

1 / 53

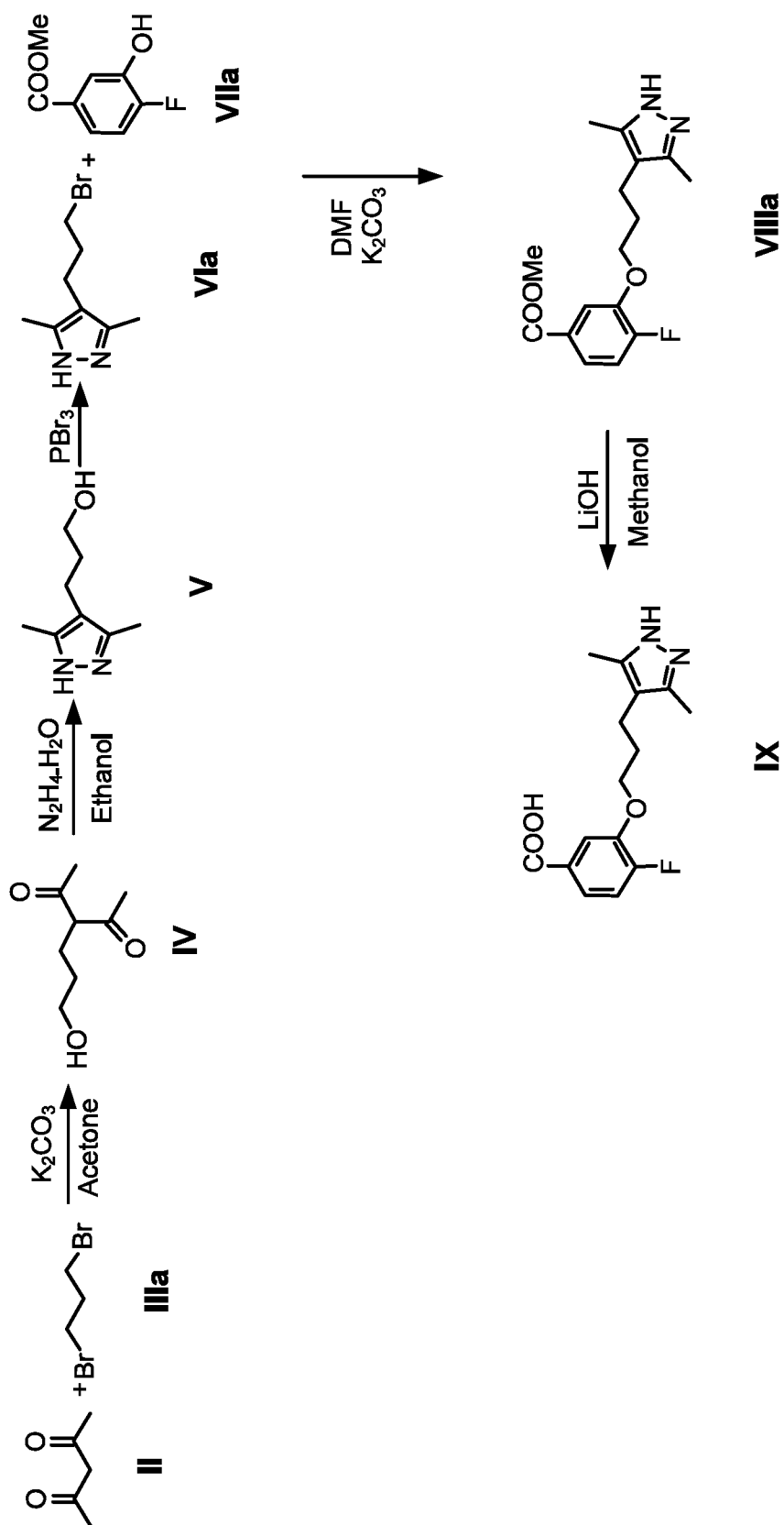
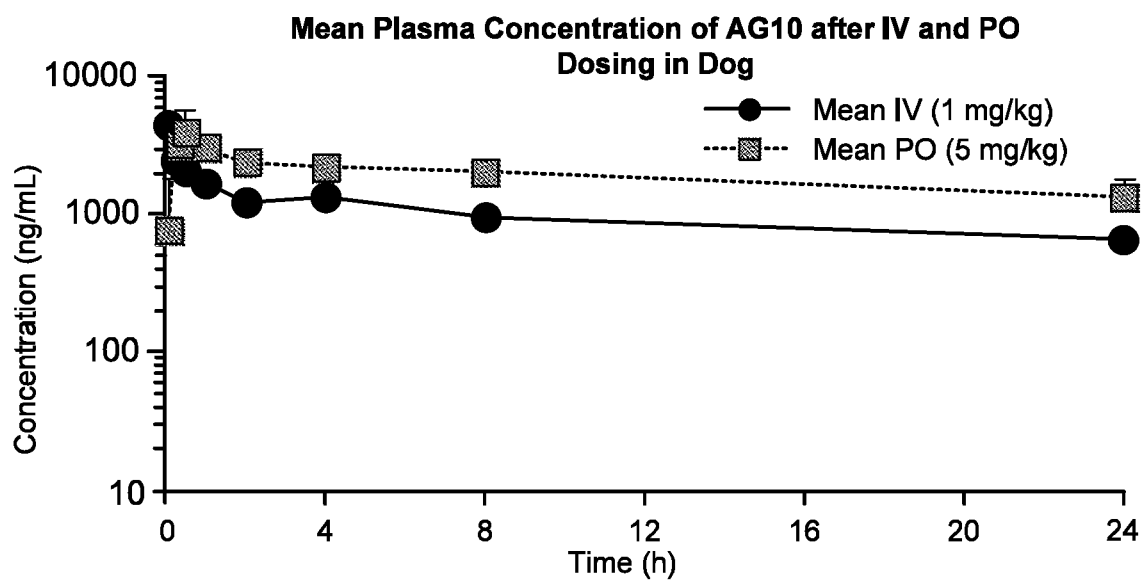


FIG. 1

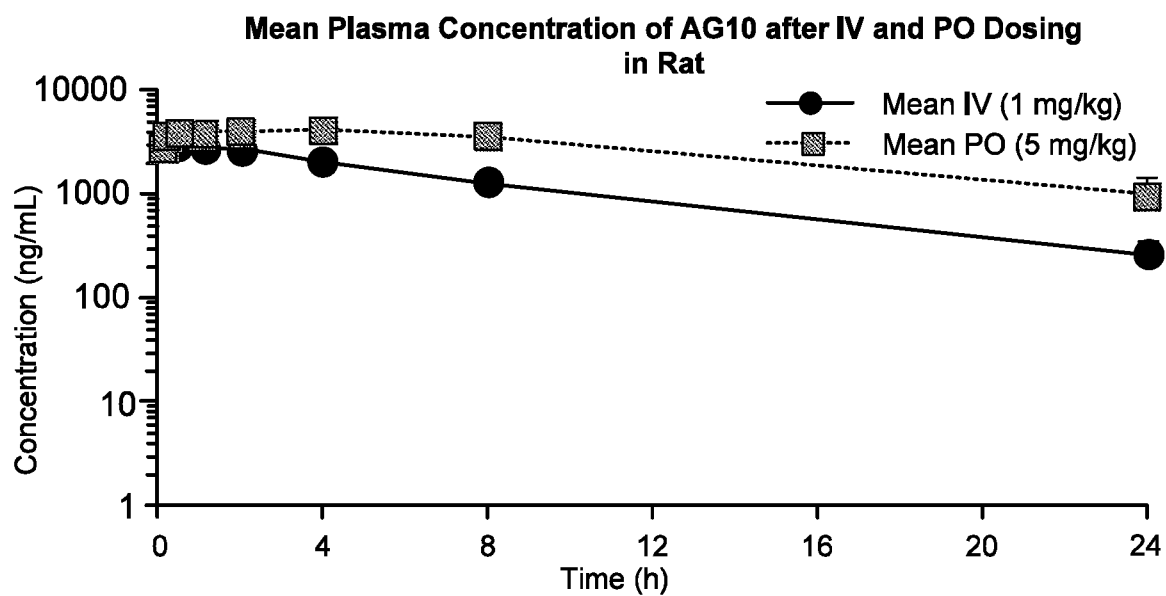
2 / 53

**FIG. 2A**

Dog		
IV 1mg/kg		
$T_{1/2}$	V_{dss}	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

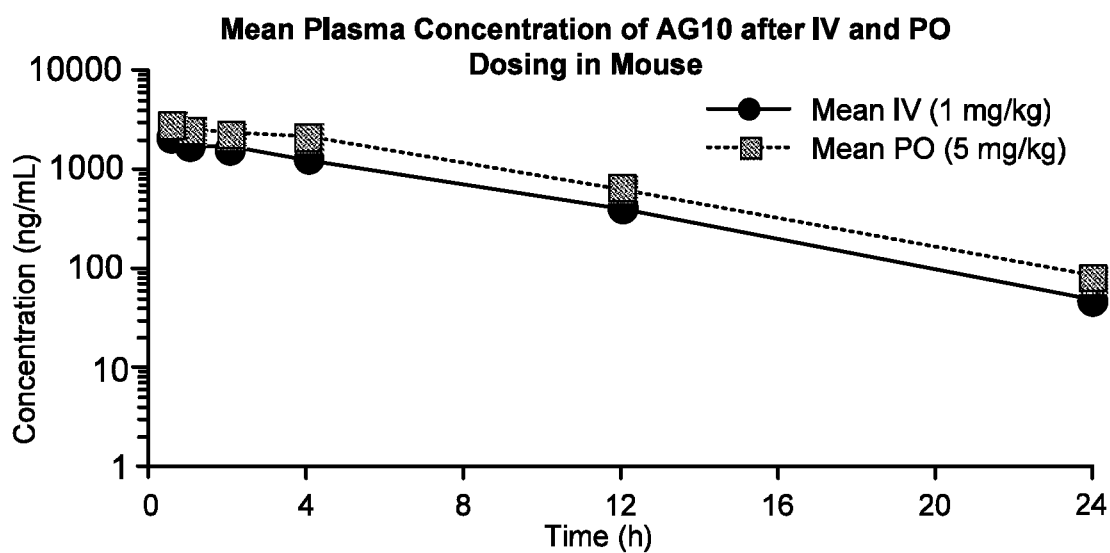
3 / 53

**FIG. 2C**

Rat		
IV 1mg/kg		
$T_{1/2}$	V_{dss}	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

