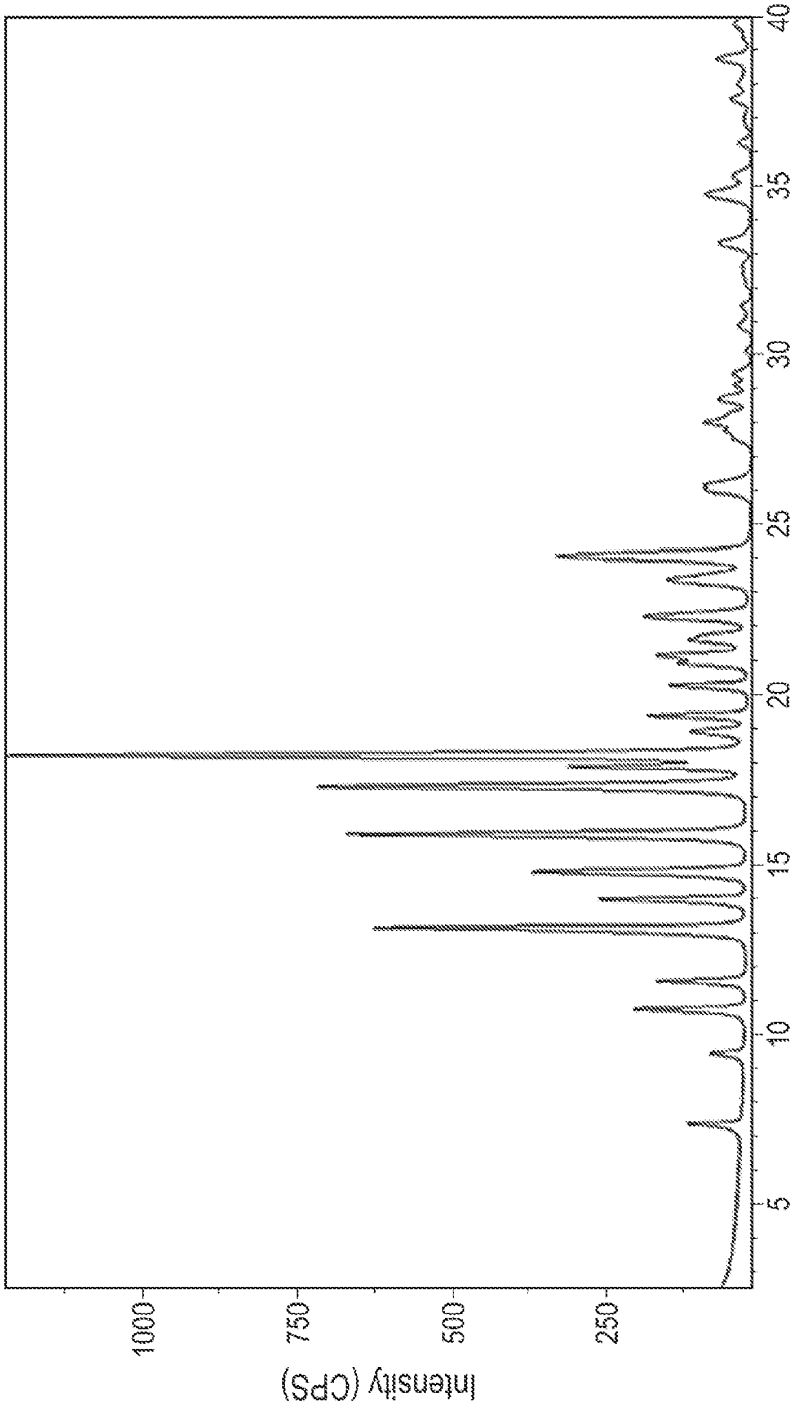


FIG. 1A

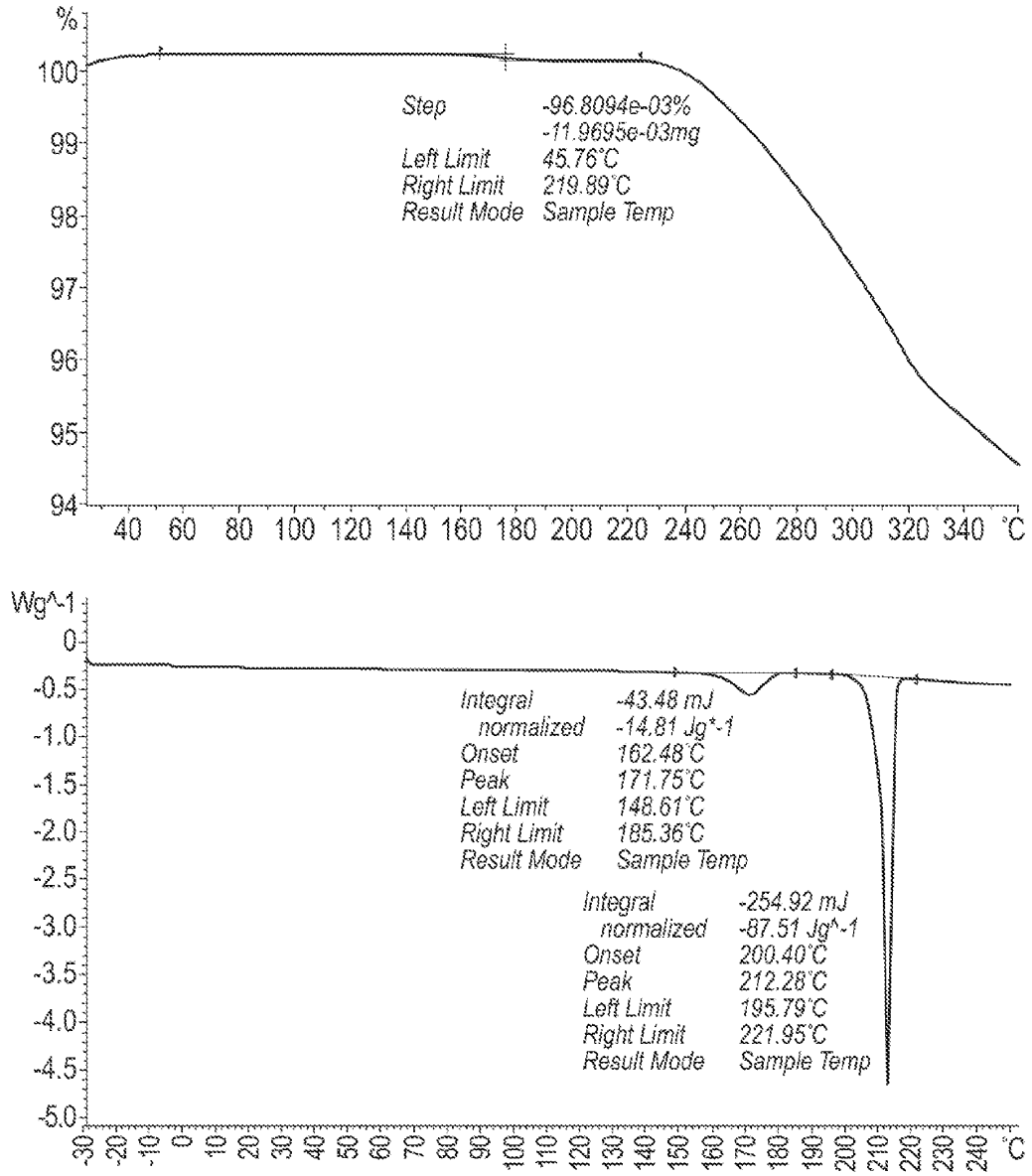

%
100
99
98
97
96
95
94
40 60 80 100 120 140 160 180 200 220 240 260 280 300 320 340 °C
Step -96.8094e-03%
-11.9695e-03mg
Left Limit 45.76°C
Right Limit 219.89°C
Result Mode Sample Temp
Wg^-1
0
-0.5
-1.0
-1.5
-2.0
-2.5
-3.0
-3.5
-4.0
-4.5
-5.0
-30 -20 -10 0 10 20 30 40 50 60 70 80 90 100 110 120 130 140 150 160 170 180 190 200 210 220 230 240 °C
Integral -43.48 mJ
normalized -14.81 Jg^-1
Onset 162.48°C
Peak 171.75°C
Left Limit 148.61°C
Right Limit 185.36°C
Result Mode Sample Temp
Integral -254.92 mJ
normalized -87.51 Jg^-1
Onset 200.40°C
Peak 212.28°C
Left Limit 195.79°C
Right Limit 221.95°C
Result Mode Sample Temp

