Conventional T Cells (2)
Virgin Professional Regulatory Cells (<1% Total) (3)
Naive T Cell Population (1)
Regulatory Composition
2
Activated Professional Regulatory Cells (4) (>50% of Total)
Treated T Cell Population (6)
FIG_1A
2
4
6
Conventional T Cells
T Cell Activator
4
2
Cytokine-producing Regulatory T Cells (5)
FIG_1B
2
Conventional T Cell Population (7)
Regulatory Composition
5
FIG_1C

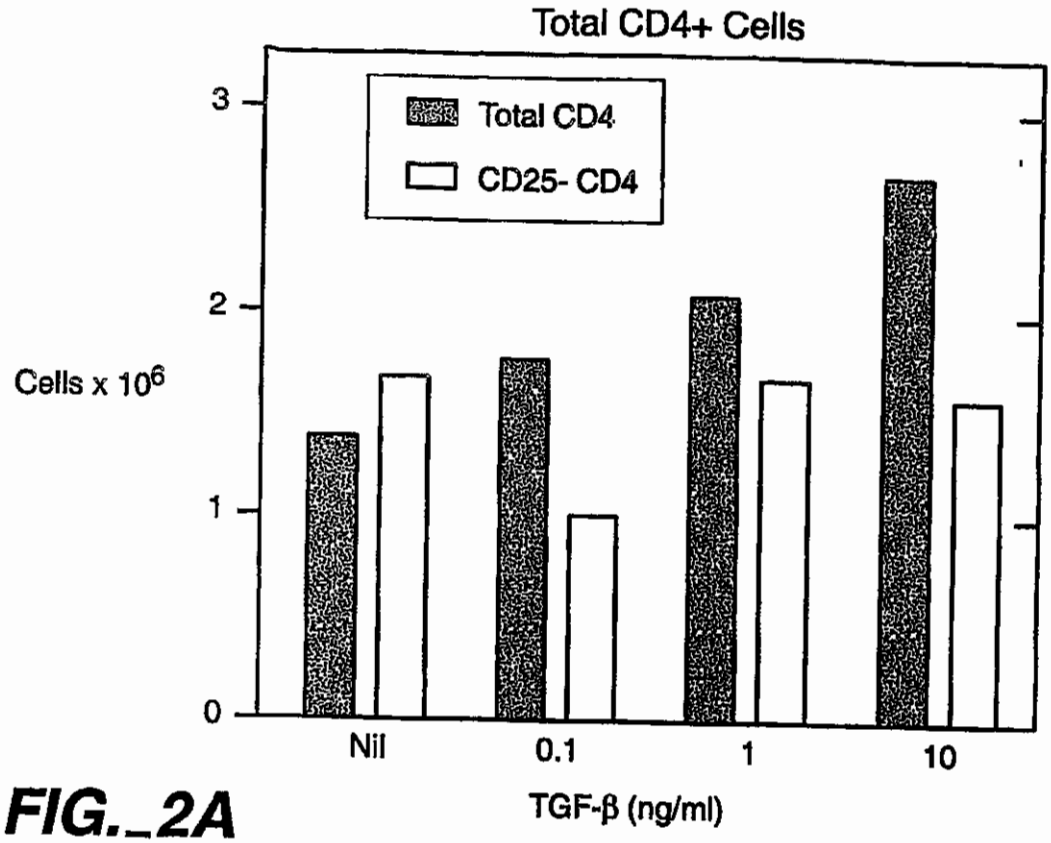

Total CD4+ Cells
Total CD4
CD25- CD4
Cells x 10^6
3
2
1
0
Nil
0.1
1
10
TGF-β (ng/ml)

FIG._2A

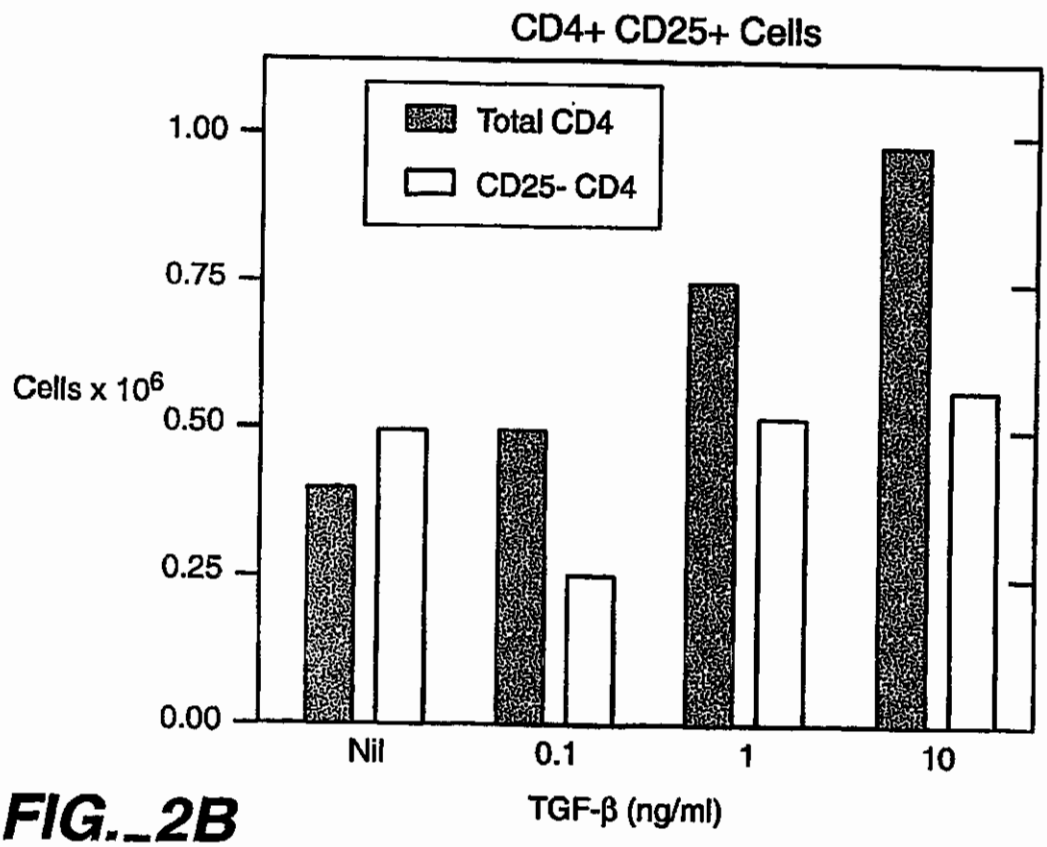

CD4+ CD25+ Cells
Total CD4
CD25- CD4
Cells x 10^6
1.00
0.75
0.50
0.25
0.00
Nil
0.1
1
10
TGF-β (ng/ml)

FIG._2B

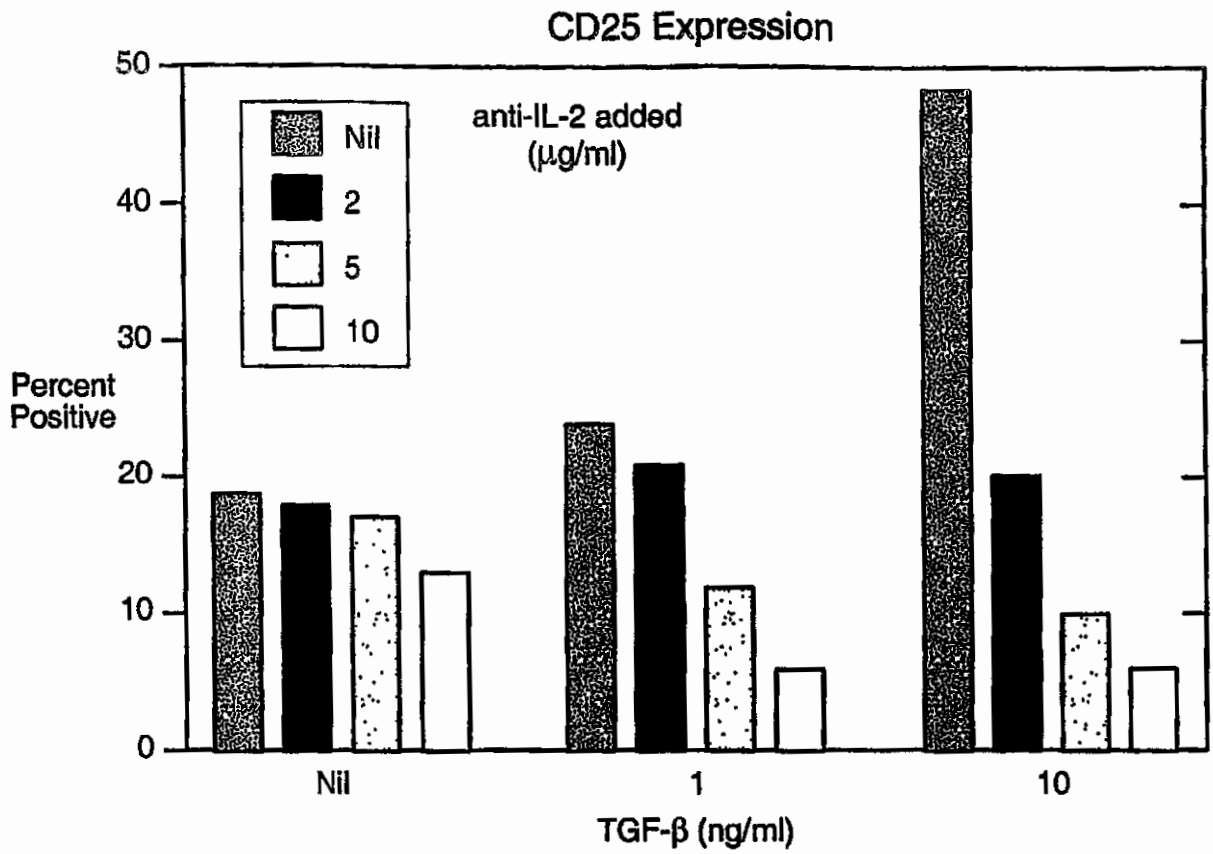

CD25 Expression
anti-IL-2 added
(μg/ml)
Nil
2
5
10
Percent
Positive
50
40
30
20
10
0
Nil
1
10
TGF-β (ng/ml)

