

1 / 53

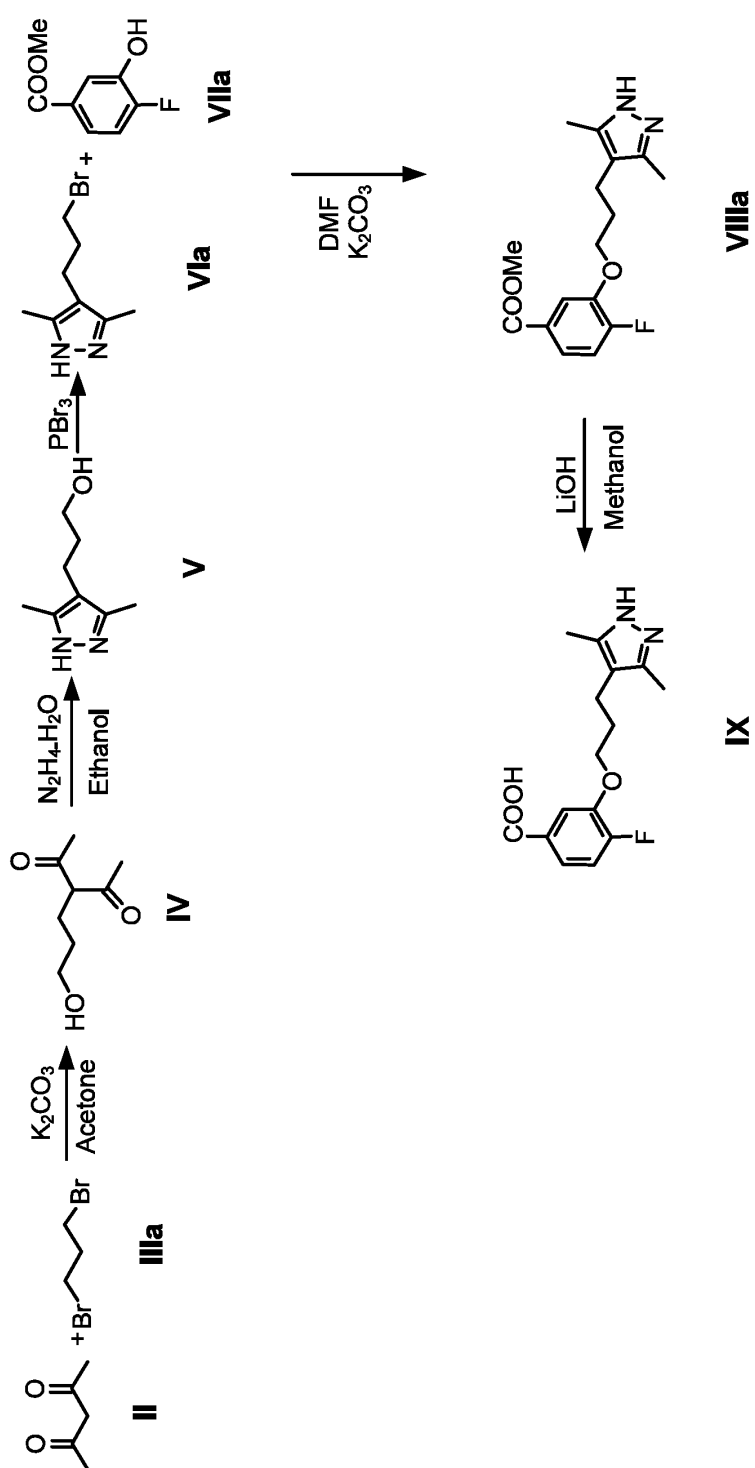


FIG. 1

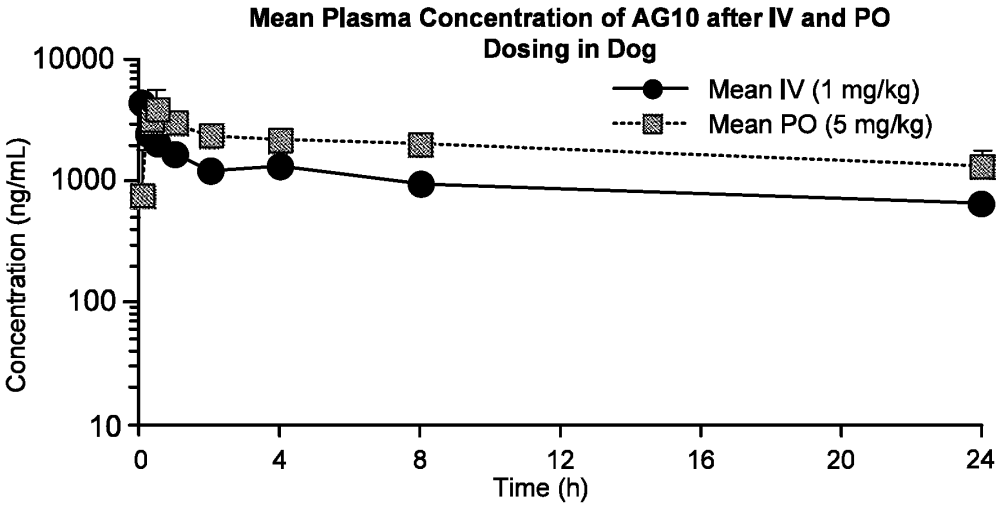


FIG. 2A

Dog		
IV 1mg/kg		
T _{1/2}	Vd _{ss}	Cl
(h)	(L/kg)	(mL/min/kg)
19.42 ± 7.64	0.660 ± 0.13	0.38 ± 0.05
PO 5mg/kg		
%F		
(%)		
39.6		

FIG. 2B

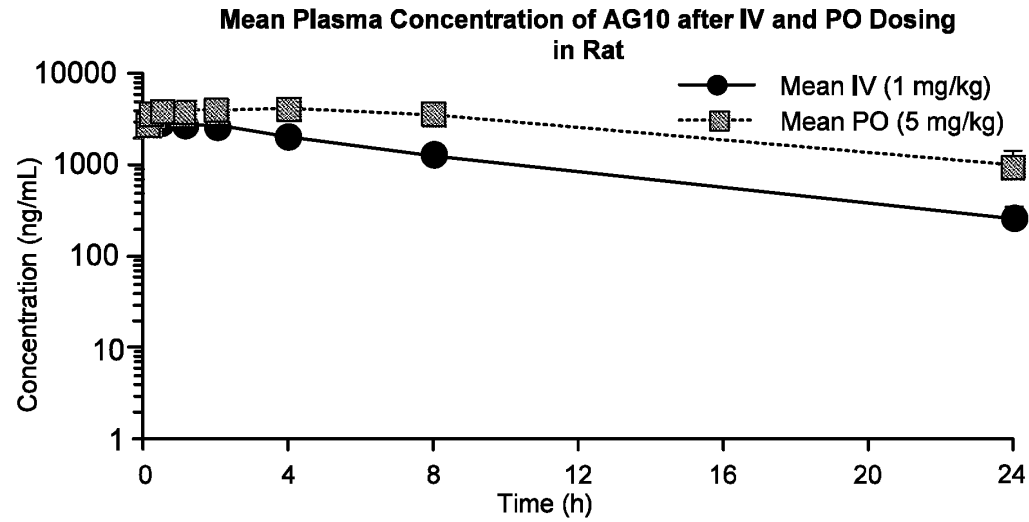
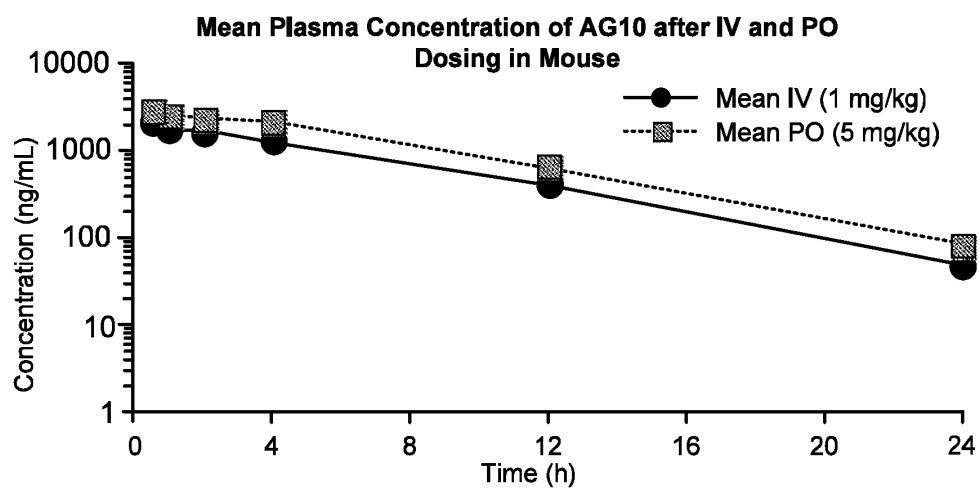


FIG. 2C

Rat		
IV 1mg/kg		
T _{1/2}	Vd _{ss}	Cl
(h)	(L/kg)	(mL/min/kg)
11.92 ± 1.75	0.41 ± 0.02	0.49 ± 0.11
PO 5mg/kg		
%F		
(%)		
60.5		

FIG. 2D

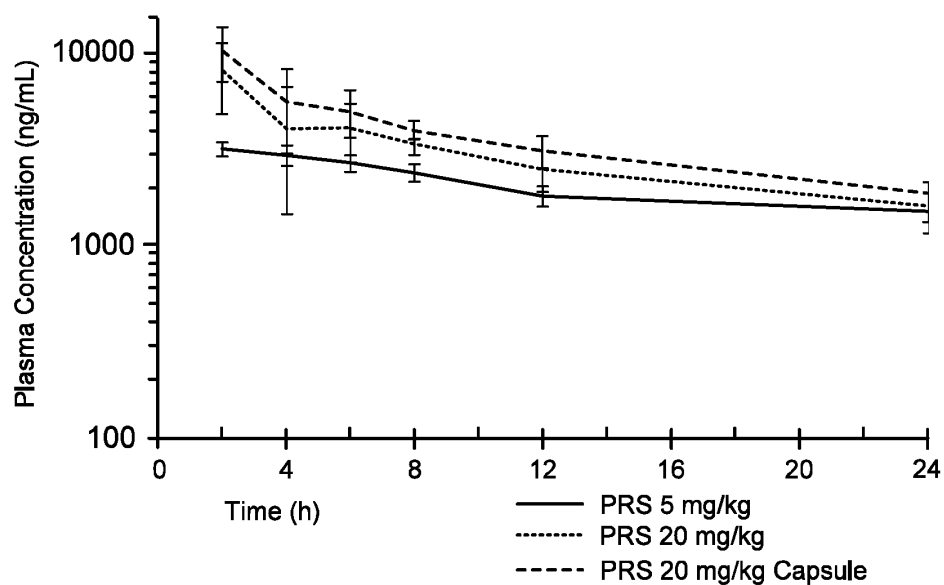
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**FIG. 2E**

Mouse		
IV 1mg/kg		
$T_{1/2}$	V_{dss}	Cl
(h)	(L/kg)	(mL/min/kg)
5.32 ± 0.62	0.47 ± 0.13	1.13 ± 0.26
PO 5mg/kg		
%F		
(%)		
30.4		

FIG. 2F

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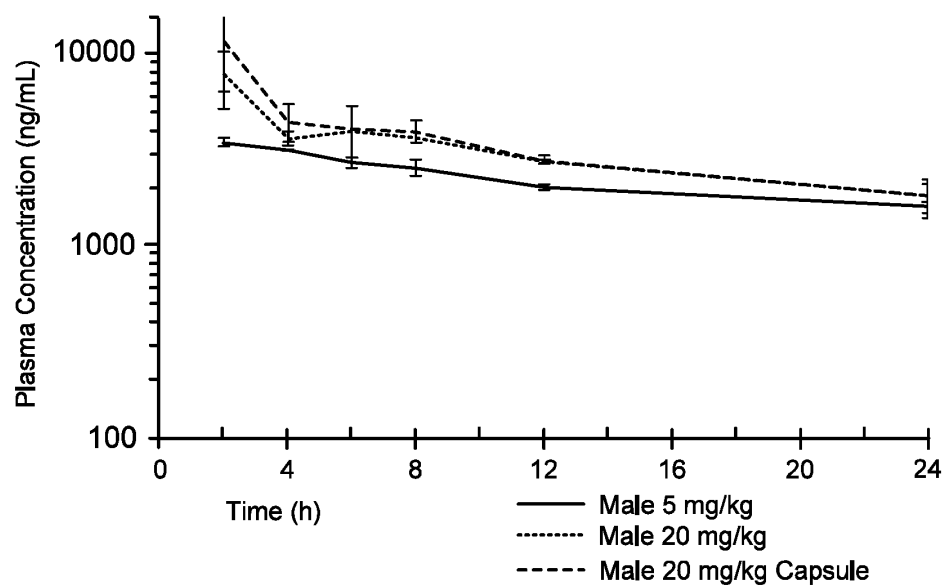
PRS Dog 1 Day Combined**FIG. 3A**

PRS Dog 1 Day		
C _{max}	C _{min} (24h)	AUC (24hr)
(ng/mL)	(ng/mL)	ng·hr/mL
n=4, ± SD	n=4, ± SD	n=4
PO 5mg/kg		
3143 ± 257.7	1490 ± 352.4	44785 ± 5078.4
PO 20mg/kg		
7903 ± 3511.5	1573 ± 323.9	63343 ± 5387.8
PO 20mg/kg Capsule		
10288 ± 3171.2	1853 ± 264.6	79043 ± 10019.9

*AUC does not include expected T_{max} at 1 hr
 AUC calculated by MS Excel

FIG. 3B

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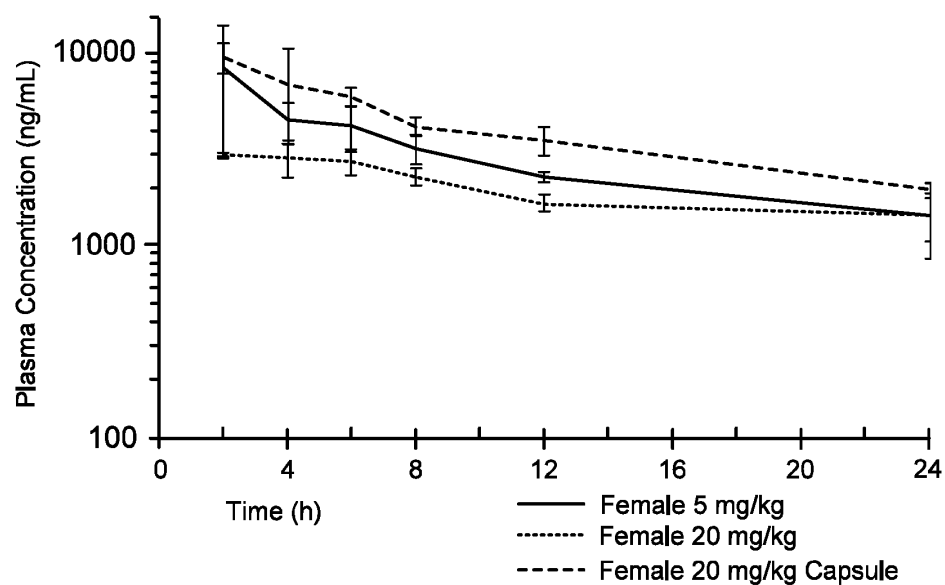
PRS Dog 1 Day Male**FIG. 3C**

PRS Dog 1 Day Male		
C _{max}	C _{min} (24h)	AUC (24hr)
(ng/mL)	(ng/mL)	ng·hr/mL
n=2, ± SD	n=2, ± SD	n=2
PO 5mg/kg		
3140 ± 198.0	1540 ± 99.0	46940 ± 1244.5
PO 20mg/kg		
7490 ± 2474.9	1735 ± 289.9	65060 ± 6010.4
PO 20mg/kg Capsule		
11090 ± 4963.9	1750 ± 396.0	70925 ± 5805.3

*AUC does not include expected T_{max} at 1 hr
AUC calculated by MS Excel

FIG. 3D

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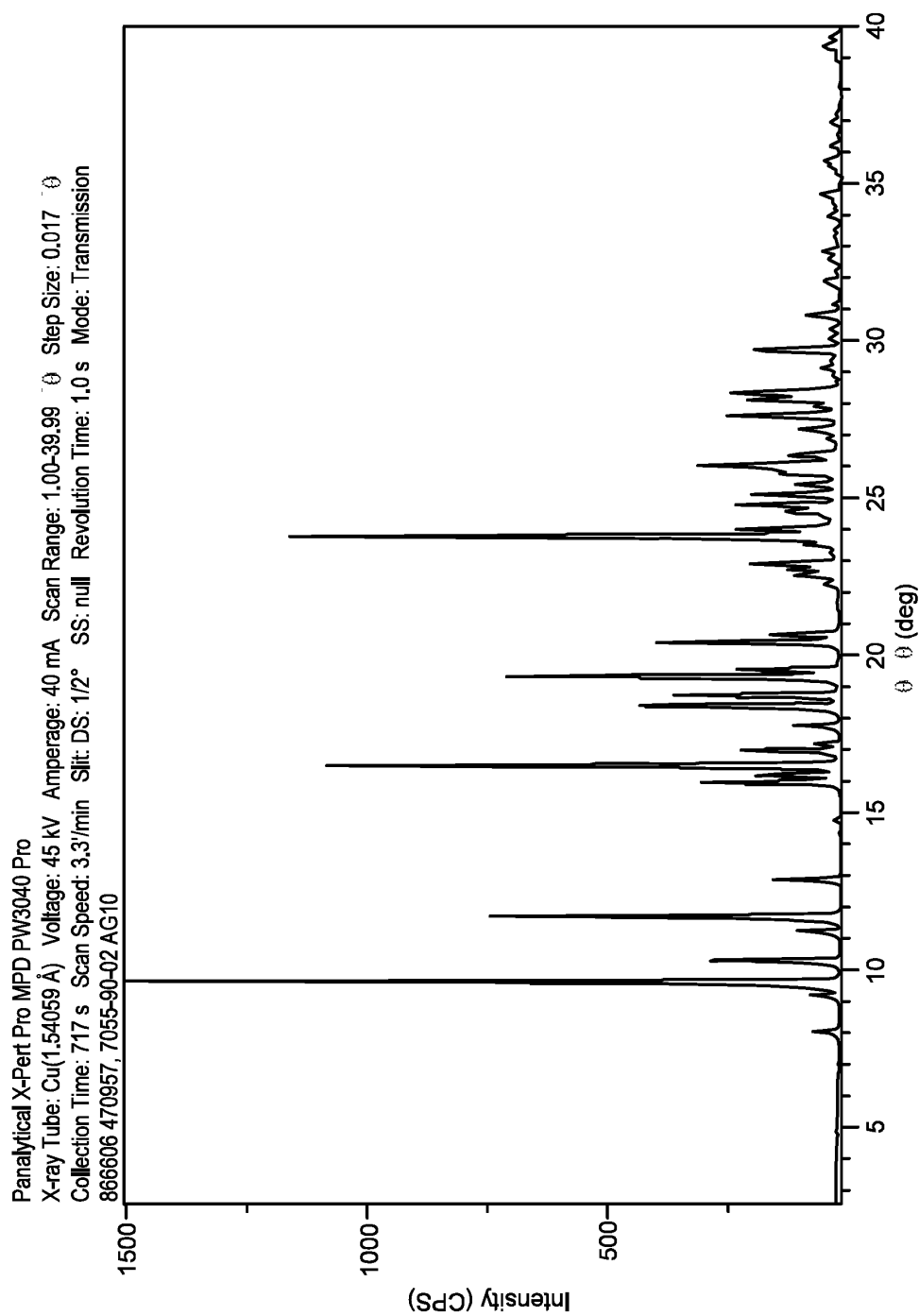
PRS Dog 1 Day Female**FIG. 3E**

PRS Dog 1 Day Female		
C_{max}	C_{min} (24h)	AUC (24hr)
(ng/mL)	(ng/mL)	ng-hr/mL
n=2, $\pm$ SD	n=2, $\pm$ SD	n=2
PO 5mg/kg		
2945 $\pm$ 63.6	1440 $\pm$ 594.0	42630 $\pm$ 7566.0
PO 20mg/kg		
8315 $\pm$ 5494.2	1410 $\pm$ 353.6	61625 $\pm$ 6257.9
PO 20mg/kg Capsule		
9485 $\pm$ 1718.3	1955 $\pm$ 106.1	871603 $\pm$ 1979.9

