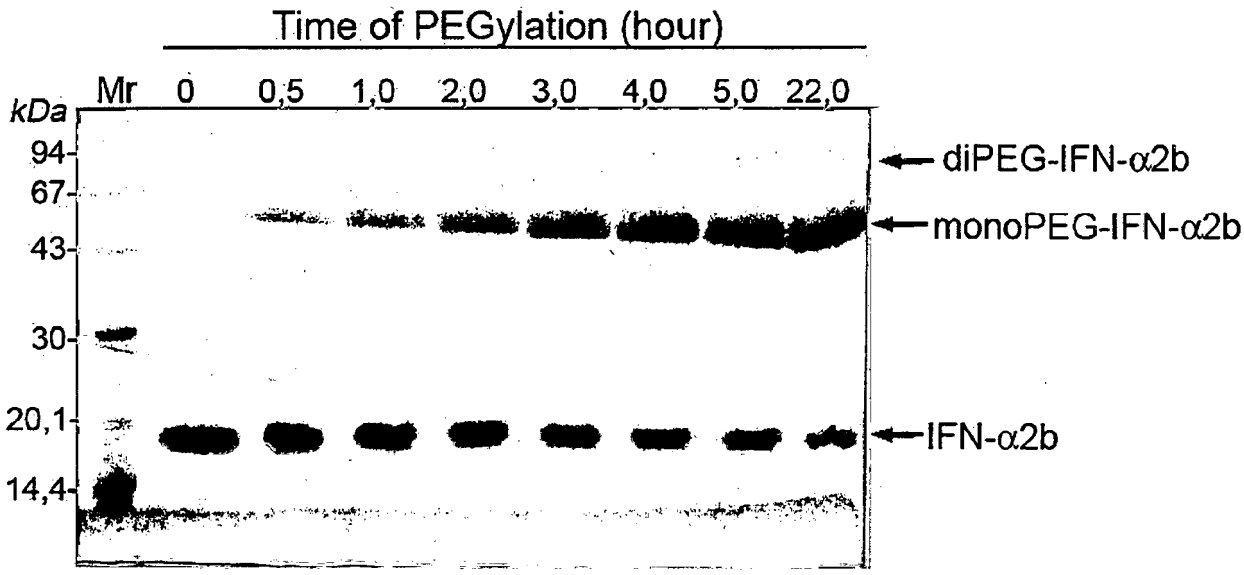
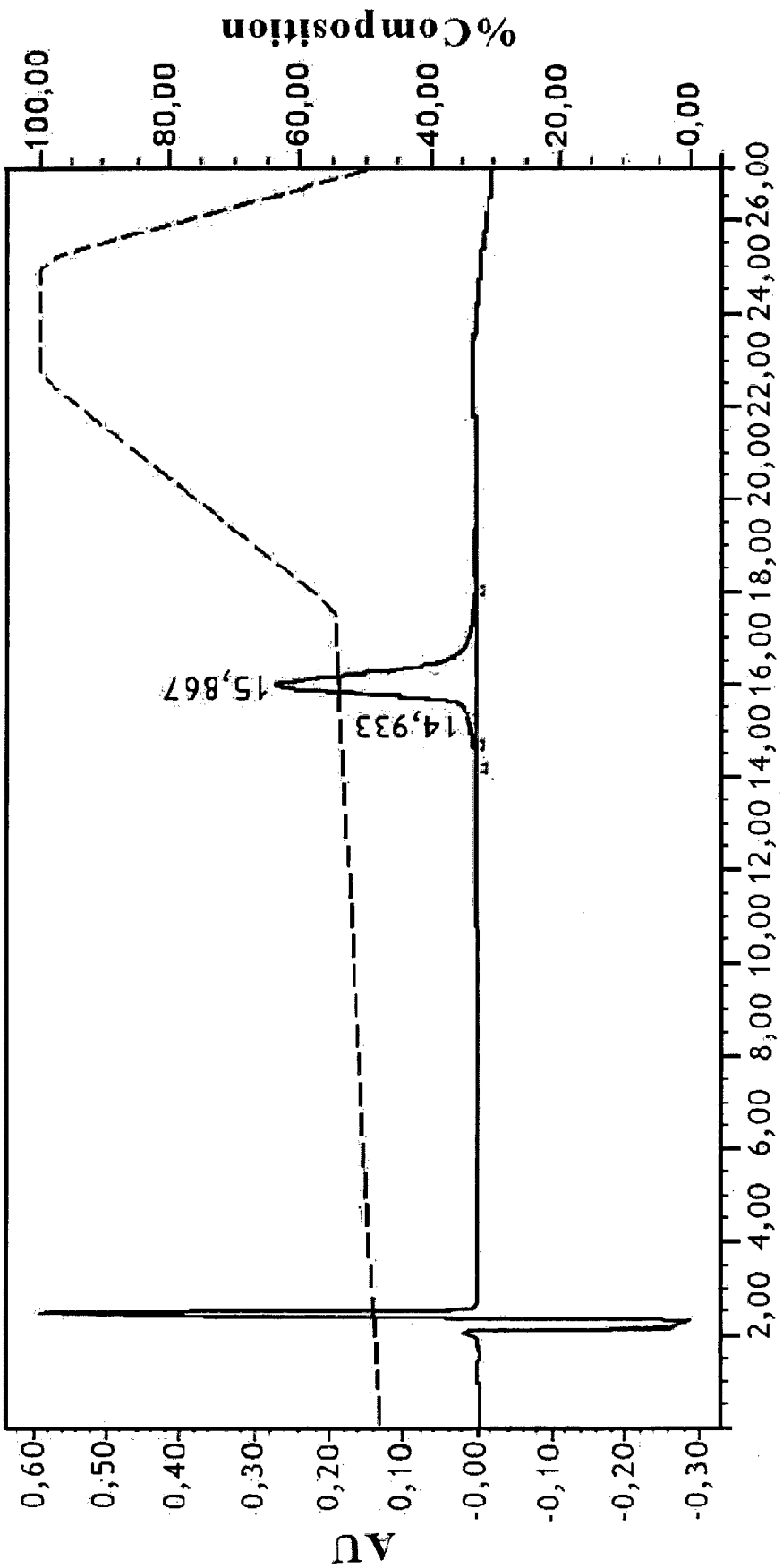