FIG._3

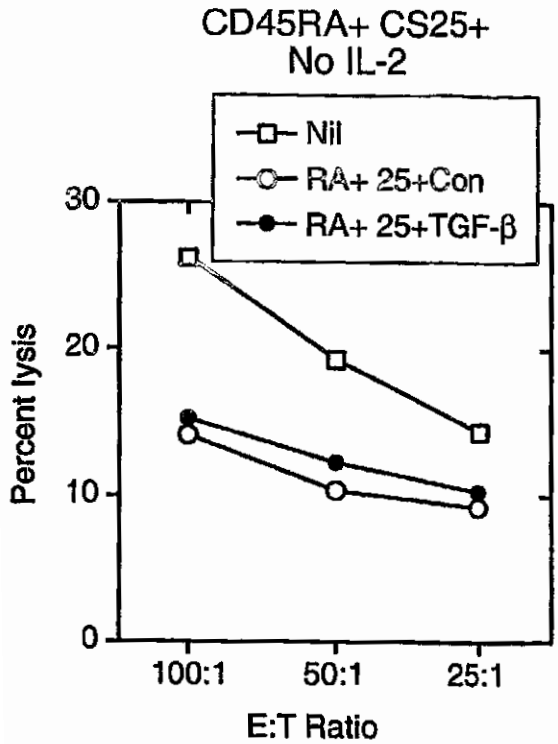


FIG._4A

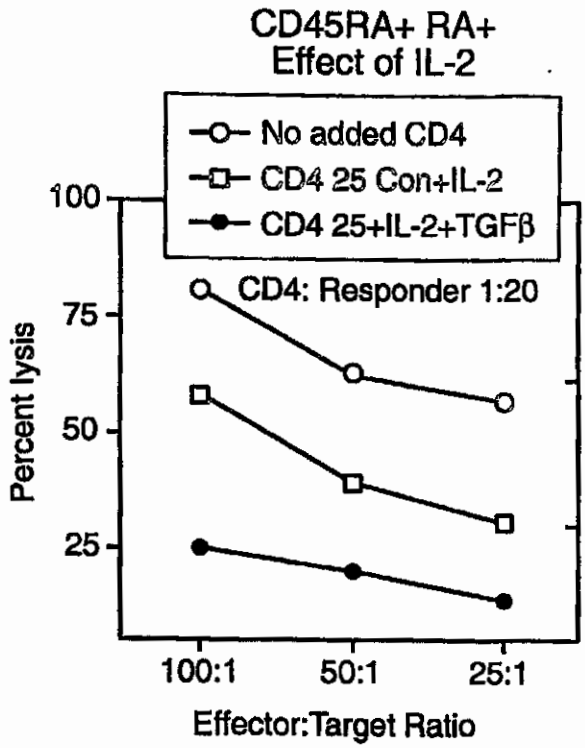


FIG._4B

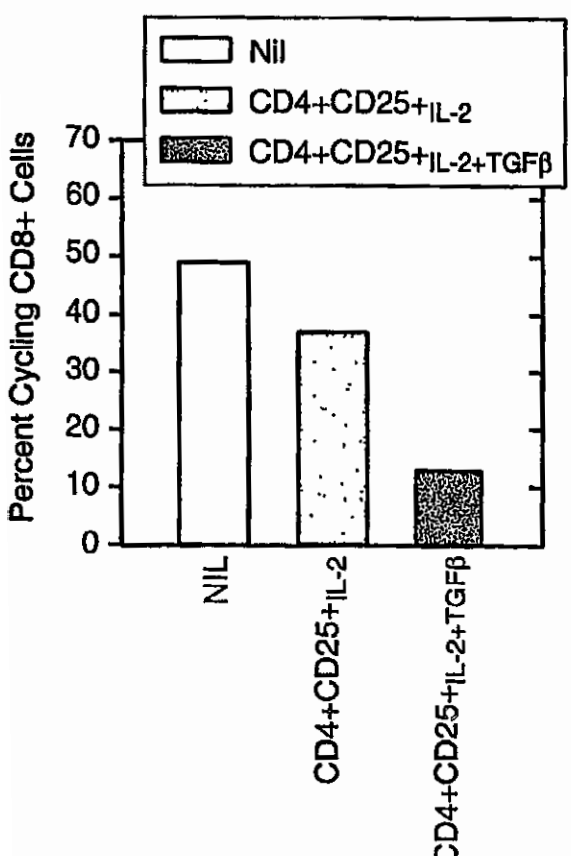


FIG._4C

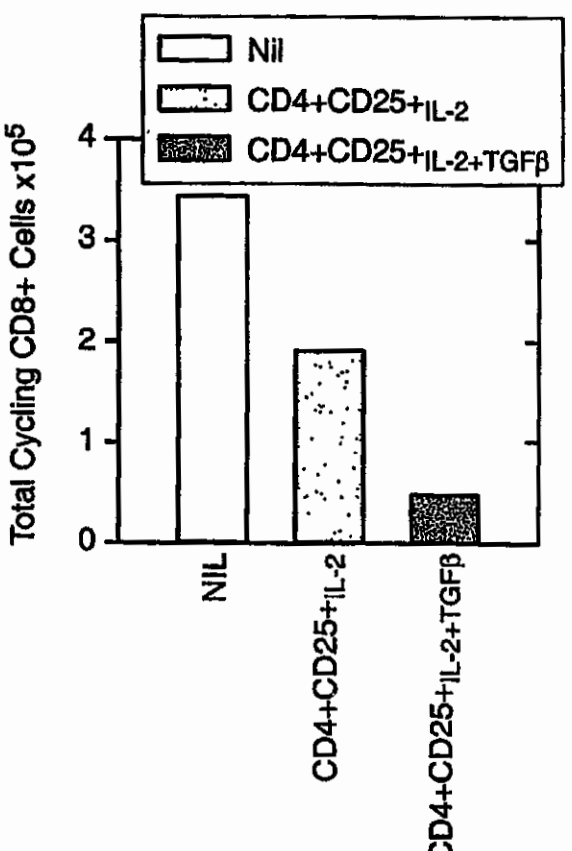


FIG._4D

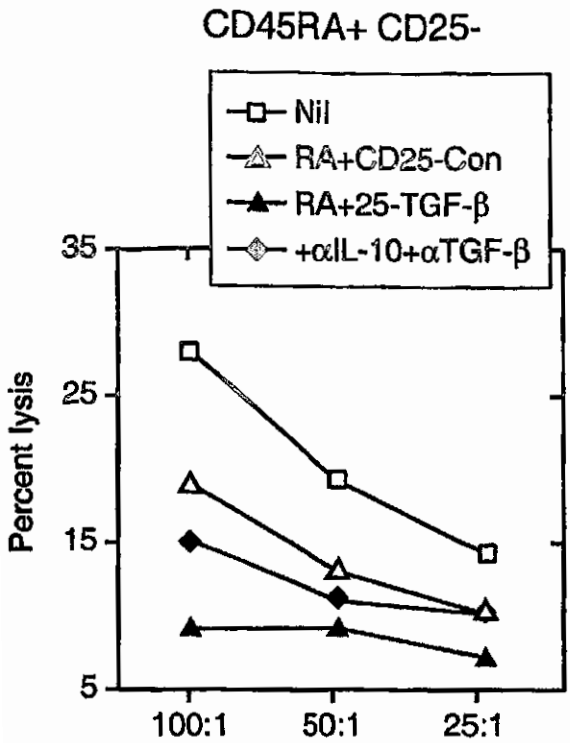


FIG._4E

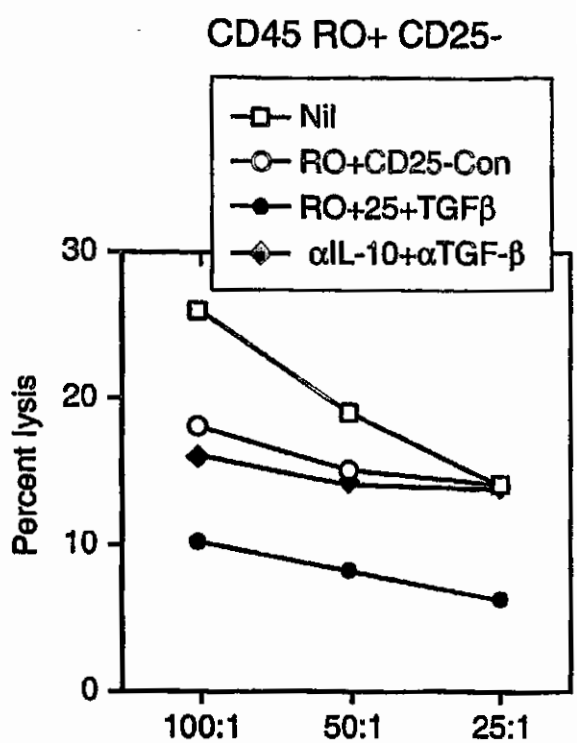


FIG._4F

