

Supporting Information

Challenges for the self-assembly of poly (ethylene glycol)–poly (lactic acid) (PEG-PLA) into polymersomes: beyond the hydrophilic volume fraction (*f*)

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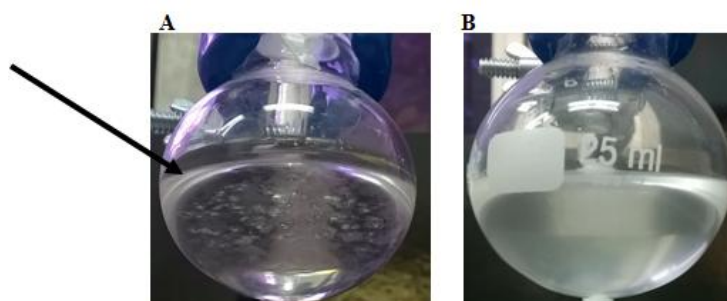


Figure S1. Aspects of poly(ethylene glycol)–poly(lactic acid) (PEG-PLA) system formed by film hydration by A) orbital stirring at 150 RPM and B) magnetic stirring at 400 RMP.

Table S1. Dynamic Light Scattering profile and respective polydispersity index (PDI) of nanostructures formed by poly(ethylene glycol)–poly(lactic acid) (PEG-PLA) by film hydration under magnetic stirring at 400 RPM after overnight.

Copolymer	Scattering Intensity (nm)	Scattering by Number (nm)	PDI
PEG ₄₅ PLA ₆₉	466 (100%)	416.1 (100%)	1
PEG ₁₁₄ PLA ₁₅₃	558.4 (85.1 %) and 155.4 (14.9%)	200.9 (100 %)	0.771
PEG ₁₁₄ PLA ₁₈₀	439.9 (96.9%) and 5331 (3.1%)	185.7 (100%)	0.609

Table S2. Dynamic Light Scattering profile and respective polydispersity index (PDI) of nanostructures formed by poly(ethylene glycol)-poly(lactic acid) (PEG-PLA) by film hydration under magnetic stirring at 400 RPM and different times of sonication.

Copolymer	Sonication by 20 min		Sonication by 50 min	
	Scattering Intensity (nm)	Scattering by Number (nm)	Scattering Intensity (nm)	Scattering by Number (nm)
PEG ₄₅ -PLA ₆₉ (2000:5000 MW)	600.6 (46 %) 380(45.2%) and 186 (8.8%)	165(50.6%) and 400 (49.4%)	376 (58 %), 1050 (30%) and 4379 (12%)	341 (94 %) and 1024 (6%)
PEG ₁₁₄ PLA ₁₅₃ (5000:11000 MW)	1081 (62.8%), 276.5 (28.7 %) and 4788 (8,5 %)	952.1 (7.8%), 226.5 (92.1%) and 4601 (0.1%)	659.4 (57%), 245.4 (35.3%) and 5059 (7.8%)	232 (99%) and 5005 (0.1%)
PEG ₁₁₄ PLA ₁₈₀ (5000:13000 MW)	446(99 %) and 5560 (1%)	218 (100%)	454 (95.4%), 5298 (2.6%) and 86 (2%)	204 (70%) and 75 (30%)

Table S3. Dynamic Light Scattering profile and respective polydispersity index (PDI) of nanostructures formed by PEG₄₅-PLA₆₉ (2000:5000 MW) by film hydration under magnetic stirring at 400 RPM at different times and temperatures.

PEG ₄₅ -PLA ₆₉ (2000:5000 MW)	Scattering Intensity	Scattering by Number
Agitation for 24 hours at room temperature	572 (100%)	491.6 (100%)
Agitation for 24 hours at 40°C	390.2 (96%), 122.6 (3%) and 5560 (1%)	106.6 (31%) and 326 (69%)
Agitation for 48 hours at room temperature	467 (97%) and 155 (3%)	415 (64%) and 164 (36%)
Agitation for 48 hours at 40°C	460 (95%), 4919(4%) and 108 (1%)	392 (70%) and 90 (30%)
Agitation for 72 hours at room temperature	752 (56%) and 335 (44%)	324 (100%)
Agitation for 72 hours at 40°C	758 (62%), 270 (37%) and 5296 (1%)	286 (100%)

Table S4. Dynamic Light Scattering profile and respective polydispersity index (PDI) of nanostructures formed by PEG₁₁₄PLA₁₅₃ (5000:11000 MW) by film hydration under magnetic stirring at 400 RPM at different times and temperatures.

