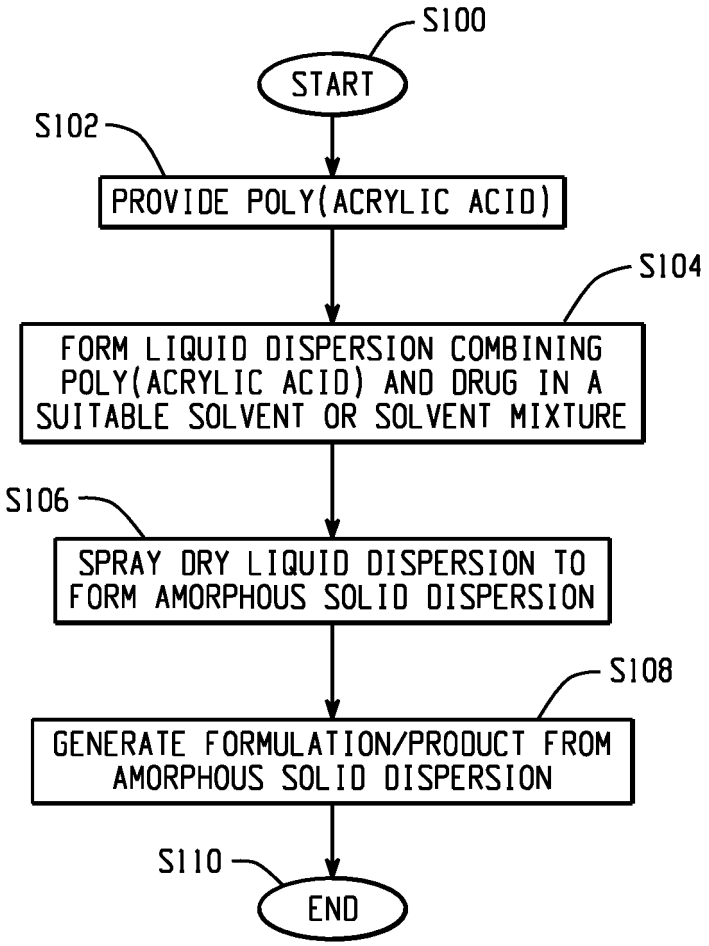

S100
START
S102
PROVIDE POLY(ACRYLIC ACID)
S104
FORM LIQUID DISPERSION COMBINING POLY(ACRYLIC ACID) AND DRUG IN A SUITABLE SOLVENT OR SOLVENT MIXTURE
S106
SPRAY DRY LIQUID DISPERSION TO FORM AMORPHOUS SOLID DISPERSION
S108
GENERATE FORMULATION/PRODUCT FROM AMORPHOUS SOLID DISPERSION
S110
END

