

## **Supplementary Information**

### **A Novel Injectable pH-Temperature Sensitive Hydrogel Contained Chitosan-Insulin Electrosprayed Nanospheres Composites As Insulin Delivery System In Type I Diabetes Treatment**

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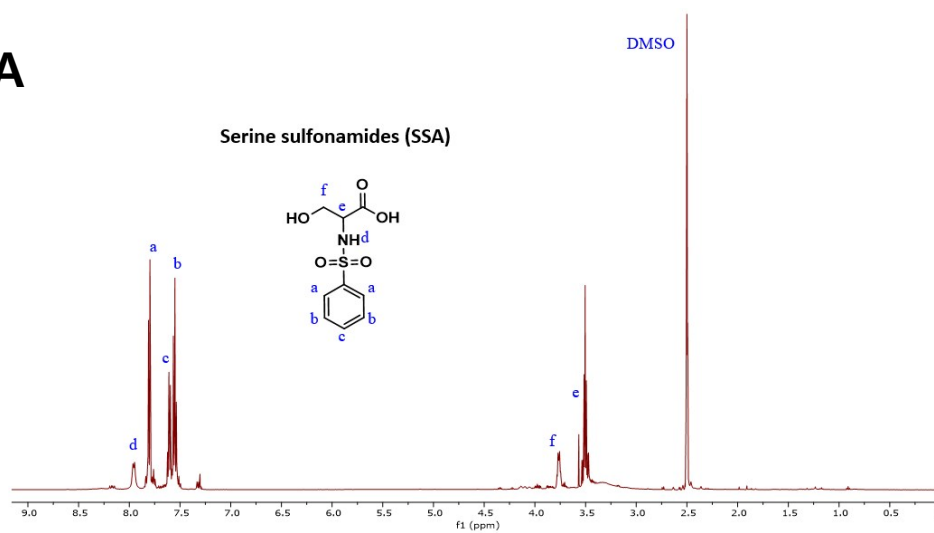
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*\* CORRESPONDING AUTHORS:*

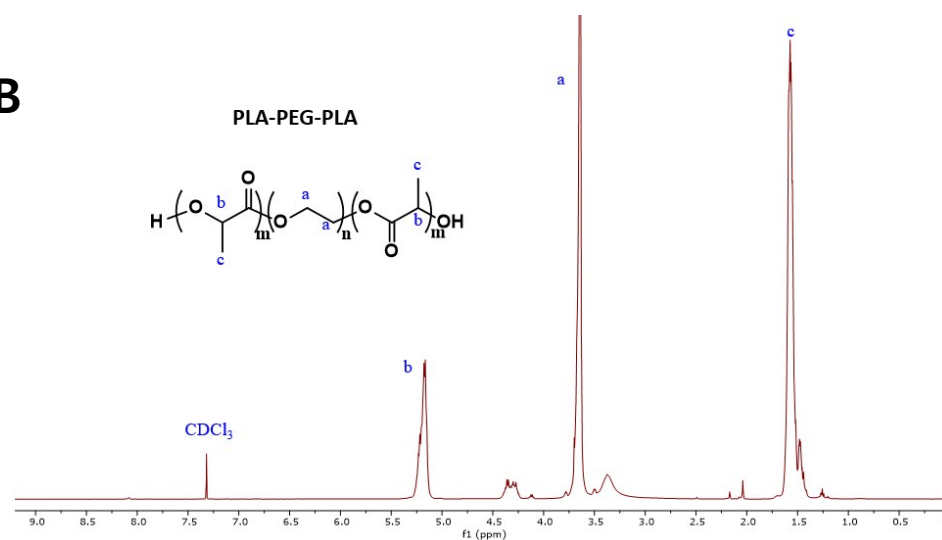
Dai Phu Huynh (hdphu@hcmut.edu.vn )

