Fig. 1

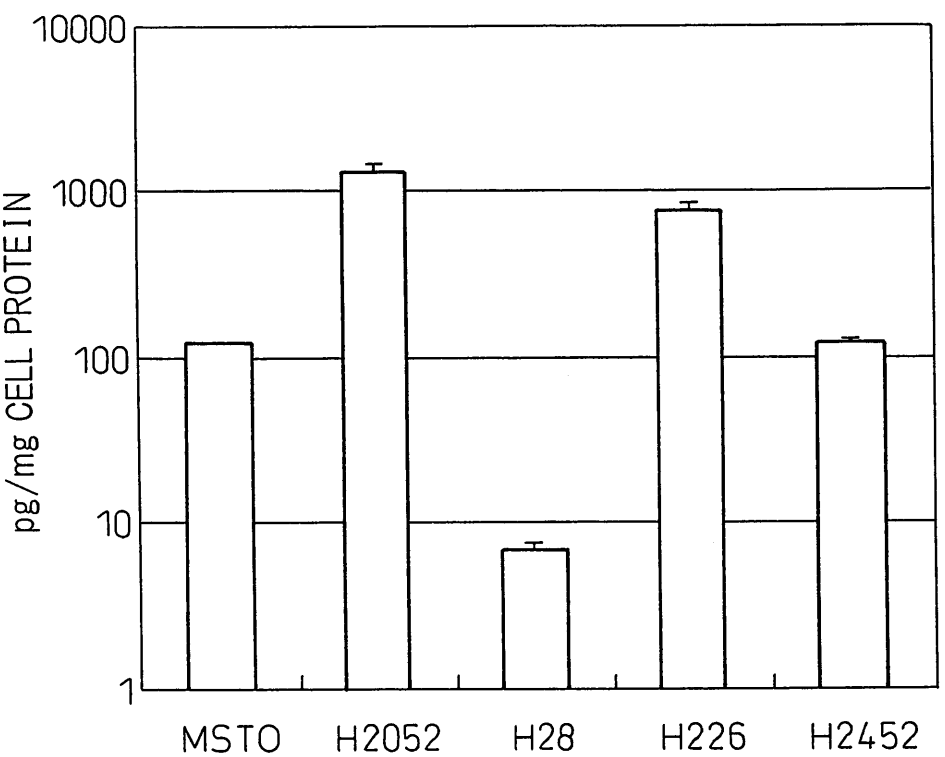

10000
1000
100
10
1
pg/mg CELL PROTEIN
MSTO
H2052
H28
H226
H2452

Fig.2

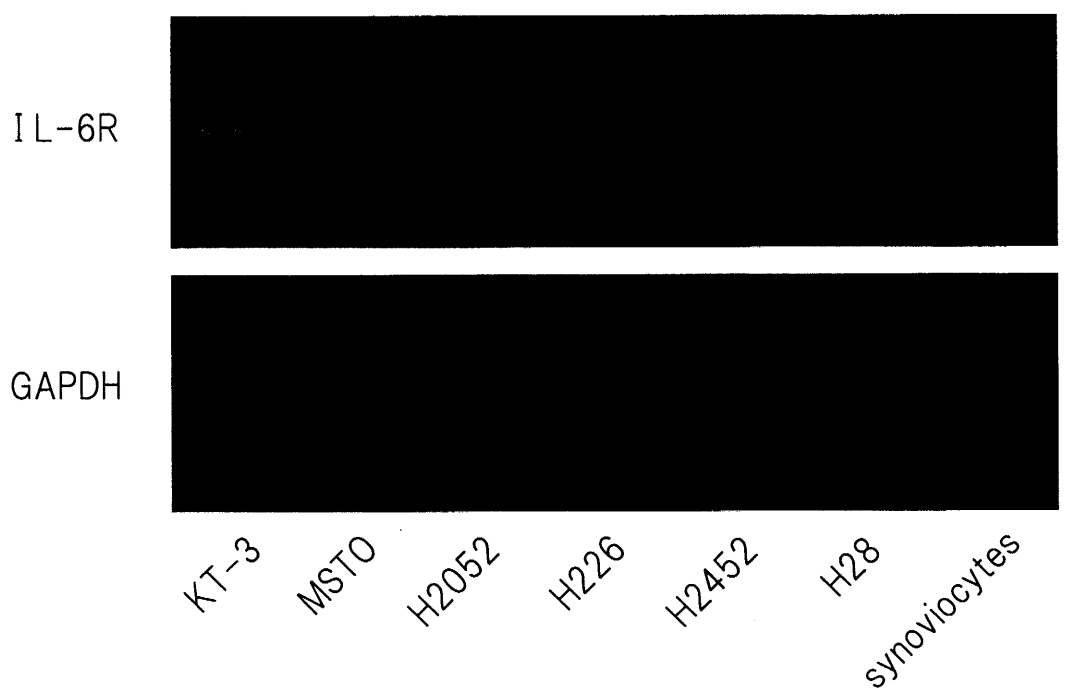


Fig.3

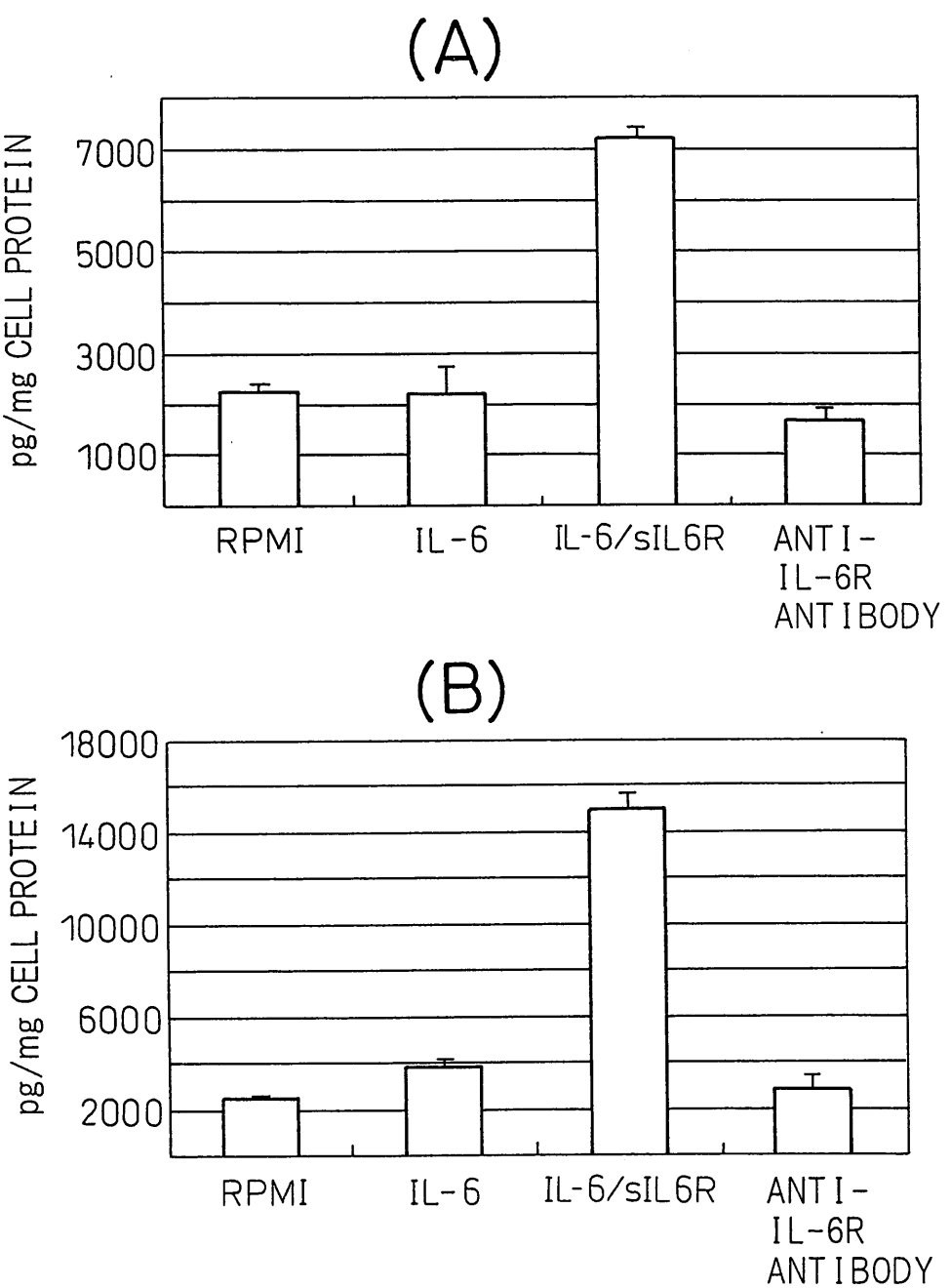

(A)
pg/mg CELL PROTEIN
7000
5000
3000
1000
RPMI
IL-6
IL-6/sIL6R
ANTI-IL-6R ANTIBODY
(B)
pg/mg CELL PROTEIN
18000
14000
10000
6000
2000
RPMI
IL-6
IL-6/sIL6R
ANTI-IL-6R ANTIBODY