*Fig. 1*

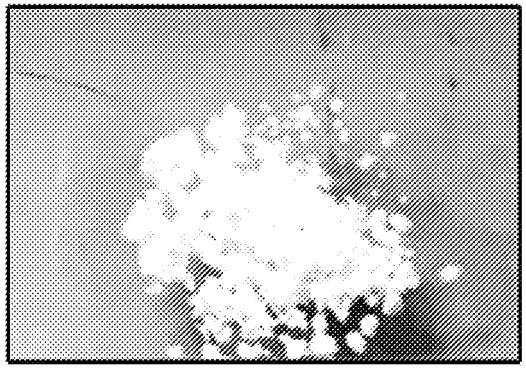


Fig. 2

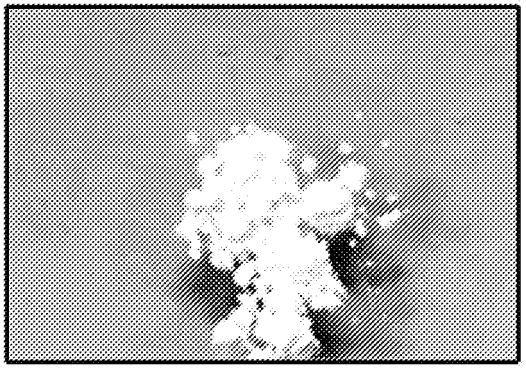


Fig. 3

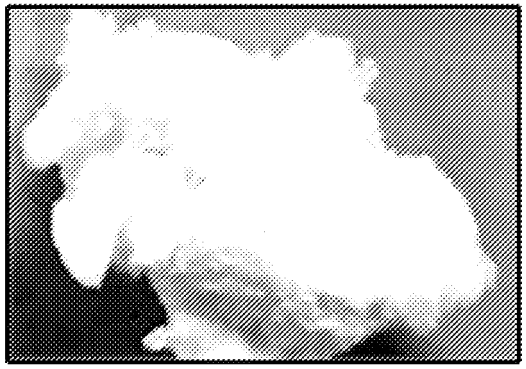


Fig. 4



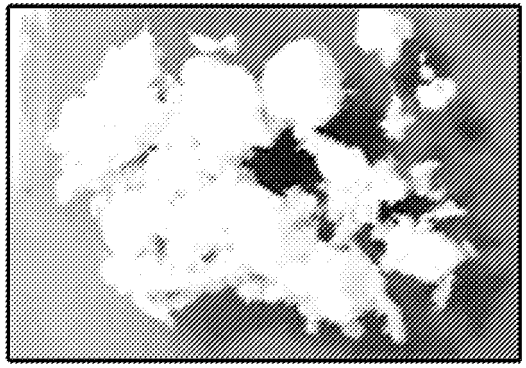


Fig. 5

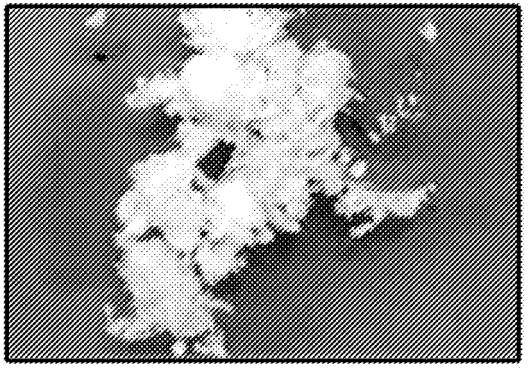


Fig. 6

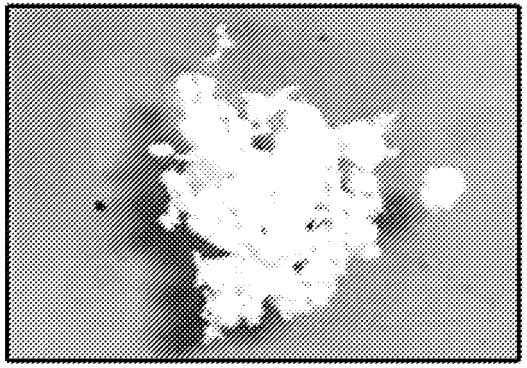


Fig. 7

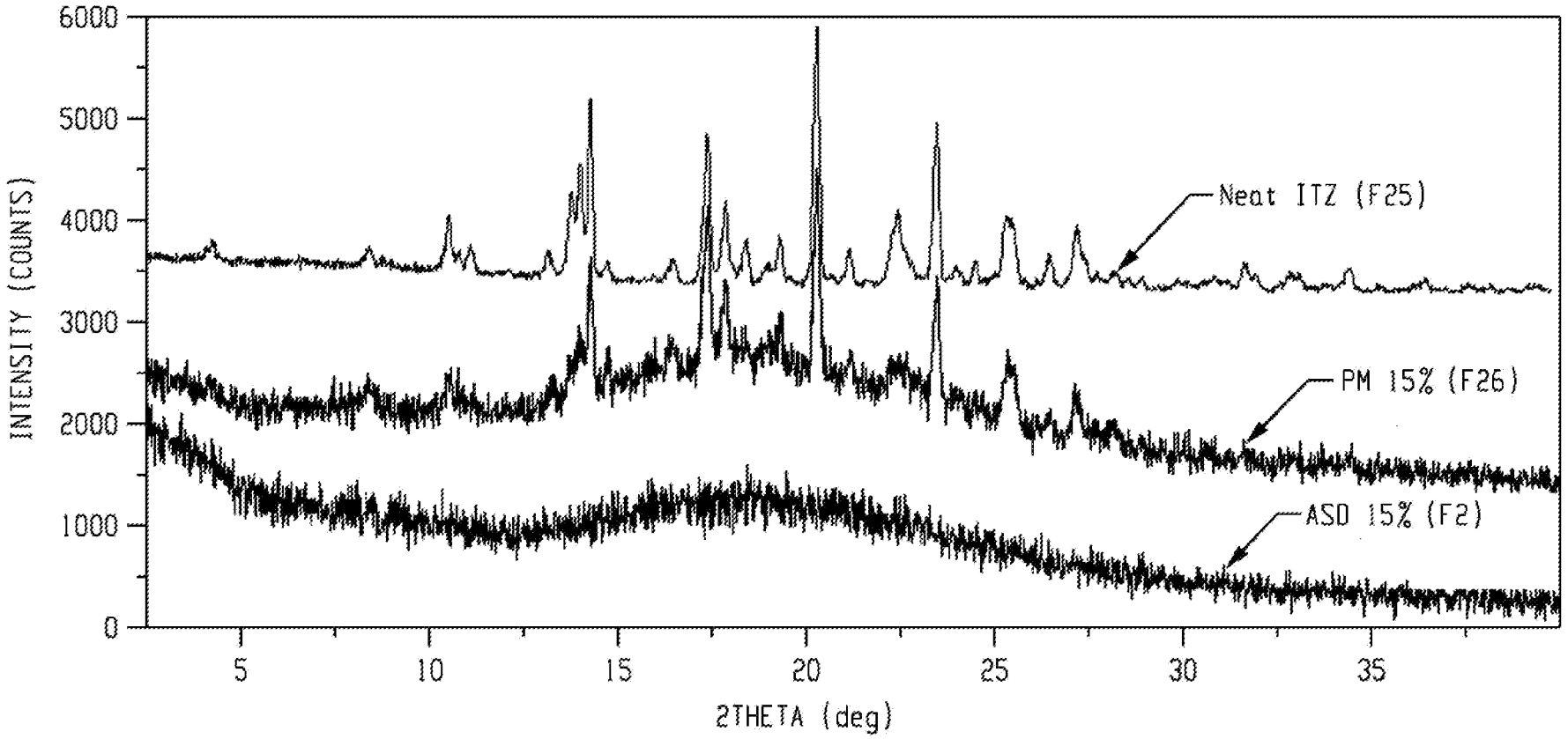

6000
5000
4000
3000
2000
1000
0
INTENSITY (COUNTS)
5
10
15
20
25
30
35
2THETA (deg)
Neat ITZ (F25)
PM 15% (F26)
ASD 15% (F2)