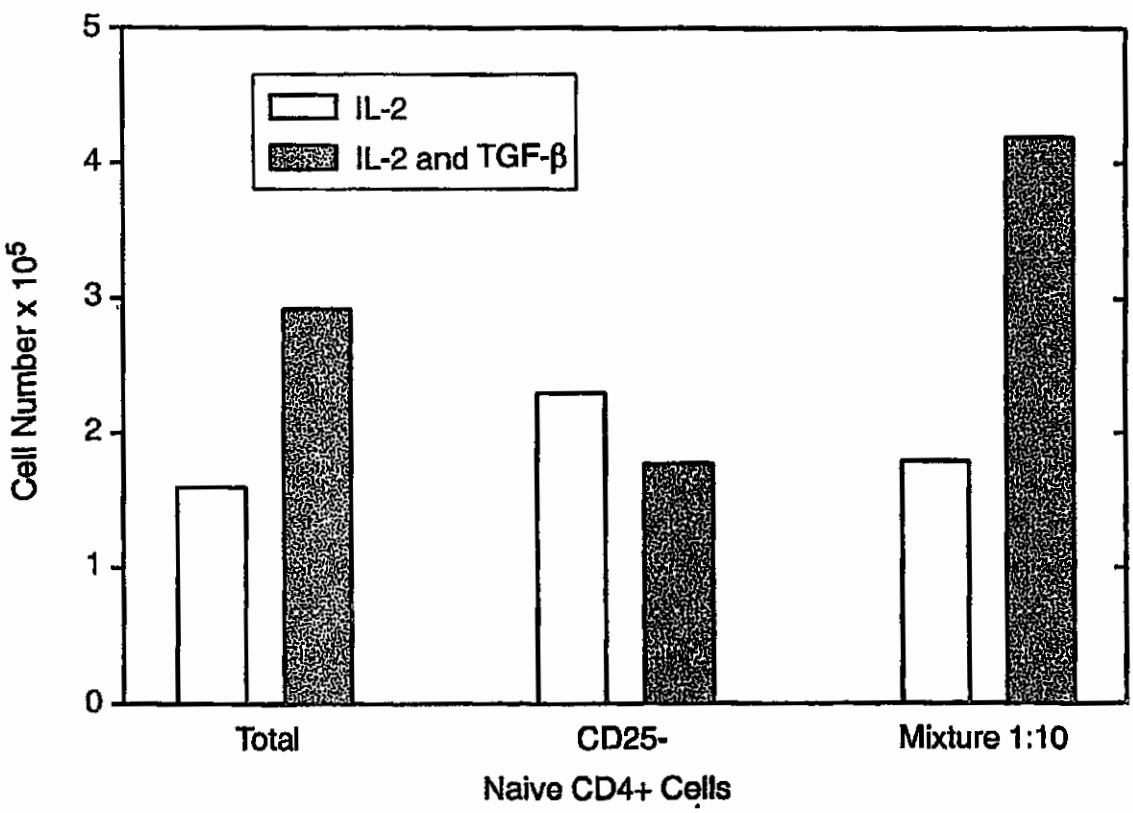


FIG._5A

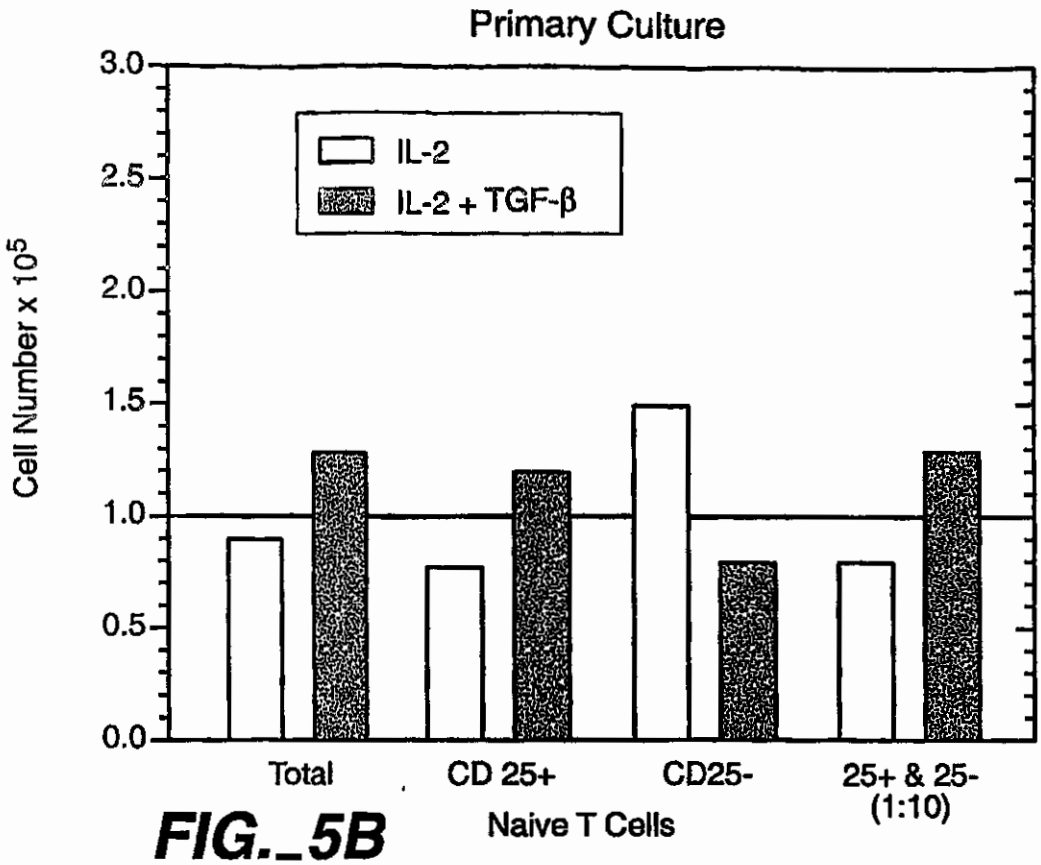


FIG._5B

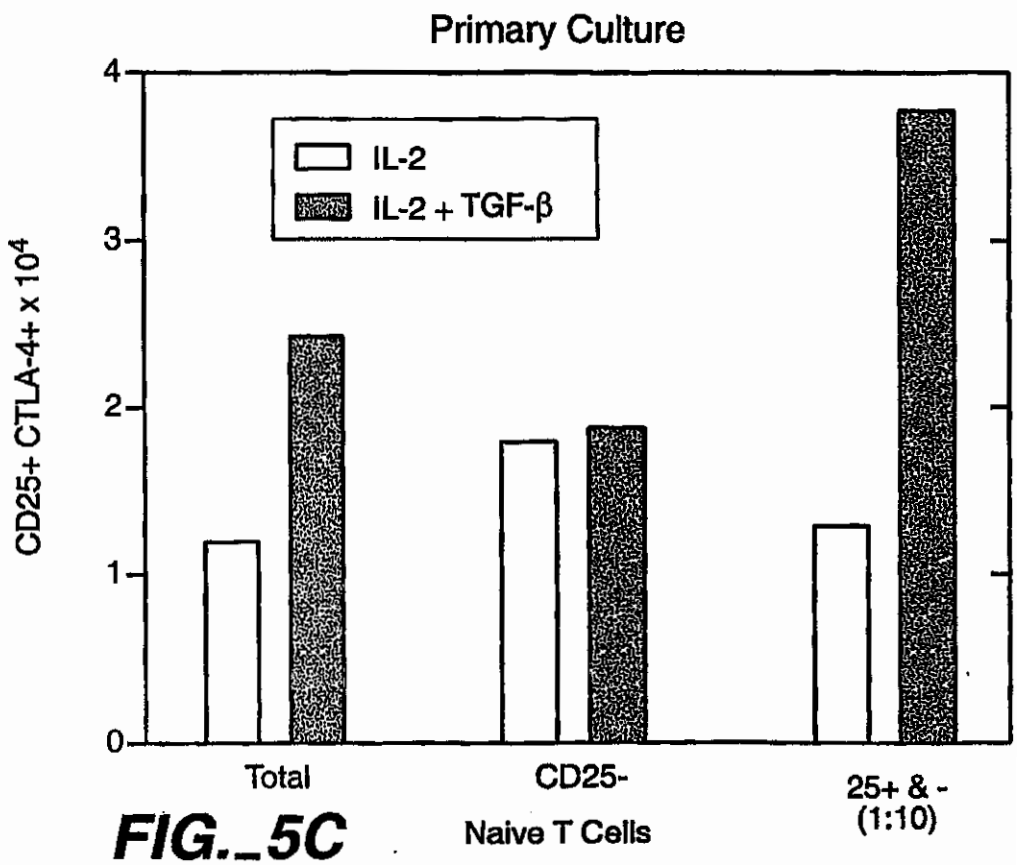


FIG._5C

SUBSTITUTE SHEET (RULE 26)

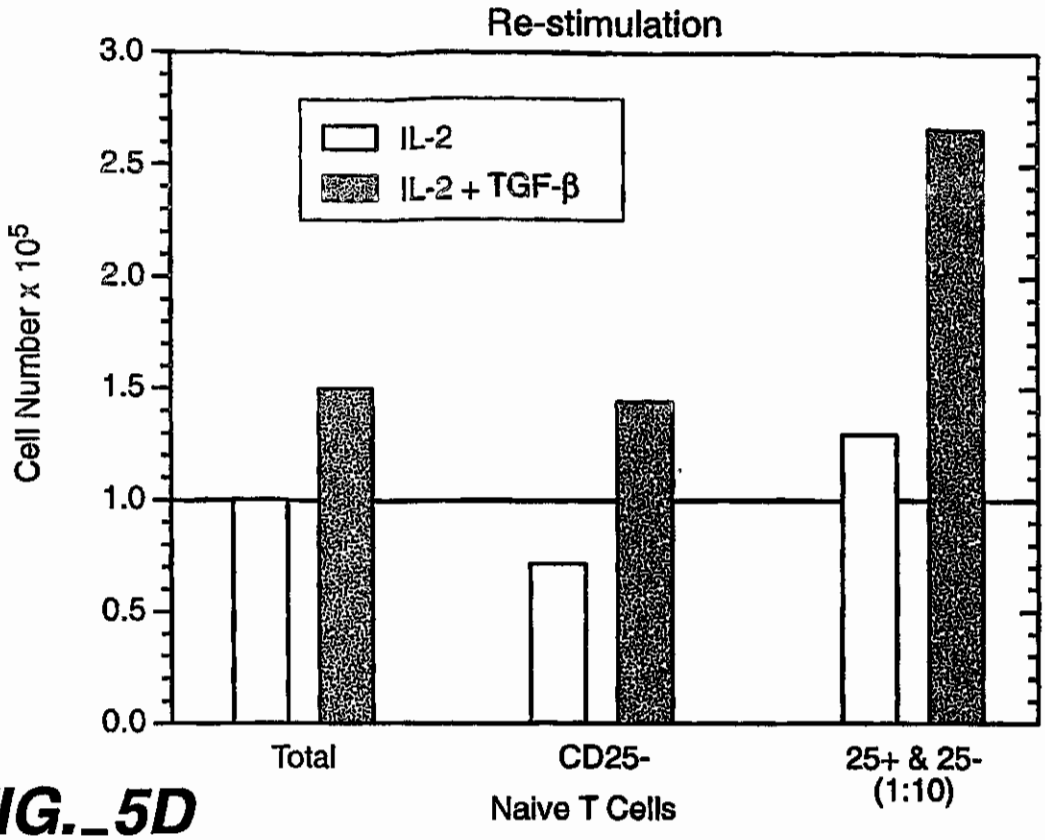

Re-stimulation
IL-2
IL-2 + TGF-β
Cell Number x 10^5
3.0
2.5
2.0
1.5
1.0
0.5
0.0
Total
CD25-
25+ & 25-
(1:10)
Naive T Cells