Fig.4

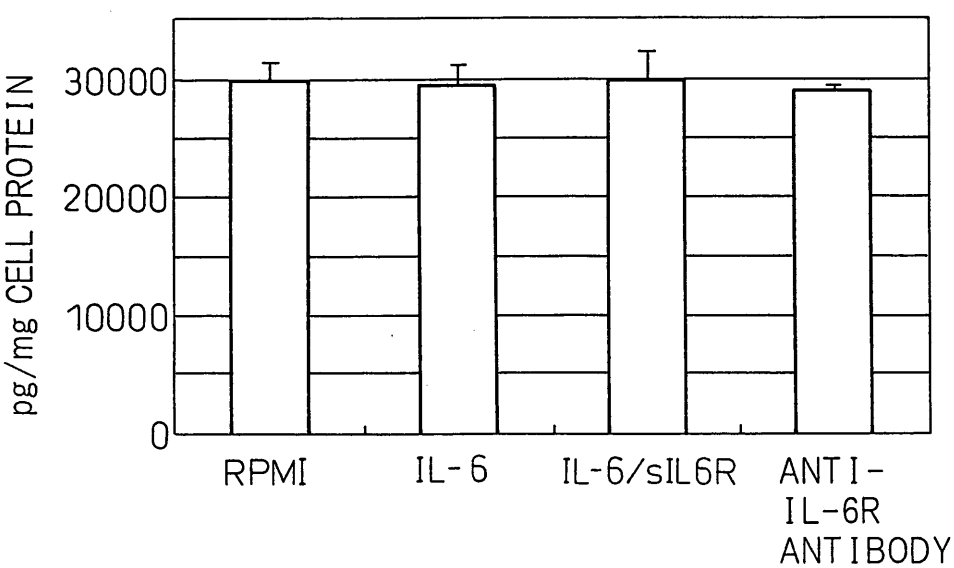

pg/mg CELL PROTEIN
30000
20000
10000
0
RPMI
IL-6
IL-6/sIL6R
ANTI-IL-6R ANTIBODY

Fig.5

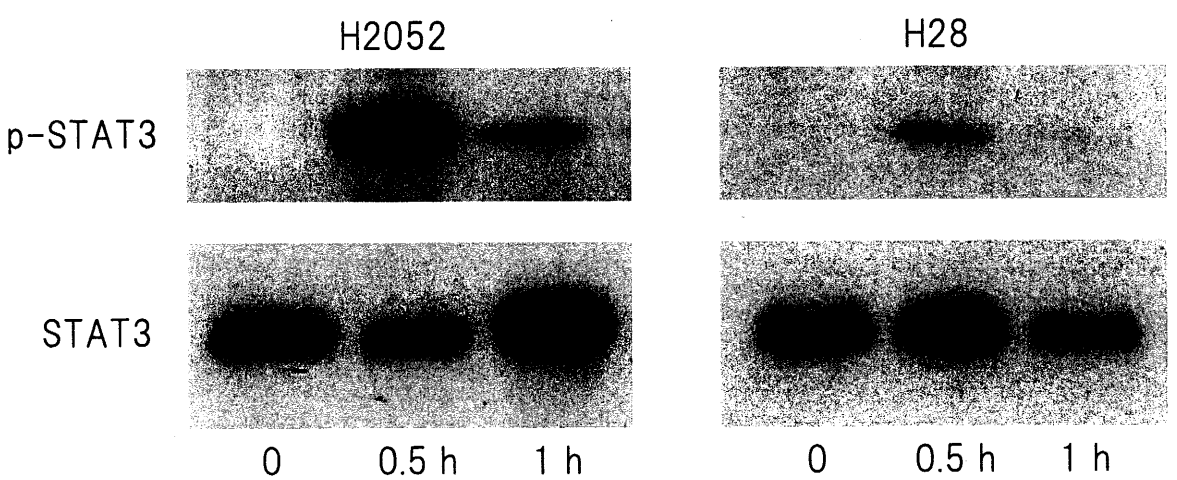


Fig.6

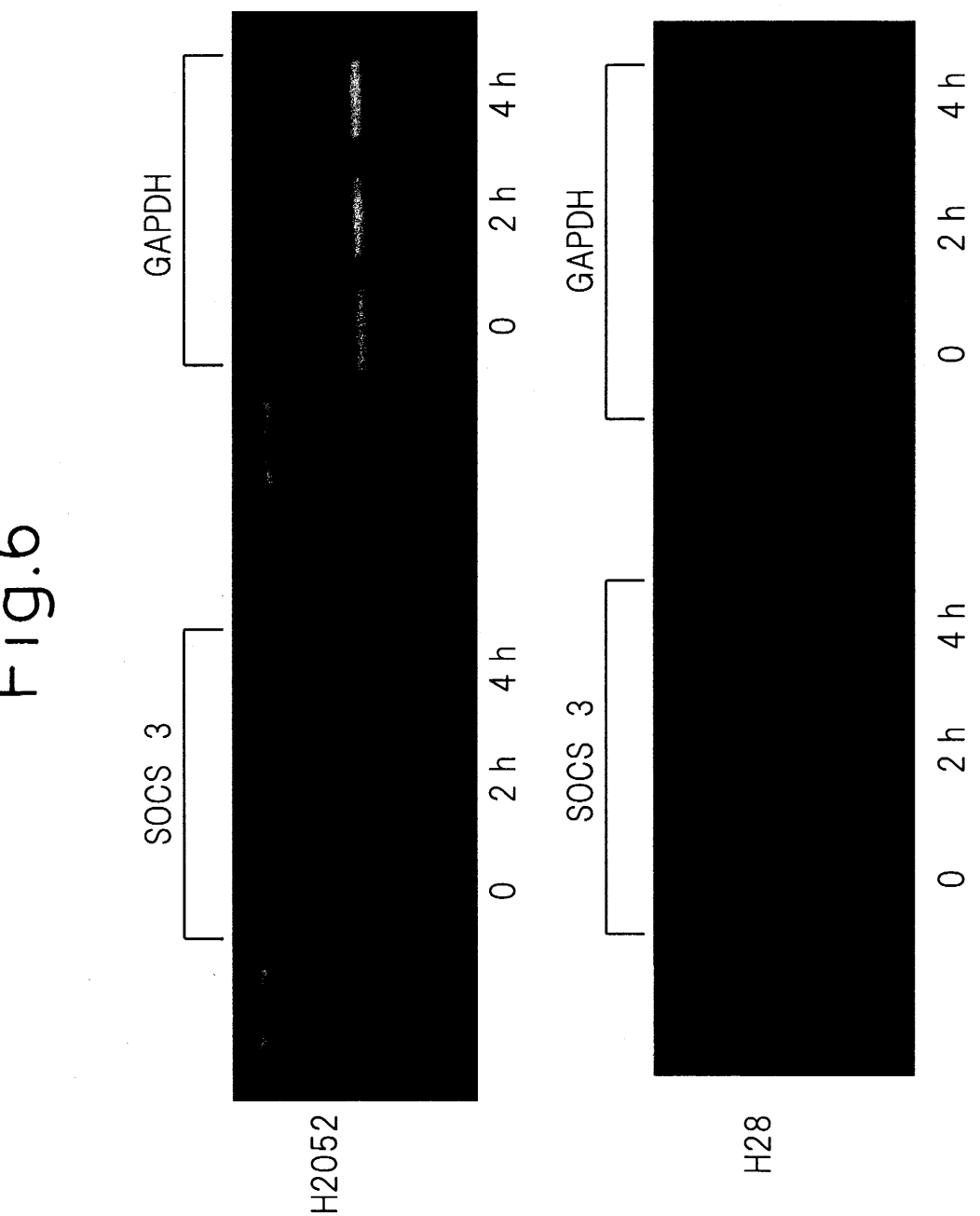


Fig.7

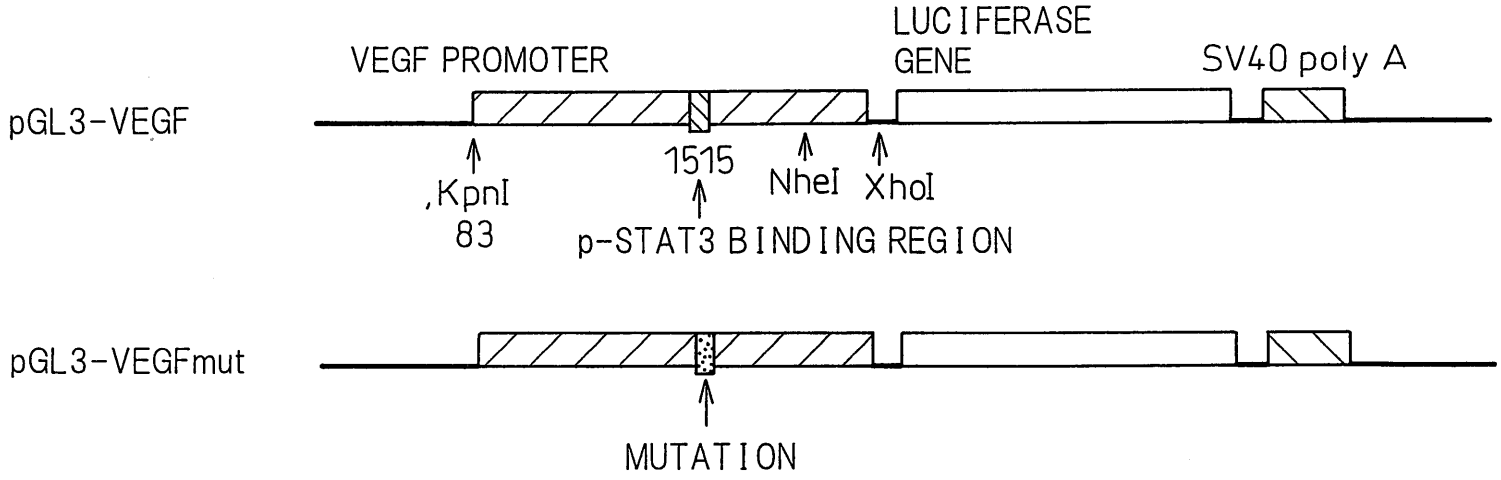