Time of PEGylation (hour)
Mr
0
0,5
1,0
2,0
3,0
4,0
5,0
22,0
kDa
94
67
43
30
20,1
14,4
diPEG-IFN-α2b
monoPEG-IFN-α2b
IFN-α2b

Fig.1



	RT (min)	Area (V'sec)	% Area	Height (V)	% Height
1	14,933	31174	0,33	1876	0,70
2	15,867	9433441	99,67	267135	99,30

Fig. 2

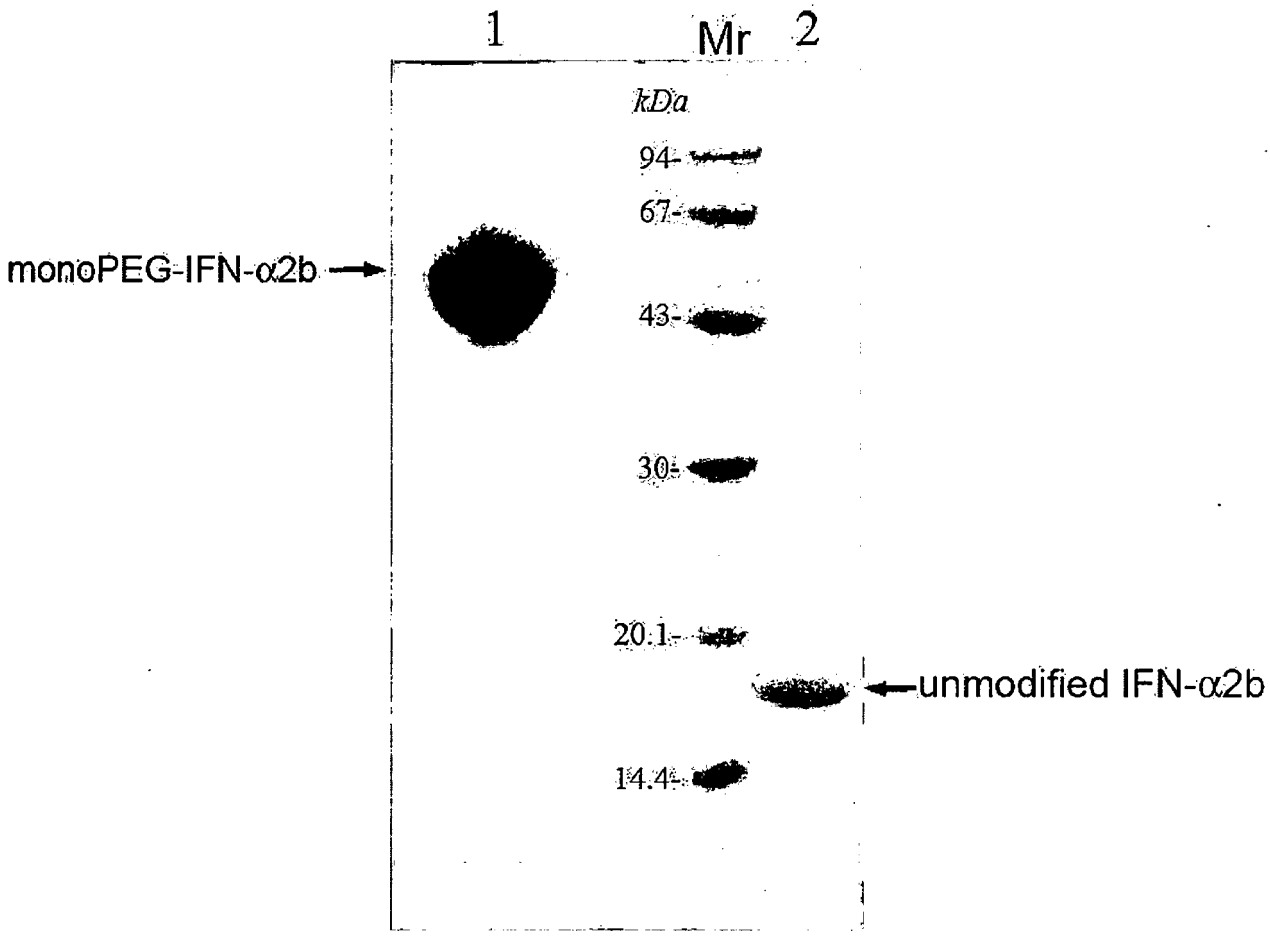


Fig. 3

Mr – molecular weight markers;

1 – 40 mkg of PEG-IFN-α2b conjugate was applied per lane;

2 – 40 mkg of unmodified IFN-α2b was applied per lane.