FIG._5D

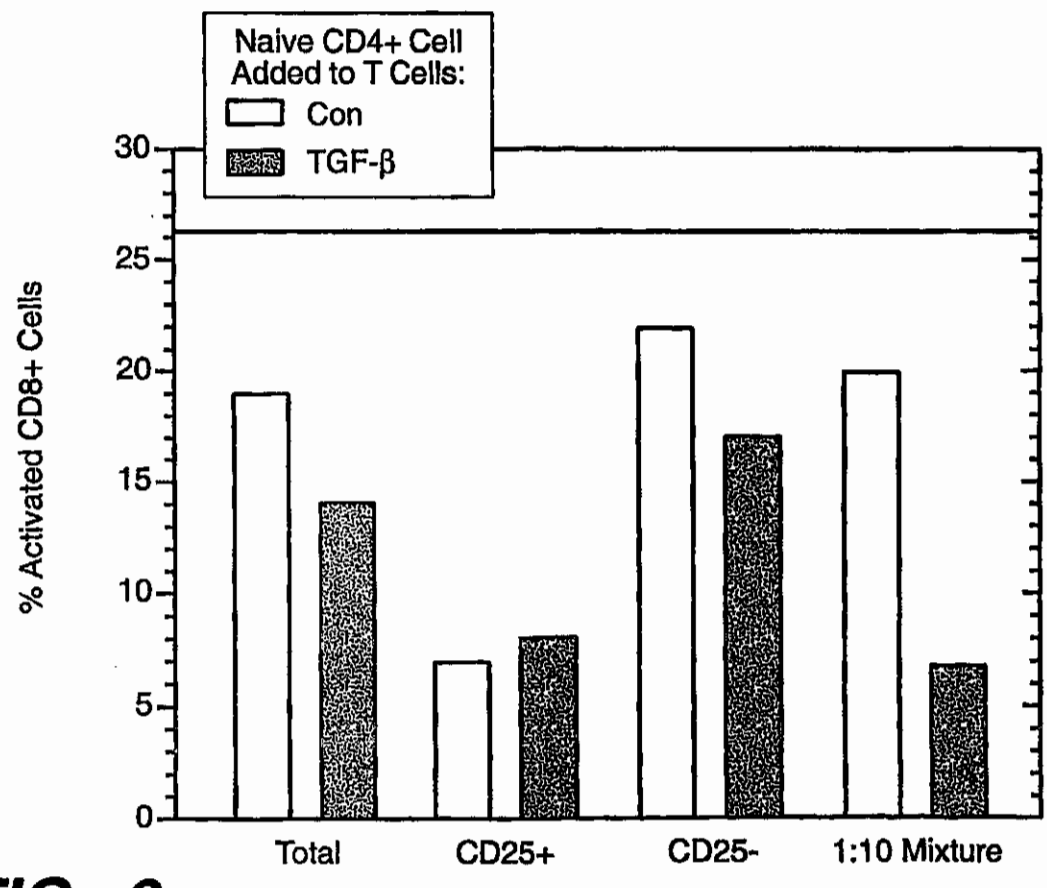

Naive CD4+ Cell
Added to T Cells:
Con
TGF-β
% Activated CD8+ Cells
30
25
20
15
10
5
0
Total
CD25+
CD25-
1:10 Mixture

FIG._6

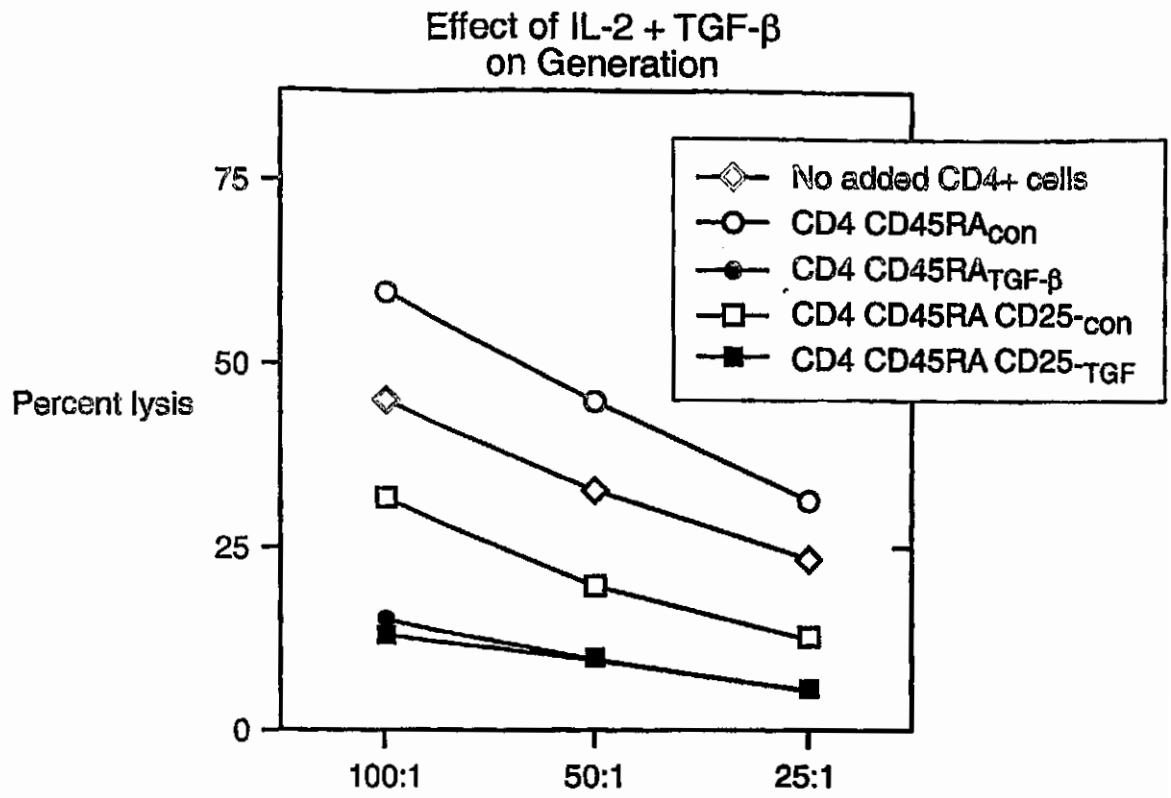

Effect of IL-2 + TGF-β
on Generation
Percent lysis
75
50
25
0
100:1
50:1
25:1
Effector to Target Ratio
No added CD4+ cells
CD4 CD45RA$_{con}$
CD4 CD45RA$_{TGF-β}$
CD4 CD45RA CD25-$_{con}$
CD4 CD45RA CD25-$_{TGF}$

FIG._7A

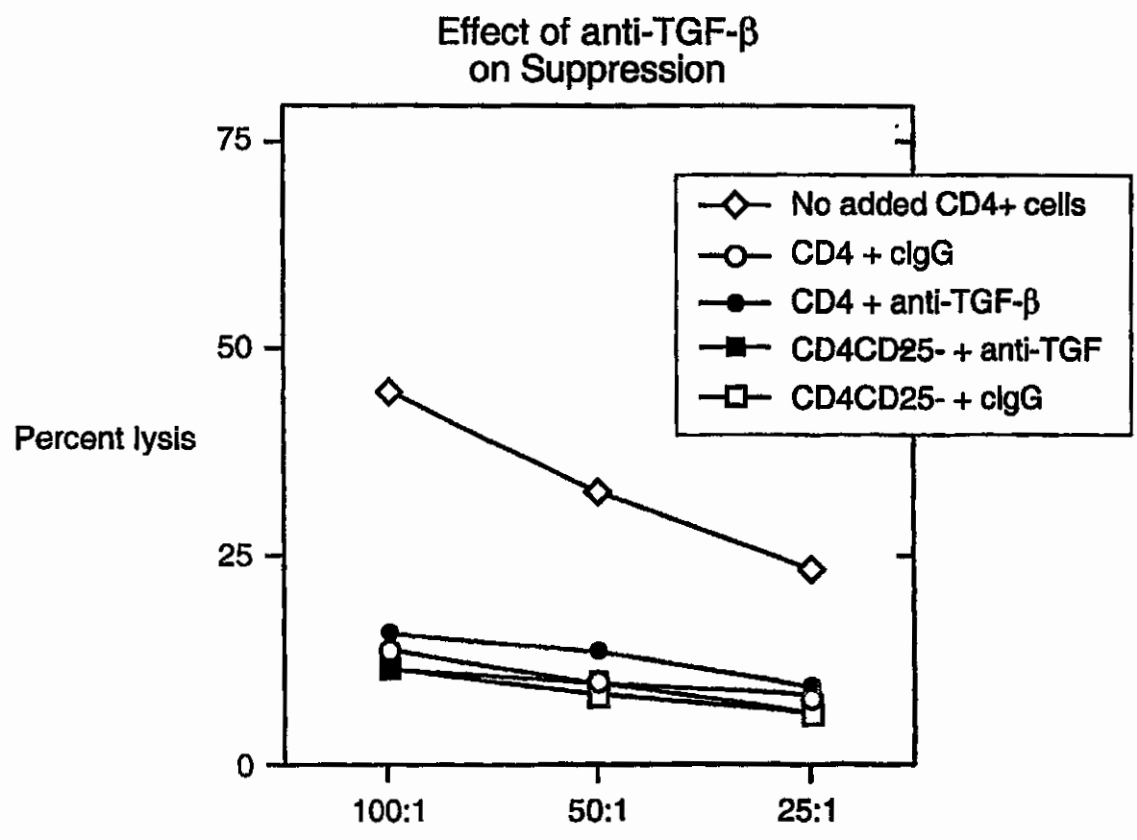

Effect of anti-TGF-β
on Suppression
Percent lysis
75
50
25
0
100:1
50:1
25:1
Effector to Target Ratio
No added CD4+ cells
CD4 + cIgG
CD4 + anti-TGF-β
CD4CD25- + anti-TGF
CD4CD25- + cIgG