VEGF PROMOTER
LUCIFERASE GENE
SV40 poly A
pGL3-VEGF
,KpnI 83
1515
NheI
XhoI
p-STAT3 BINDING REGION
pGL3-VEGFmut
MUTATION

Fig.8

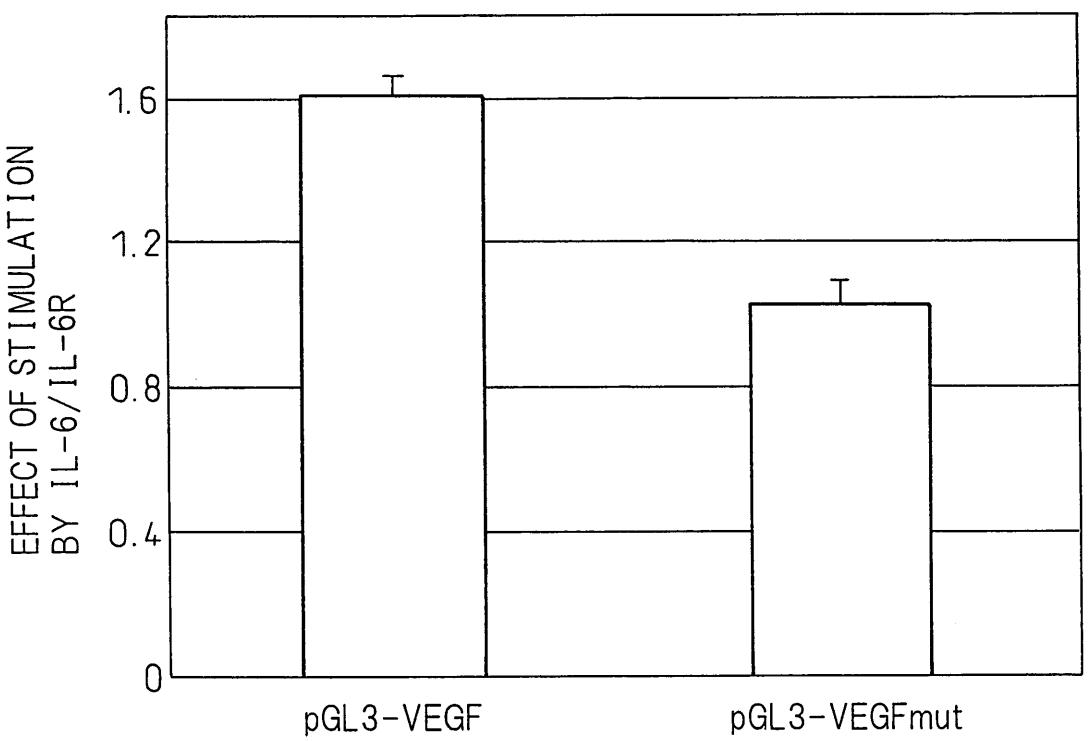

EFFECT OF STIMULATION
BY IL-6/IL-6R
1.6
1.2
0.8
0.4
0
pGL3-VEGF
pGL3-VEGFmut