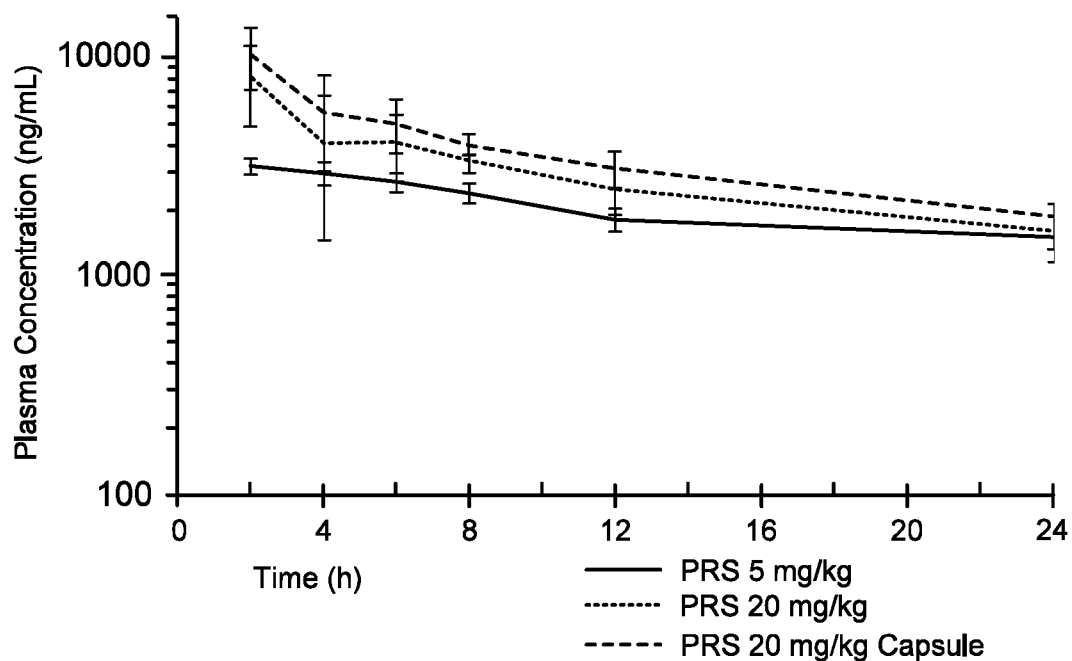
4 / 53

**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

5 / 53

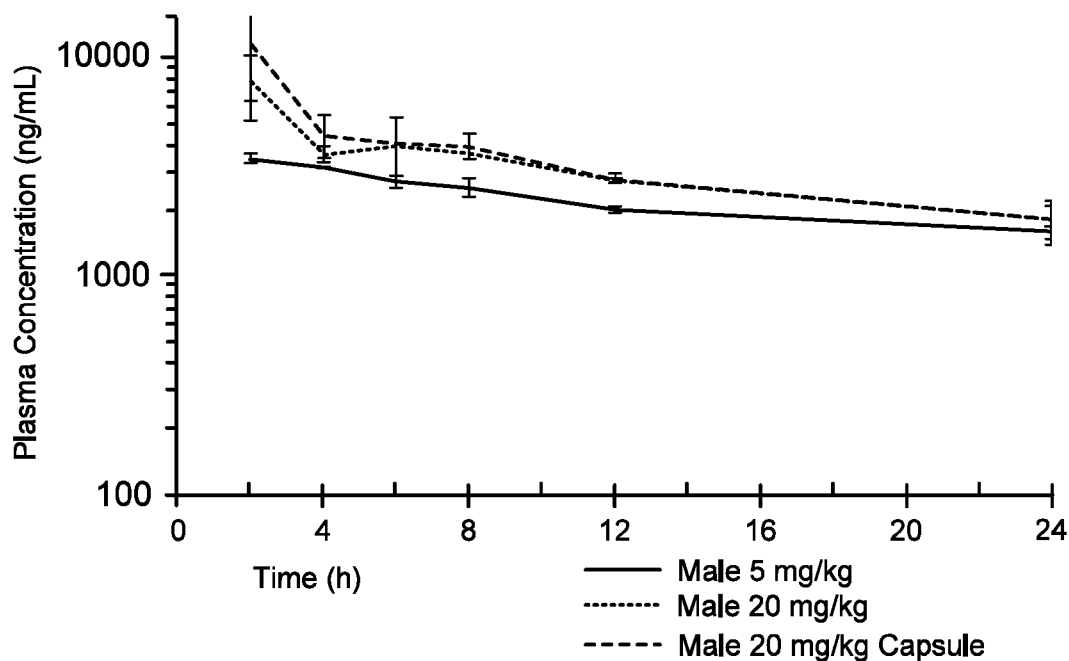
PRS Dog 1 Day Combined**FIG. 3A**

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

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

FIG. 3B

6 / 53

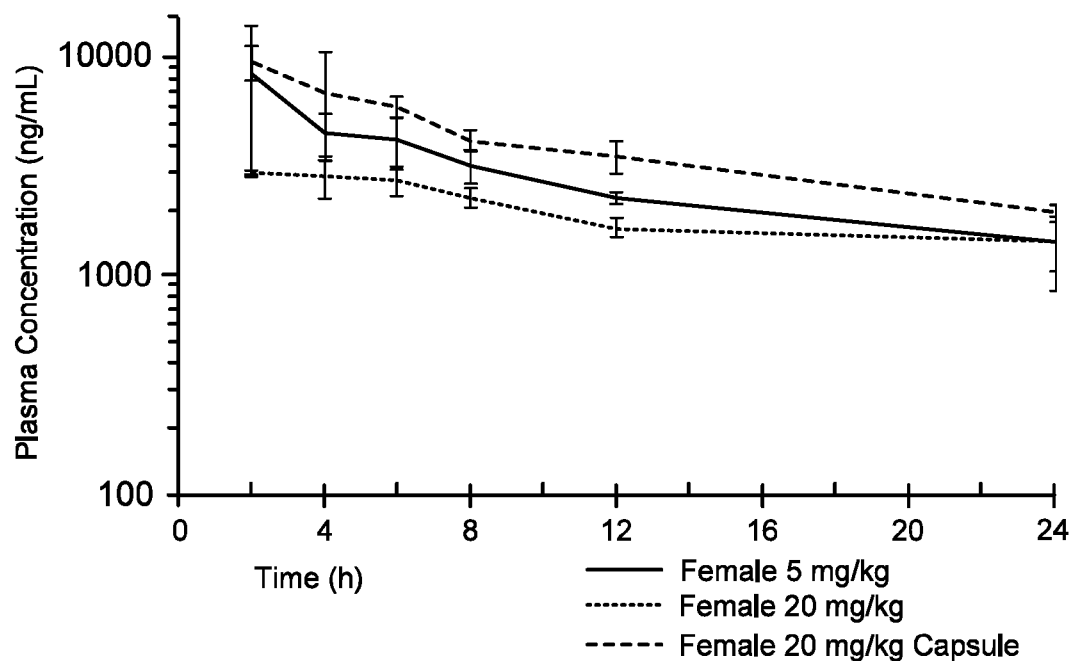
PRS Dog 1 Day Male**FIG. 3C**

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

7 / 53

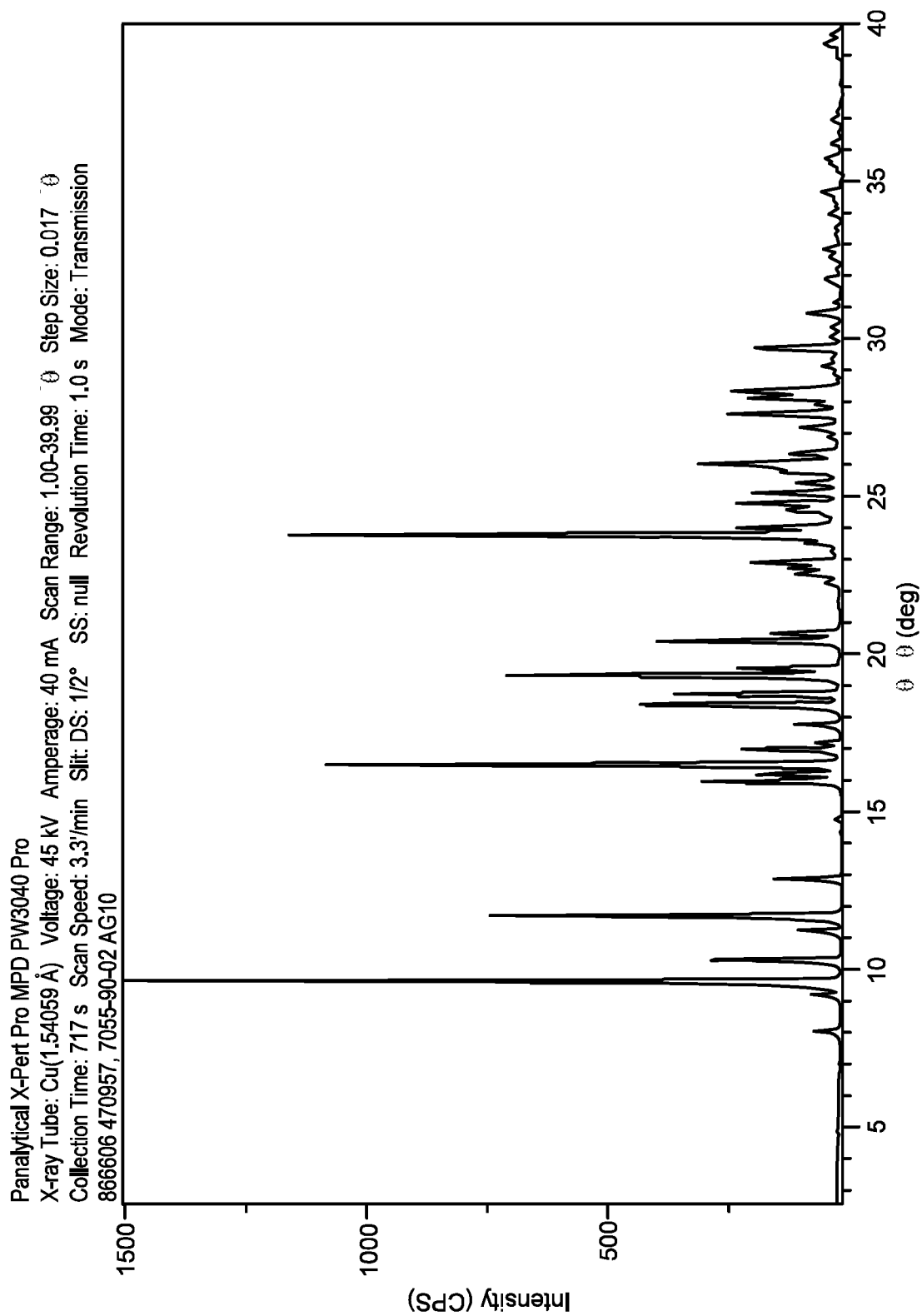
PRS Dog 1 Day Female**FIG. 3E**

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

8 / 53

**FIG. 4**

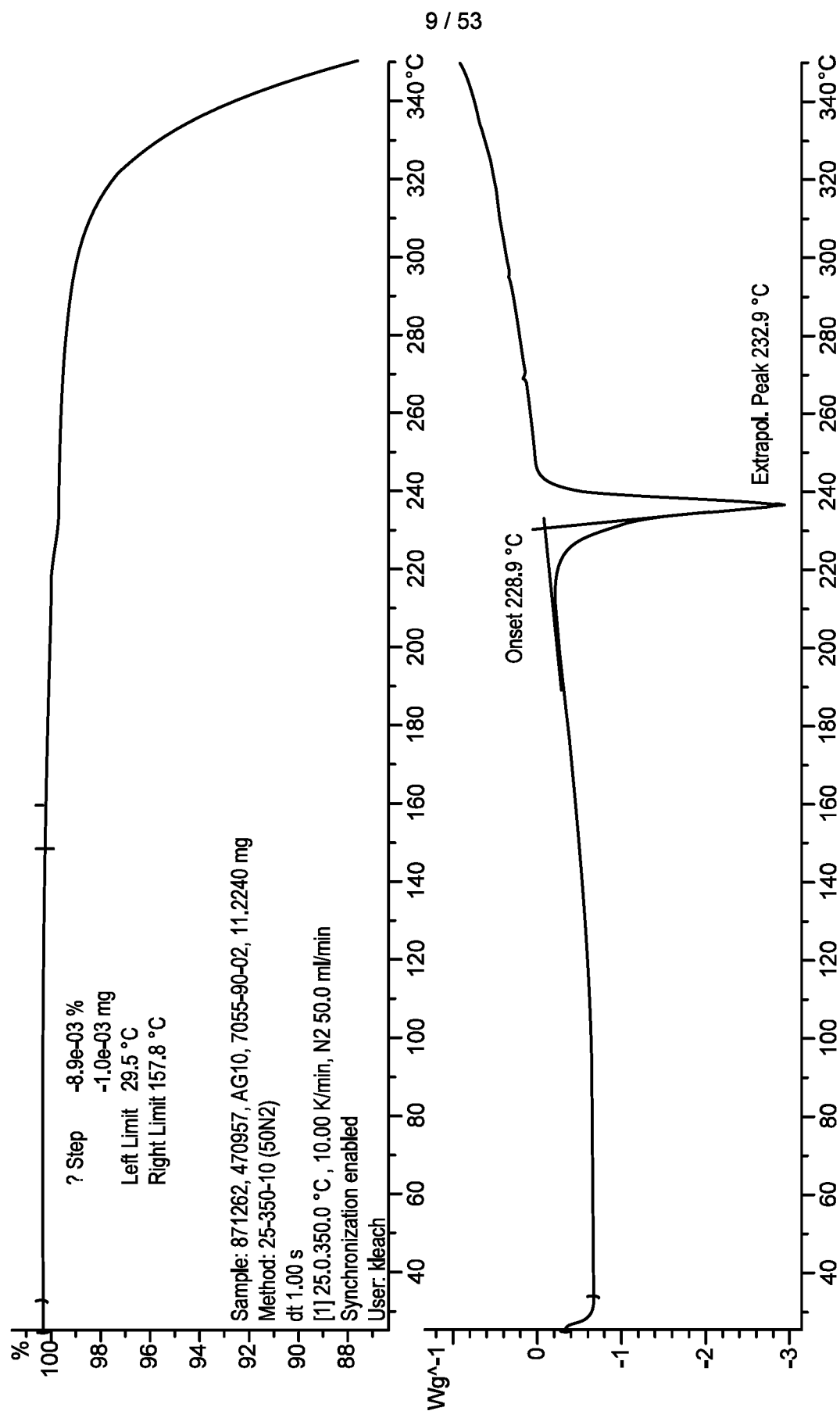
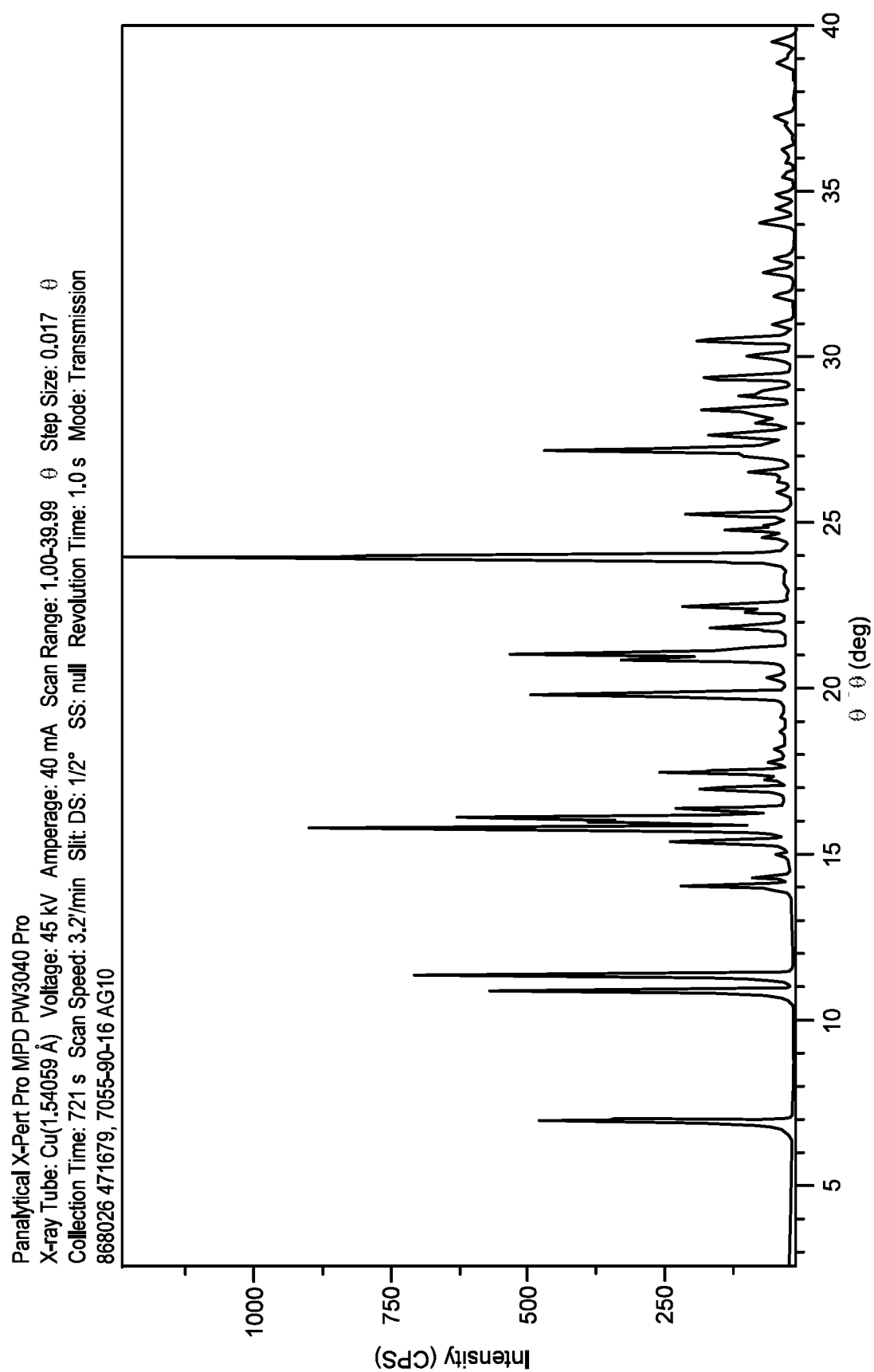
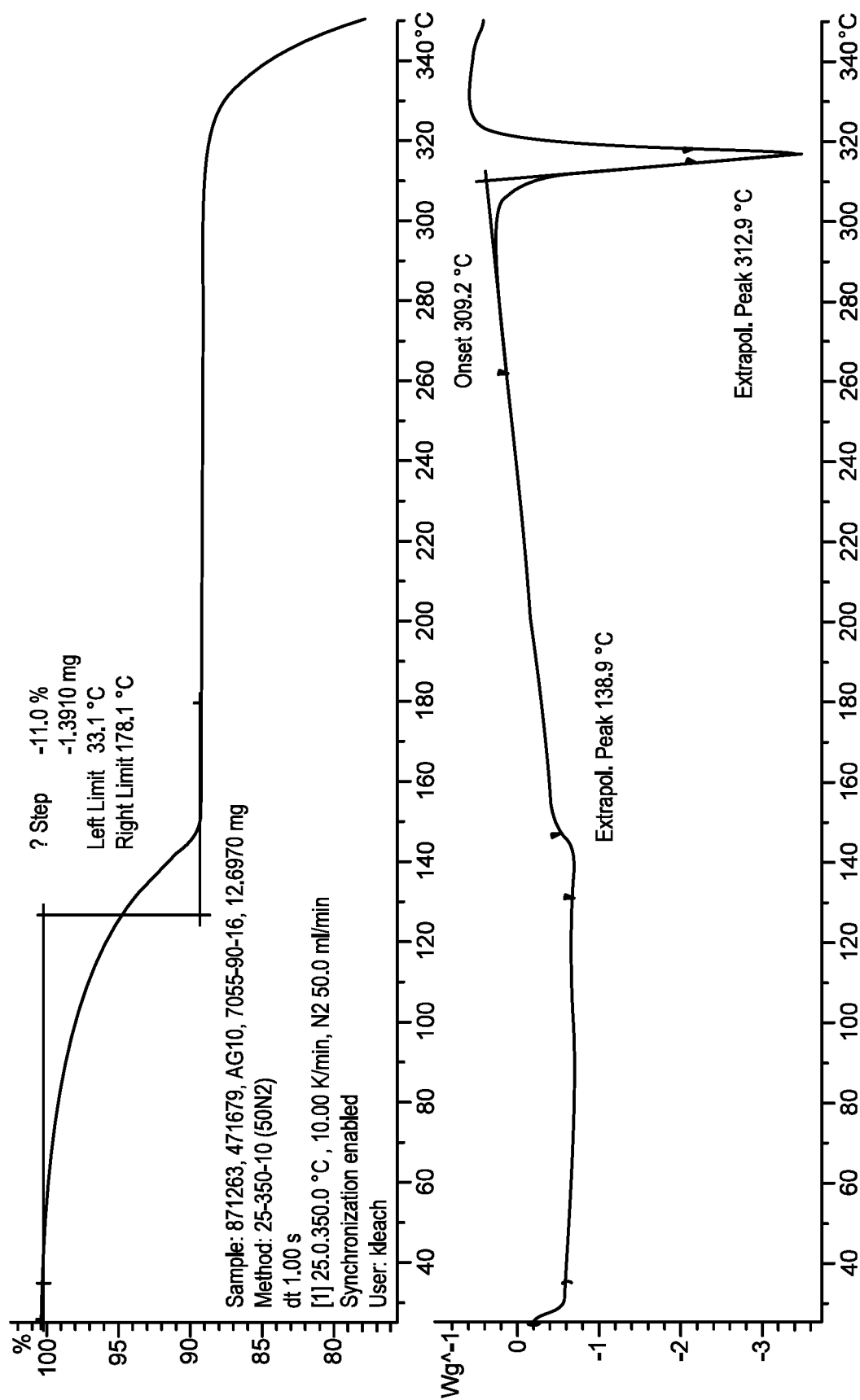