*Fig. 8*

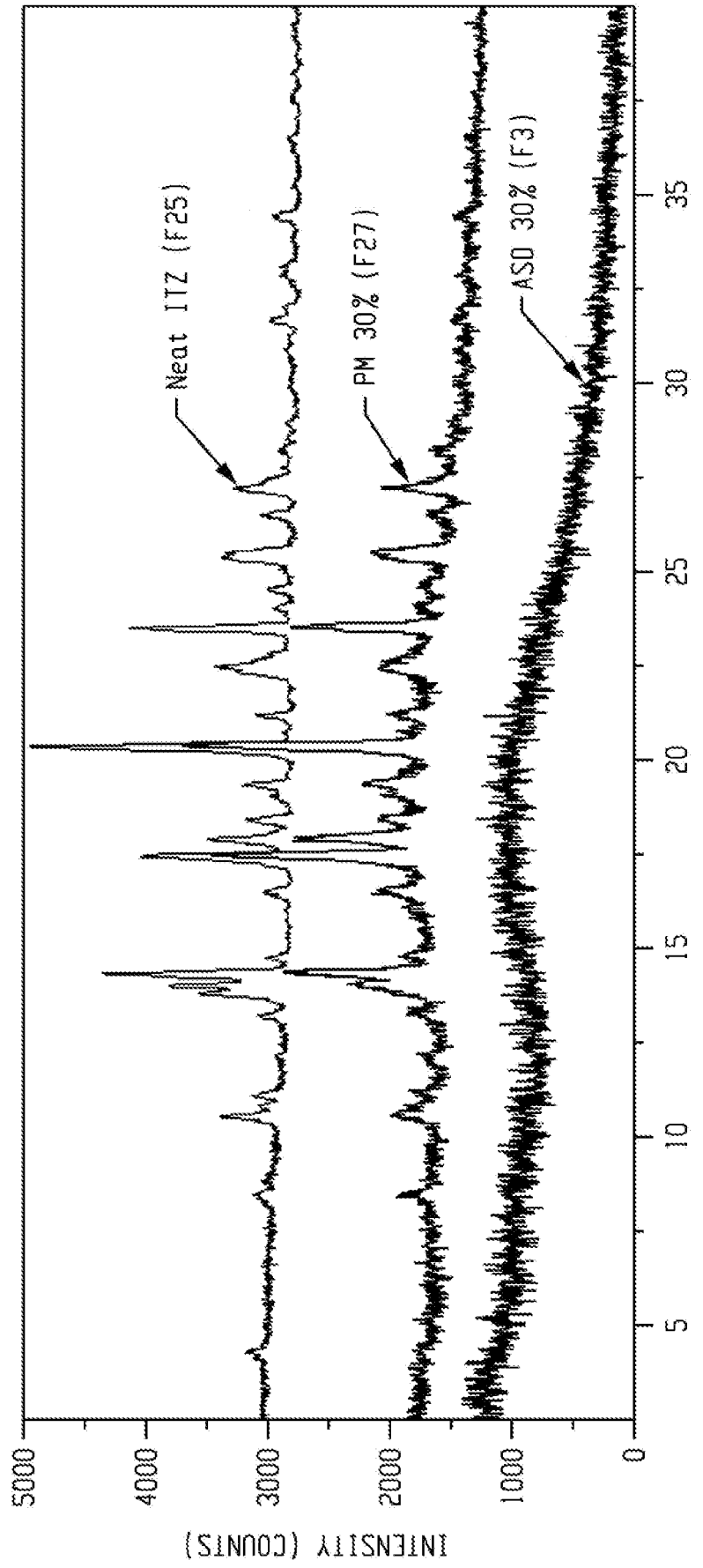

Neat ITZ (F25)
PM 30% (F27)
ASD 30% (F3)
INTENSITY (COUNTS)
2THETA (deg)
0
1000
2000
3000
4000
5000
5
10
15
20
25
30
35

*Fig. 9*

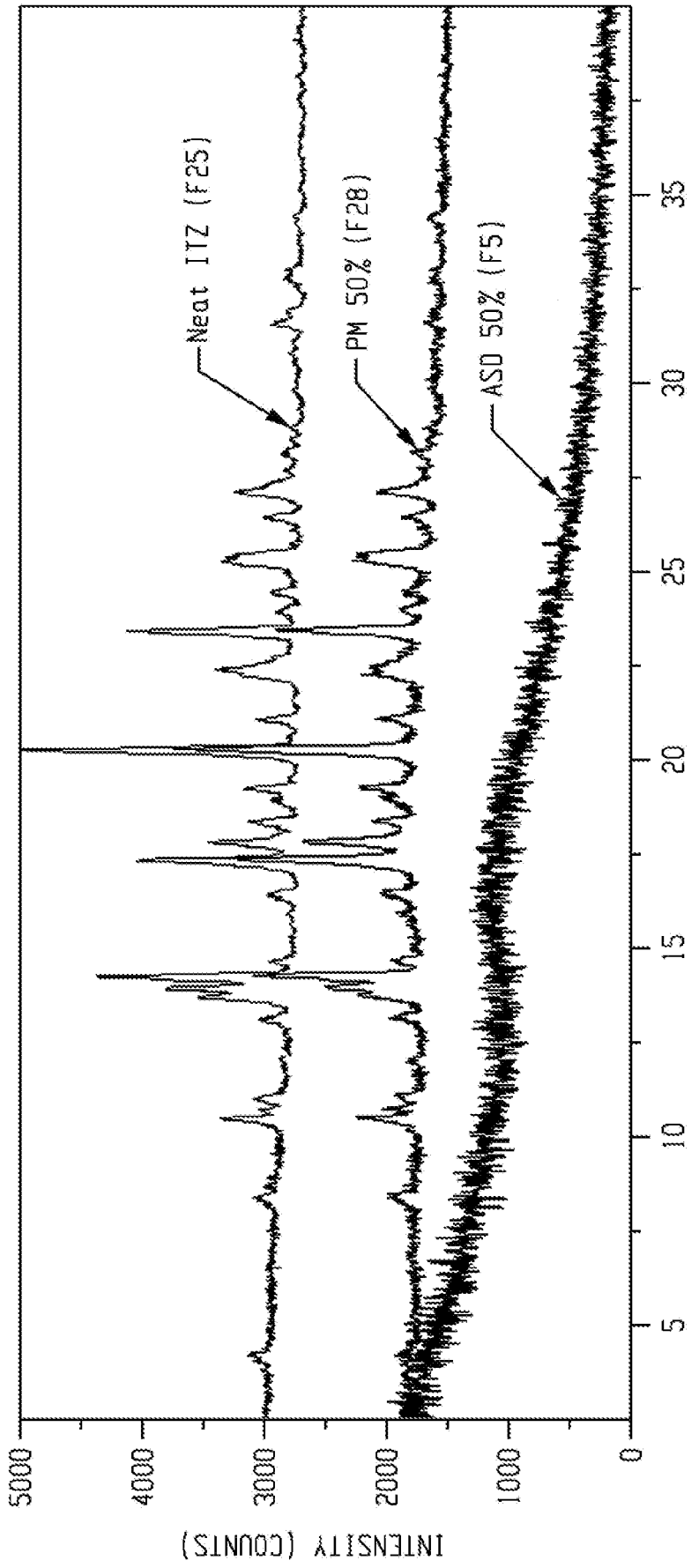

Neat ITZ (F25)
PM 50% (F28)
ASD 50% (F5)
INTENSITY (COUNTS)
2THETA (deg)
0
1000
2000
3000
4000
5000
5
10
15
20
25
30
35

*Fig. 10*

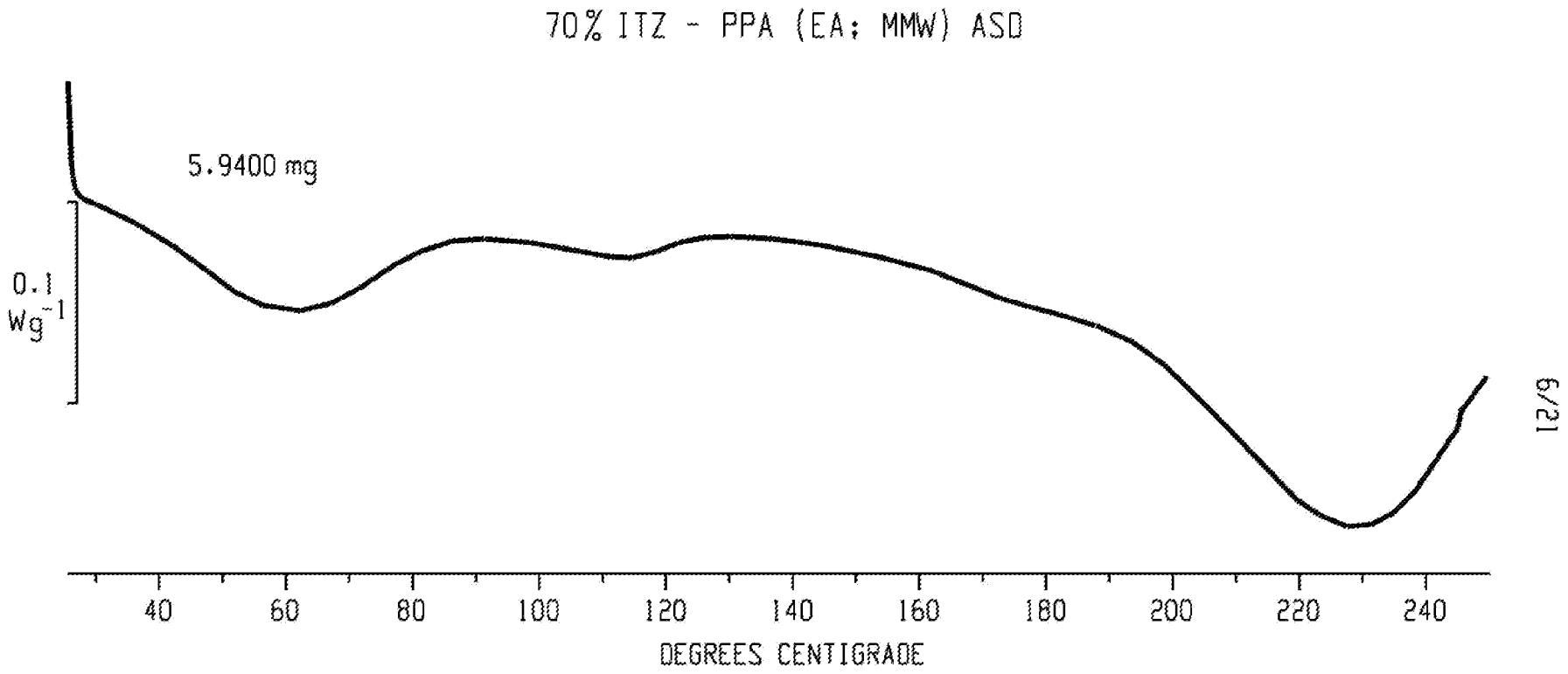

70% ITZ - PPA (EA; MMW) ASD
5.9400 mg
0.1
Wg-1
40
60
80
100
120
140
160
180
200
220
240
DEGREES CENTIGRADE



*Fig. 11*

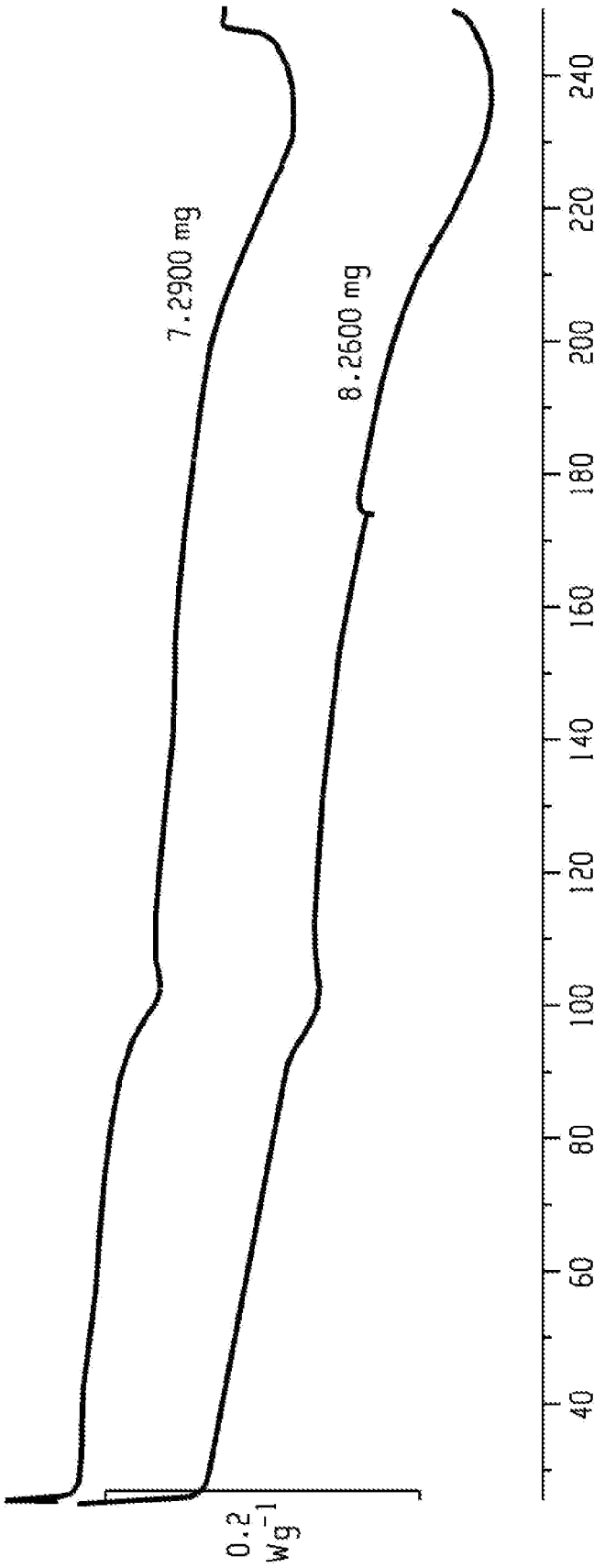

80% ITZ - PPA (EA: MMW) ASD
7.2900 mg
8.2600 mg
0.2
Wg-1
40
60
80
100
120
140
160
180