Fig.9

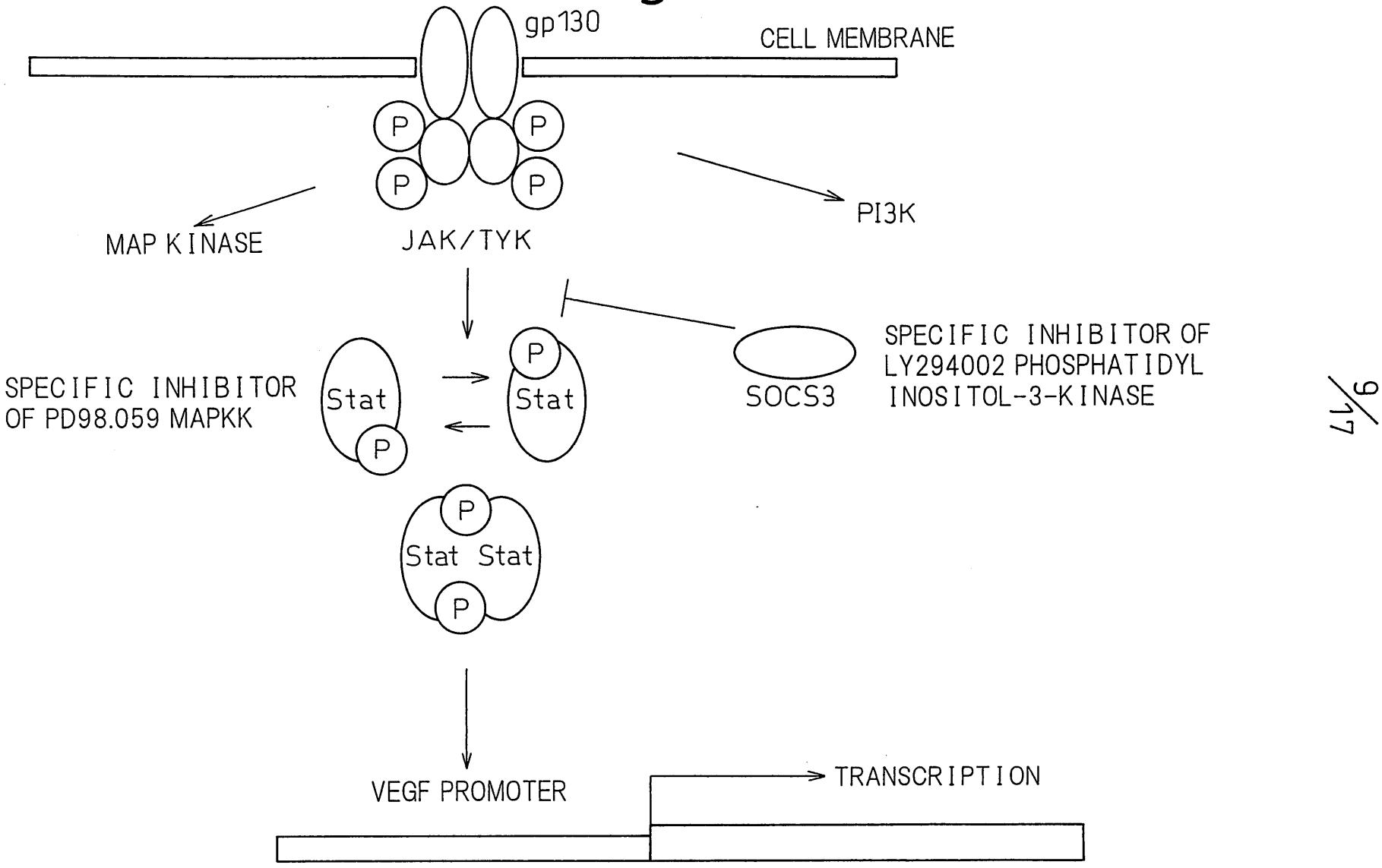

gp130
CELL MEMBRANE
P
P
P
P
MAP KINASE
JAK/TYK
PI3K
SPECIFIC INHIBITOR
OF PD98.059 MAPKK
Stat
P
P
Stat
SOCS3
SPECIFIC INHIBITOR OF
LY294002 PHOSPHATIDYL
INOSITOL-3-KINASE
P
Stat Stat
P
VEGF PROMOTER
TRANSCRIPTION