FIG. 1B

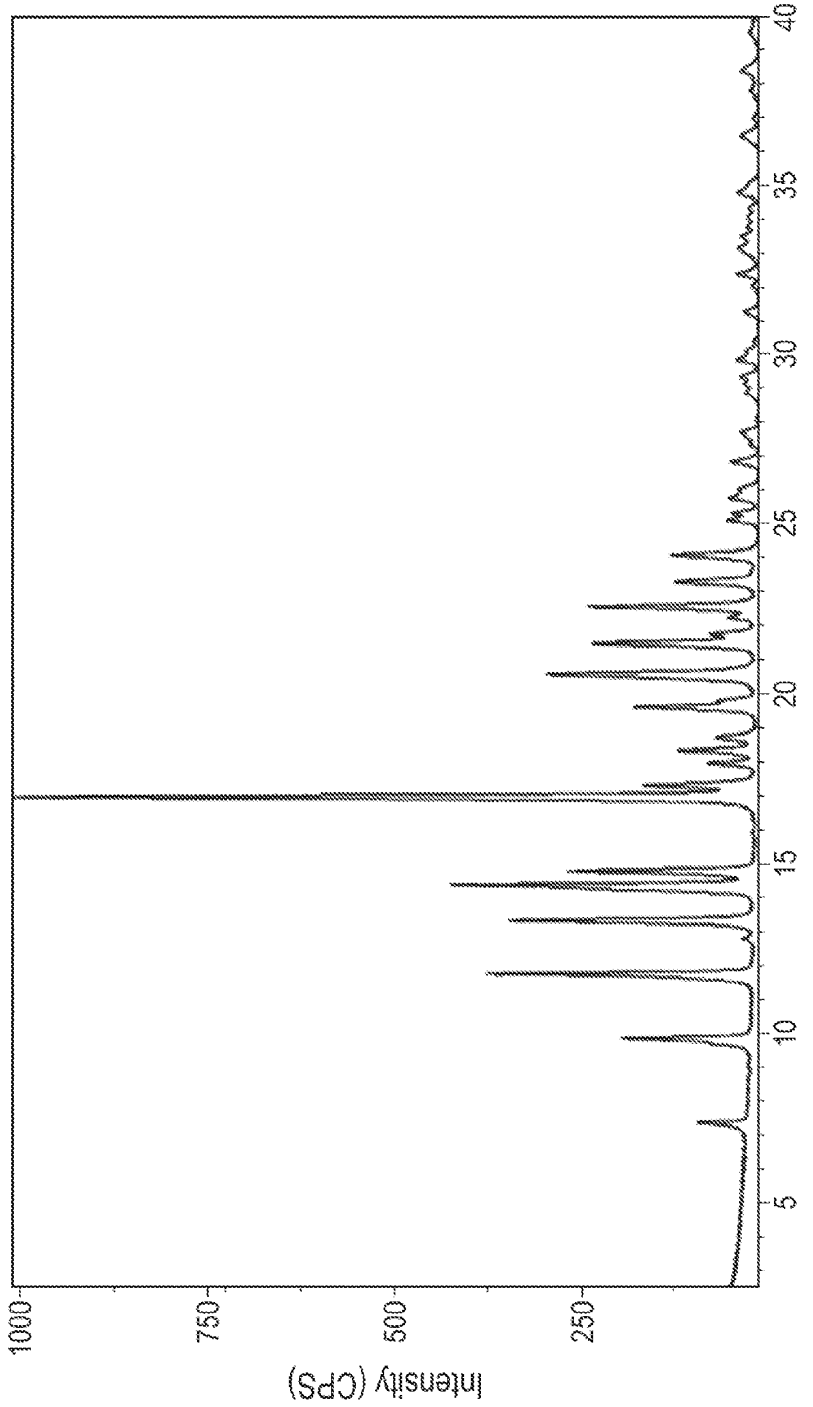


FIG. 2A

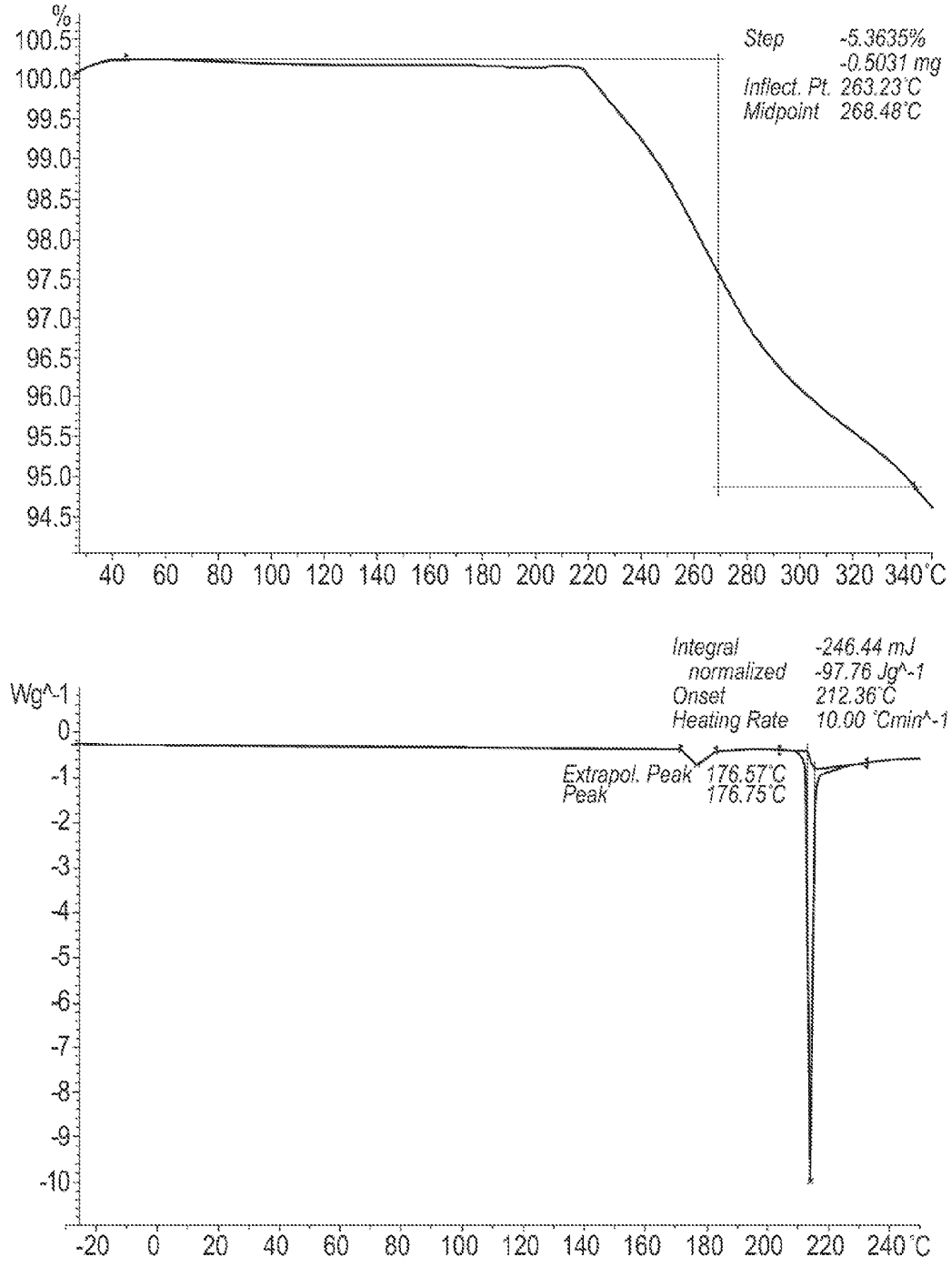

%
100.5
100.0
99.5
99.0
98.5
98.0
97.5
97.0
96.5
96.0
95.5
95.0
94.5
Step -5.3635%
-0.5031 mg
Inflect. Pt. 263.23°C
Midpoint 268.48°C
40 60 80 100 120 140 160 180 200 220 240 260 280 300 320 340°C
Integral -246.44 mJ
normalized -97.76 Jg^-1
Onset 212.36°C
Heating Rate 10.00 °Cmin^-1
Wg^-1
0
-1
-2
-3
-4
-5
-6
-7
-8
-9
-10
Extrapol. Peak 176.57°C
Peak 176.75°C
-20 0 20 40 60 80 100 120 140 160 180 200 220 240°C

FIG. 2B

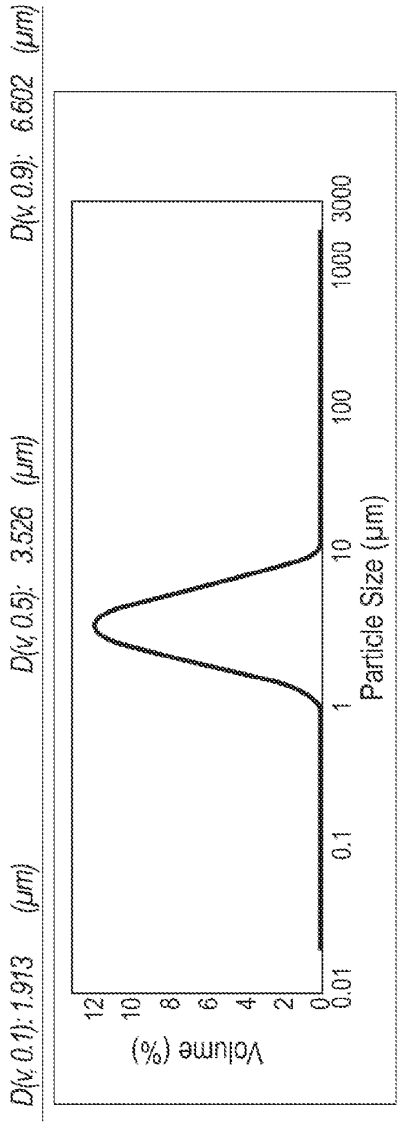

D(v, 0.1): 1.913 (µm)
D(v, 0.5): 3.526 (µm)
D(v, 0.9): 6.602 (µm)
Volume (%)
0
2
4
6
8
10
12
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 4A