200
220
240
DEGREES CENTIGRADE

*Fig. 12*

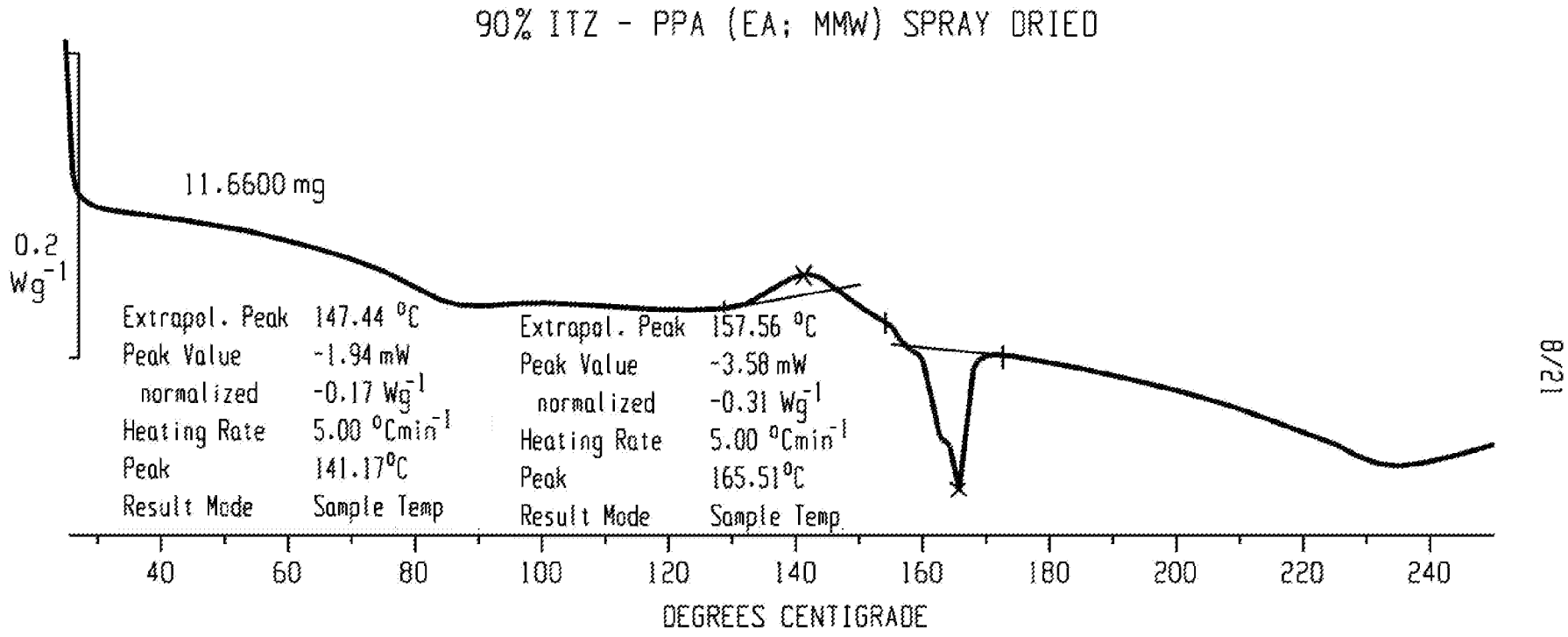

90% ITZ - PPA (EA; MMW) SPRAY DRIED
11.6600 mg
0.2
Wg-1
Extrapol. Peak 147.44 °C
Peak Value -1.94 mW
normalized -0.17 Wg-1
Heating Rate 5.00 °Cmin-1
Peak 141.17°C
Result Mode Sample Temp
Extrapol. Peak 157.56 °C
Peak Value -3.58 mW
normalized -0.31 Wg-1
Heating Rate 5.00 °Cmin-1
Peak 165.51°C
Result Mode Sample Temp
40
60
80
100
120
140
160
180
200
220
240
DEGREES CENTIGRADE

*Fig. 13*

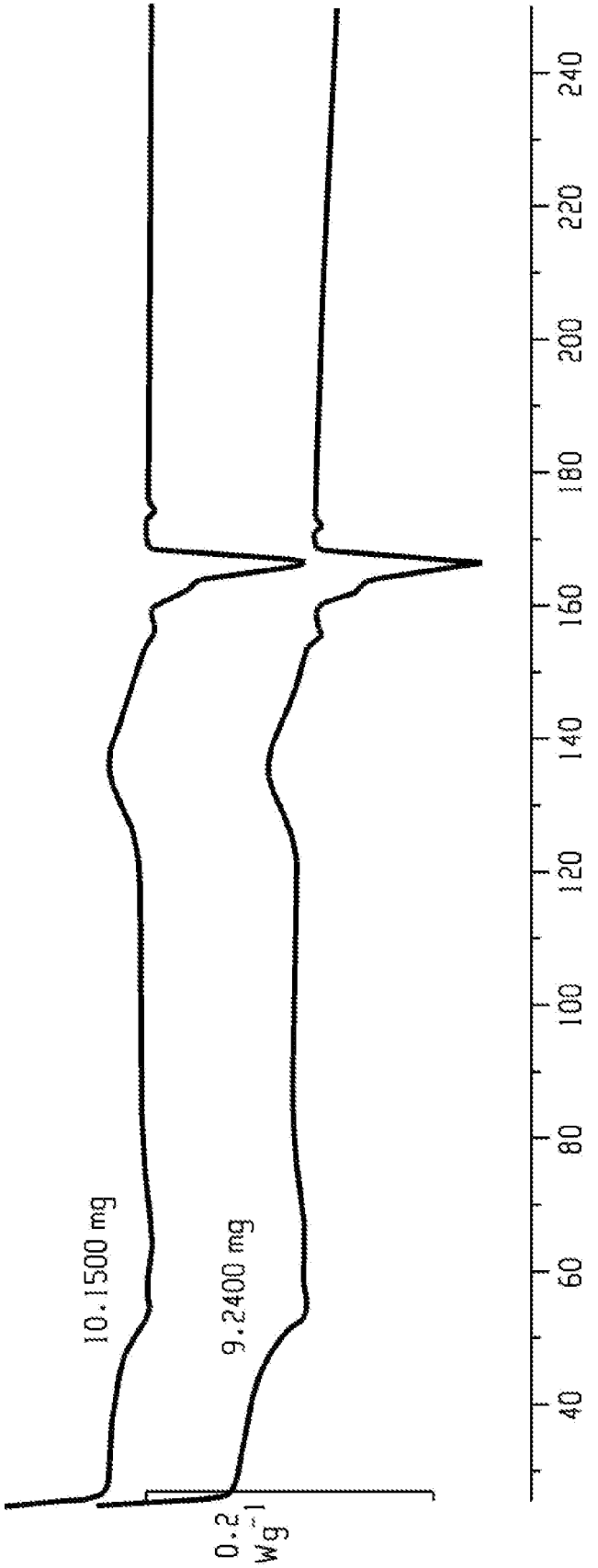

80% ITZ/SOLUPLUS SPRAY DRIED MATERIAL
10.1500 mg
9.2400 mg
0.2 Wg⁻¹
40
60
80
100
120
140
160
180
200
220
240
DEGREES CENTIGRADE