FIG. 5

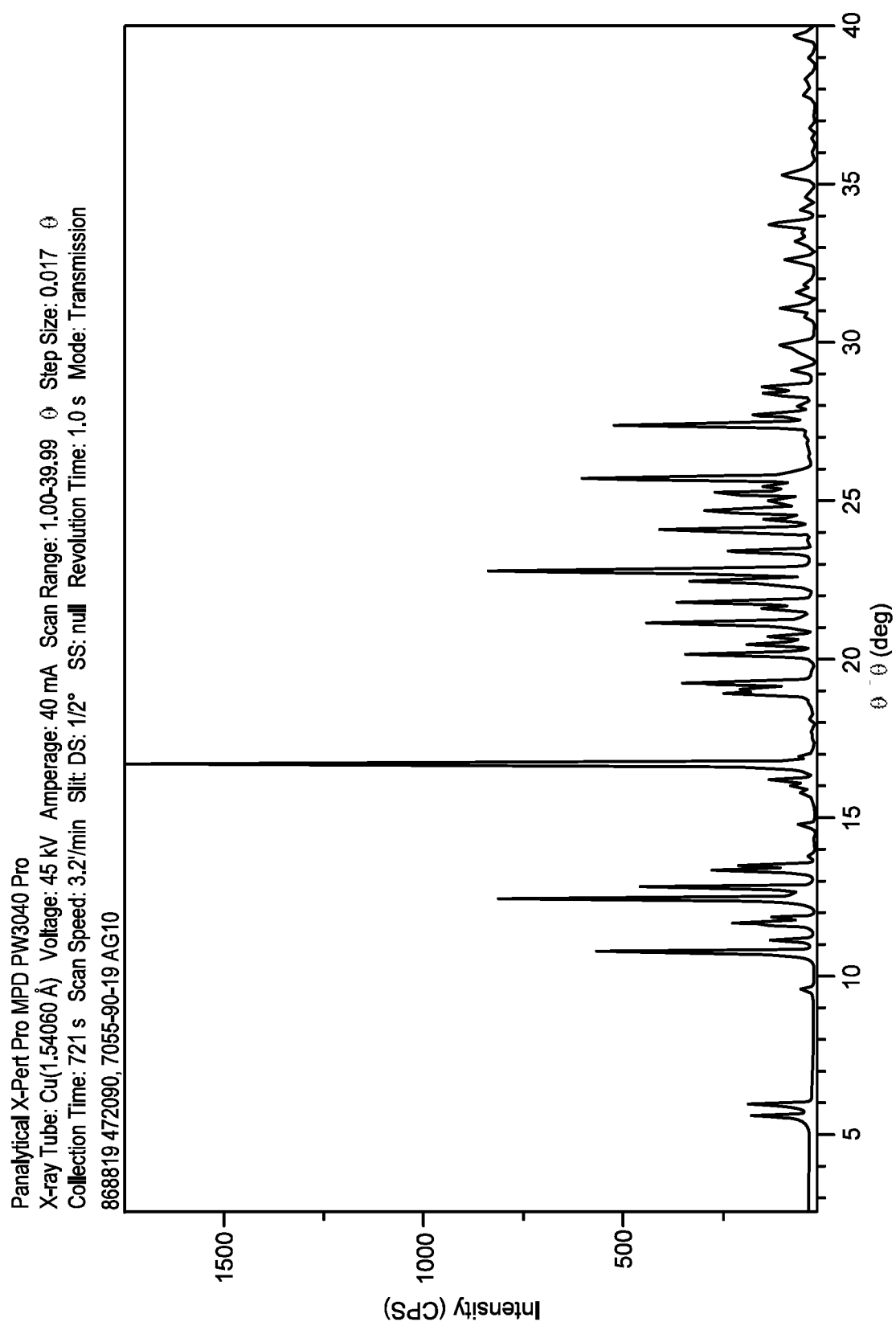
10 / 53

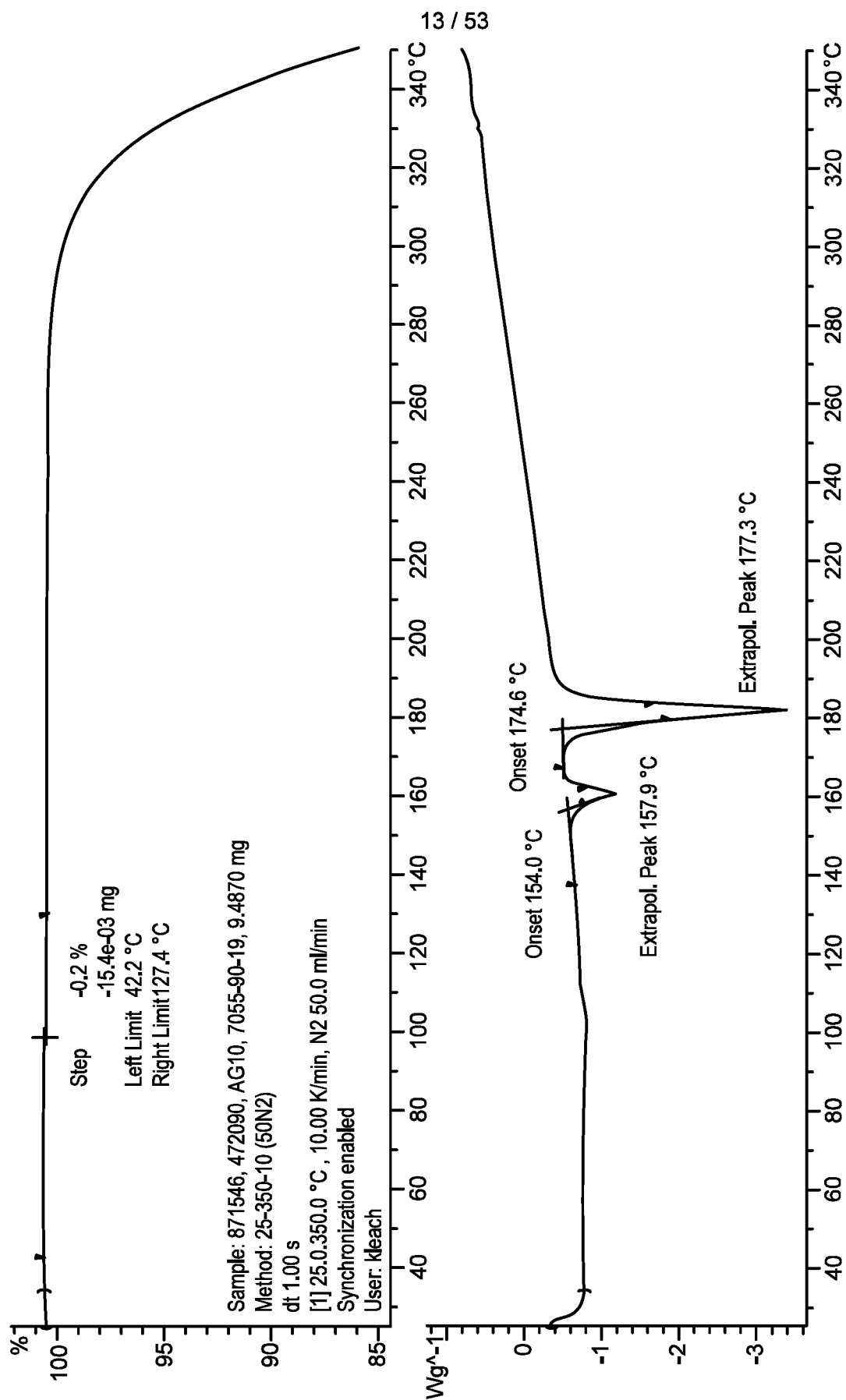
**FIG. 6**

11 / 53

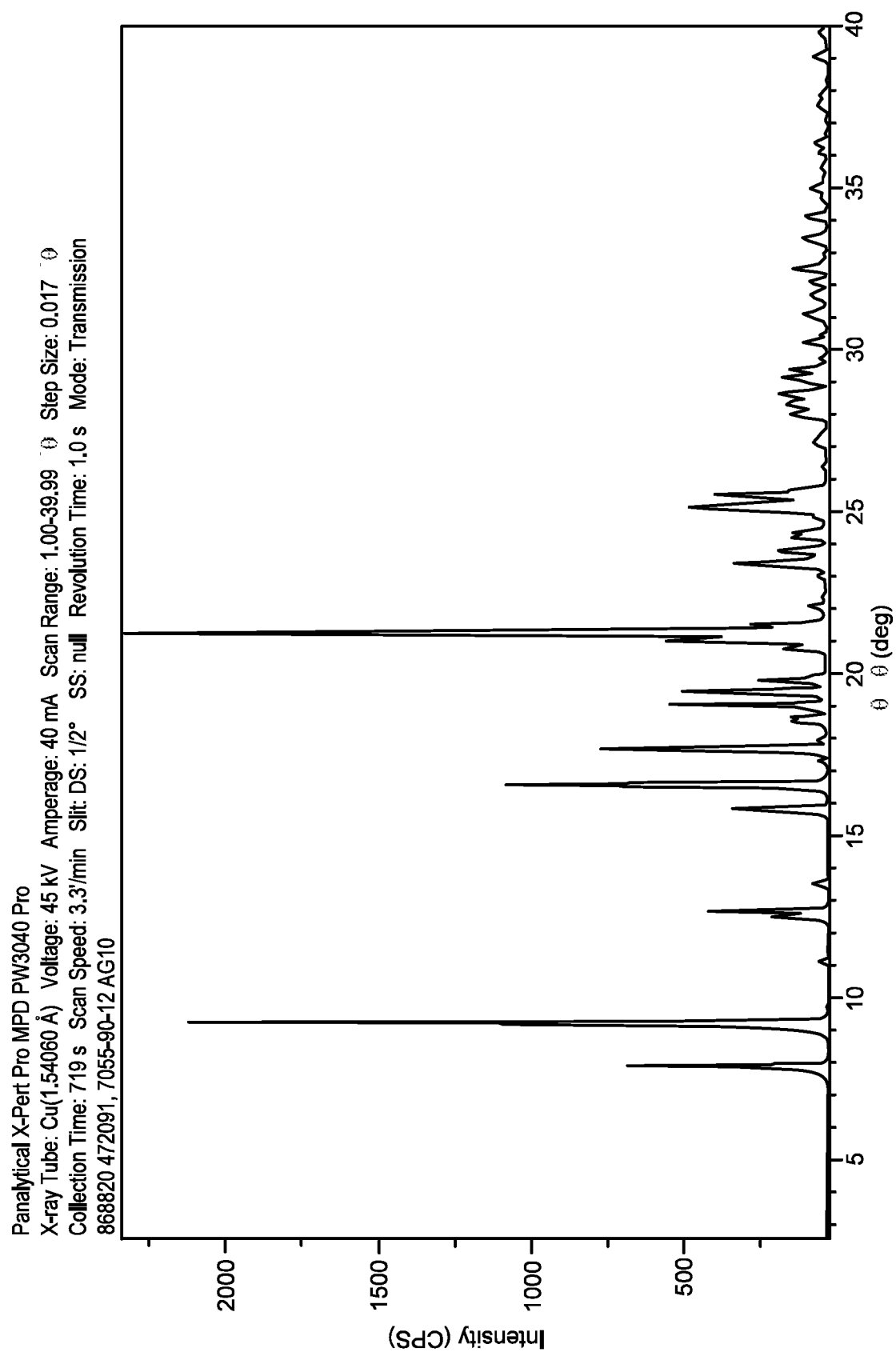
**FIG. 7**

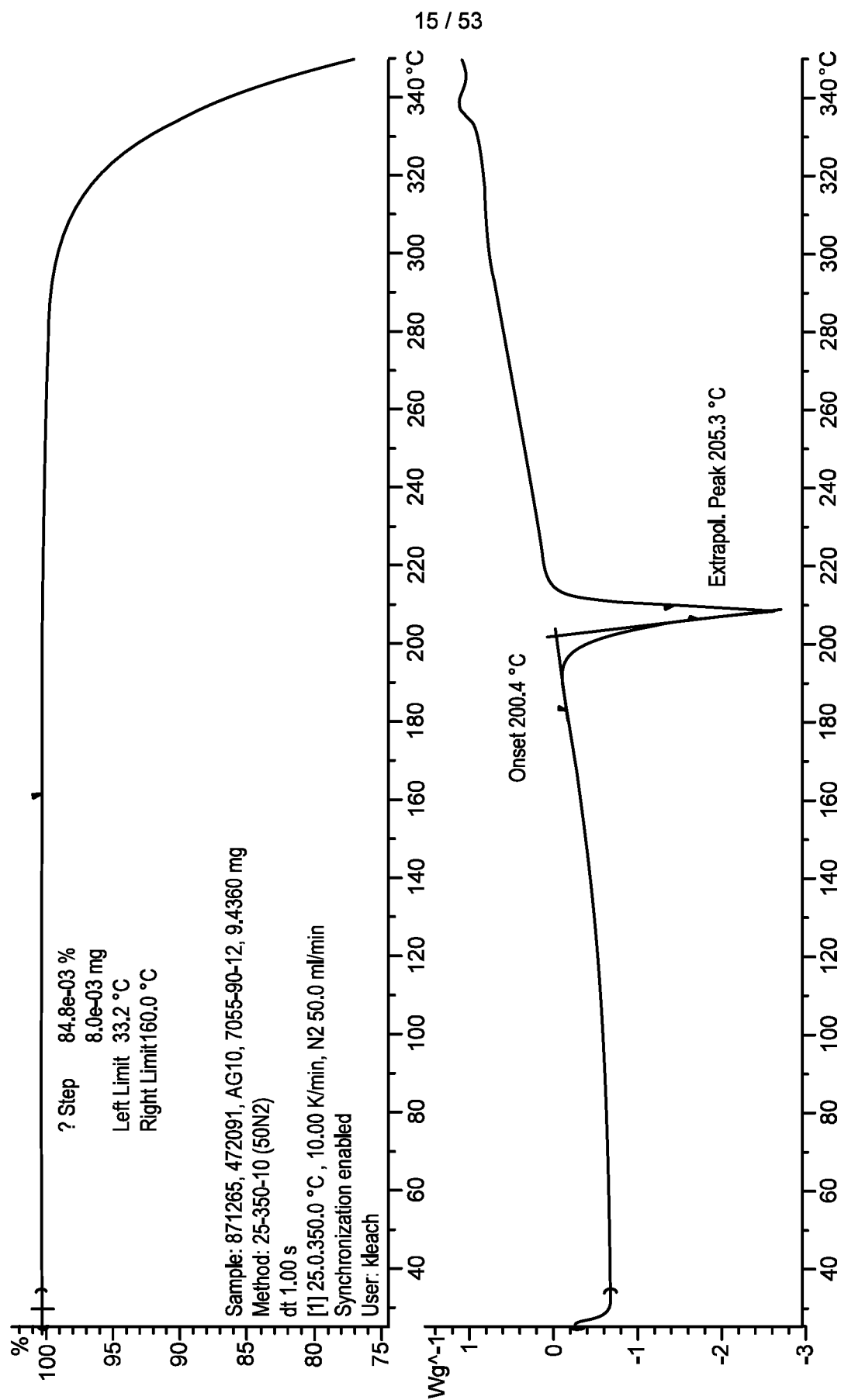
12 / 53

**FIG. 8**

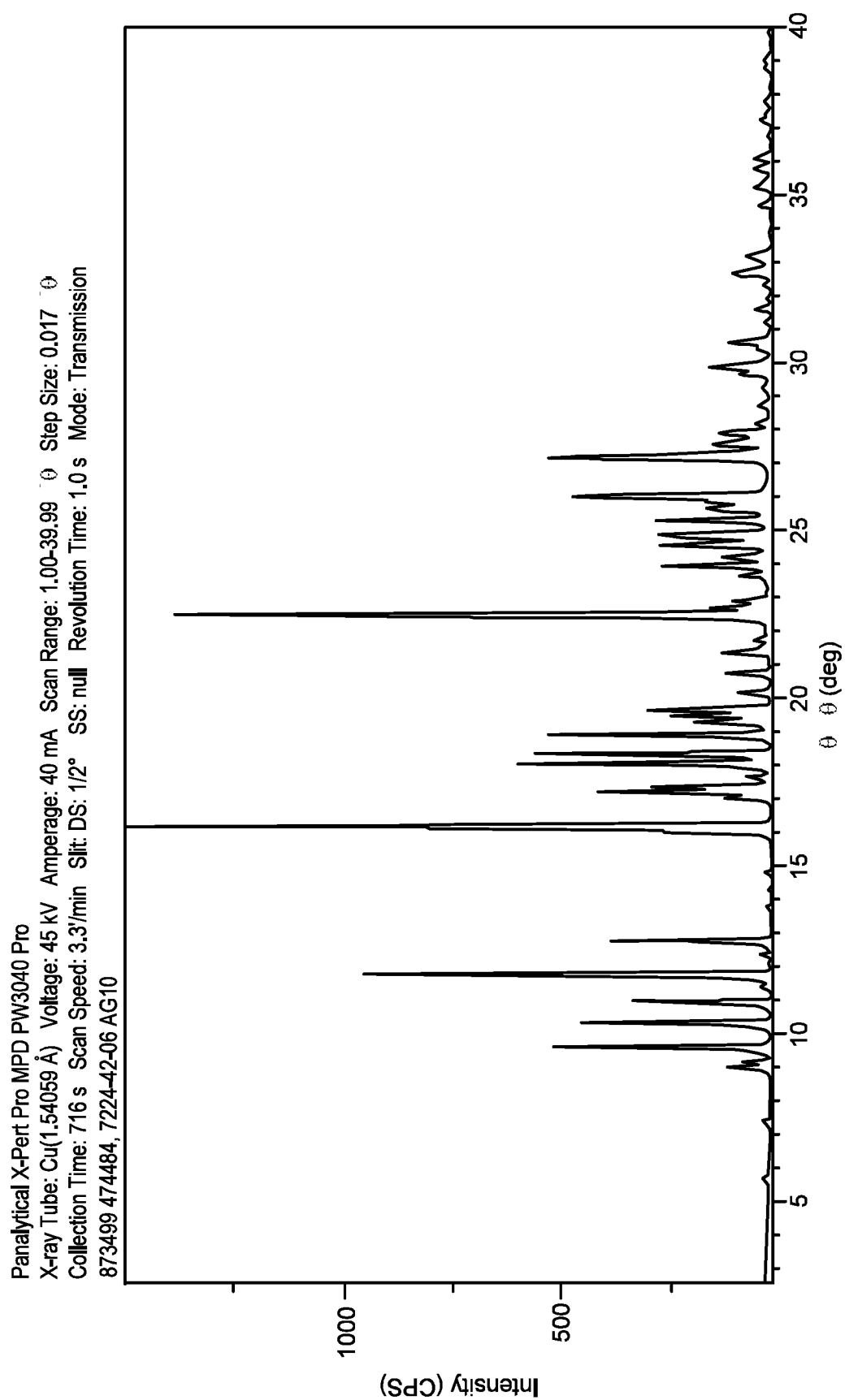
**FIG. 9**

14 / 53

**FIG. 10**

**FIG. 11**

16 / 53

**FIG. 12**

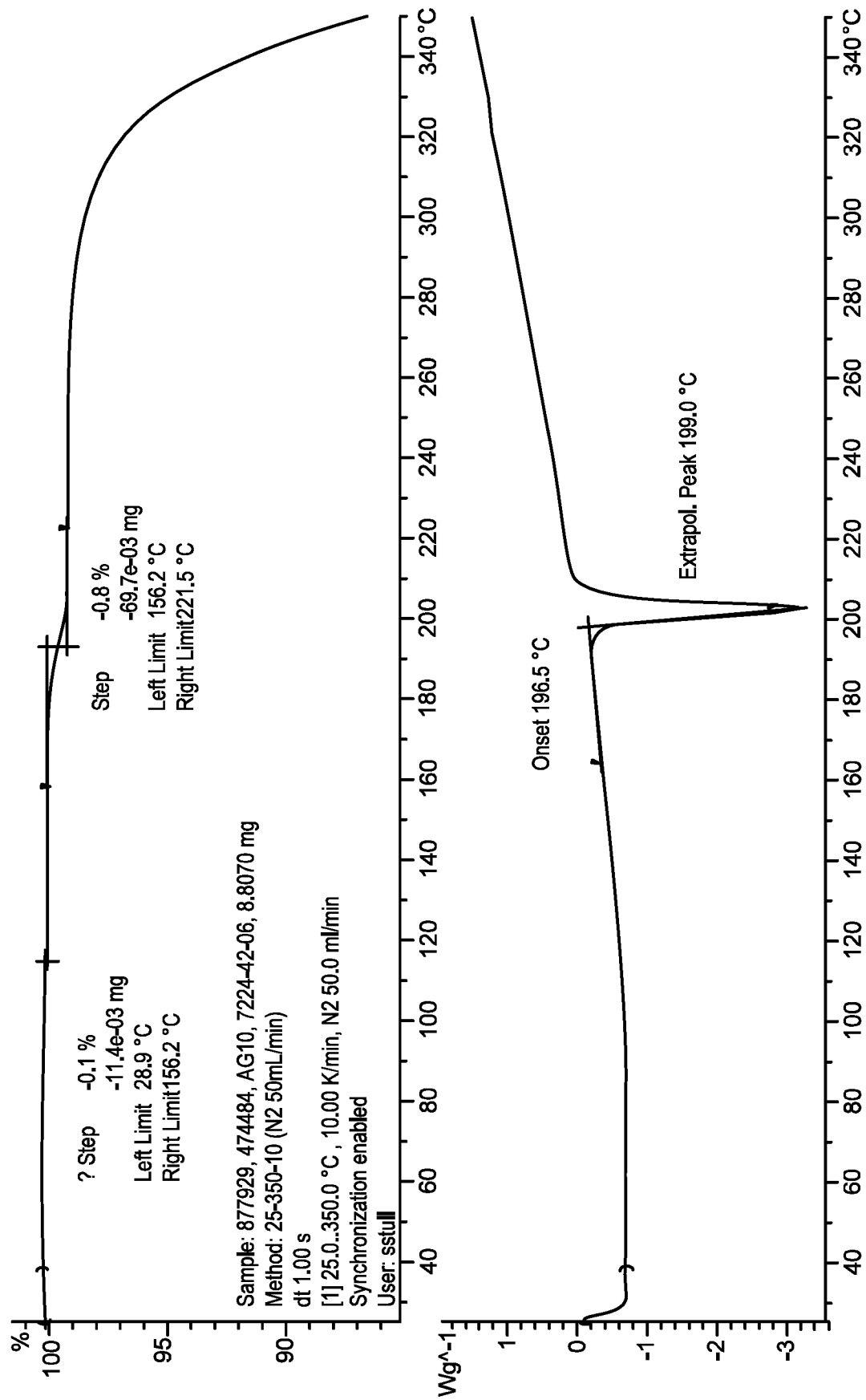
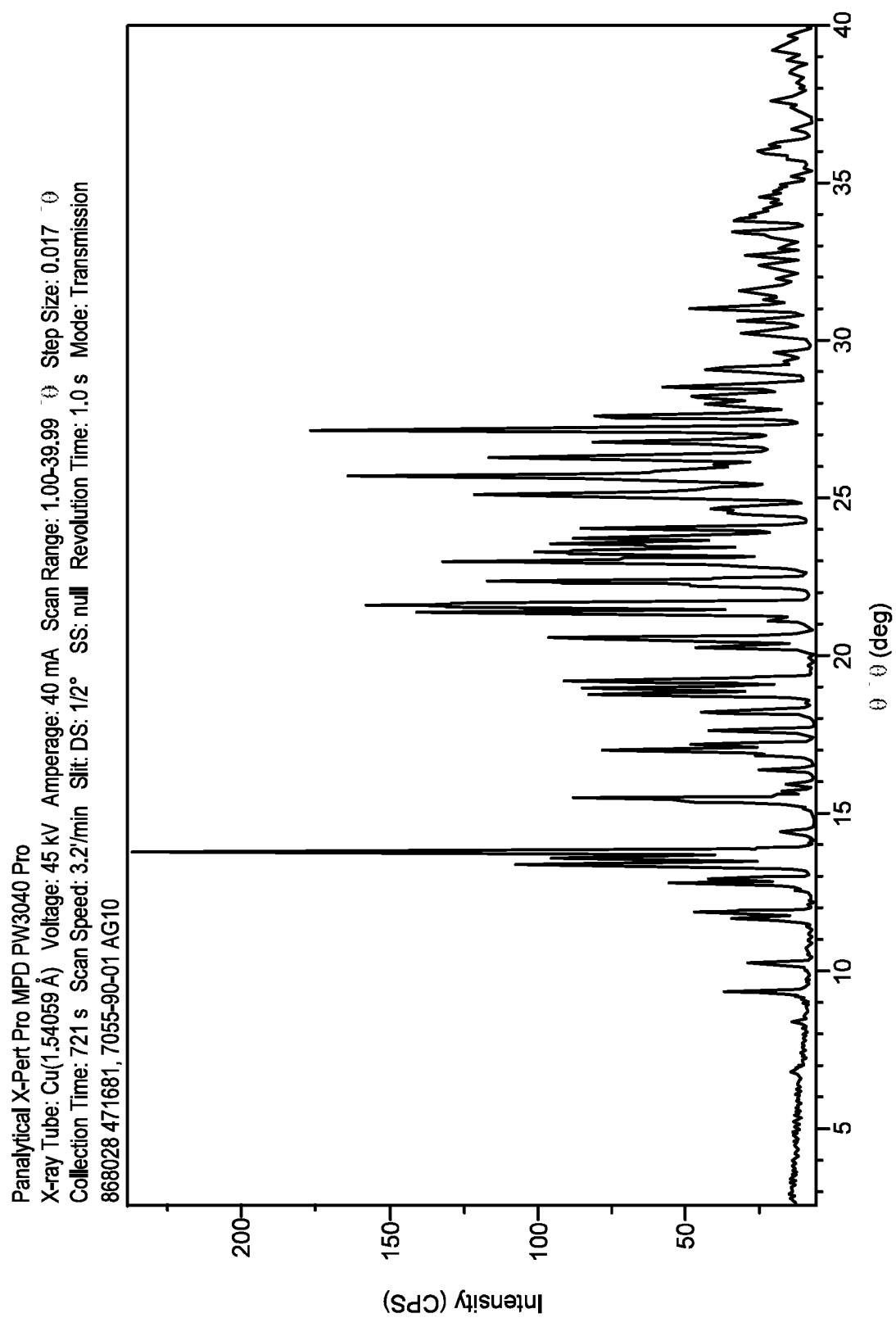
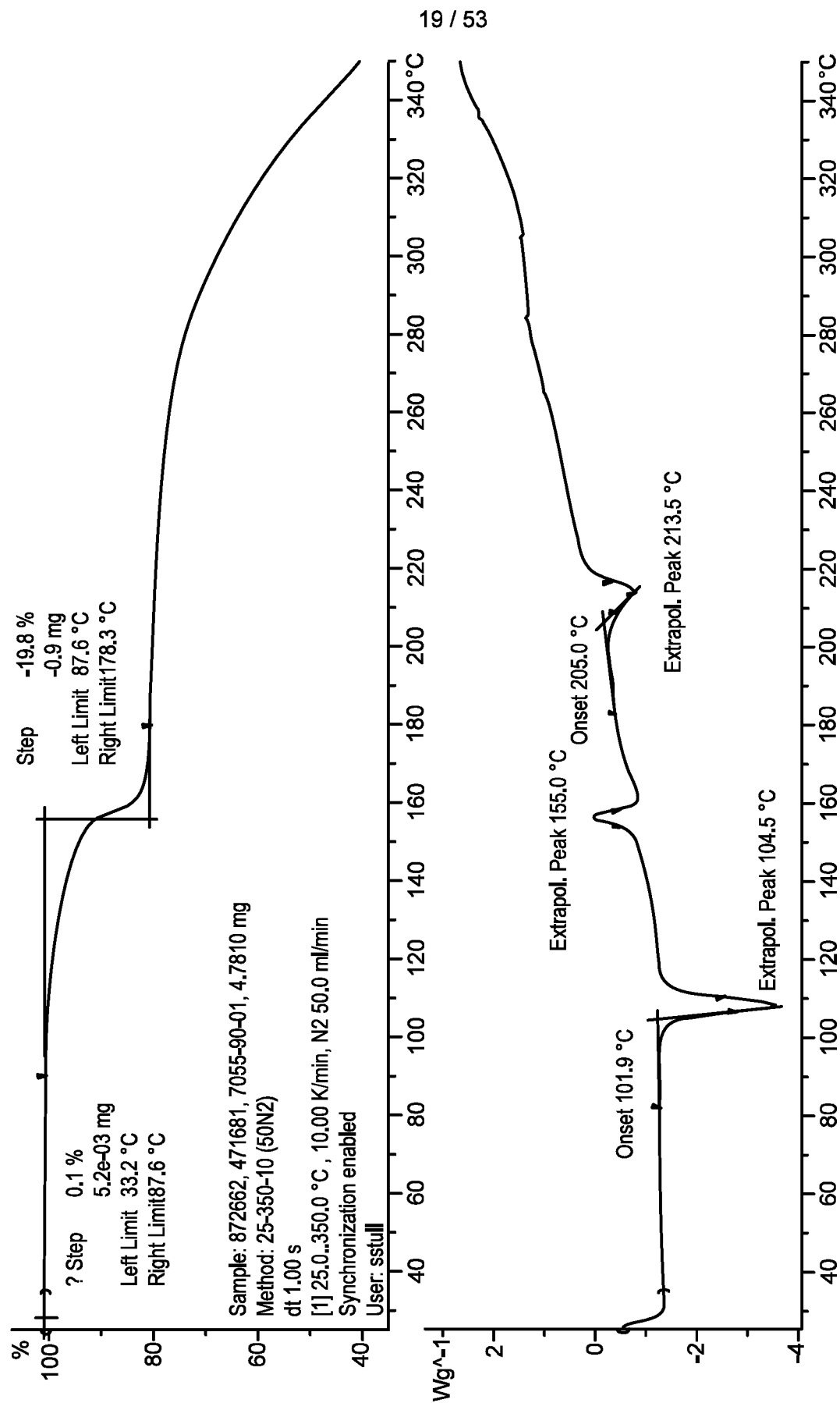


FIG. 13

18 / 53

**FIG. 14**

**FIG. 15**

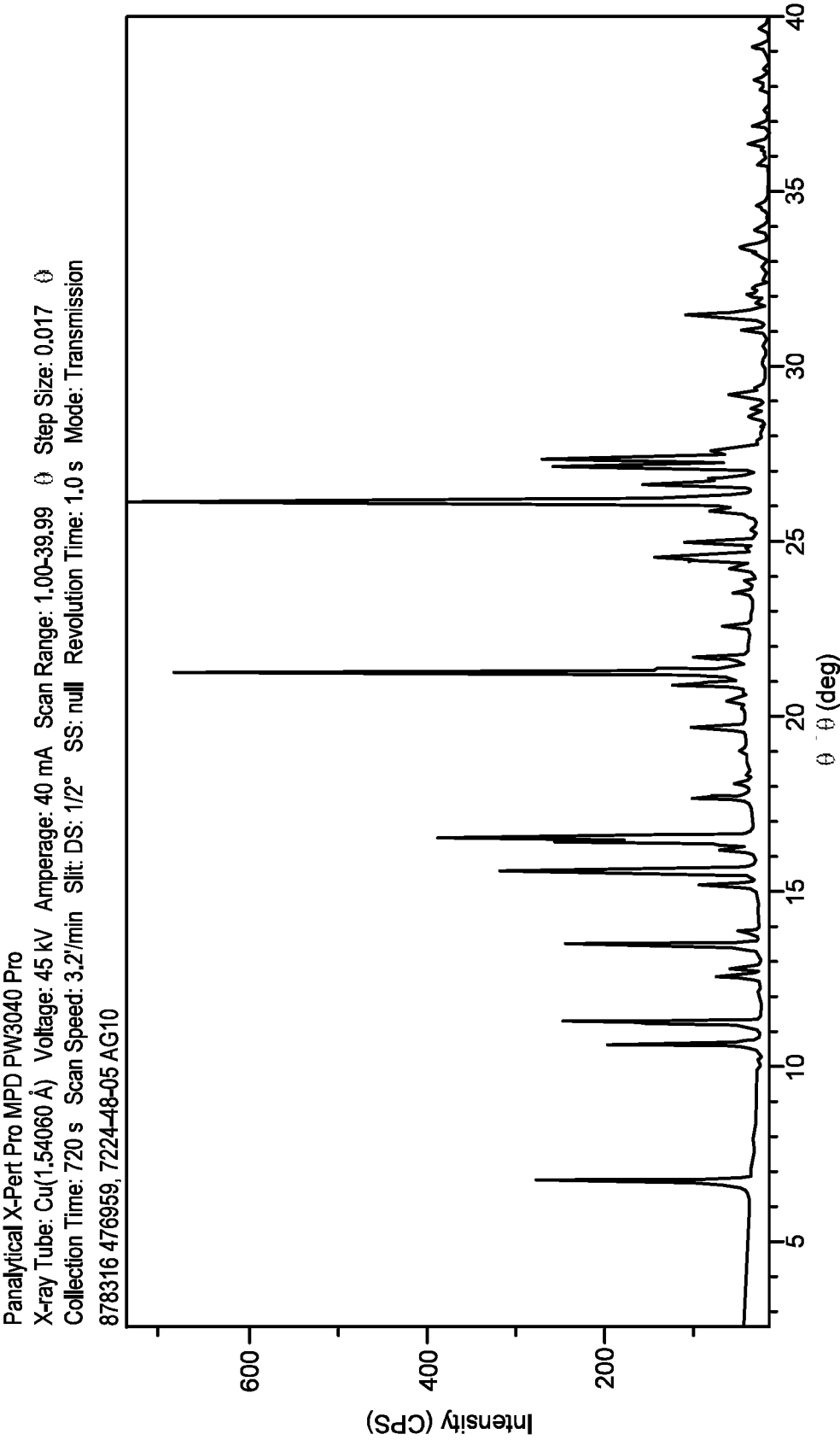


FIG. 16

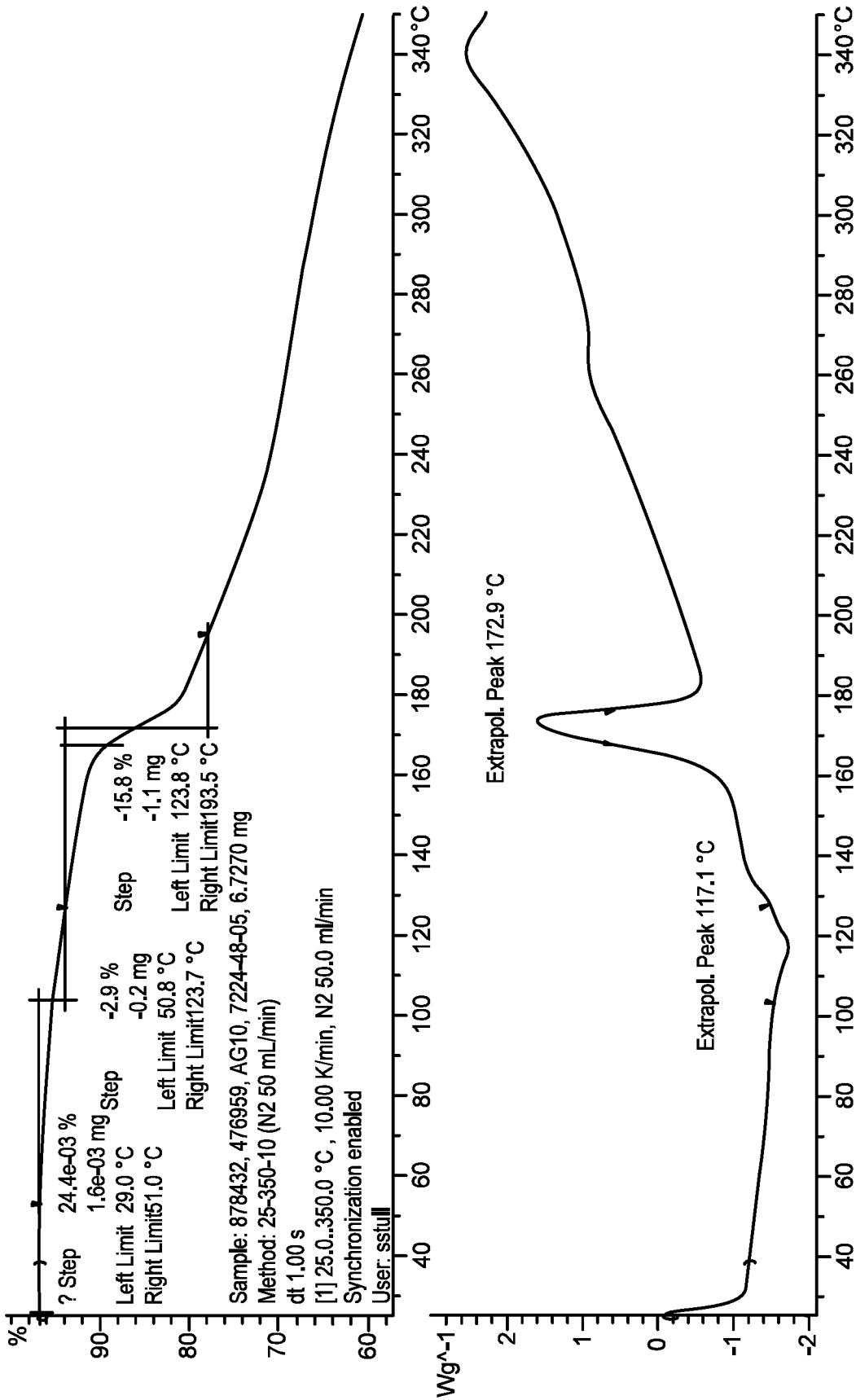
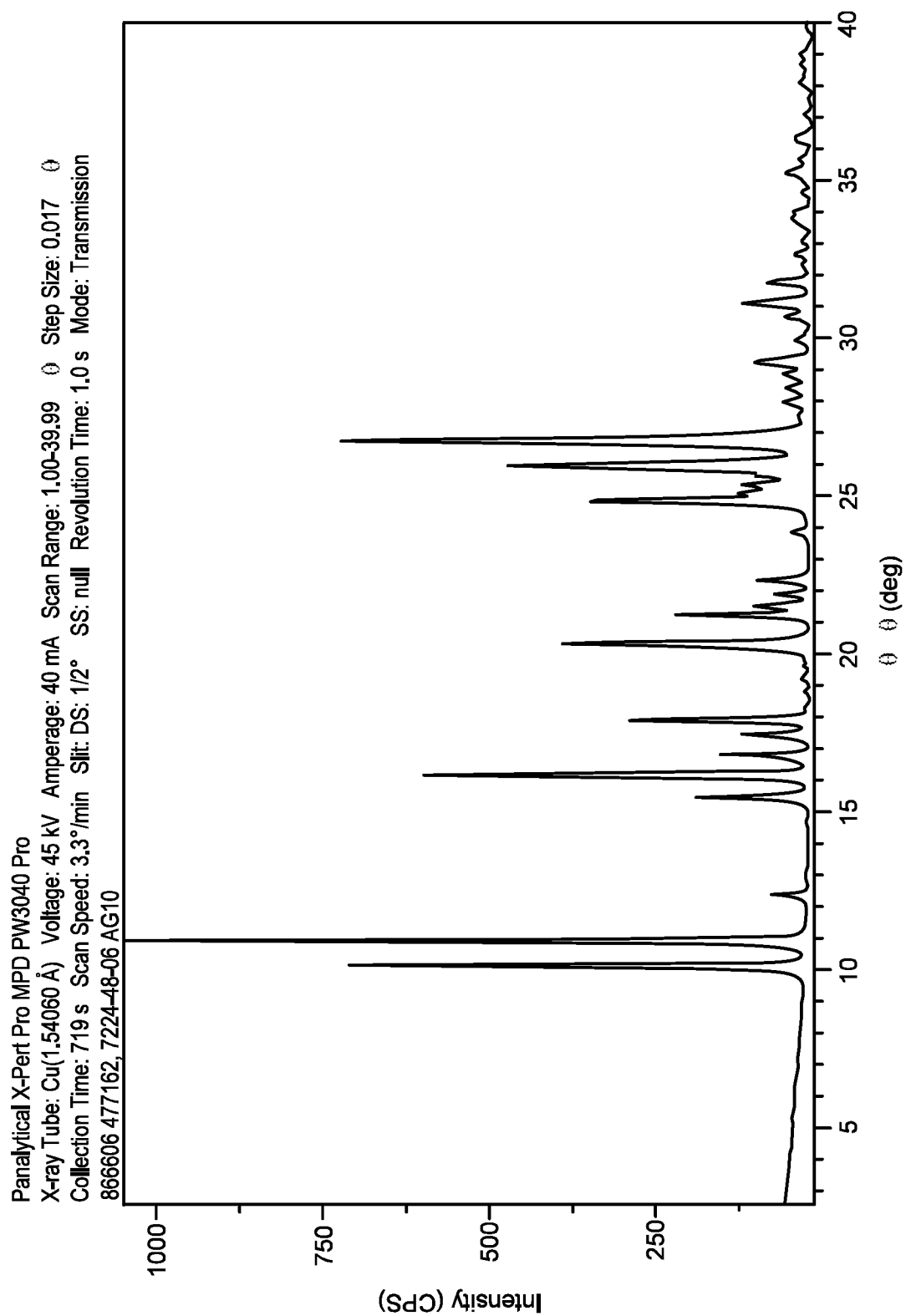
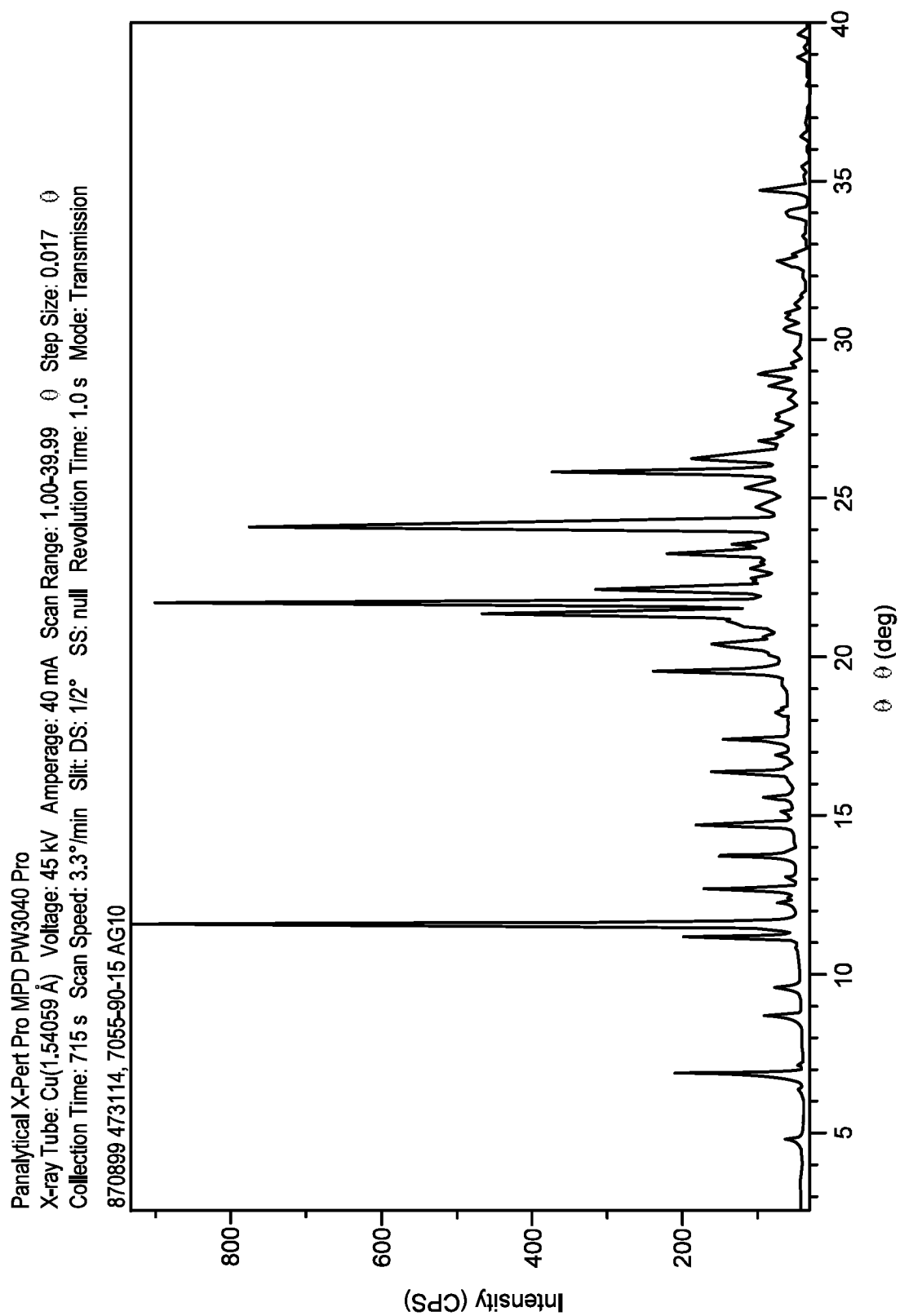