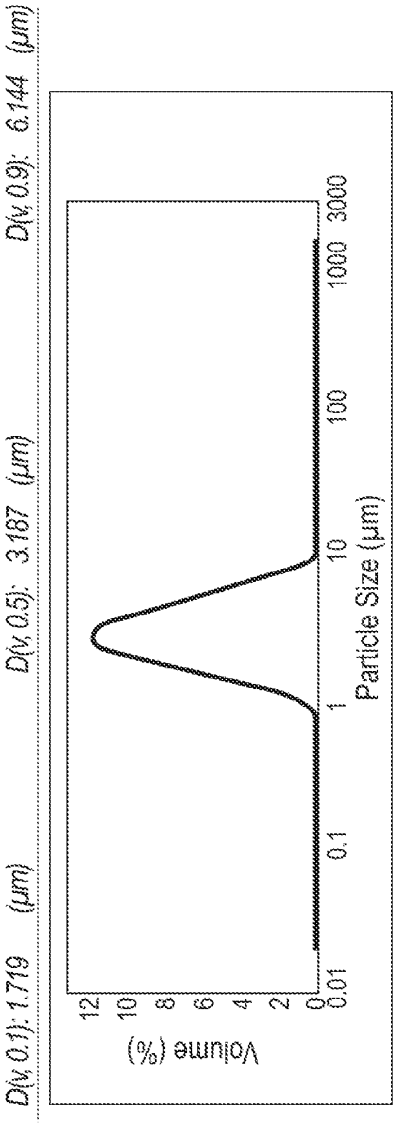

D(v, 0.1): 1.719 (µm)
D(v, 0.5): 3.187 (µm)
D(v, 0.9): 6.144 (µm)
Volume (%)
0
2
4
6
8
10
12
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 4B

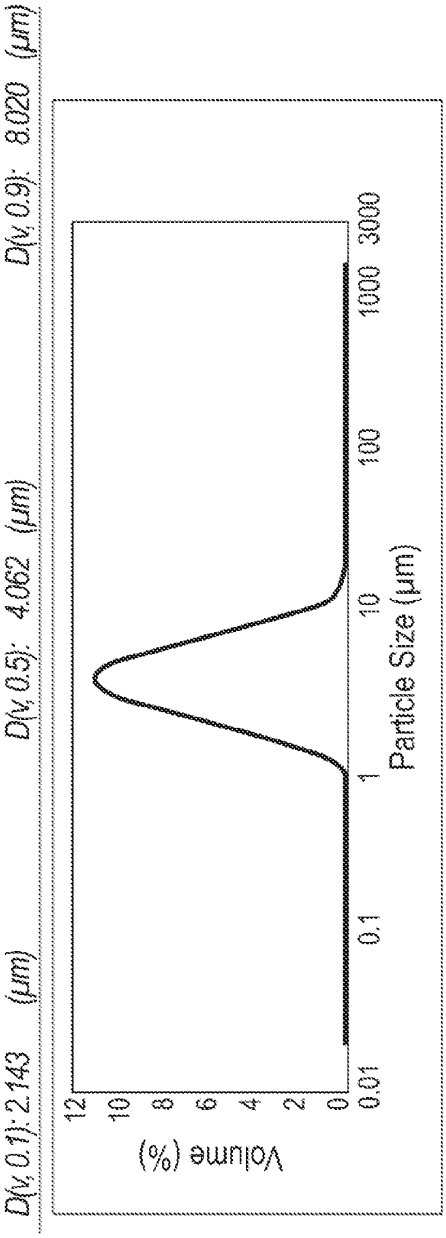

D(v,0.1): 2.143 (µm)
D(v,0.5): 4.062 (µm)
D(v,0.9): 8.020 (µm)
Volume (%)
0
2
4
6
8
10
12
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 4C

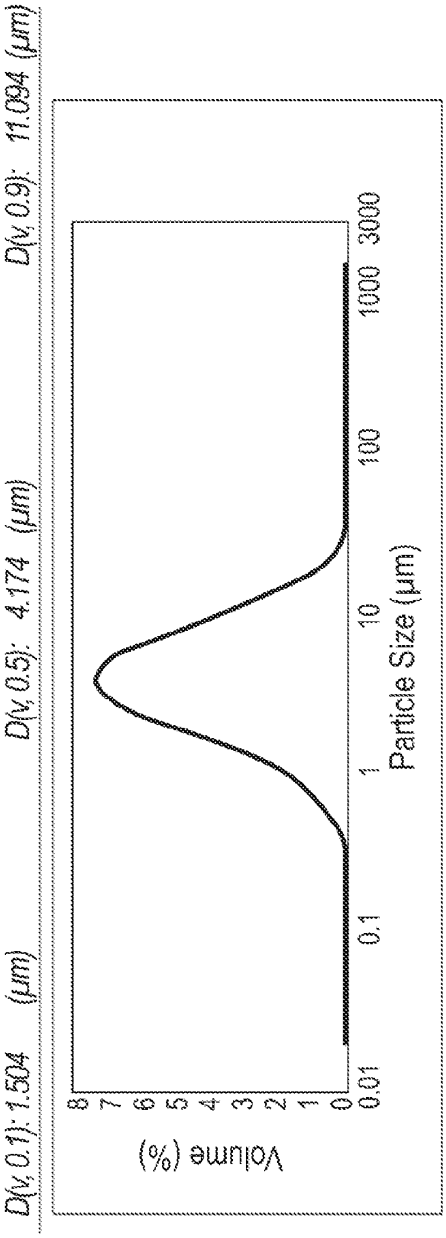

D(v,0.1): 1.504 (µm)
D(v,0.5): 4.174 (µm)
D(v,0.9): 11.094 (µm)
Volume (%)
0
1
2
3
4
5
6
7
8
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 5

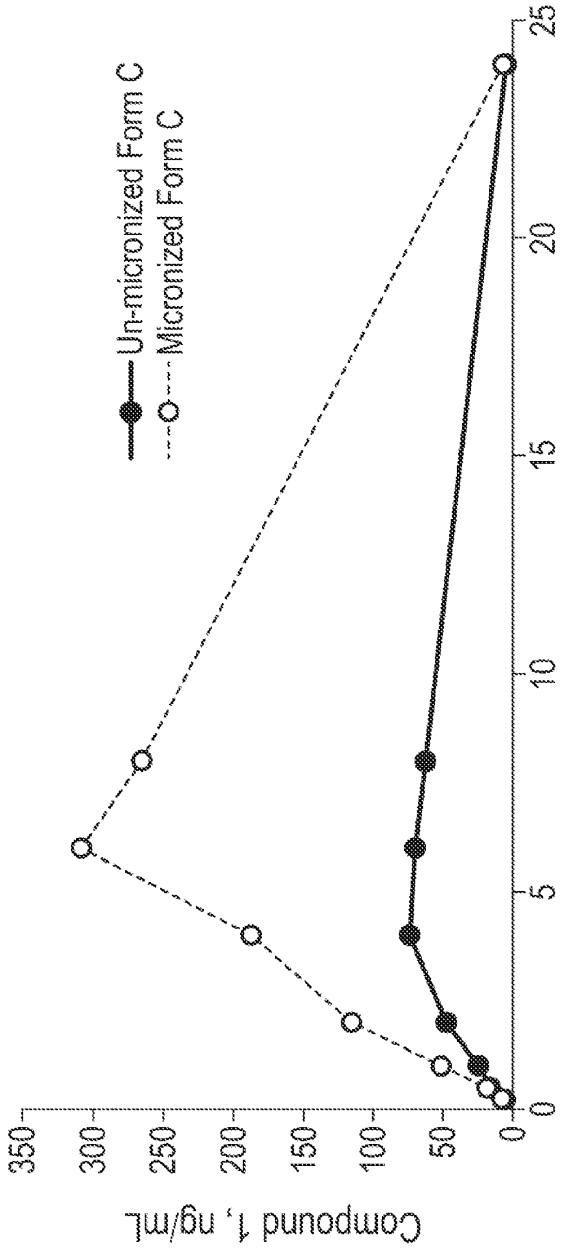

Un-micronized Form C
Micronized Form C
Compound 1, ng/mL
0
50
100
150
200
250
300
350
0
5
10
15
20
25
Time (h)

FIG. 6

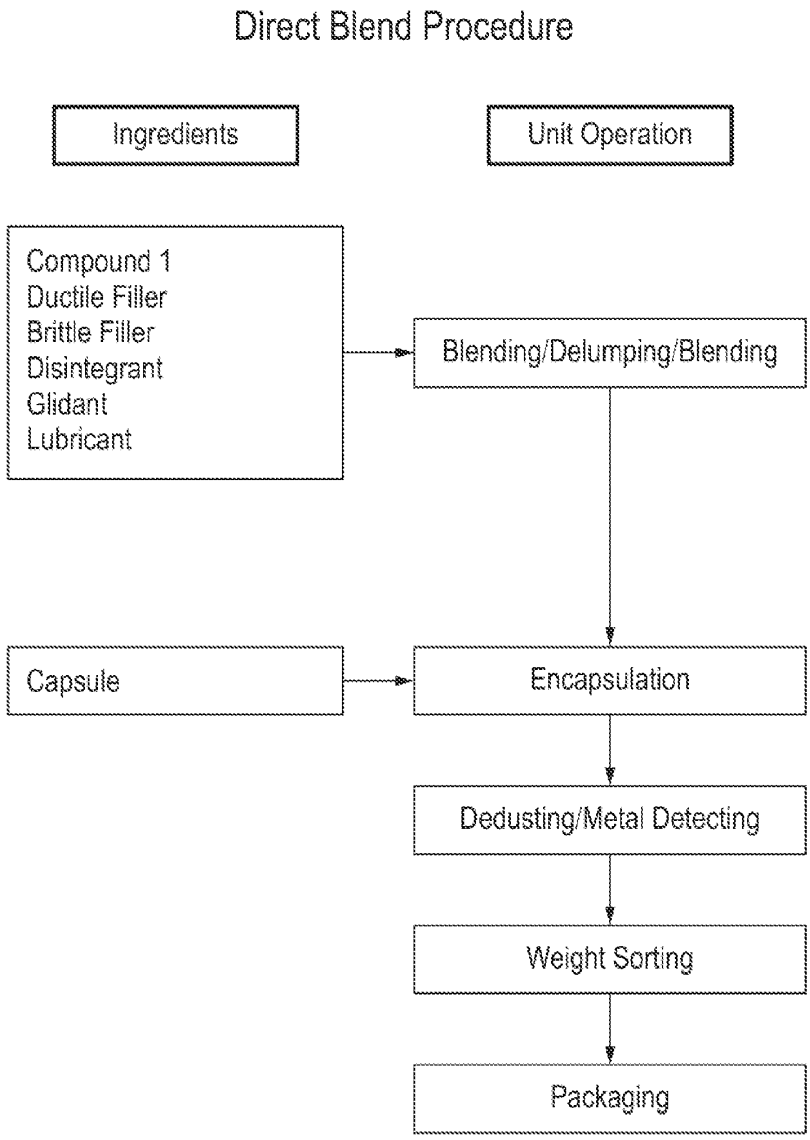

Direct Blend Procedure
Ingredients
Unit Operation
Compound 1
Ductile Filler
Brittle Filler
Disintegrant
Glidant
Lubricant
Blending/Delumping/Blending
Capsule
Encapsulation
Dedusting/Metal Detecting
Weight Sorting
Packaging