*Fig. 14*

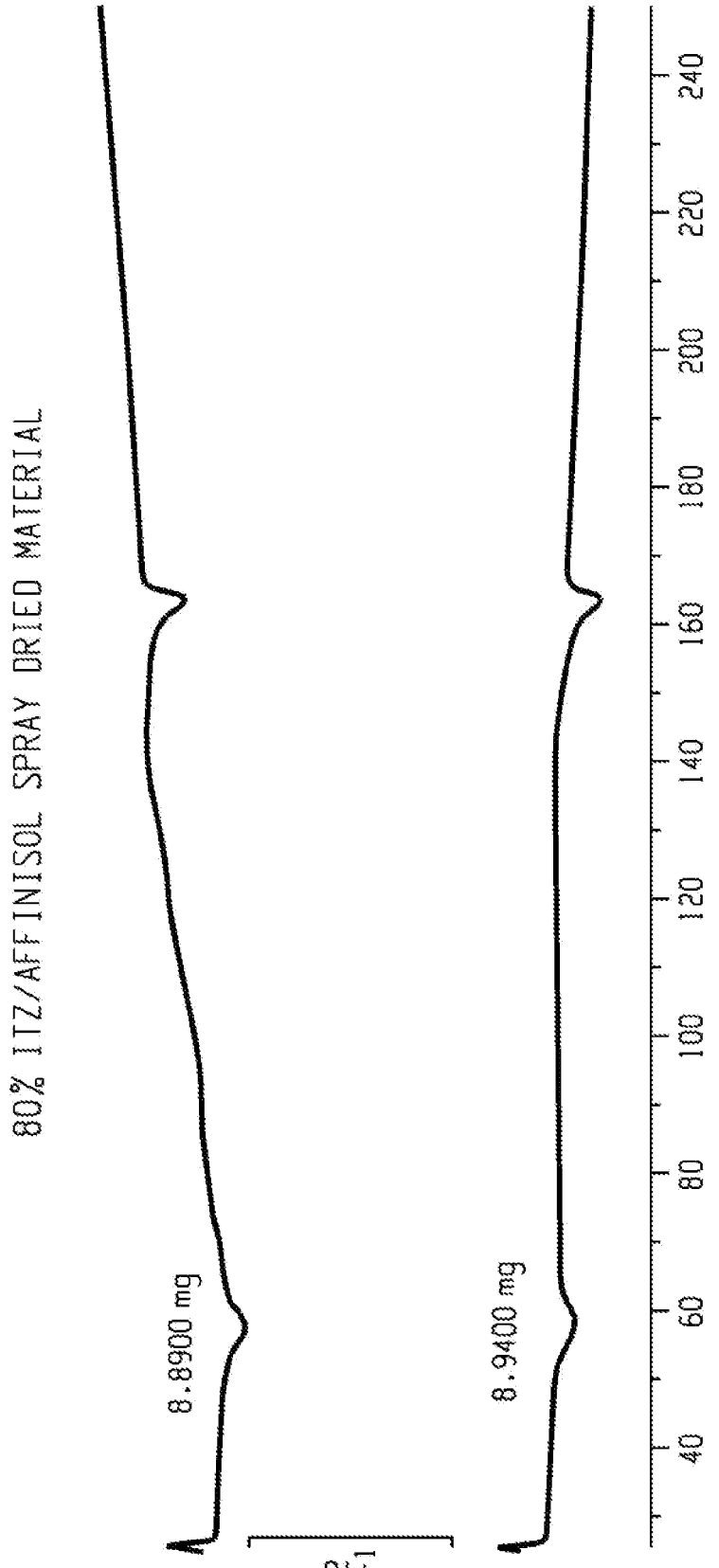

80% ITZ/AFFINISOL SPRAY DRIED MATERIAL
8.8900 mg
8.9400 mg
0.2 Wg⁻¹
40
60
80
100
120
140
160
180
200
220
240
DEGREES CENTIGRADE

*Fig. 15*

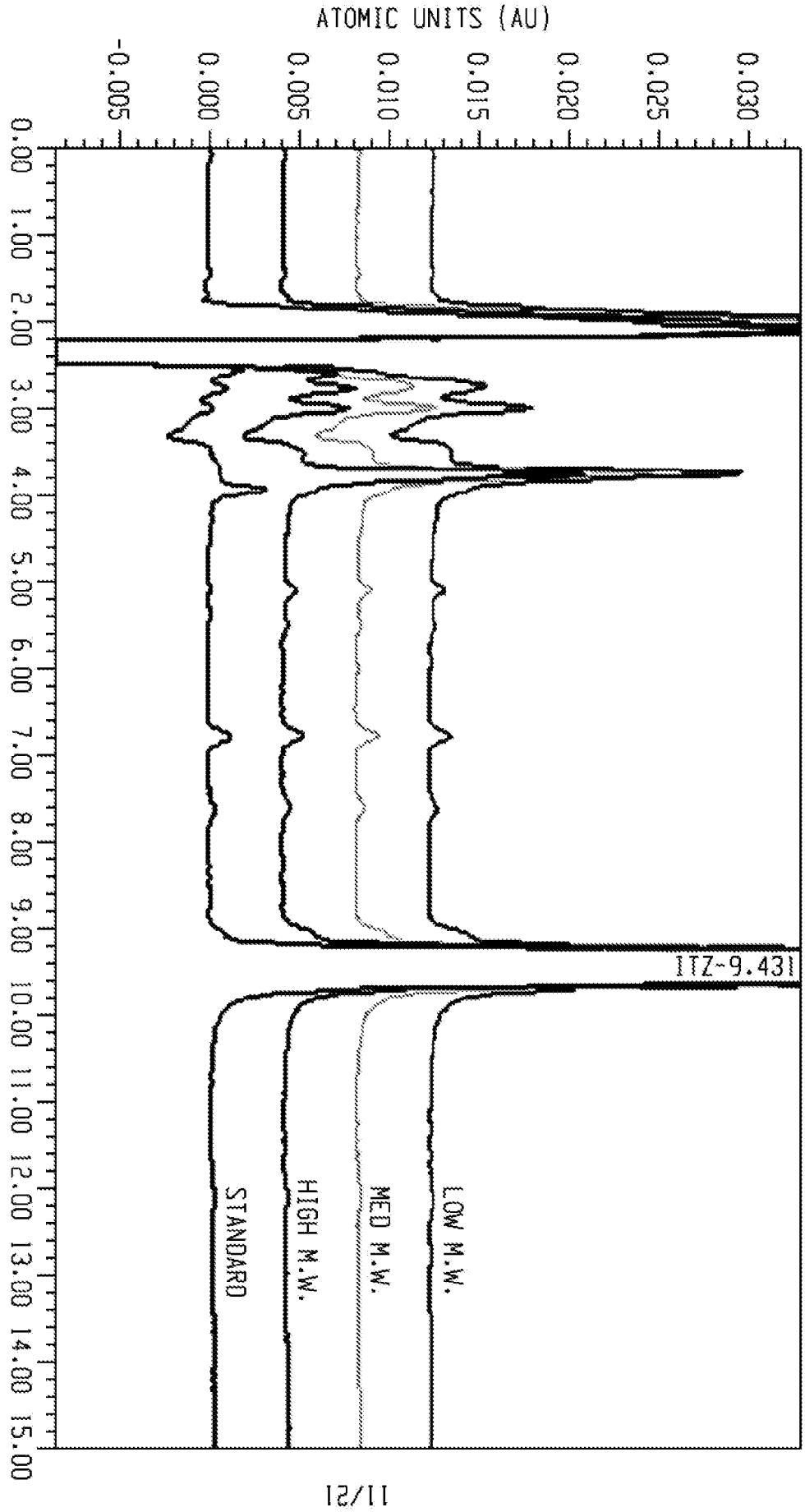

ATOMIC UNITS (AU)
-0.005
0.000
0.005
0.010
0.015
0.020
0.025
0.030
0.00
1.00
2.00
3.00
4.00
5.00
6.00
7.00
8.00
9.00
10.00
11.00
12.00
13.00
14.00
15.00
MINUTES
ITZ-9.431
STANDARD
HIGH M.W.
MED M.W.
LOW M.W.

