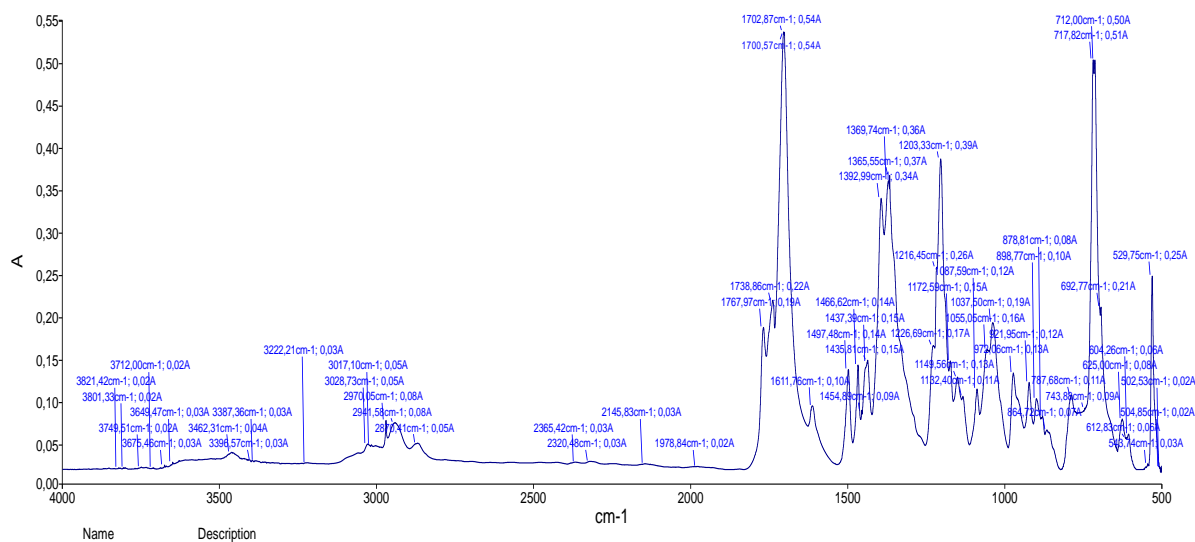


Figure S8. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-[(isoindoline-1,3-dione)propoxy]-pillar[5]arene (**11**)

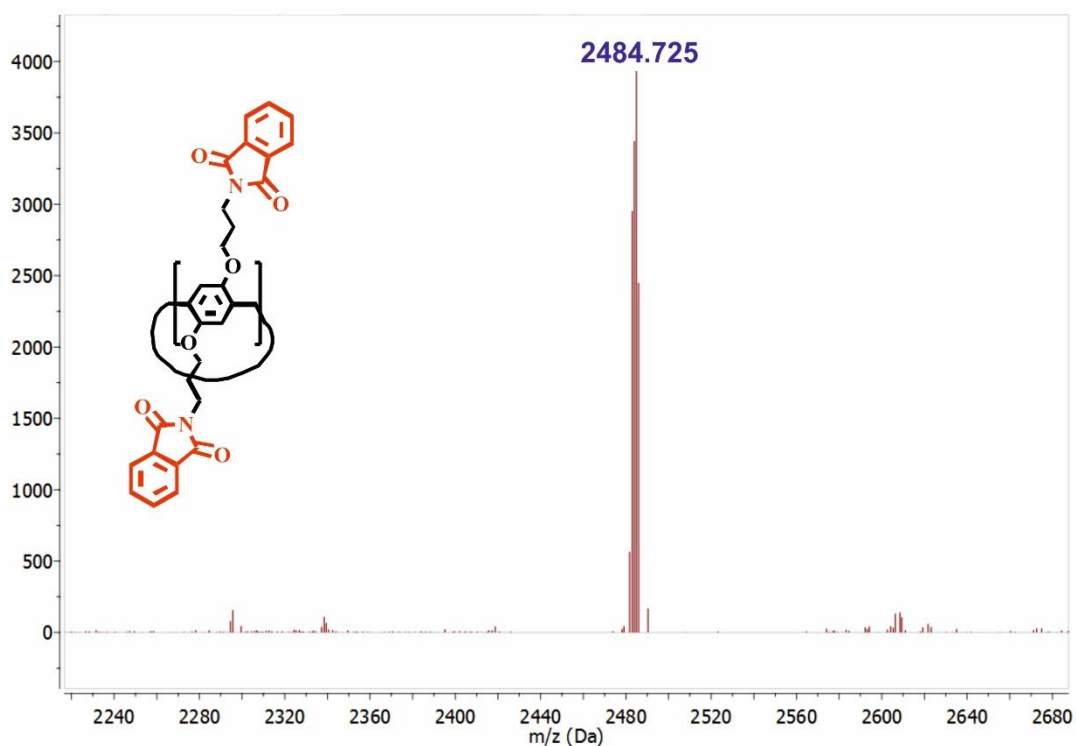


Figure S9. ^1H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(pyrrolidin-1-yl)ethoxy]-pillar[5]arene (7), CDCl_3 , 298 K, 400 MHz.

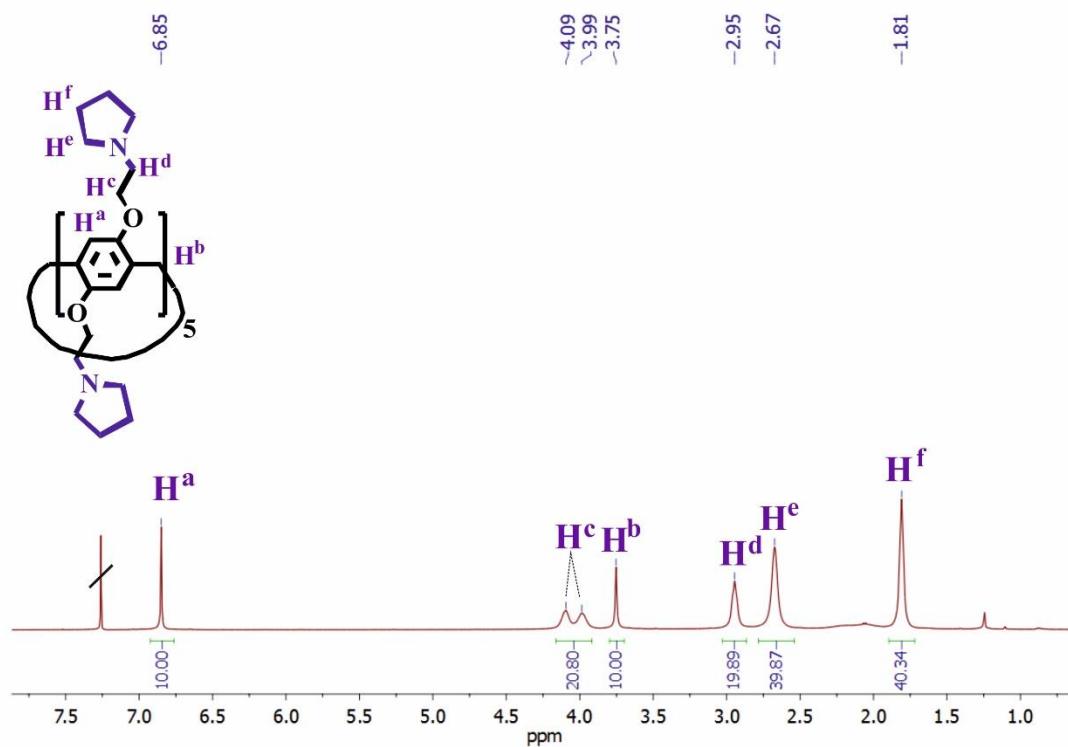


Figure S10. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(pyrrolidin-1-yl)ethoxy]-pillar[5]arene (7), CDCl_3 , 298 K, 100 MHz.

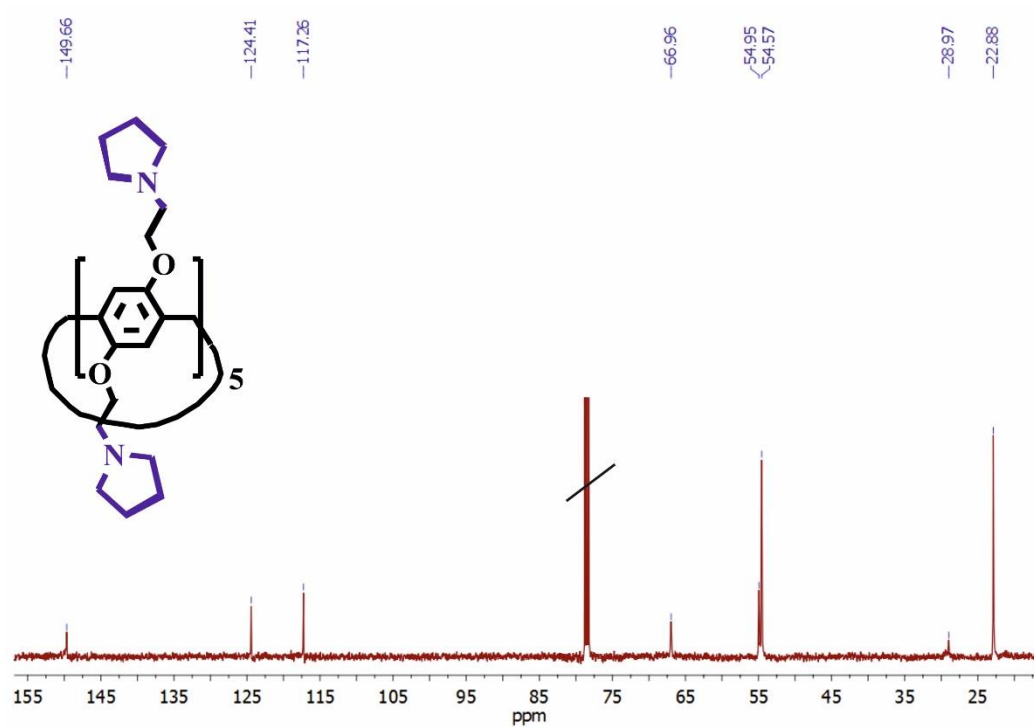


Figure S11. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-[2-(pyrrolidin-1-yl)ethoxy]-pillar[5]arene (7).

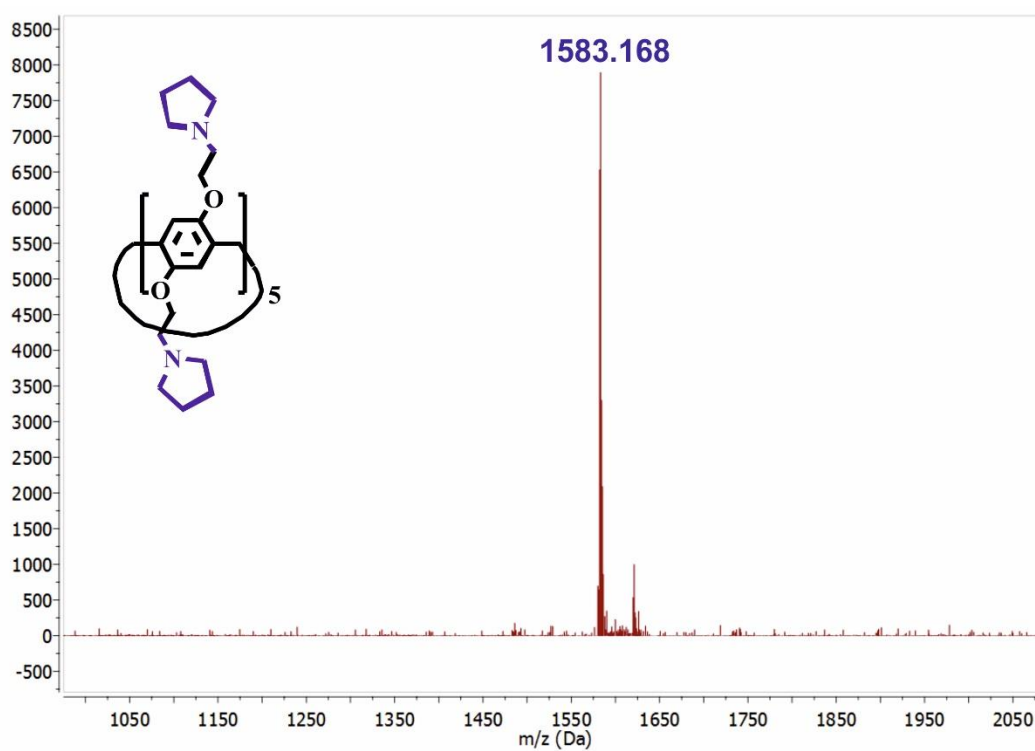


Figure S12. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(pyrrolidin-1-yl)ethoxy]-pillar[5]arene (7).

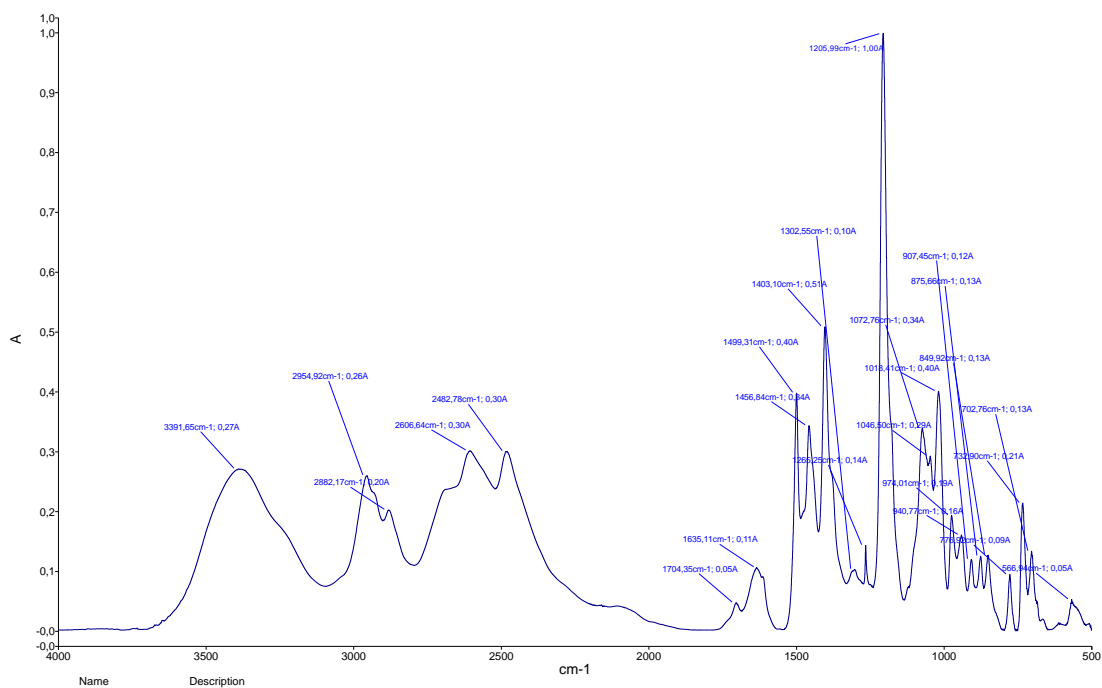


Figure S13. ^1H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(piperidin-1-yl)ethoxy]-pillar[5]arene (8), CDCl_3 , 298 K, 400 MHz.