*AUC does not include expected T_{max} at 1 hr
 AUC calculated by MS Excel

FIG. 3F

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**FIG. 4**

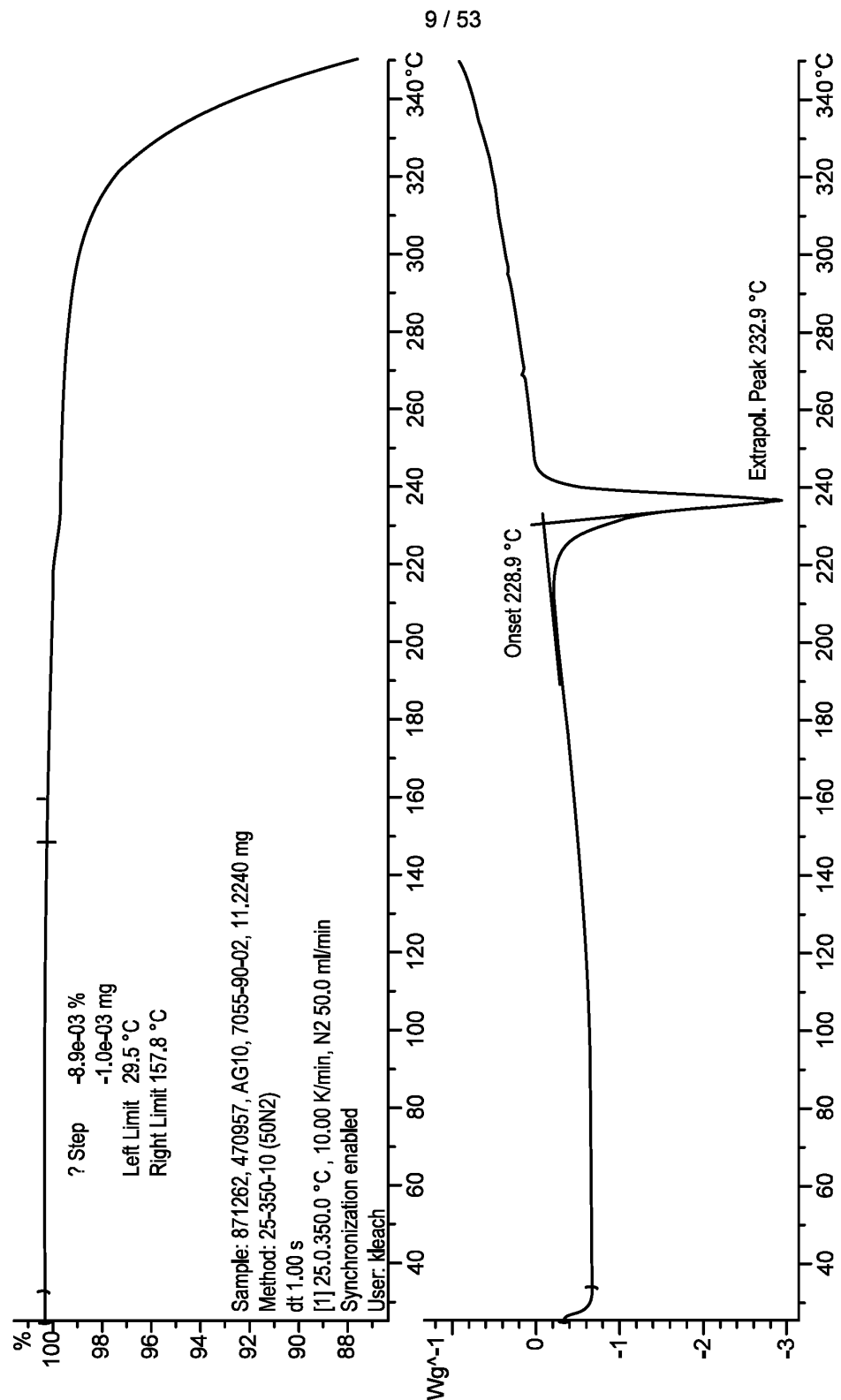


FIG. 5

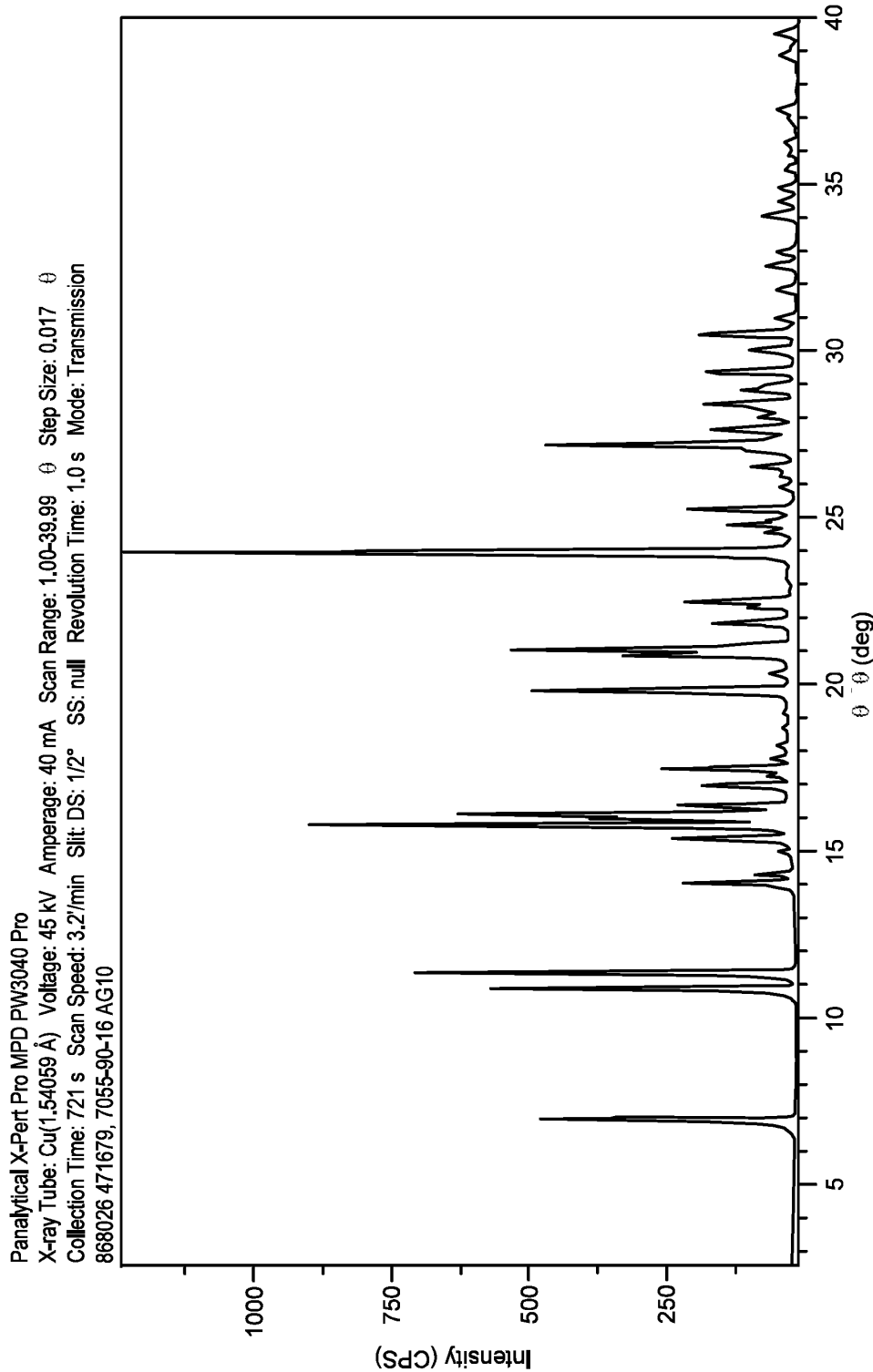


FIG. 6

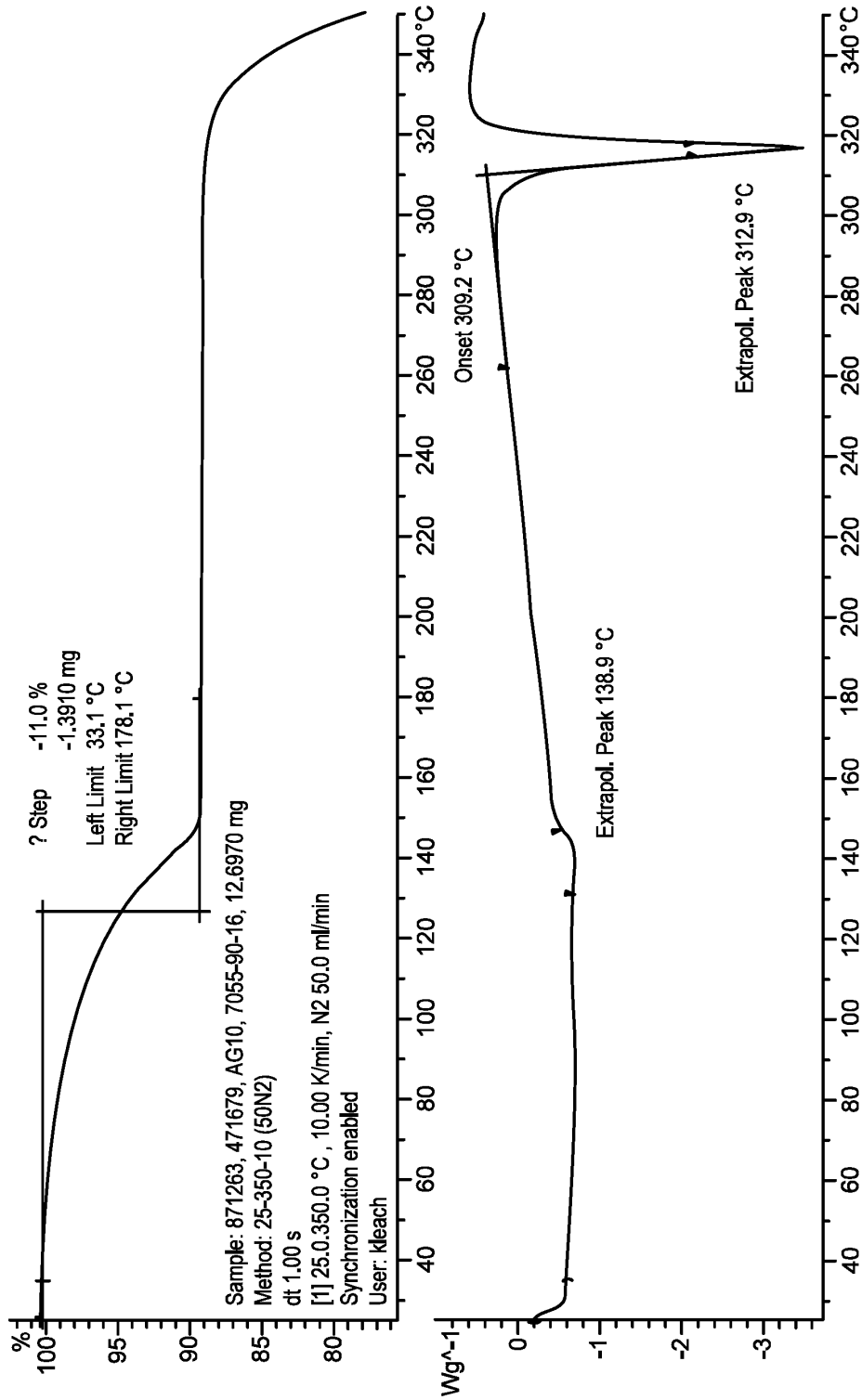
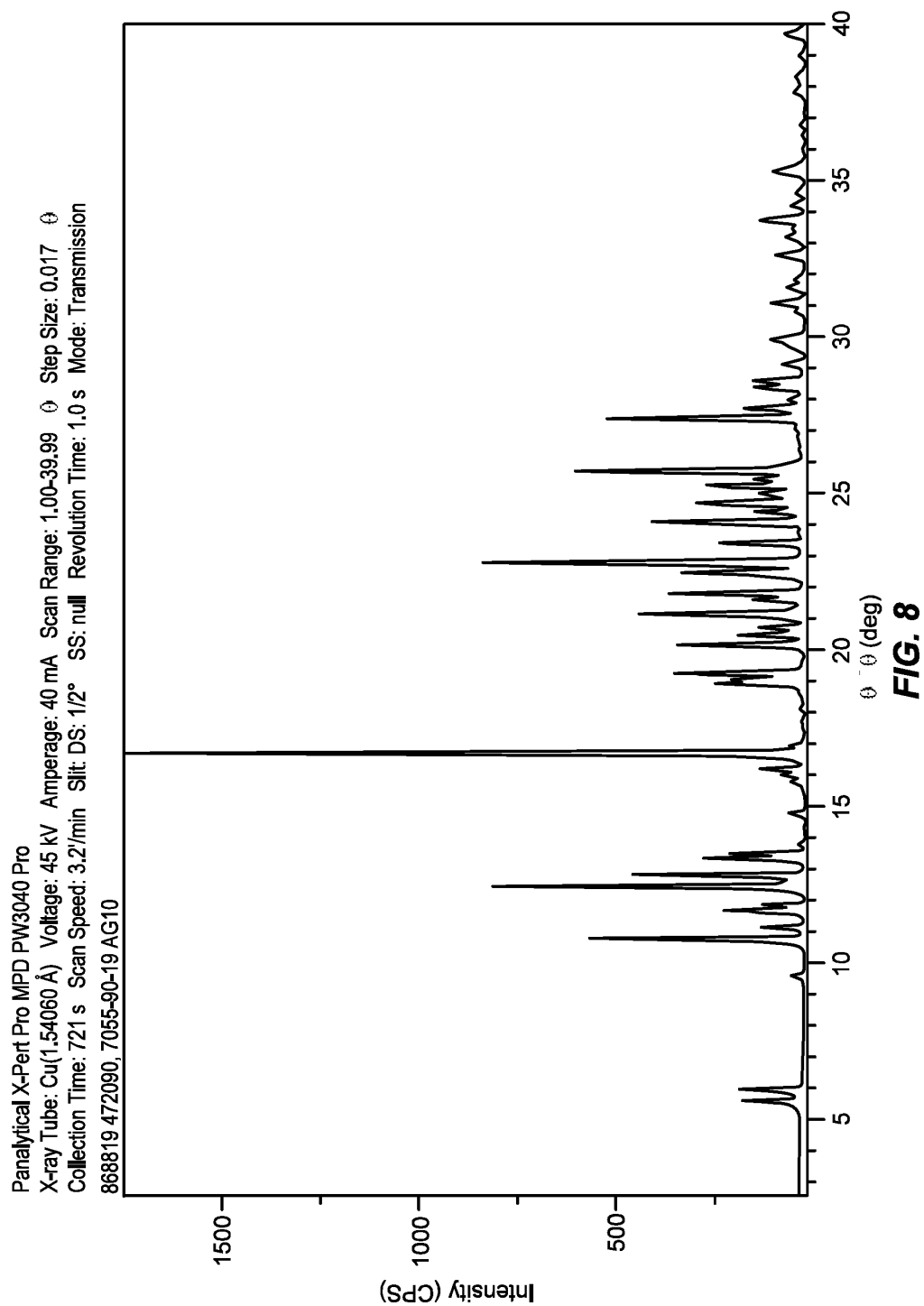


FIG. 7

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SUBSTITUTE SHEET (RULE 26)

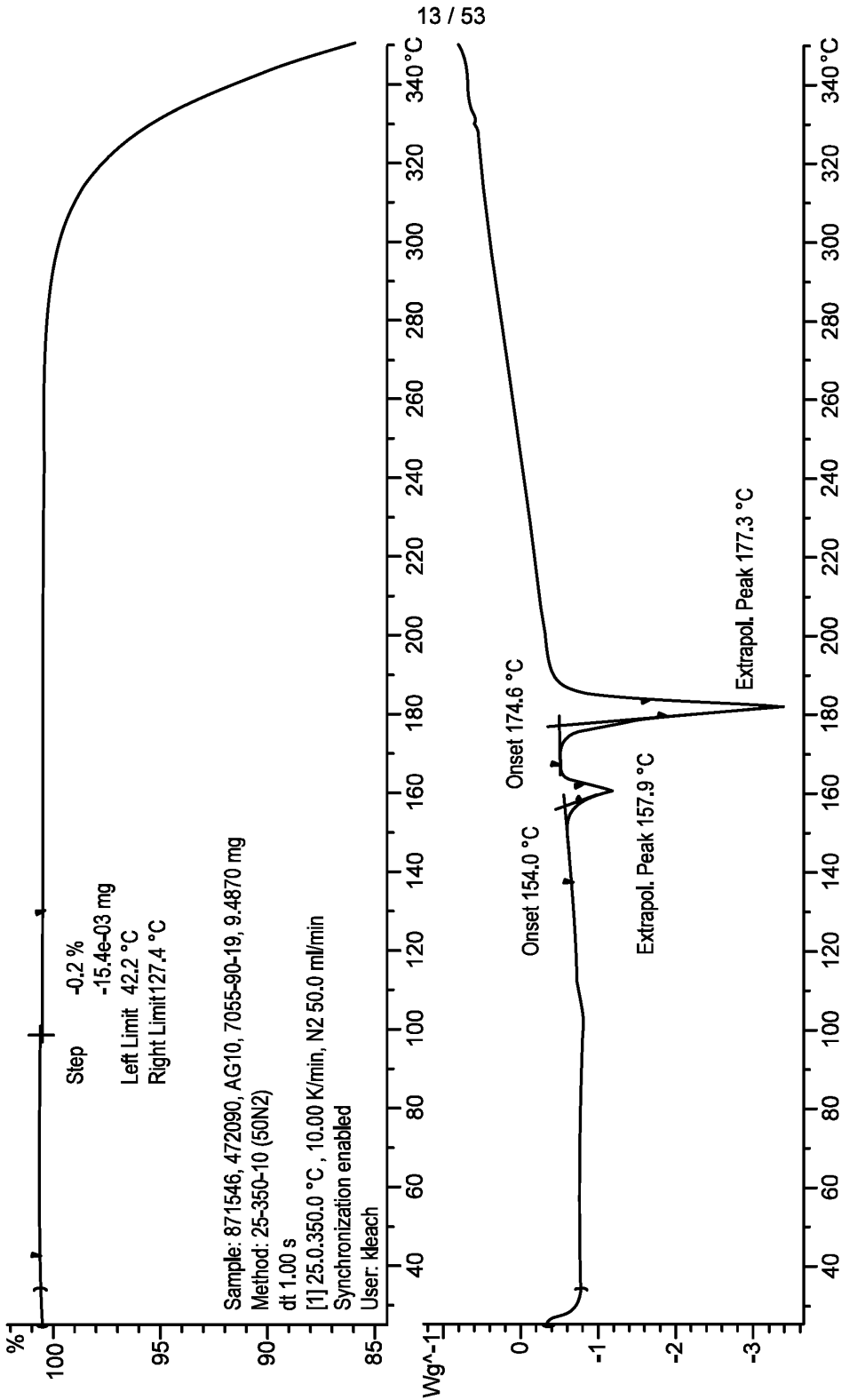


FIG. 9

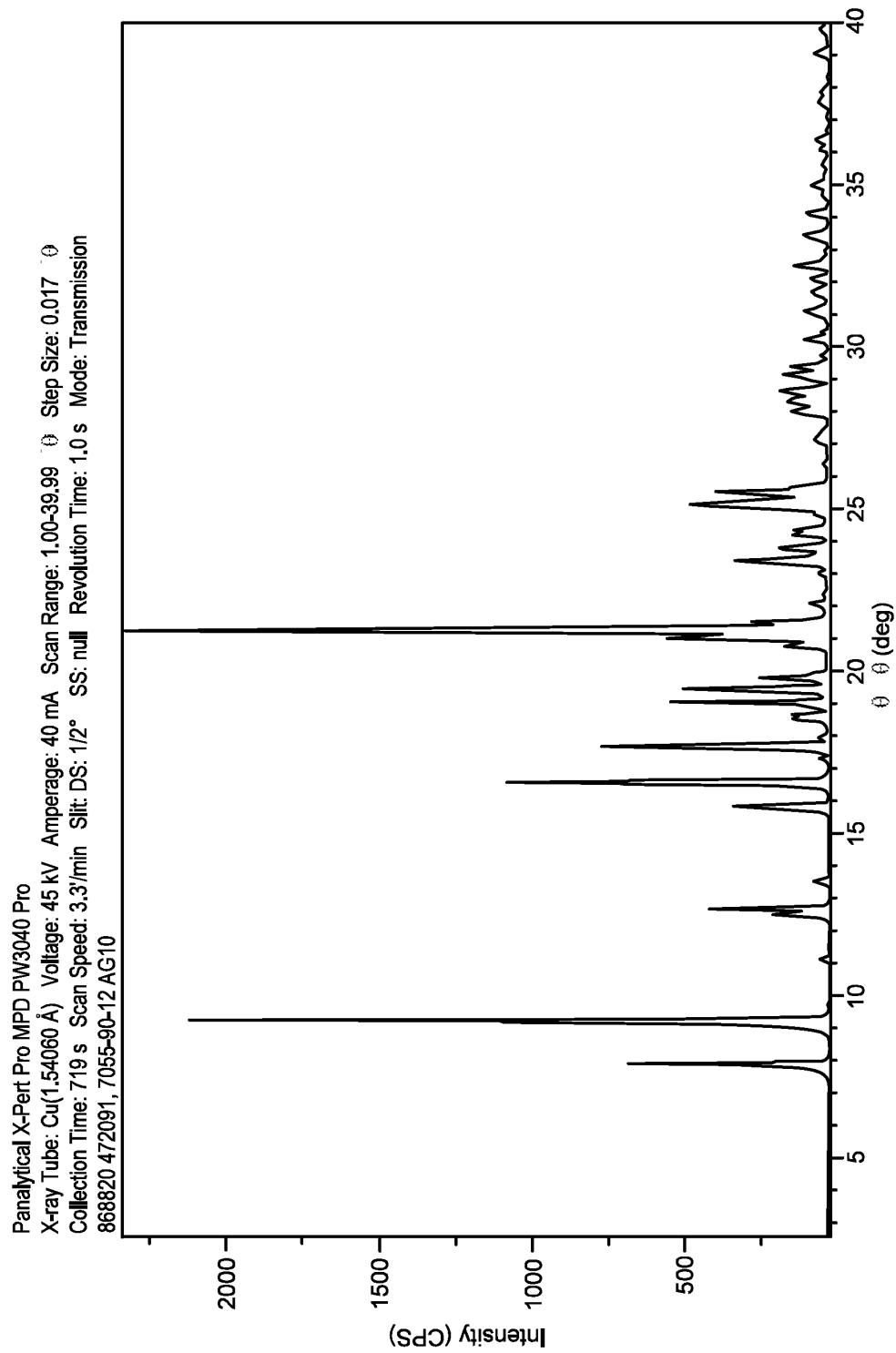


FIG. 10

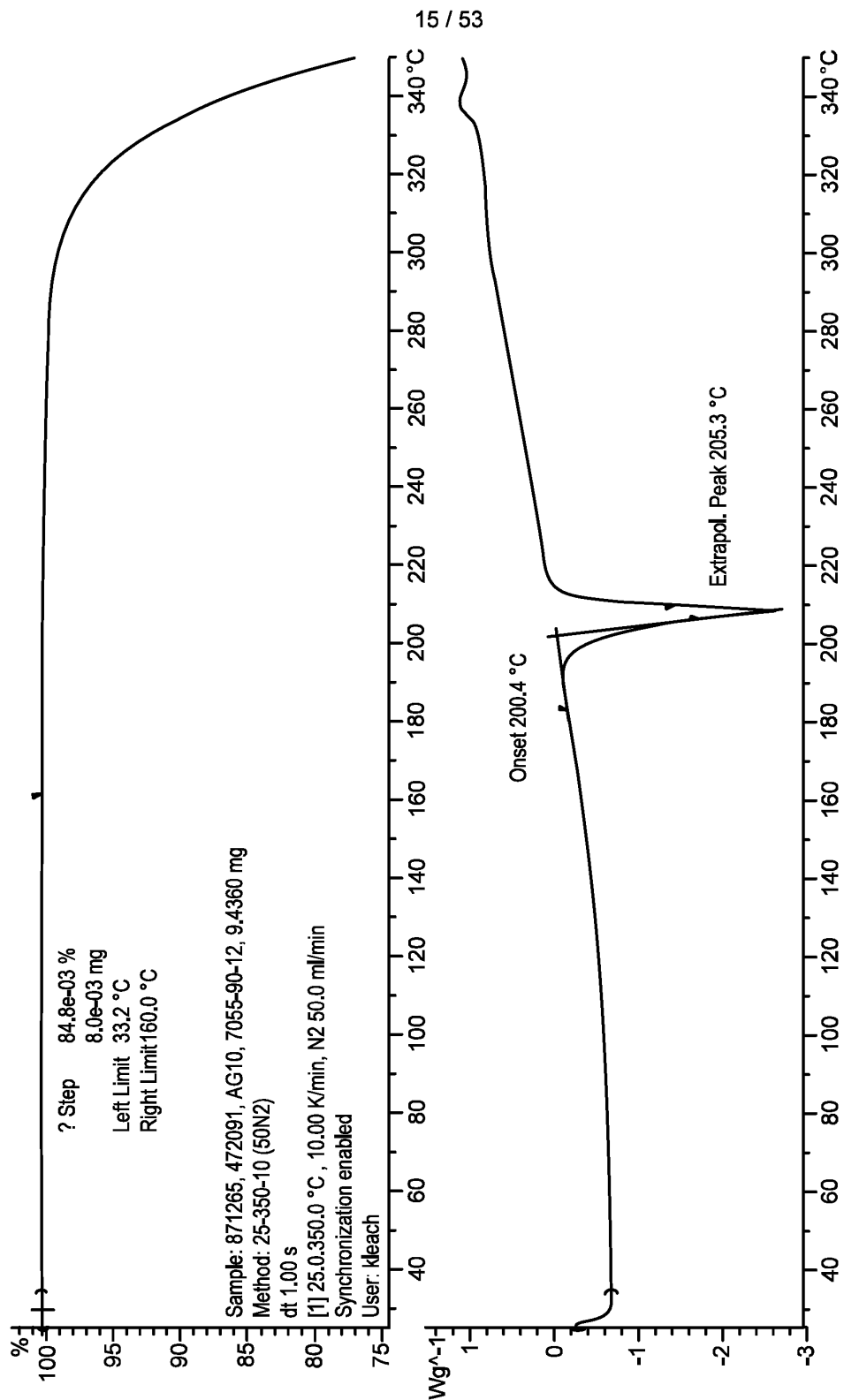
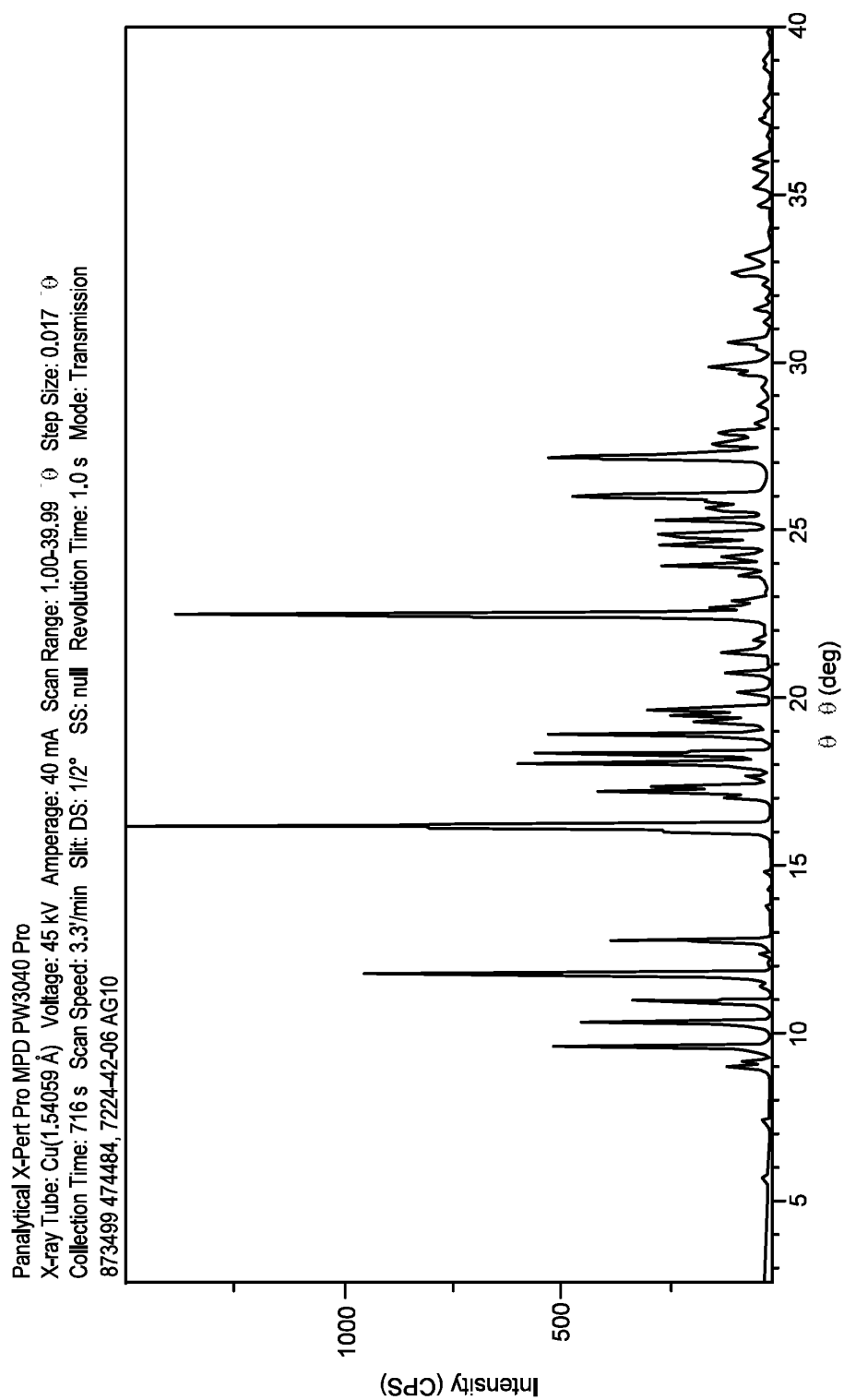