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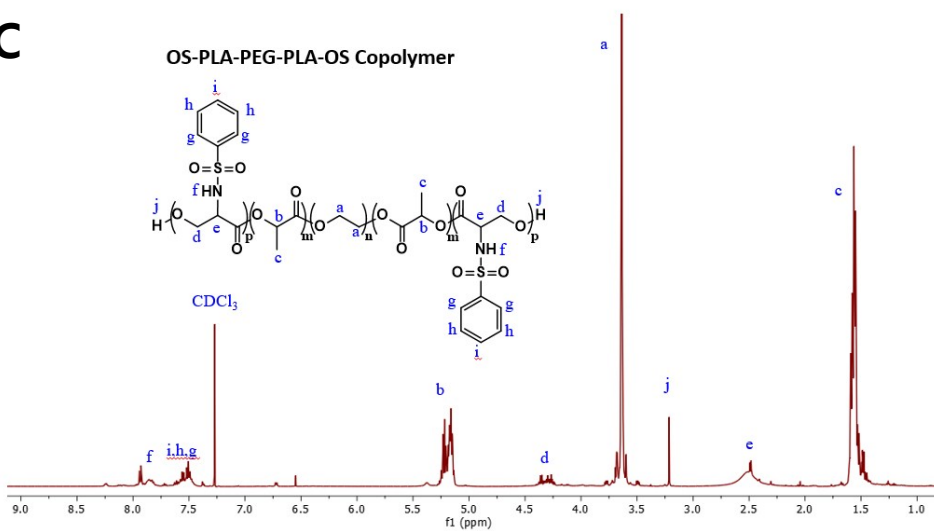