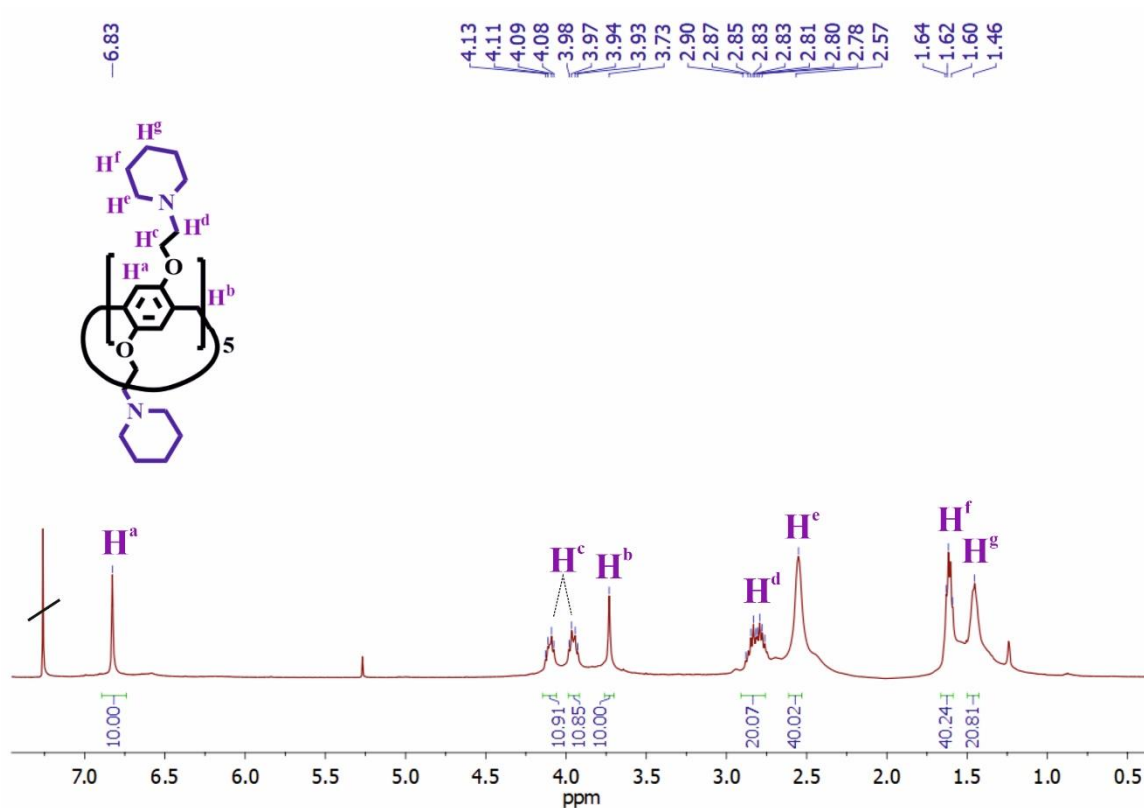


Figure S14. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(piperidin-1-yl)ethoxy]-pillar[5]arene (**8**), CDCl_3 , 298 K, 100 MHz.

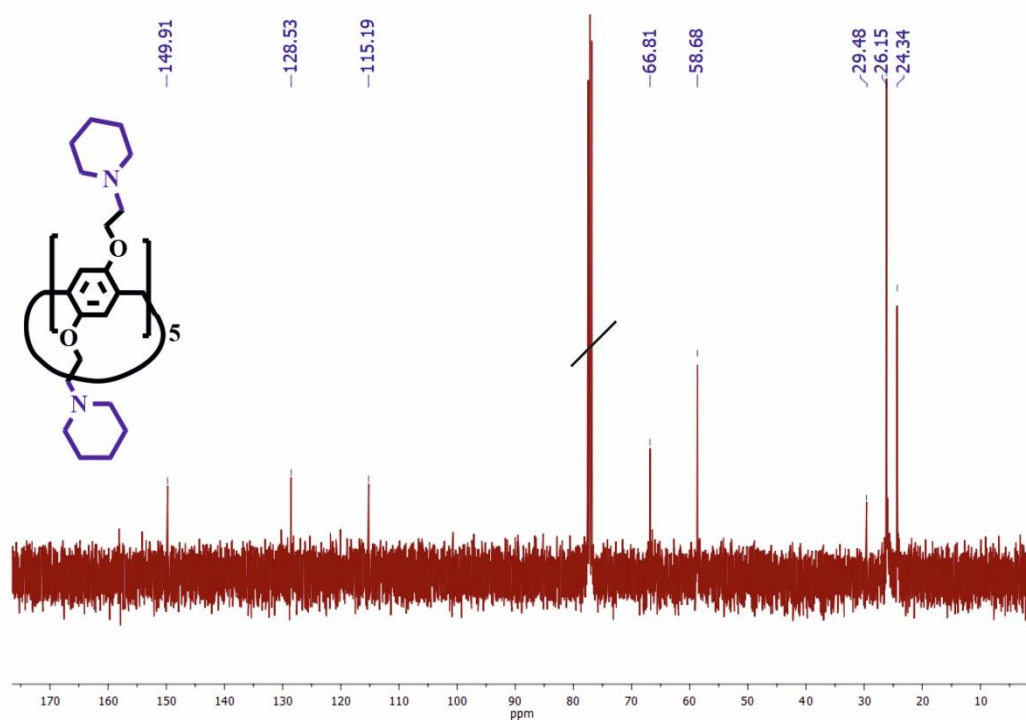


Figure S15. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-[2-(piperidin-1-yl)ethoxy]-pillar[5]arene (**8**).

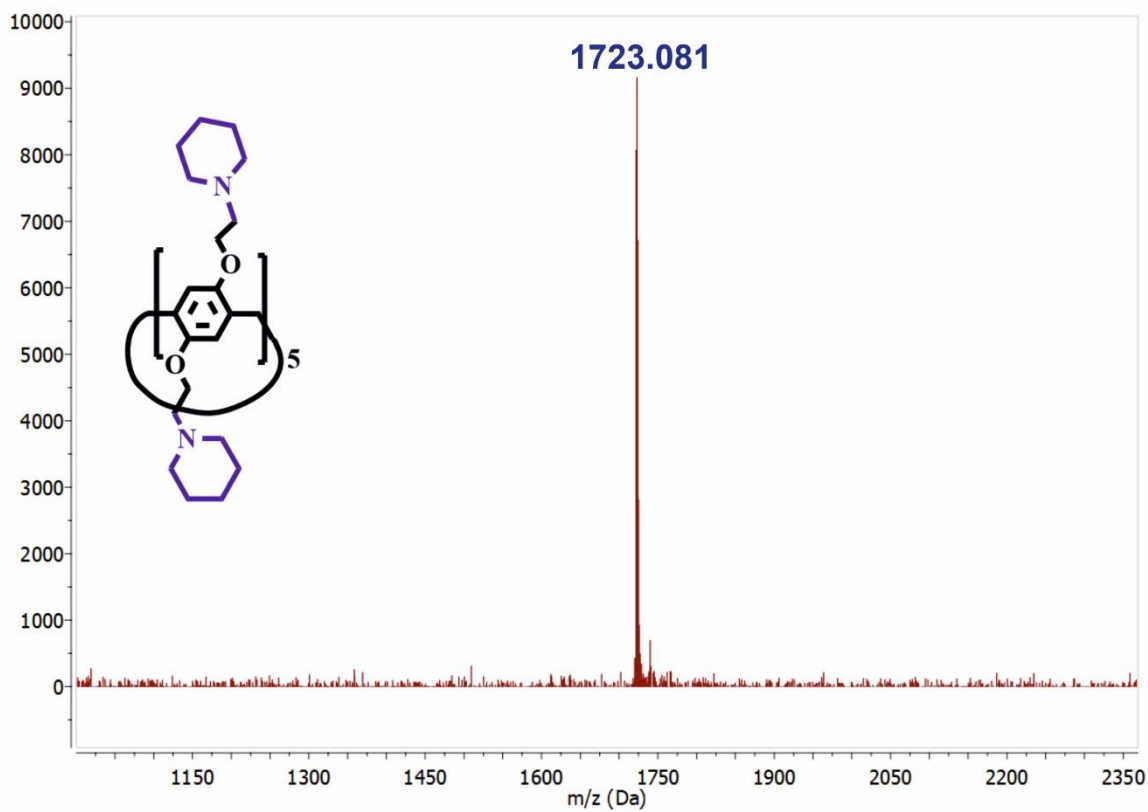


Figure S16. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-[2-(piperidin-1-yl)ethoxy]-pillar[5]arene (8).

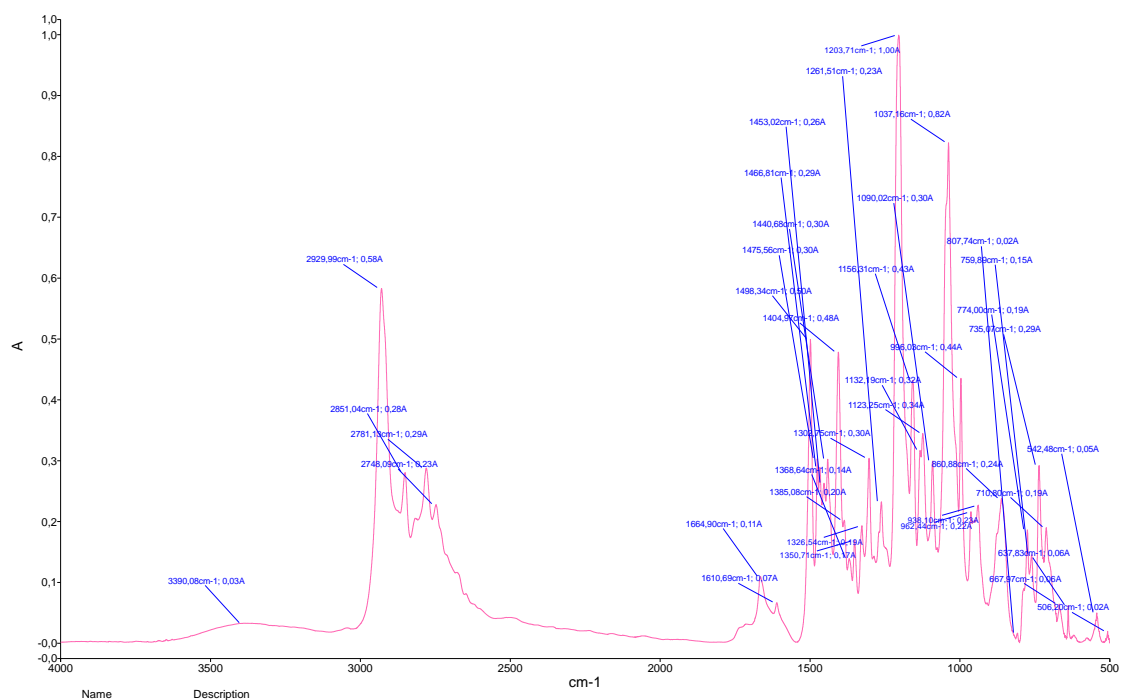


Figure S17. ¹H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(2-morpholinoethoxy)-pillar[5]arene (9), CDCl₃, 298 K, 400 MHz.

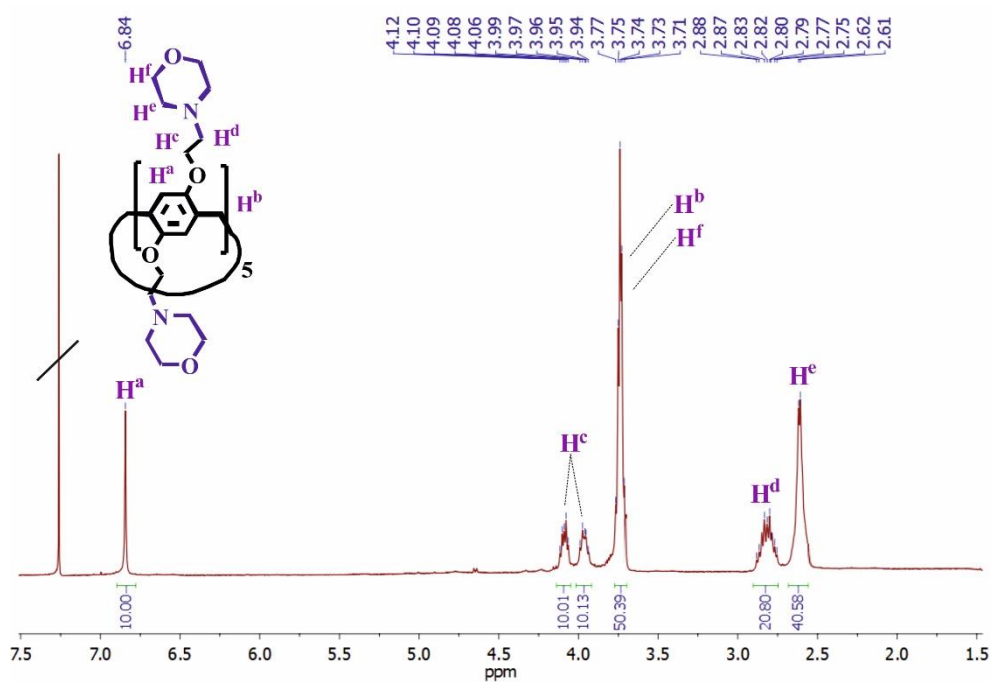


Figure S18. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(2-morpholinoethoxy)-pillar[5]arene (**9**), CDCl_3 , 298 K, 100 MHz.