Fig.10

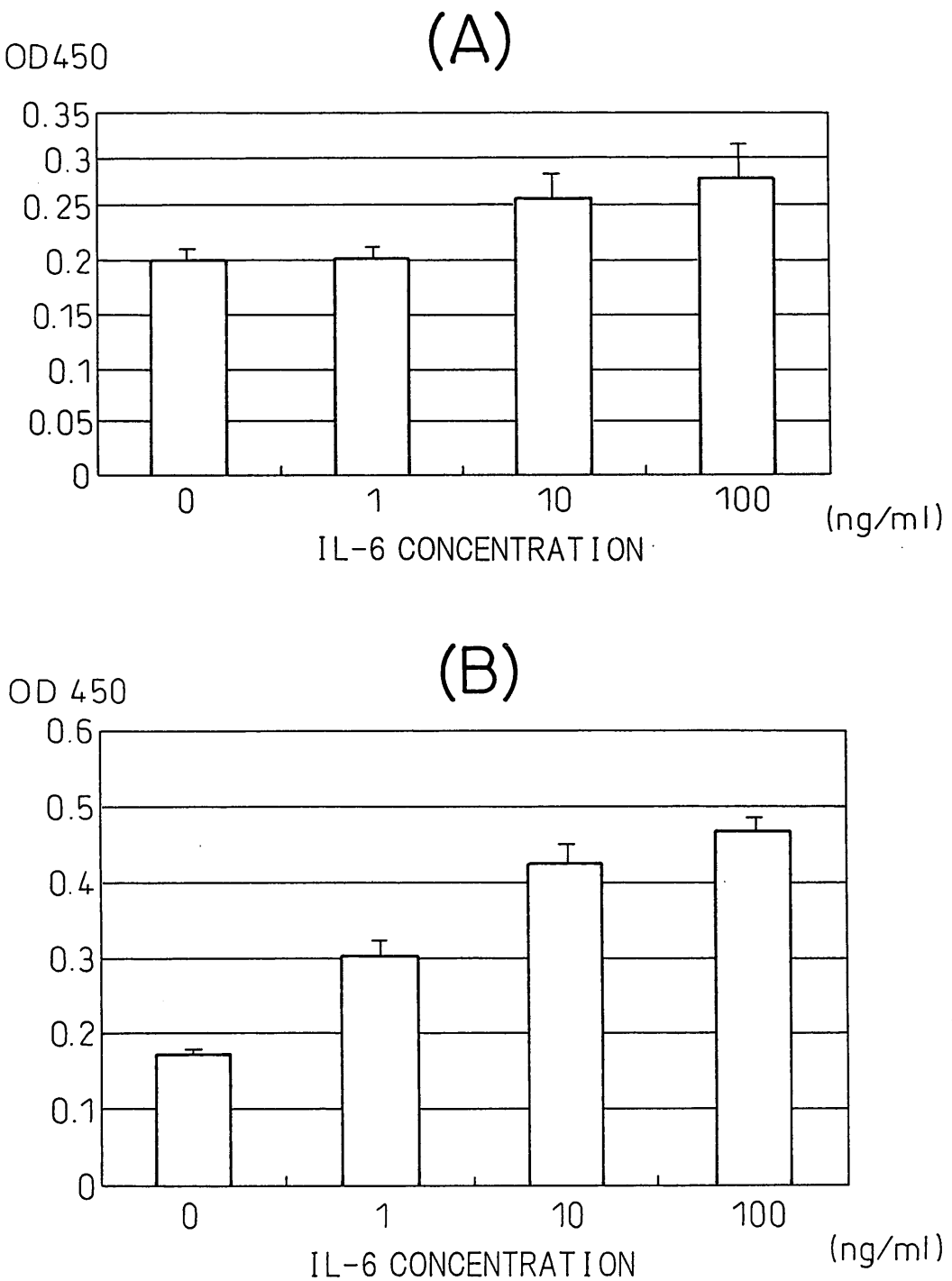
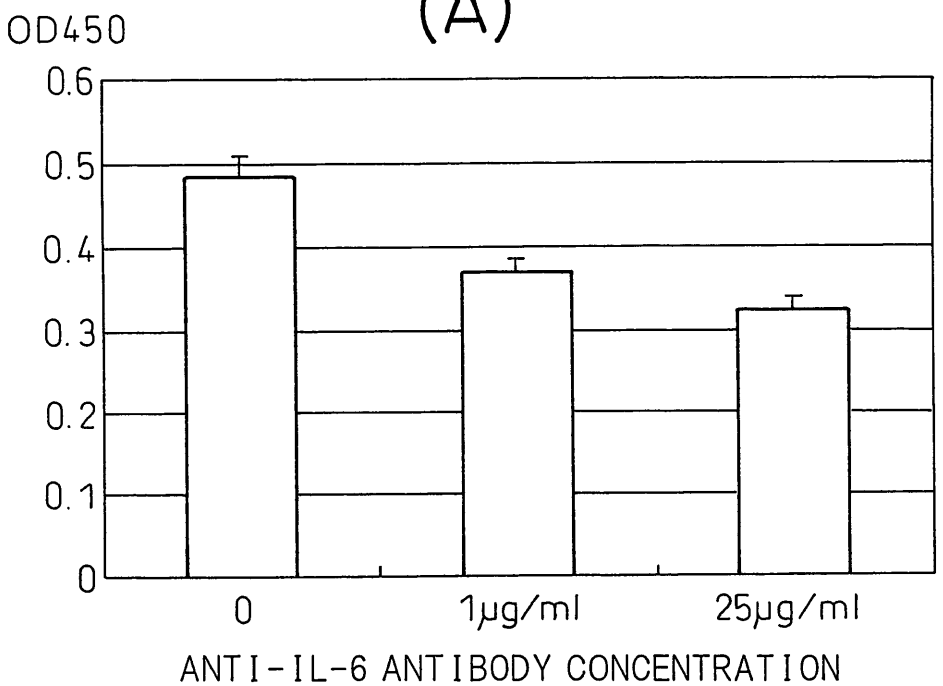

(A)
OD450
0.35
0.3
0.25
0.2
0.15
0.1
0.05
0
0
1
10
100
(ng/ml)
IL-6 CONCENTRATION
(B)
OD 450
0.6
0.5
0.4
0.3
0.2
0.1
0
0
1
10
100
(ng/ml)
IL-6 CONCENTRATION

Fig. 11

(A)


OD450
0.6
0.5
0.4
0.3
0.2
0.1
0
0
1μg/ml
25μg/ml
ANTI-IL-6 ANTIBODY CONCENTRATION

(B)