FIG. 11

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**FIG. 12**

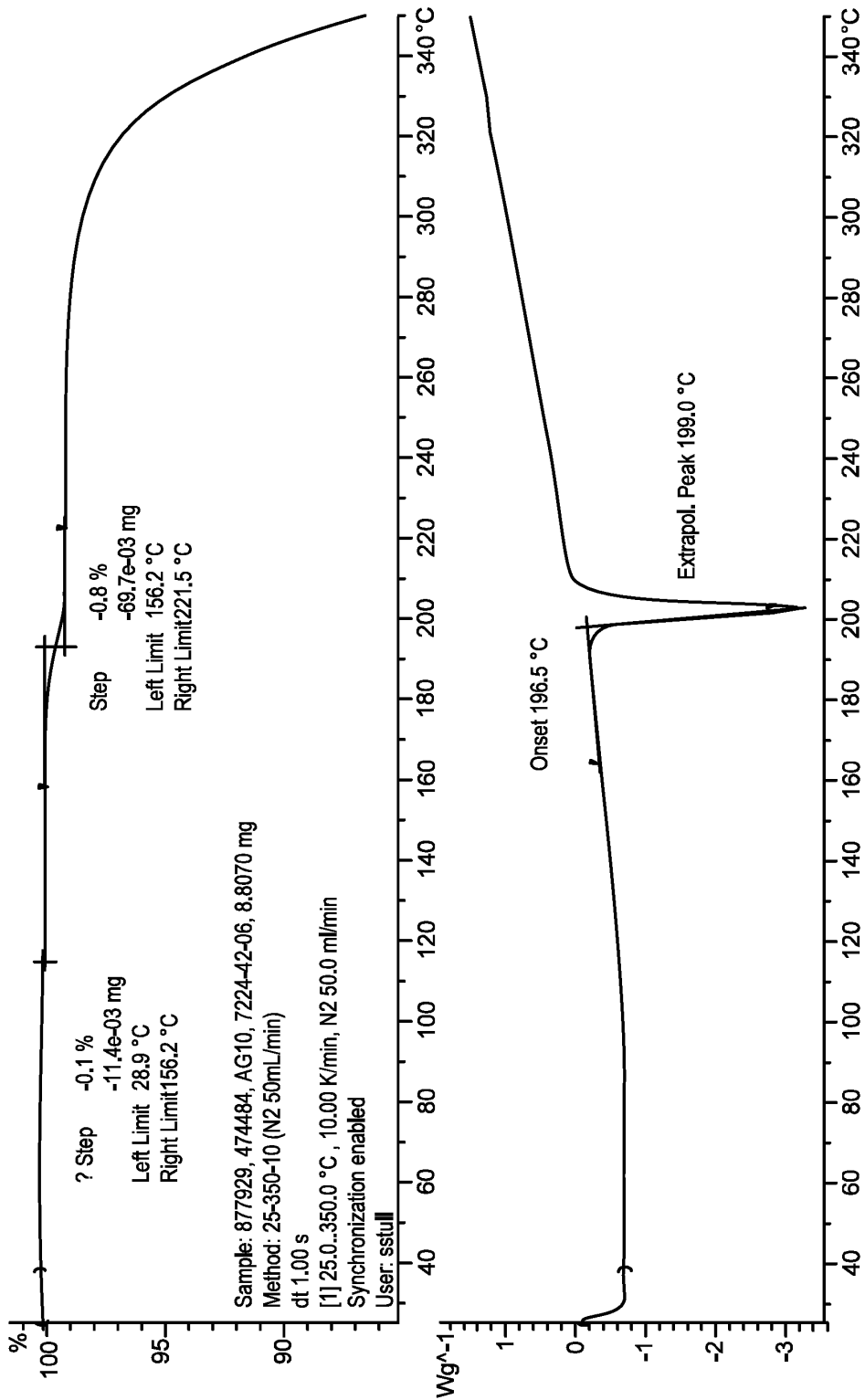
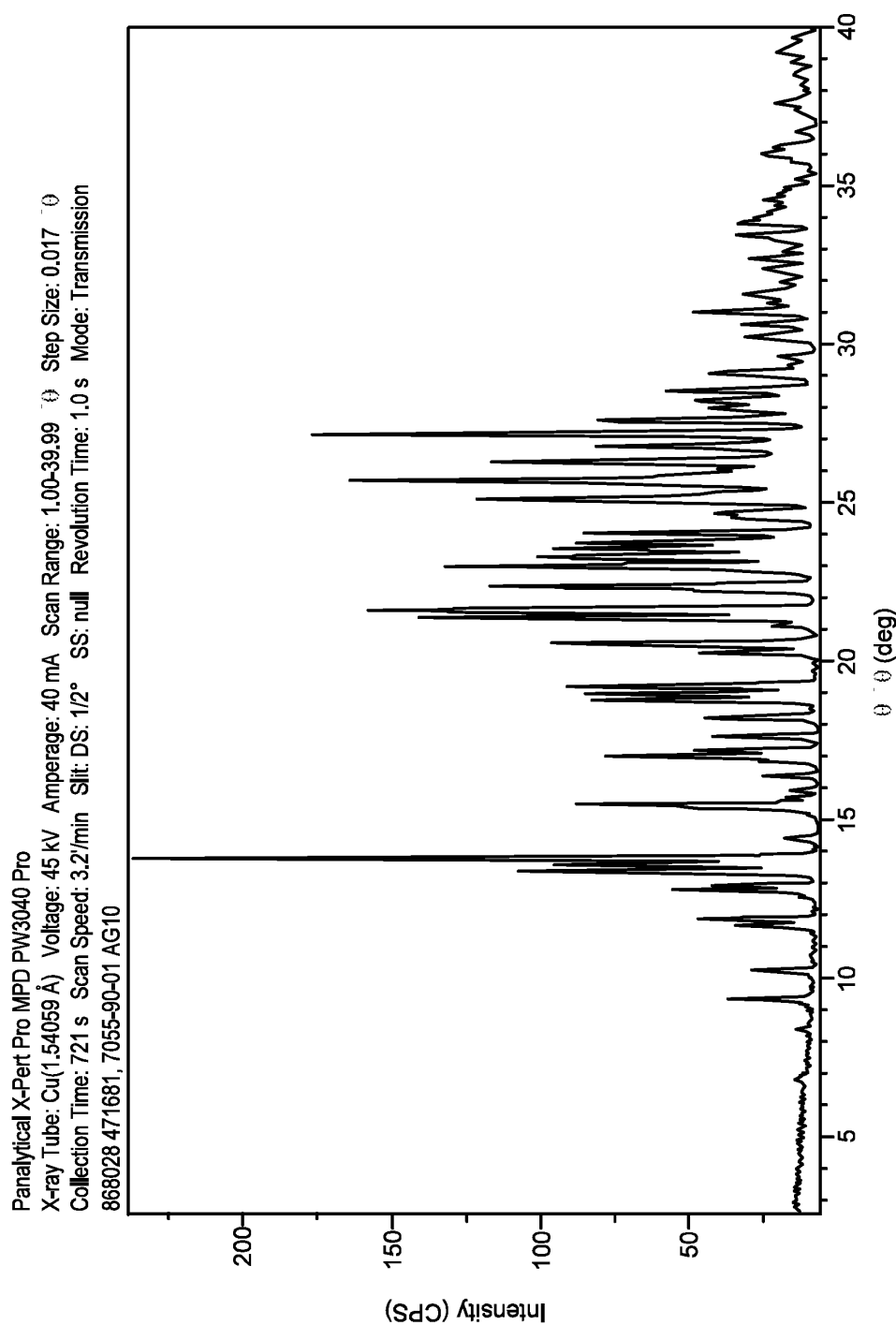


FIG. 13

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**FIG. 14**

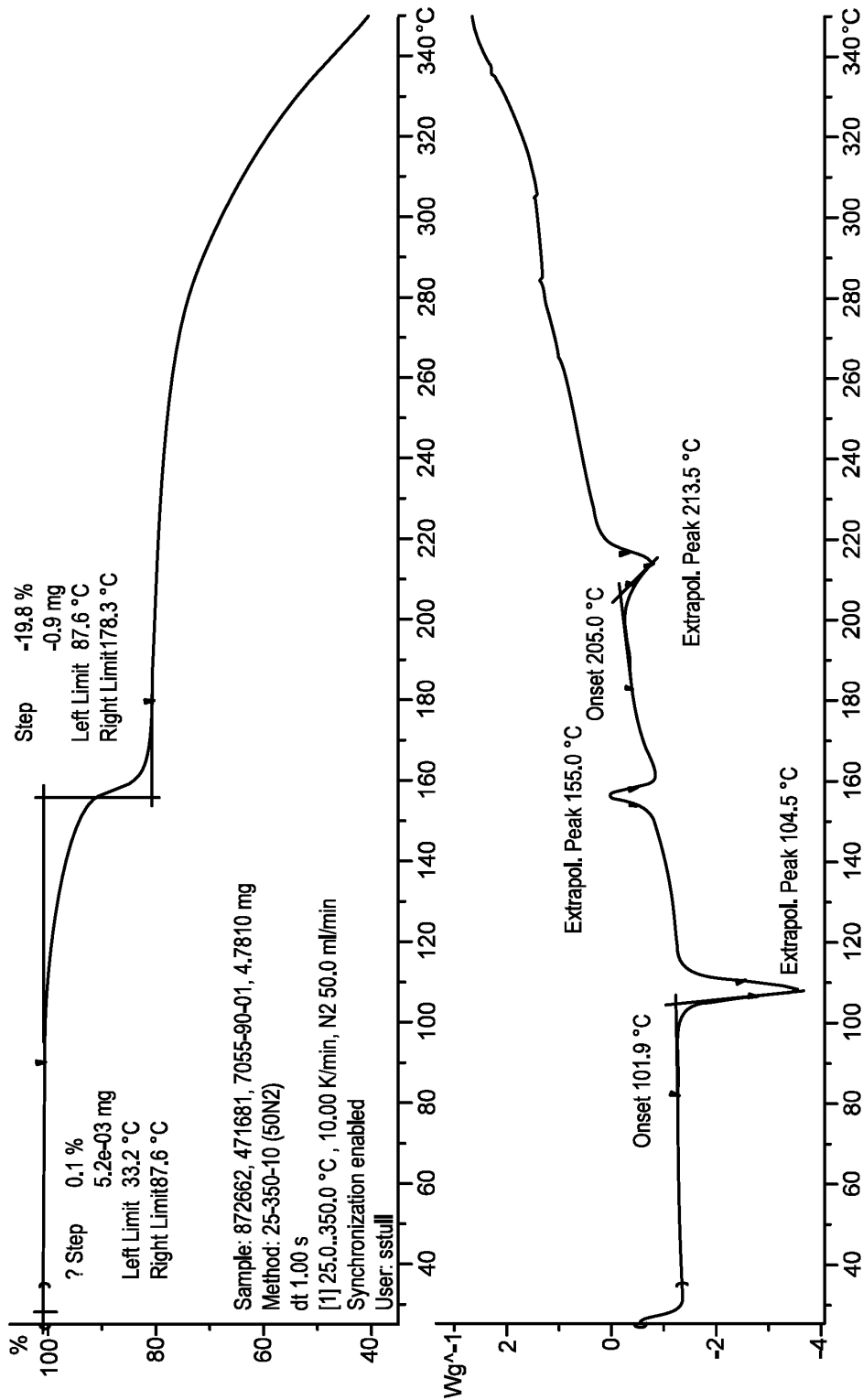


FIG. 15

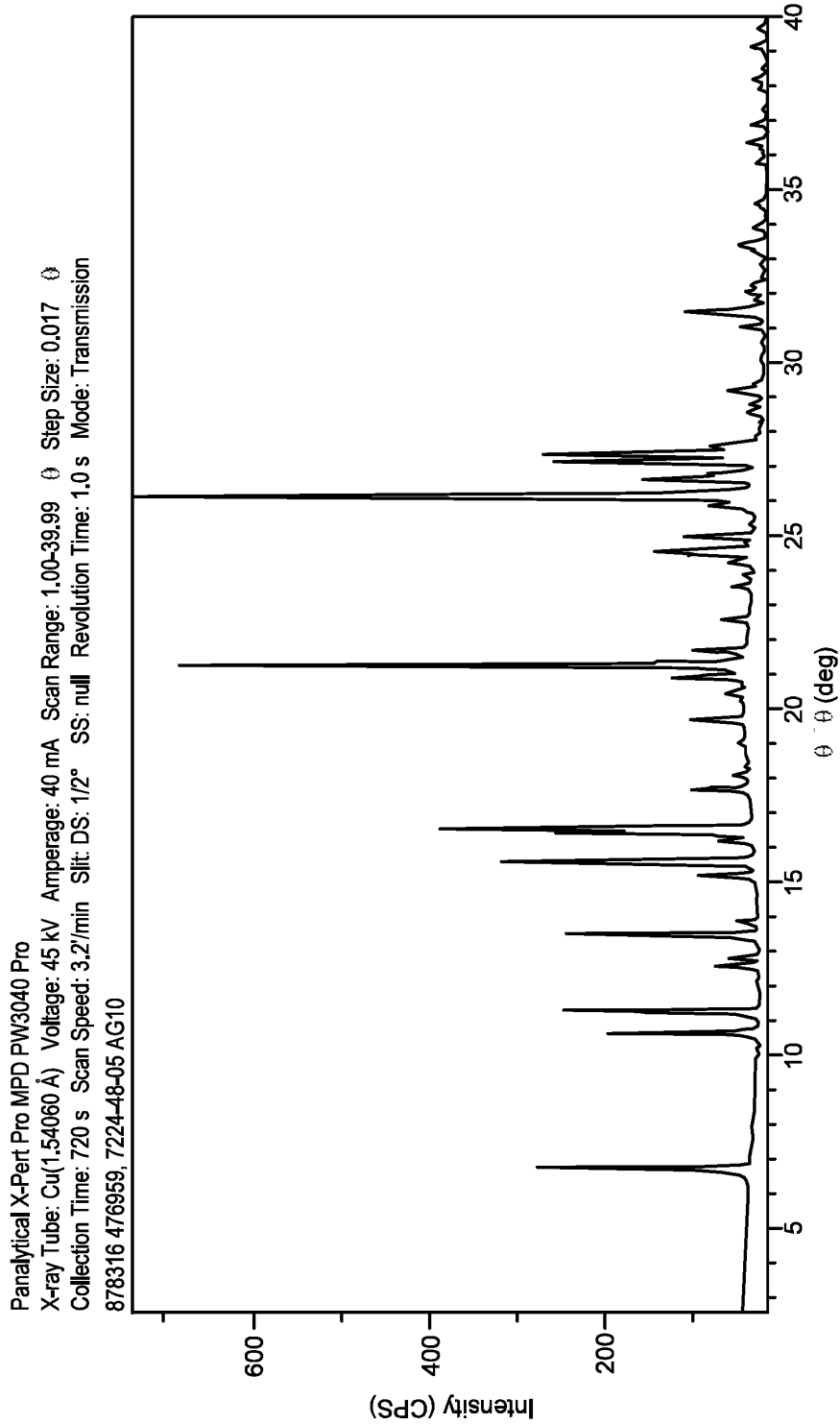


FIG. 16

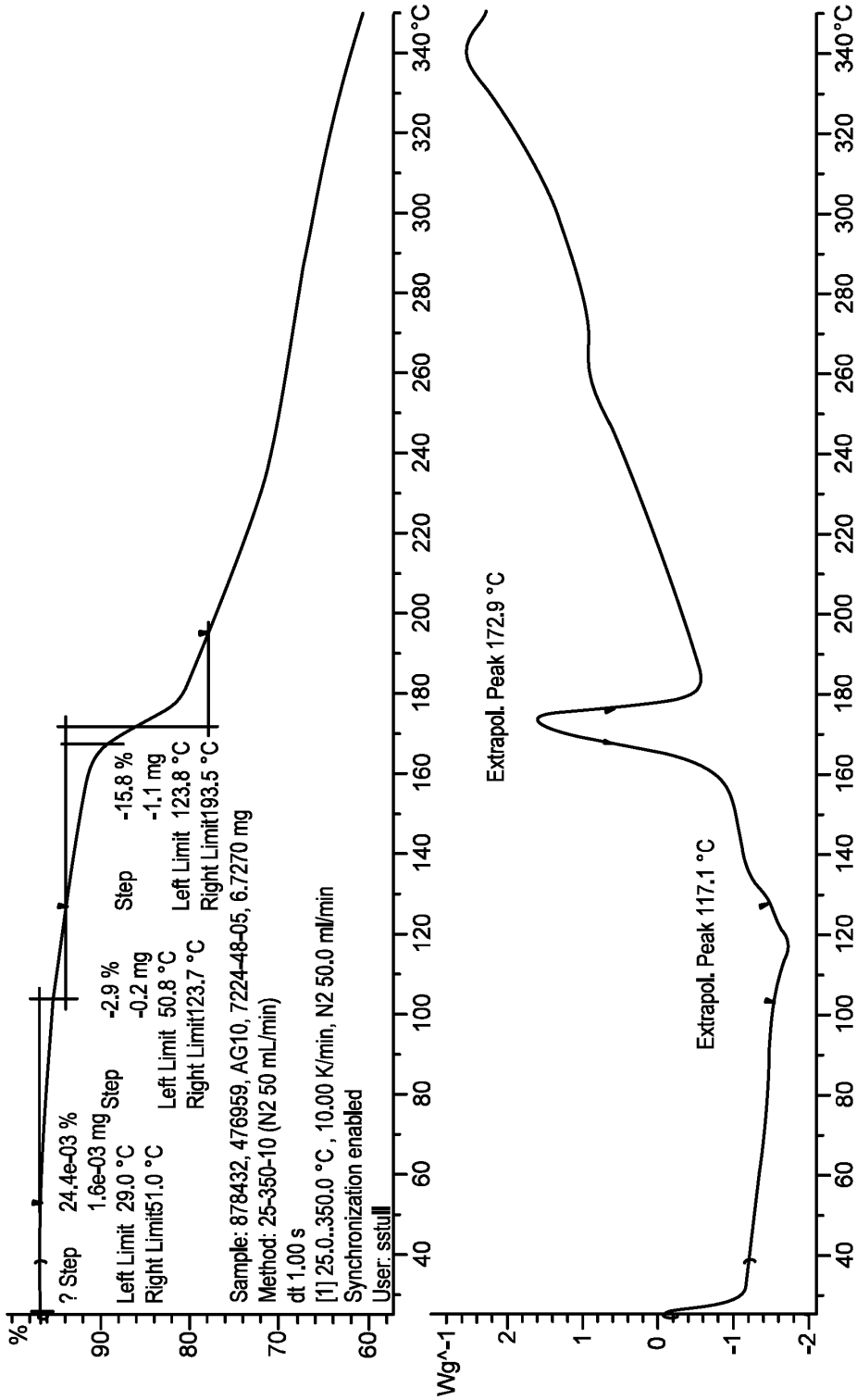
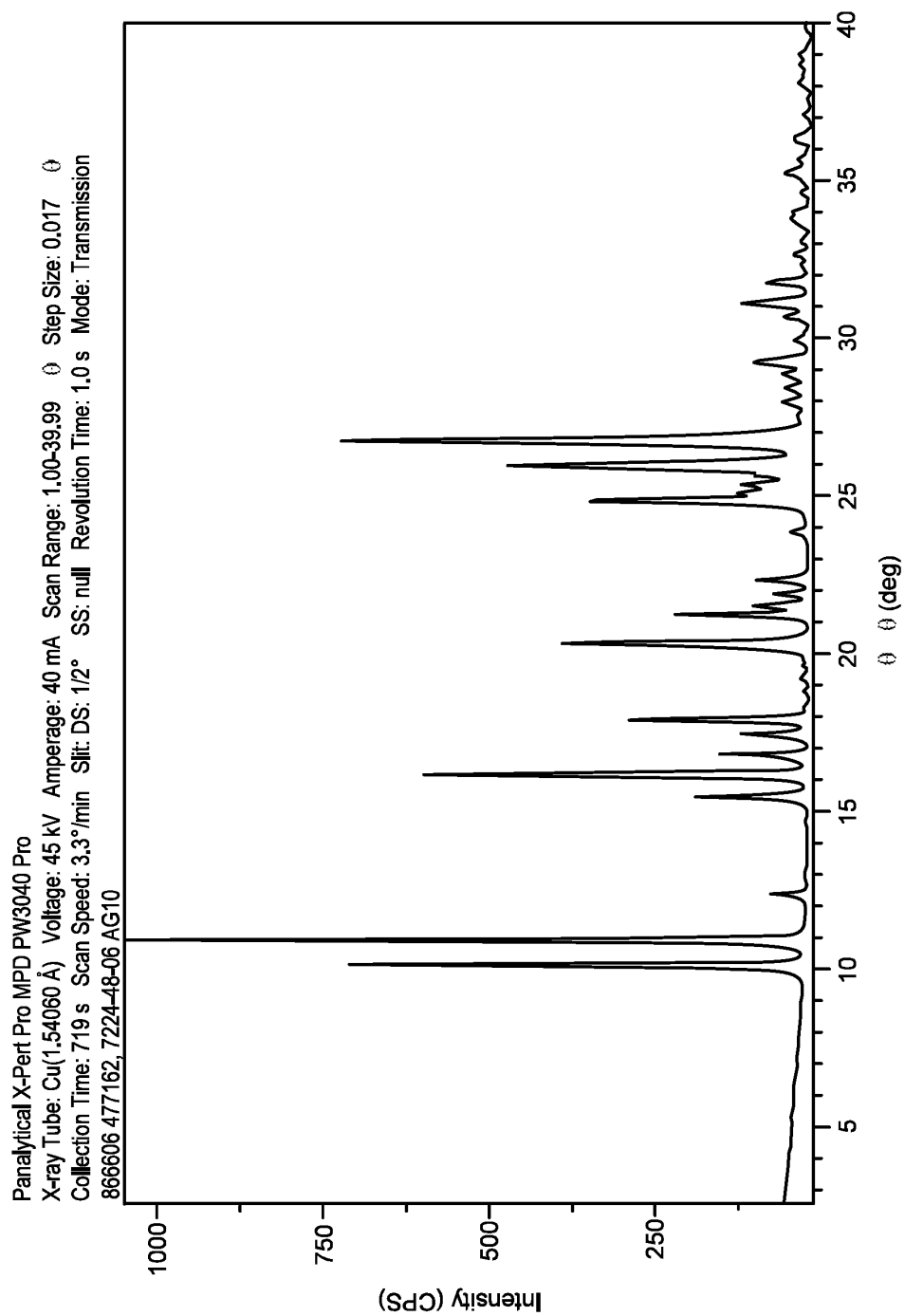