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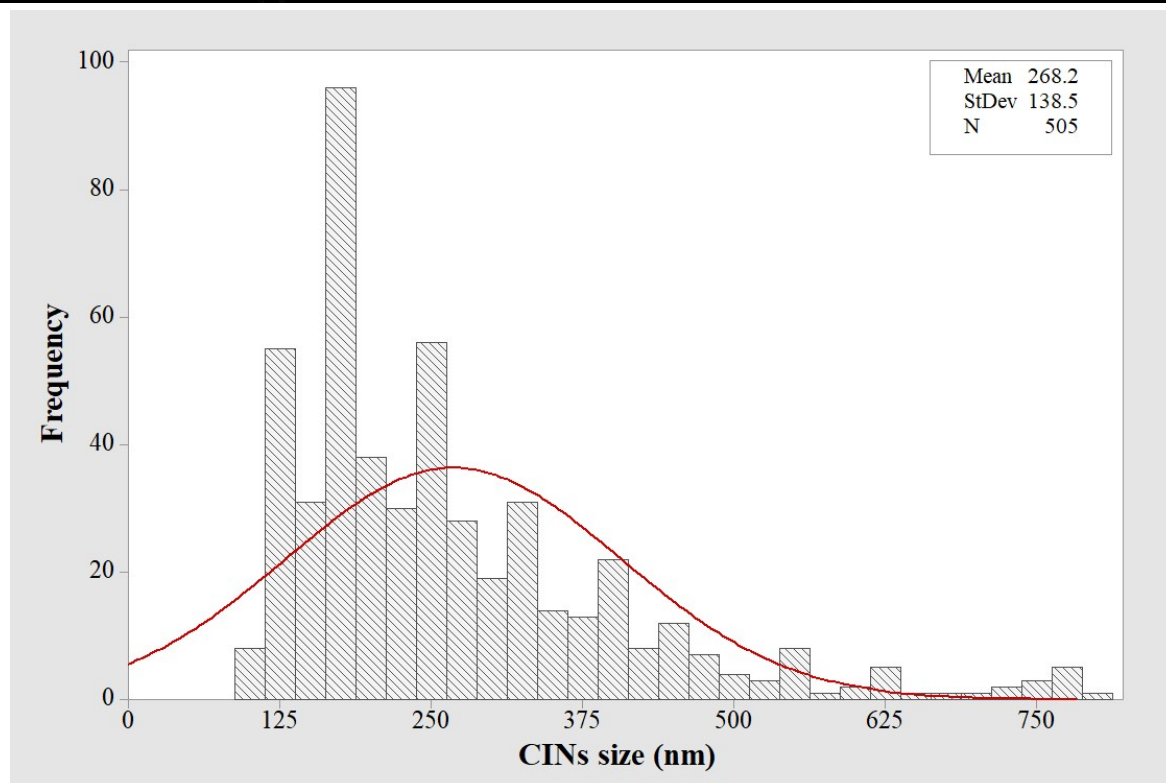
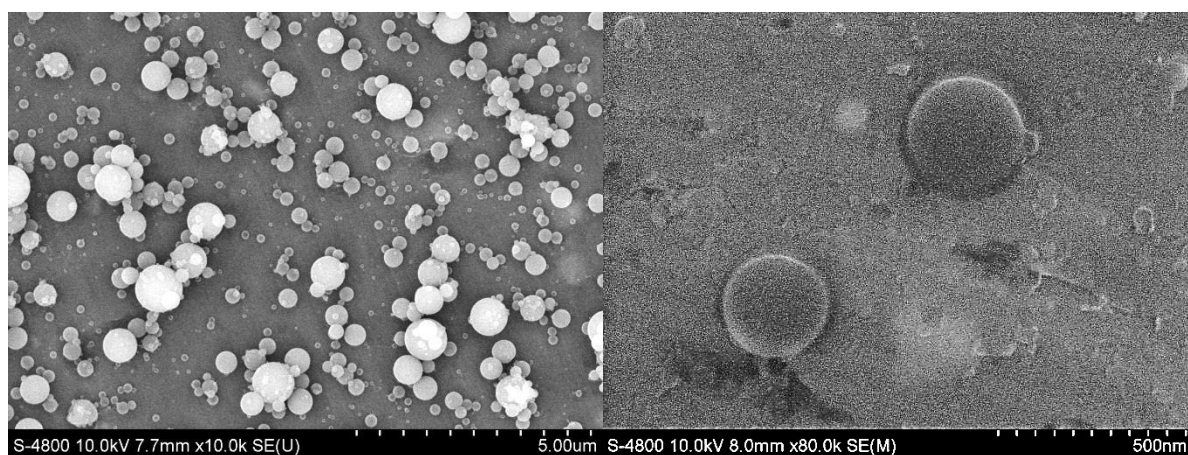
**B**