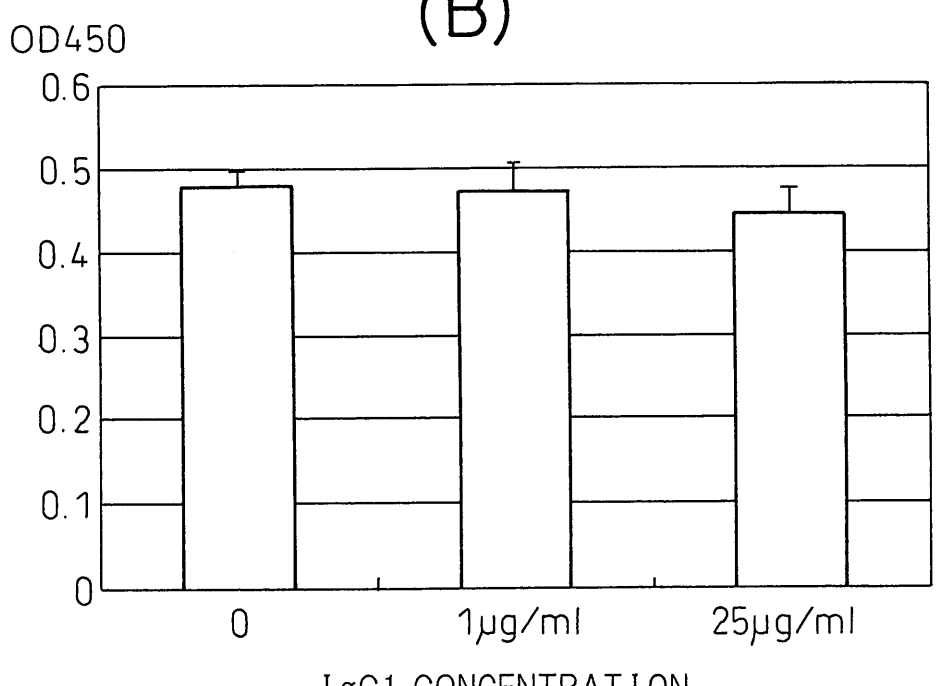

OD450
0.6
0.5
0.4
0.3
0.2
0.1
0
0
1μg/ml
25μg/ml
IgG1 CONCENTRATION

Fig. 12

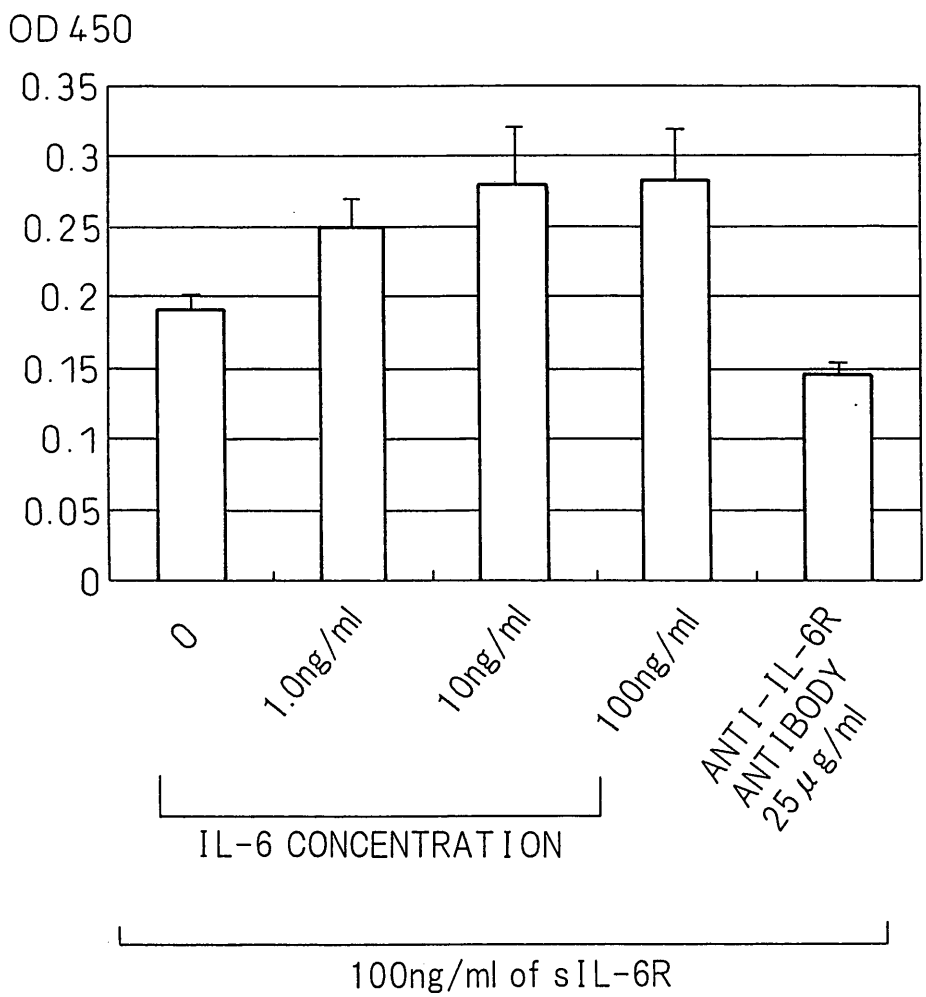

OD 450
0.35
0.3
0.25
0.2
0.15
0.1
0.05
0
0
1.0ng/ml
10ng/ml
100ng/ml
ANTI-IL-6R ANTIBODY 25μg/ml
IL-6 CONCENTRATION
100ng/ml of sIL-6R

Fig.13

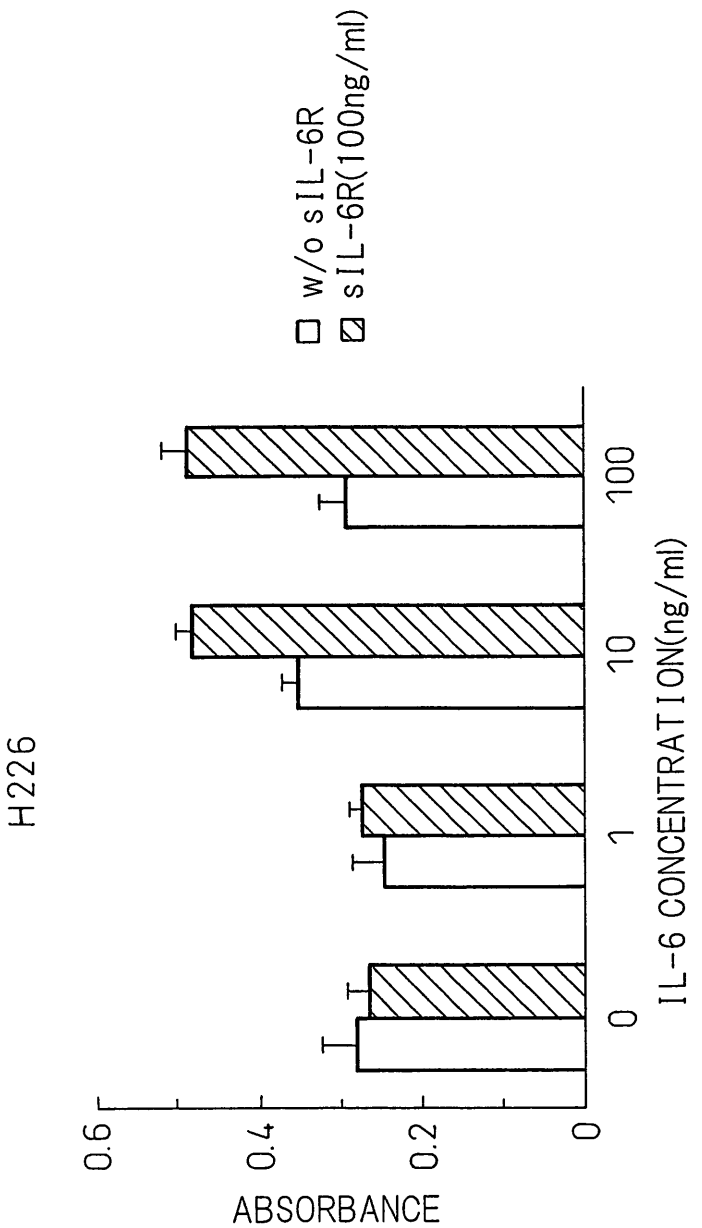

H226
w/o sIL-6R
sIL-6R(100ng/ml)
ABSORBANCE
0
0.2
0.4
0.6
0
1
10
100
IL-6 CONCENTRATION(ng/ml)