FIG. 7

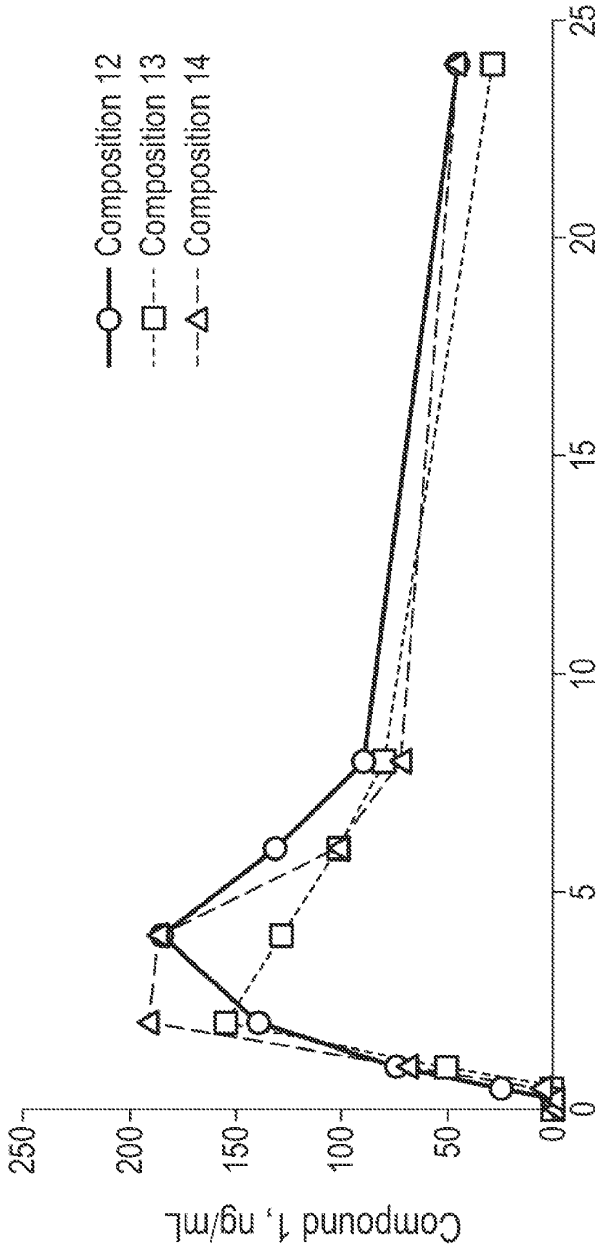

Composition 12
Composition 13
Composition 14
Compound 1, ng/mL
250
200
150
100
50
0
0
5
10
15
20
25
Time (h)

FIG. 8

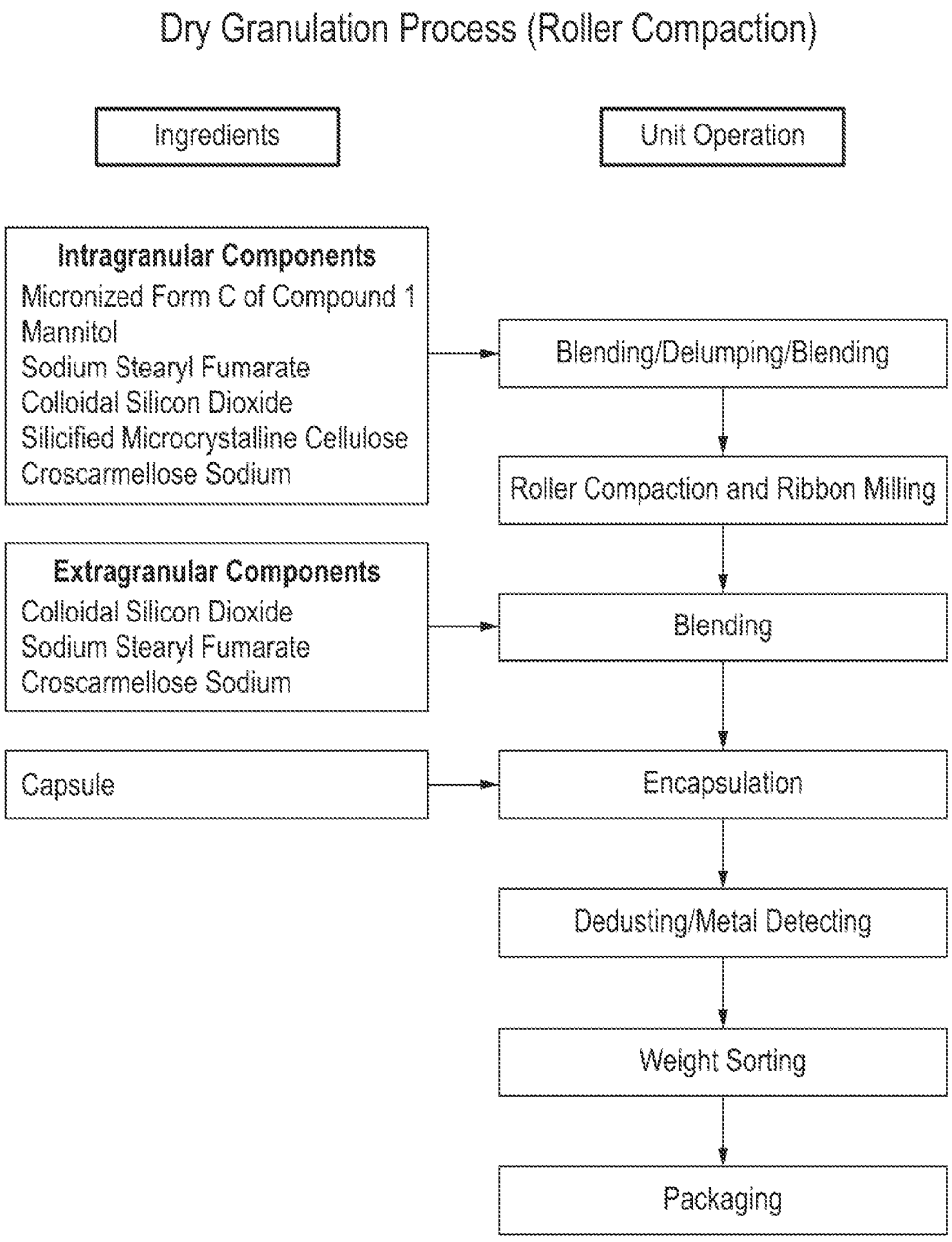

Dry Granulation Process (Roller Compaction)
Ingredients
Unit Operation
Intragranular Components
Micronized Form C of Compound 1
Mannitol
Sodium Stearyl Fumarate
Colloidal Silicon Dioxide
Silicified Microcrystalline Cellulose
Croscarmellose Sodium
Blending/Delumping/Blending
Roller Compaction and Ribbon Milling
Extragranular Components
Colloidal Silicon Dioxide
Sodium Stearyl Fumarate
Croscarmellose Sodium
Blending
Capsule
Encapsulation
Dedusting/Metal Detecting
Weight Sorting
Packaging