**C**

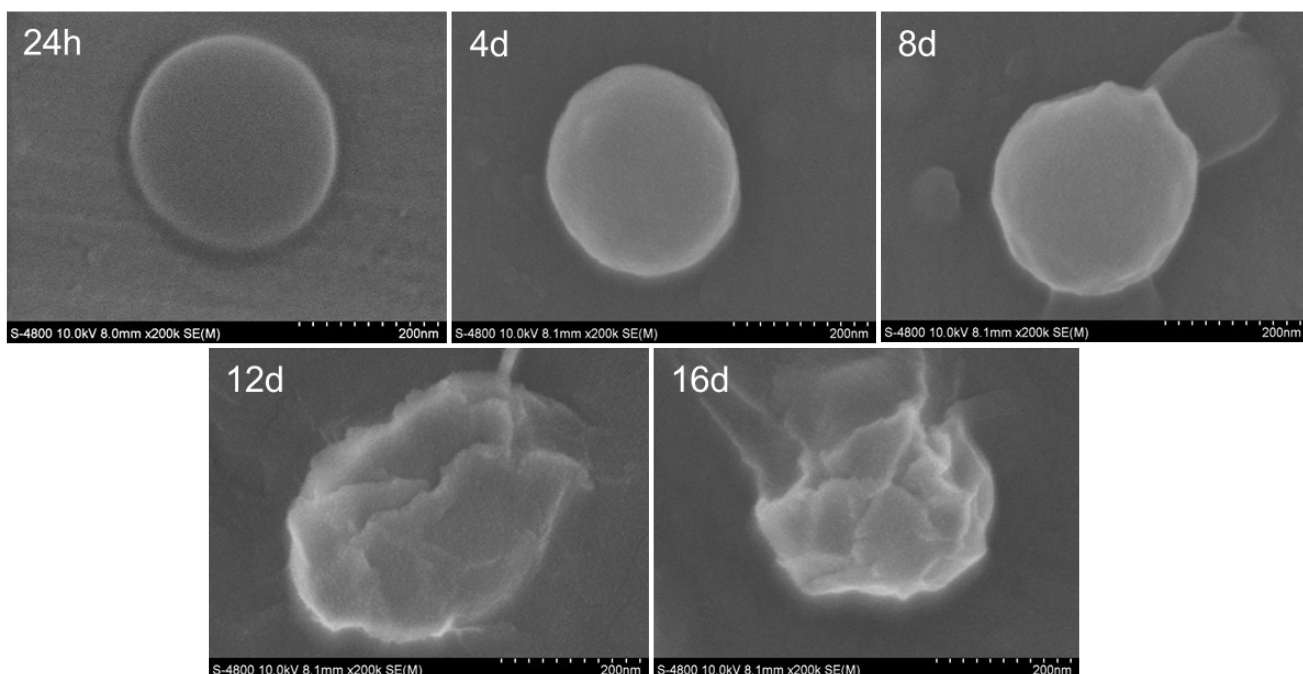


**Figure S1.** <sup>1</sup>H NMR spectra of (A) SSA, (B) PLA-PEG-PLA, and (C) PeCo1 copolymers.

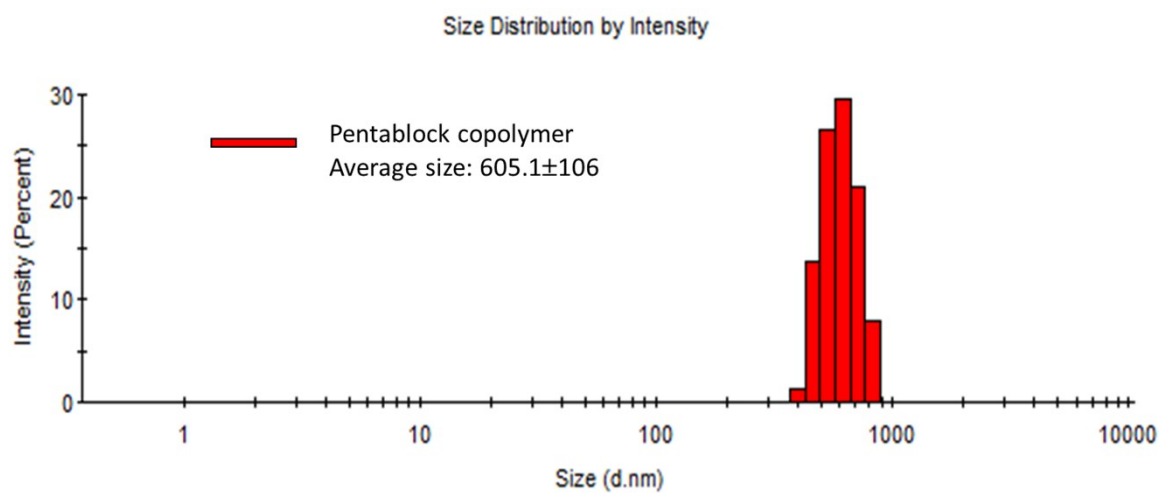


**Figure S2.** SEM images of electrospayed CINs and the CINs size distribution

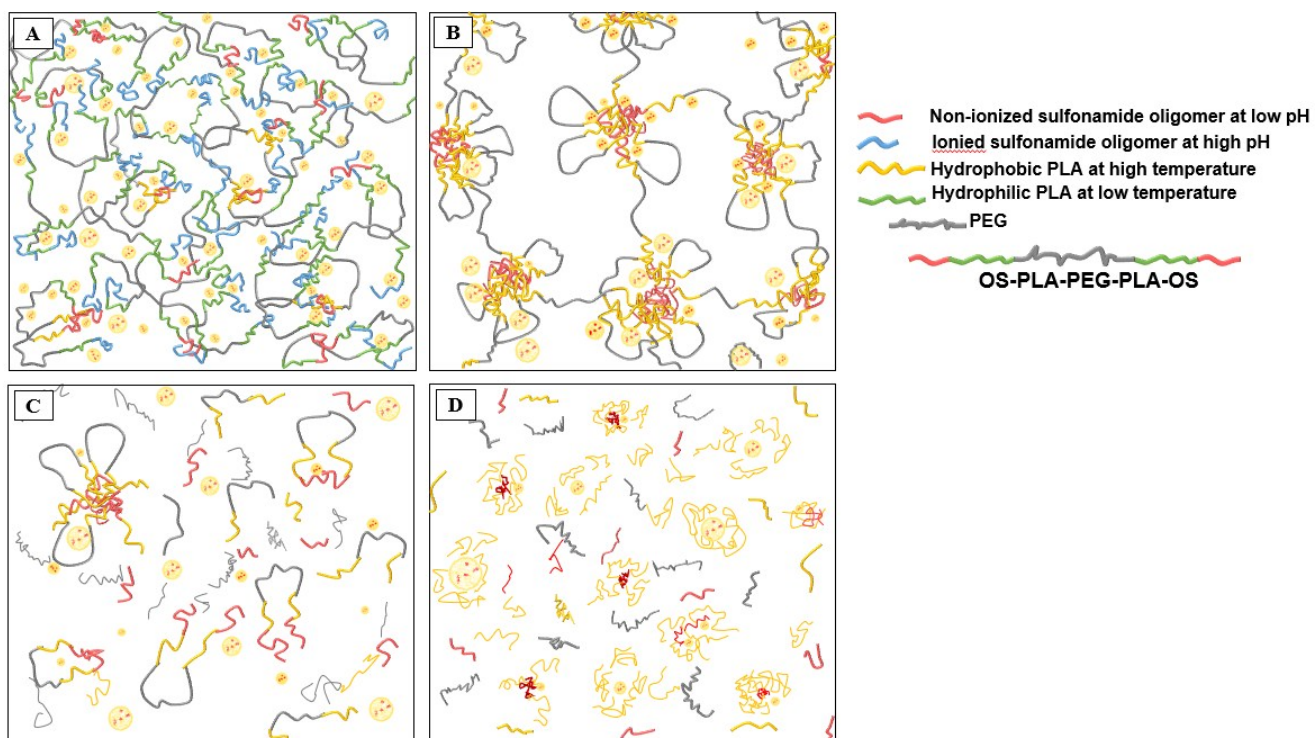
$C_{\text{Chitosan}}=3,25\%$ ;  $U=12\text{kV}$ ;  $Q=0,2\text{ml/h}$ ;  $L=12,5\text{cm}$ ;  $\text{Insulin}=20\% \text{ wt.}$



**Figure S3.** SEM images of electrospayed CINs incubated in PBS pH 7.4 (0.5 wt% tween 20) at 37 °C for 24h, 4d, 8d, 12d, 16d.



**Figure S4.