Fig.14

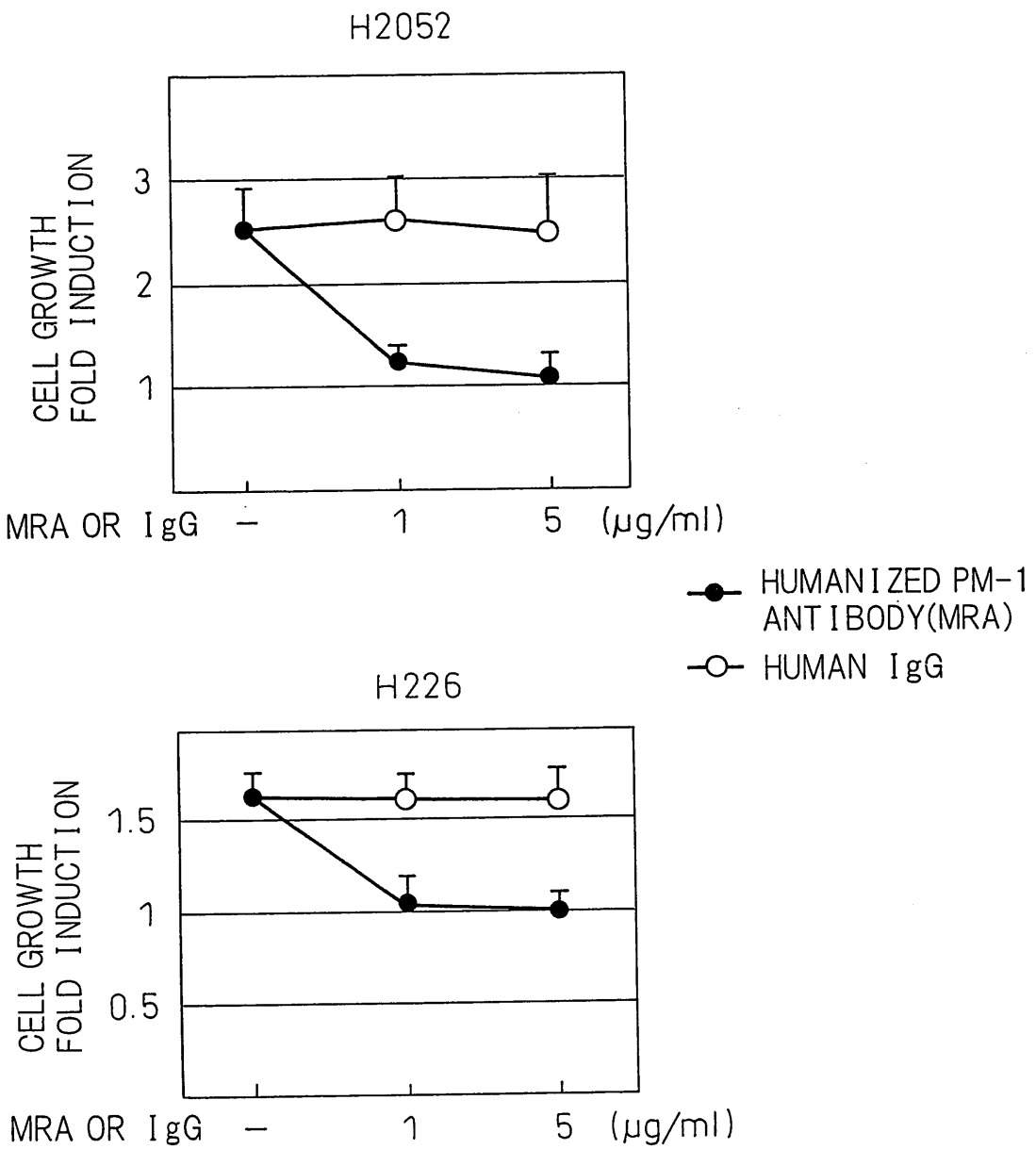

H2052
CELL GROWTH FOLD INDUCTION
3
2
1
MRA OR IgG
—
1
5
(μg/ml)
HUMANIZED PM-1 ANTIBODY(MRA)
HUMAN IgG
H226
CELL GROWTH FOLD INDUCTION
1.5
1
0.5
MRA OR IgG
—
1
5
(μg/ml)

Fig.15

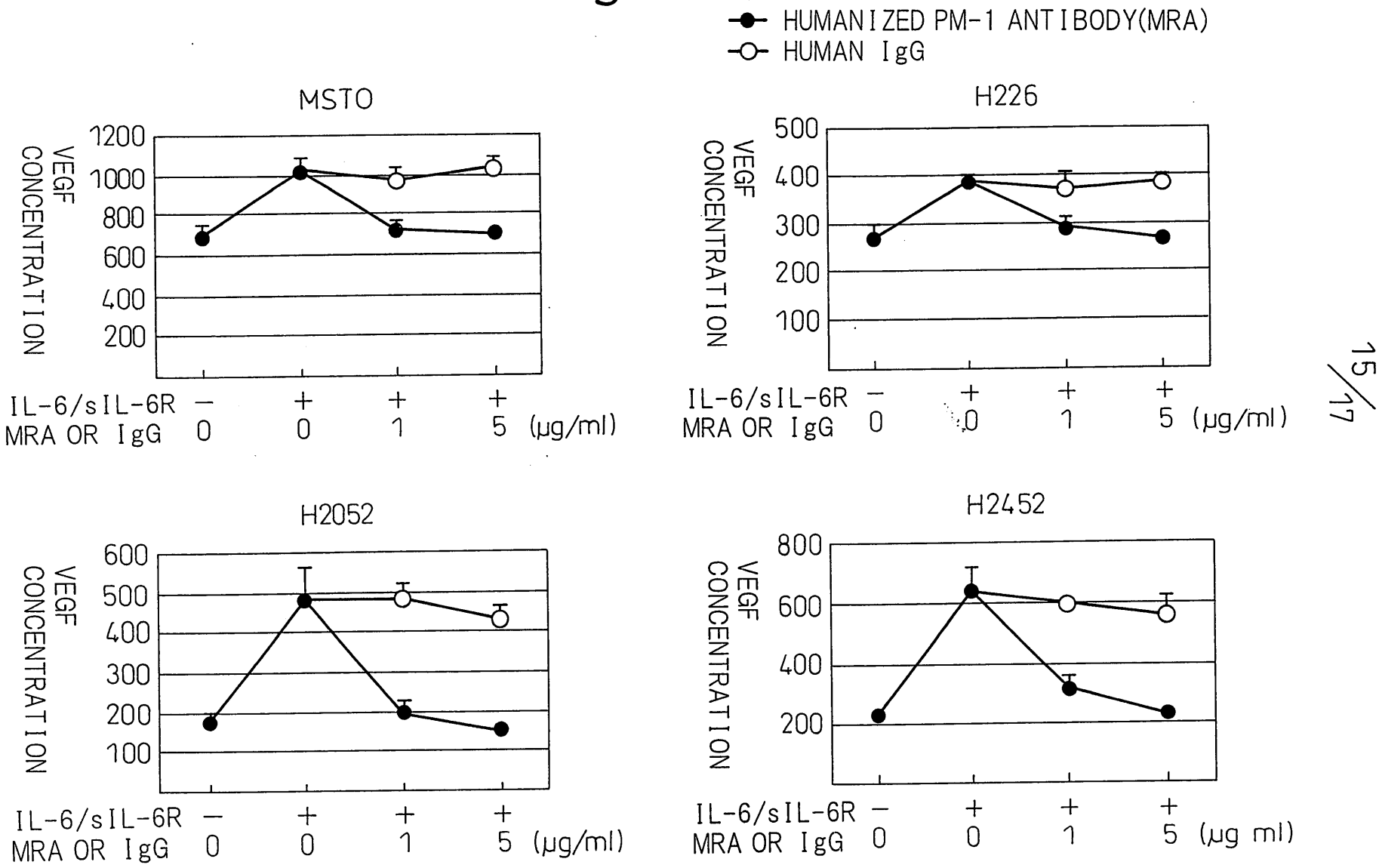

HUMANIZED PM-1 ANTIBODY(MRA)
HUMAN IgG
MSTO
H226
H2052
H2452
VEGF CONCENTRATION
IL-6/sIL-6R
MRA OR IgG
(μg/ml)