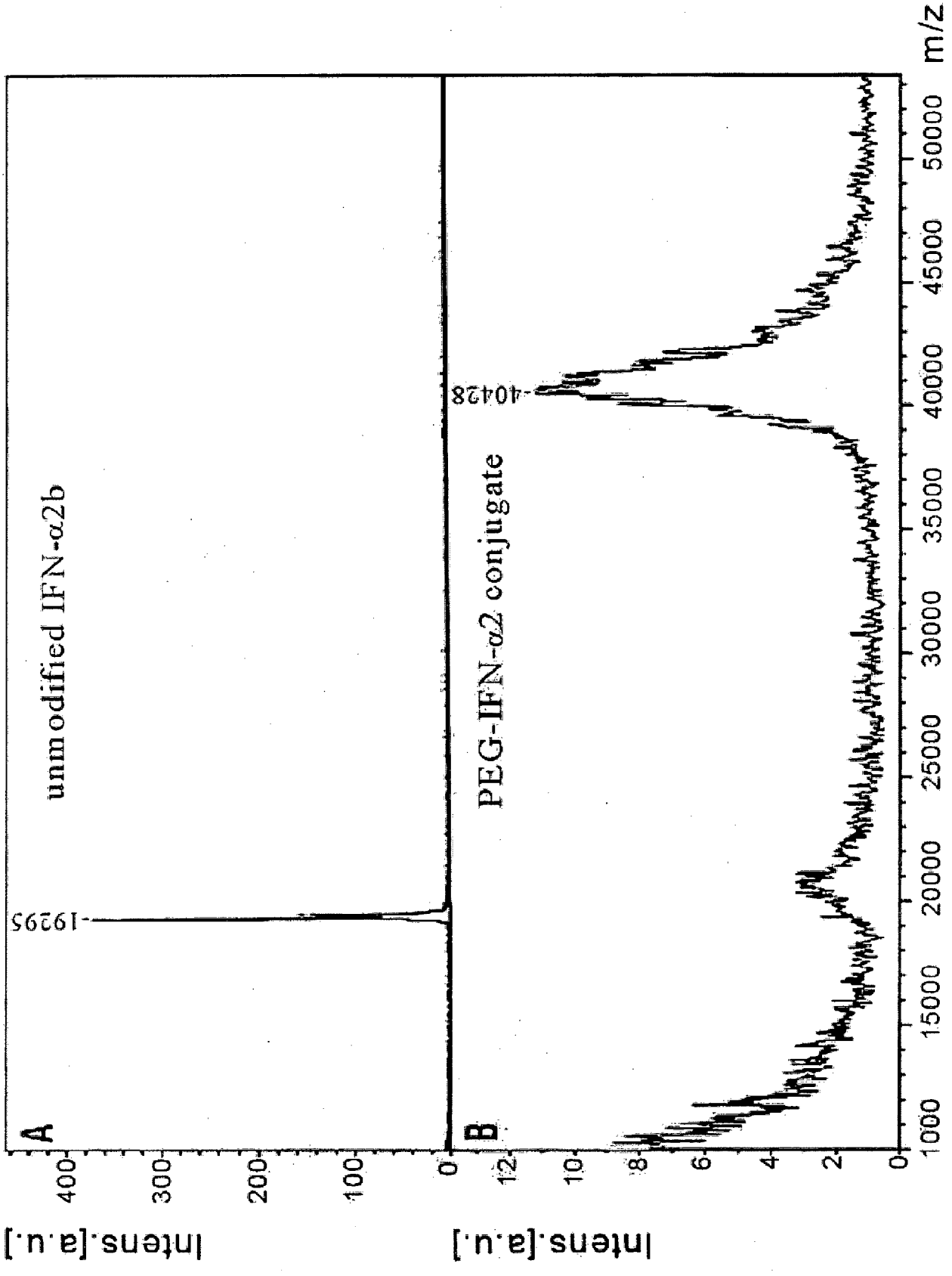

A
unmodified IFN-α2b
19295
Intens.[a.u.]
400
300
200
100
0
B
PEG-IFN-α2 conjugate
40428
Intens.[a.u.]
12
10
8
6
4
2
0
1000
15000
20000
25000
30000
35000
40000
45000
50000
m/z