Fig. 16

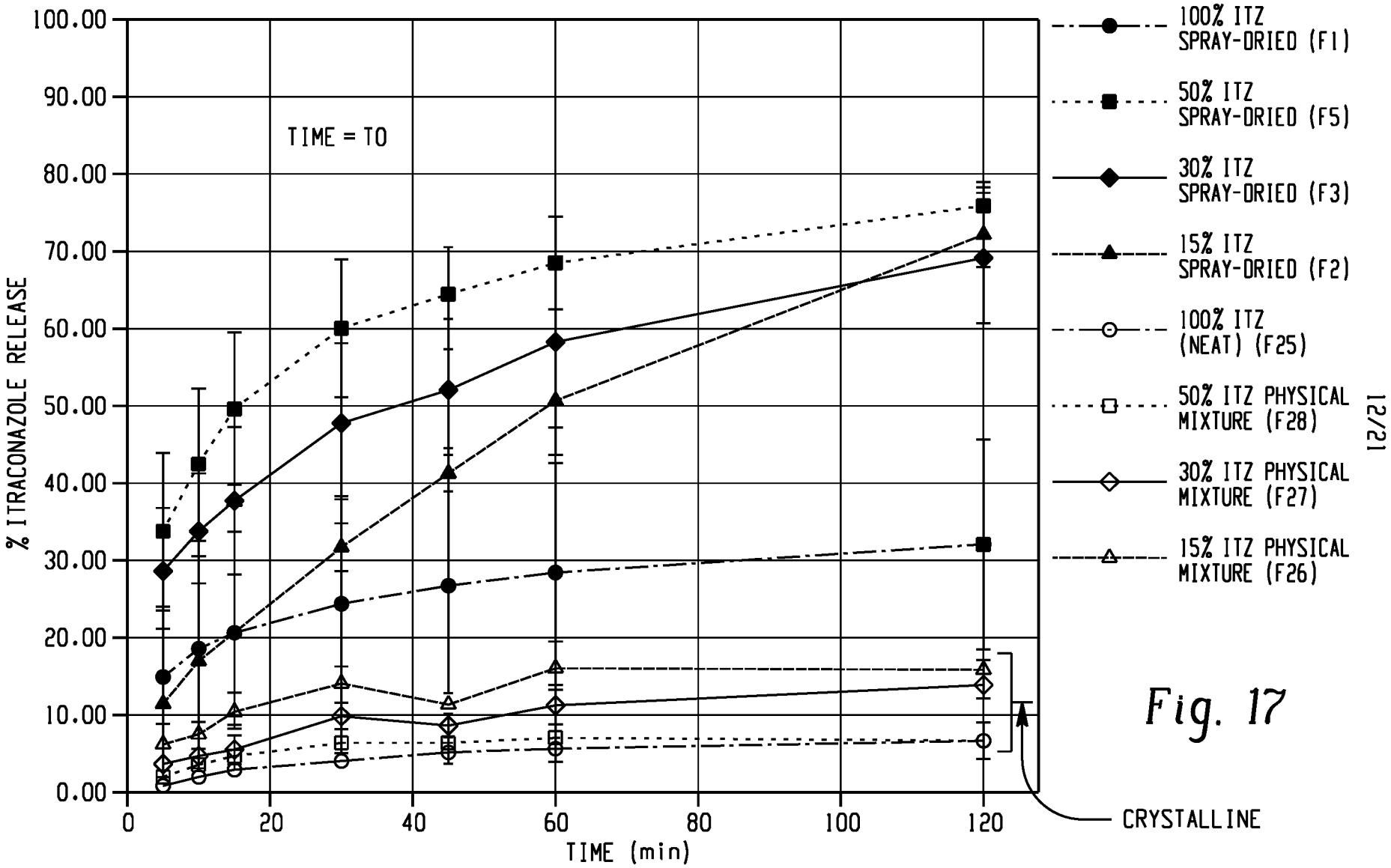

100.00
90.00
80.00
70.00
60.00
50.00
40.00
30.00
20.00
10.00
0.00
% ITRACONAZOLE RELEASE
TIME = TO
0
20
40
60
80
100
120
TIME (min)
100% ITZ SPRAY-DRIED (F1)
50% ITZ SPRAY-DRIED (F5)
30% ITZ SPRAY-DRIED (F3)
15% ITZ SPRAY-DRIED (F2)
100% ITZ (NEAT) (F25)
50% ITZ PHYSICAL MIXTURE (F28)
30% ITZ PHYSICAL MIXTURE (F27)
15% ITZ PHYSICAL MIXTURE (F26)
CRYSTALLINE

Fig. 17

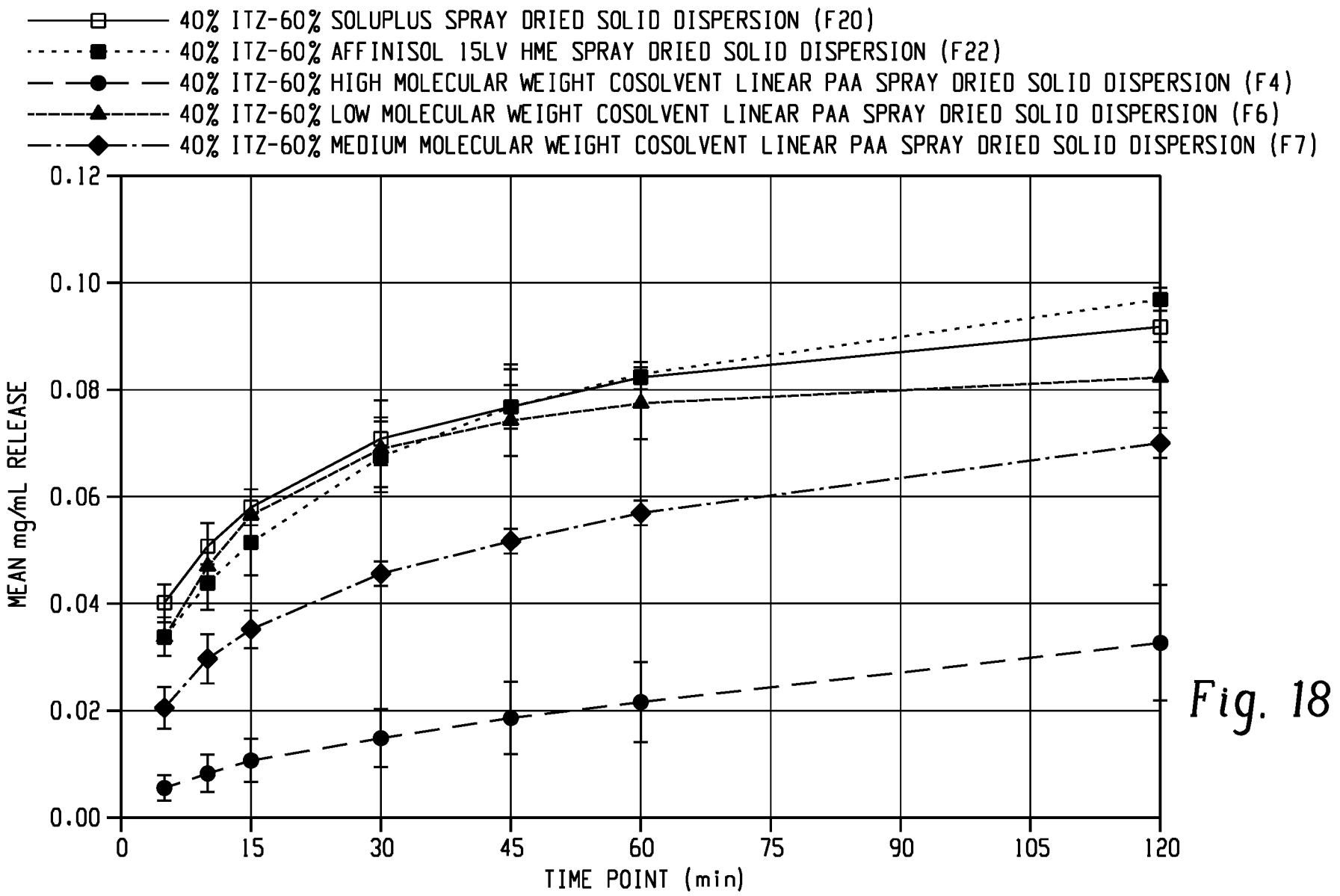

40% ITZ-60% SOLUPLUS SPRAY DRIED SOLID DISPERSION (F20)
40% ITZ-60% AFFINISOL 15LV HME SPRAY DRIED SOLID DISPERSION (F22)
40% ITZ-60% HIGH MOLECULAR WEIGHT COSOLVENT LINEAR PAA SPRAY DRIED SOLID DISPERSION (F4)
40% ITZ-60% LOW MOLECULAR WEIGHT COSOLVENT LINEAR PAA SPRAY DRIED SOLID DISPERSION (F6)
40% ITZ-60% MEDIUM MOLECULAR WEIGHT COSOLVENT LINEAR PAA SPRAY DRIED SOLID DISPERSION (F7)
MEAN mg/mL RELEASE
0.12
0.10
0.08
0.06
0.04
0.02
0.00
0
15
30
45
60
75
90
105
120
TIME POINT (min)