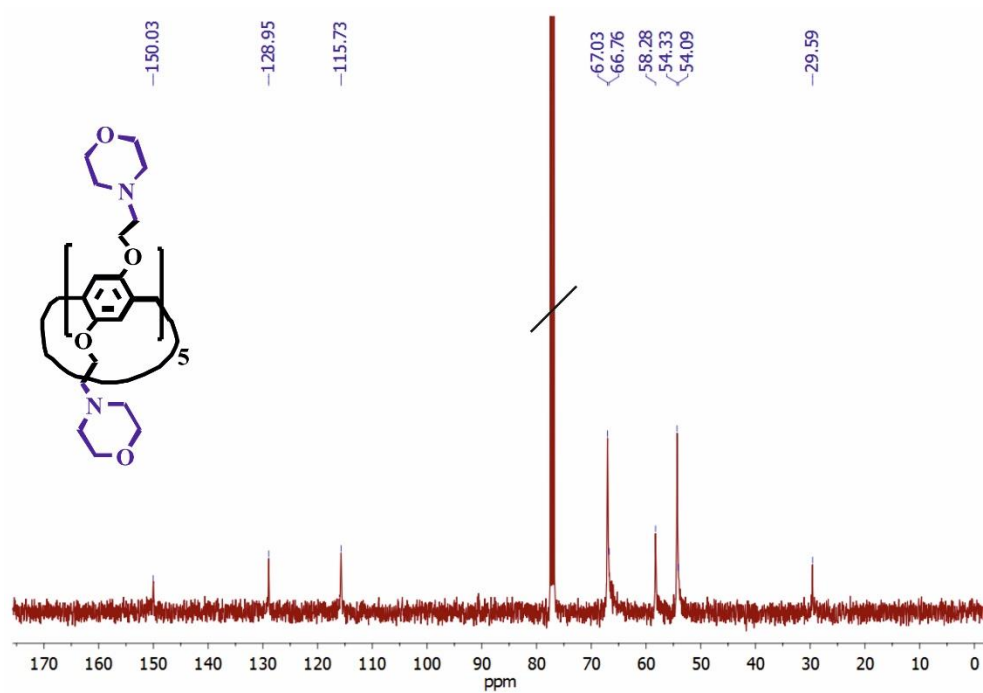


Figure S19. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-(2-morpholinoethoxy)-pillar[5]arene (**9**).

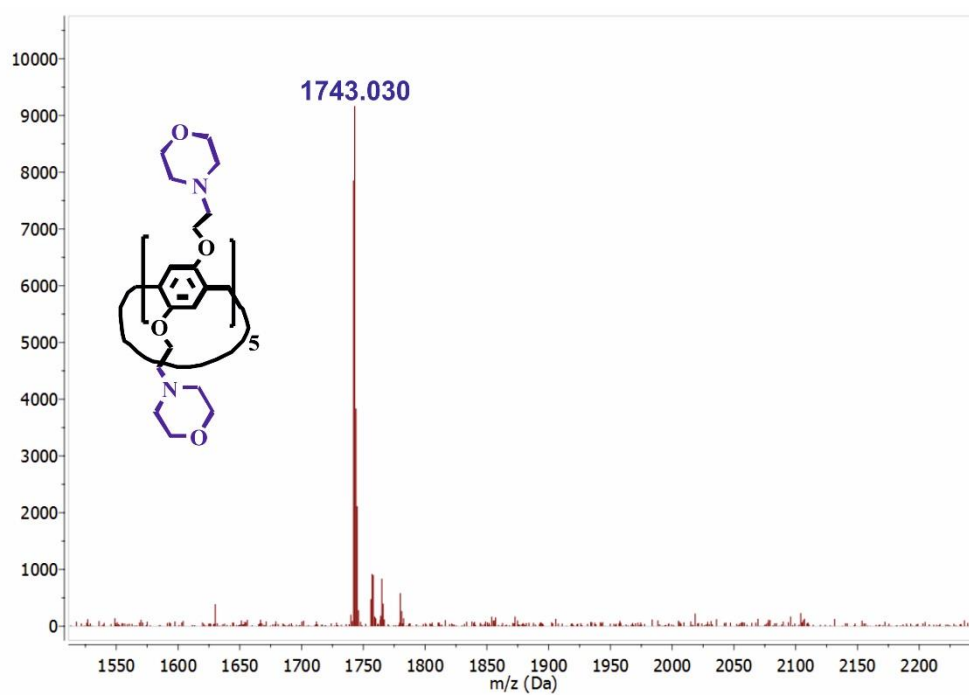


Figure S20. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(2-morpholinoethoxy)-pillar[5]arene (9).

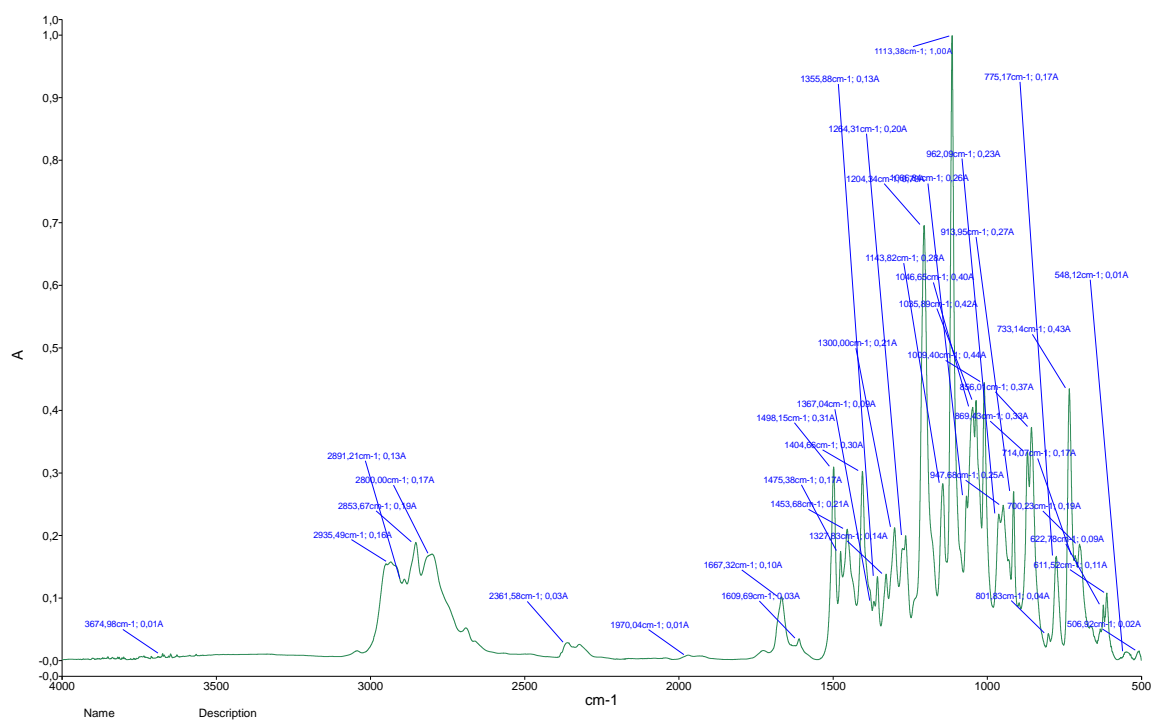


Figure S21. ^1H NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(aminopropoxy)-pillar[5]arene (**12**), D_2O , 298 K, 400 MHz.

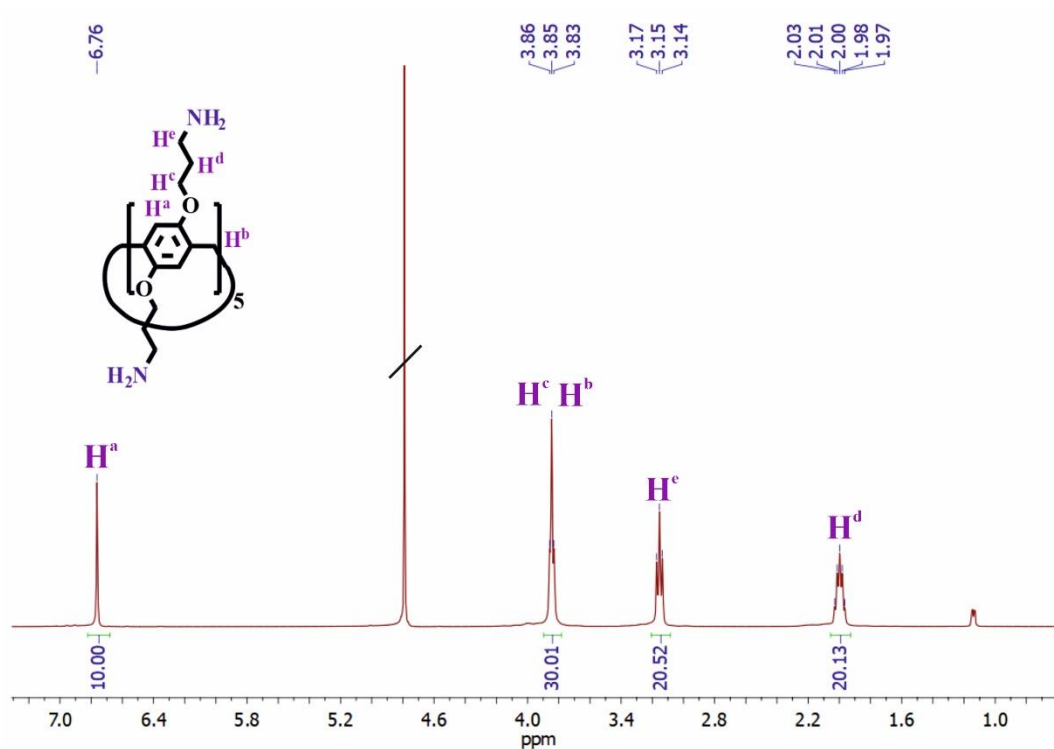


Figure S22. ^{13}C NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(aminopropoxy)-pillar[5]arene (**12**), D_2O , 298 K, 100 MHz.

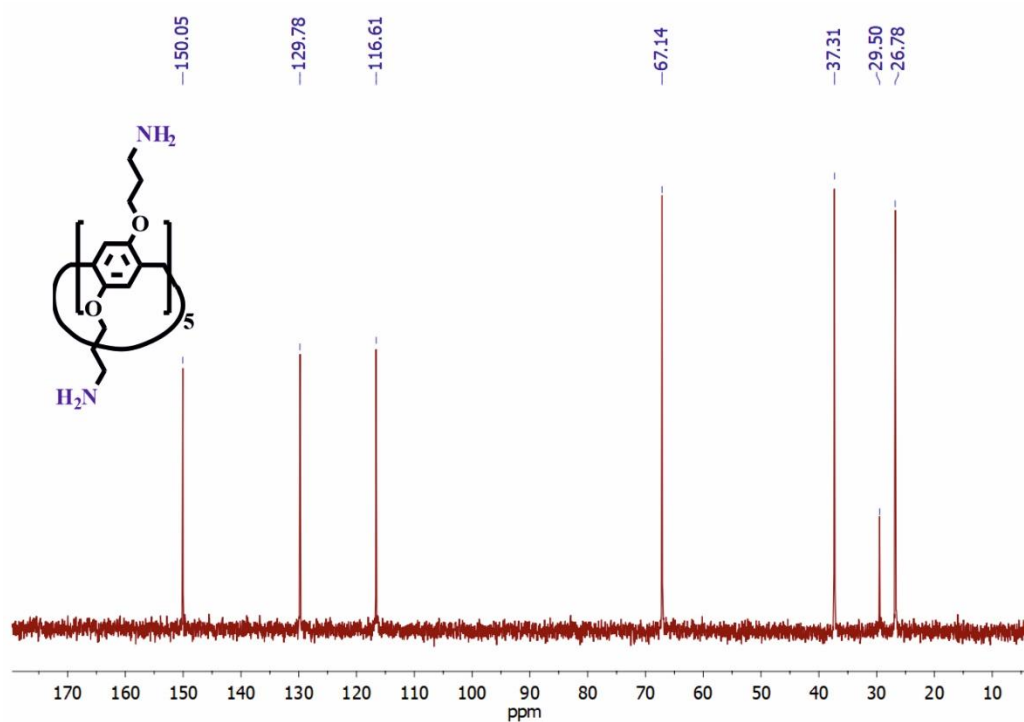


Figure S23. Mass spectrum (MALDI-TOF, 4-nitroaniline matrix) of 4,8,14,18,23,26,28,31,32,35-deca-(aminopropoxy)-pillar[5]arene (**12**).

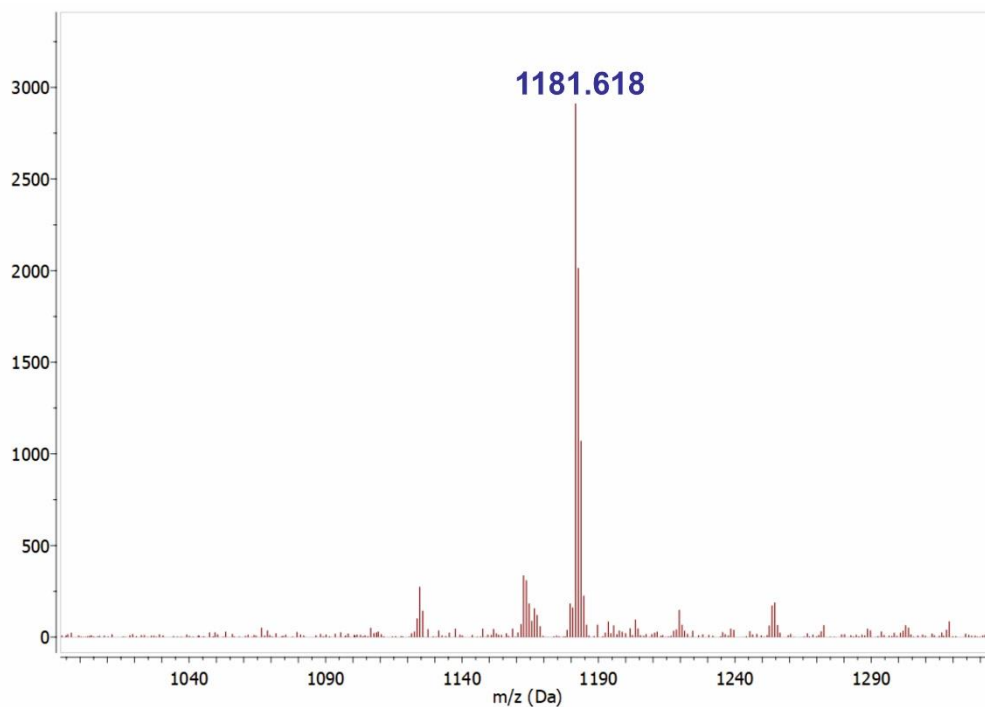


Figure S24. IR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(aminopropoxy)-pillar[5]arene (**12**).