FIG._7B

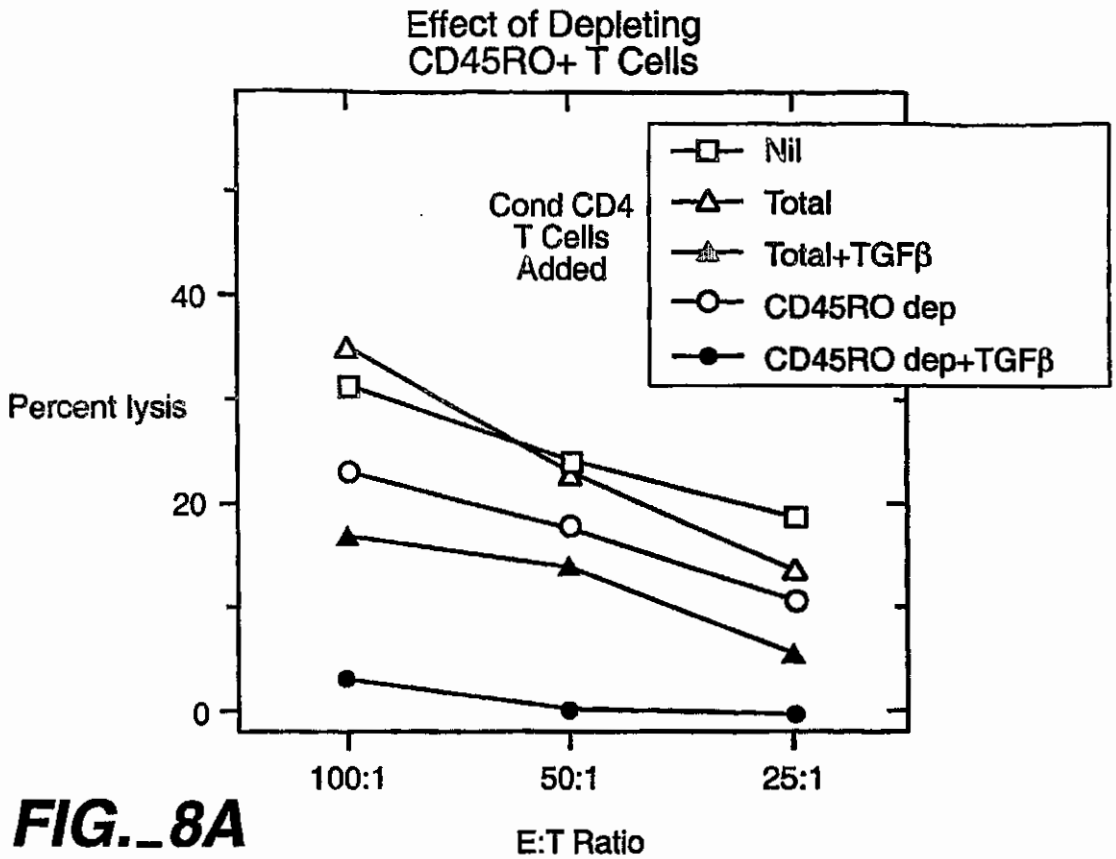

Effect of Depleting
CD45RO+ T Cells
Cond CD4
T Cells
Added
Nil
Total
Total+TGFβ
CD45RO dep
CD45RO dep+TGFβ
Percent lysis
40
20
0
100:1
50:1
25:1
E:T Ratio

FIG._8A

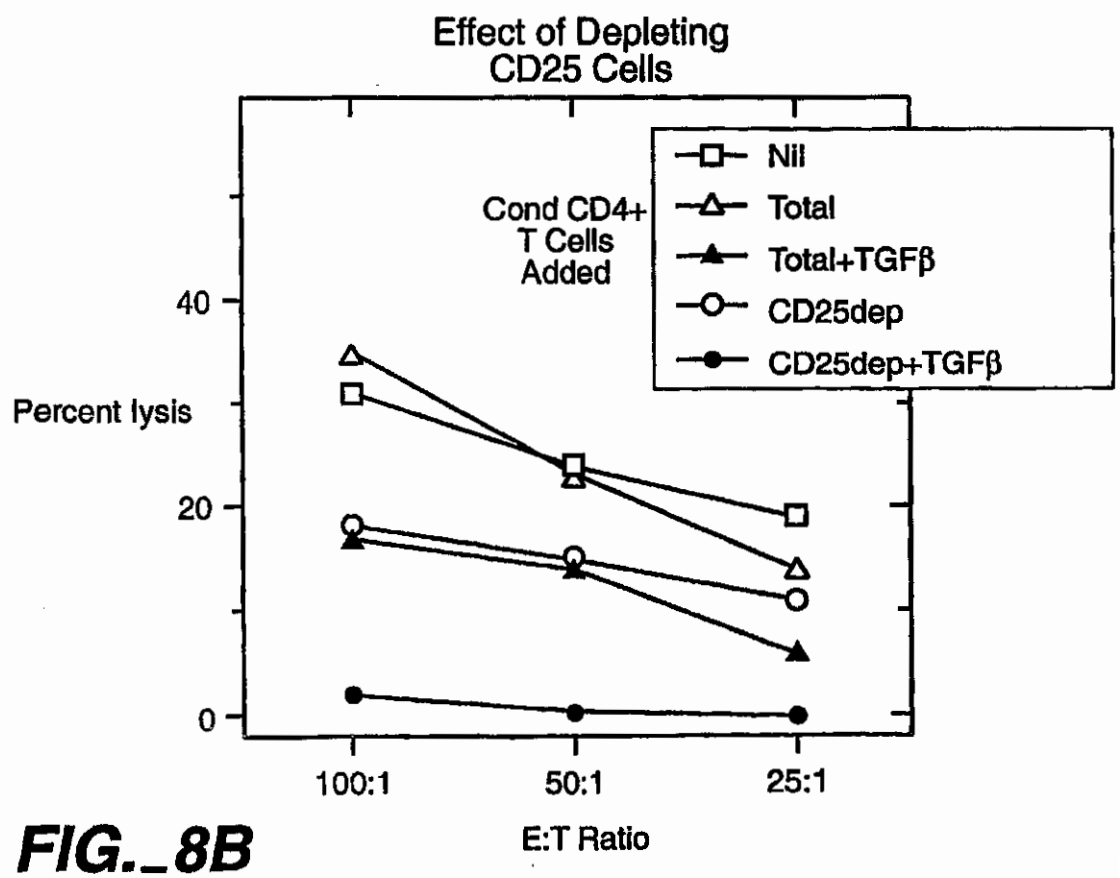

Effect of Depleting
CD25 Cells
Cond CD4+
T Cells
Added
Nil
Total
Total+TGFβ
CD25dep
CD25dep+TGFβ
Percent lysis
40
20
0
100:1
50:1
25:1
E:T Ratio