FIG. 9

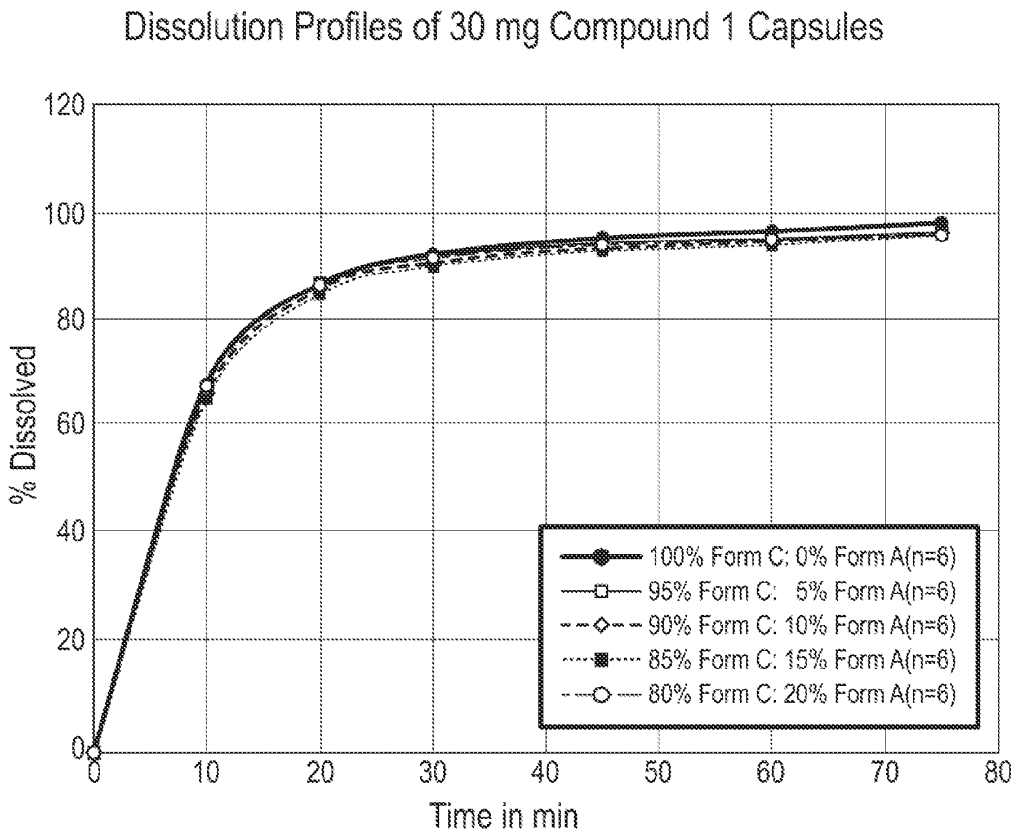

Dissolution Profiles of 30 mg Compound 1 Capsules
% Dissolved
Time in min
0
20
40
60
80
100
120
0
10
20
30
40
50
60
70
80
100% Form C: 0% Form A(n=6)
95% Form C: 5% Form A(n=6)
90% Form C: 10% Form A(n=6)
85% Form C: 15% Form A(n=6)
80% Form C: 20% Form A(n=6)

FIG. 10

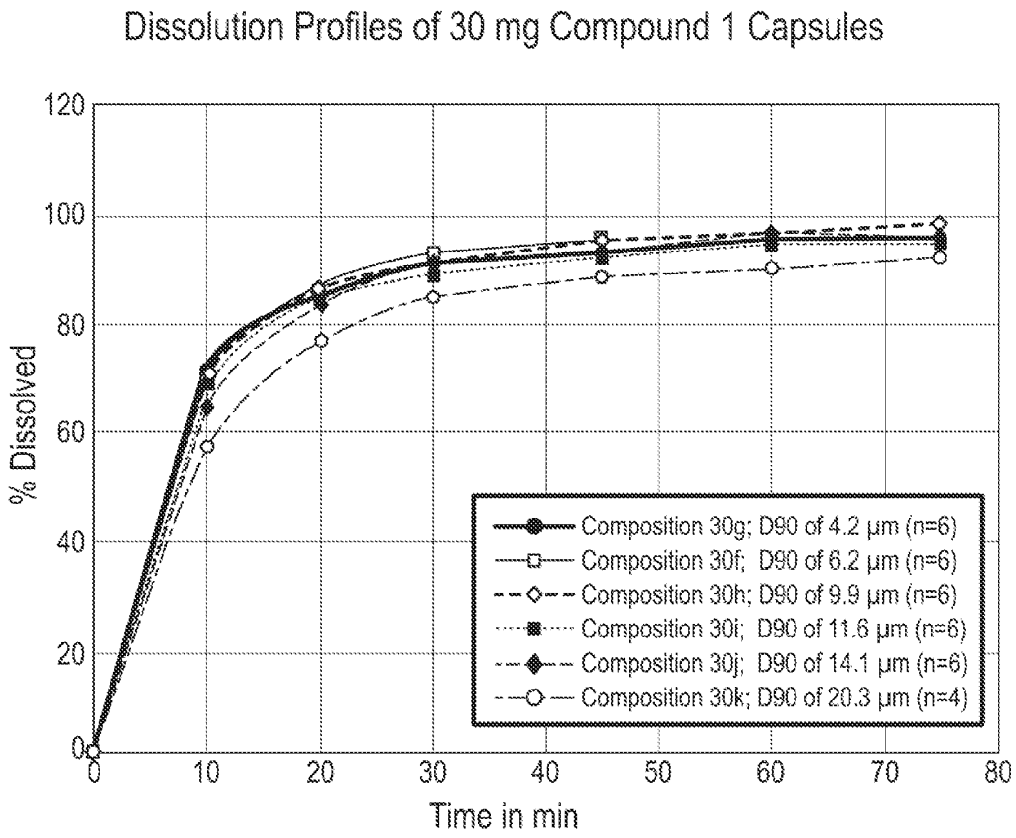

Dissolution Profiles of 30 mg Compound 1 Capsules
% Dissolved
Time in min
0
20
40
60
80
100
120
0
10
20
30
40
50
60
70
80
Composition 30g; D90 of 4.2 µm (n=6)
Composition 30f; D90 of 6.2 µm (n=6)
Composition 30h; D90 of 9.9 µm (n=6)
Composition 30i; D90 of 11.6 µm (n=6)
Composition 30j; D90 of 14.1 µm (n=6)
Composition 30k; D90 of 20.3 µm (n=4)

FIG. 11

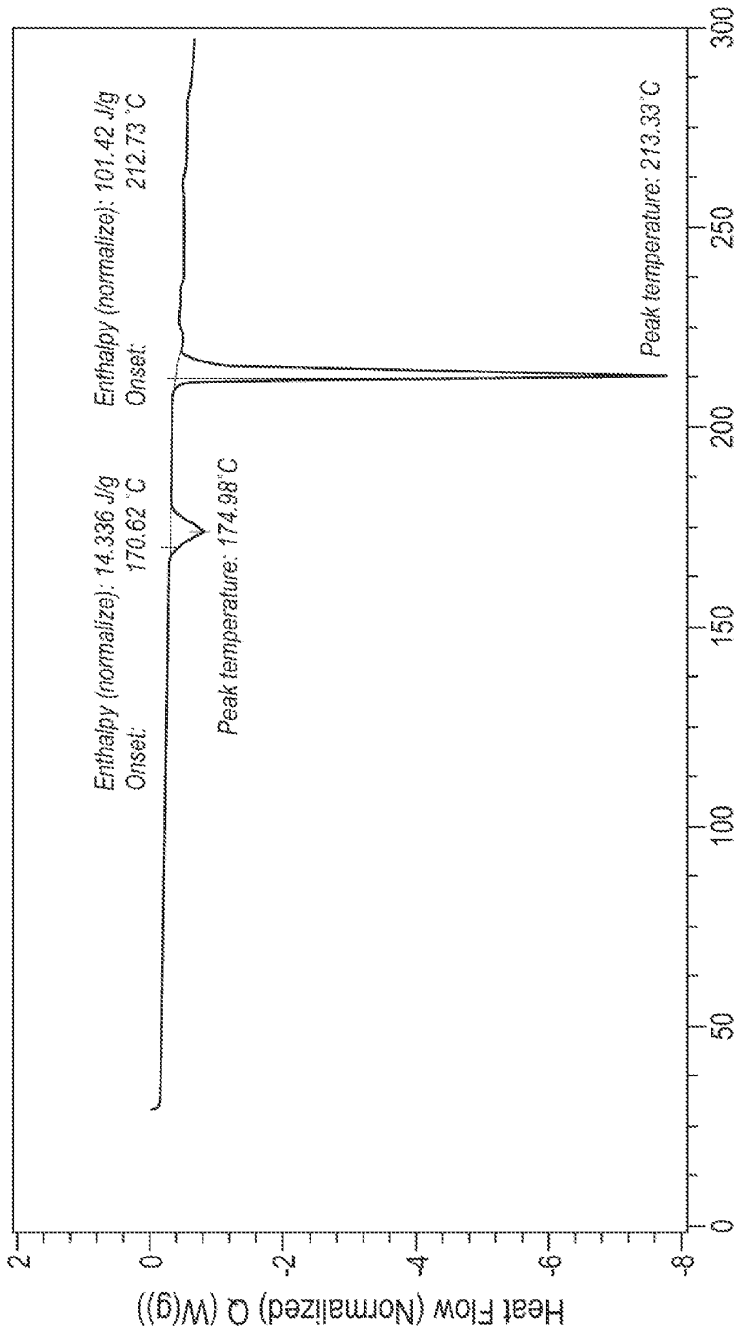

Heat Flow (Normalized) Q (W/g)
2
0
-2
-4
-6
-8
0
50
100
150
200
250
300
Temperature (°C)
Enthalpy (normalize): 14.336 J/g
Onset: 170.62 °C
Peak temperature: 174.98°C
Enthalpy (normalize): 101.42 J/g
Onset: 212.73 °C
Peak temperature: 213.33°C