FIG. 17

22 / 53

**FIG. 18**

23 / 53

**FIG. 19**

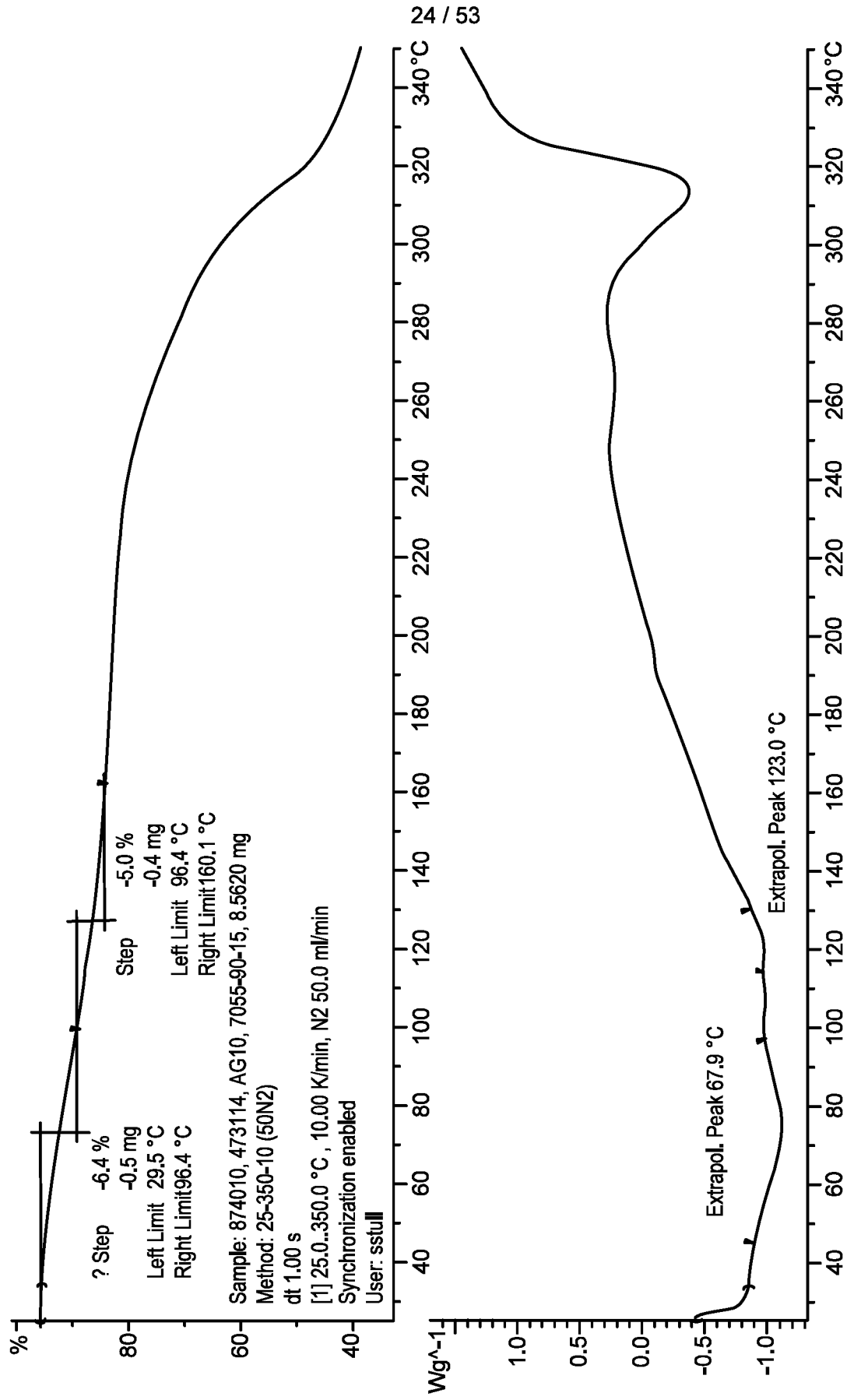
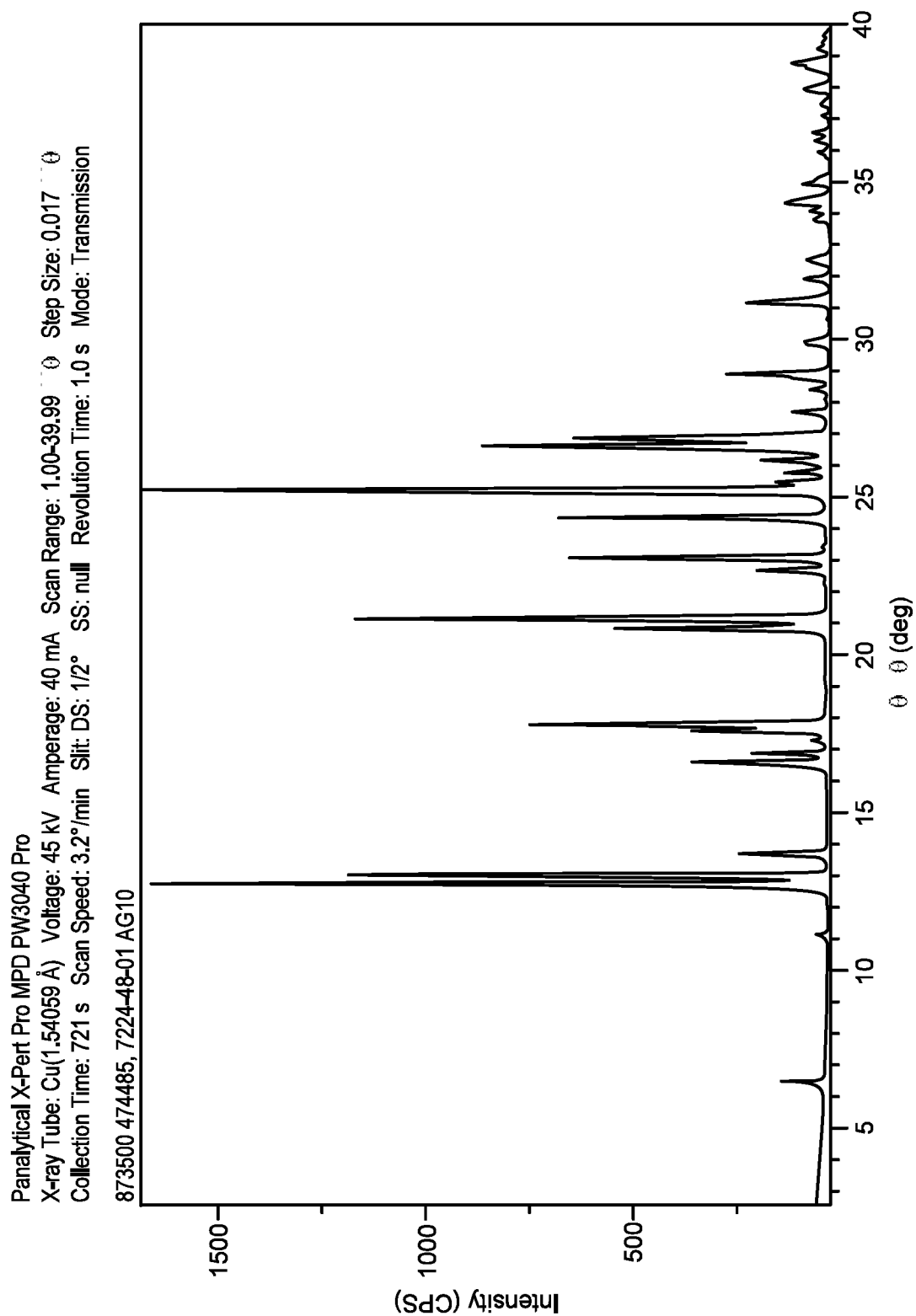


FIG. 20

25 / 53

**FIG. 21**

26 / 53

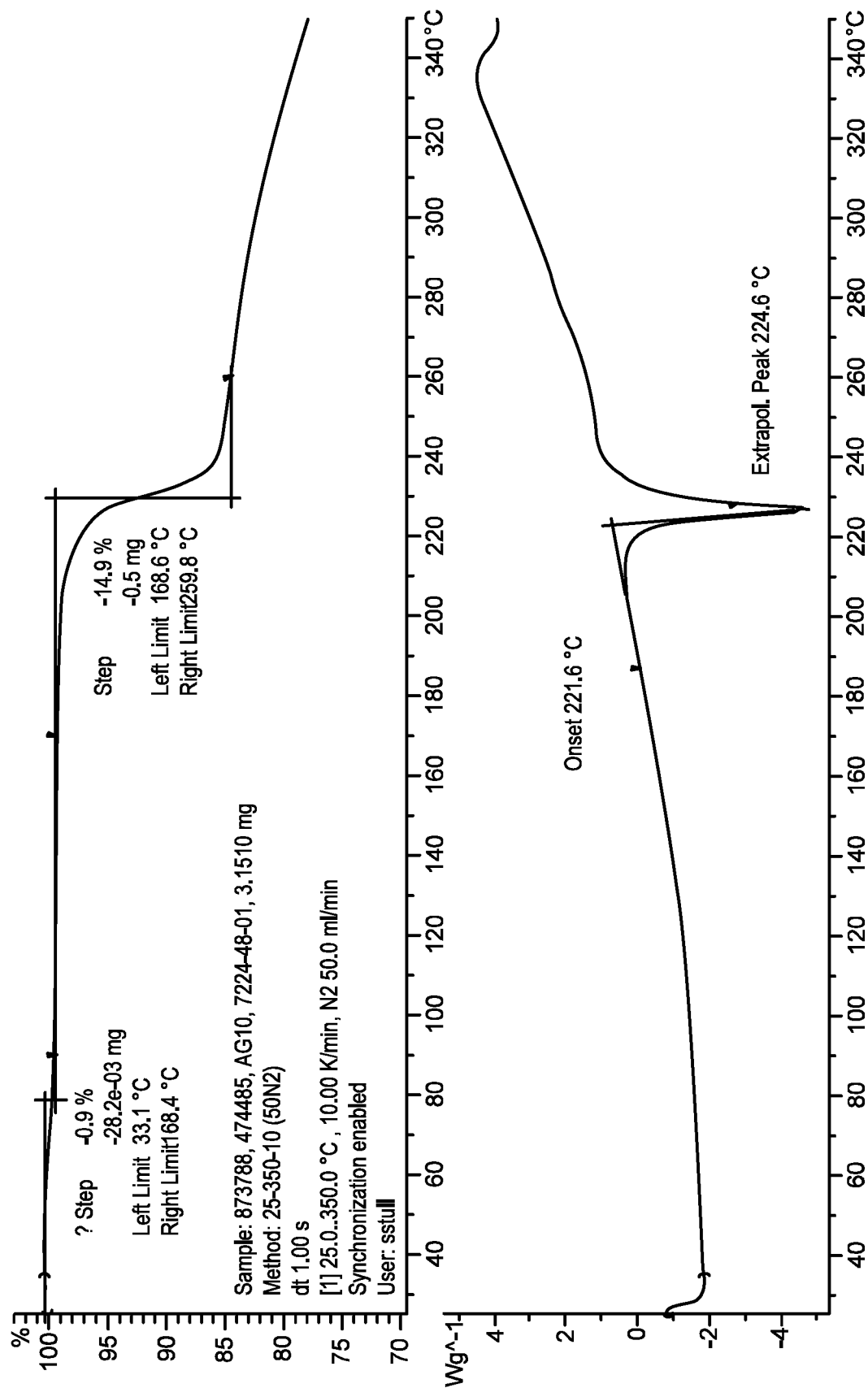
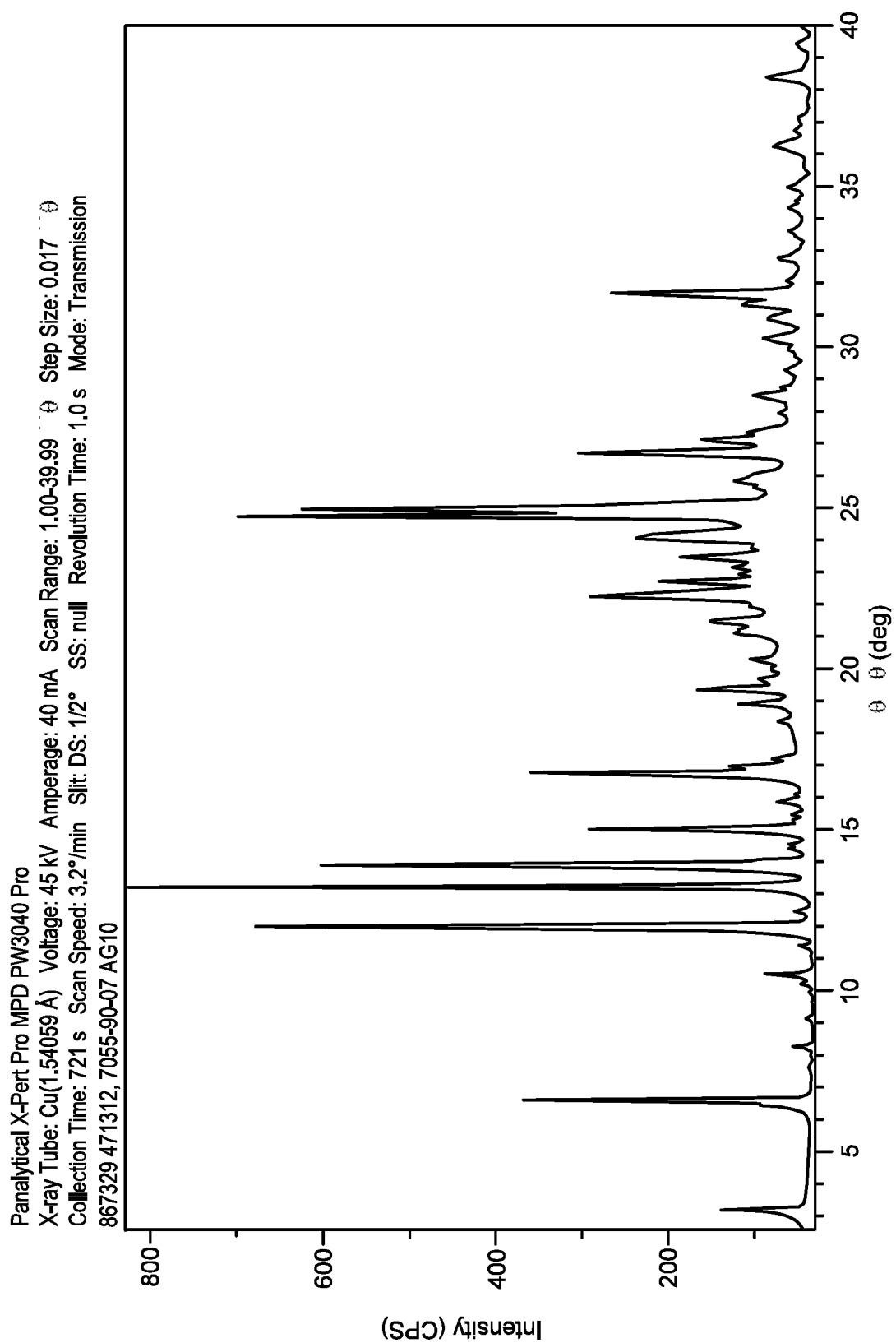
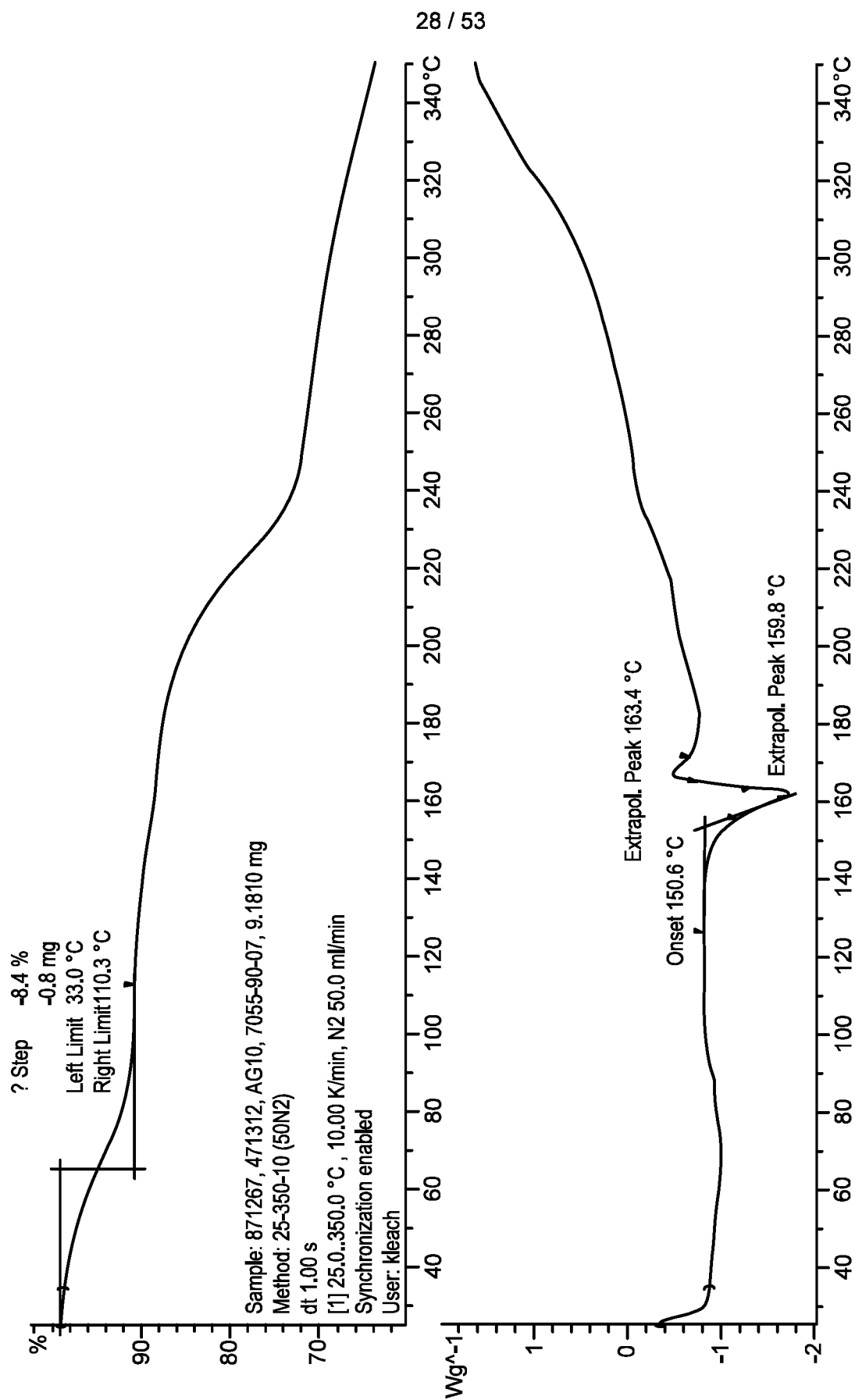


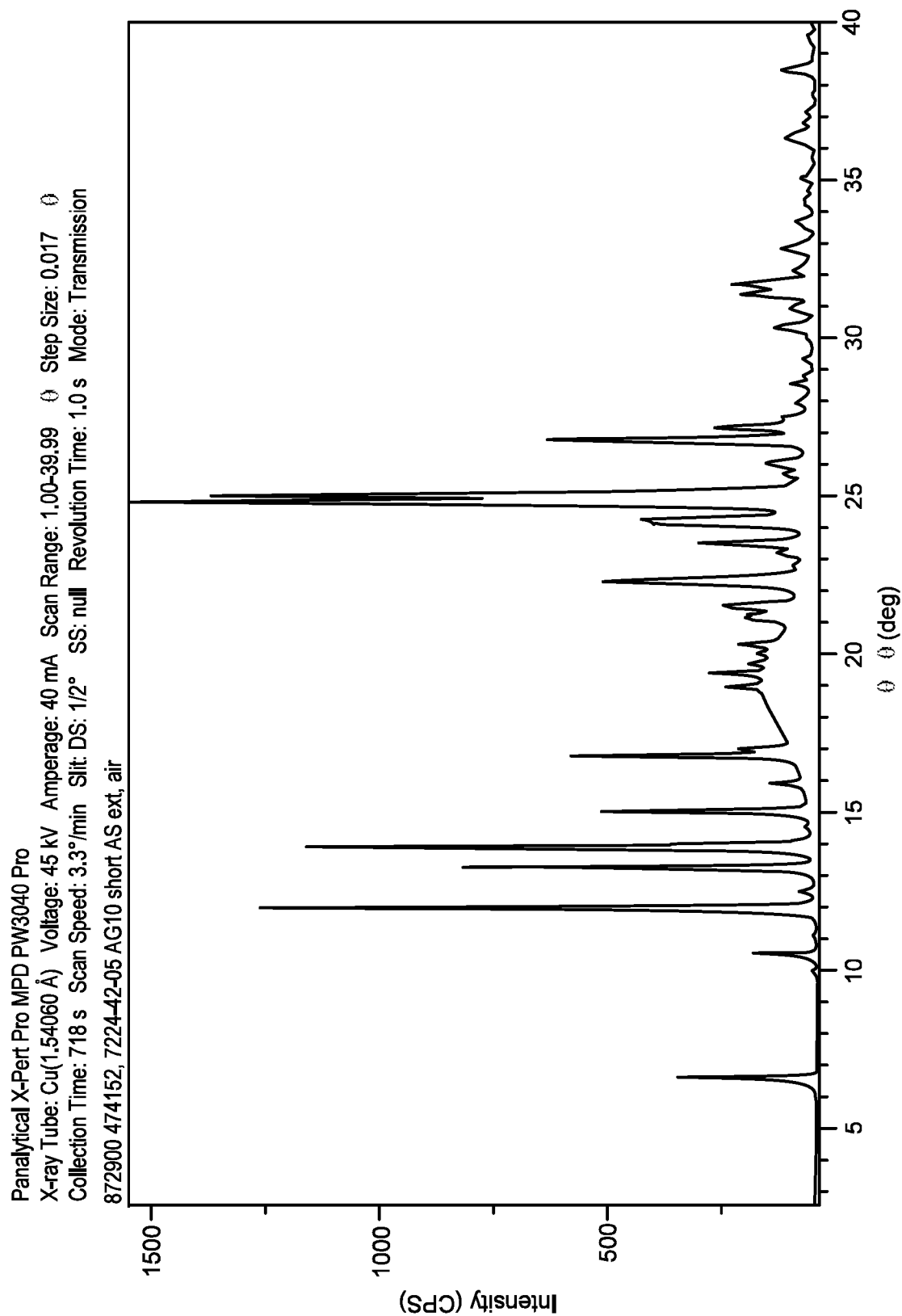
FIG. 22

27 / 53

**FIG. 23**

**FIG. 24**

29 / 53

**FIG. 25**

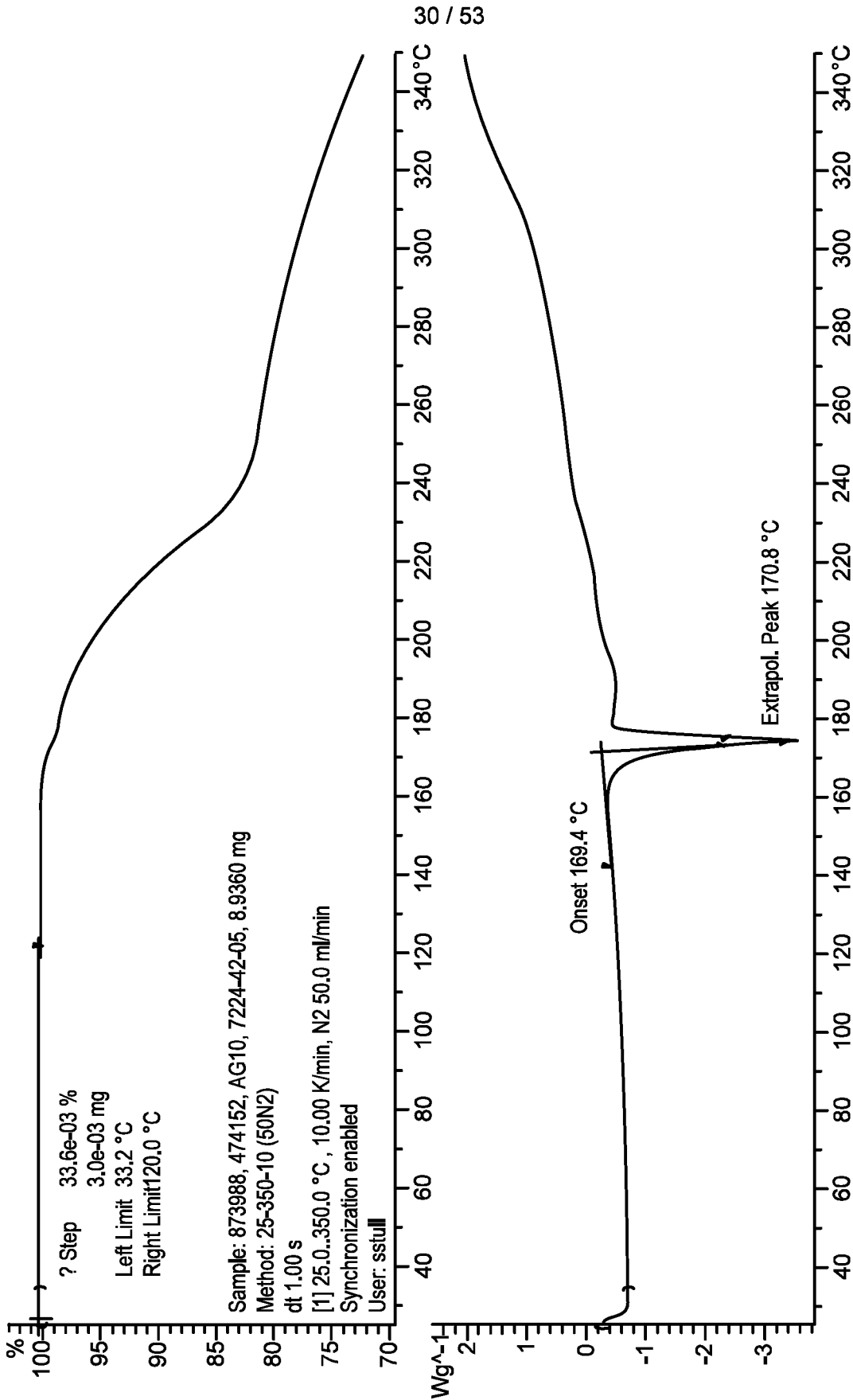
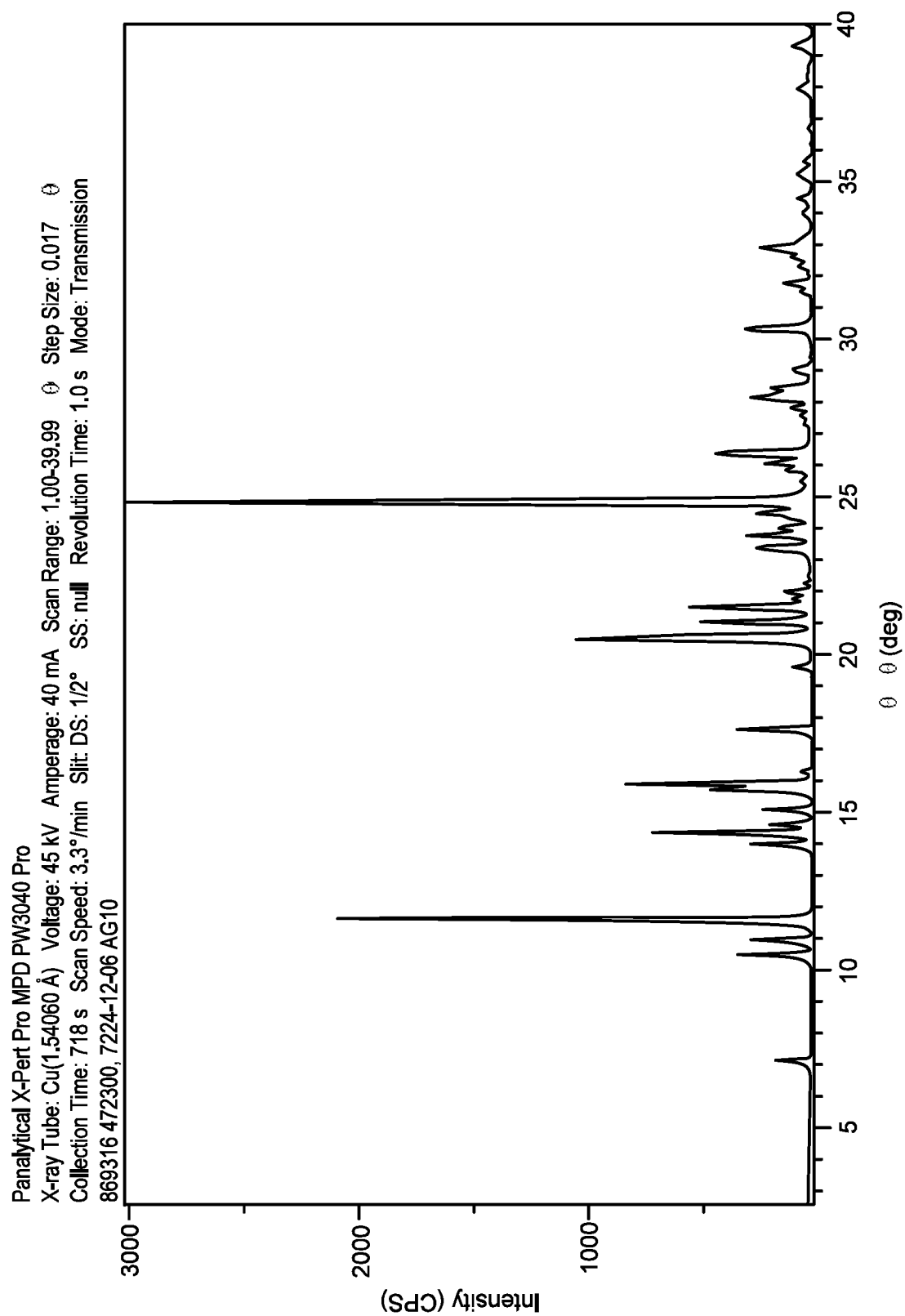