Fig. 4

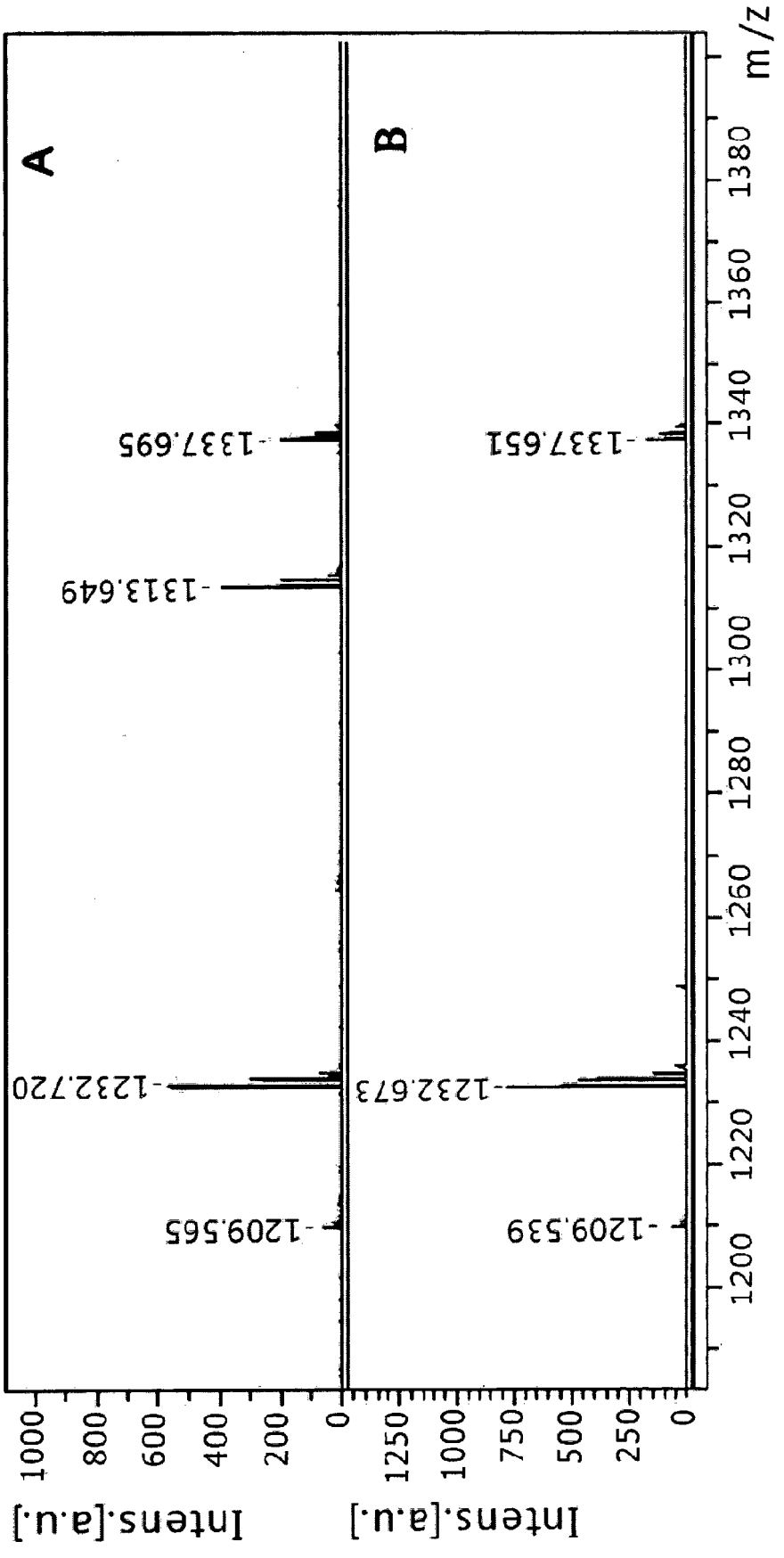

A
B
m/z
Intens.[a.u.]
1337.695
1313.649
1232.720
1209.565
1337.651
1232.673
1209.539