FIG. 12A

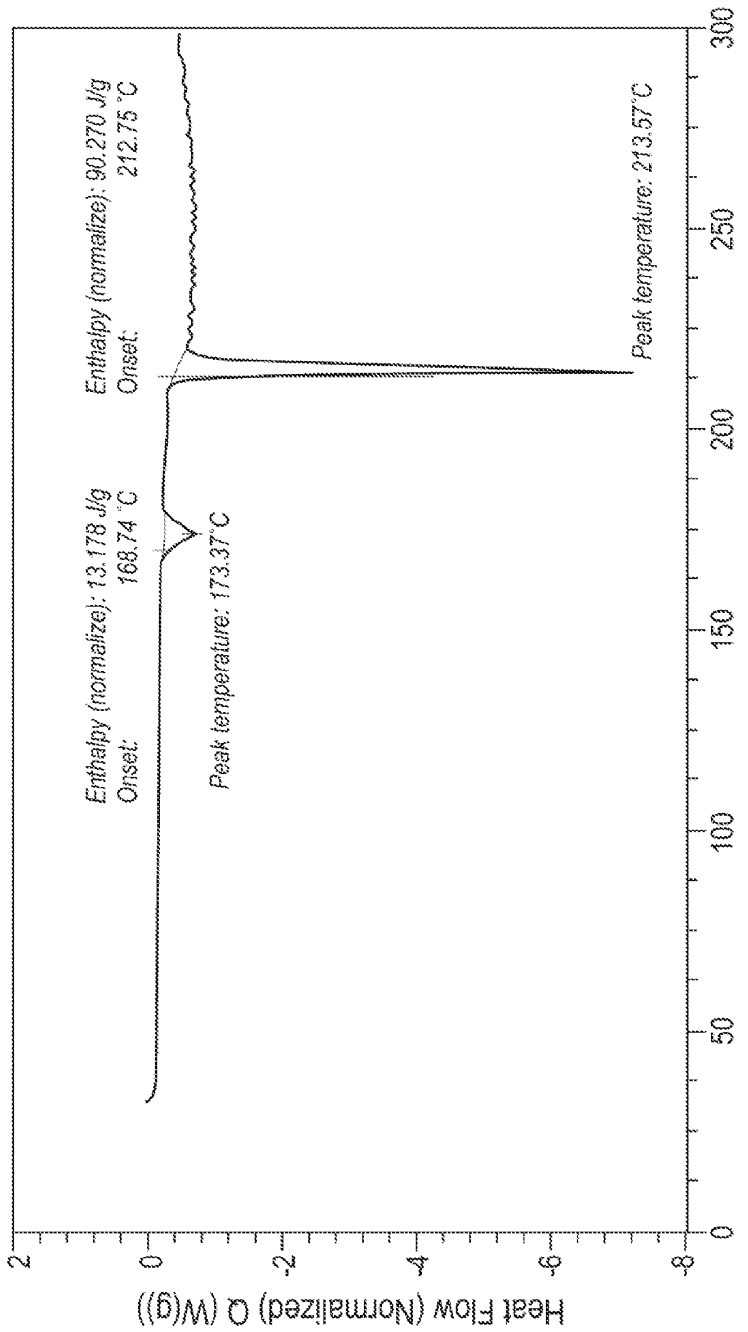

Heat Flow (Normalized) Q (W/g)
2
0
-2
-4
-6
-8
0
50
100
150
200
250
300
Temperature (°C)
Enthalpy (normalize): 13.178 J/g
Onset: 168.74 °C
Peak temperature: 173.37°C
Enthalpy (normalize): 90.270 J/g
Onset: 212.75 °C
Peak temperature: 213.57°C

FIG. 12B

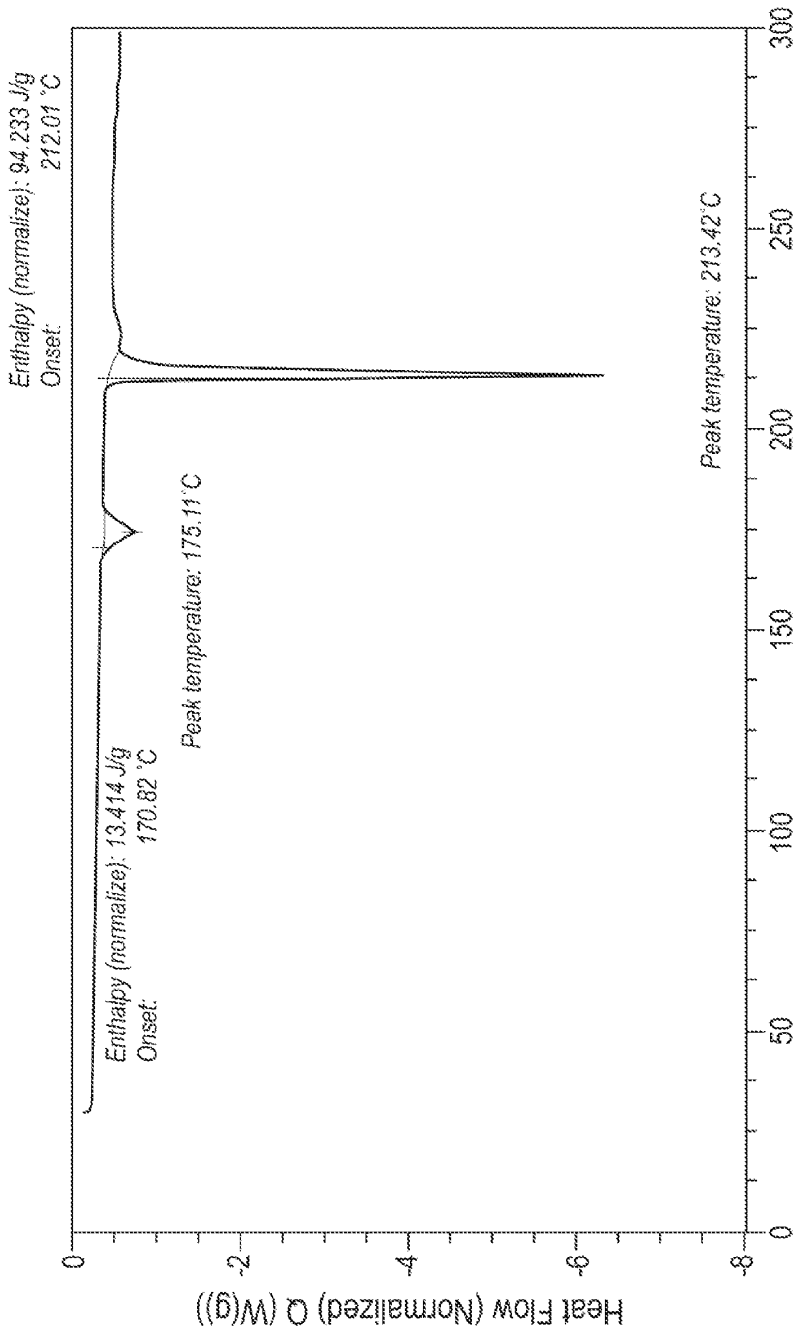

Enthalpy (normalize): 94.233 J/g
Onset: 212.01 °C
Enthalpy (normalize): 13.414 J/g
Onset: 170.82 °C
Peak temperature: 175.11 °C
Peak temperature: 213.42°C
Heat Flow (Normalized) Q (W/g))
Temperature (°C)
0
-2
-4
-6
-8
0
50
100
150
200
250
300

FIG. 12C

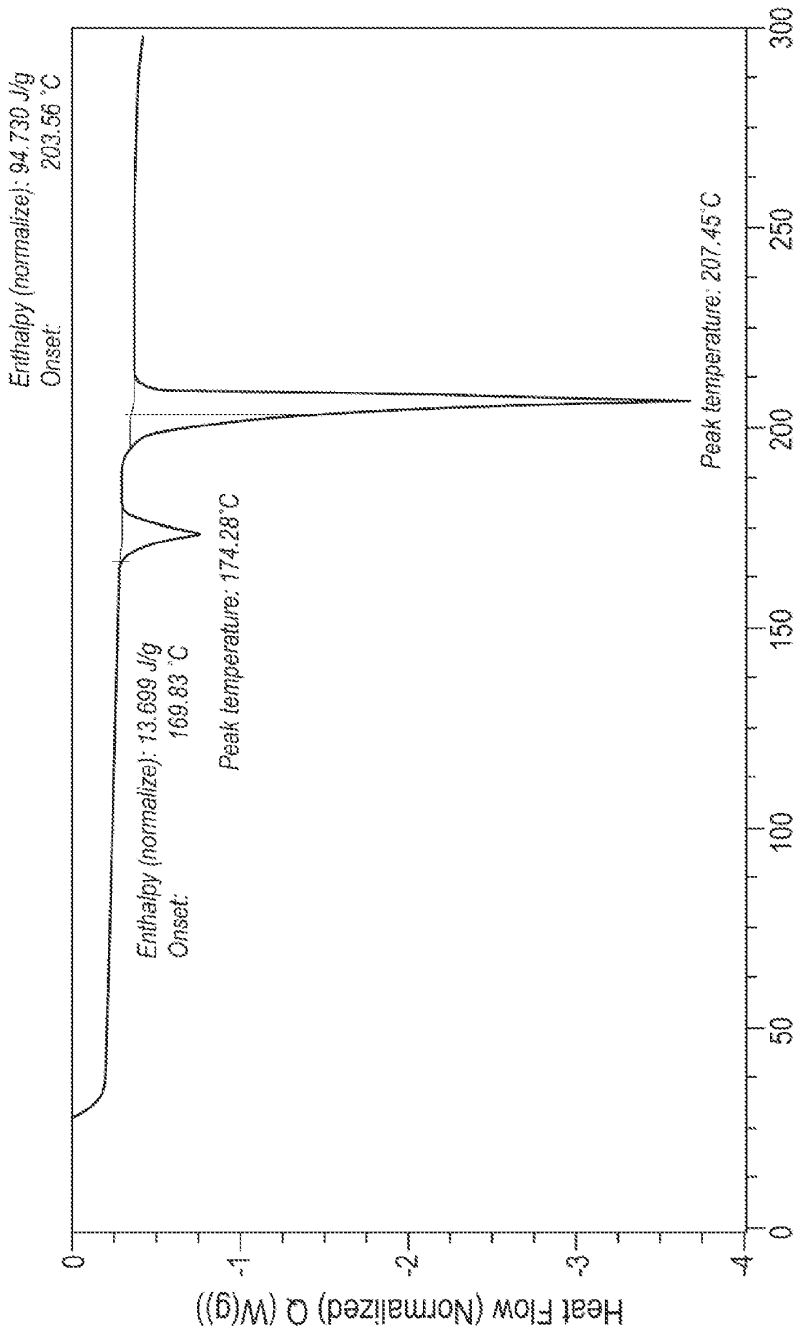

Enthalpy (normalize): 94.730 J/g
Onset: 203.56 °C
Enthalpy (normalize): 13.699 J/g
Onset: 169.83 °C
Peak temperature: 174.28°C
Peak temperature: 207.45°C
Heat Flow (Normalized) Q (W/g))
Temperature (°C)
0
-1
-2
-3
-4
0
50
100
150
200
250
300