FIG. 26

31 / 53

**FIG. 27**

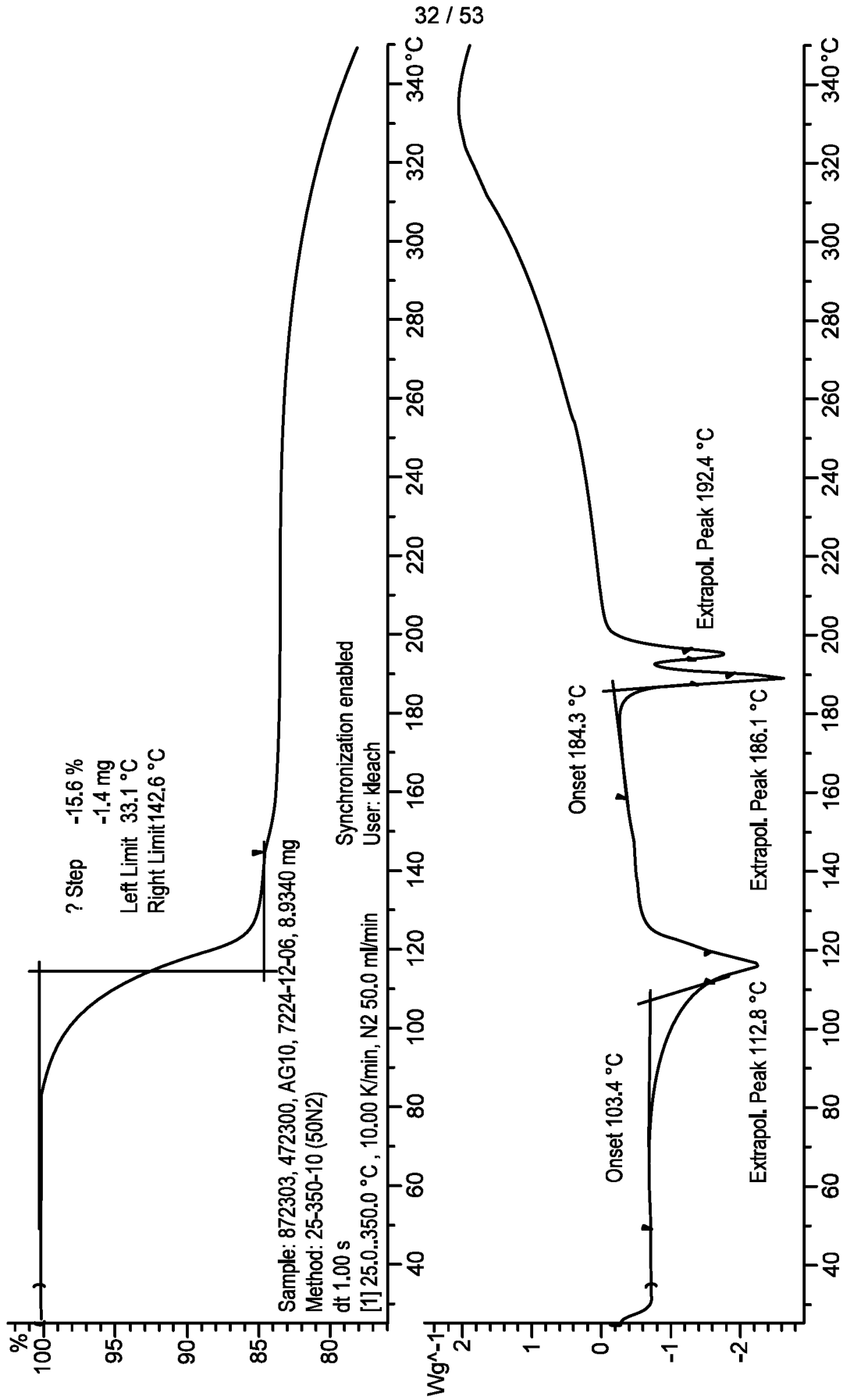
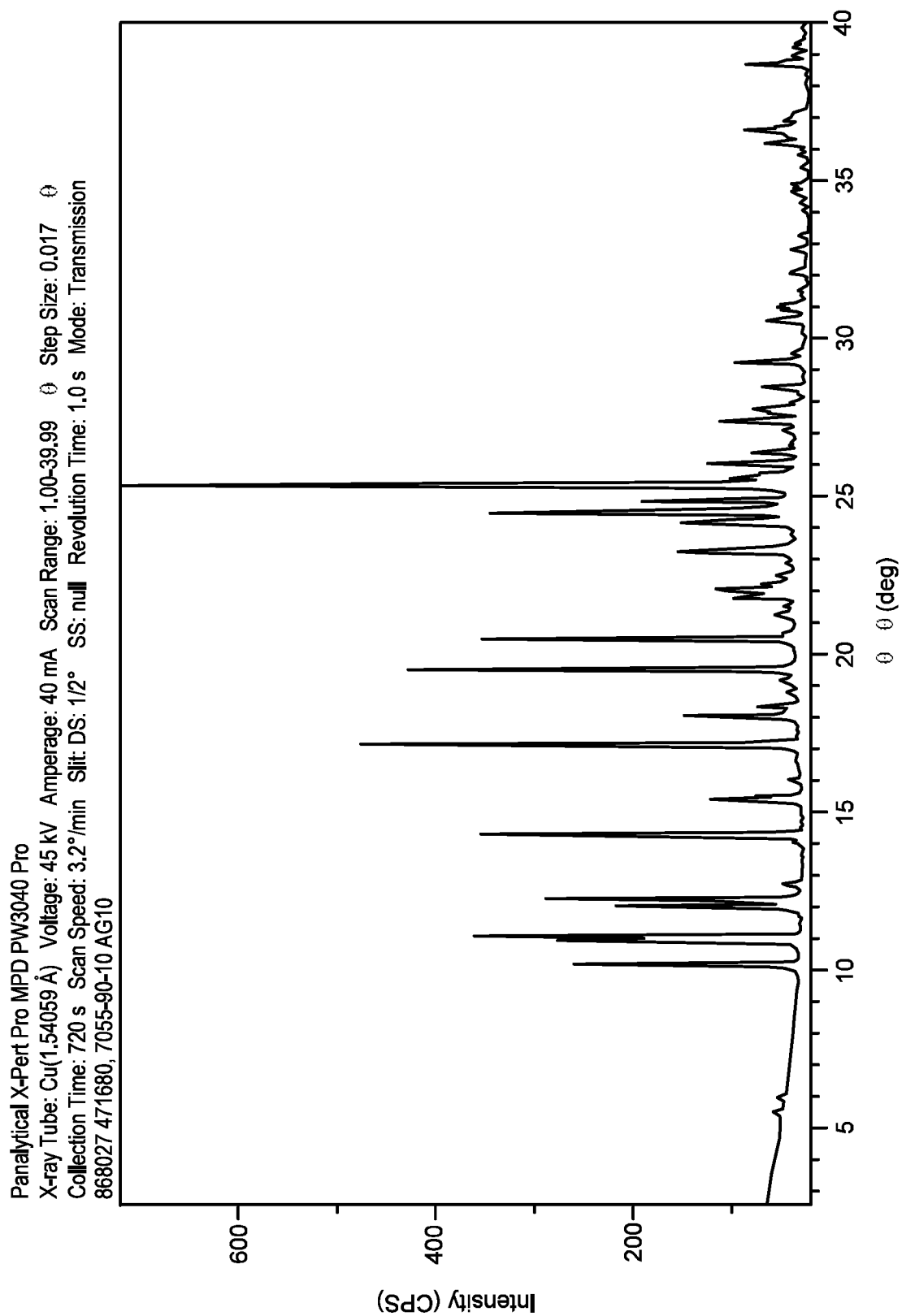


FIG. 28

33 / 53

**FIG. 29**

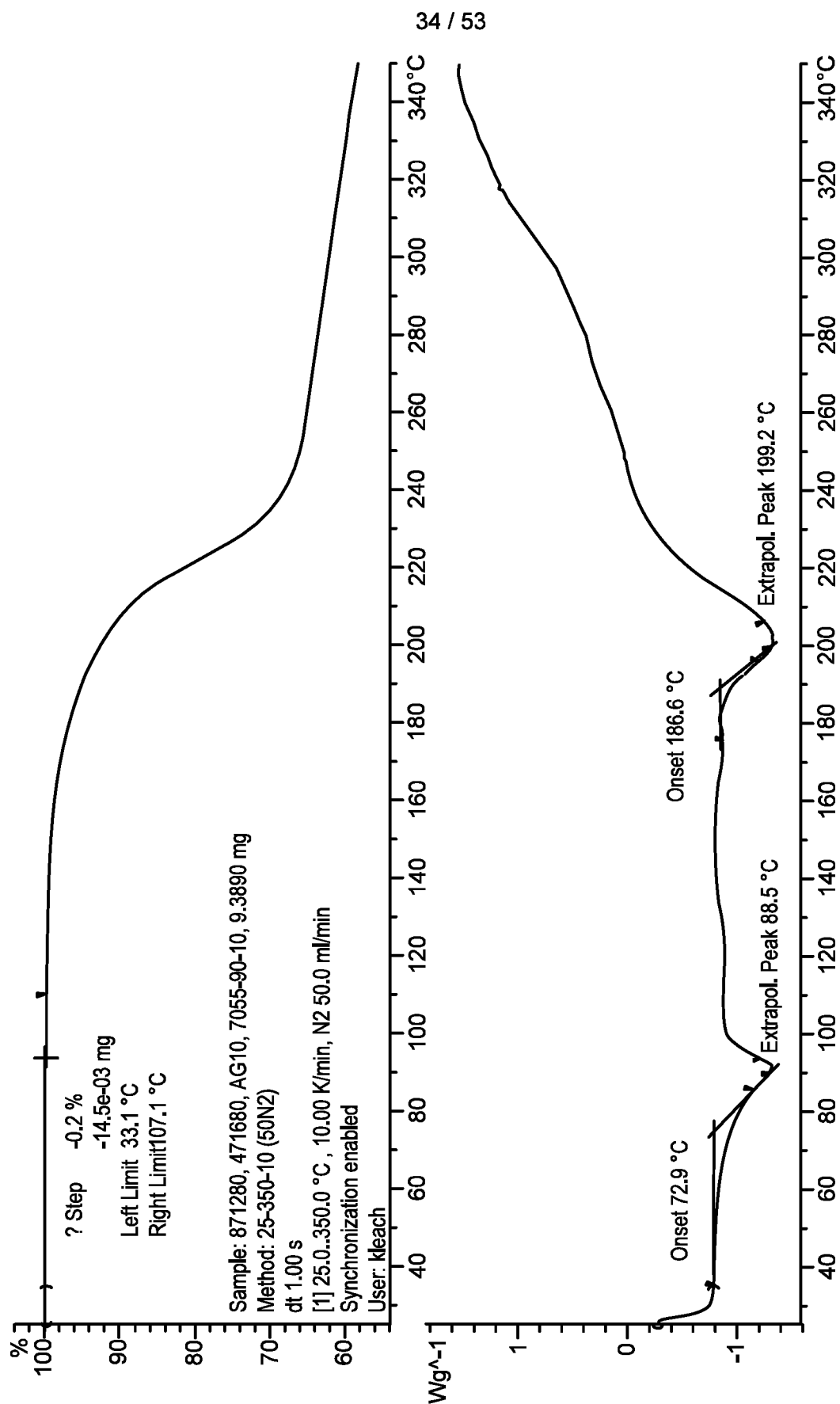
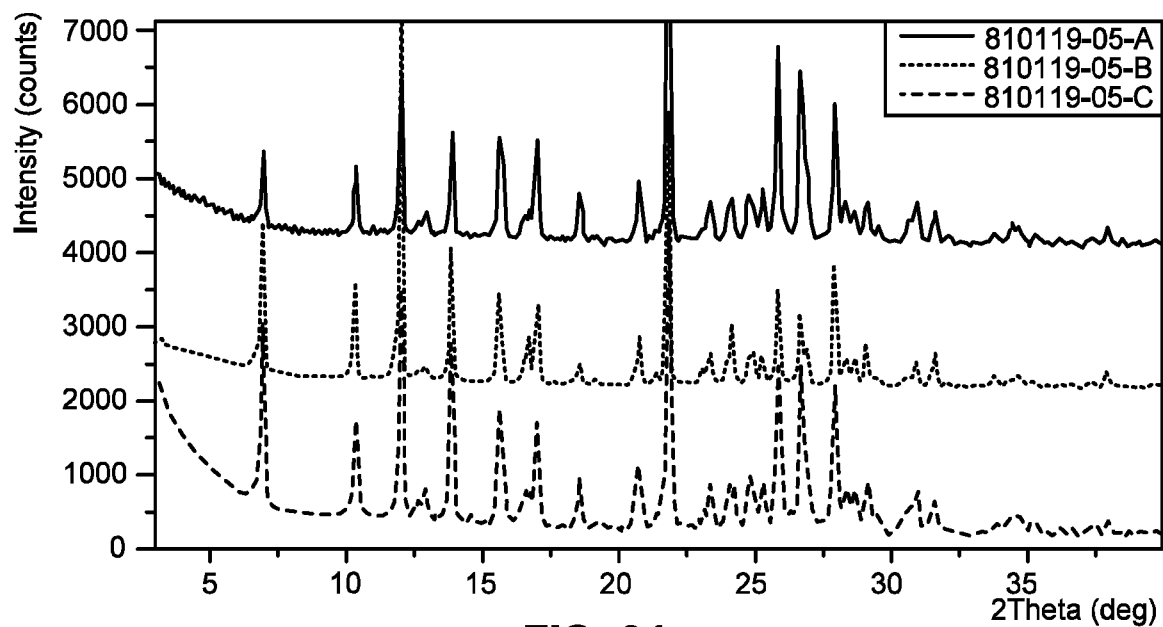
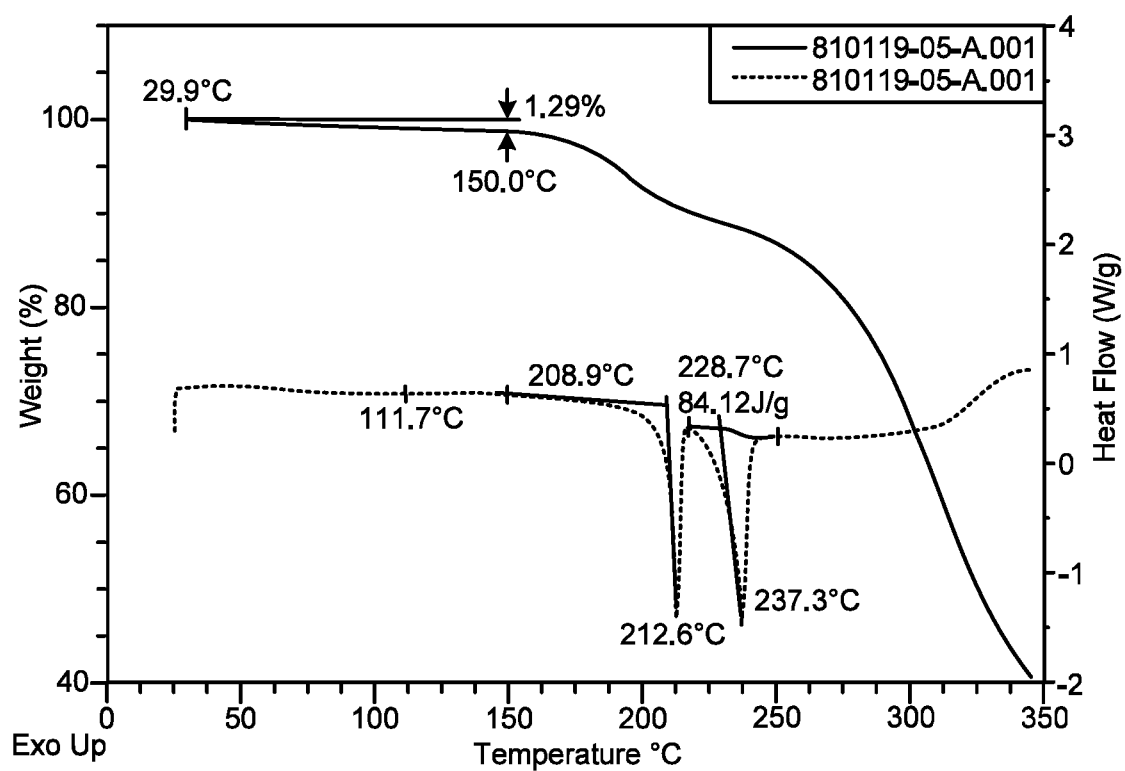
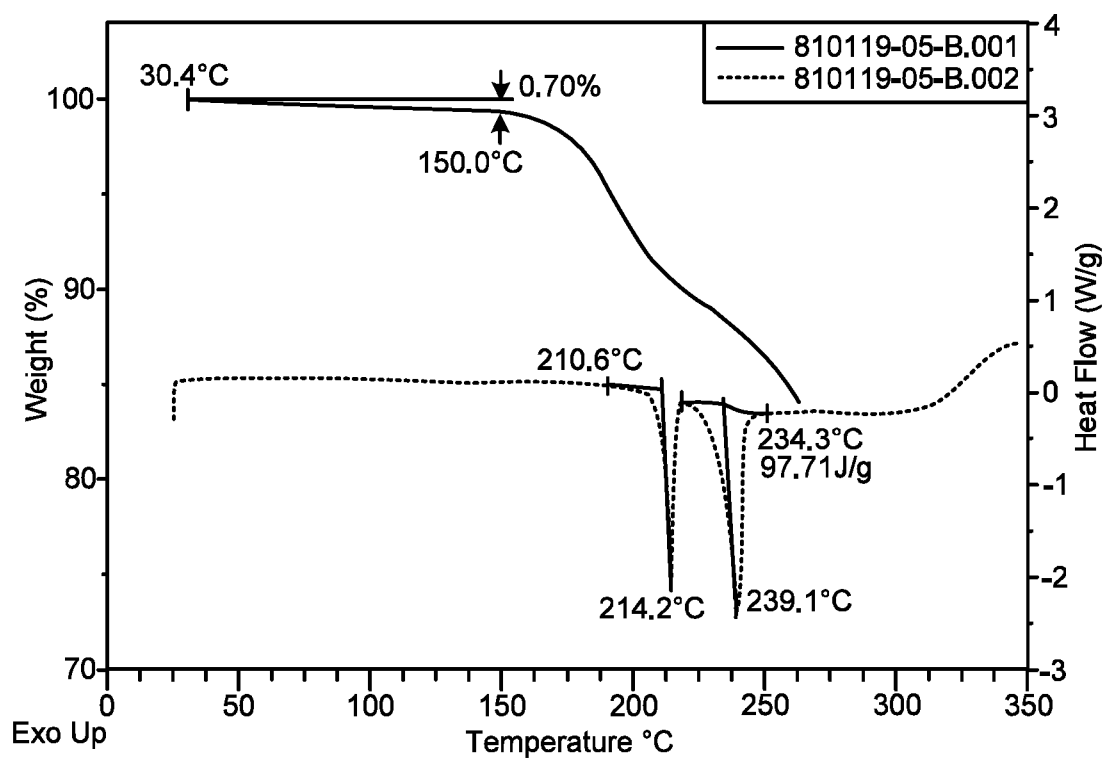
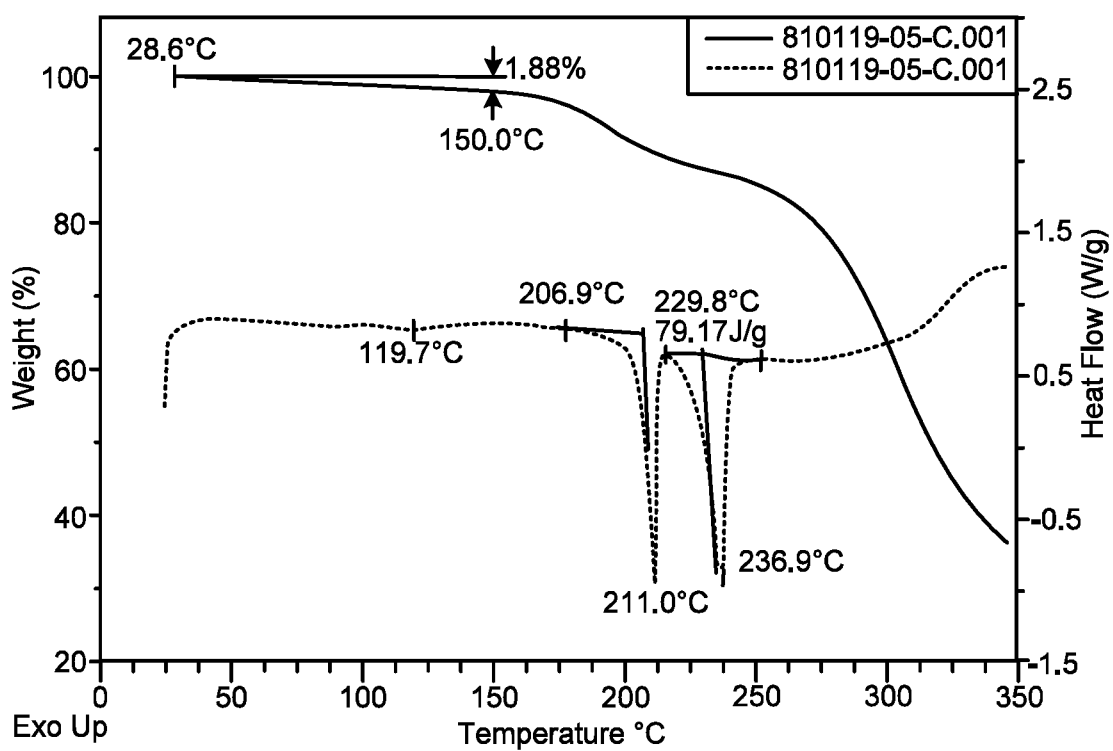