Fig. 18

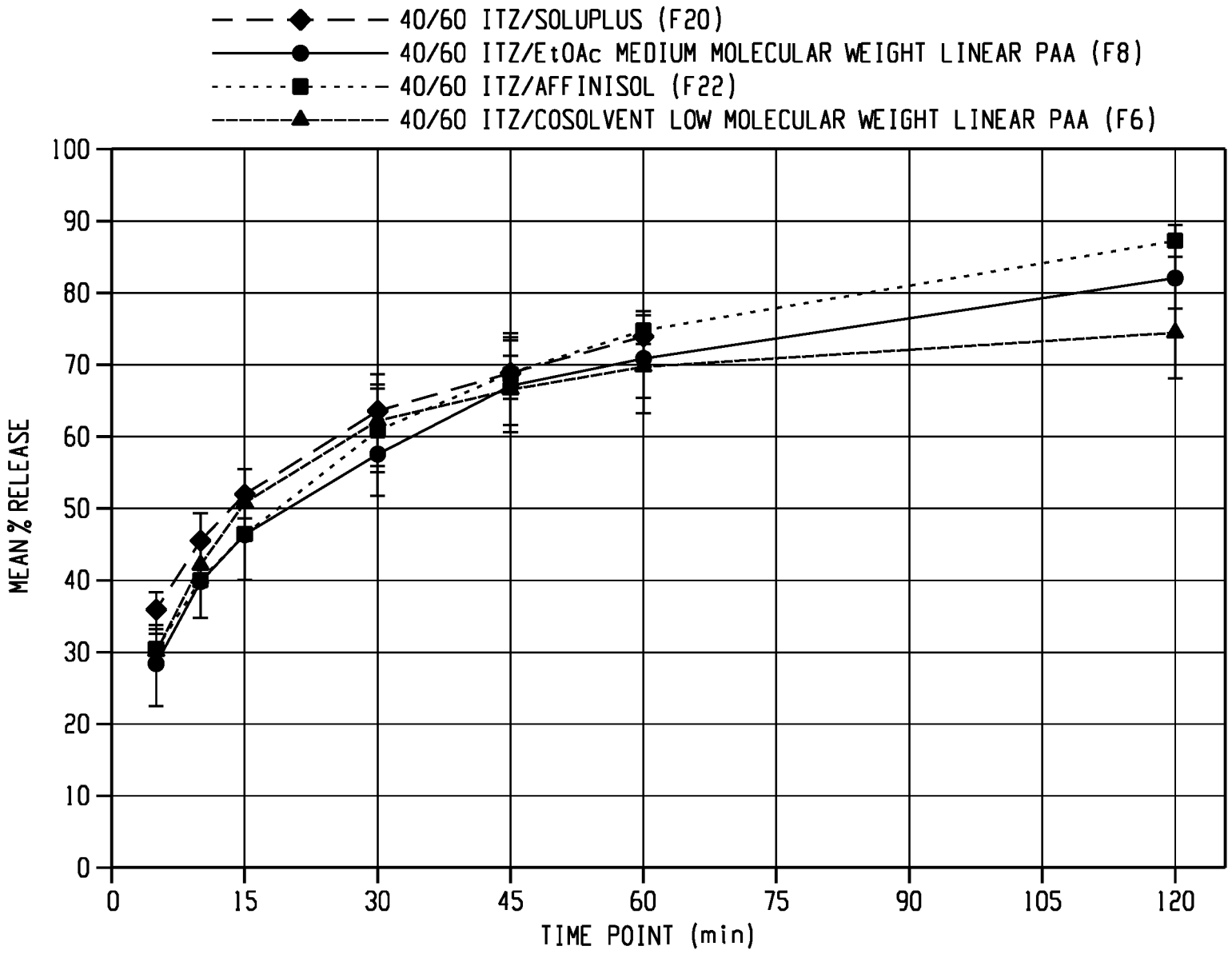

40/60 ITZ/SOLUPLUS (F20)
40/60 ITZ/EtOAc MEDIUM MOLECULAR WEIGHT LINEAR PAA (F8)
40/60 ITZ/AFFINISOL (F22)
40/60 ITZ/COSOLVENT LOW MOLECULAR WEIGHT LINEAR PAA (F6)
MEAN % RELEASE
TIME POINT (min)
0
10
20
30
40
50
60
70
80
90
100
0
15
30
45
60
75
90
105
120

Fig. 19

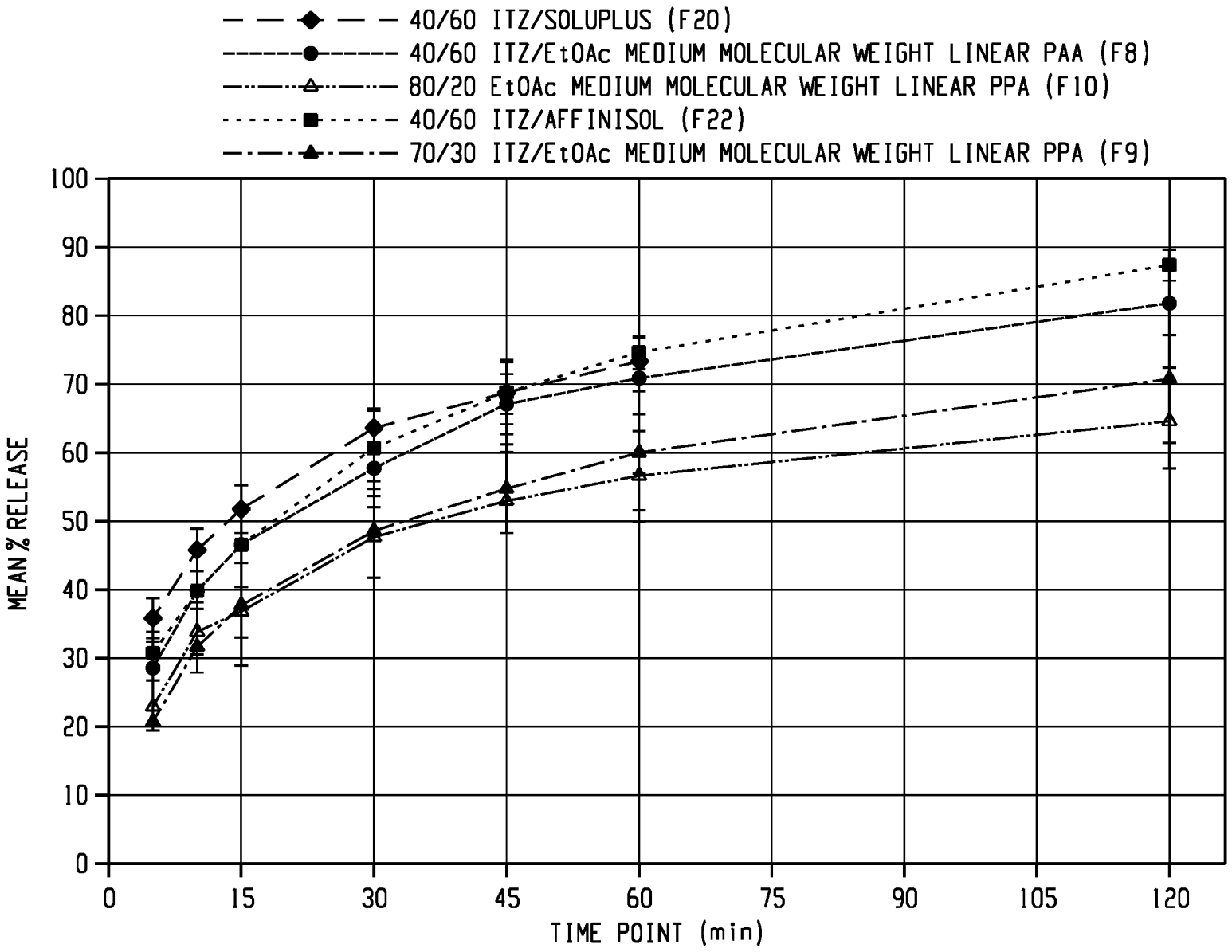

40/60 ITZ/SOLUPLUS (F20)
40/60 ITZ/EtOAc MEDIUM MOLECULAR WEIGHT LINEAR PAA (F8)
80/20 EtOAc MEDIUM MOLECULAR WEIGHT LINEAR PPA (F10)
40/60 ITZ/AFFINISOL (F22)
70/30 ITZ/EtOAc MEDIUM MOLECULAR WEIGHT LINEAR PPA (F9)
MEAN % RELEASE
TIME POINT (min)
0
10
20
30
40
50
60
70
80
90
100
0
15
30
45
60
75
90
105
120