FIG. 12D

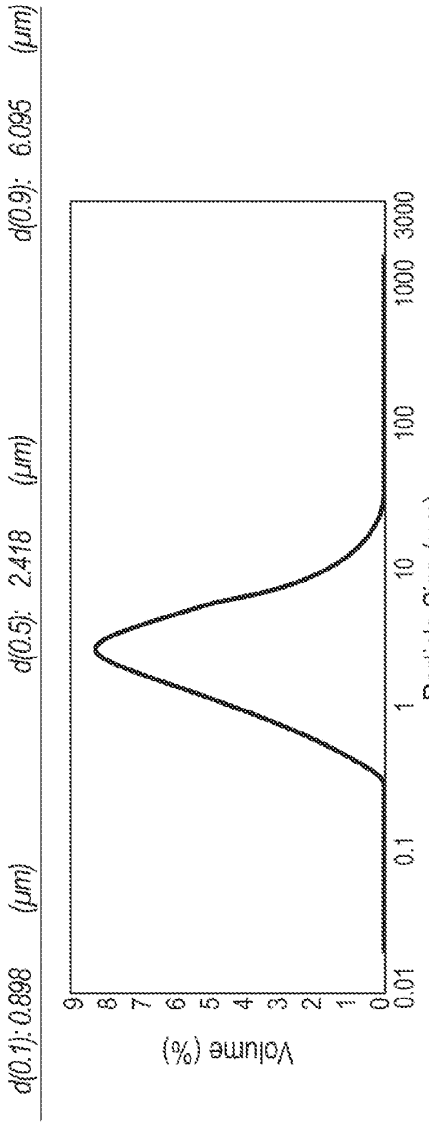


FIG. 13A

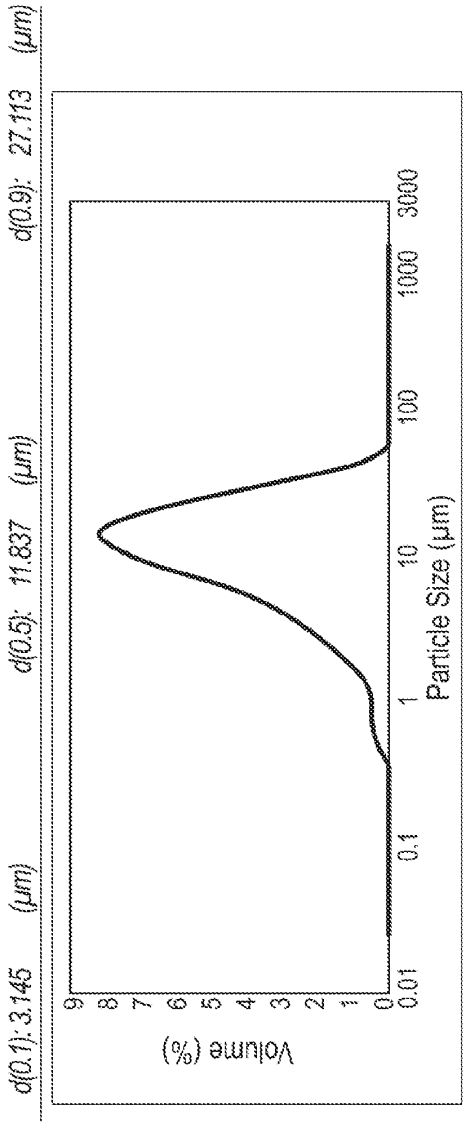


FIG. 13B

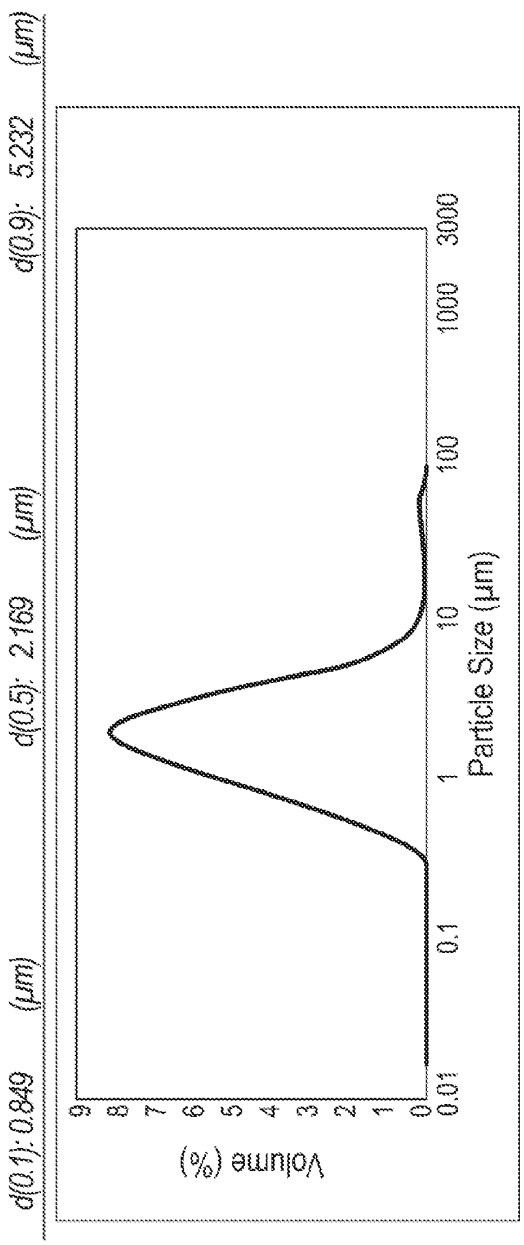

d(0.1): 0.849 (µm)
d(0.5): 2.169 (µm)
d(0.9): 5.232 (µm)
Volume (%)
0
1
2
3
4
5
6
7
8
9
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 14A

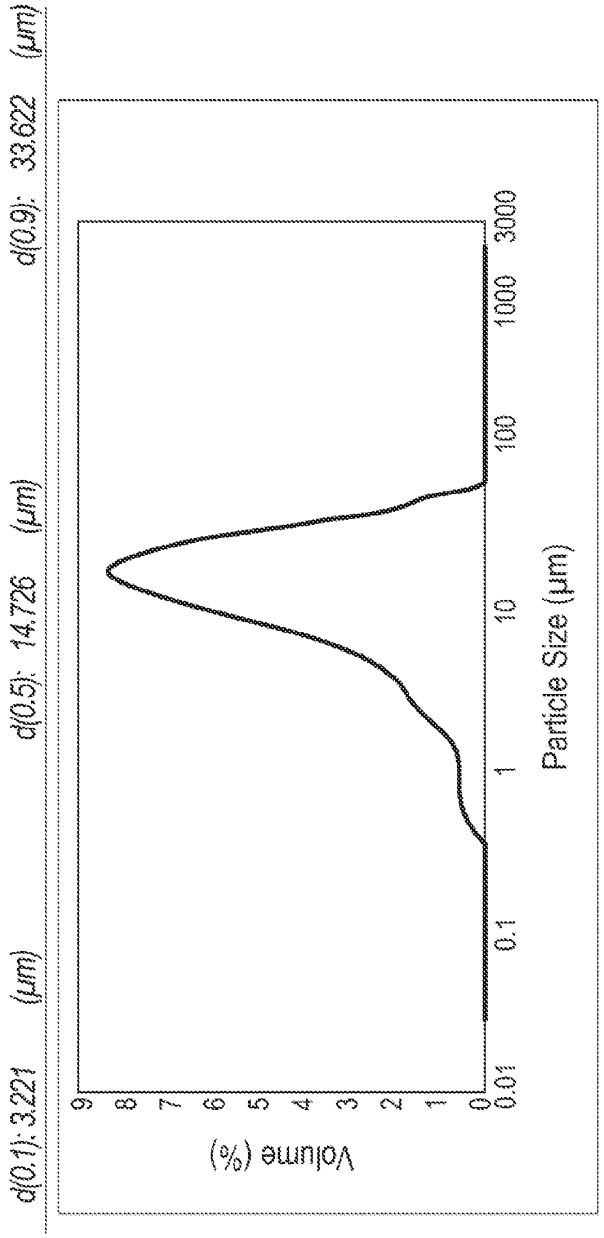

d(0.1): 3.221 (µm)
d(0.5): 14.726 (µm)
d(0.9): 33.622 (µm)
Volume (%)
0
1
2
3
4
5
6
7
8
9
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 14B