Fig. 5

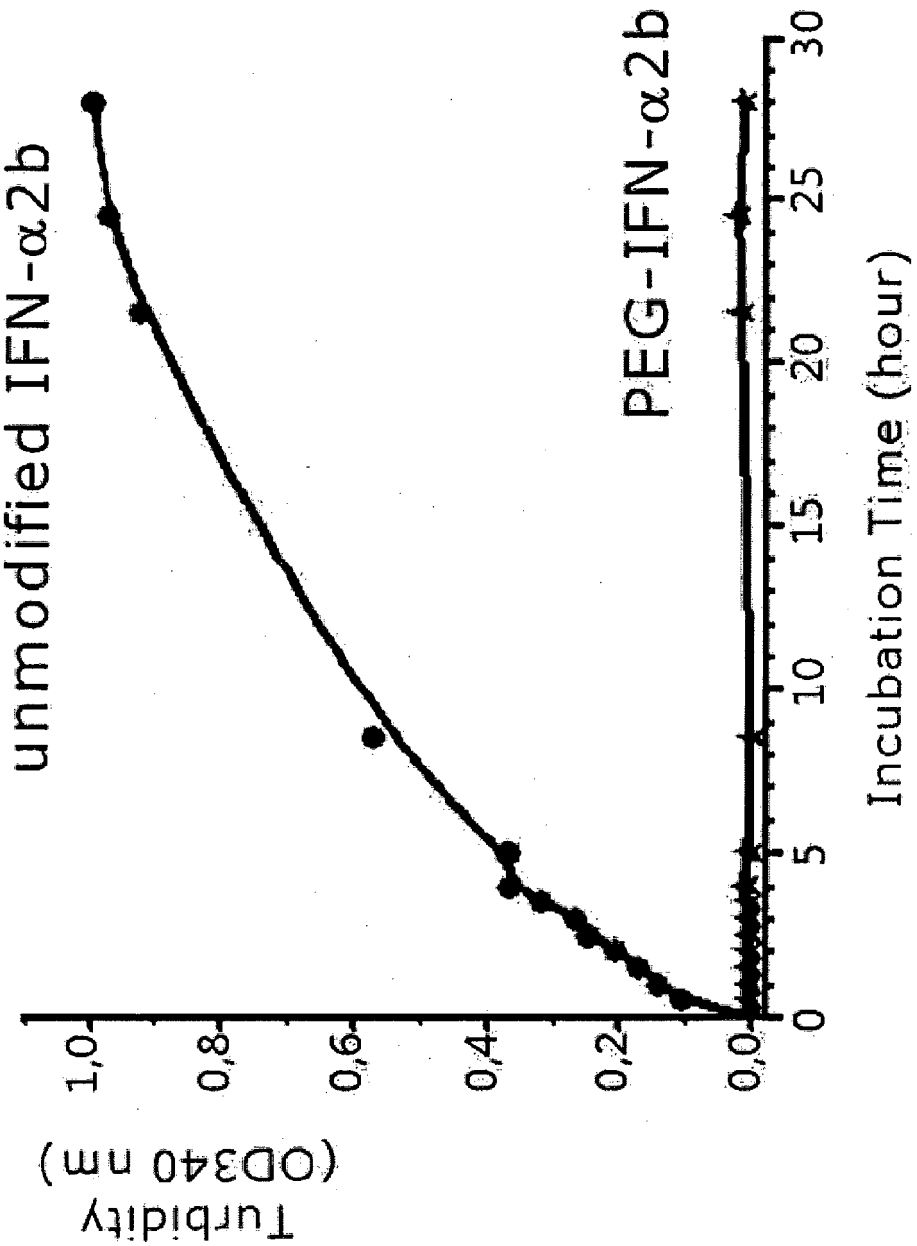

unmodified IFN-α2b
PEG-IFN-α2b
Turbidity (OD340 nm)
1,0
0,8
0,6
0,4
0,2
0,0
0
5
10
15
20
25
30
Incubation Time (hour)