FIG._8B

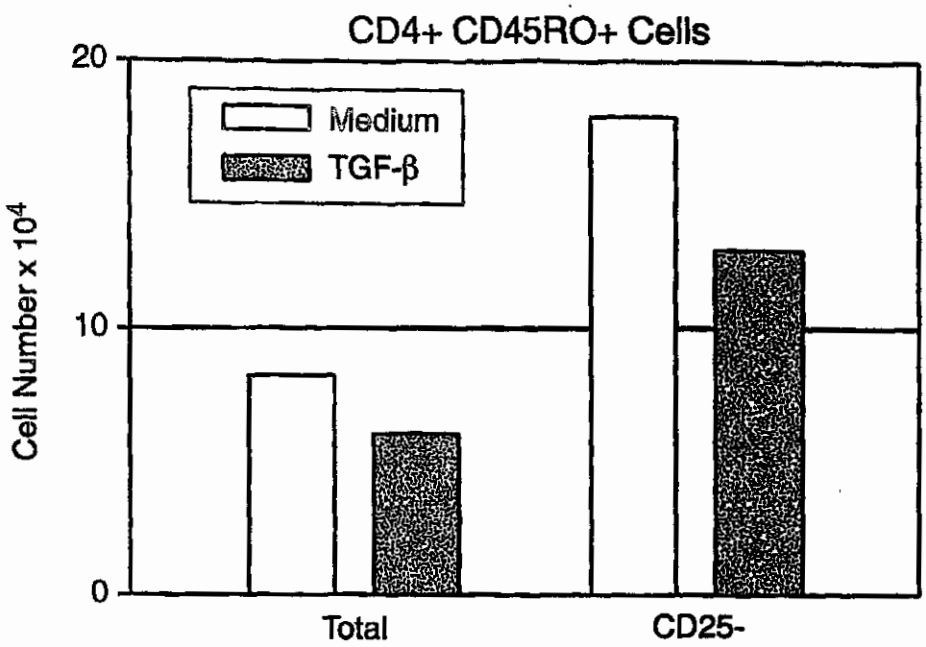

CD4+ CD45RO+ Cells
20
10
0
Cell Number x 10^4
Medium
TGF-β
Total
CD25-

FIG._8C

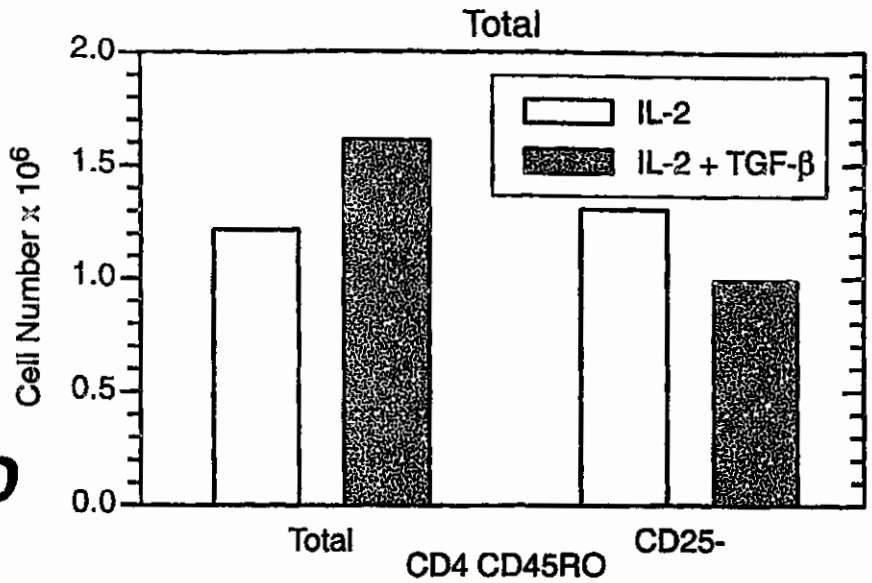


FIG._8D

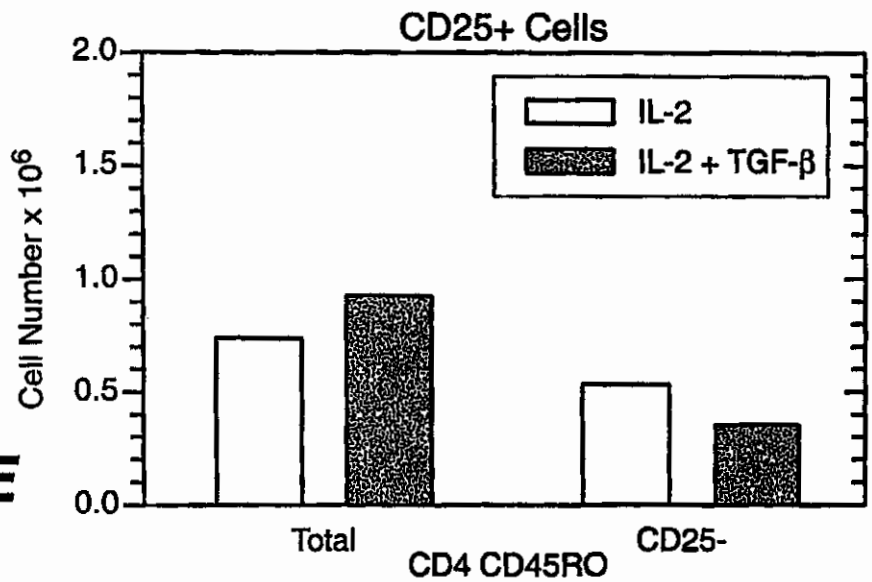


FIG._8E

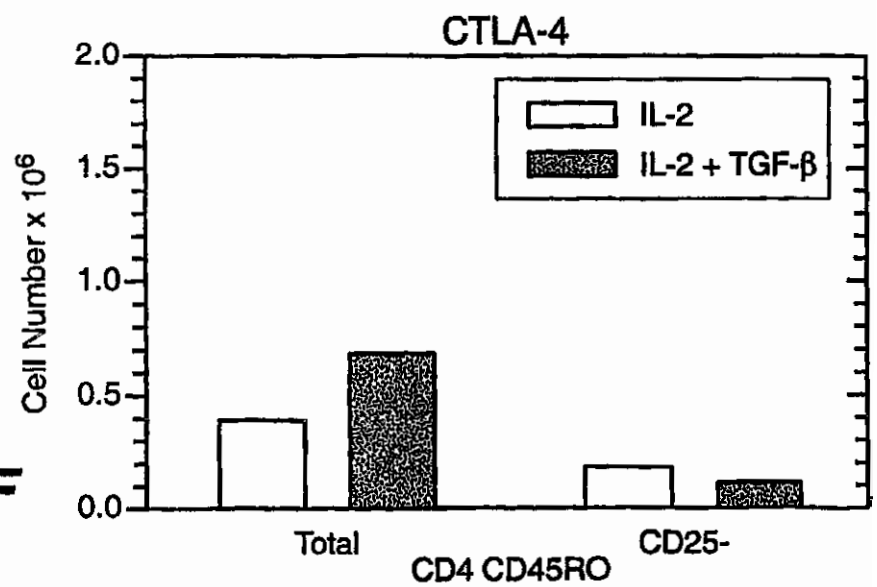


FIG._8F

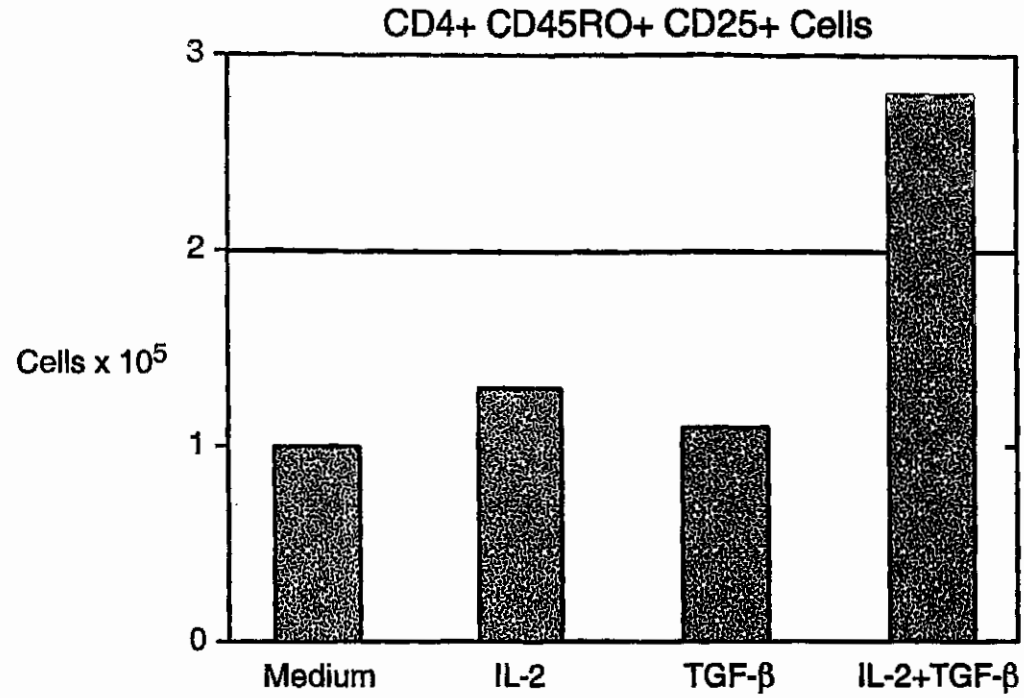

CD4+ CD45RO+ CD25+ Cells
Cells x 10^5
3
2
1
0
Medium
IL-2
TGF-β
IL-2+TGF-β

FIG._8G

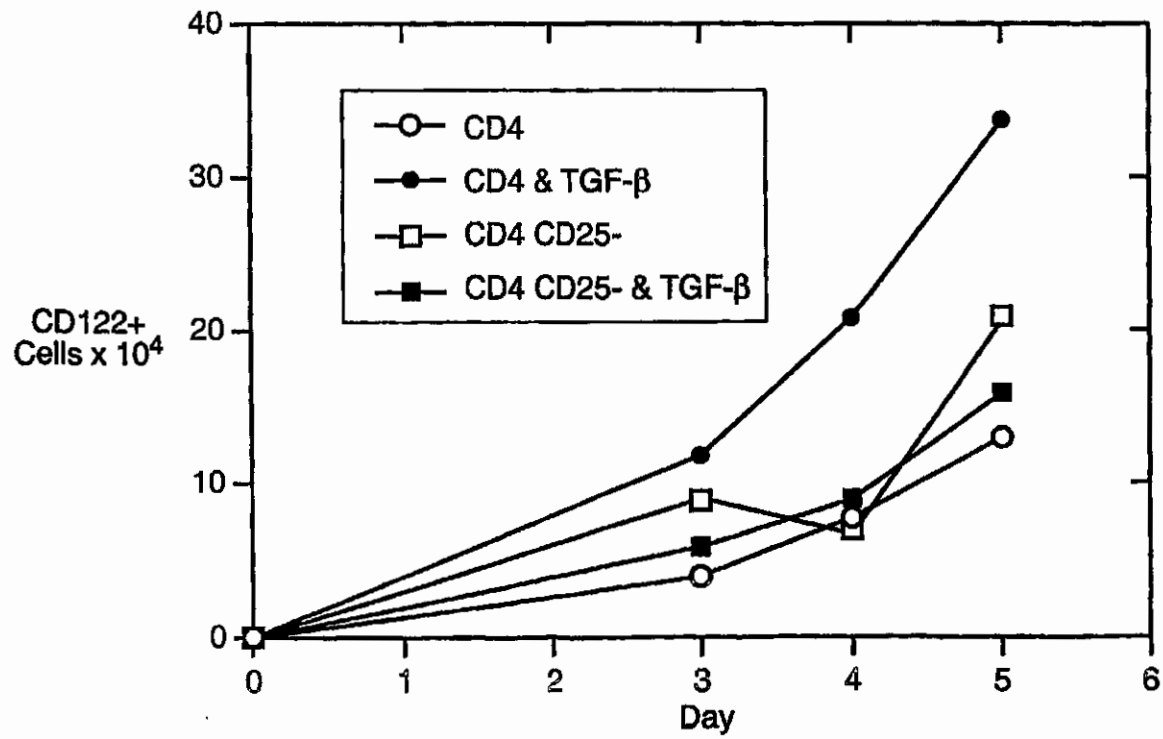

CD122+ Cells x 10^4
40
30
20
10
0
CD4
CD4 & TGF-β
CD4 CD25-
CD4 CD25- & TGF-β
0
1
2
3
4
5
6
Day