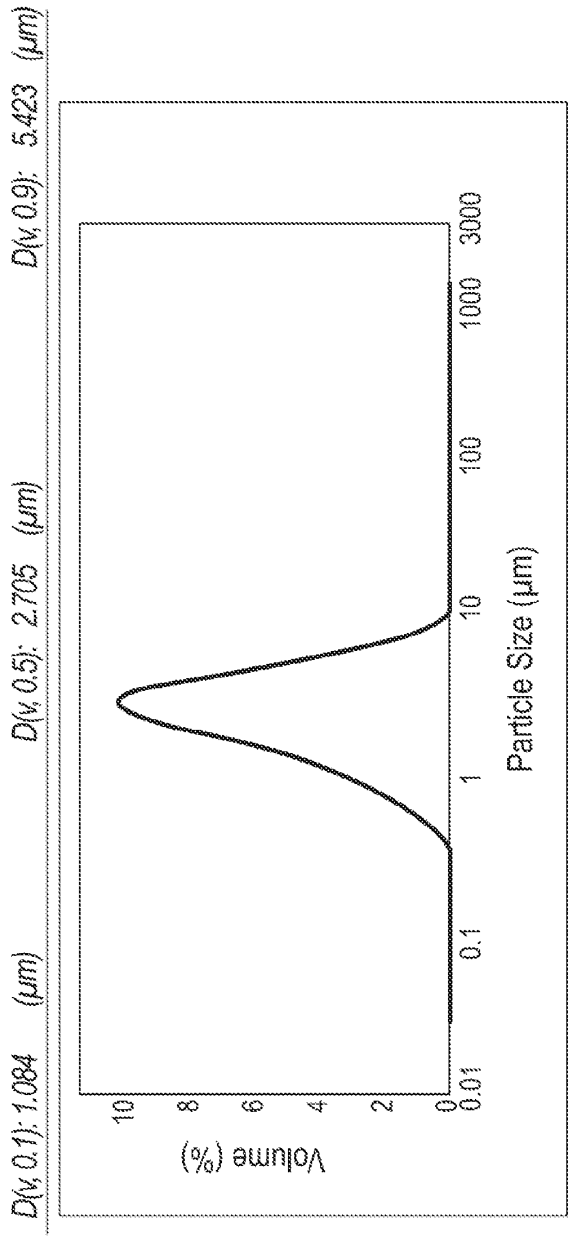

D(v, 0.1): 1.084 (µm)
D(v, 0.5): 2.705 (µm)
D(v, 0.9): 5.423 (µm)
Volume (%)
0
2
4
6
8
10
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 15A

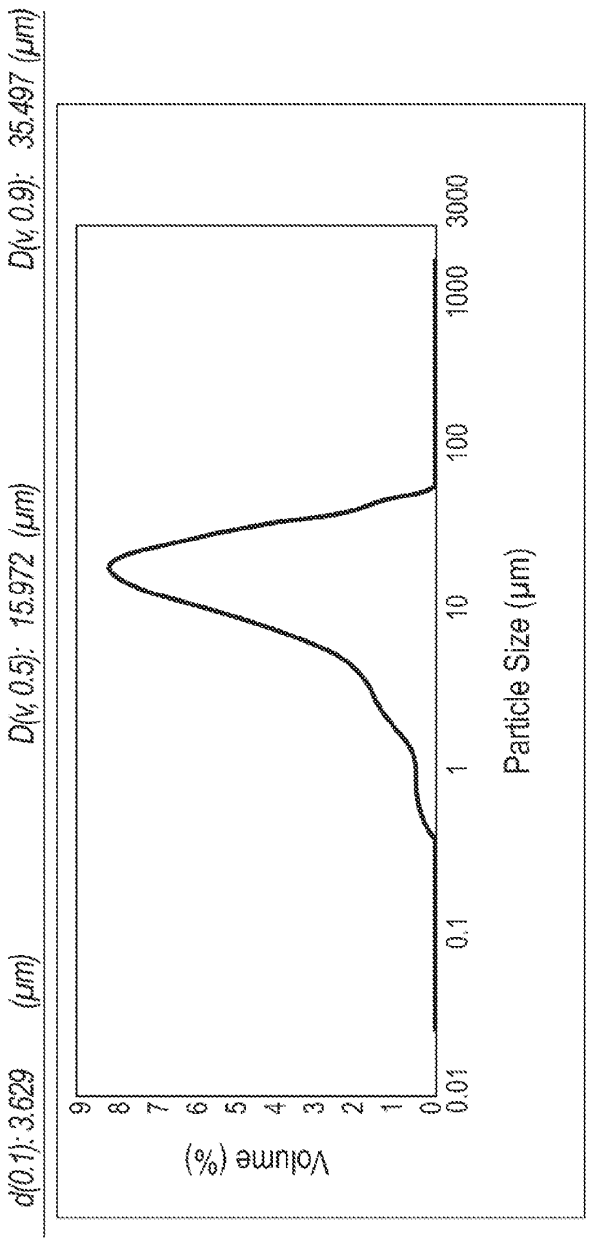

d(0.1): 3.629 (µm)
D(v, 0.5): 15.972 (µm)
D(v, 0.9): 35.497 (µm)
Volume (%)
0
1
2
3
4
5
6
7
8
9
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 15B

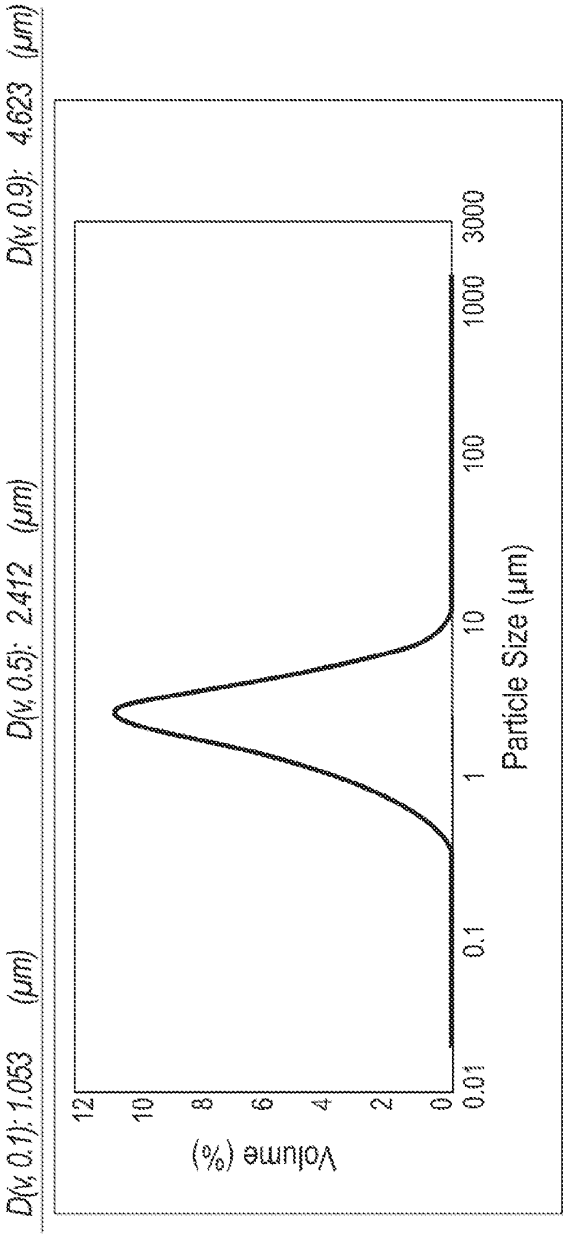

D(v, 0.1): 1.053 (µm)
D(v, 0.5): 2.412 (µm)
D(v, 0.9): 4.623 (µm)
Volume (%)
0
2
4
6
8
10
12
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 16A

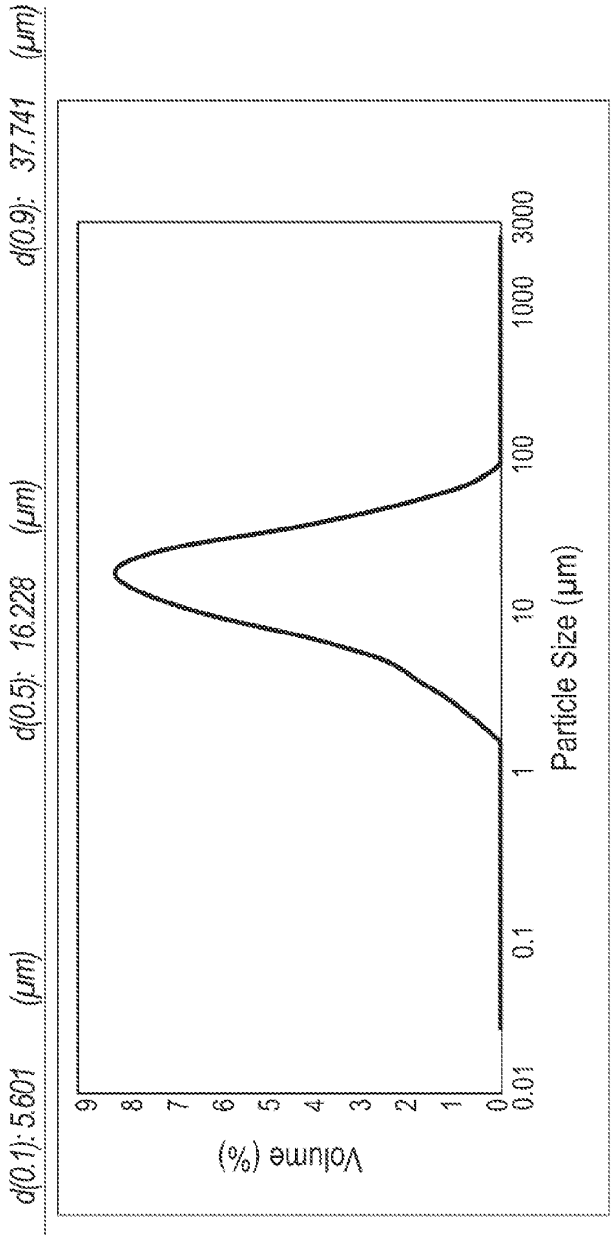

d(0.1): 5.601 (µm)
d(0.5): 16.228 (µm)
d(0.9): 37.741 (µm)
Volume (%)
0
1
2
3
4
5
6
7
8
9
0.01
0.1
1
10
100
1000
3000
Particle Size (µm)

FIG. 16B