FIG. 30

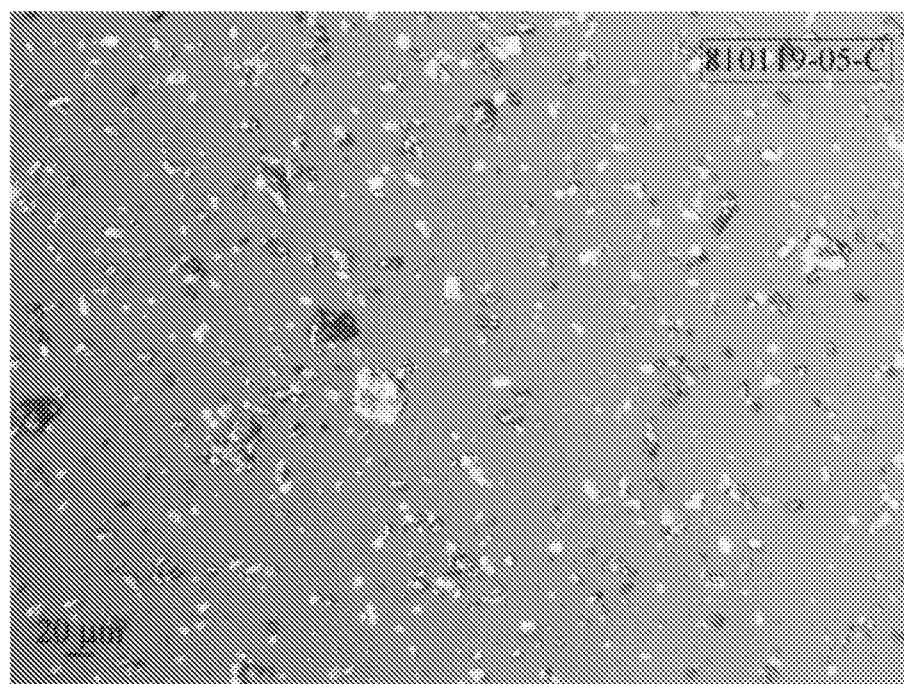
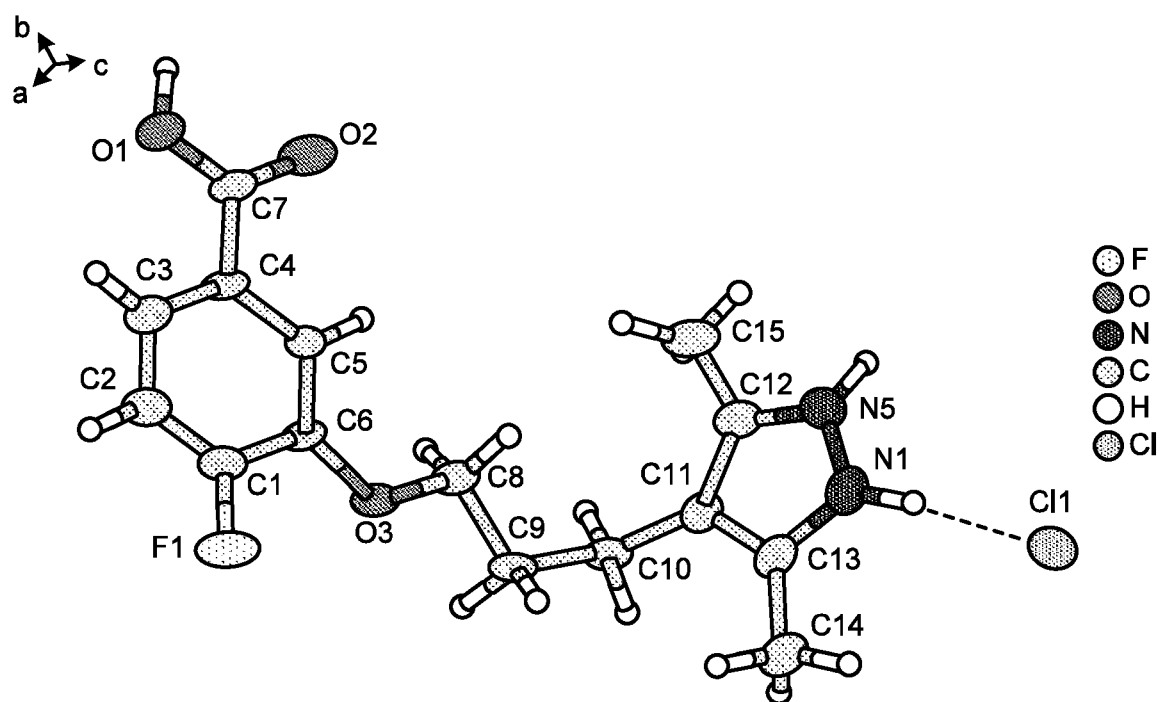
35 / 53

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

36 / 53

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

37 / 53

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

38 / 53

DVS Isotherm Plot

Date: 16 Mar 2017

Time: 13-31

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

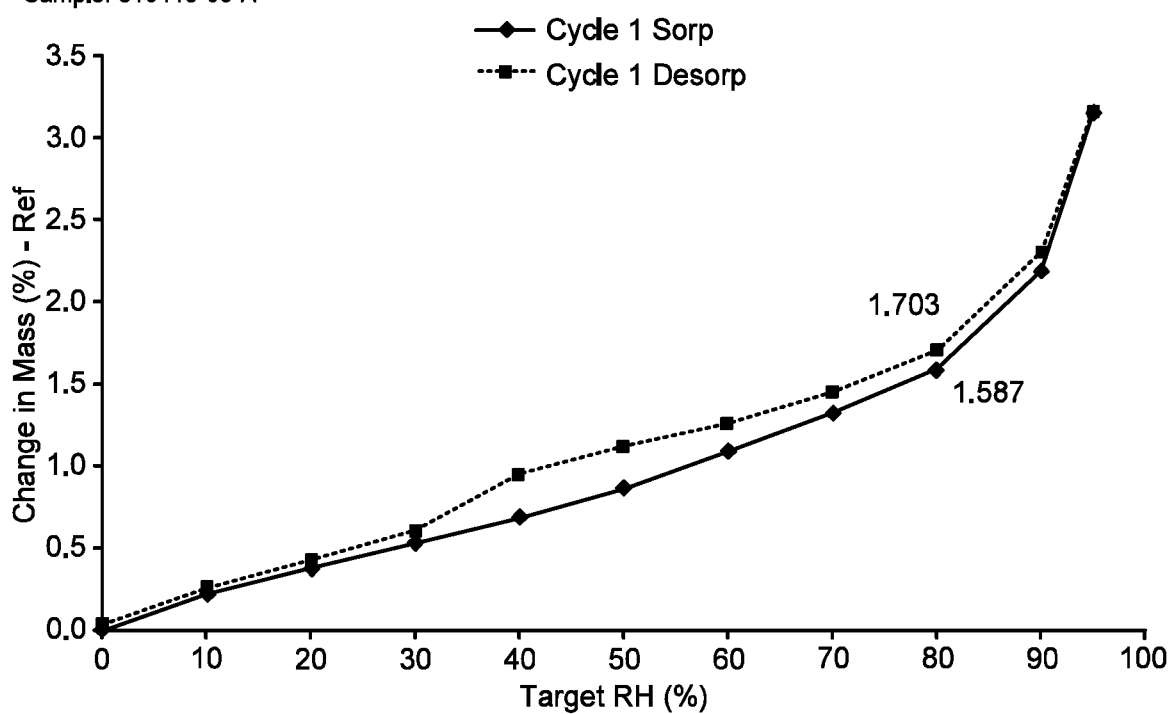
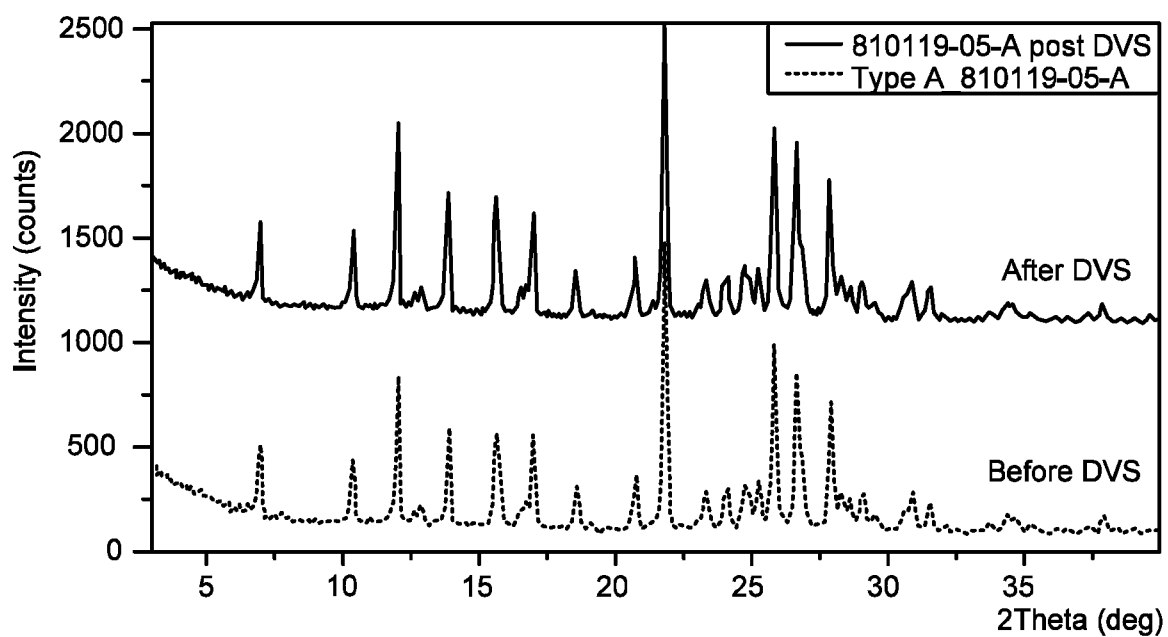
Sample: 810119-05-A

Temp: 25.0 °C

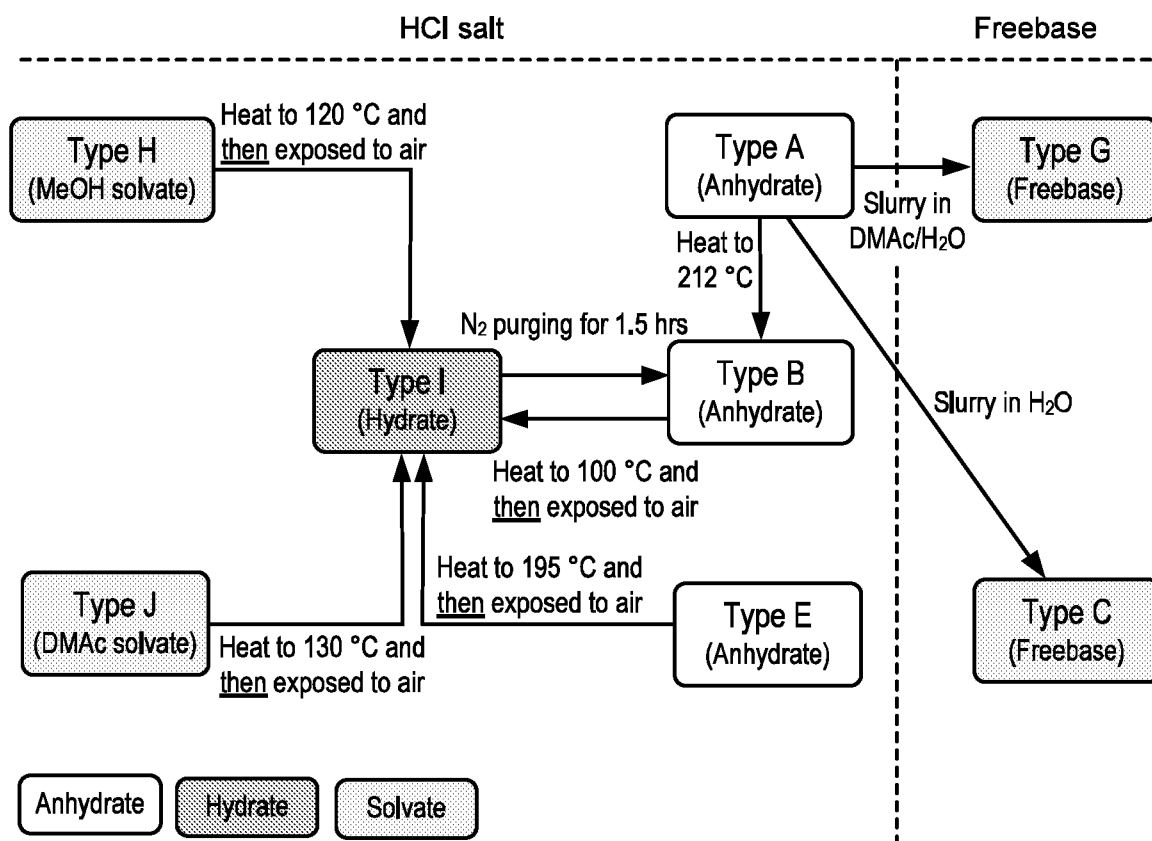
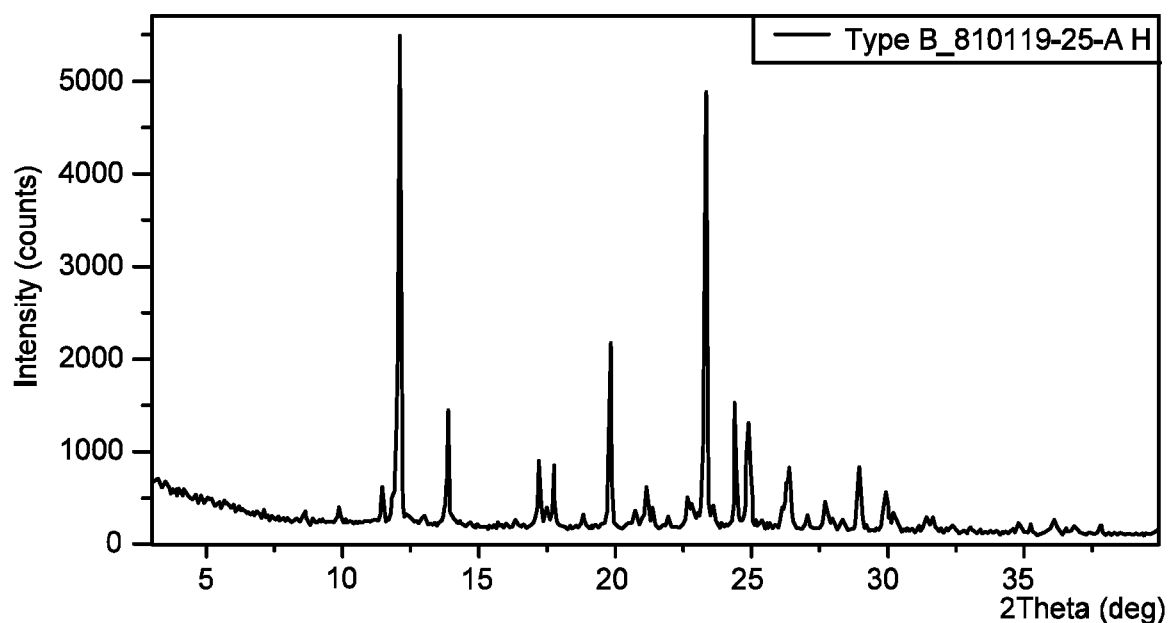
Meth: 0-95-0 CP3677 LONG

METHOD.sao

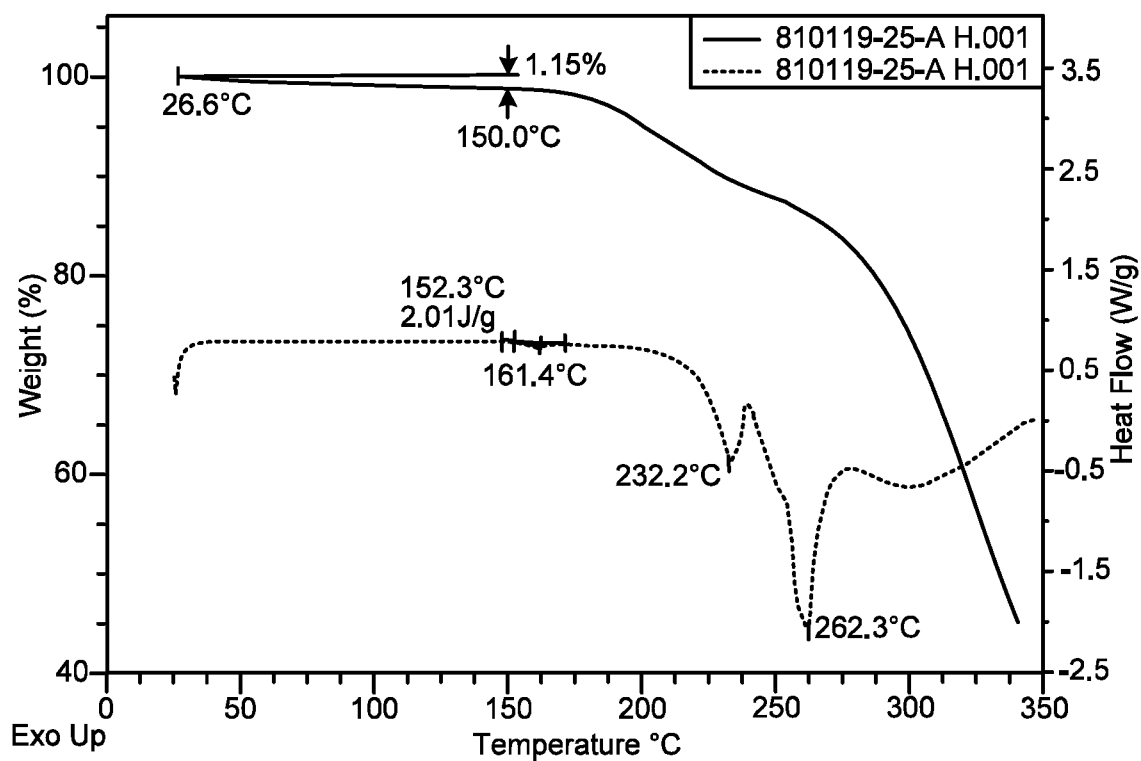
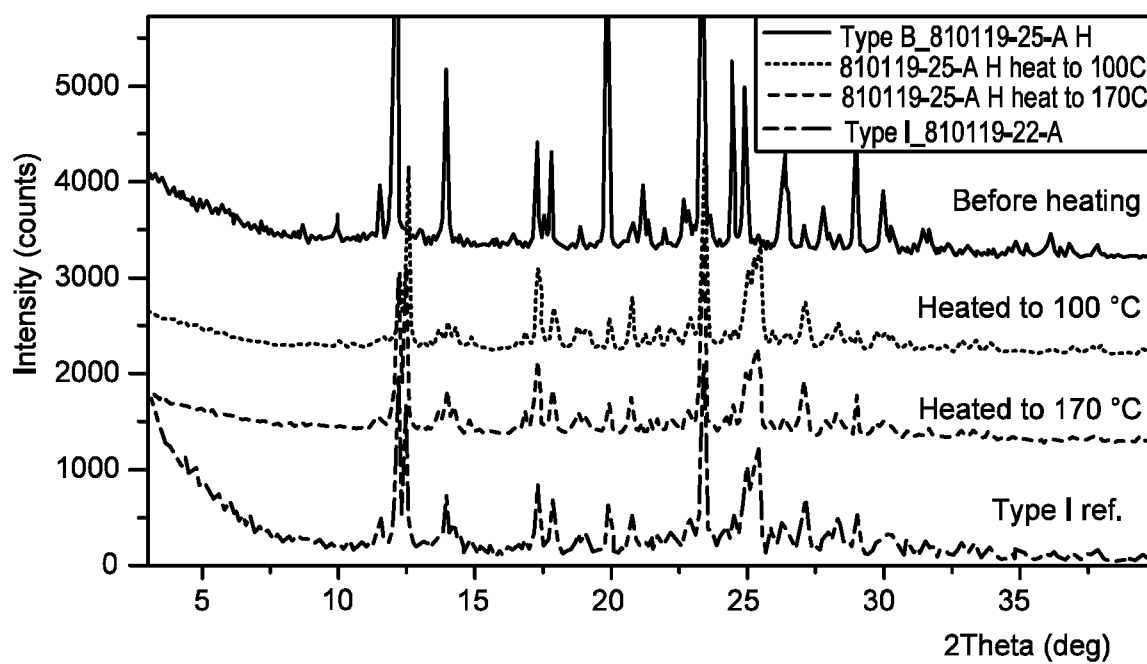
Mref: 26.8199

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

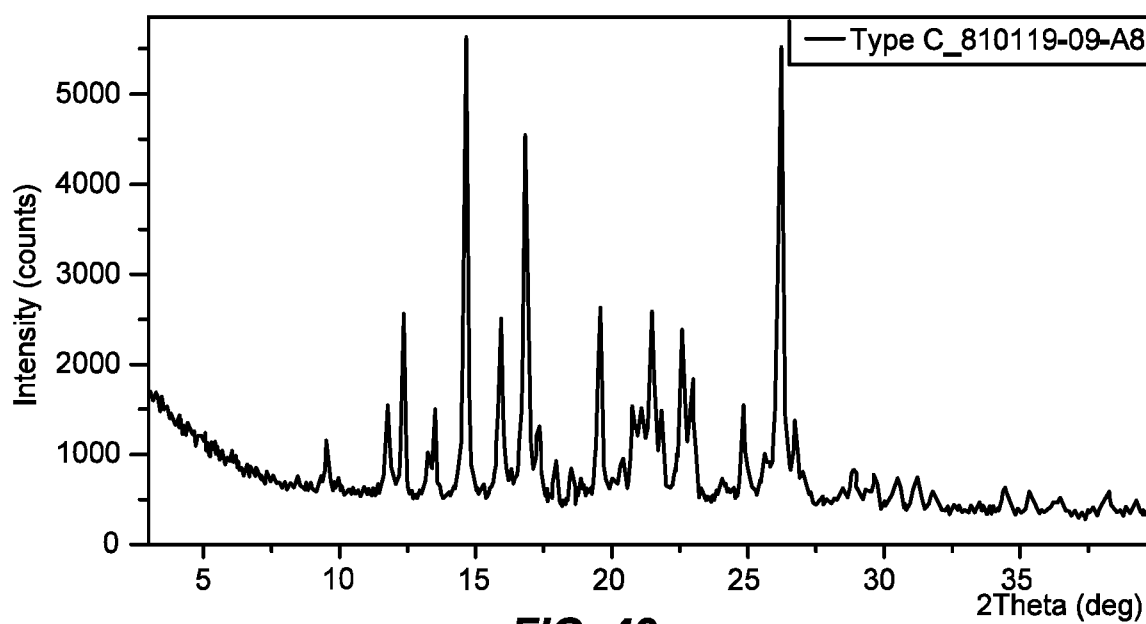
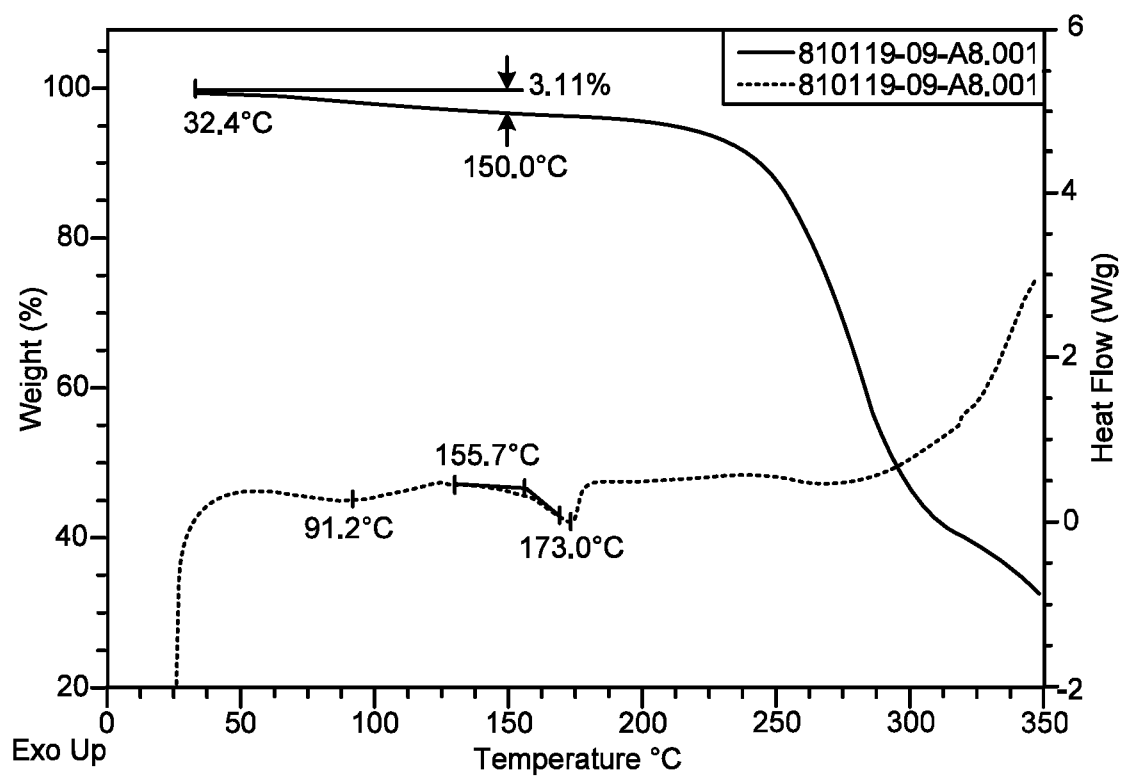
39 / 53