** Size distribution of pentablock copolymer micelles in PBS pH 7.4



**Figure S5.** Schematic concept of hydrogels-CINs composite injection and the release of insulin in molecular scale (A) Copolymer solution containing CINs. (B) Hydrogel-CINs composite matrix. (C) Degradation of matrix and the release of CINs. (D) The release of insulin



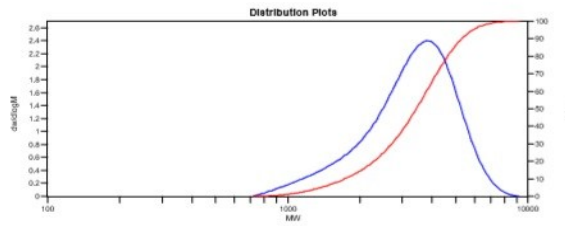
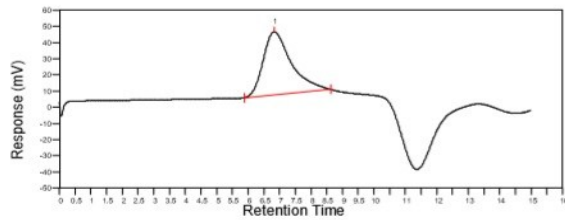
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### Cirrus GPC Sample Injection Report

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Workbook: C:\Cirrus Workbooks\MESOPORE-035-6\MESOPORE-035-6.plw