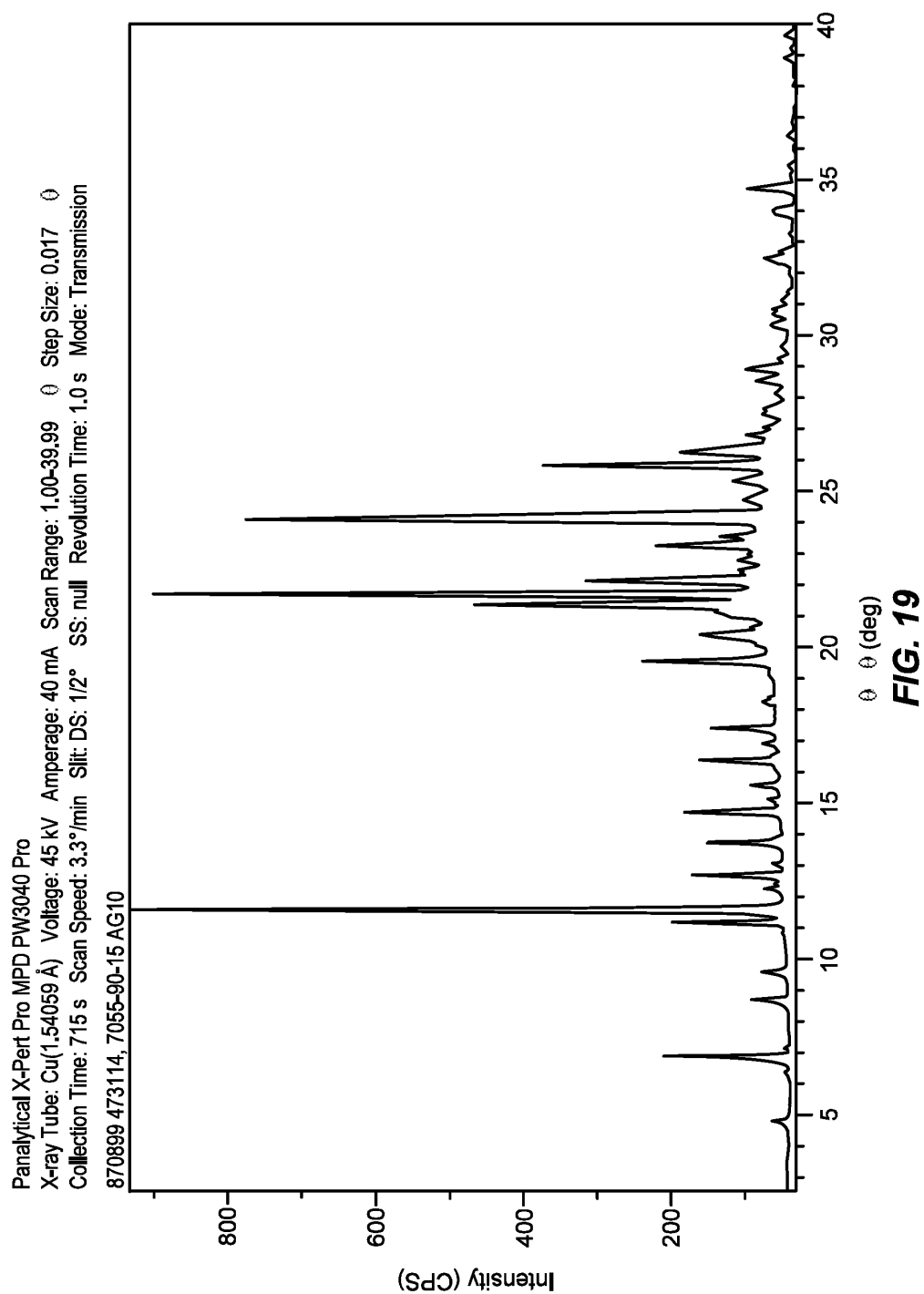
FIG. 17

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**FIG. 18**

SUBSTITUTE SHEET (RULE 26)

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SUBSTITUTE SHEET (RULE 26)

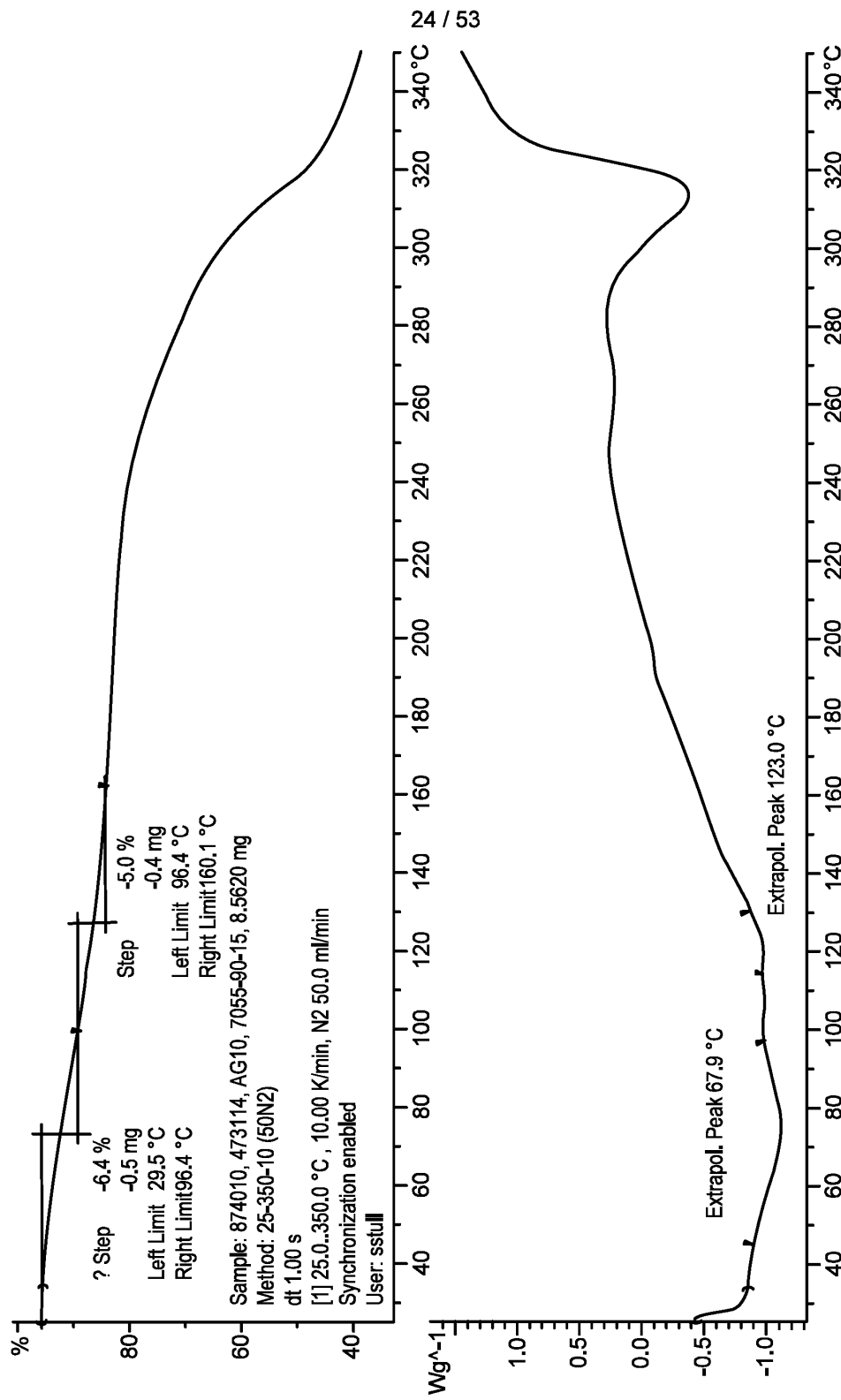


FIG. 20

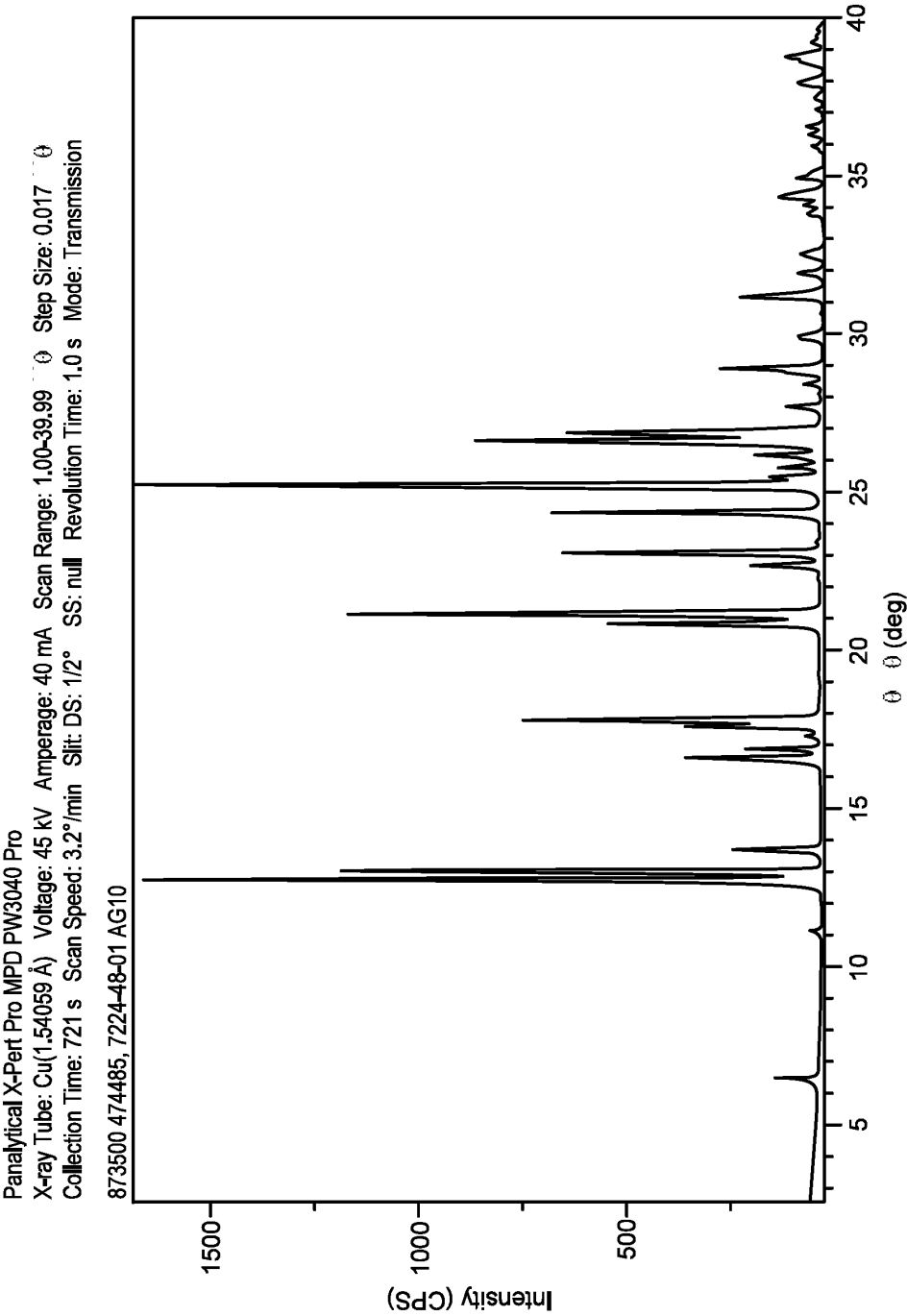


FIG. 21

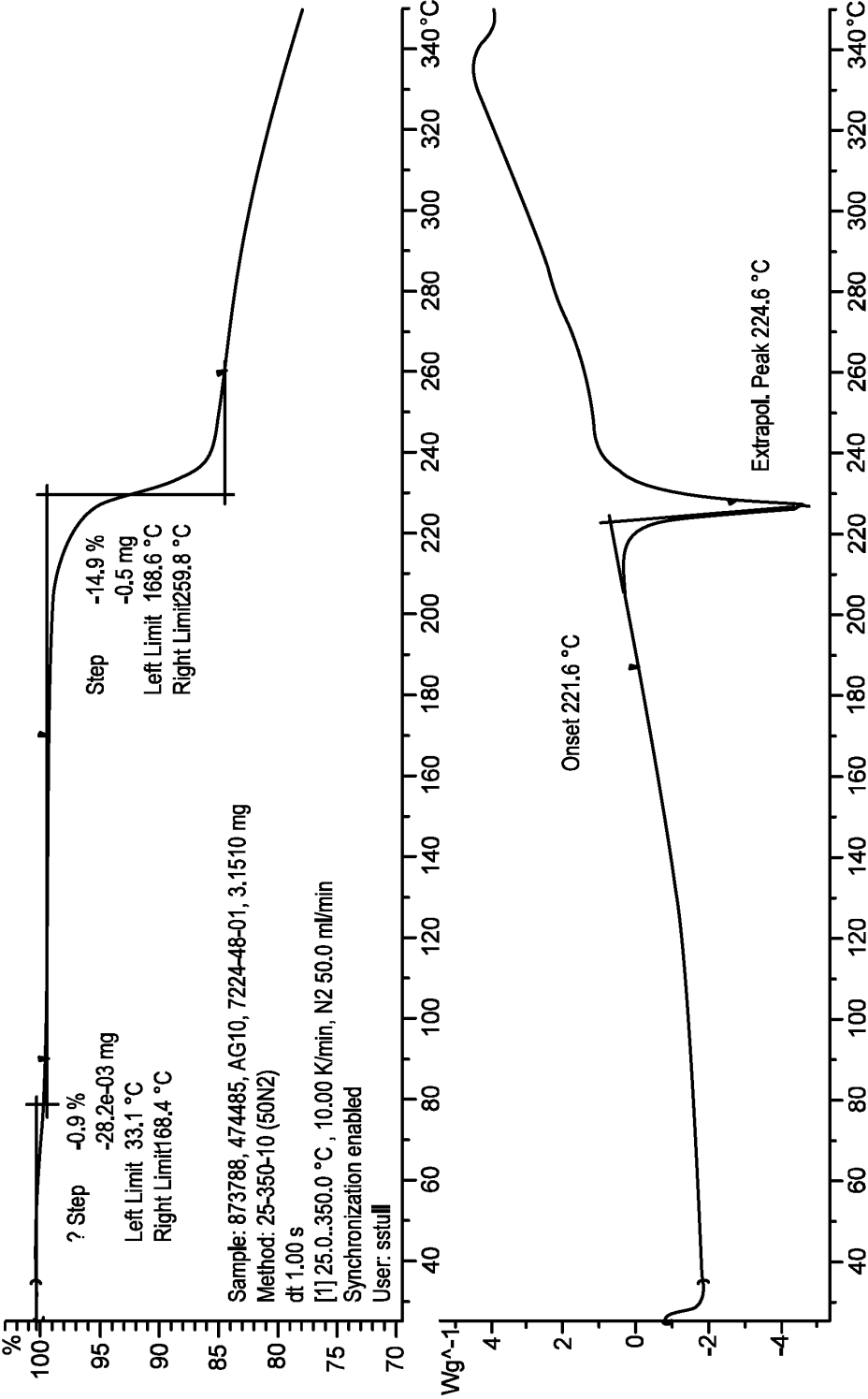
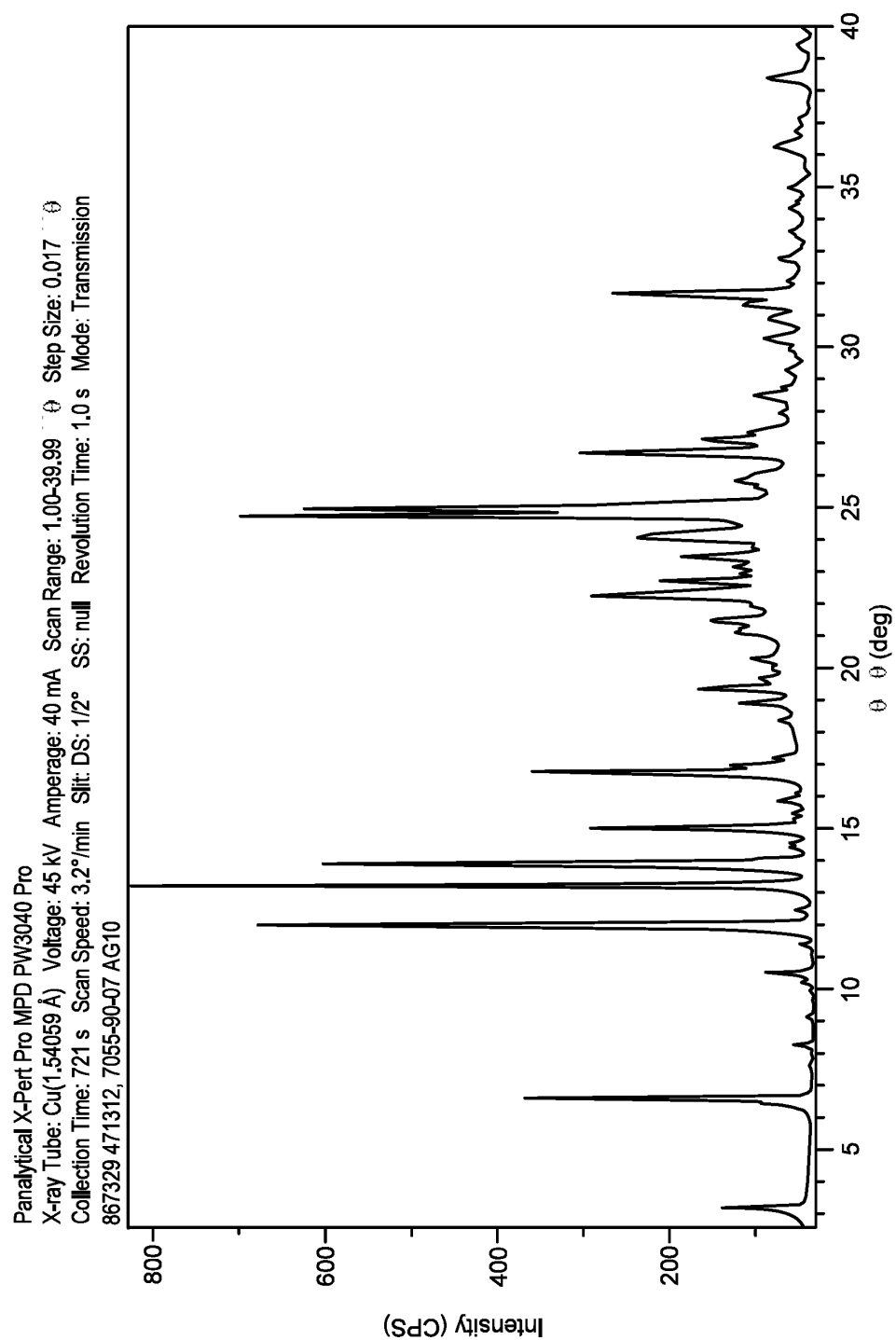


FIG. 22

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**FIG. 23**

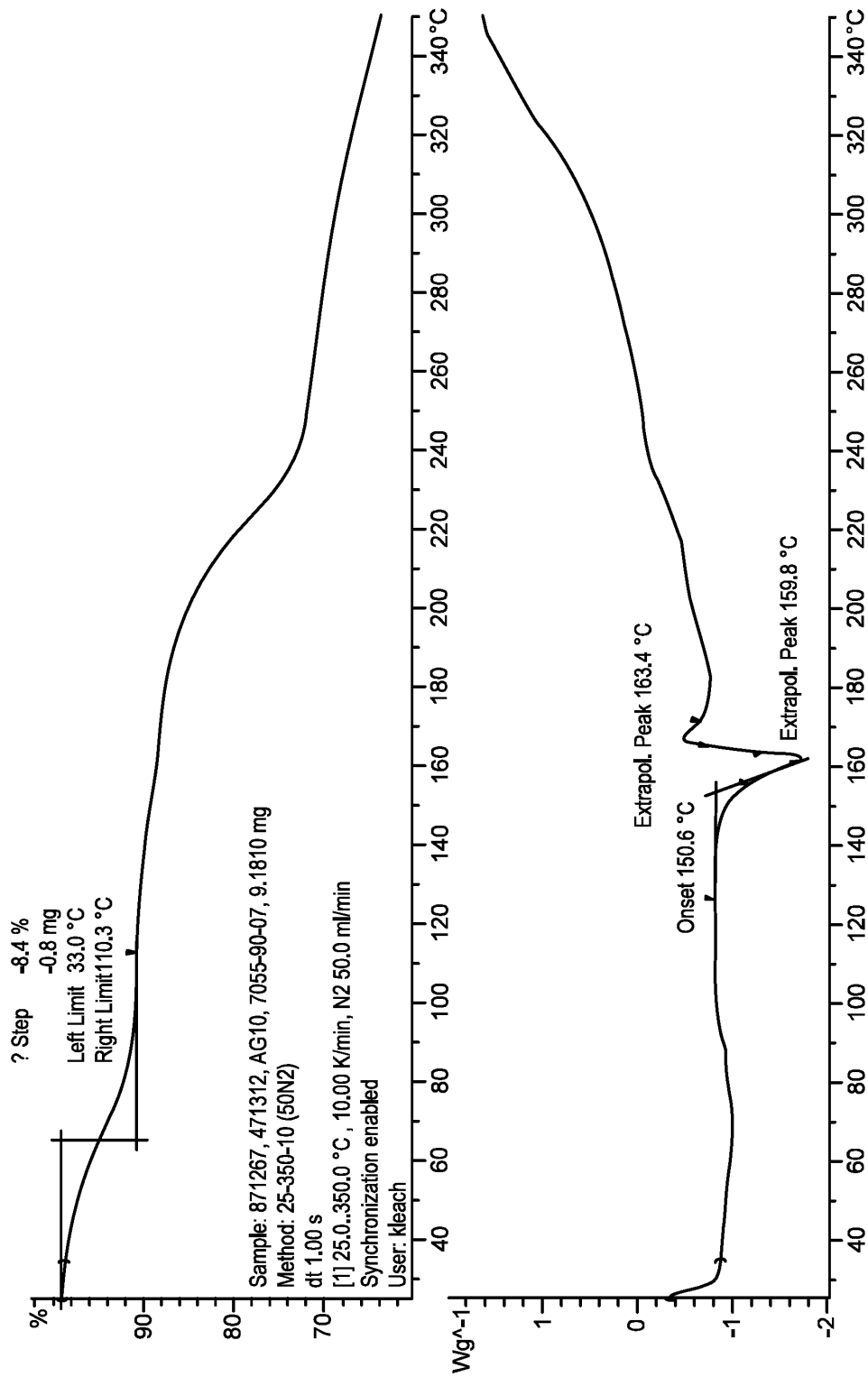
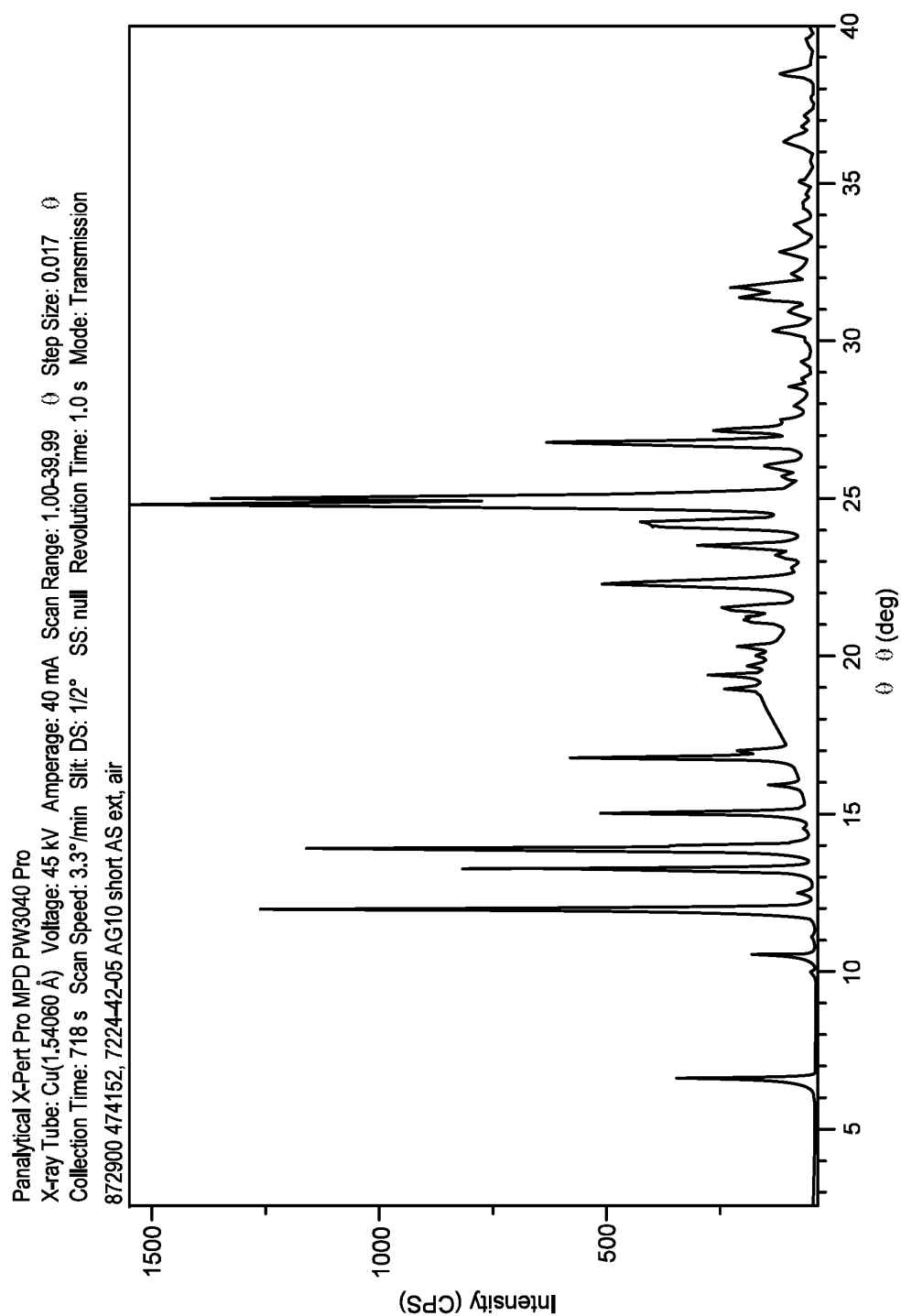