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

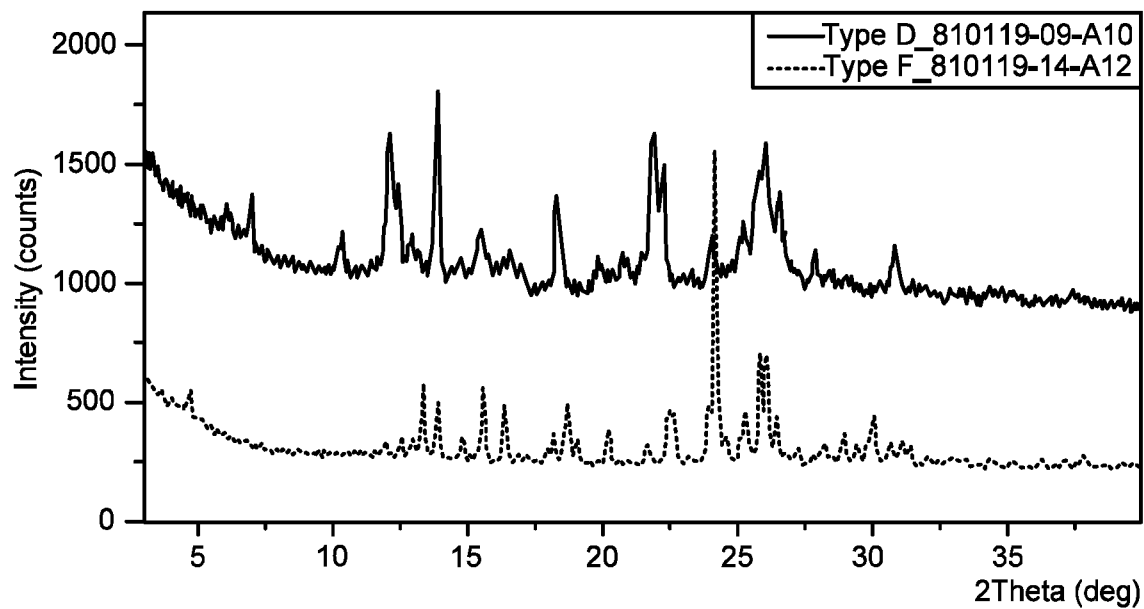
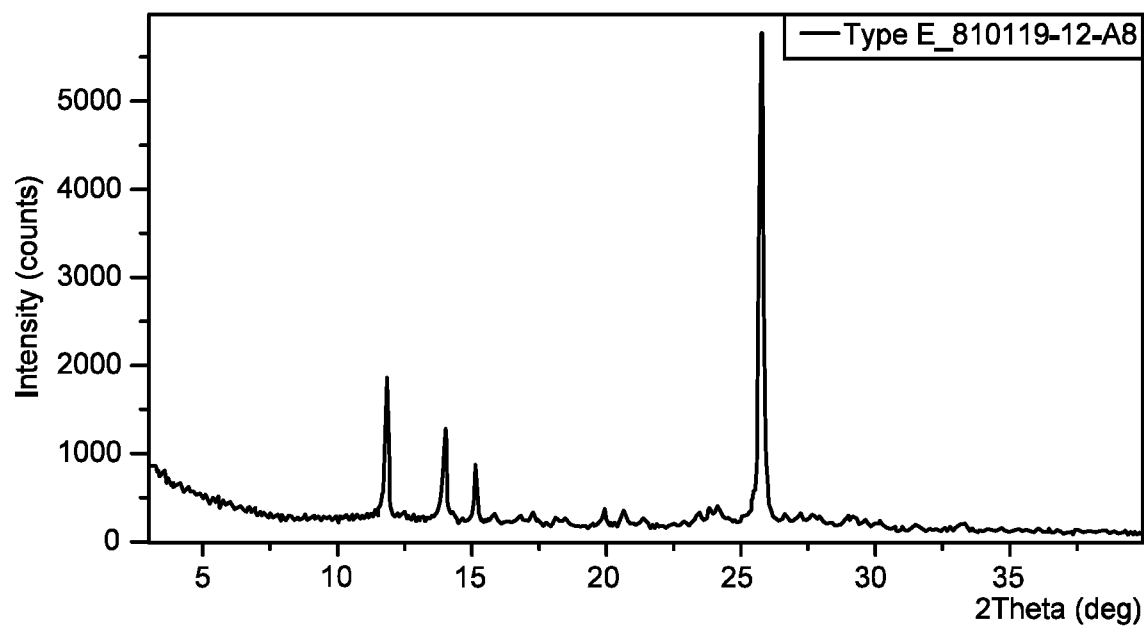
40 / 53

**FIG. 41****FIG. 42**

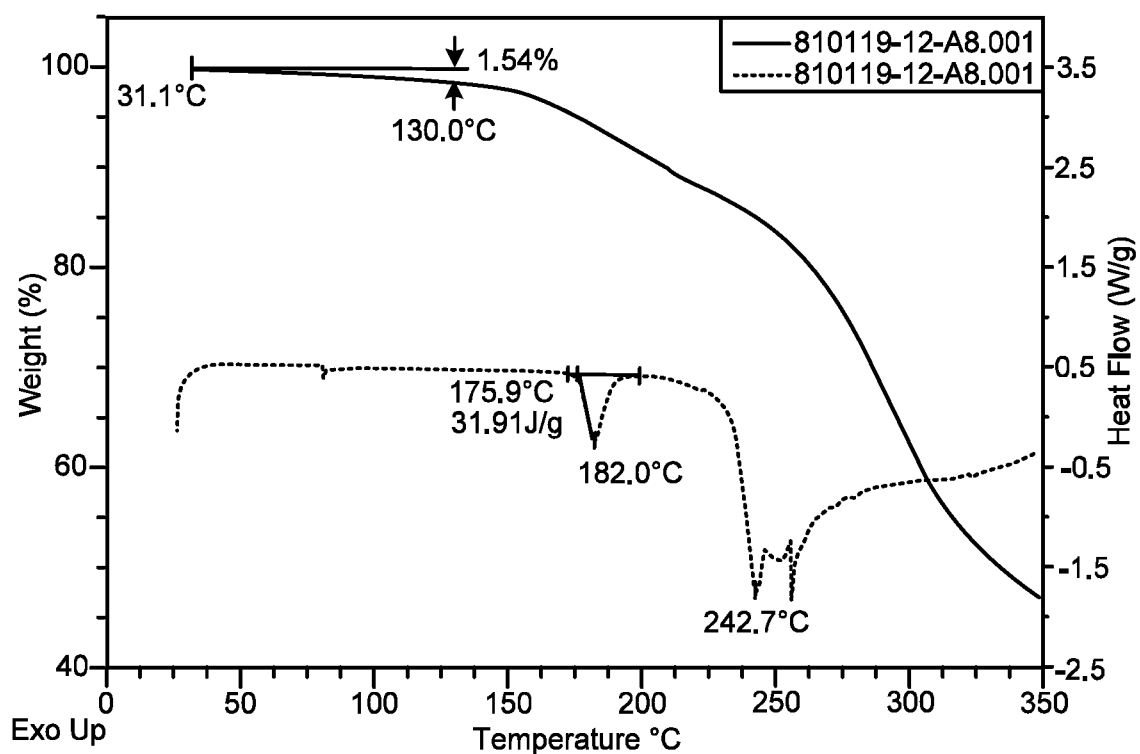
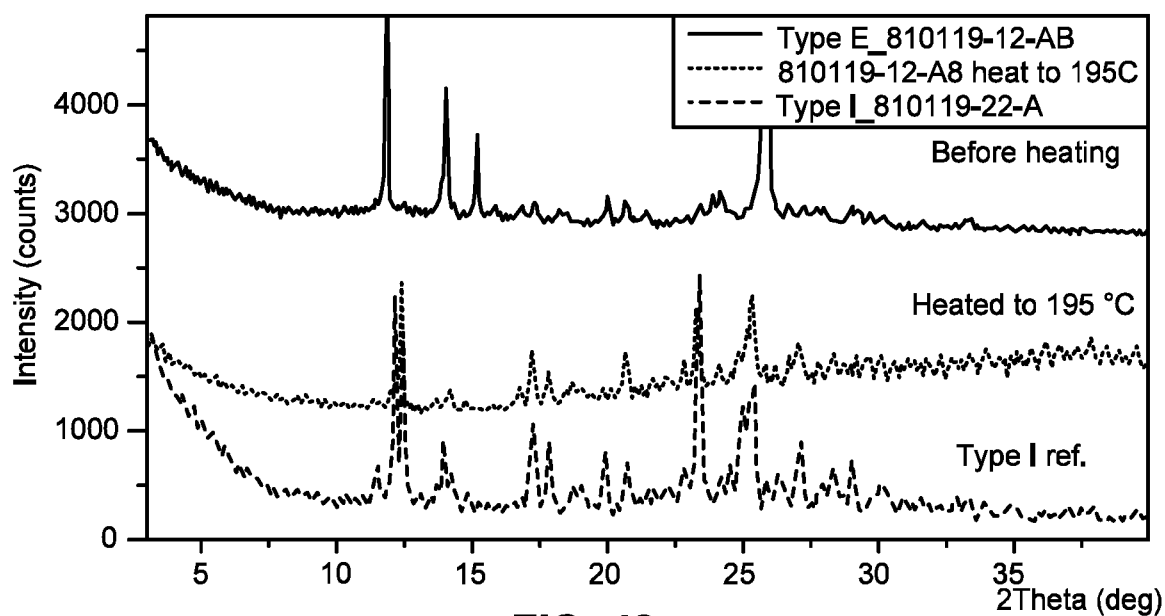
41 / 53

**FIG. 43****FIG. 44**

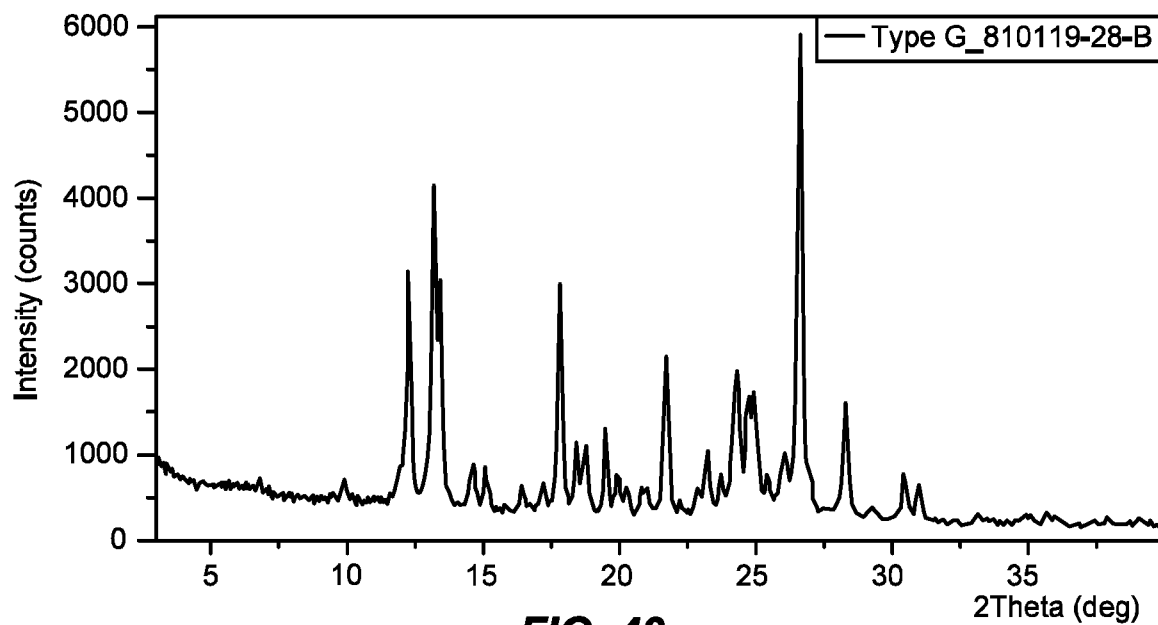
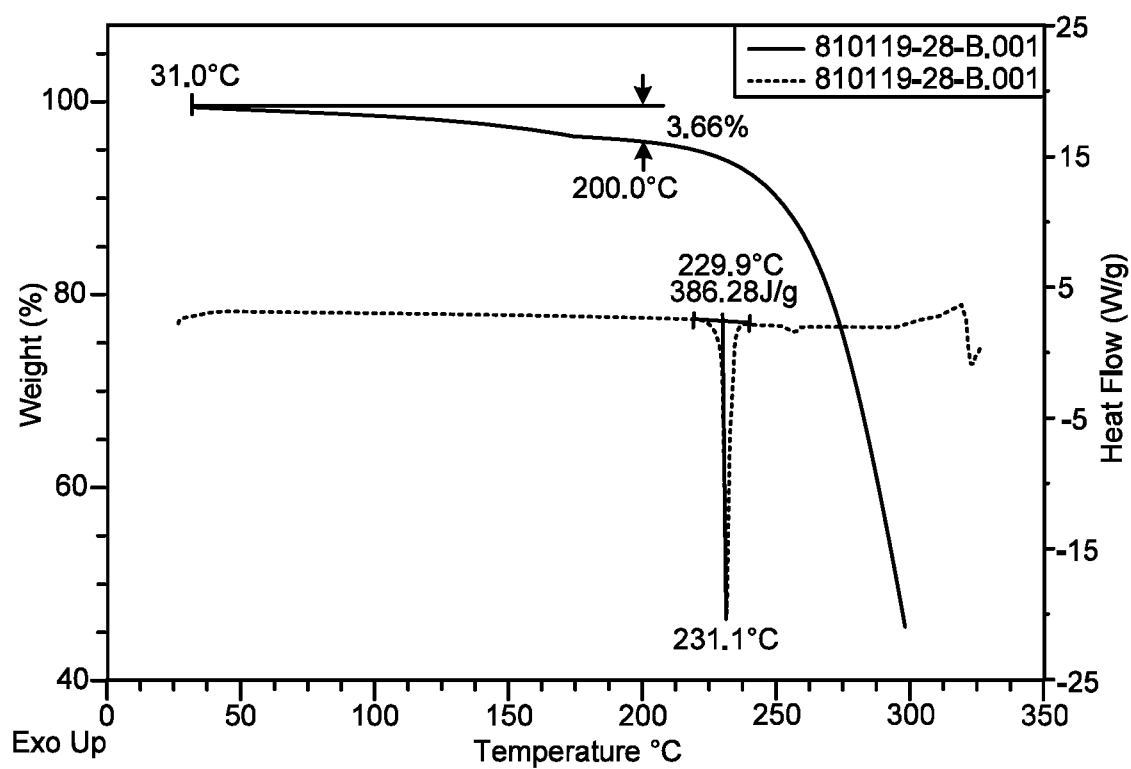
42 / 53

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

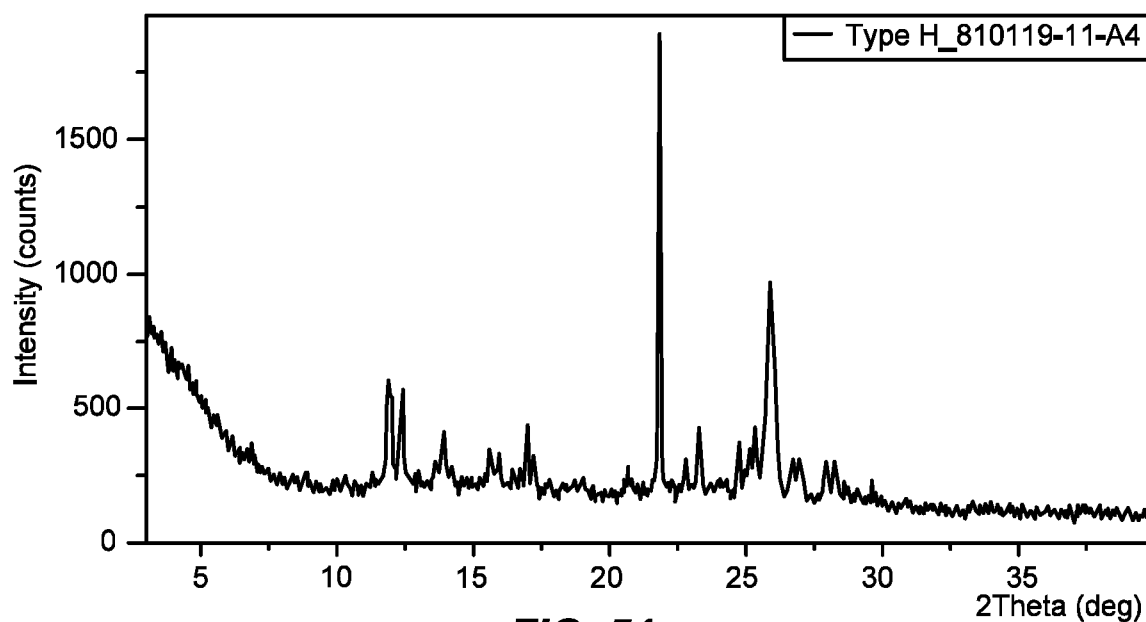
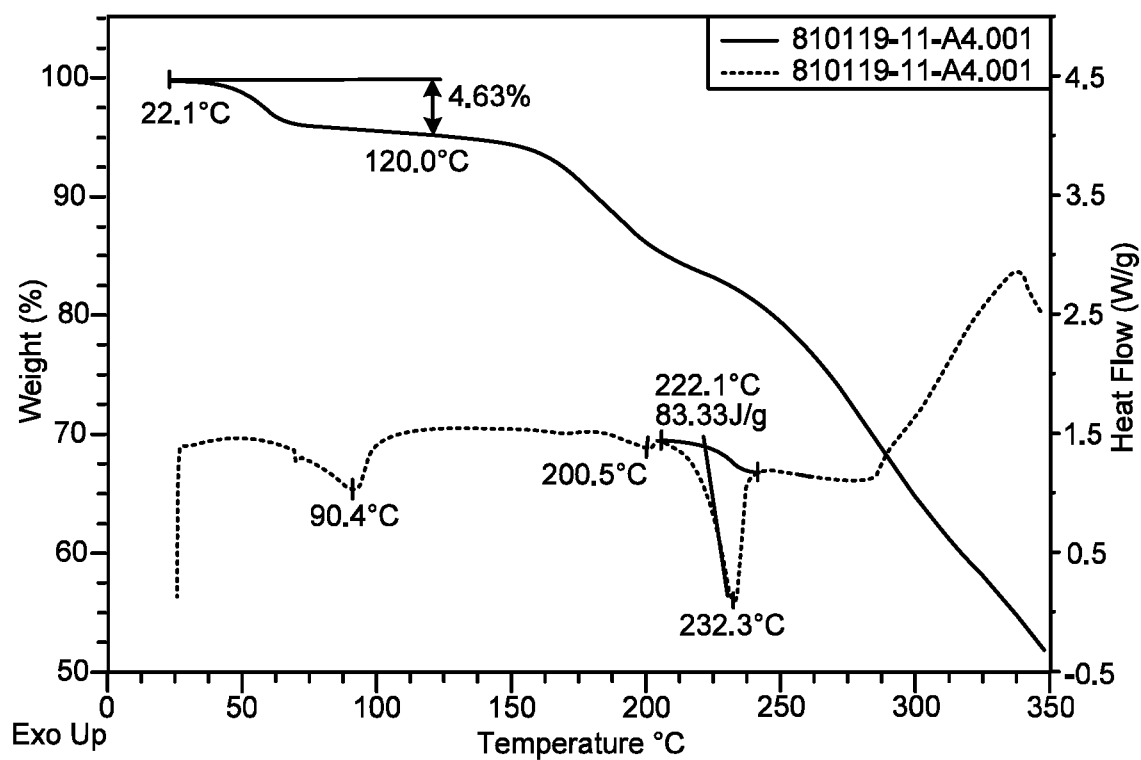
43 / 53