#### Sample Details

Sample Name: PEG2050-PLA2 Batch Name: LINH25102018  
Chloroform 1.000000 Concentration: 0.10 mg/ml  
Injection Volume: 20.0 ul K of Sample: 14.1000 Alpha of Sample: 0.7000  
Analysis Using Method: PEG



#### MW Averages

Peak No	Mp	Mn	Mw	Mz	Mz+1	Mv	PD
1	3807	2877	3477	4006	4464	3395	1.20855

Peak No	Name	Start RT (mins)	Max RT (mins)	End RT (mins)	Pk Height (mV)	% Height	Area (mV.secs)	% Area
1		5.88	6.83	8.83	39.0477	100	2429.59	100

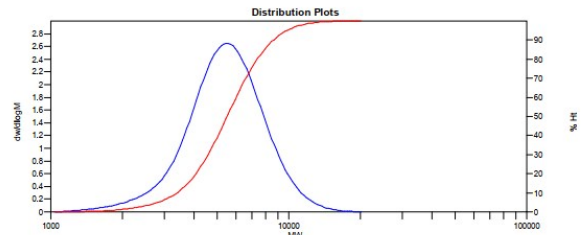
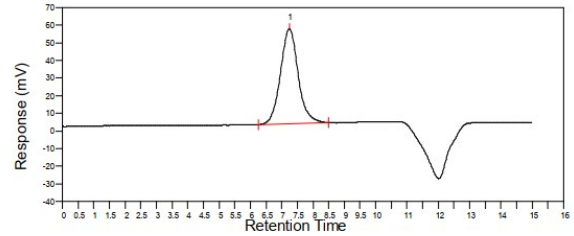
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### Cirrus GPC Sample Injection Report

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#### Sample Details

Sample Name: PEG2050-2-4PLA Batch Name: NVVLINH\_-FEB2019  
Chloroform 1.000000 Concentration: 0.10 mg/ml  
Injection Volume: 40.0 ul K of Sample: 14.1000 Alpha of Sample: 0.7000  
Analysis Using Method: PEG-13022019



#### MW Averages

Peak No	Mp	Mn	Mw	Mz	Mz+1	Mv	PD
1	5446	4971	5747	6547	7409	5632	1.15611

Peak No	Name	Start RT (mins)	Max RT (mins)	End RT (mins)	Pk Height (mV)	% Height	Area (mV.secs)	% Area
1		6.27	7.25	8.50	53.978	100	2120.47	100

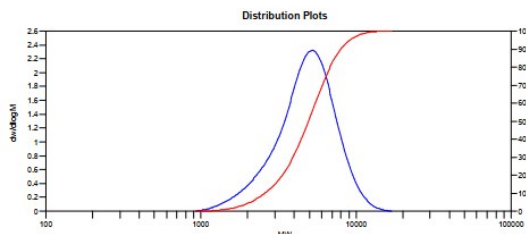
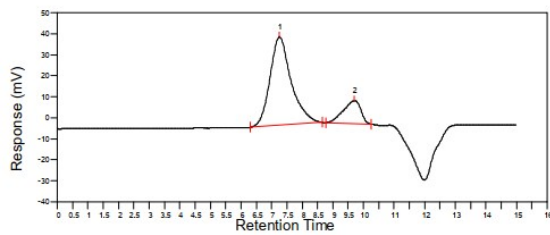
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### Cirrus GPC Sample Injection Report

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#### Sample Details

Sample Name: PEG2050-2 PLA-OS12 Batch Name: NVVLINH\_-FEB2019  
Chloroform 1.000000 Concentration: 0.10 mg/ml  
Injection Volume: 40.0 ul K of Sample: 14.1000 Alpha of Sample: 0.7000  
Analysis Using Method: PS



#### MW Averages

Peak No	Mp	Mn	Mw	Mz	Mz+1	Mv	PD
1	5232	4218	5122	6007	6879	4962	1.21432
2	247	274	303	339	379	298	1.10584

Peak No	Name	Start RT (mins)	Max RT (mins)	End RT (mins)	Pk Height (mV)	% Height	Area (mV.secs)	% Area
1		6.30	7.25	8.85	42.0778	79.3526	2024.61	82.3676
2		8.77	9.70	10.25	10.9486	20.6474	433.407	17.6324