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**FIG. 25**

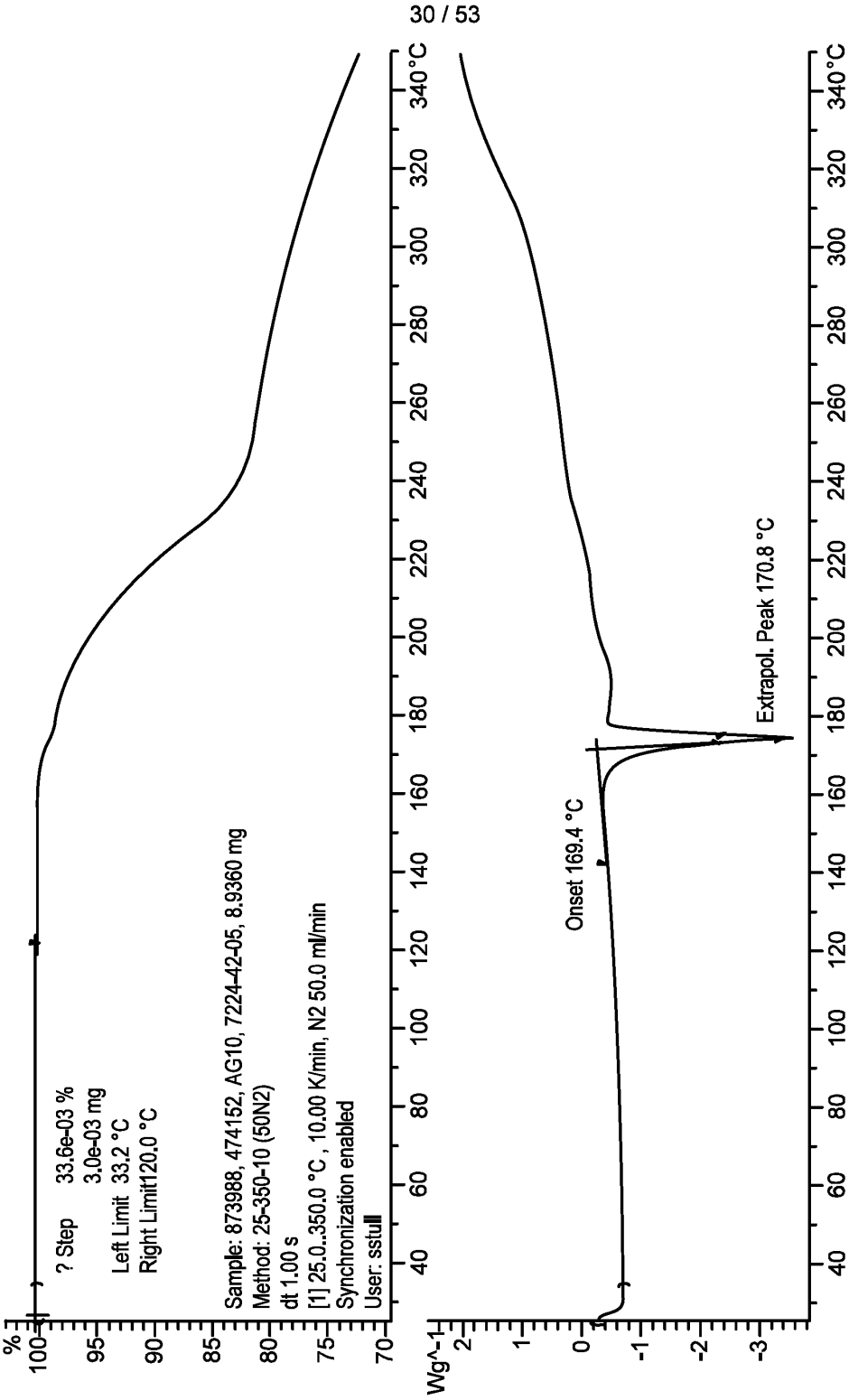
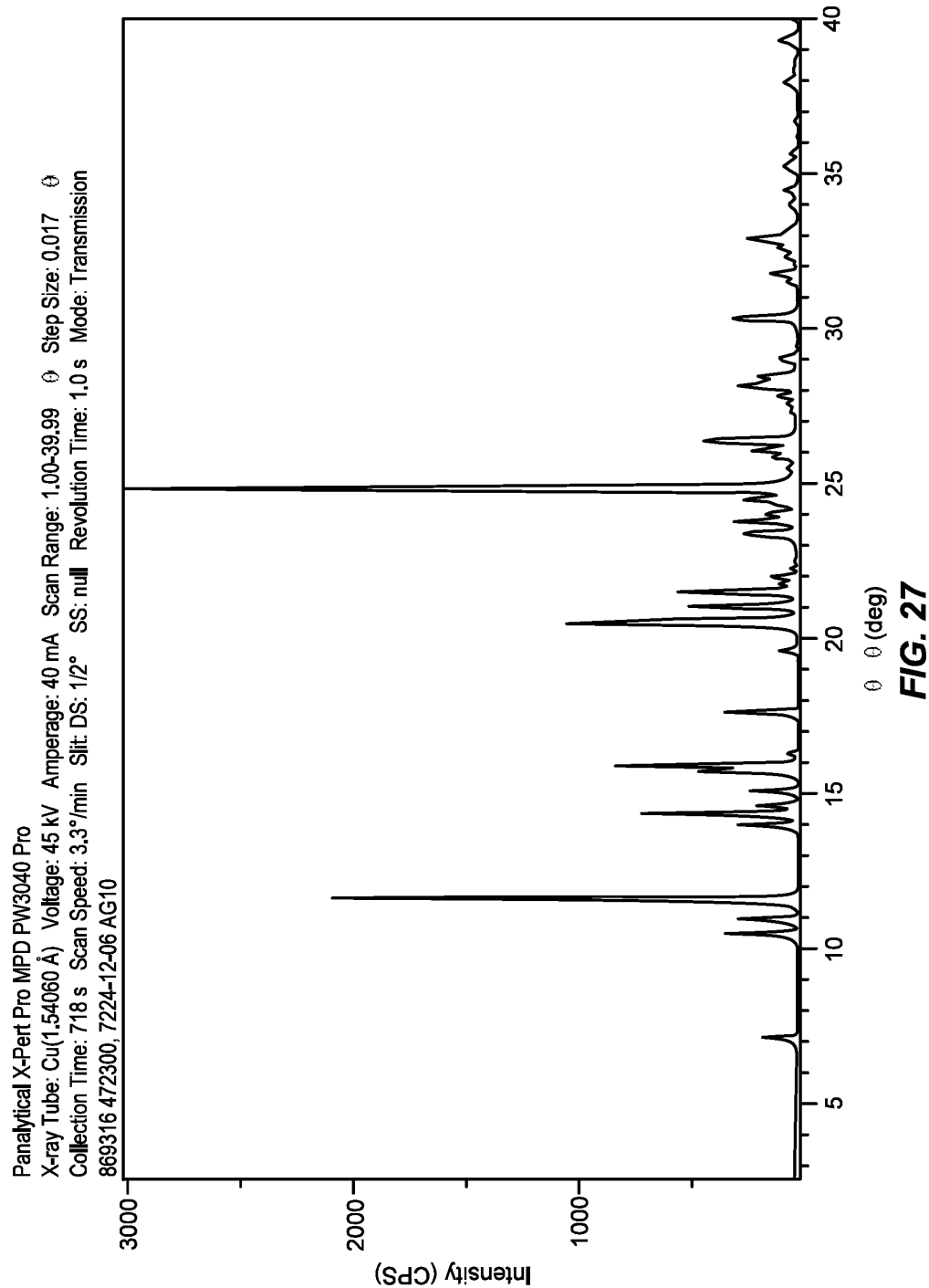


FIG. 26



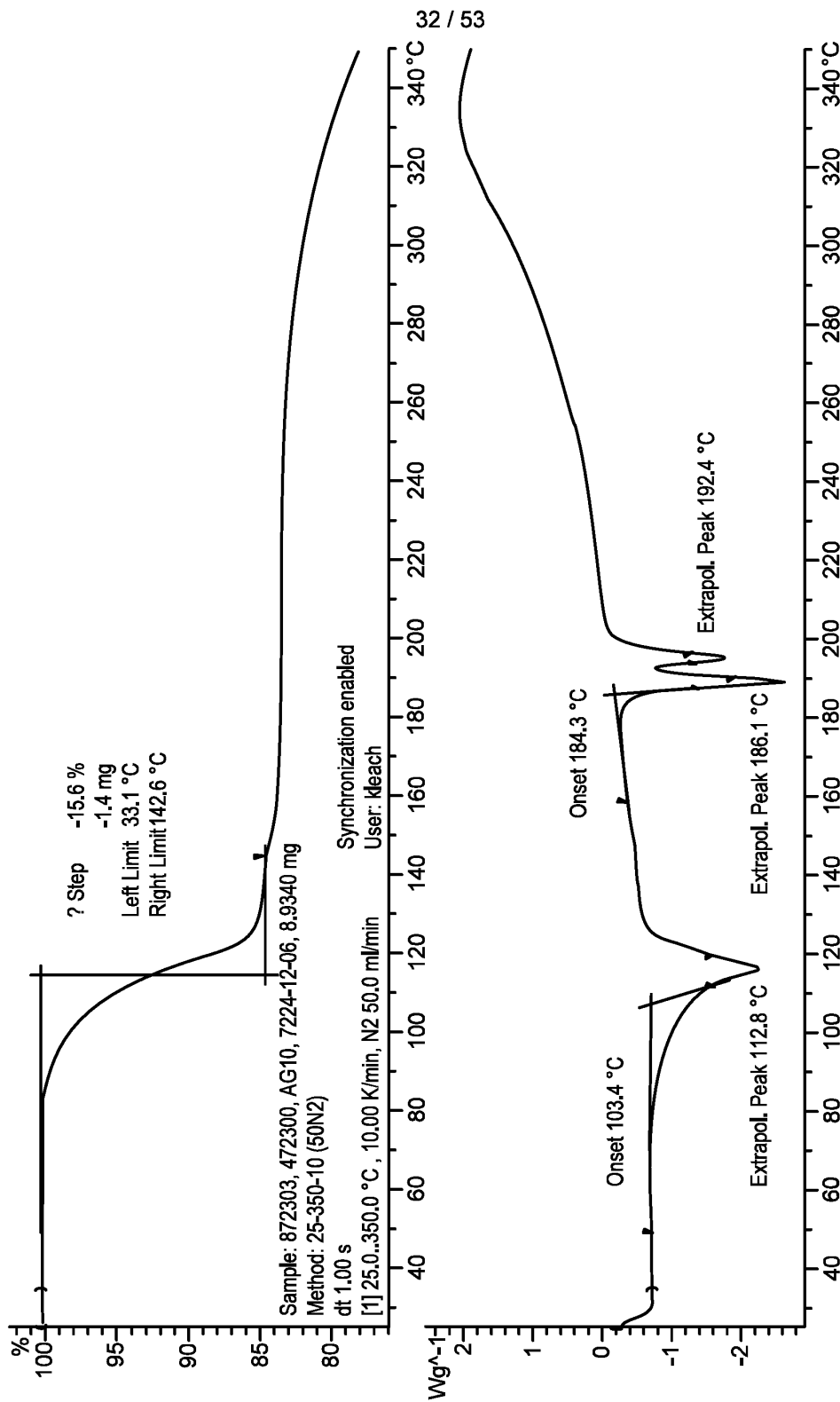
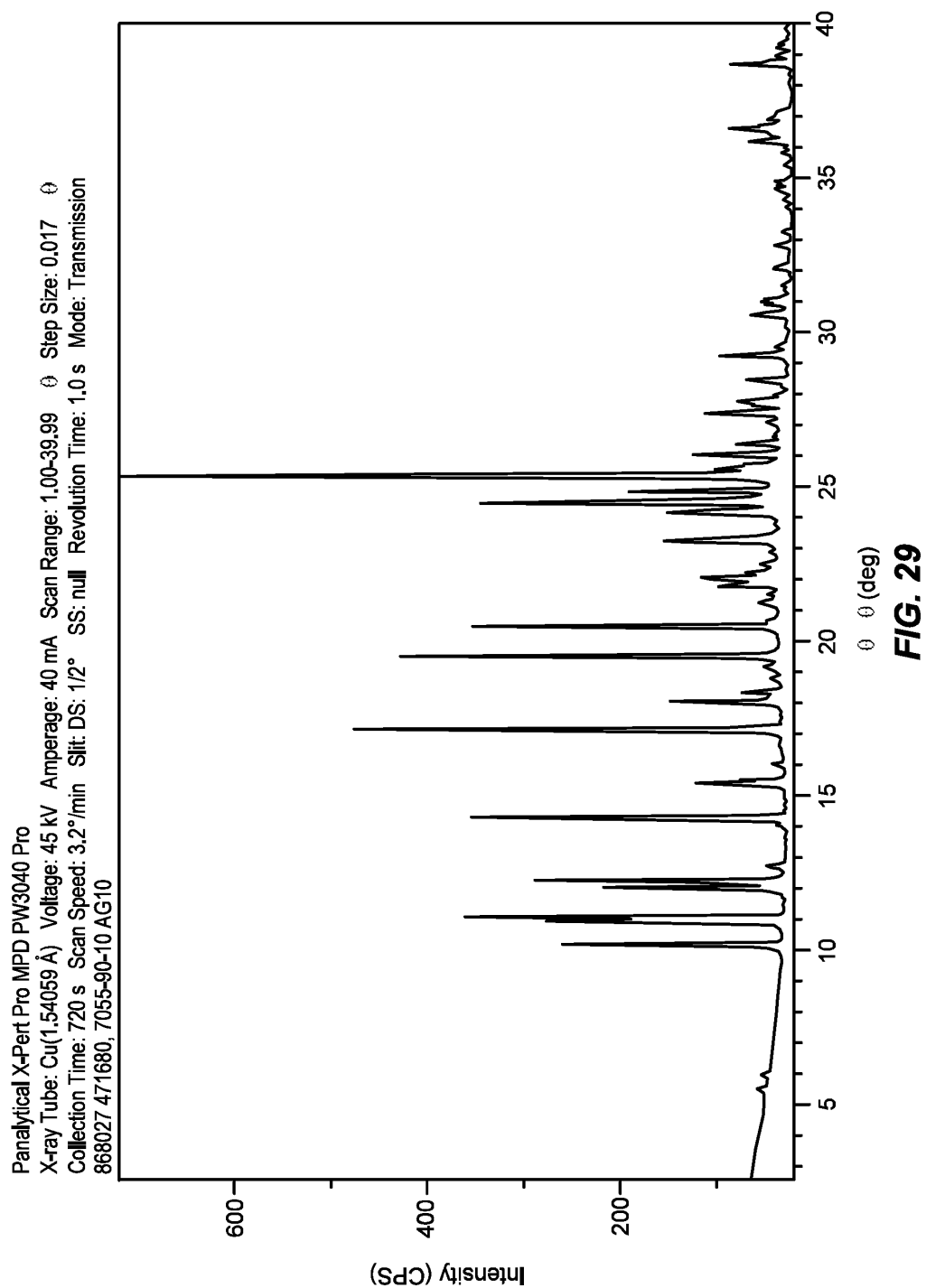


FIG. 28

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SUBSTITUTE SHEET (RULE 26)