Fig. 6

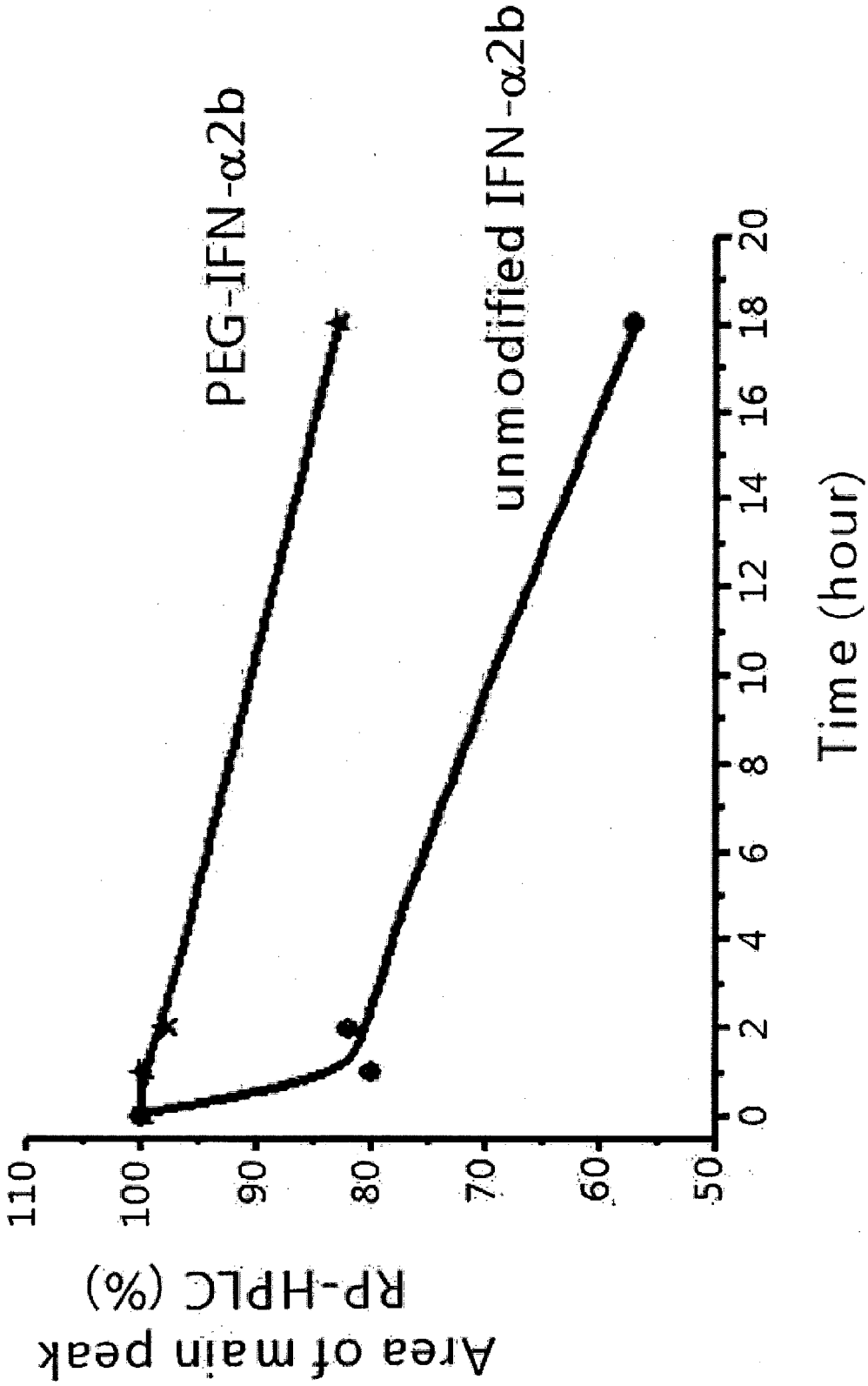

PEG-IFN-α2b
unmodified IFN-α2b
Area of main peak
RP-HPLC (%)
110
100
90
80
70
60
50
0
2
4
6
8
10
12
14
16
18
20
Time (hour)