Fig.16

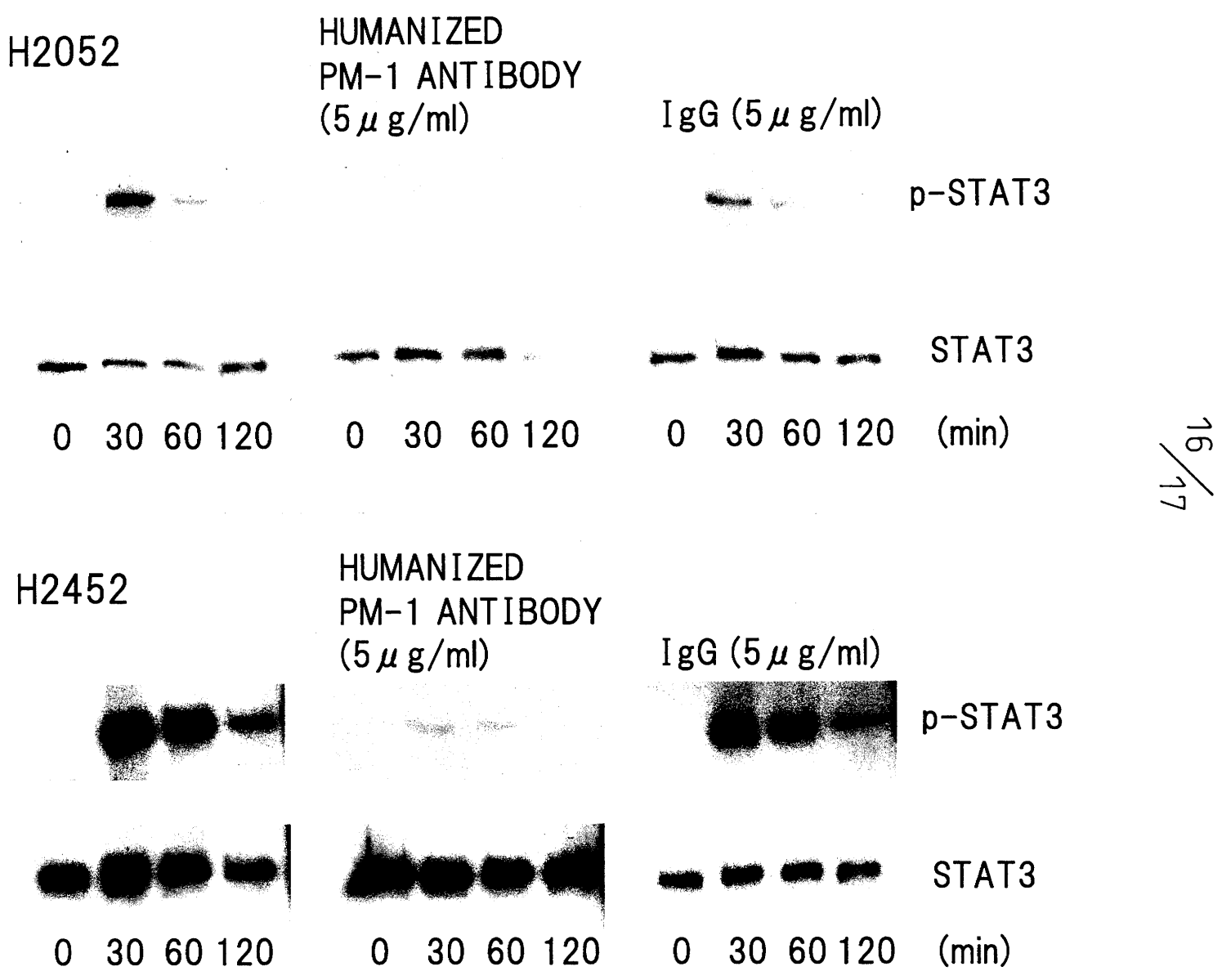
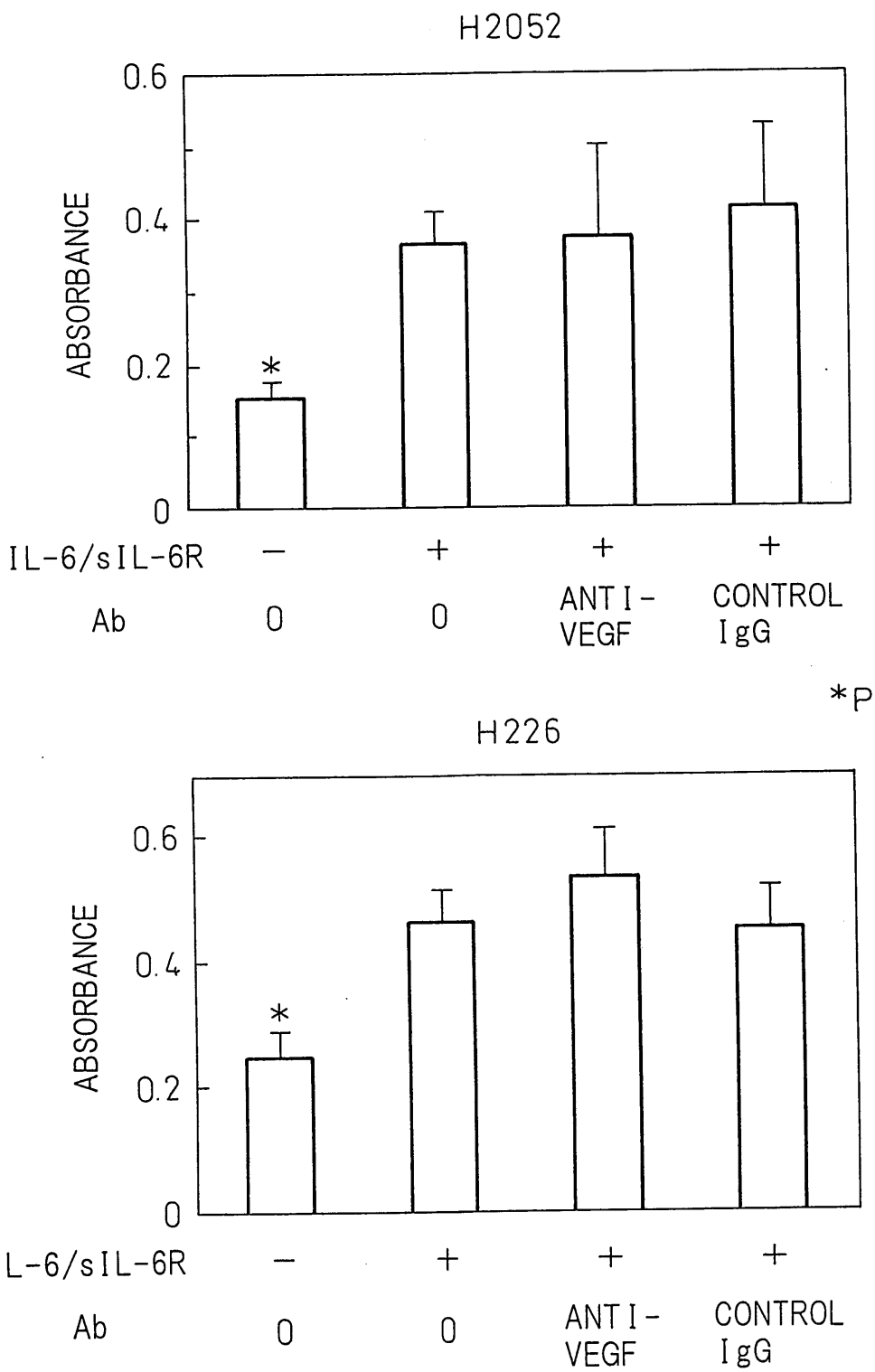

H2052
HUMANIZED
PM-1 ANTIBODY
(5μg/ml)
IgG (5μg/ml)
p-STAT3
STAT3
0 30 60 120
0 30 60 120
0 30 60 120
(min)
H2452
HUMANIZED
PM-1 ANTIBODY
(5μg/ml)
IgG (5μg/ml)
p-STAT3
STAT3
0 30 60 120
0 30 60 120
0 30 60 120
(min)

Fig.17


H2052
ABSORBANCE
0.6
0.4
0.2
0
*
IL-6/sIL-6R
−
+
+
+
Ab
0
0
ANTI-VEGF
CONTROL IgG
*P<0.05
H226
ABSORBANCE
0.6
0.4
0.2
0
*
IL-6/sIL-6R
−
+
+
+
Ab
0
0
ANTI-VEGF
CONTROL IgG