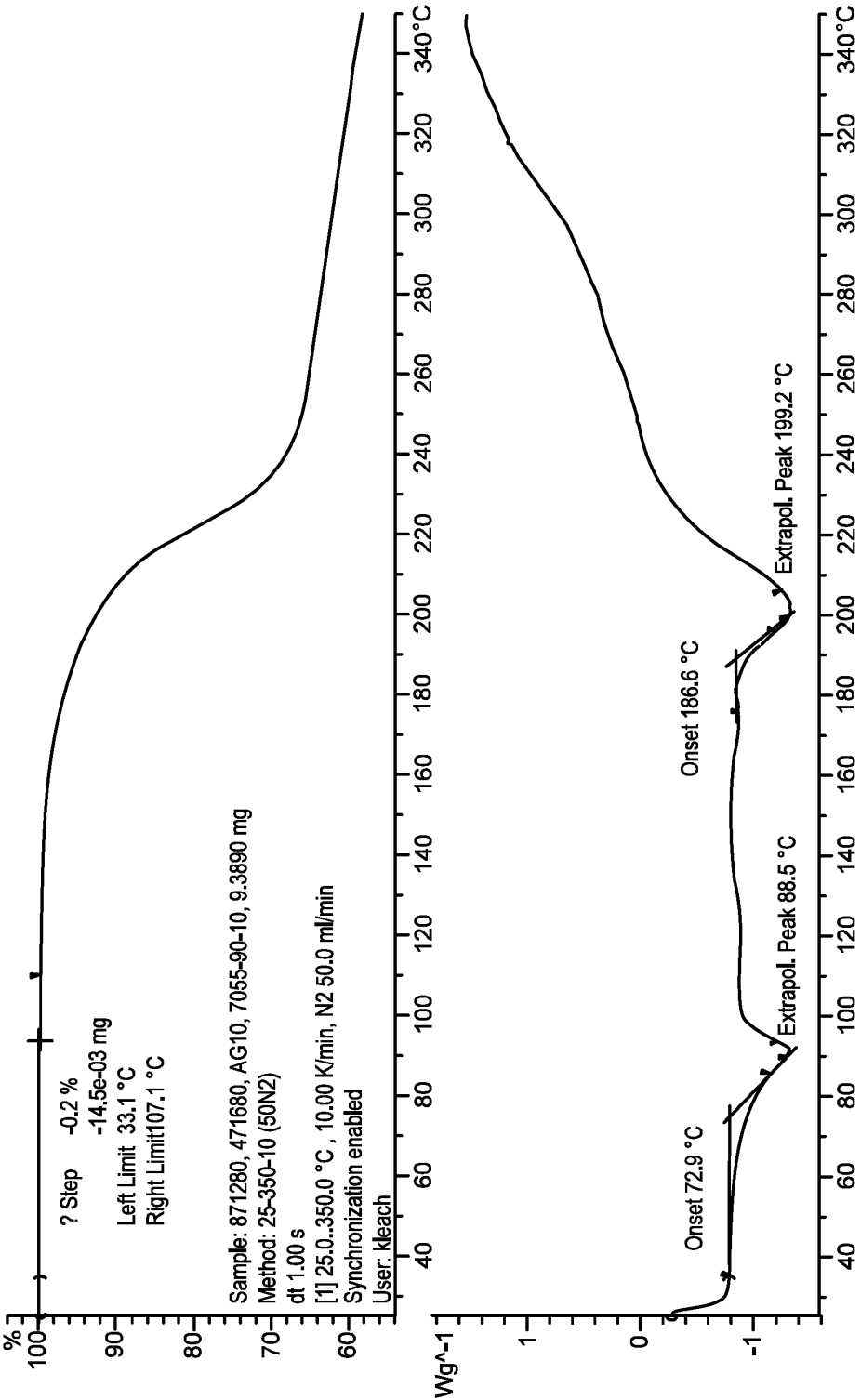
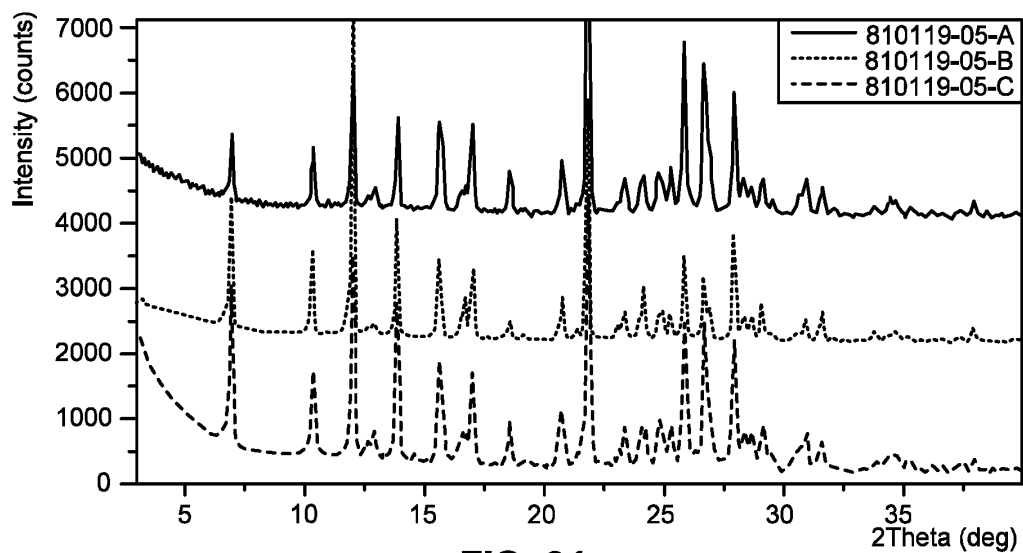
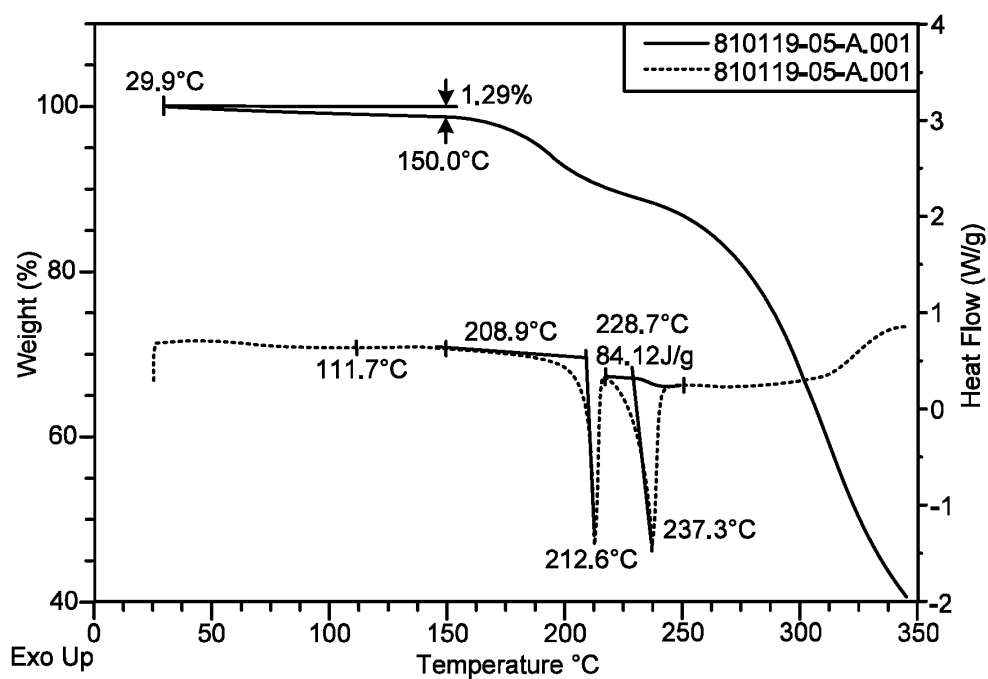
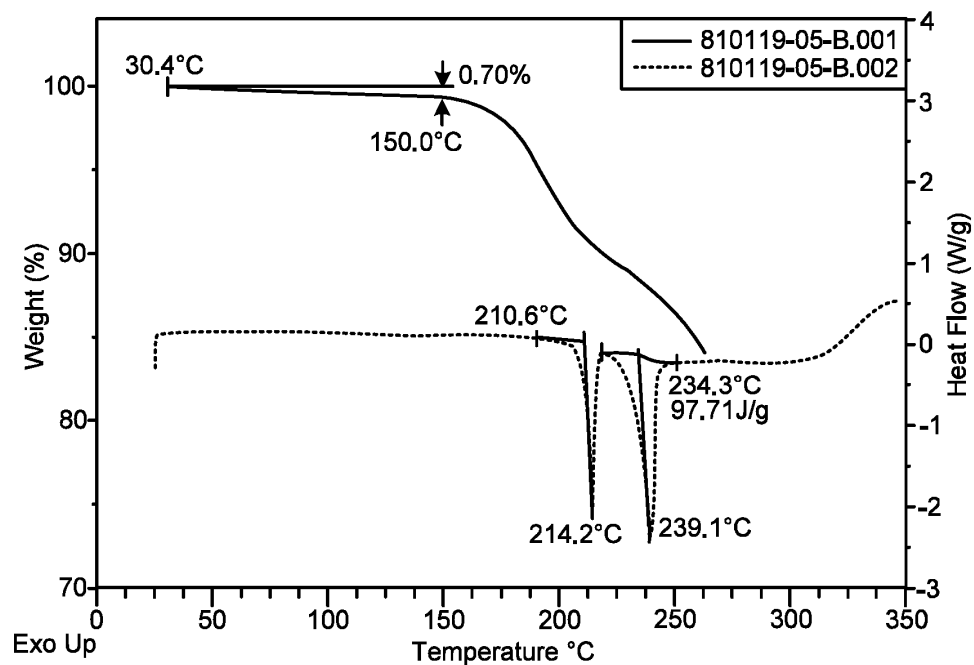
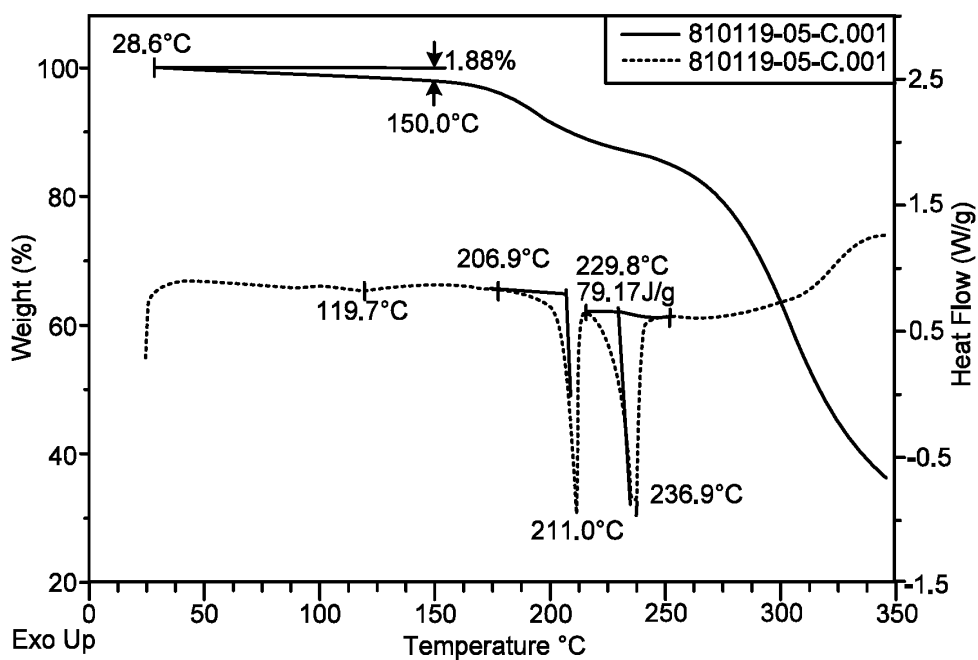


FIG. 30

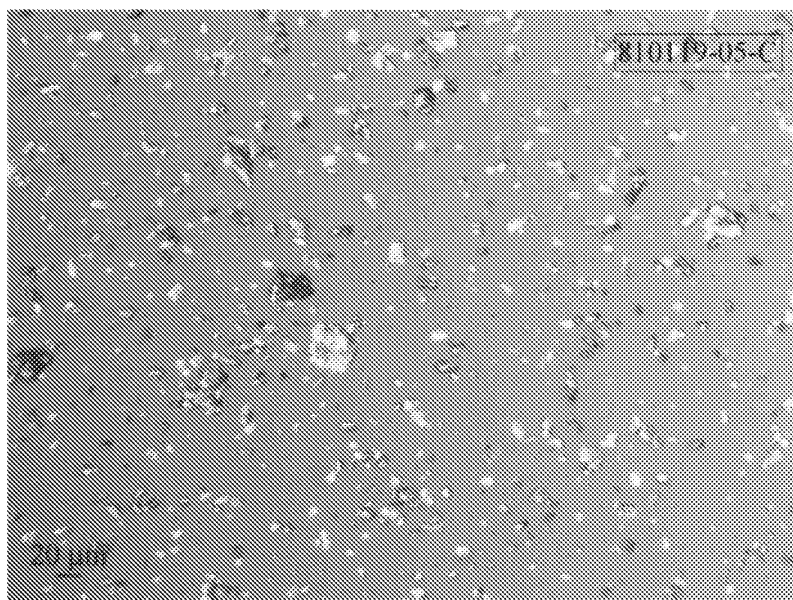
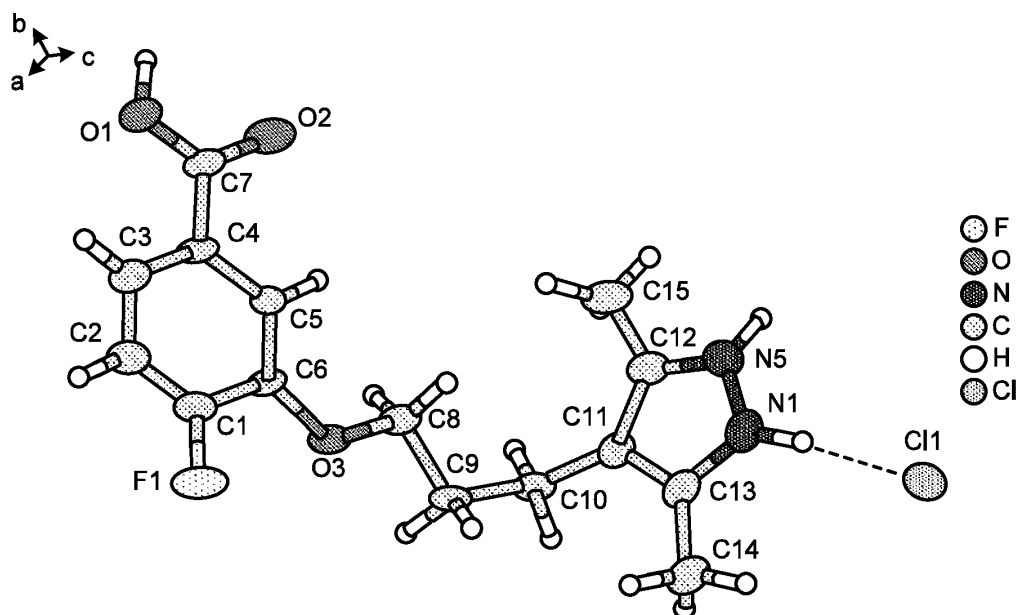
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**FIG. 31****FIG. 32**

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**FIG. 33****FIG. 34**

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**FIG. 35****FIG. 36**

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DVS Isotherm Plot

Date: 16 Mar 2017

Time: 13-31

File: 810119-05-A - Thu 16 Mar 2017 13-31-27.xls

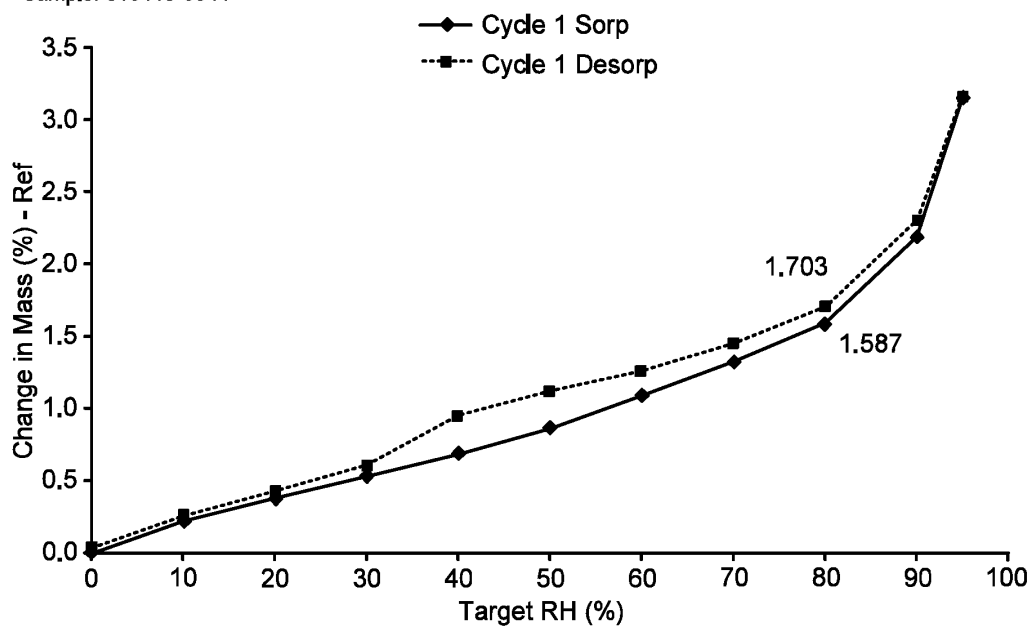
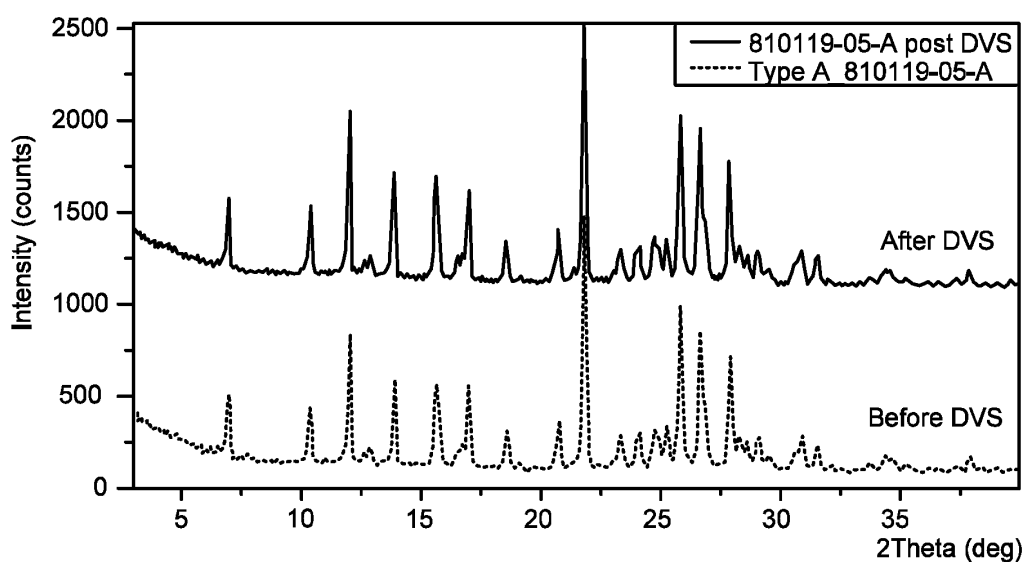
Sample: 810119-05-A

Temp: 25.0 °C

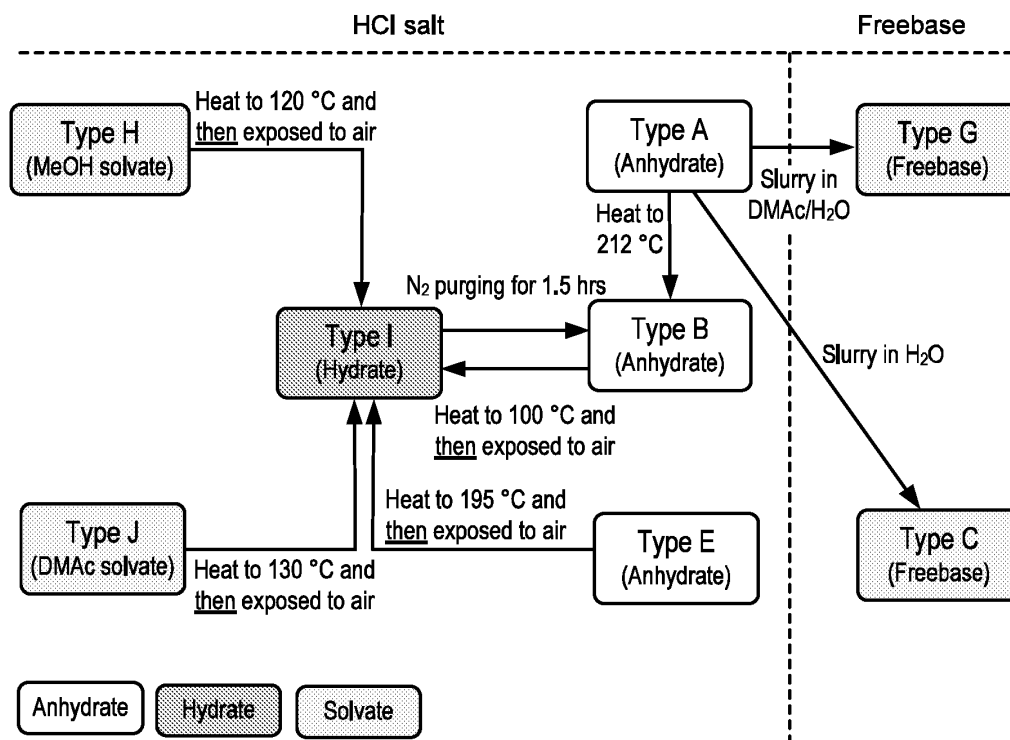
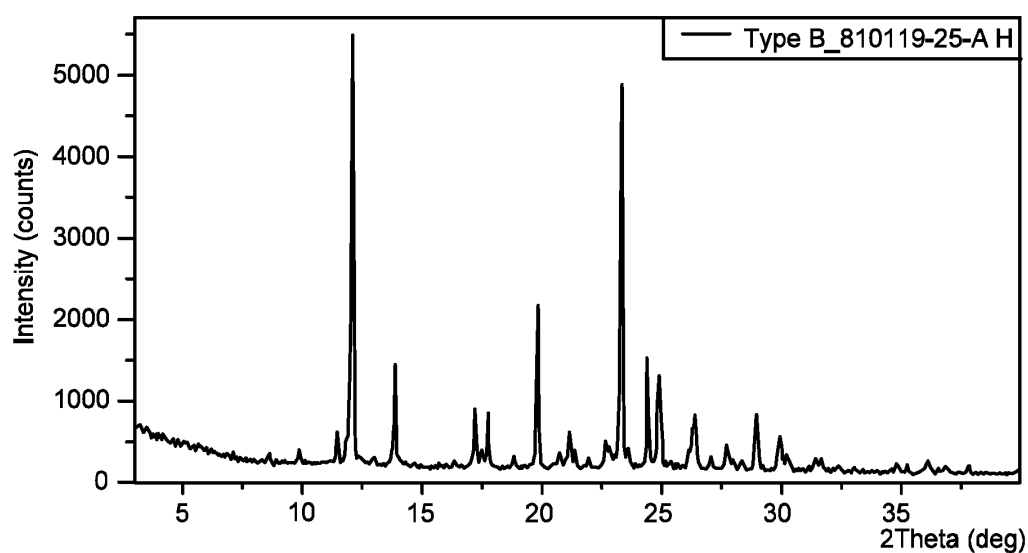
Meth: 0-95-0 CP3677 LONG

METHOD.sao

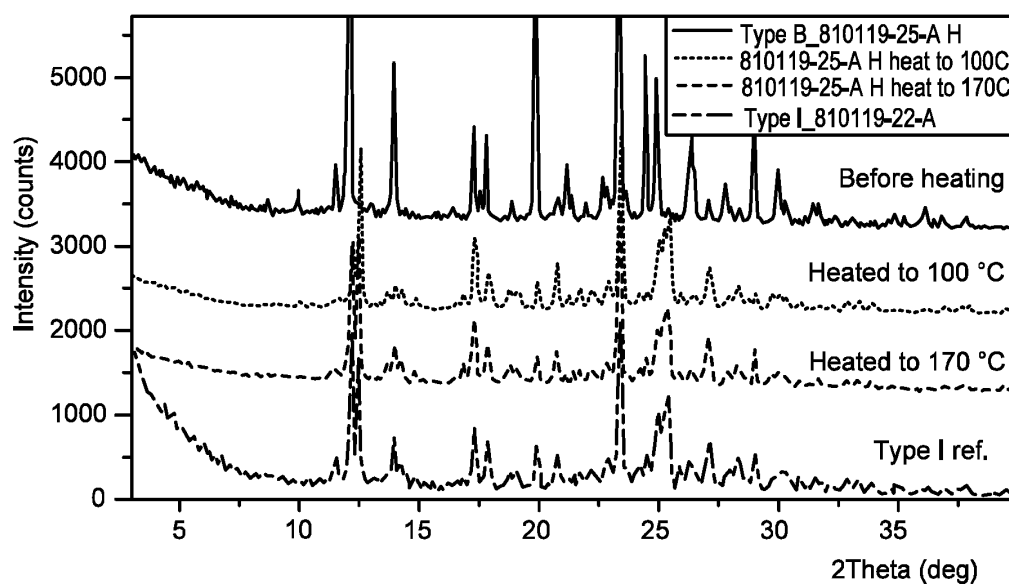
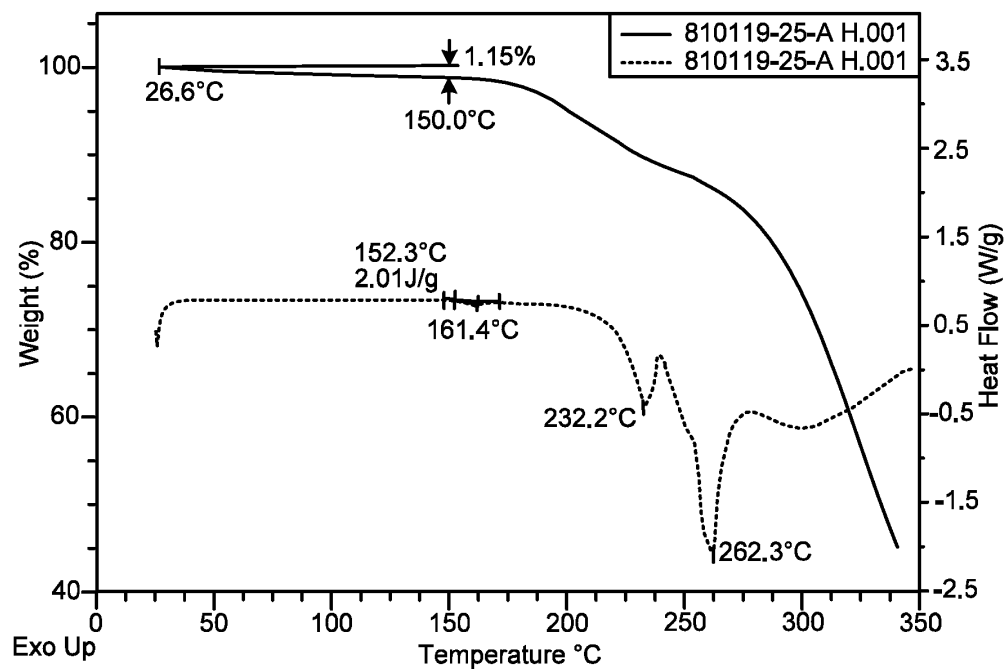
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**FIG. 37****FIG. 38**