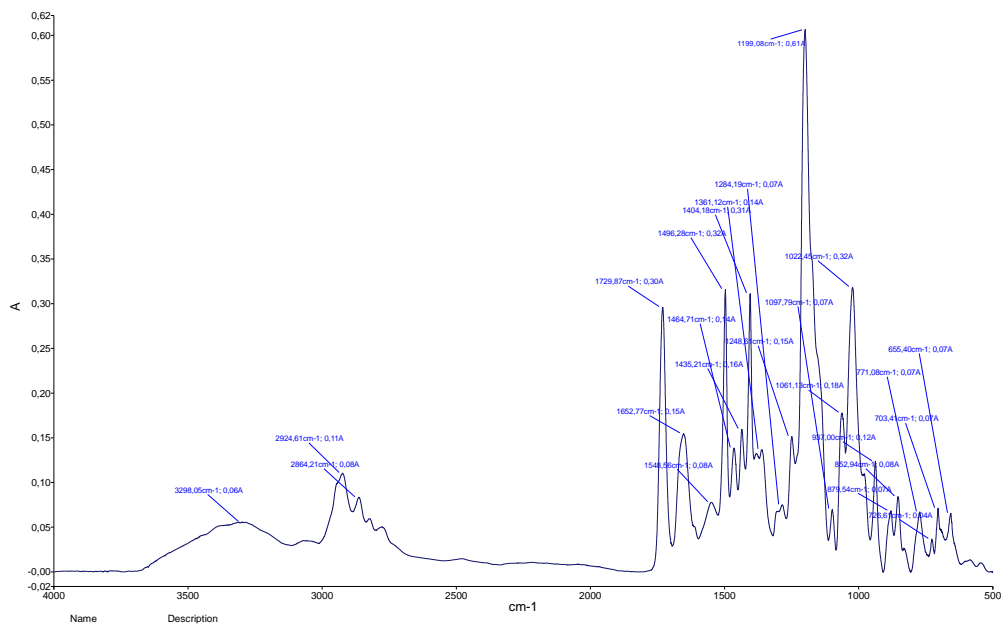


Figure S25. ^1H - ^{13}C HSQC NMR spectrum of 4,8,14,18,23,26,28,31,32,35-deca-(4-methylbenzylsulfonate-1-ethoxy)-pillar[5]arene (**6**), CDCl_3 , 298 K, 400 MHz.

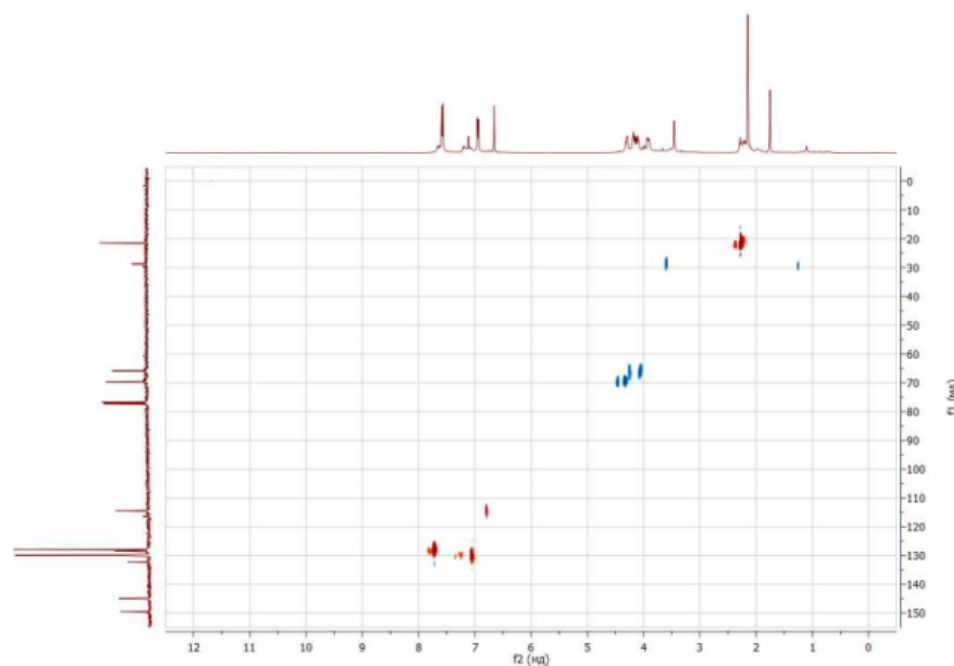


Table S1. Reaction conditions for the synthesis of target macrocycles **6** and **11** from starting compounds **5** and **10**, respectively.

Conditions	Compound s (equiv.)			Solvent	Temp. (°C)	Time (h)	Yield (%)
	Starting compound	Methylene Bridge reagent	Catalyst				
1	5 (1 equiv.)	PFA (3 equiv.)	$\text{BF}_3 \times \text{Et}_2\text{O}$ (1 equiv.)	$\text{CH}_2\text{Cl}-\text{CH}_2\text{Cl}$	0-85	0.5-24	40-64
2	5 (1 equiv.)	PFA (3 equiv.)	$\text{BF}_3 \times \text{Et}_2\text{O}$ (1 equiv.)	CHCl_3	0-50	2-24	5-20 ¹
3	5 (1 equiv.)	PFA (3 equiv.)	$\text{BF}_3 \times \text{Et}_2\text{O}$ (1 equiv.)	CH_2Cl_2	0-30	2-24	0
4	5 (1 equiv.)	PFA (3 equiv.)	AlBr_3 (1 equiv.)	$\text{CH}_2\text{Cl}-\text{CH}_2\text{Cl}$	0-85	0.5-24	2-25
5	5 (1 equiv.)	PFA (3 equiv.)	AlBr_3 (1 equiv.)	CHCl_3	0-50	2-24	4-40 ¹
6	5 (1 equiv.)	PFA (3 equiv.)	AlBr_3 (1 equiv.)	CH_2Cl_2	0-30	2-24	0
7	5 (1 equiv.)	PFA (3 equiv.)	$\text{CF}_3\text{SO}_3\text{H}$ (1 equiv.)	$\text{CH}_2\text{Cl}-\text{CH}_2\text{Cl}$	0-85	0.5-5	35-85
8	5	PFA	$\text{CF}_3\text{SO}_3\text{H}$	CHCl_3	0-50	0.5-5	7-45 ¹

	(1 equiv.)	(3 equiv.)	(1 equiv.)				
9	5 (1 equiv.)	PFA (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CH ₂ Cl ₂	0-30	0.5-5	0
10	5 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-5	1-10
11	5 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CHCl ₃	0-50	2-24	1-15 ¹
12	5 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CH ₂ Cl ₂	0-30	2-24	0
13	5 (1 equiv.)	Paraldehyde (3 equiv.)	BF ₃ ×Et ₂ O (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	5-16
14	5 (1 equiv.)	Paraldehyde (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	10-20
15	10 (1 equiv.)	PFA (3 equiv.)	BF ₃ ×Et ₂ O (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	21-53
16	10 (1 equiv.)	PFA (3 equiv.)	BF ₃ ×Et ₂ O (1 equiv.)	CHCl ₃	0-50	2-24	1-15 ¹
17	10 (1 equiv.)	PFA (3 equiv.)	BF ₃ ×Et ₂ O (1 equiv.)	CH ₂ Cl ₂	0-30	2-24	0
18	10 (1 equiv.)	PFA (3 equiv.)	AlBr ₃ (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	1-17
19	10 (1 equiv.)	PFA (3 equiv.)	AlBr ₃ (1 equiv.)	CHCl ₃	0-50	2-24	1-10 ¹
20	10 (1 equiv.)	PFA (3 equiv.)	AlBr ₃ (1 equiv.)	CH ₂ Cl ₂	0-30	2-24	0
21	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-5	15-74
22	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CHCl ₃	0-50	0.5-5	1-17 ¹
23	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CH ₂ Cl ₂	0-30	0.5-5	0
24	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-5	0
25	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CHCl ₃	0-50	2-24	0
26	10 (1 equiv.)	PFA (3 equiv.)	CF ₃ COOH (1 equiv.)	CH ₂ Cl ₂	0-30	2-24	0
27	10	Paraldehyde	BF ₃ ×Et ₂ O	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	5-10

	(1 equiv.)	(3 equiv.)	(1 equiv.)				
28	10 (1 equiv.)	Paraldehyde (3 equiv.)	CF ₃ SO ₃ H (1 equiv.)	CH ₂ Cl-CH ₂ Cl	0-85	0.5-24	3-12

¹ According to data of NMR spectroscopy.

2. Crystal data

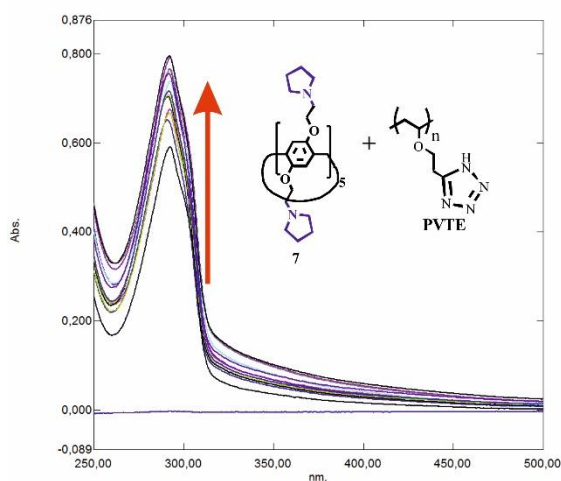
Table S2. Crystal data and structure refinement for **6**, **7** and **11**.

Compound	6	7	11
Formula	C ₁₂₉ H ₁₃₆ N ₂ O ₄₀ S ₁₀	C ₉₇ H ₁₄₃ N ₁₁ O ₁₀	C ₁₅₀ H ₁₂₇ Cl ₃ N ₁₂ O ₃₀
<i>D</i> _{calc.} / g cm ⁻³	1.392	1.167	1.082
μ /mm ⁻¹	2.315	0.596	1.056
Formula Weight	2674.99	1623.22	2683.98
Colour	clear brown	colourless	yellow
Shape	prism	plate	plate
Size/mm ³	0.27×0.18×0.11	0.15×0.09×0.02	0.42×0.23×0.15
<i>T</i> /K	99.9(2)	100.00(10)	99.99(10)
Crystal System	monoclinic	triclinic	triclinic
Space Group	<i>C</i> 2/ <i>c</i>	<i>P</i> -1	<i>P</i> -1
<i>a</i> /Å	27.2340(4)	12.4634(4)	19.9245(3)
<i>b</i> /Å	17.5786(3)	20.2507(12)	20.8844(3)
<i>c</i> /Å	26.8160(5)	20.7444(10)	21.69742(17)
α /°	90	63.291(6)	104.8530(11)
β /°	95.9945(14)	81.416(4)	98.5060(10)
γ /°	90	88.833(4)	104.3794(13)
<i>V</i> /Å ³	12767.6(4)	4618.2(4)	8234.8(2)
<i>Z</i>	4	2	2
<i>Z</i> '	0.5	1	1
Wavelength/Å	1.54184	1.54184	1.54184
Radiation type	Cu K α	Cu K α	Cu K α
θ _{min} /°	2.997	2.414	2.163
θ _{max} /°	77.501	77.237	71.992
Measured Refl.	66822	63746	265502
Independent Refl.	13185	18772	32032
Reflections with <i>I</i> > 2(<i>I</i>)	11003	9154	22857
<i>R</i> _{int}	0.0400	0.1296	0.1000
Parameters	822	1093	1758
Restraints	0	0	63

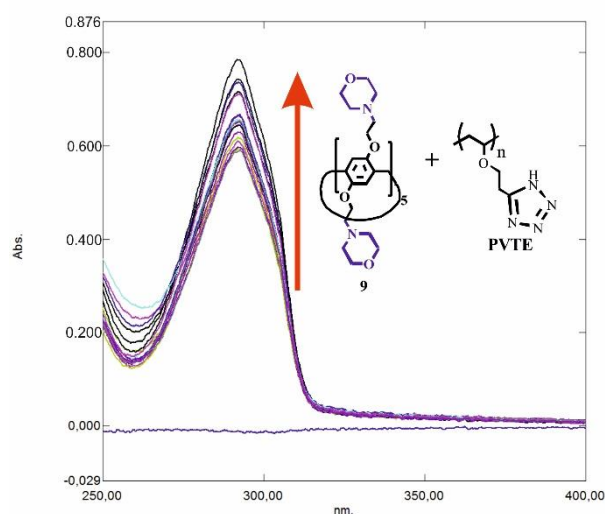
Largest Peak	0.664	1.041	1.004
Deepest Hole	-0.431	-0.397	-1.328
GooF	1.075	1.023	1.436
wR_2 (all data)	0.2288	0.2831	0.3620
wR_2	0.2181	0.2235	0.3401
R_1 (all data)	0.0778	0.1672	0.1259
R_1	0.0706	0.0875	0.1096
CCDC Refcode	2027115	2027117	2027116

3. Figure S26. UV spectra and Bindfit (Fit data to 1:1, 1:2 and 2:1 Host-Guest equilibria)

UV-vis spectra of pillar[5]arene 7 (1×10^{-5} M) at different concentrations of PVTE.



UV-vis spectra of pillar[5]arene 9 (1×10^{-5} M) at different concentrations of PVTE.



UV-vis spectra of Fluorescein (1×10^{-5} M) at different concentrations of pillar[5]arene 7.

UV-vis spectra of Fluorescein (1×10^{-5} M) at different concentrations of pillar[5]arene 9.

Fluorescence spectra of Fluorescein (1×10^{-5} M) at different concentrations of pillar[5]arene 7.

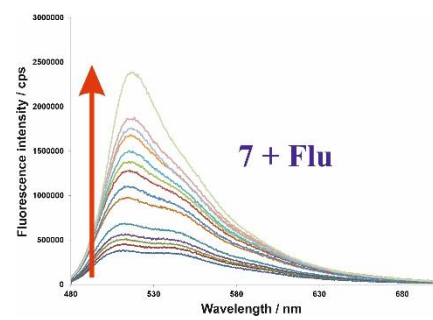
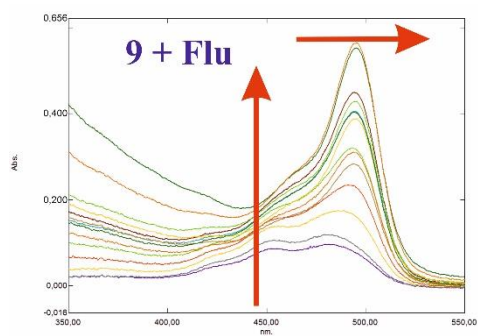
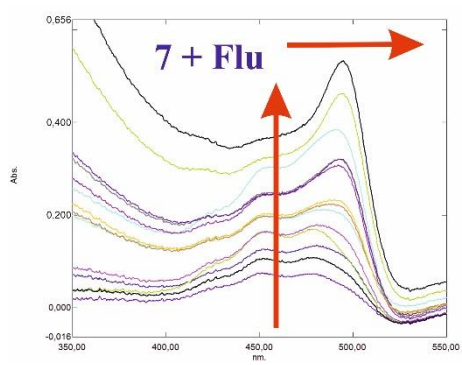


Figure S27. Screenshots taken from the summary window of the website supramolecular.org. This screenshots shows the raw data for UV-vis titration of **7** with PVTE, the data fitted to 1:1 binding model (A), 1:2 binding model (B) and 2:1 binding model (C).