Fig. 20

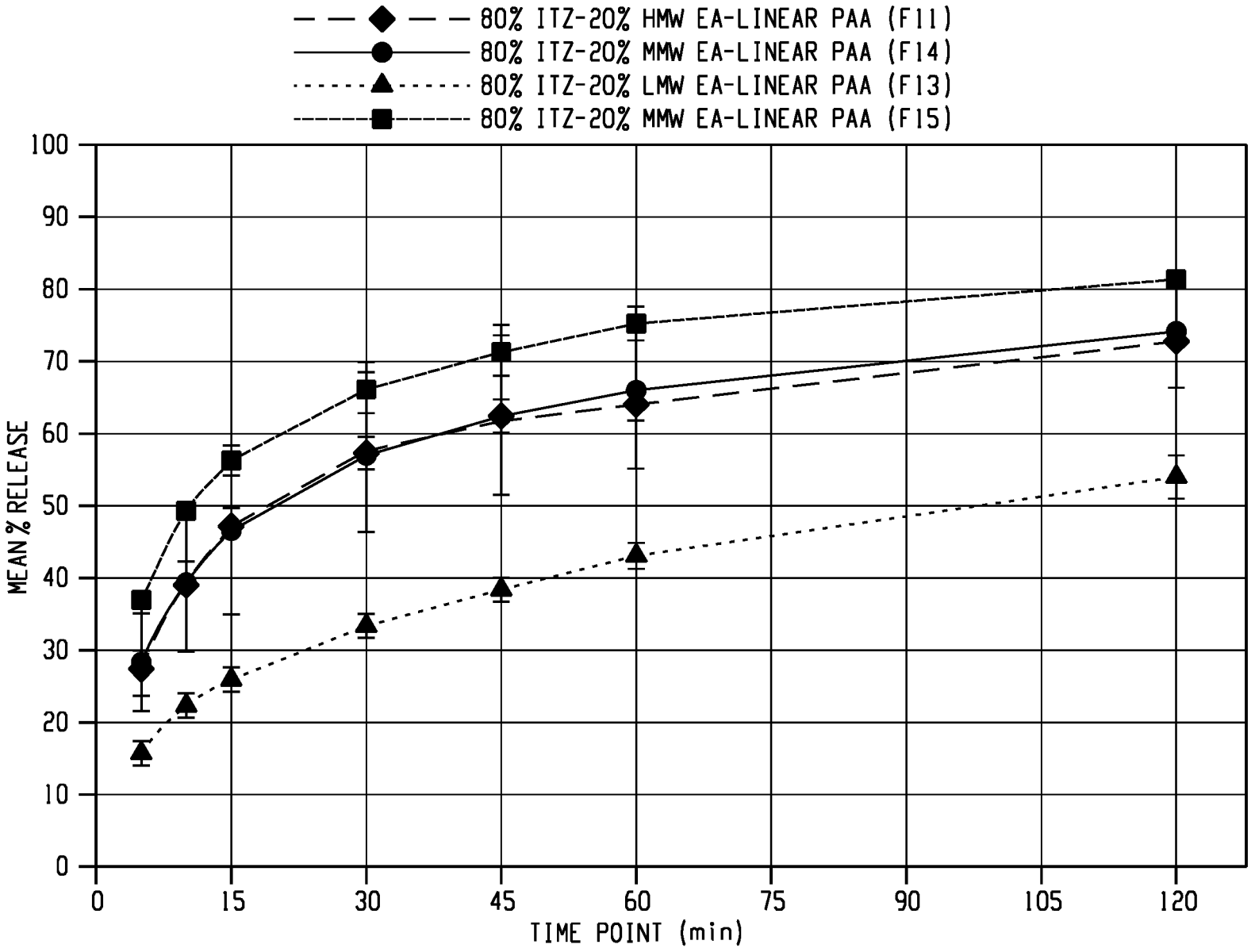

80% ITZ-20% HMW EA-LINEAR PAA (F11)
80% ITZ-20% MMW EA-LINEAR PAA (F14)
80% ITZ-20% LMW EA-LINEAR PAA (F13)
80% ITZ-20% MMW EA-LINEAR PAA (F15)
MEAN % RELEASE
TIME POINT (min)
0
10
20
30
40
50
60
70
80
90
100
0
15
30
45
60
75
90
105
120

Fig. 21

● 80% ITZ-20% PAA 0 MONTHS
○ 80% ITZ-20% PAA 6 MONTHS 40C/75% RH
%ITZ
TIME (min)
0
10
20
30
40
50
60
70
80
90
100
0
20
40
60
80
100
120
140

Fig. 22

▲ 40% ITZ-60% SOLUPLUS 0 MONTHS
△ 40% ITZ-60% SOLUPLUS 6 MONTHS 40C/75% RH
%ITZ
TIME (min)
0
10
20
30
40
50
60
70
80
90
100
0
20
40
60
80
100
120
140

Fig. 23

■ 40% ITZ-60% AFFINISOL 0 MONTHS
□ 40% ITZ-60% AFFINISOL 6 MONTHS 40C/75% RH
%ITZ
TIME (min)
0
10
20
30
40
50
60
70
80
90
100
0
20
40
60
80
100
120
140

Fig. 24

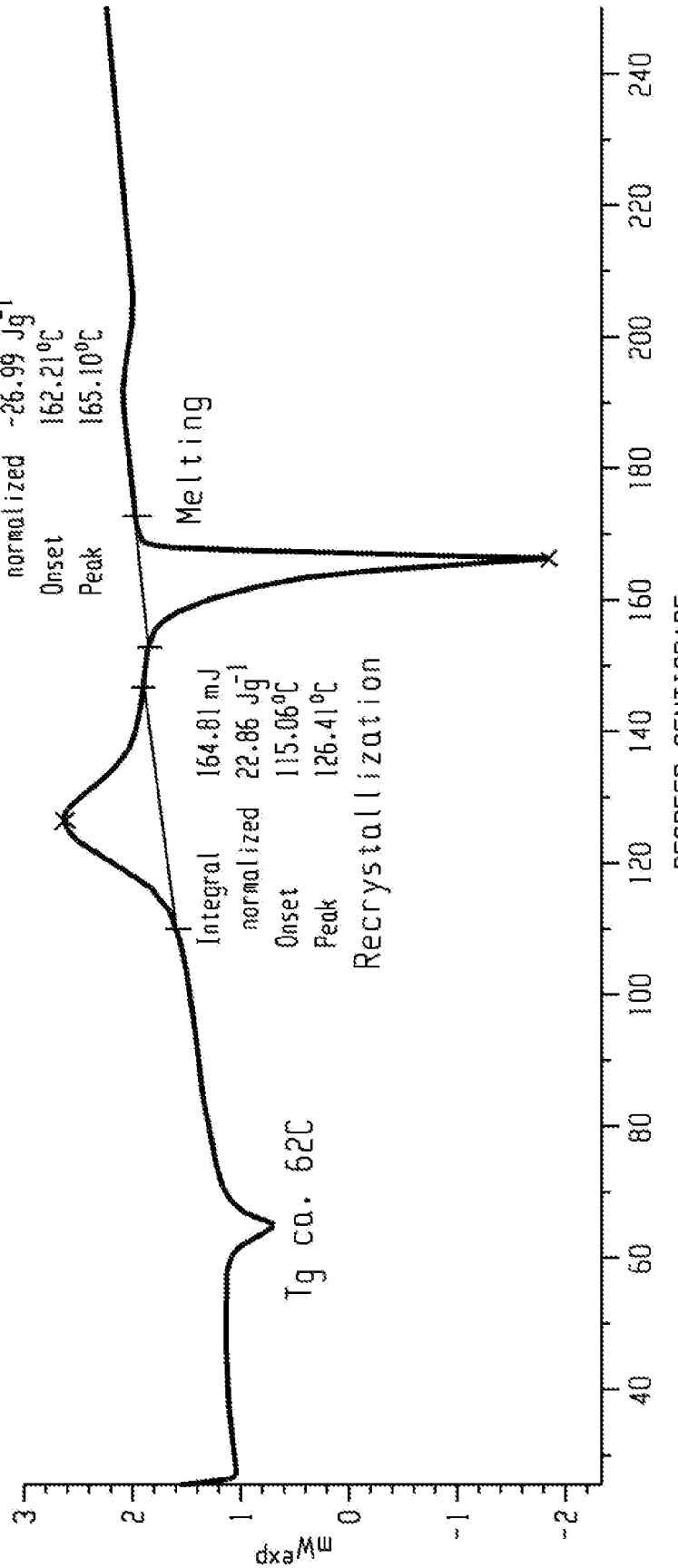

Integral -194.62 mJ
normalized -26.99 Jg-1
Onset 162.21°C
Peak 165.10°C
Melting
Integral 164.81 mJ
normalized 22.86 Jg-1
Onset 115.06°C
Peak 126.41°C
Recrystallization
Tg ca. 62C
mW exo
DEGREES CENTIGRADE
3
2
1
0
-1
-2
40
60
80
100
120
140
160
180
200
220
240

Fig. 25

Integral -259.08 mJ
normalized -43.40 Jg-1
Onset 161.76°C
Peak 165.80°C
Melting
Recrystallization
Integral 260.00 mJ
normalized 43.57 Jg-1
Onset 117.56°C
Peak 126.01°C
Tg ca. 62C
mWexp
DEGREES CENTIGRADE

*Fig. 26*