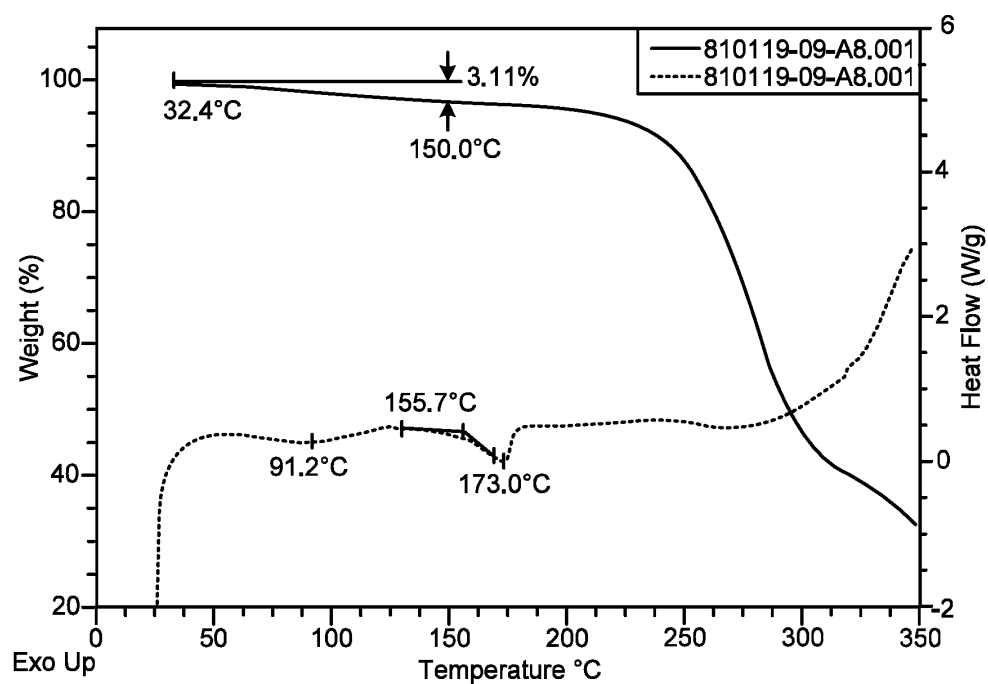
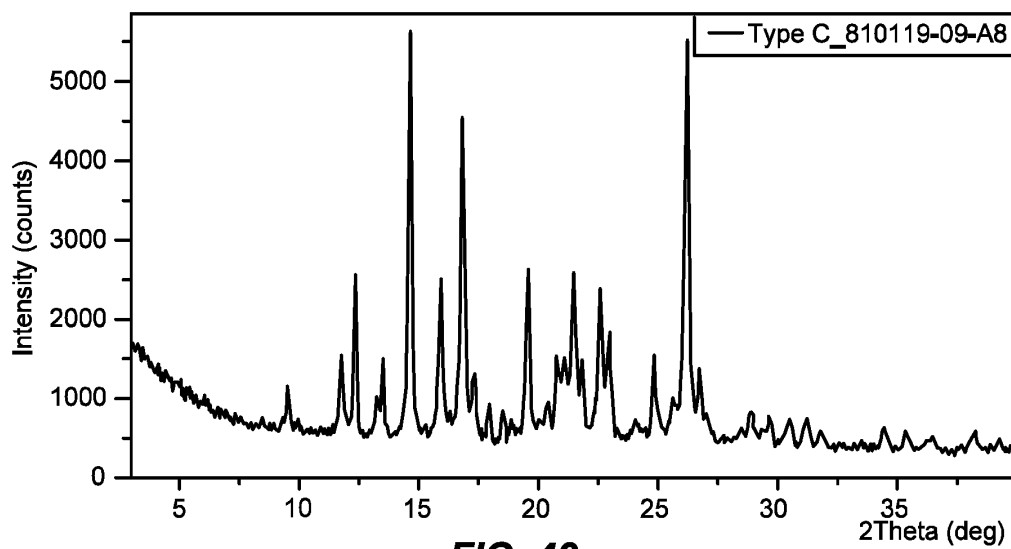
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**FIG. 39****FIG. 40**

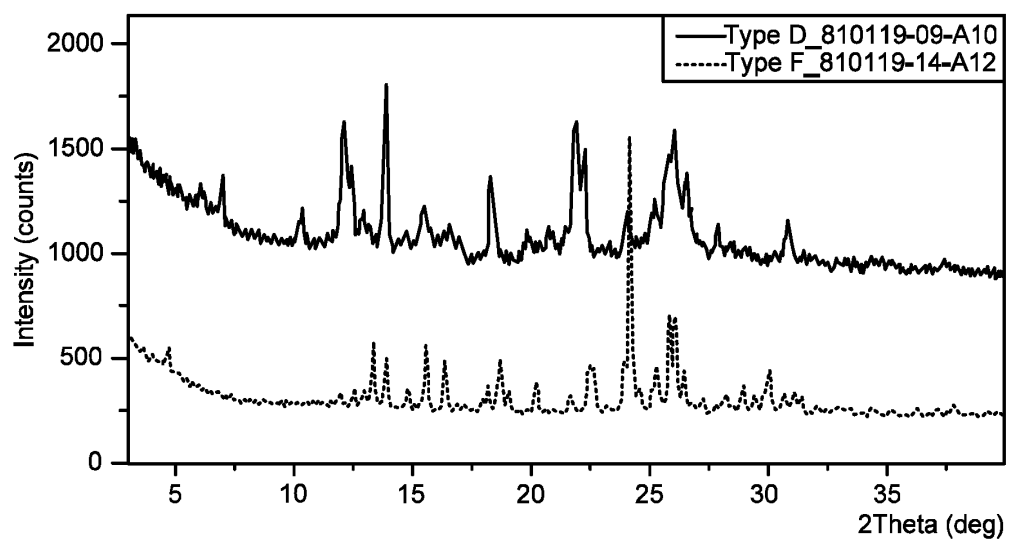
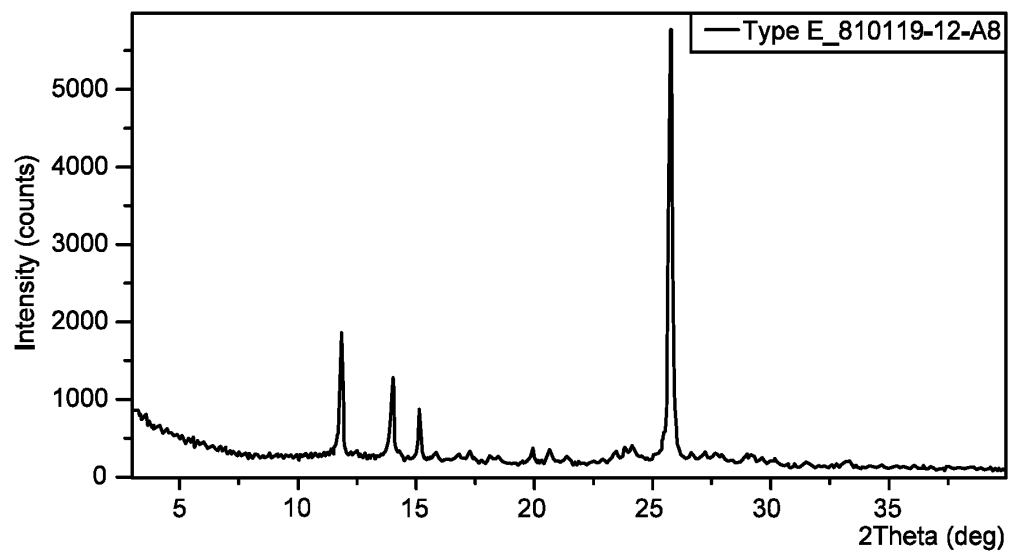
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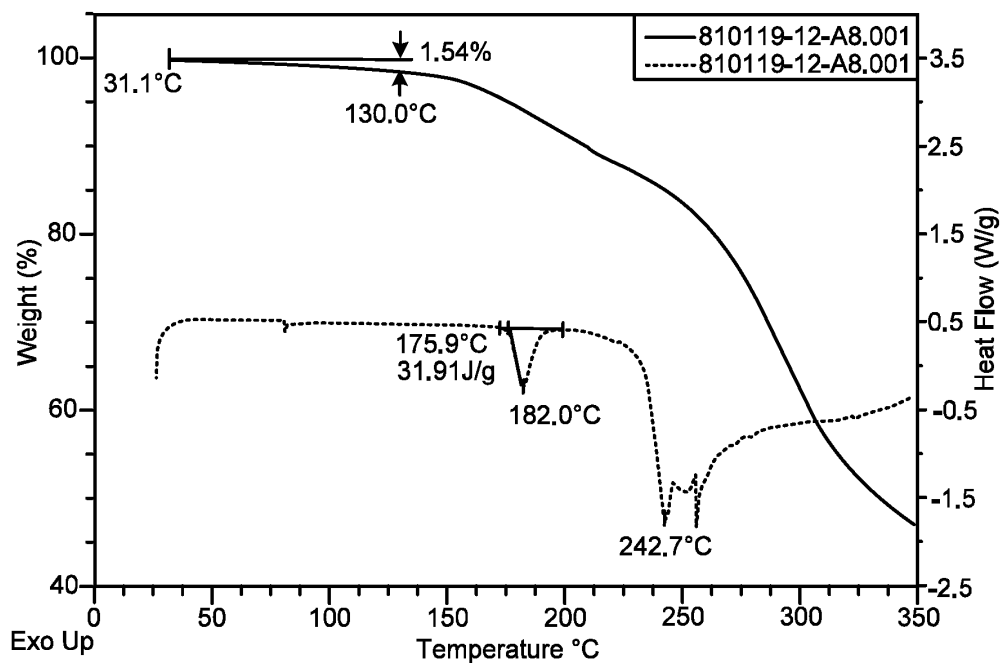
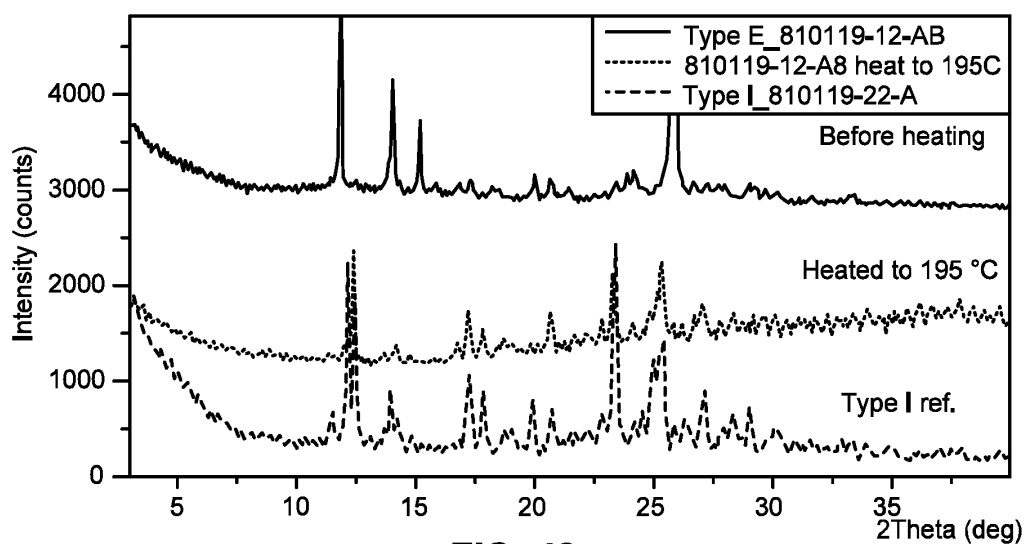
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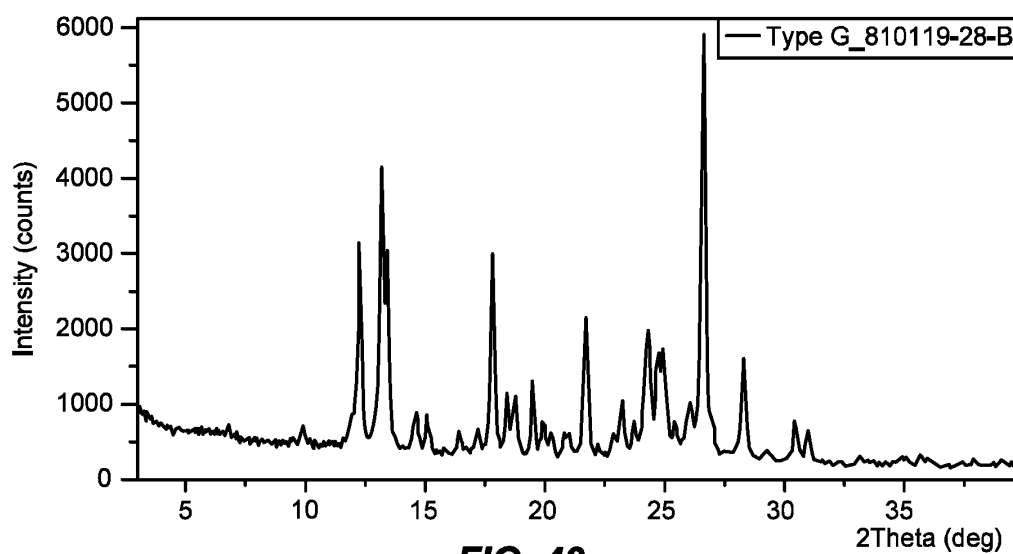
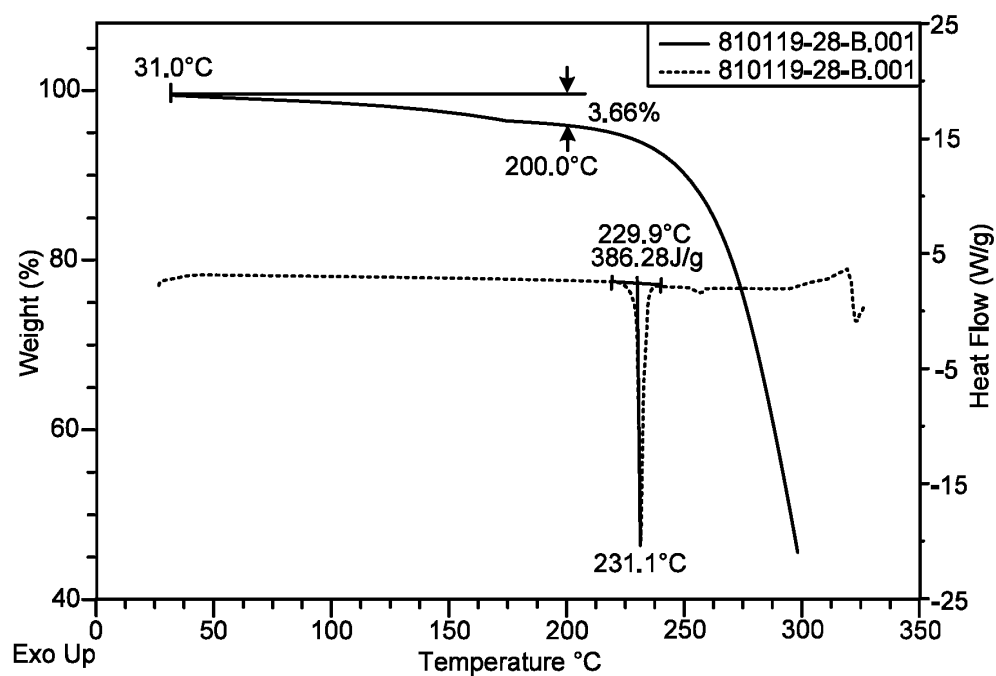
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**FIG. 45****FIG. 46**

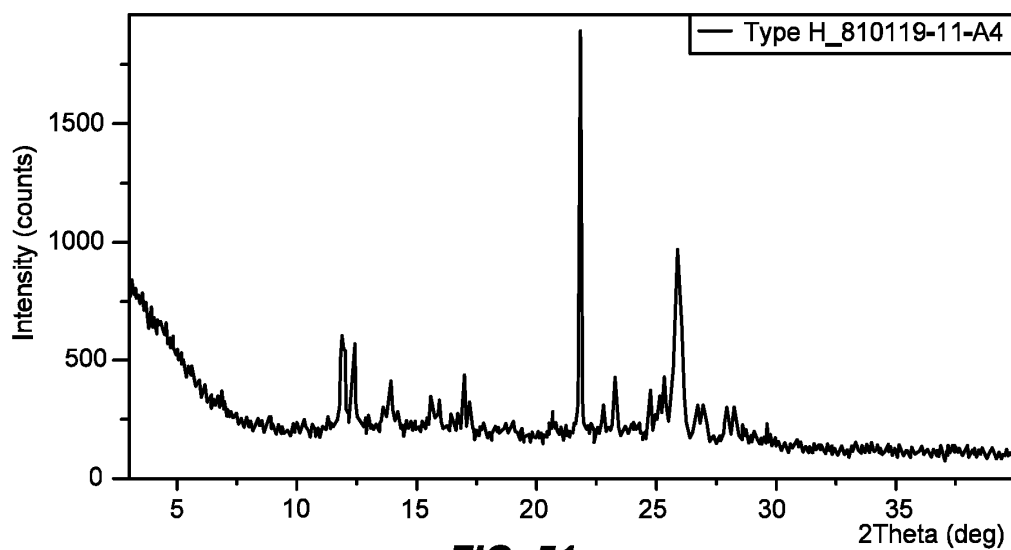
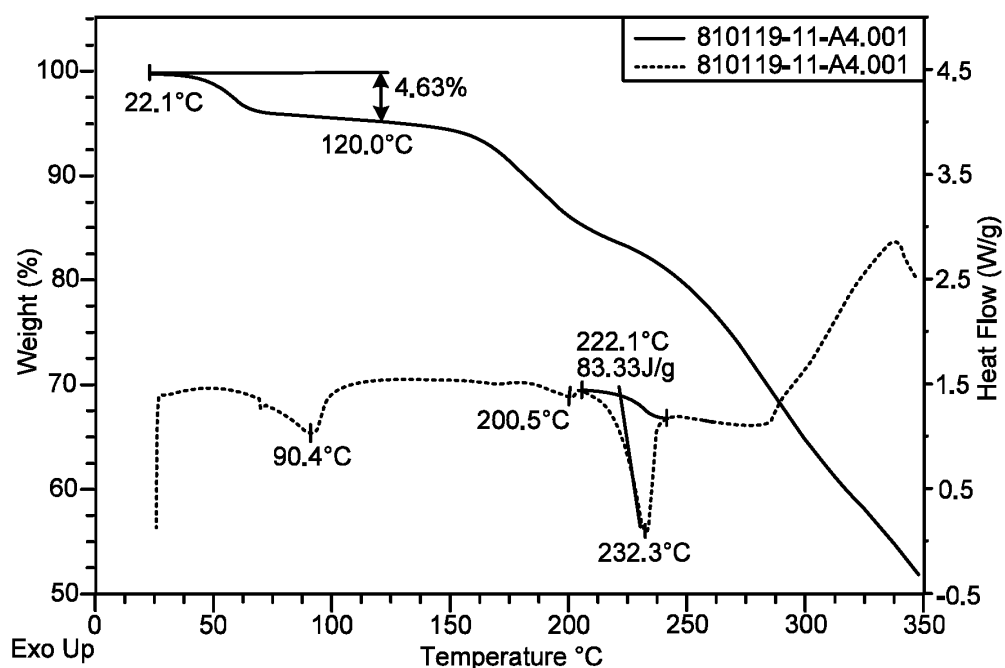
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**FIG. 47****FIG. 48**