Fig. 7

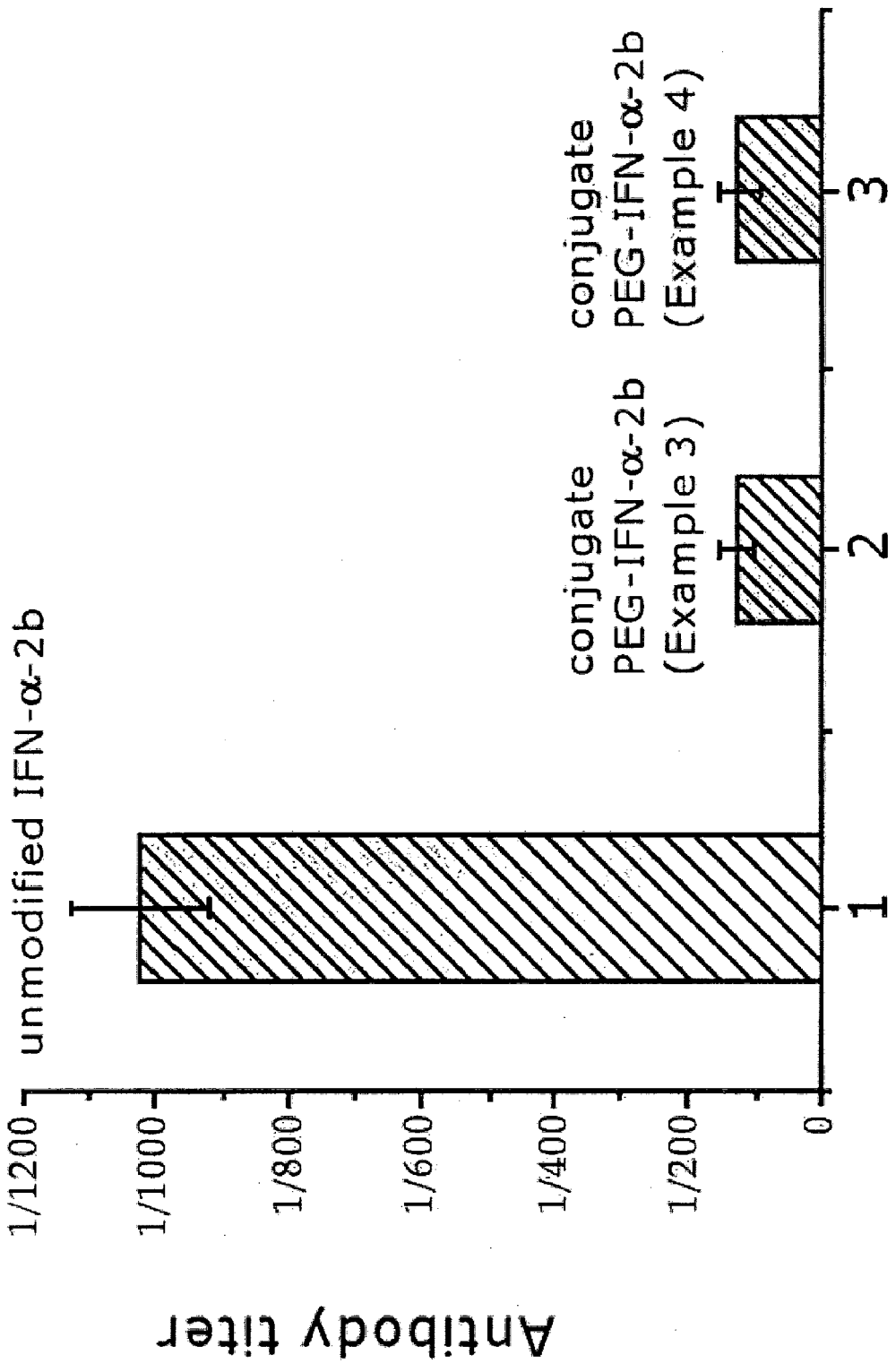

Antibody titer
1/1200
1/1000
1/800
1/600
1/400
1/200
0
1
2
3
unmodified IFN-α-2b
conjugate PEG-IFN-α-2b (Example 3)
conjugate PEG-IFN-α-2b (Example 4)