FIG._8H

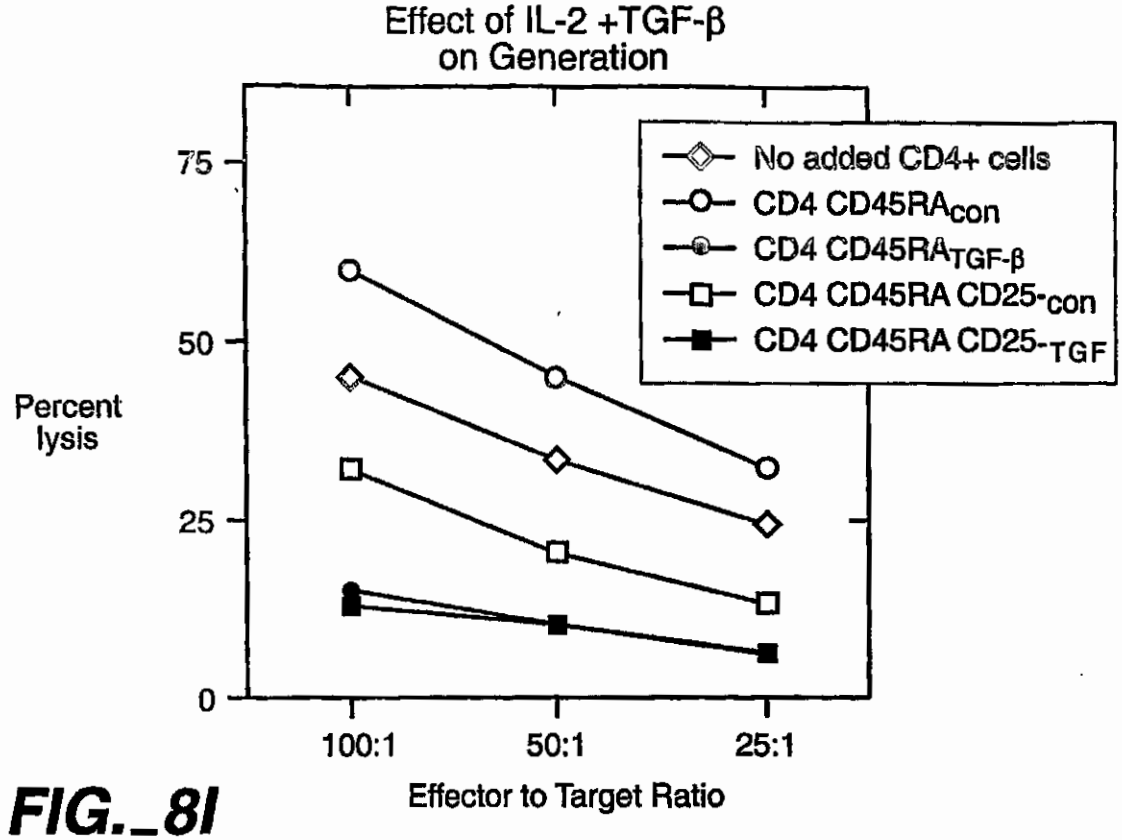

Effect of IL-2 +TGF-β
on Generation
No added CD4+ cells
CD4 CD45RAcon
CD4 CD45RATGF-β
CD4 CD45RA CD25-con
CD4 CD45RA CD25-TGF
Percent lysis
75
50
25
0
100:1
50:1
25:1
Effector to Target Ratio

FIG._8I

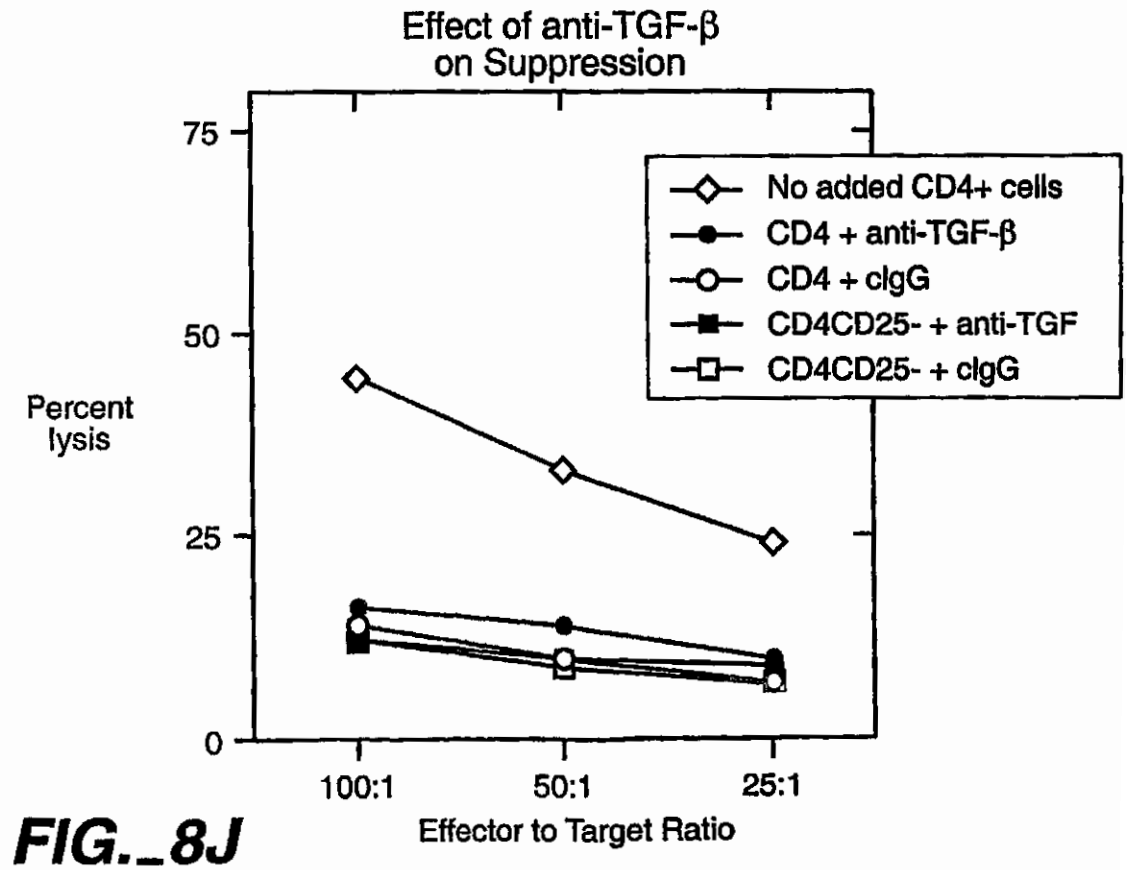

Effect of anti-TGF-β
on Suppression
No added CD4+ cells
CD4 + anti-TGF-β
CD4 + clgG
CD4CD25- + anti-TGF
CD4CD25- + clgG
Percent lysis
75
50
25
0
100:1
50:1
25:1
Effector to Target Ratio