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**FIG. 49****FIG. 50**

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**FIG. 51****FIG. 52**

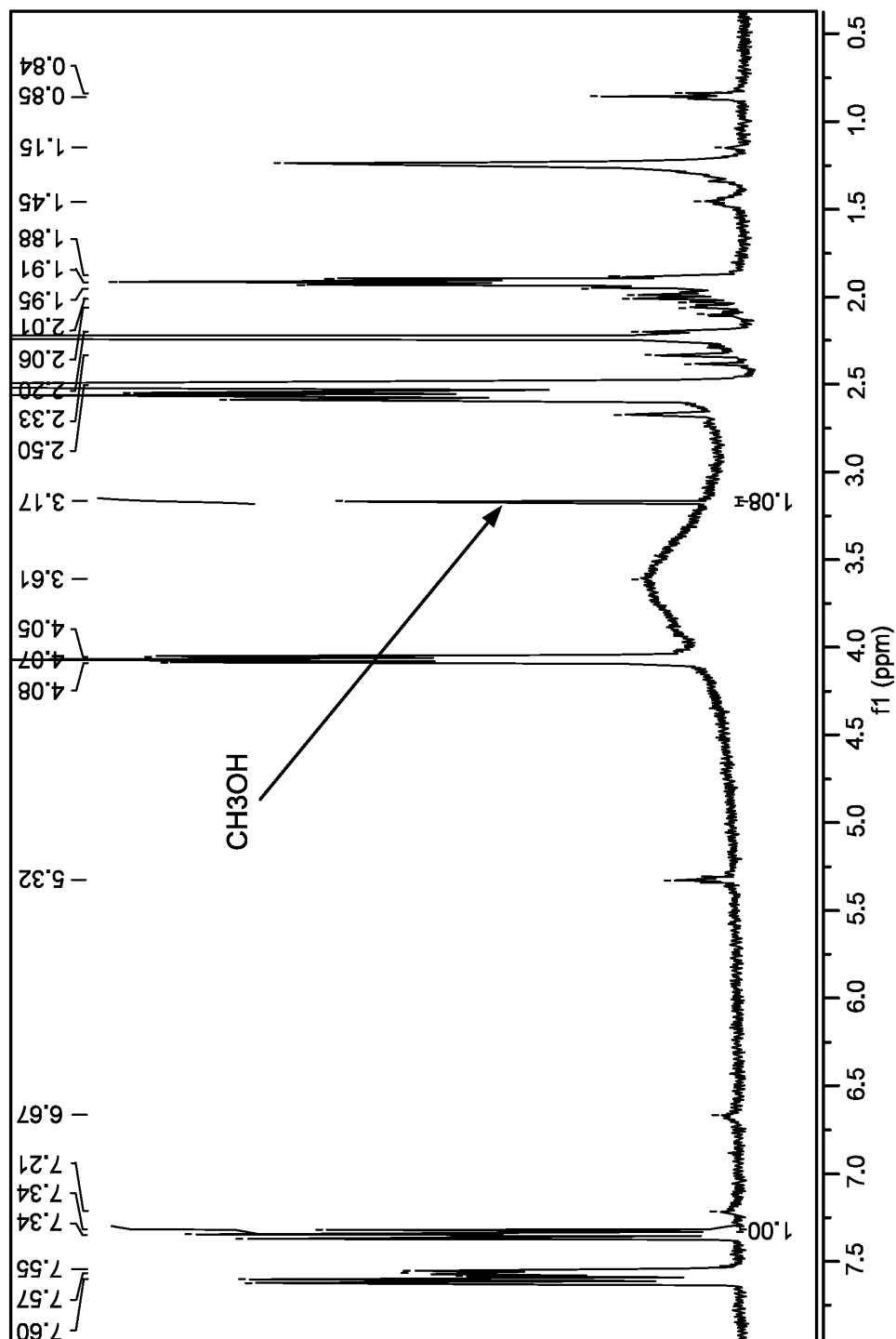


FIG. 53

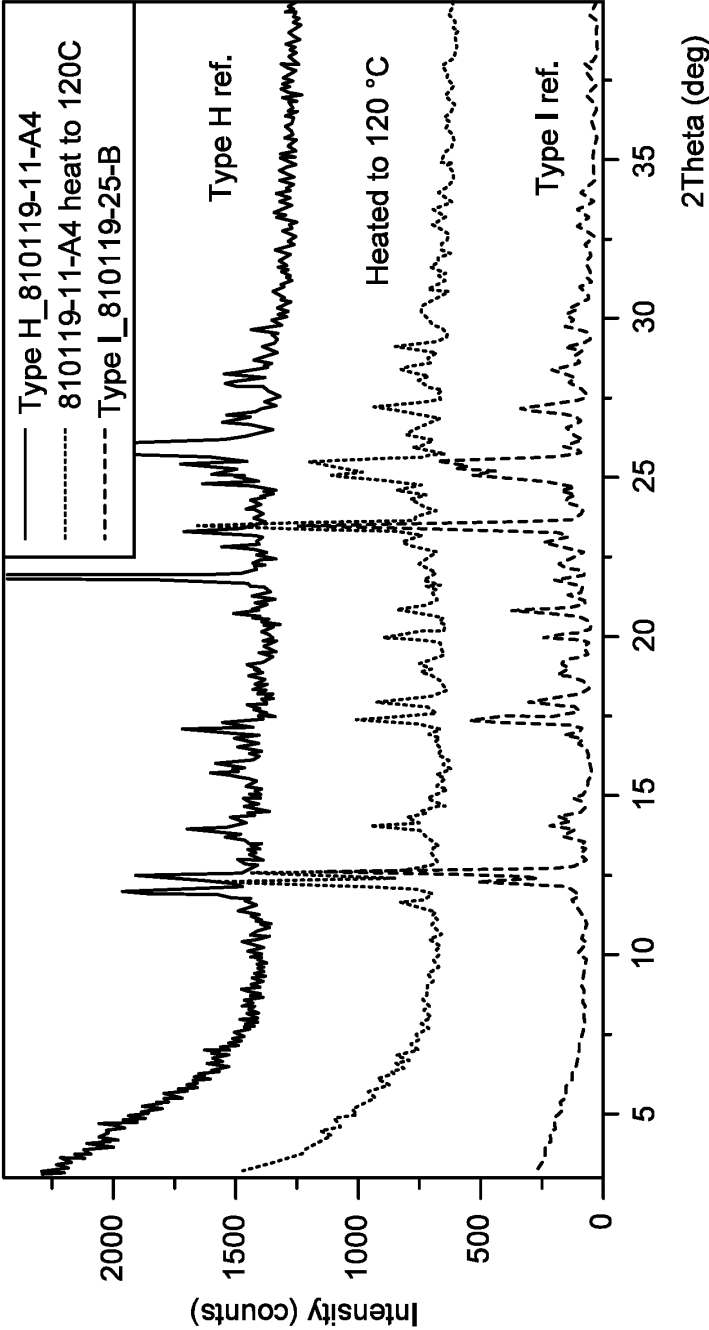
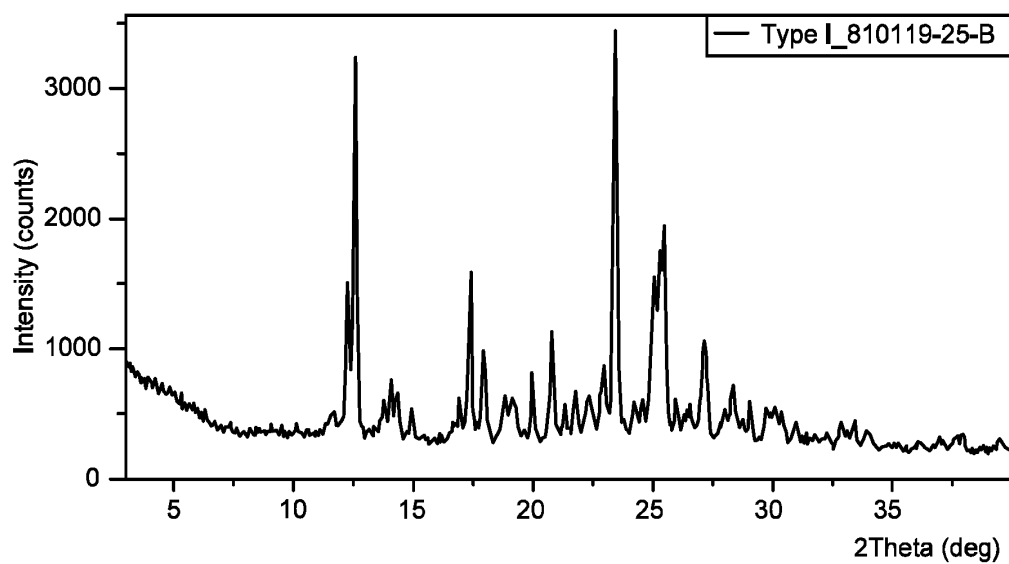
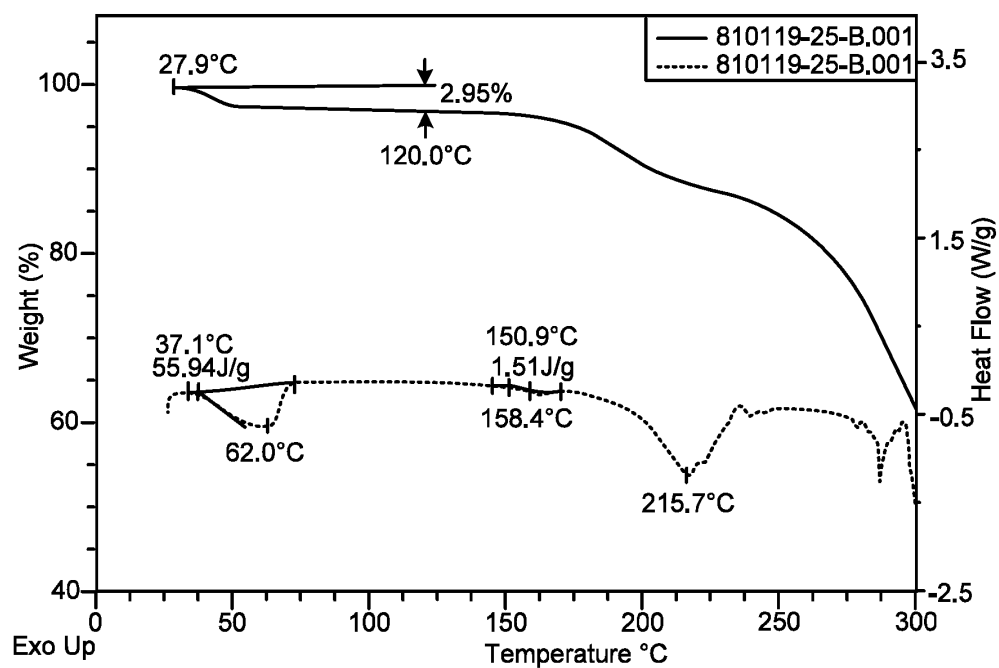
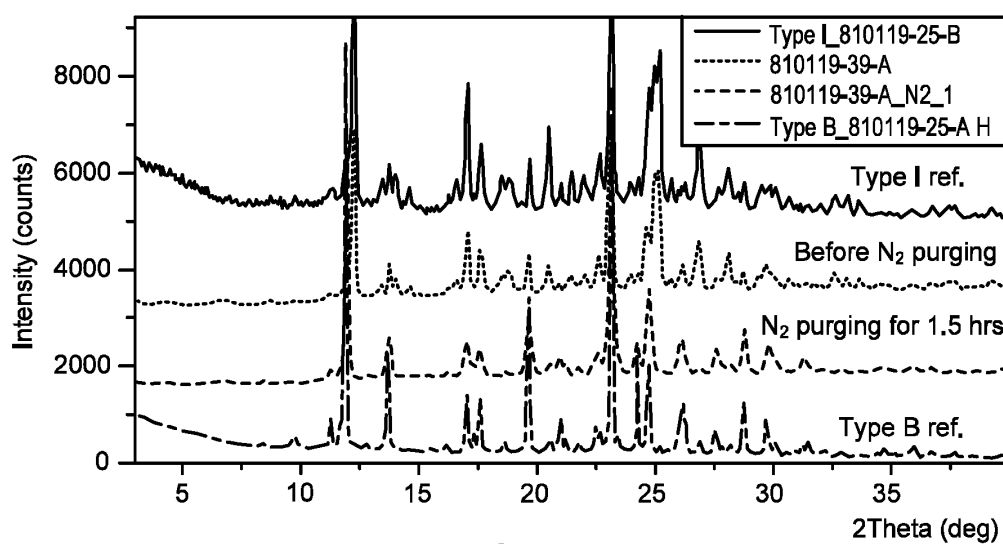
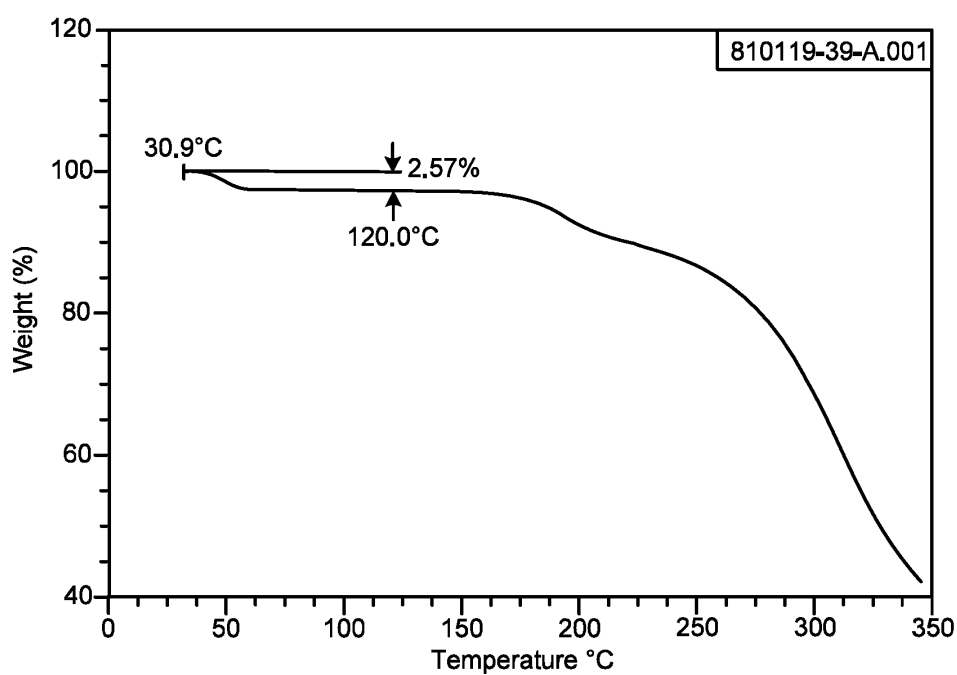


FIG. 54

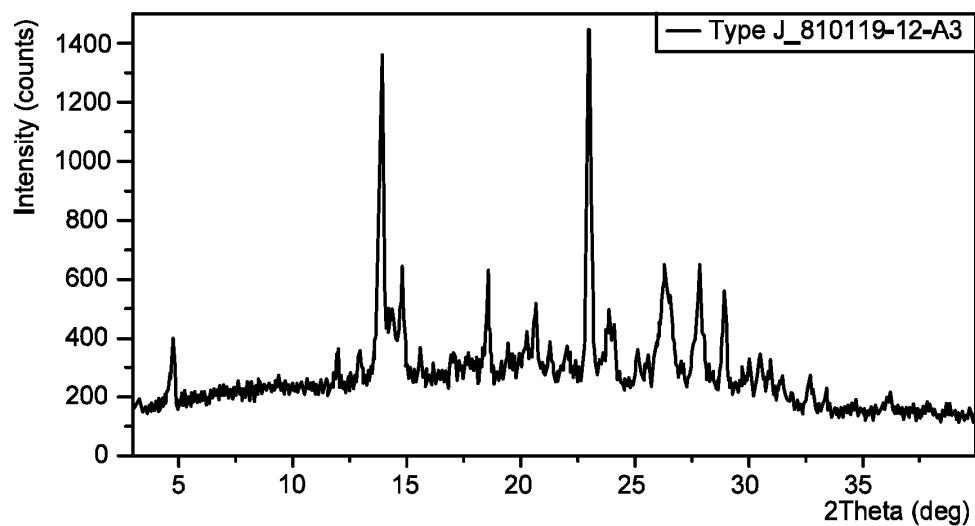
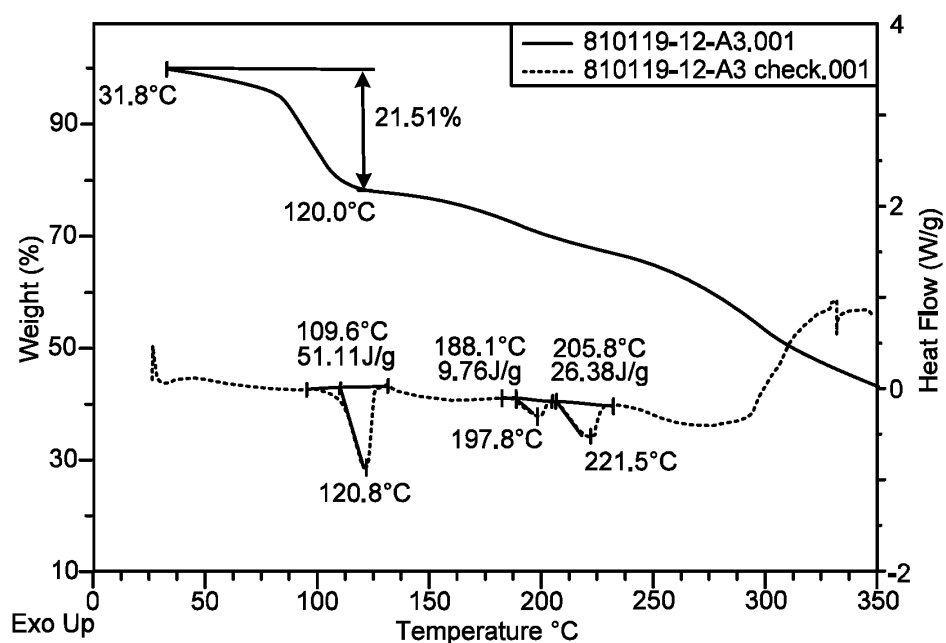
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**FIG. 55****FIG. 56**

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**FIG. 57****FIG. 58**

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**FIG. 59****FIG. 60**

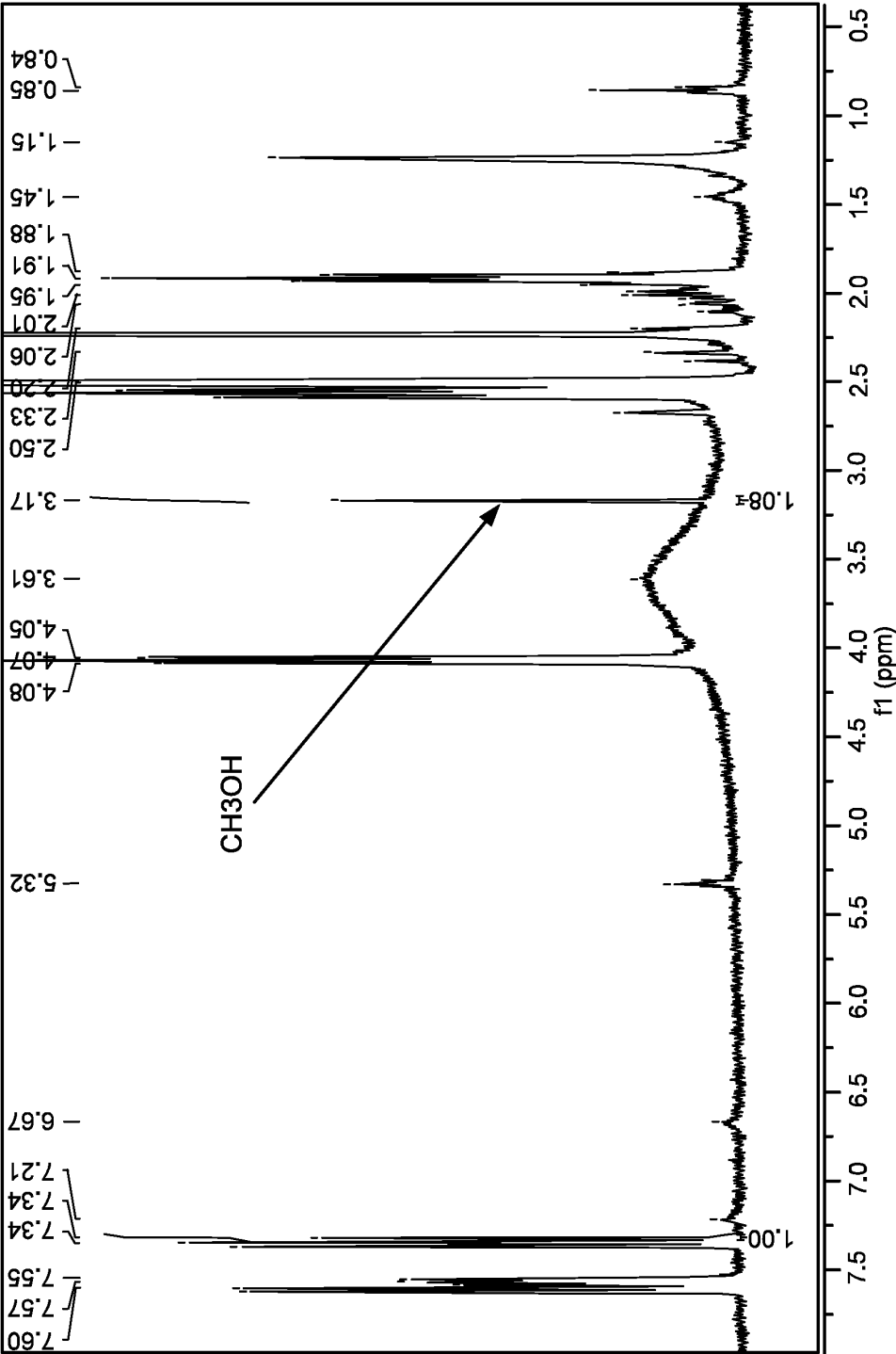


FIG. 61

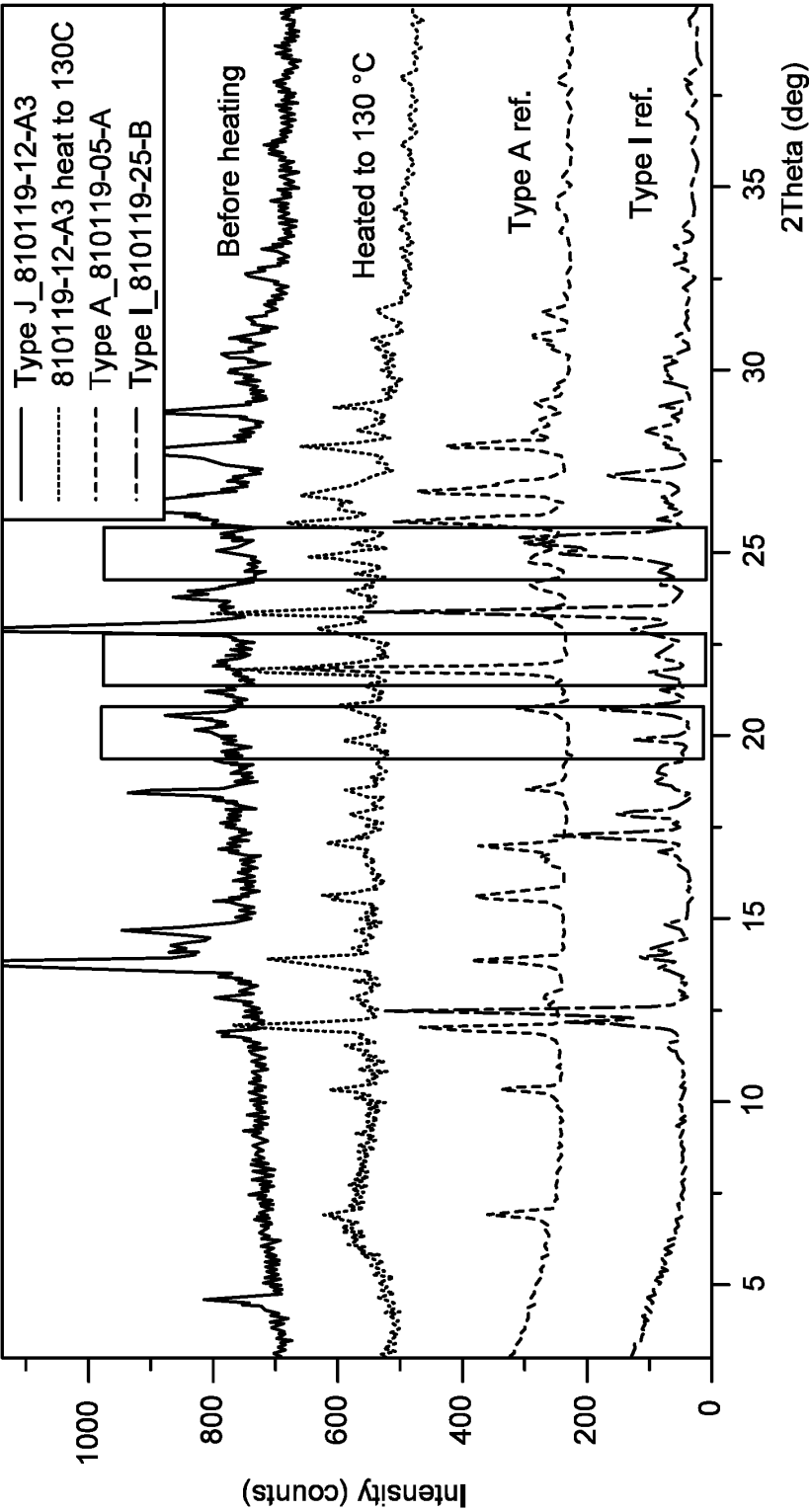
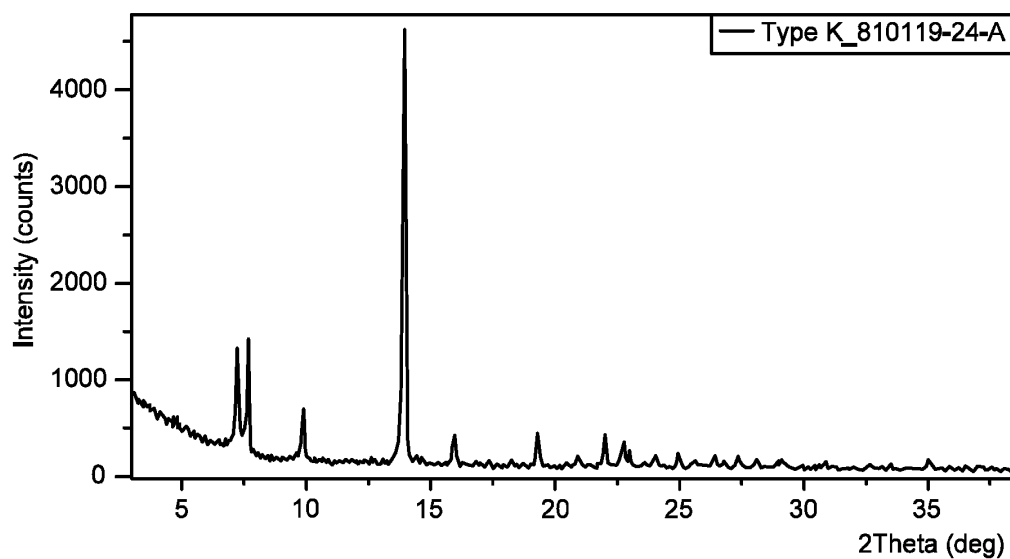
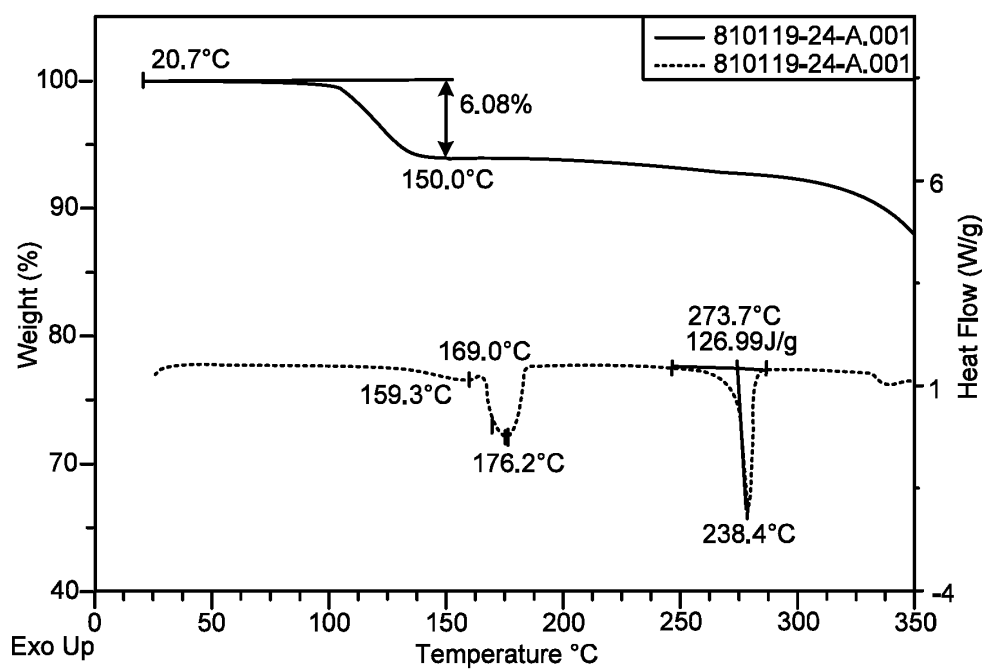


FIG. 62

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**FIG. 63****FIG. 64**