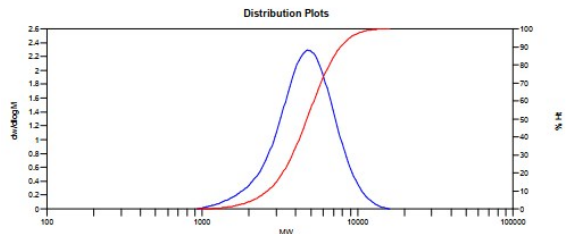
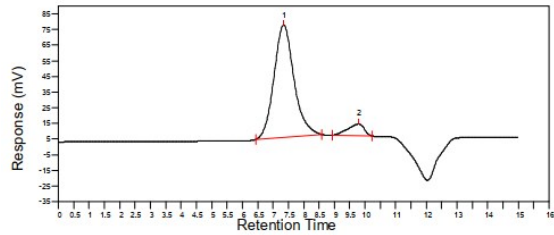
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Workbook: C:\Cirrus Workbooks\MESOPORE-34-72-CHLOROFORM\MESOPORE-34-72-CHLOROFORM.plw

#### Sample Details

Sample Name: PEG 2050-2PLA-OS6 Batch Name: NVVLINH\_-FEB2019  
Chloroform 1.000000 Concentration: 0.10 mg/ml  
Injection Volume: 40.0 ul K of Sample: 14.1000 Alpha of Sample: 0.7000  
Analysis Using Method: PEG-13022019



#### MW Averages

Peak No	Mp	Mn	Mw	Mz	Mz+1	Mv	PD
1	4769	4130	4988	5828	6714	4843	1.20291
2	189	210	233	261	292	230	1.10952

Peak No	Name	Start RT (mins)	Max RT (mins)	End RT (mins)	Pk Height (mV)	% Height	Area (mV.secs)	% Area
1		6.43	7.35	8.58	71.6953	90.699	3261.79	92.2121
2		8.93	9.78	10.22	7.35218	9.30096	275.481	7.78795

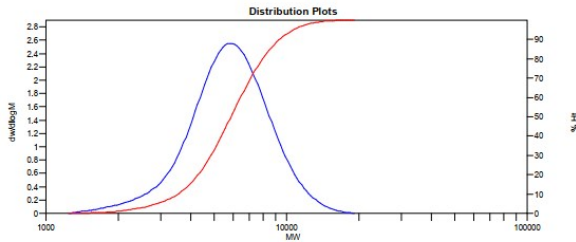
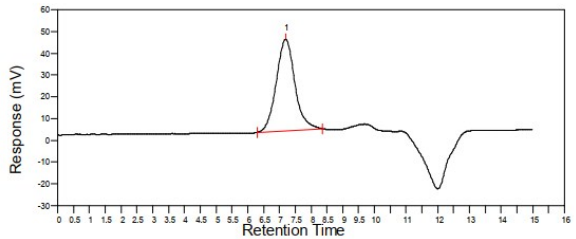
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Cirrus GPC Sample Injection Report

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Sample Details

Sample Name: PEG2050-2.4PLA-OS6      Batch Name: NVVLINH\_-FEB2019  
Chloroform      1.000000      Concentration: 0.10 mg/ml  
Injection Volume: 40.0 ul      K of Sample: 14.1000      Alpha of Sample: 0.7000  
Analysis Using Method: PEG-13022019



MW Averages

Peak No	Mp	Mn	Mw	Mz	Mz+1	Mv	PD
1	5820	5272	6148	7045	7987	6018	1.16616

Peak No	Name	Start RT (mins)	Max RT (mins)	End RT (mins)	Pk Height (mV)	% Height	Area (mV.secs)	% Area
1		6.30	7.20	8.37	42.0874	100	1717.78	100

**Figure S6.** GPC results of (A) Triblock copolymer PLA-PEG-PLA (PEG/LA=1/2) (B) Triblock copolymer PLA-PEG-PLA (PEG/LA=1/2.4) (C) Pentablock copolymer PeCo1 (D) Pentablock copolymer PeCo2 (E) Pentablock copolymer PeCo3