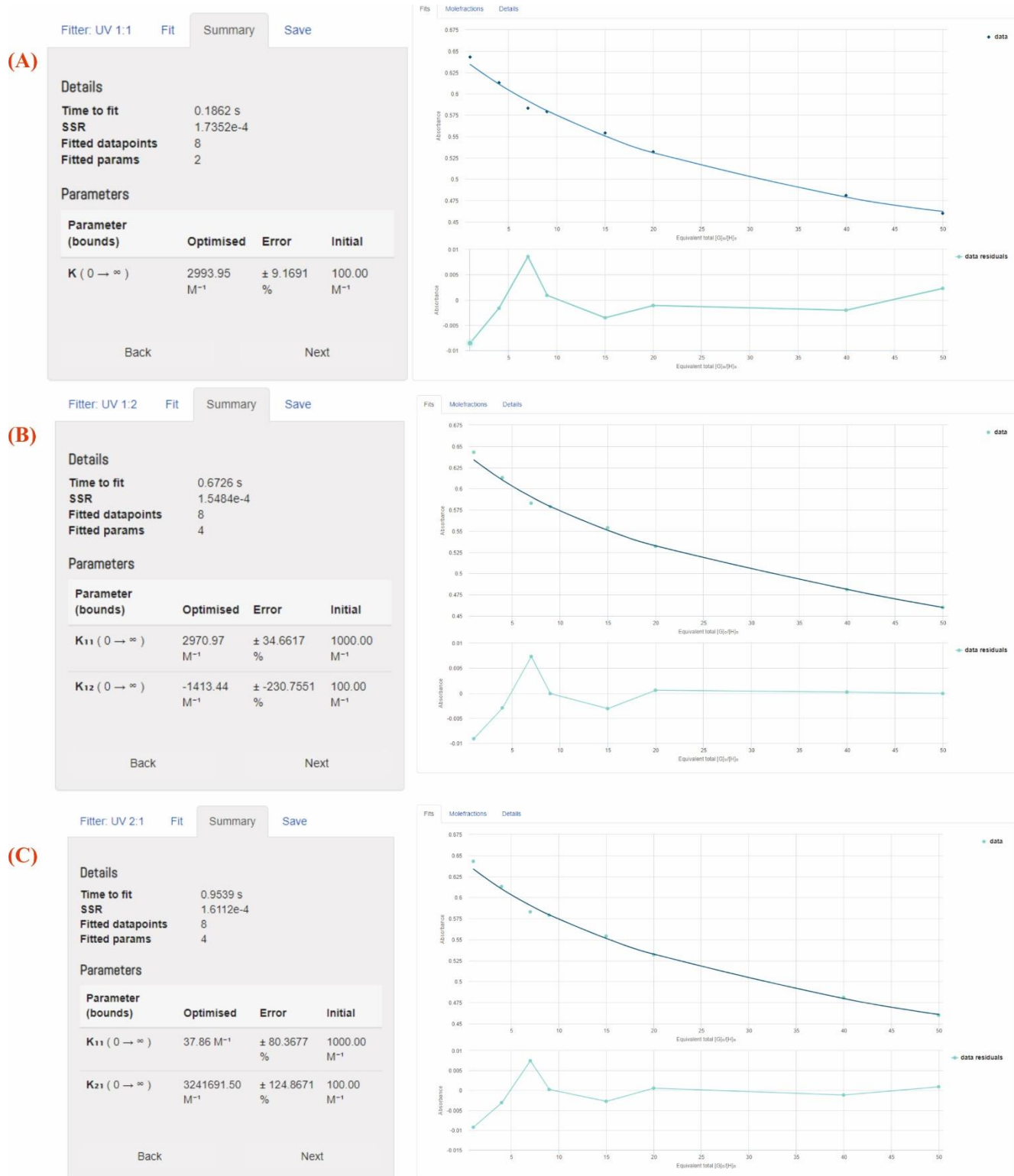


Figure S28. Screenshots taken from the summary window of the website supramolecular.org. This screenshots shows the raw data for UV-vis titration of **9** with PVTE, the data fitted to 1:1 binding model (A), 1:2 binding model (B) and 2:1 binding model (C).

Fitter: UV 1:1 Fit Summary Save

(A)

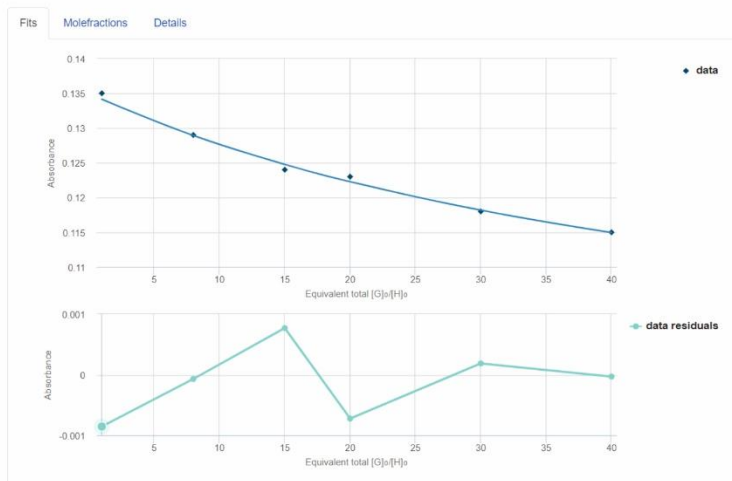
Details

Time to fit 0.1815 s
 SSR 1.8832e-6
 Fitted datapoints 6
 Fitted params 2

Parameters

Parameter (bounds)	Optimised	Error	Initial
$K (0 \rightarrow \infty)$	1898.15 M ⁻¹	± 10.7670 %	100.00 M ⁻¹

Back Next



Fitter: UV 1:2 Fit Summary Save

(B)

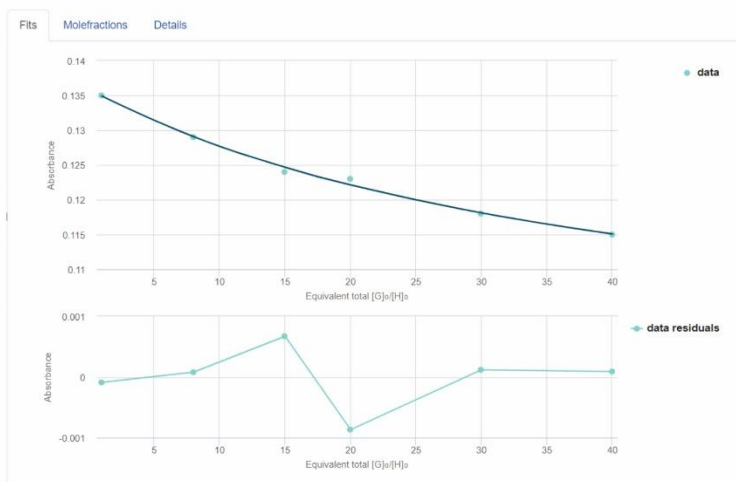
Details

Time to fit 0.8484 s
 SSR 1.2385e-6
 Fitted datapoints 6
 Fitted params 4

Parameters

Parameter (bounds)	Optimised	Error	Initial
$K_{11} (0 \rightarrow \infty)$	152009581692571008.00 M ⁻¹	± 1327040882.8891 %	1000. M ⁻¹
$K_{12} (0 \rightarrow \infty)$	2400.13 M ⁻¹	± 25.2300 %	100.0 M ⁻¹

Back Next



Fitter: UV 2:1 Fit Summary Save

(C)

Details

Time to fit 0.8791 s
 SSR 1.0864e-6
 Fitted datapoints 6
 Fitted params 4

Parameters

Parameter (bounds)	Optimised	Error	Initial
$K_{11} (0 \rightarrow \infty)$	3662.85 M ⁻¹	± 330.1660 %	1000.00 M ⁻¹
$K_{12} (0 \rightarrow \infty)$	2338451.31 M ⁻¹	± 354.3563 %	100.00 M ⁻¹

Back Next

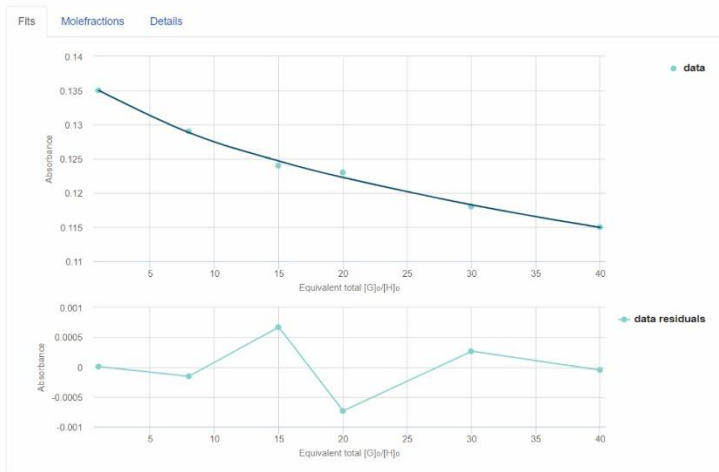


Figure S29. Screenshots taken from the summary window of the website supramolecular.org. This screenshots shows the raw data for UV-vis titration of **7** with Fluorescein, the data fitted to 1:1 binding model (A), 1:2 binding model (B) and 2:1 binding model (C).

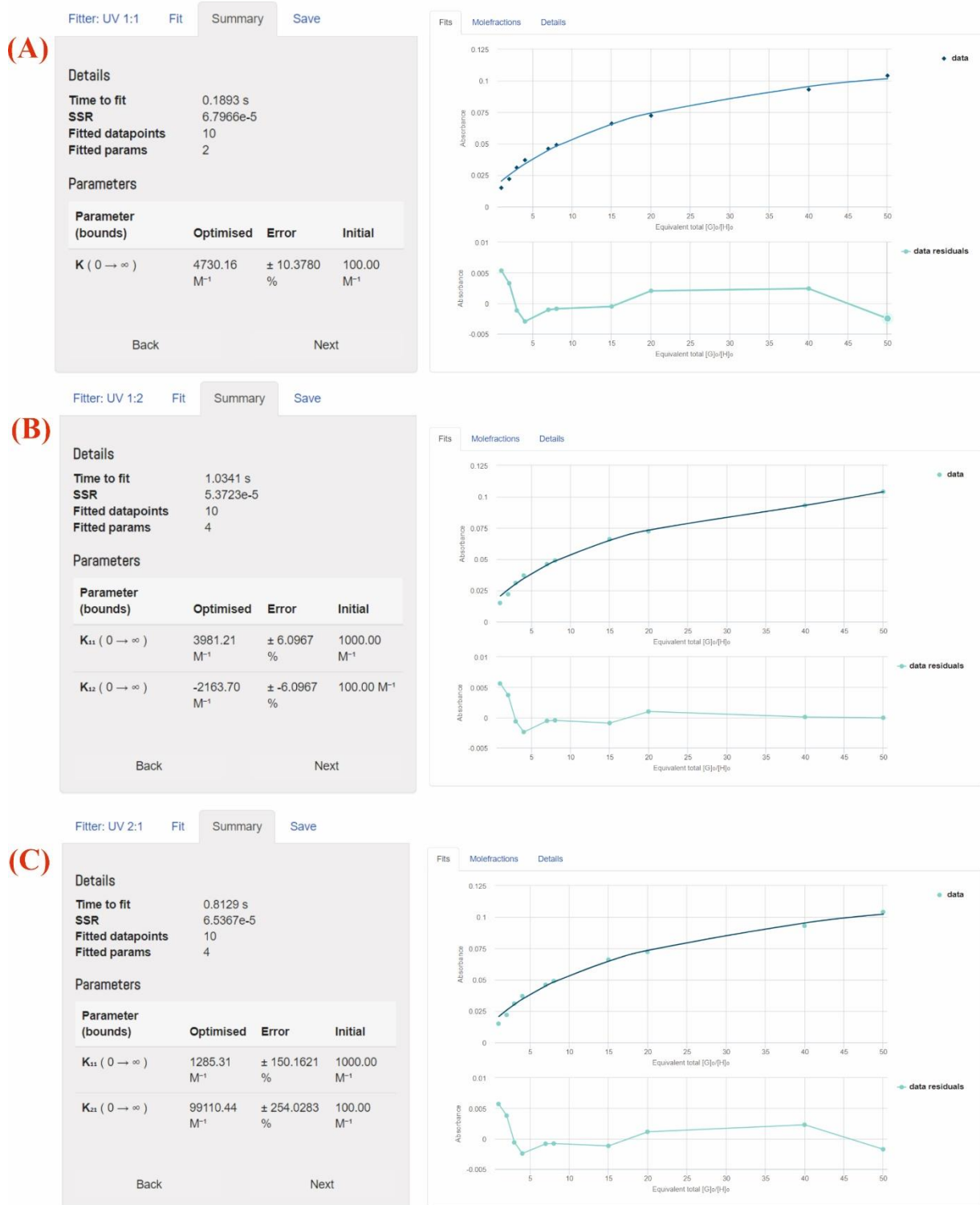


Figure S30. Screenshots taken from the summary window of the website supramolecular.org. This screenshots shows the raw data for UV-vis titration of **9** with Fluorescein, the data fitted to 1:1 binding model (A), 1:2 binding model (B) and 2:1 binding model (C).