Fig. 8

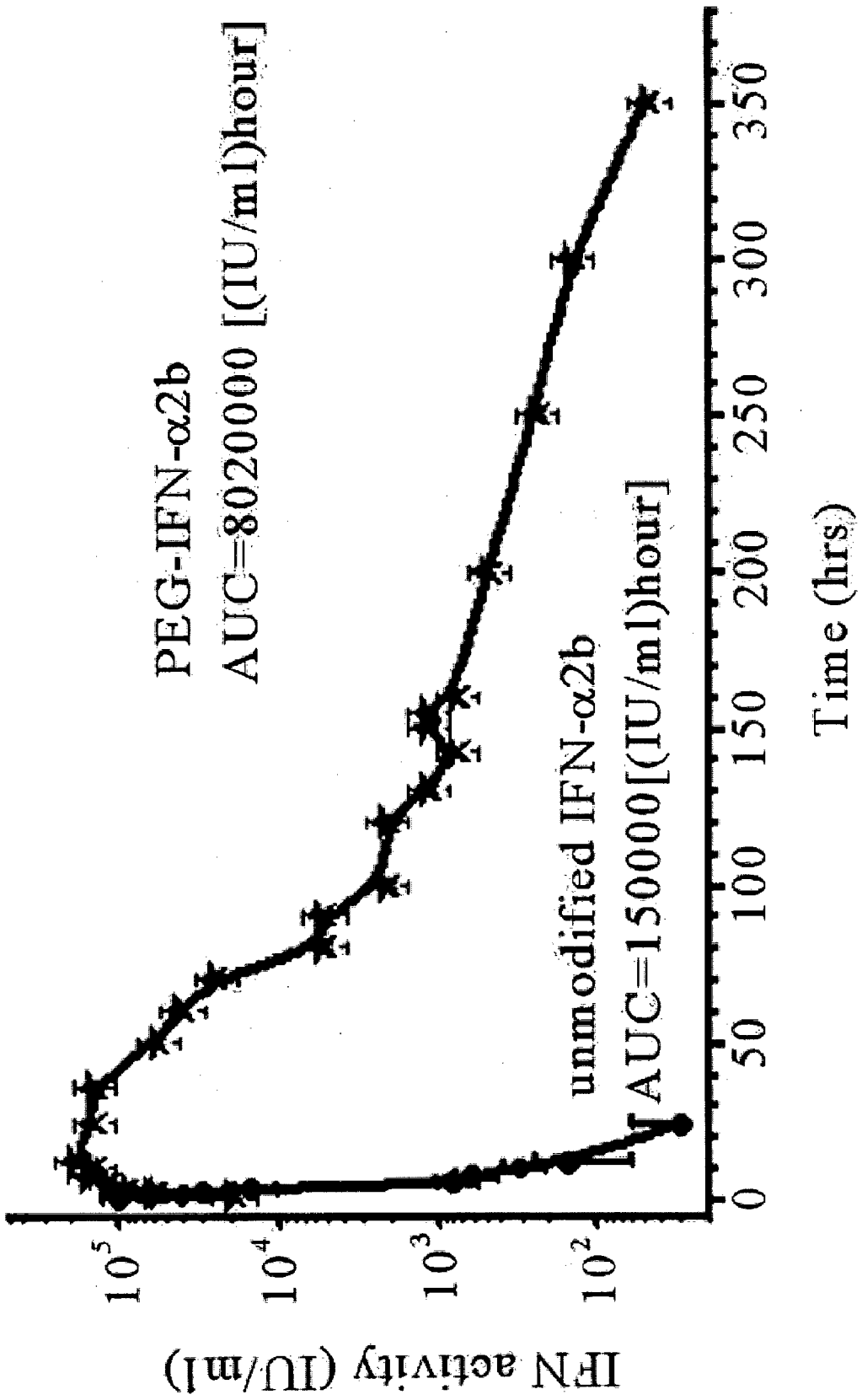

PEG-IFN-α2b
AUC=8020000 [(IU/ml)hour]
unmodified IFN-α2b
AUC=150000[(IU/ml)hour]
IFN activity (IU/ml)
Time (hrs)
10^5
10^4
10^3
10^2
0
50
100
150
200
250
300
350

Fig. 9