PEG ₁₁₄ PLA ₁₅₃ (5000:11000 MW)	Intensity	Number
Agitation for 24 hours at room temperature	362 (100%)	110 (51%) and 283 (49%)
Agitation for 24 hours at 40°C	349 (93%) and 109 (7%)	280 (37%) and 93 (63%)
Agitation for 48 hours at room temperature	357.5 (100 %)	140 (19%) and 294 (81%)
Agitation for 48 hours at 40°C	567 (73%) and 176 (27%)	520 (6%) and 138 (94%)
Agitation for 48 hours and sonication for 50 min at 40°C	309 (92%) and 4224 (8%)	66.44 (30%) and 142 (70%)
Agitation for 72 hours at room temperature	401 (94%) and 5321 (6%)	136 (100%)
Agitation for 72 hours at 40°C	574 (52%), 234 (47%) and 5560(1%)	192 (100%)

Table S5. Dynamic Light Scattering profile and respective polydispersity index (PDI) of nanostructures formed by PEG₁₁₄PLA₁₈₀ (5000:13000 MW) by film hydration under magnetic stirring at 400 RPM at different times and temperatures.

PEG ₁₁₄ PLA ₁₈₀ (5000:13000 MW)	Intensity	Number
Agitation for 24 hours at room temperature	745 (59%), 276 (40%) and 5526 (1%)	246 (86%) and 670 (14%)
Agitation for 24 hours at 40°C	200.6 (100 %) and 5448 (1%)	128 (80%) and 53 (20%)
Agitation for 48 hours at room temperature	596 (98) and 167(2%)	376(84 %) and 160 (16%)
Agitation for 48 hours at 40°C	229 (83%) and 623.4 (17%)	115 (25%) and 218 (75 %)
Agitation for 72 hours at room temperature	450 (60%), 216 (36%) and 5156 (4%)	177 (100%)
Agitation for 72 hours at 40°C	325 (78.5%) and 129.5 (21%)	119 (100%)

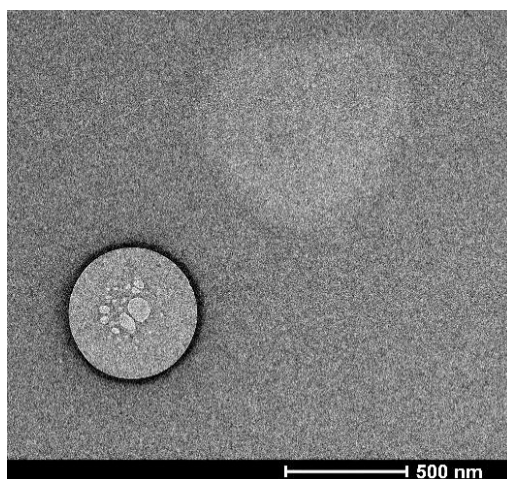


Figure S2. Polymersomes from poly(ethylene glycol)-poly(lactic acid) PEG₁₁₄PLA₁₅₃ after extrusion through 0.4 μm pores.