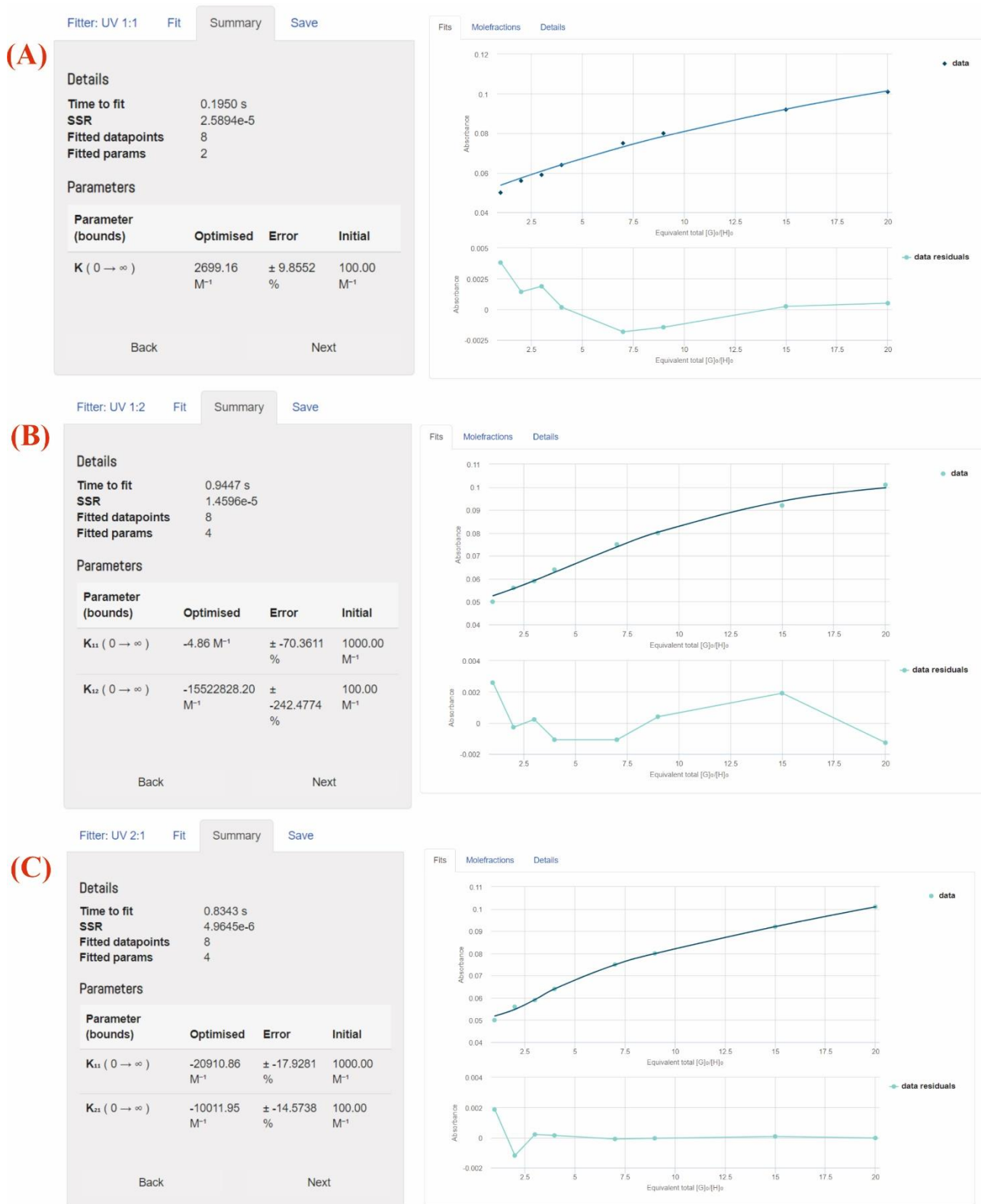


Figure S31. Screenshots taken from the summary window of the website supramolecular.org. This screenshots shows the raw data for fluorescence titration of 7 with Fluorescein, the data fitted to 1:1 binding model (A), 1:2 binding model (B) and 2:1 binding model (C).

(A)

Filter: UV 1:1 Fit Summary Save

Details

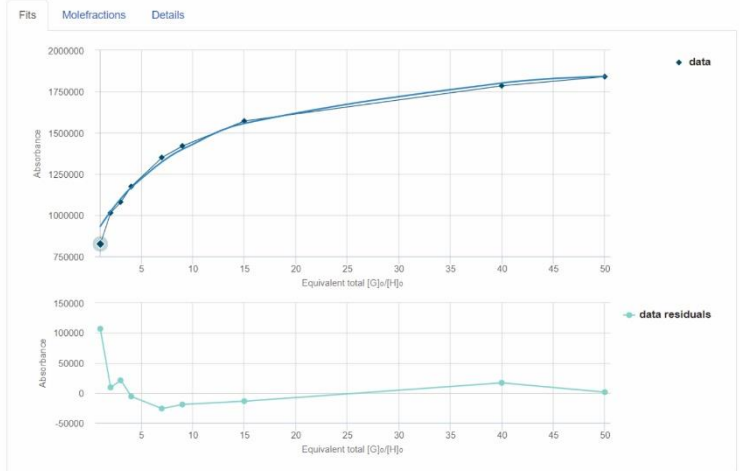
Time to fit 0.1942 s
SSR 13495131713.9540
Fitted datapoints 9
Fitted params 2

Parameters

Parameter (bounds)	Optimised	Error	Initial
K (0 → ∞)	10614.09 M ⁻¹	± 10.0111 %	100.00 M ⁻¹

Back

Next



(B)

Filter: UV 1:2 Fit Summary Save

Details

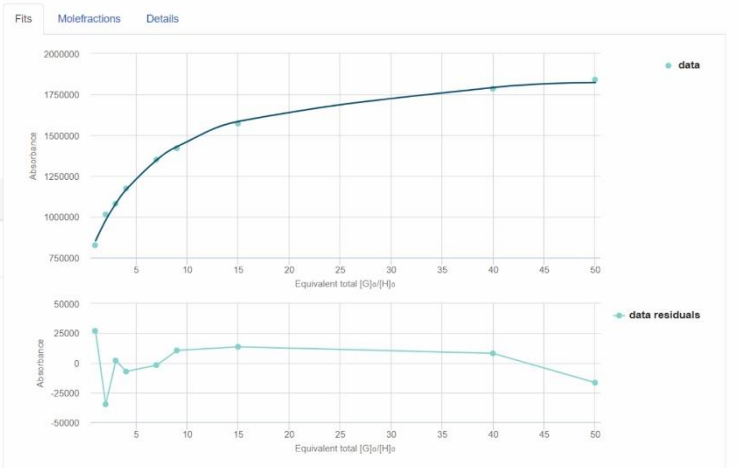
Time to fit 0.9734 s
SSR 2642467558.3104
Fitted datapoints 9
Fitted params 4

Parameters

Parameter (bounds)	Optimised	Error	Initial
K ₁₁ (0 → ∞)	38234790356612352.00 M ⁻¹	± 67084809.1027 %	1000.00 M ⁻¹
K ₁₂ (0 → ∞)	14527.24 M ⁻¹	± 11.7228 %	100.00 M ⁻¹

Back

Next



(C)

Filter: UV 2:1 Fit Summary Save

Details

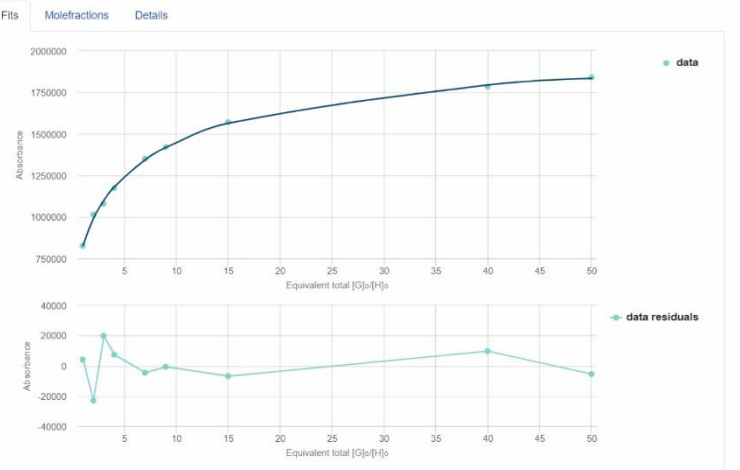
Time to fit 0.9015 s
SSR 1186825413.6849
Fitted datapoints 9
Fitted params 4

Parameters

Parameter (bounds)	Optimised	Error	Initial
K ₁₁ (0 → ∞)	227365.07 M ⁻¹	± 96.7122 %	1000.00 M ⁻¹
K ₁₂ (0 → ∞)	1005266.52 M ⁻¹	± 97.6923 %	100.00 M ⁻¹

Back

Next



4. Table S3. Dynamic light scattering.
Aggregation of the particles for 7 / Flu and 7 / PVTE in EtOH.

Ratio 7 / Flu	V, μ l	C ₇ , M	C _{Flu} , M	Z average (d) , nm	PDI	ζ - potential, mV
1:0	1000	10 ⁻³	0	376.40±34.49	0.42±0.12	-
1:0	1000	10 ⁻⁴	0	406.60±71.45	0.35±0.10	-
1:0	1000	10 ⁻⁵	0	760.20±111.20	0.44±0.29	-
1:1	1000	10 ⁻³	10 ⁻³	428.30±8.12	0.36±0.08	3.20±0.10
1:2	1000	10 ⁻³	2×10 ⁻³	433.90±102.40	0.41 ± 0.02	-
2:1	1000	2×10 ⁻³	10 ⁻³	305.70±48.18	0.43±0.01	-
1:1	1000	10 ⁻⁴	10 ⁻⁴	336.00±13.60	0.33±0.02	1.40±0.72
1:2	1000	10 ⁻⁴	2×10 ⁻⁴	457.50±149.50	0.47±0.13	-
2:1	1000	2×10 ⁻⁴	10 ⁻⁴	456.80±131.40	0.45±0.11	-
1:1	1000	10 ⁻⁵	10 ⁻⁵	155.40±7.16	0.16±0.02	5.94±0.06
1:2	1000	10 ⁻⁵	2×10 ⁻⁵	225.40±7.12	0.23±0.02	2.50±0.58
2:1	1000	2×10 ⁻⁵	10 ⁻⁵	191.00±25.87	0.26±0.05	2.74±0.14
0:1	1000	0	10 ⁻³	460	1	-
0:1	1000	0	10 ⁻⁴	-	-	-
0:1	1000	0	10 ⁻⁵	2328	1	-
7 / PVTE	V, μ l	C ₇ , M	C _{PVTE} , M	Z average (d) , nm	PDI	ζ - potential, mV
50:1	1000	5×10 ⁻⁴	10 ⁻⁵	213.22±4.11	0.31±0.07	-5.44±0.08
10:1	1000	10 ⁻⁴	10 ⁻⁵	116.01±2.26	0.23±0.01	-9.21±0.05
5:1	1000	5×10 ⁻⁵	10 ⁻⁵	198.44±8.10	0.34±0.15	-
2:1	1000	2×10 ⁻⁵	10 ⁻⁵	302.11±1.87	0.35±0.05	-
1:1	1000	10 ⁻⁵	10 ⁻⁵	417.20±11.18	0.39±0.11	-
1:2	1000	10 ⁻⁵	2×10 ⁻⁵	405.54±10.05	0.41±0.17	-
1:5	1000	10 ⁻⁵	5×10 ⁻⁵	440.56±18.16	0.46±0.15	-
1:15	1000	10 ⁻⁵	1.5×10 ⁻⁴	315.20±5.23	0.37±0.28	-
0:1	1000	0	10 ⁻³	670.30±456.40	0.60±0.19	-
0:1	1000	0	10 ⁻⁴	108.80±21.02	0.41±0.09	-
0:1	1000	0	10 ⁻⁵	678.00±486.90	0.52±0.17	-

Aggregation of the particles for 9 / Flu and 9 / PVTE in EtOH.

Ratio 9 / Flu	V, μ l	C ₉ , M	C _{Flu} , M	Z average (d) , nm	PDI	ζ - potential, mV
1:0	1000	10 ⁻³	0	690.10±93.47	0.56±0.34	-
1:0	1000	10 ⁻⁴	0	255.80±96.04	0.58±0.24	-
1:0	1000	10 ⁻⁵	0	262.30±166.60	0.45±0.14	-
1:1	1000	10 ⁻³	10 ⁻³	733.70±209.60	0.48±0.37	-
1:2	1000	10 ⁻³	2×10 ⁻³	1448.00±72.40	0.47 ± 0.12	-
2:1	1000	2×10 ⁻³	10 ⁻³	760.80±20.55	0.49±0.13	-
1:1	1000	10 ⁻⁴	10 ⁻⁴	876.40±438.80	0.62±0.16	-
1:2	1000	10 ⁻⁴	2×10 ⁻⁴	451.10±221.10	0.44±0.21	-
2:1	1000	2×10 ⁻⁴	10 ⁻⁴	363.20±167.30	0.40±0.14	-
1:1	1000	10 ⁻⁵	10 ⁻⁵	273.50±162.90	0.31±0.16	0.05±0.03
1:2	1000	10 ⁻⁵	2×10 ⁻⁵	309.90±37.24	0.40±0.12	-
2:1	1000	2×10 ⁻⁵	10 ⁻⁵	408.80±176.60	0.45±0.14	-

9 / PVTE	V, μl	C ₉ , M	C _{PVTE} , M	Z average (d), nm	PDI	ζ - potential, mV
50:1	1000	5×10^{-4}	10^{-5}	312.25 \pm 7.12	0.31 \pm 0.07	-
10:1	1000	10^{-4}	10^{-5}	125.10 \pm 4.15	0.40 \pm 0.09	-
5:1	1000	5×10^{-5}	10^{-5}	270.21 \pm 5.14	0.38 \pm 0.23	-
2:1	1000	2×10^{-5}	10^{-5}	288.60 \pm 3.13	0.39 \pm 0.10	-
1:1	1000	10^{-5}	10^{-5}	339.45 \pm 10.21	0.40 \pm 0.08	-
1:2	1000	10^{-5}	2×10^{-5}	550.28 \pm 24.35	0.48 \pm 0.21	-
1:5	1000	10^{-5}	5×10^{-5}	1120.50 \pm 56.40	0.50 \pm 0.21	-
1:15	1000	10^{-5}	1.5×10^{-4}	-	-	-

Aggregation of the particles for 7/Flu/PVTE in EtOH.

7/Flu/PVTE	V, μl	C ₇ , M	C _{Flu} , M	C _{PVTE} , M	Z average (d), nm	PDI	ζ - potential, mV
1:1:0.1	1000	10^{-3}	10^{-3}	10^{-4}	418.24 \pm 10.11	0.41 \pm 0.12	-
1:1:1	1000	10^{-3}	10^{-3}	10^{-3}	312.15 \pm 5.10	0.35 \pm 0.14	-
1:1:5	1000	10^{-3}	10^{-3}	5×10^{-3}	214.74 \pm 10.08	0.33 \pm 0.04	-6.13 \pm 0.11
1:1:10	1000	10^{-3}	10^{-3}	10^{-2}	54.11 \pm 1.12	0.20 \pm 0.04	-11.15 \pm 0.10
1:1:0.1	1000	10^{-4}	10^{-4}	10^{-5}	502.11 \pm 20.43	0.54 \pm 0.21	-
1:1:1	1000	10^{-4}	10^{-4}	10^{-4}	395.00 \pm 66.37	0.36 \pm 0.04	-8.25 \pm 0.54
1:1:5	1000	10^{-4}	10^{-4}	5×10^{-4}	188.23 \pm 9.17	0.38 \pm 0.09	-9.16 \pm 0.36
1:1:10	1000	10^{-4}	10^{-4}	10^{-3}	51.04 \pm 2.07	0.21 \pm 0.04	-10.79 \pm 0.08
1:1:0.1	1000	10^{-5}	10^{-5}	10^{-6}	287.20 \pm 34.17	0.32 \pm 0.07	-7.14 \pm 0.24
1:1:1	1000	10^{-5}	10^{-5}	10^{-5}	114.30 \pm 2.10	0.20 \pm 0.01	-10.45 \pm 0.07
1:1:5	1000	10^{-5}	10^{-5}	5×10^{-3}	102.20 \pm 2.24	0.21 \pm 0.05	-11.04 \pm 0.17
1:1:10	1000	10^{-5}	10^{-5}	10^{-4}	48.02 \pm 1.10	0.16 \pm 0.01	-12.81 \pm 0.04

Aggregation of the particles for 7/Flu/PVTE in H₂O/EtOH (100/1).