FIG._8J

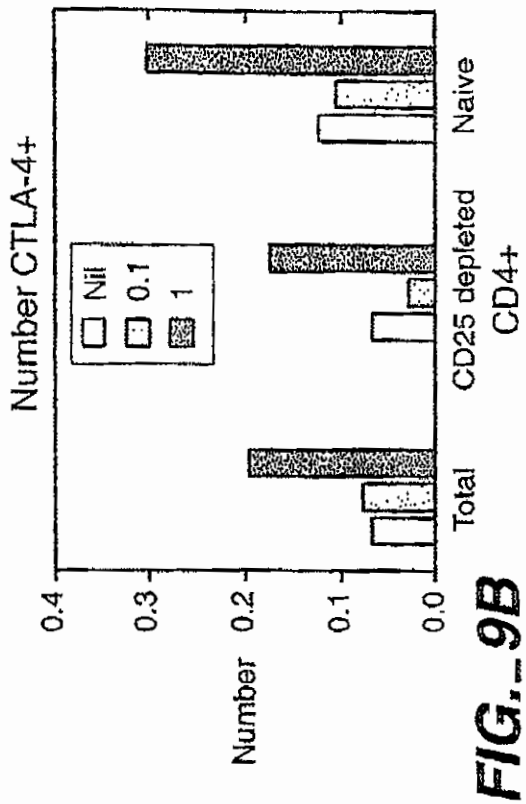


FIG._9B

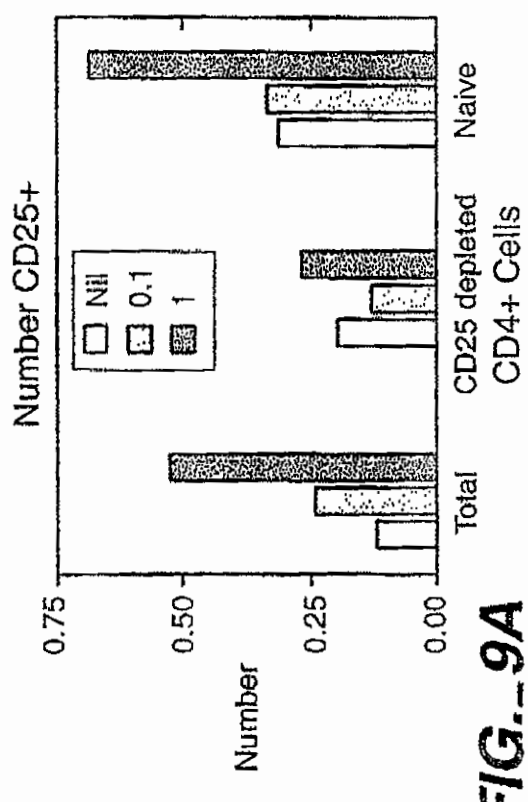


FIG._9A

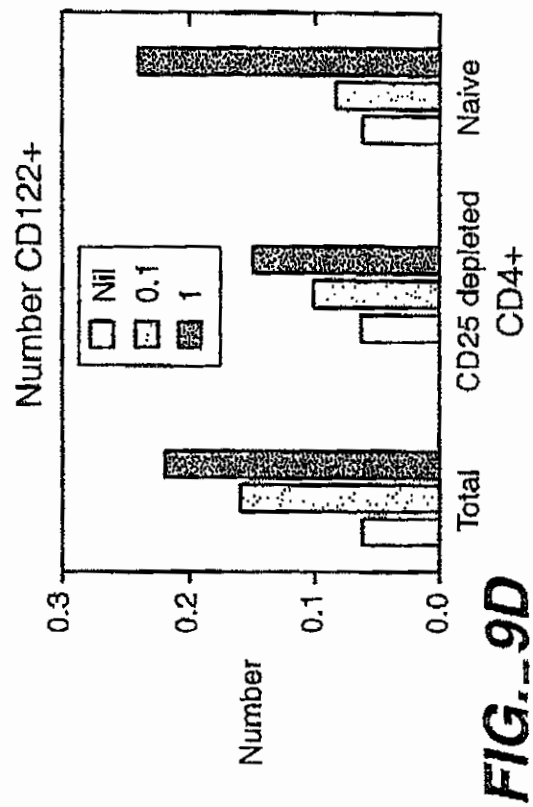


FIG._9D

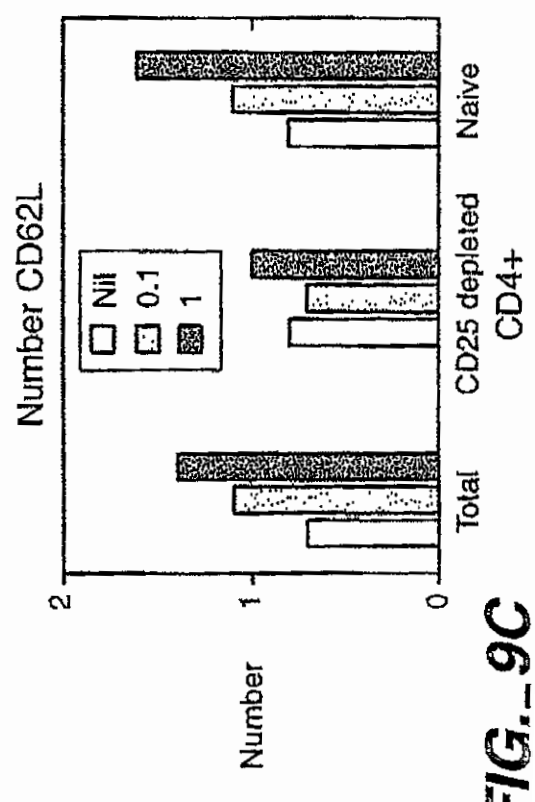


FIG._9C

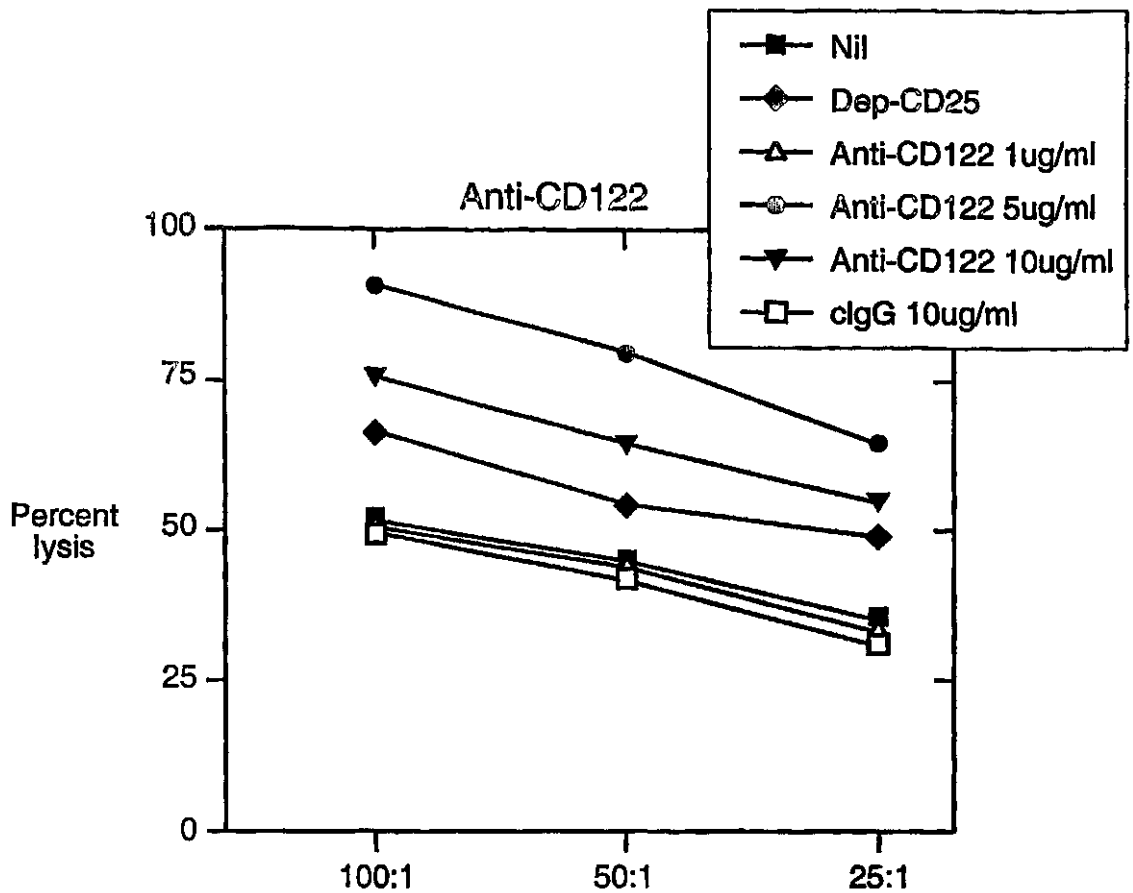

Anti-CD122
Nil
Dep-CD25
Anti-CD122 1ug/ml
Anti-CD122 5ug/ml
Anti-CD122 10ug/ml
cIgG 10ug/ml
Percent lysis
100
75
50
25
0
100:1
50:1
25:1

FIG._10

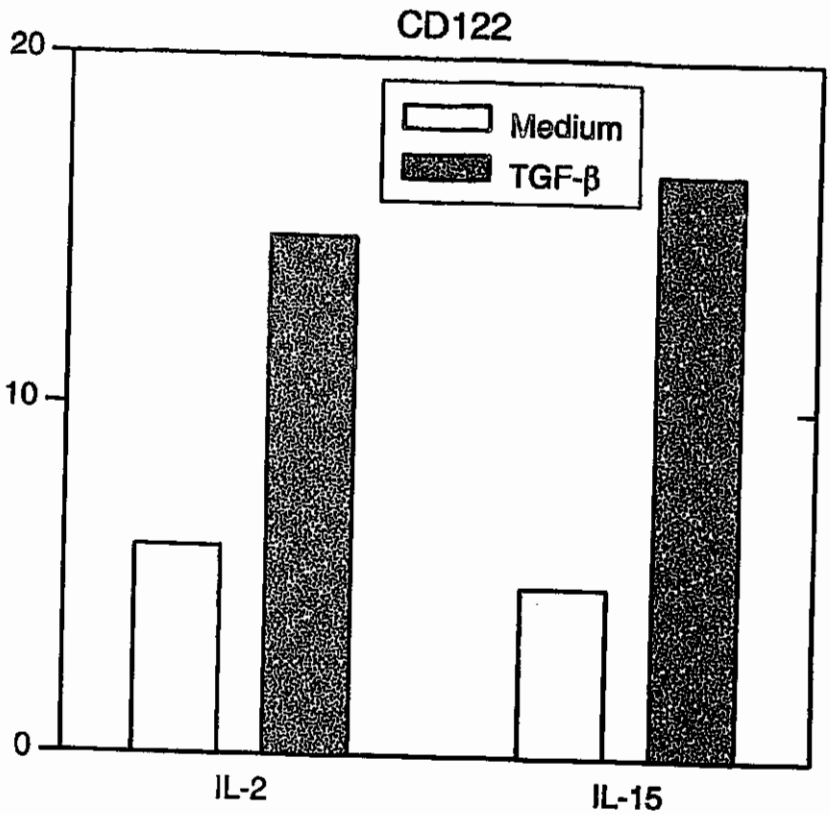


FIG._11A

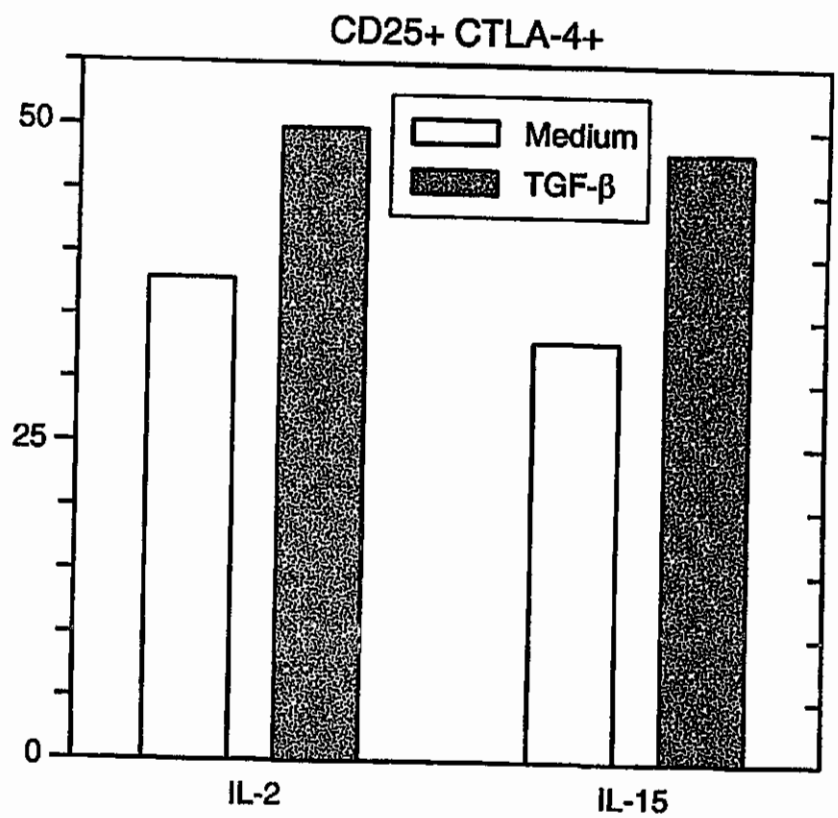


FIG._11B