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

44 / 53

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

45 / 53

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

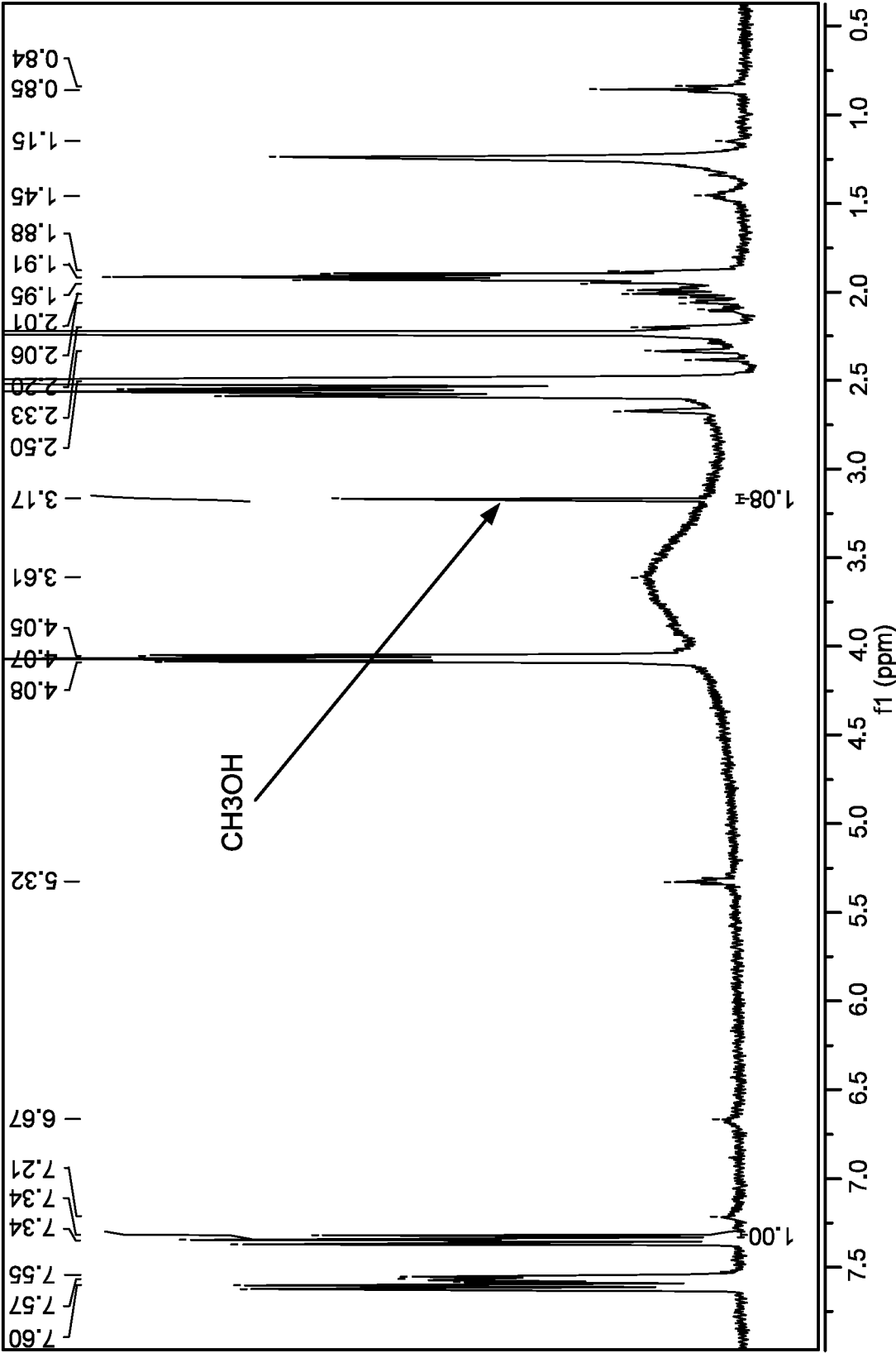


FIG. 53

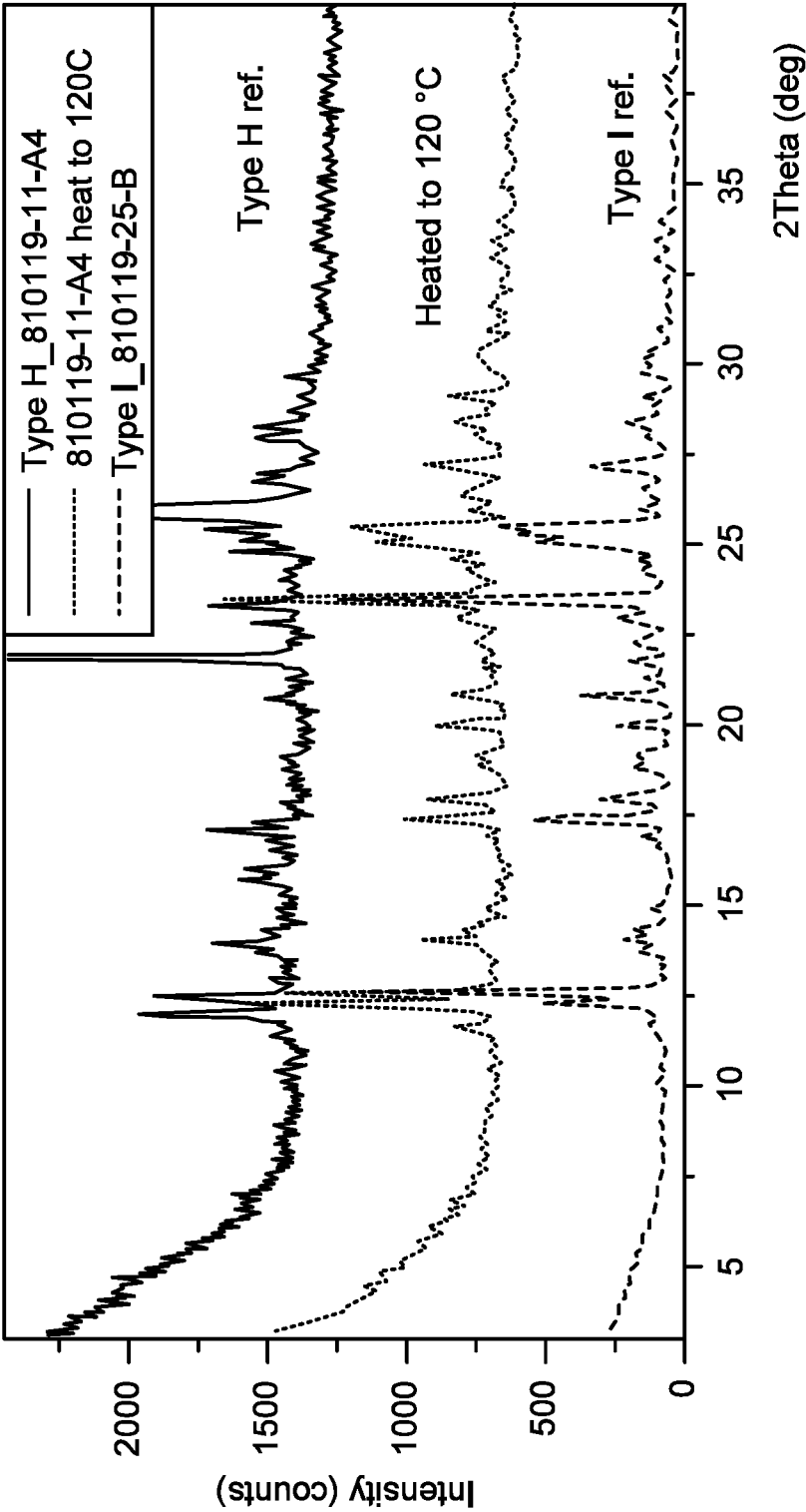
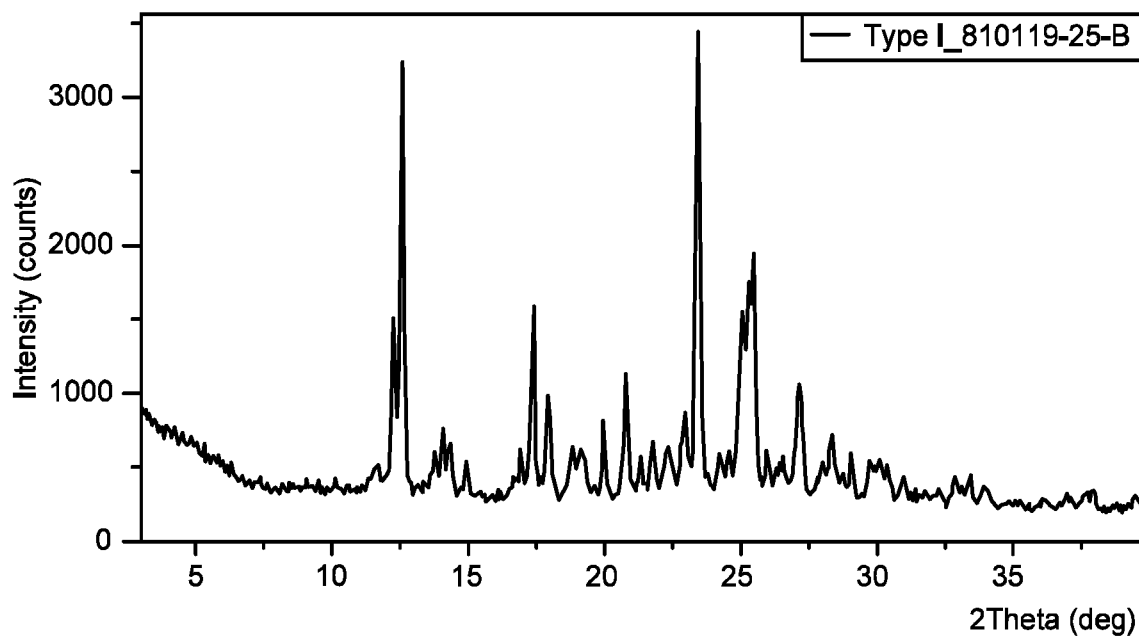
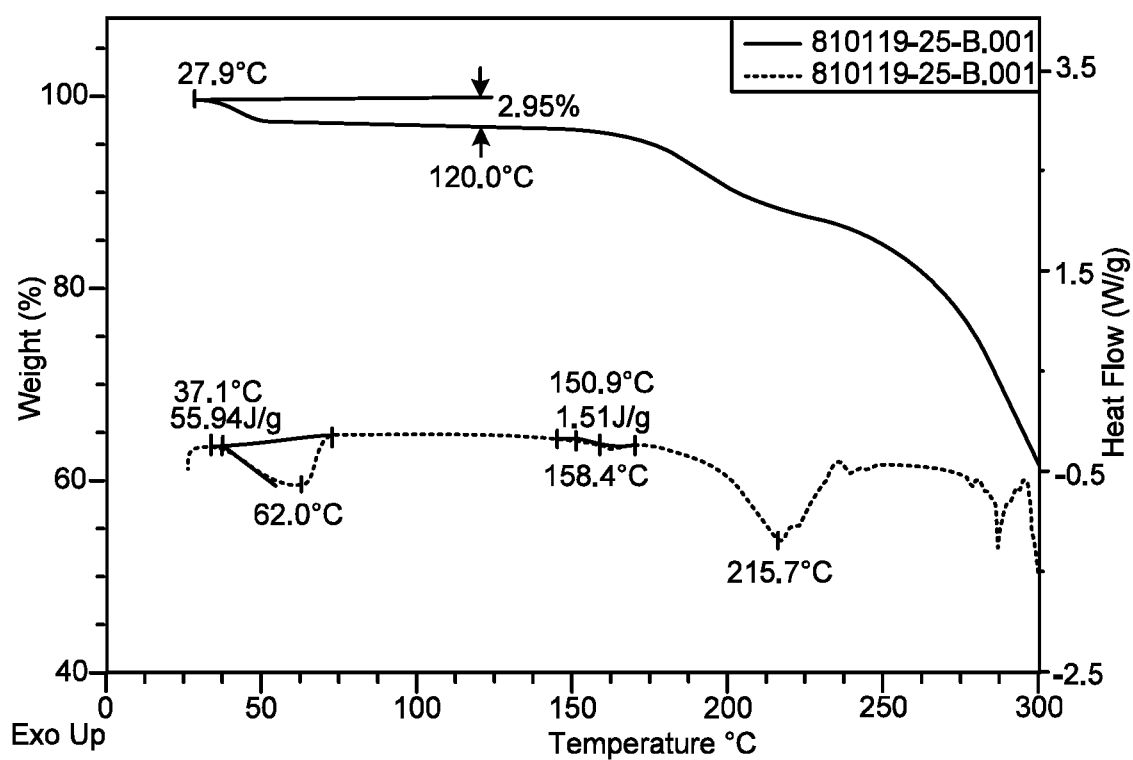
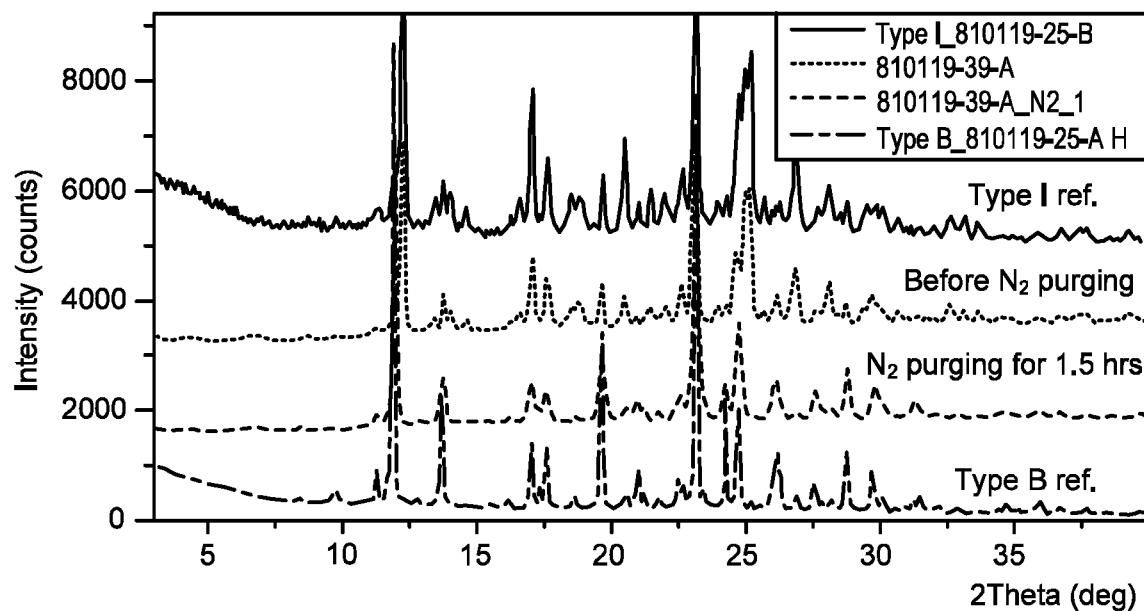
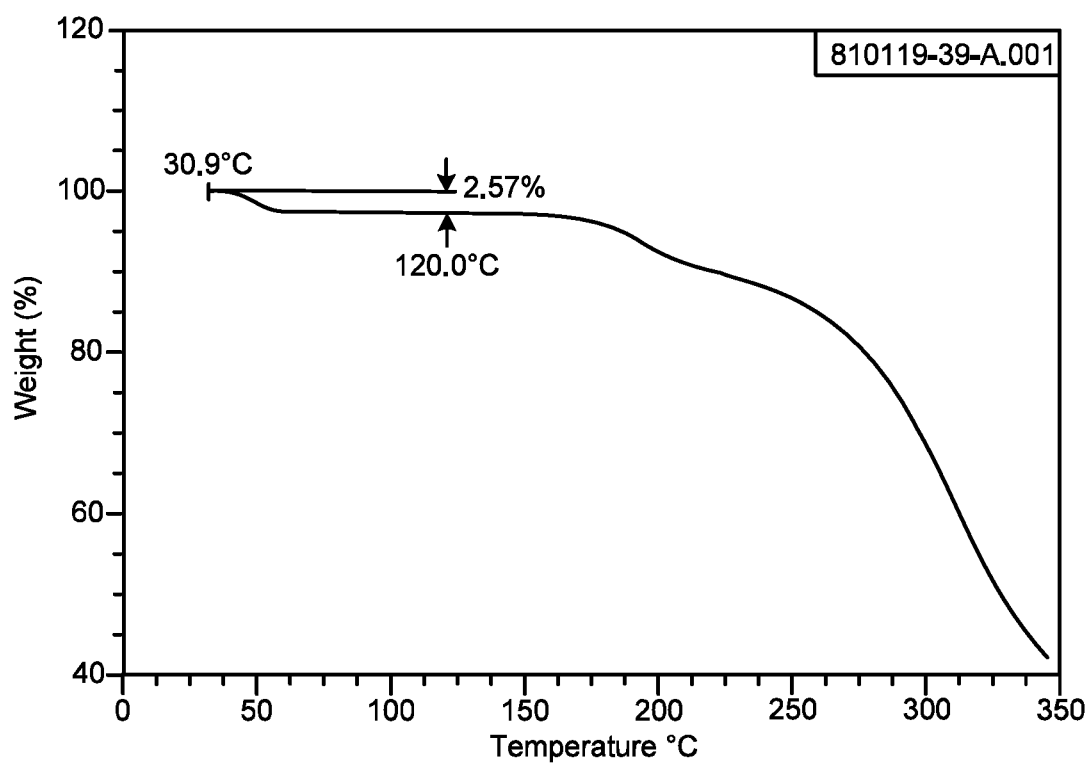


FIG. 54

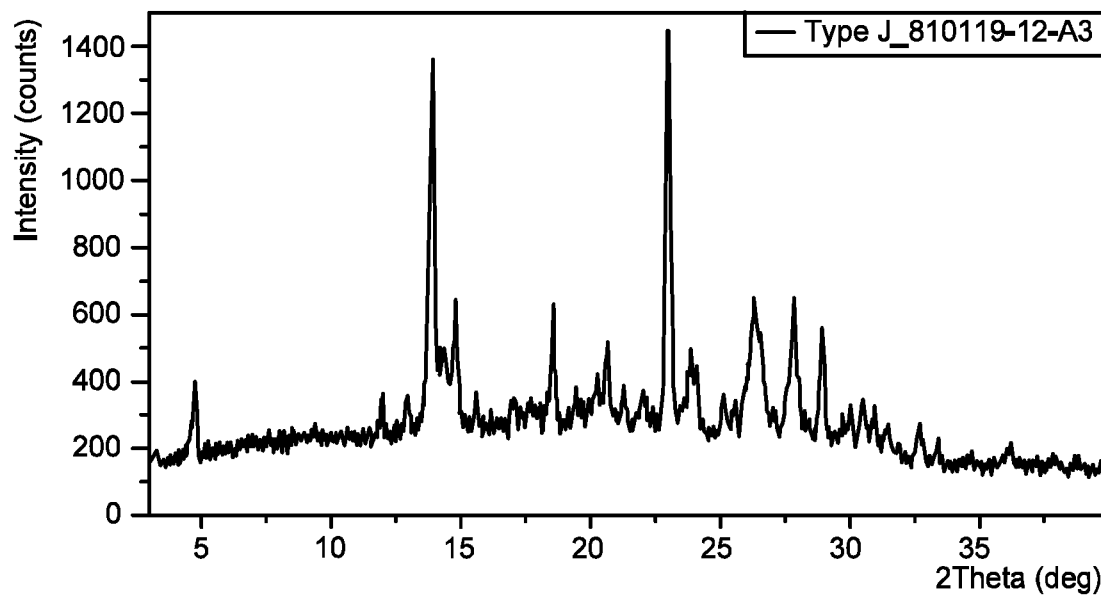
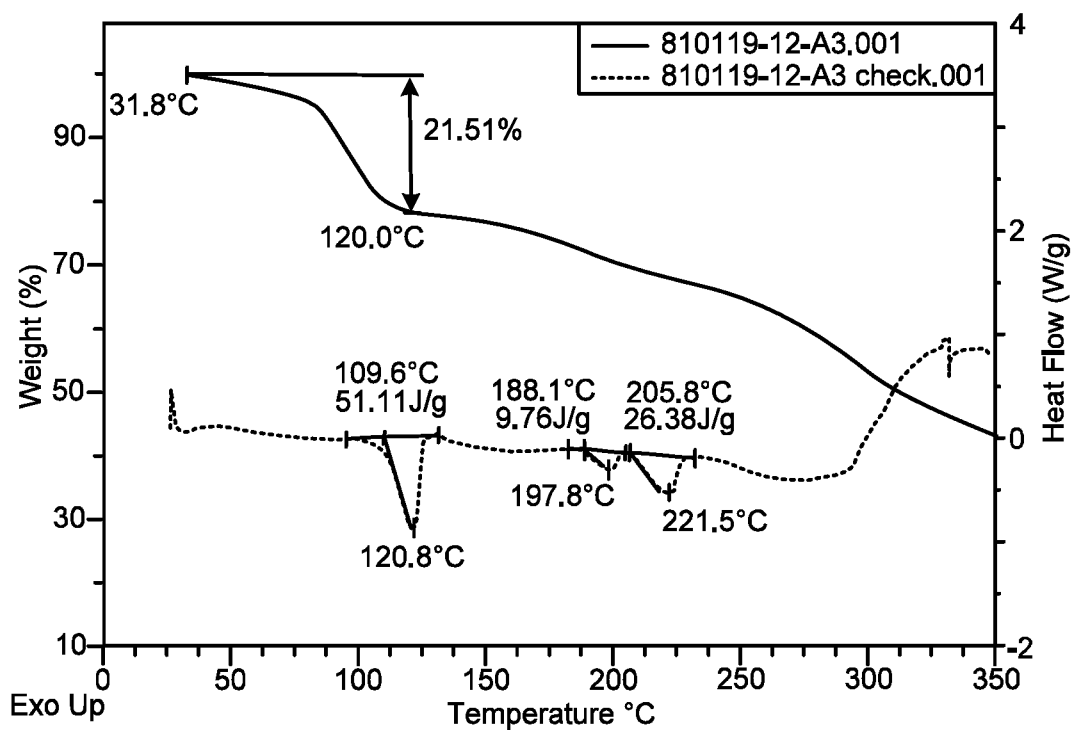
48 / 53

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

49 / 53

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

50 / 53

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

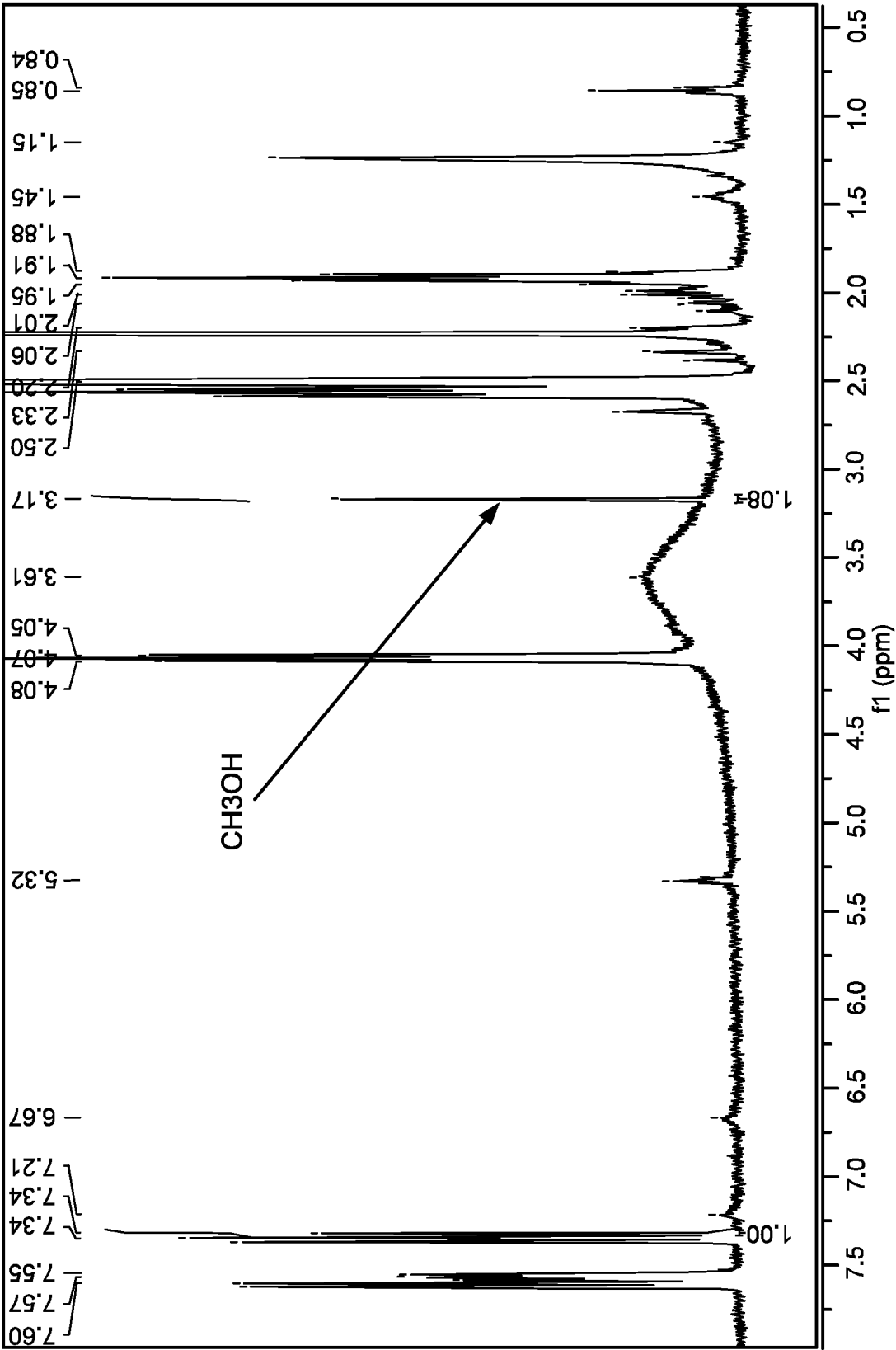


FIG. 61

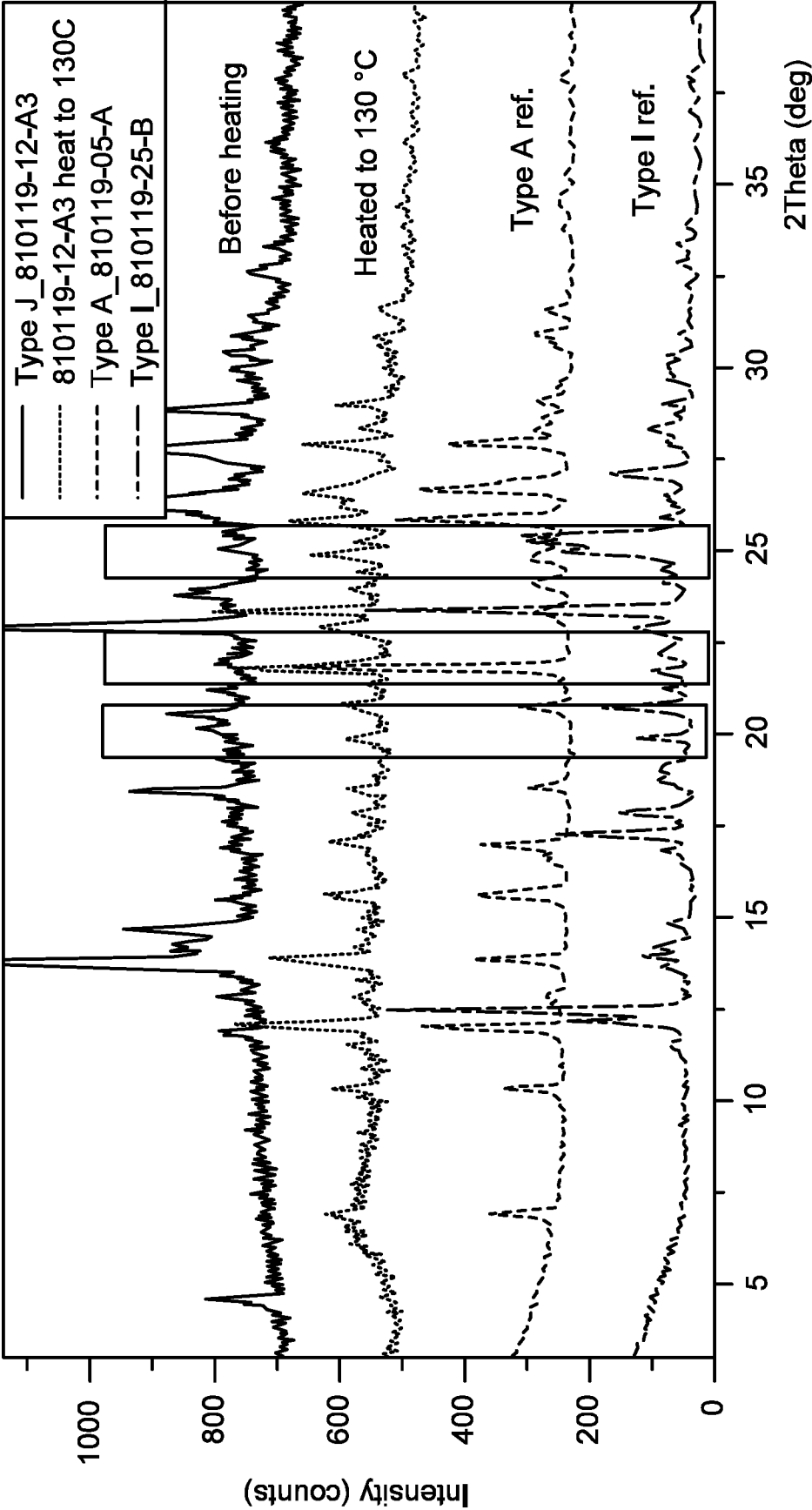
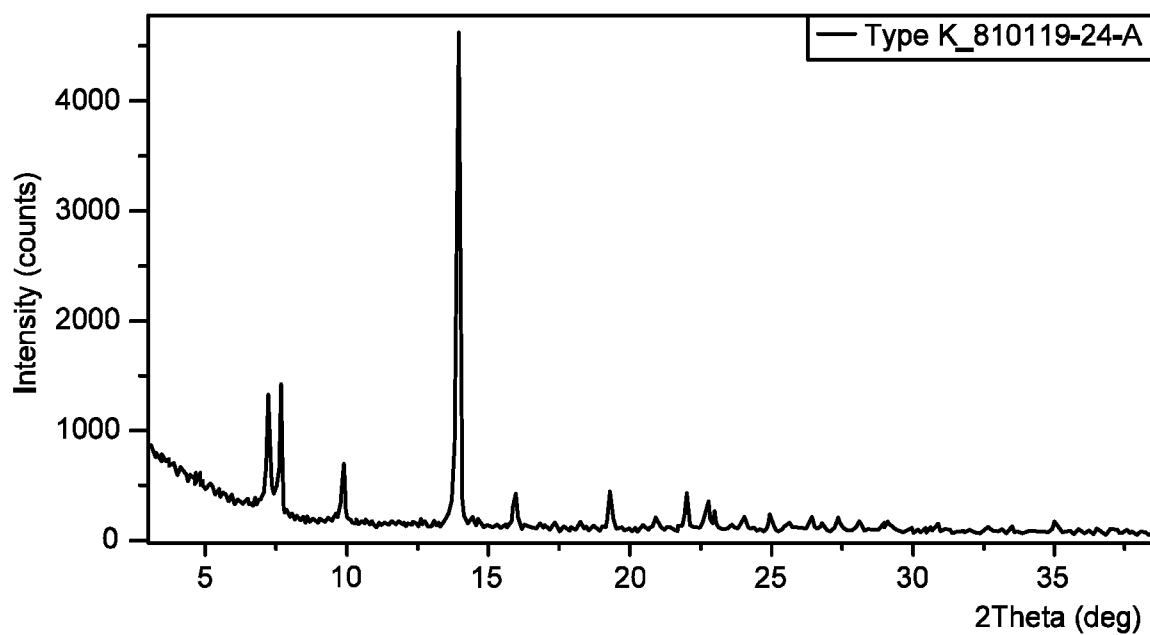
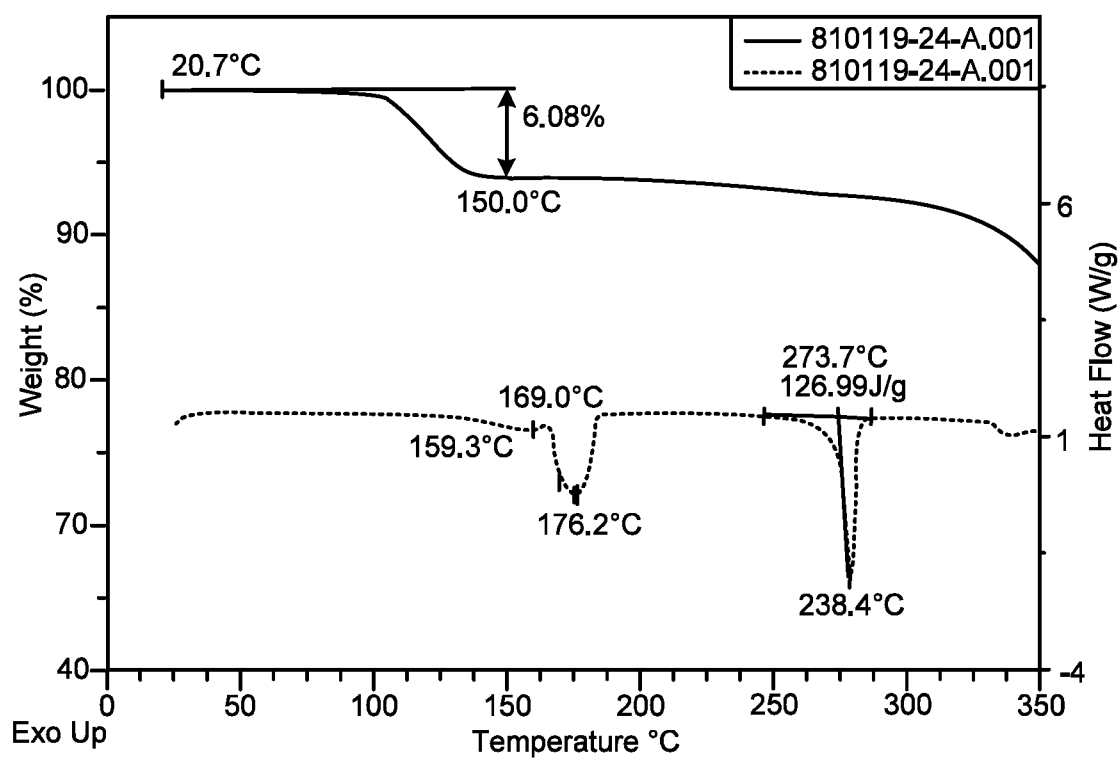


FIG. 62

53 / 53

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