7/Flu/PVTE	V _{H₂O} , μl	V _{EtOH} , μl	C ₇ , M	C _{Flu} , M	C _{PVTE} , M	Z average (d), nm	PDI	ζ - potential, mV
1:1:0.1	1000	10	10^{-3}	10^{-3}	10^{-4}	388.58 \pm 9.23	0.64 \pm 0.25	-
1:1:1	1000	10	10^{-3}	10^{-3}	10^{-3}	324.90 \pm 7.80	0.53 \pm 0.02	-
1:1:5	1000	10	10^{-3}	10^{-3}	5×10^{-3}	183.45 \pm 5.15	0.36 \pm 0.09	-9.23 \pm 1.47
1:1:10	1000	10	10^{-3}	10^{-3}	10^{-2}	68.26 \pm 1.12	0.09 \pm 0.01	-34.12 \pm 1.94
1:1:0.1	1000	10	10^{-4}	10^{-4}	10^{-5}	1560.40 \pm 145.24	1	-
1:1:1	1000	10	10^{-4}	10^{-4}	10^{-4}	417.40 \pm 54.39	0.42 \pm 0.15	-
1:1:5	1000	10	10^{-4}	10^{-4}	5×10^{-4}	245.36 \pm 2.23	0.35 \pm 0.04	-9.11 \pm 2.13
1:1:10	1000	10	10^{-4}	10^{-4}	10^{-3}	175.00 \pm 3.05	0.29 \pm 0.01	-26.40 \pm 1.45

Aggregation of the particles for 7/Flu/PVTE in buffer (pH 9-2).

7/Flu/PVTE	V_{buff}, μl (pH)	V_{EtOH}, μl	C₇ (EtOH), M	C_{Flu} (EtOH), M	C_{PVTE} (EtOH), M	Z_{average} (d), nm	PDI
1:1:10	1000 (9)	10	10 ⁻³	10 ⁻³	10 ⁻²	82.99±0.47	0.15±0.02
	1000 (7)	10	10 ⁻³	10 ⁻³	10 ⁻²	84.33±0.72	0.12±0.01
	1000 (5)	10	10 ⁻³	10 ⁻³	10 ⁻²	489.33±10.30	0.43±0.15
	1000 (4)	10	10 ⁻³	10 ⁻³	10 ⁻²	688.40±25.69	0.63±0.09
	1000 (2)	10	10 ⁻³	10 ⁻³	10 ⁻²	737.80±45.58	0.55±0.07

Figure S32. Size distribution of the particles by intensity for PVTE ($1 \times 10^{-5} \text{M}$) in ethanol.

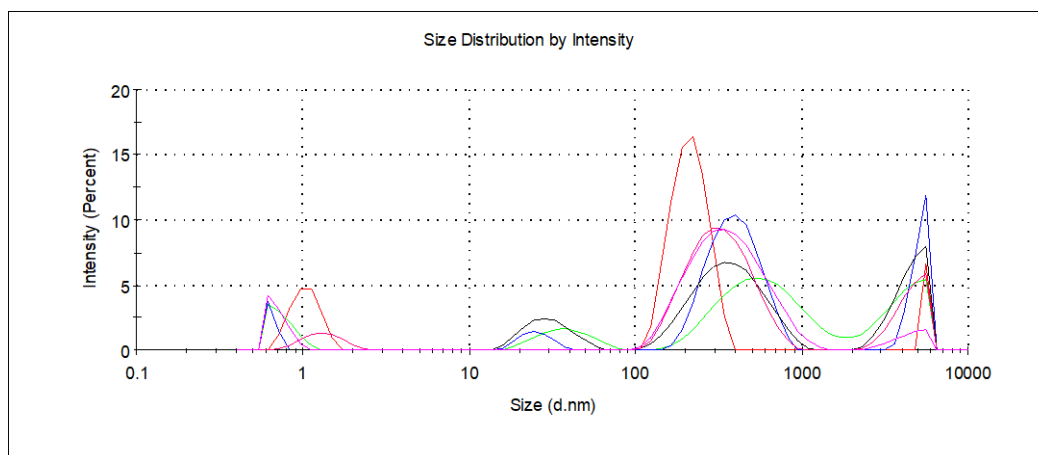


Figure S33. Size distribution of the particles by intensity for 7 ($1 \times 10^{-5} \text{M}$) in ethanol.

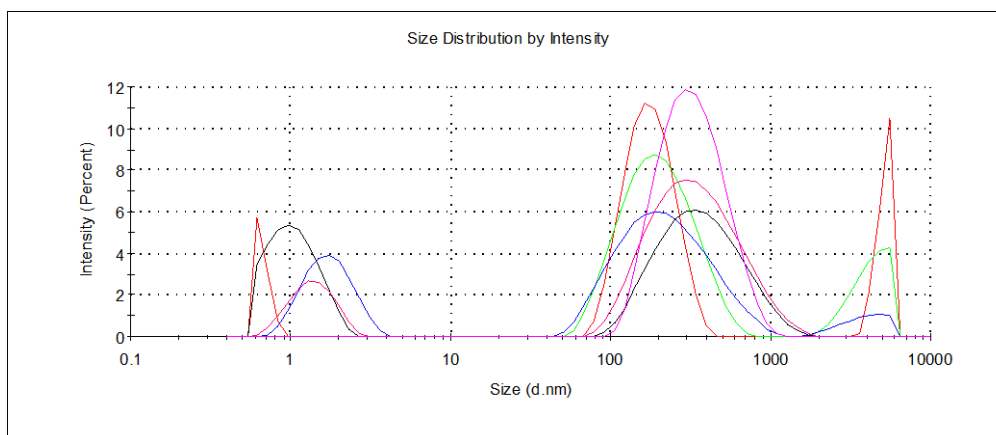


Figure S34. Size distribution of the particles by intensity for 7 ($1 \times 10^{-4} \text{M}$) / PVTE ($1 \times 10^{-5} \text{M}$) (10:1) in ethanol.

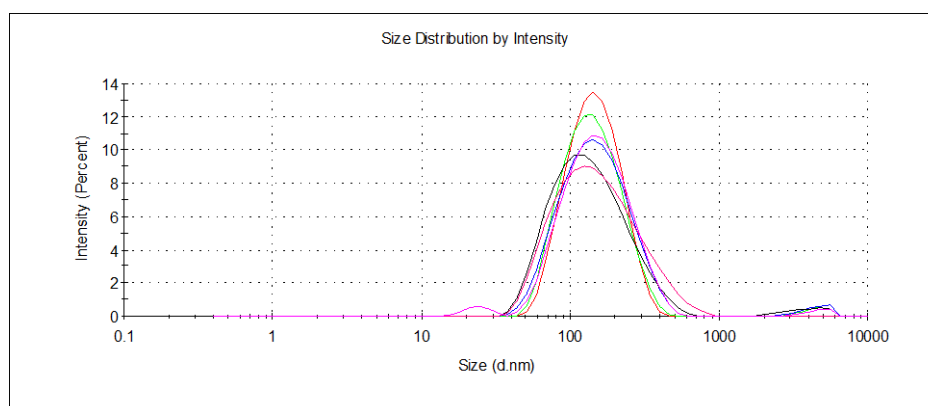
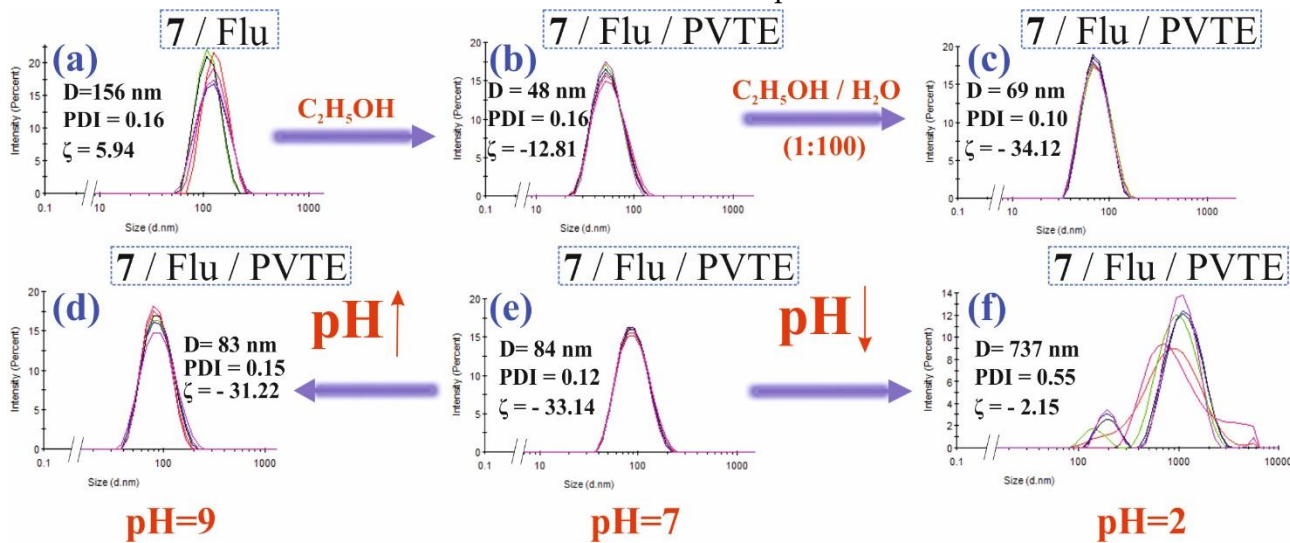


Figure S35 Size distribution of the particles by intensity for 7/Flu, 7/Flu/PVTE in ethanol and buffer at different pH.



5. Scanning electron microscopy.

Figure S36 SEM image of silicon substrate.

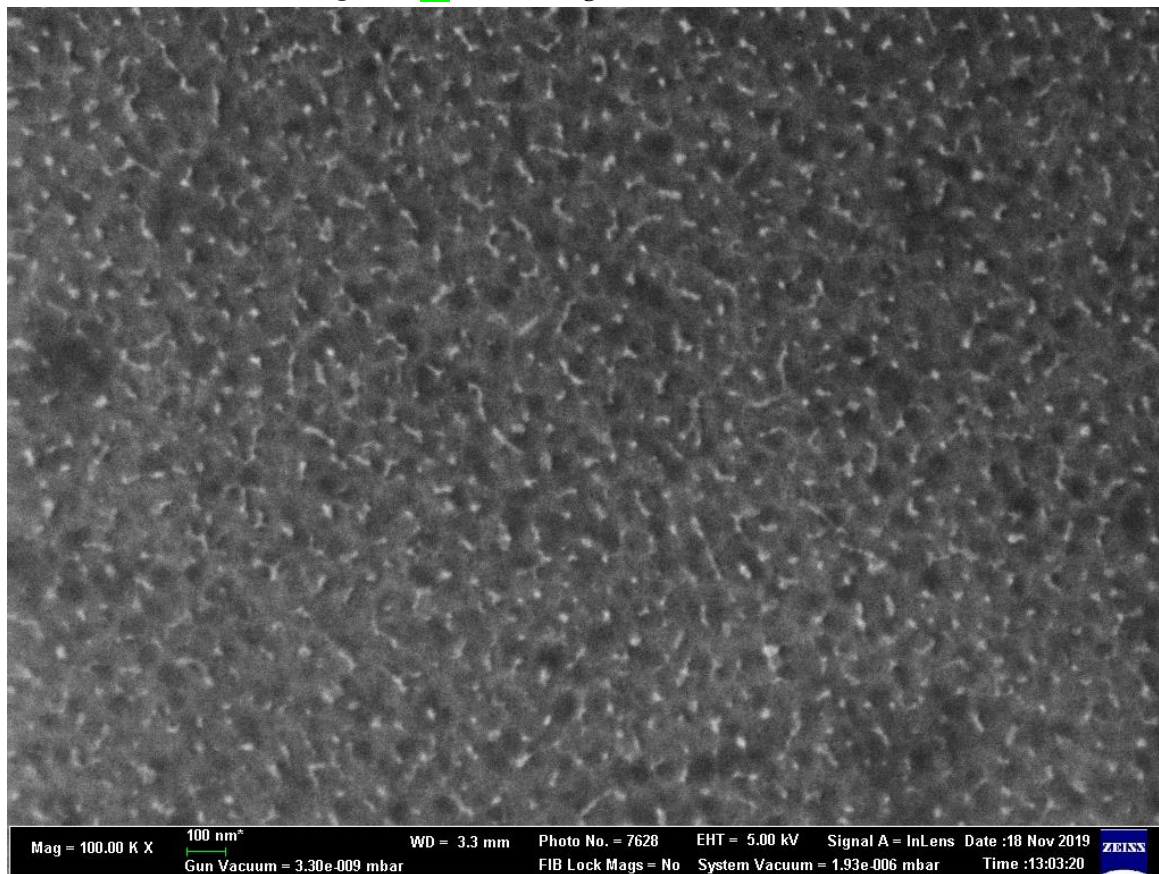


Figure S37. SEM image of 7/Flu/PVTE (1×10^{-5} M) after the solvent ($\text{H}_2\text{O}/\text{EtOH}$) evaporation.