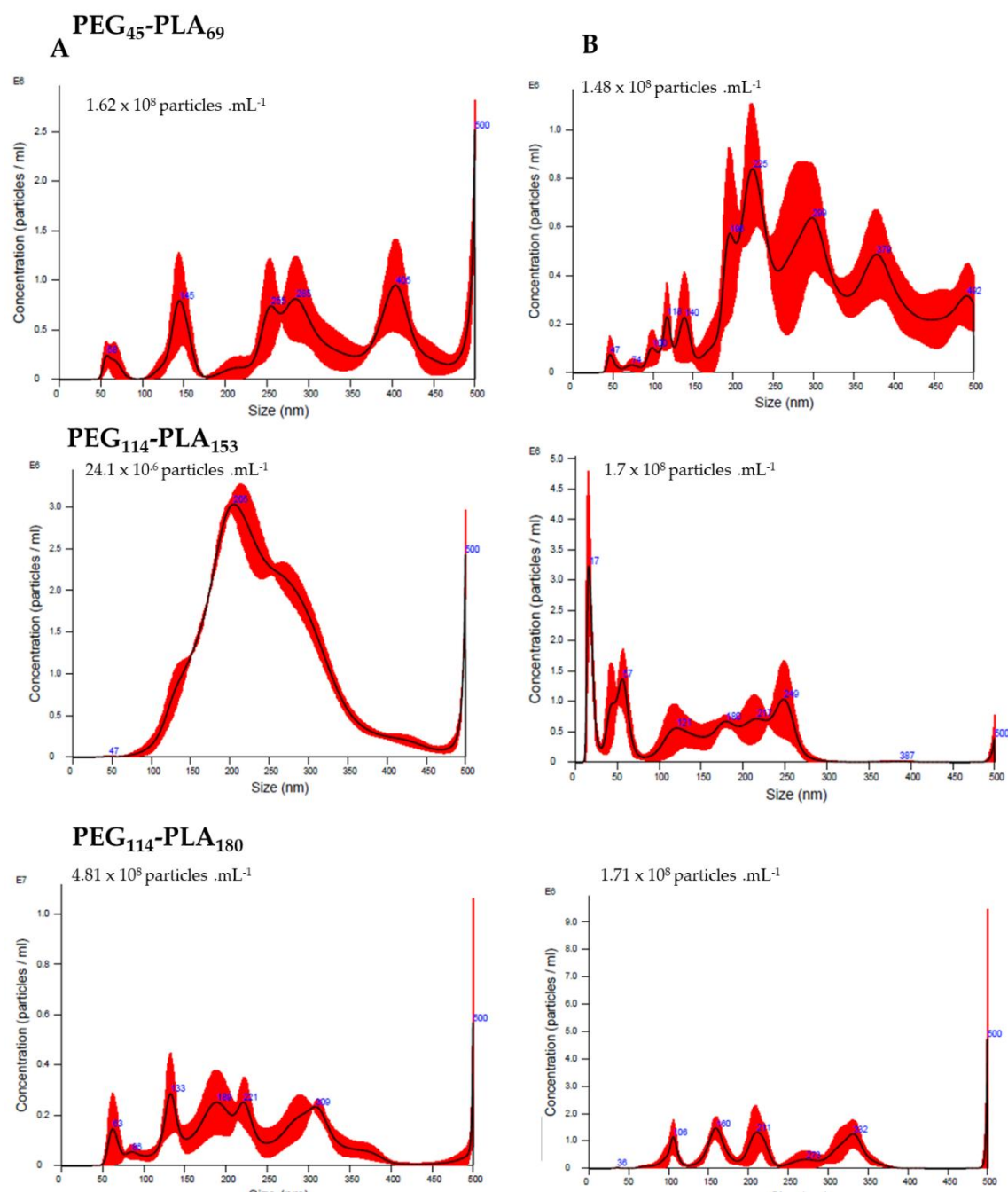


Figure S3. Nanoparticle Tracking Analysis of poly(ethylene glycol)-poly(lactic acid) (PEG-PLA) nanostructures after centrifugation: A) 0.1 % (m/v) B) At 0.03 % (m/v).

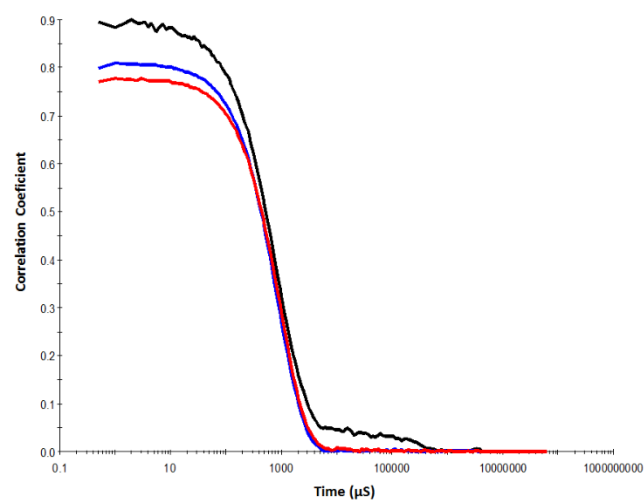


Figure S4. Correlation coefficients: Decaying time for nanostructures formed from PEG₄₅-PLA₆₉ (black), PEG₁₁₄PLA₁₅₃ (red) and PEG₁₁₄PLA₁₅₃ (blue).