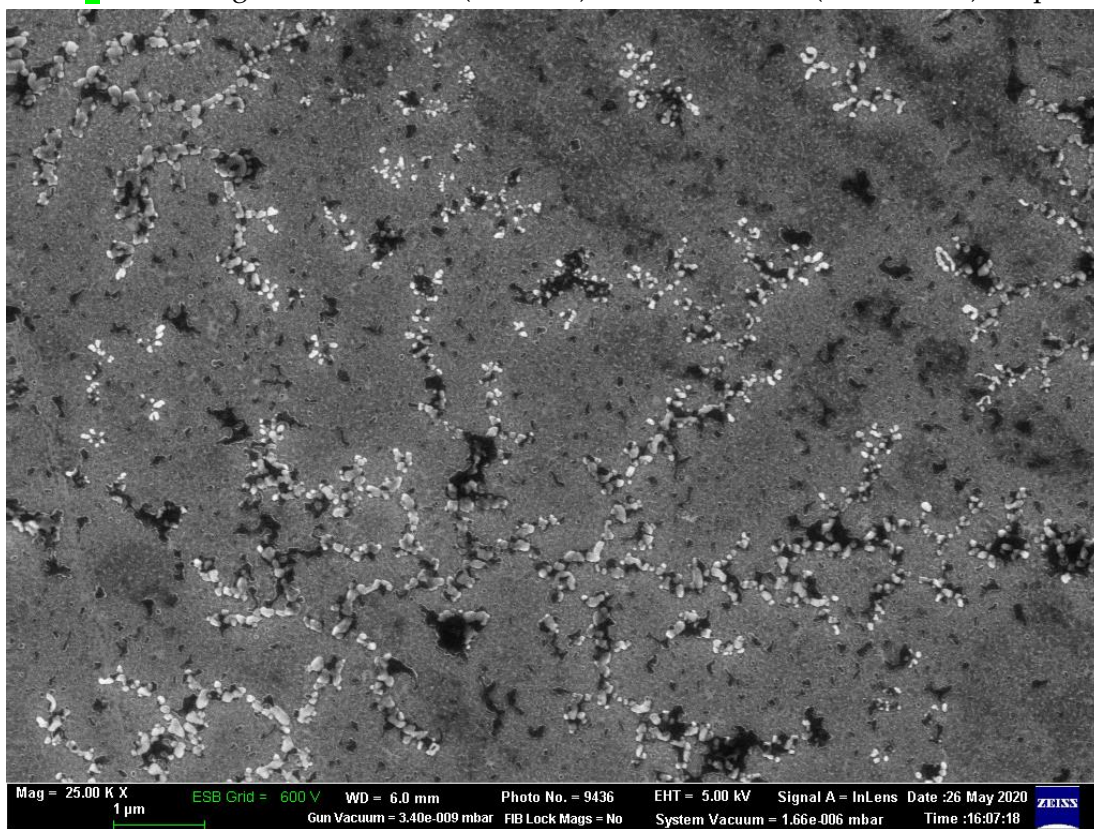


Figure S38. SEM image of 7/Flu/PVTE (1×10^{-5} M) after the solvent ($\text{H}_2\text{O}/\text{EtOH}$) evaporation.

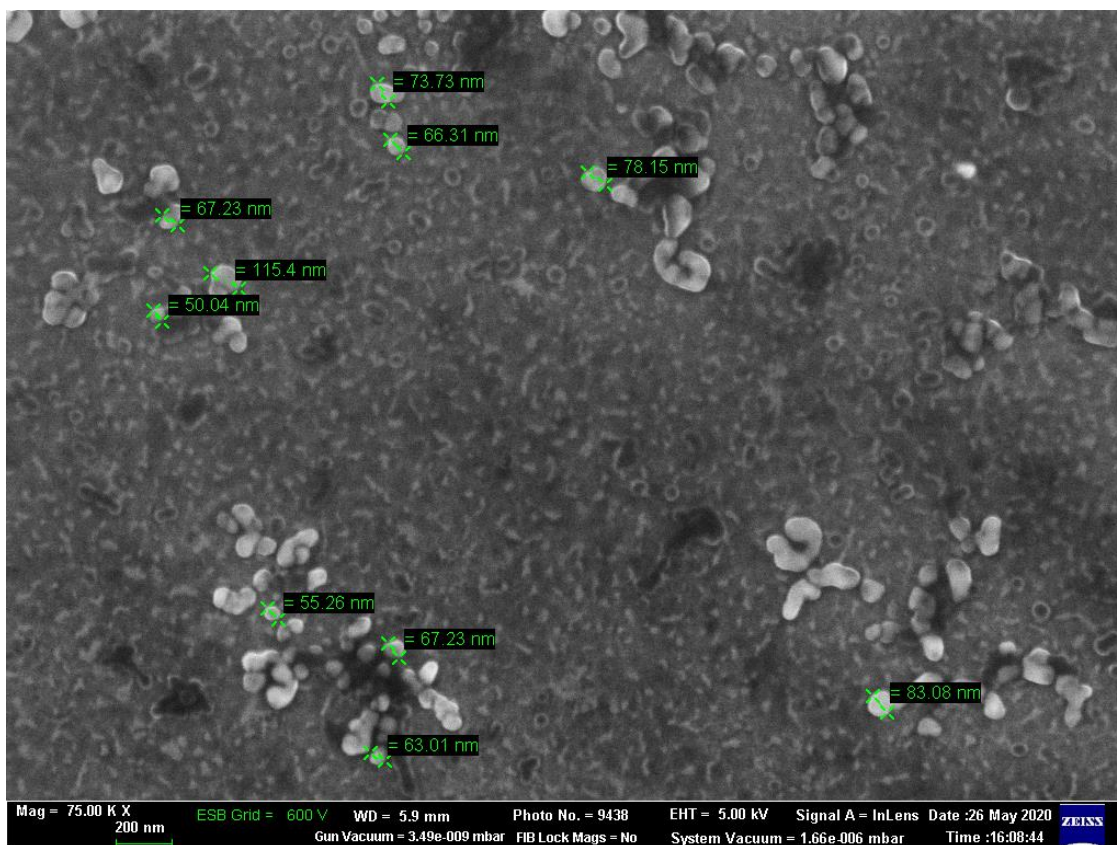


Figure S39. SEM image of 7/Flu/PVTE (1×10^{-5} M) after the solvent ($\text{H}_2\text{O}/\text{EtOH}$) evaporation.

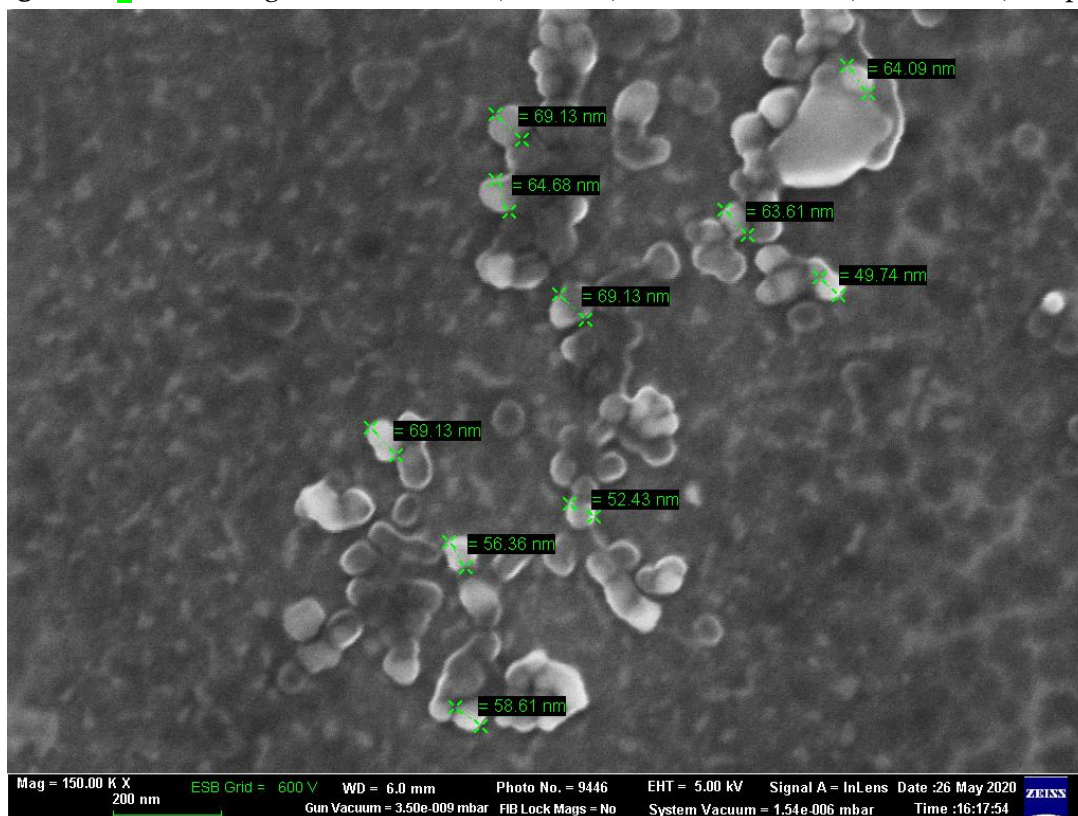


Figure S40. SEM image of 7/Flu (1×10^{-5} M) after the solvent (H₂O/EtOH) evaporation.

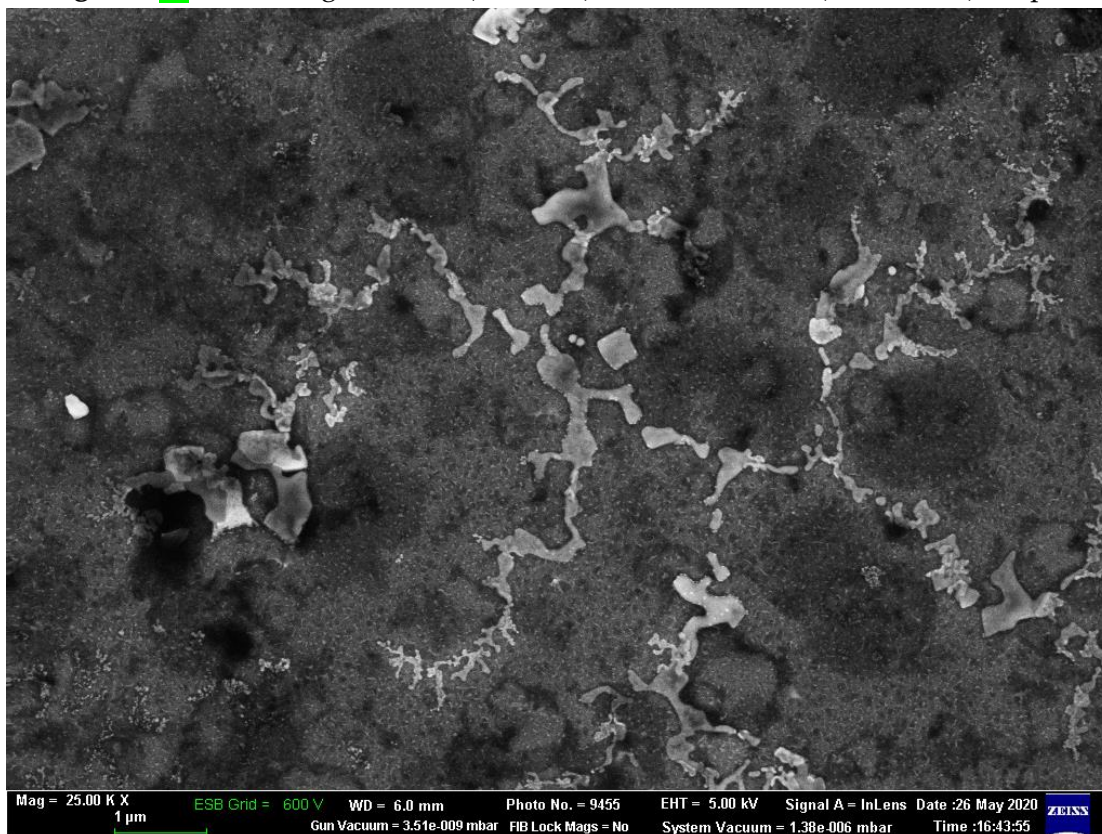
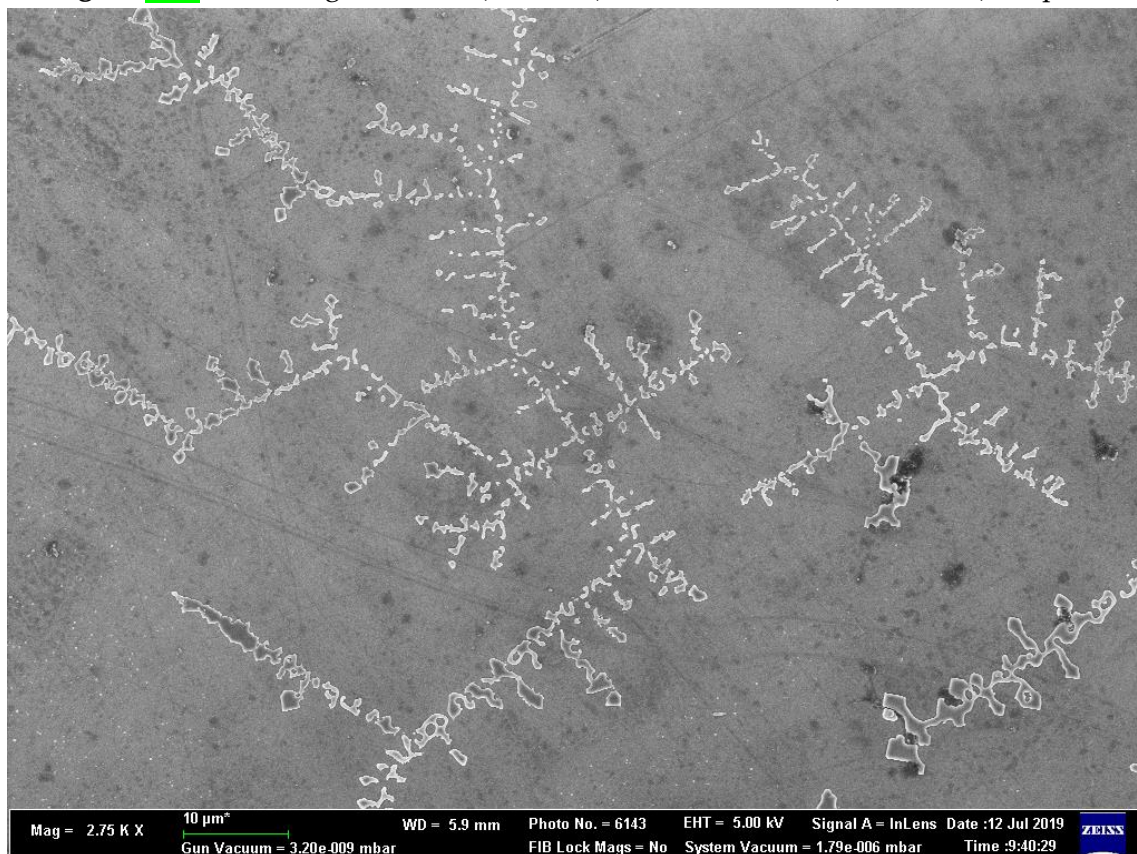


Figure S41. SEM image of PVTE (1×10^{-5} M) after the solvent (H₂O/EtOH) evaporation.



6. Fluorescence spectra

Figure S42. Fluorescence spectra of 7/Flu/PVTE (1×10^{-5} M) in buffer at different pH (9-2).

