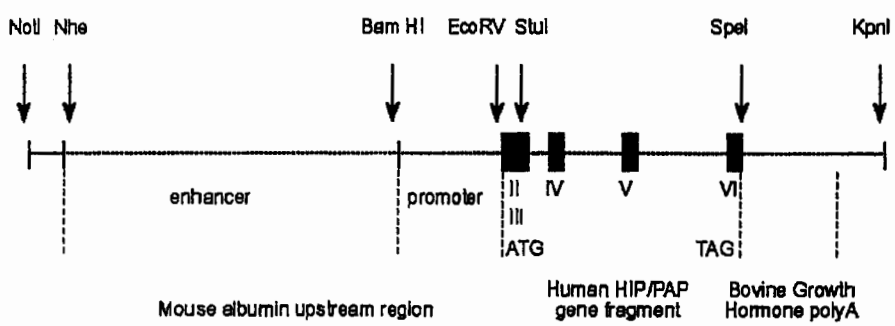

NotI
Nhe
Bam HI
EcoRV
StuI
SpeI
KpnI
enhancer
promoter
II
III
IV
V
VI
ATG
TAG
Mouse albumin upstream region
Human HIP/PAP gene fragment
Bovine Growth Hormone polyA

FIGURE 1

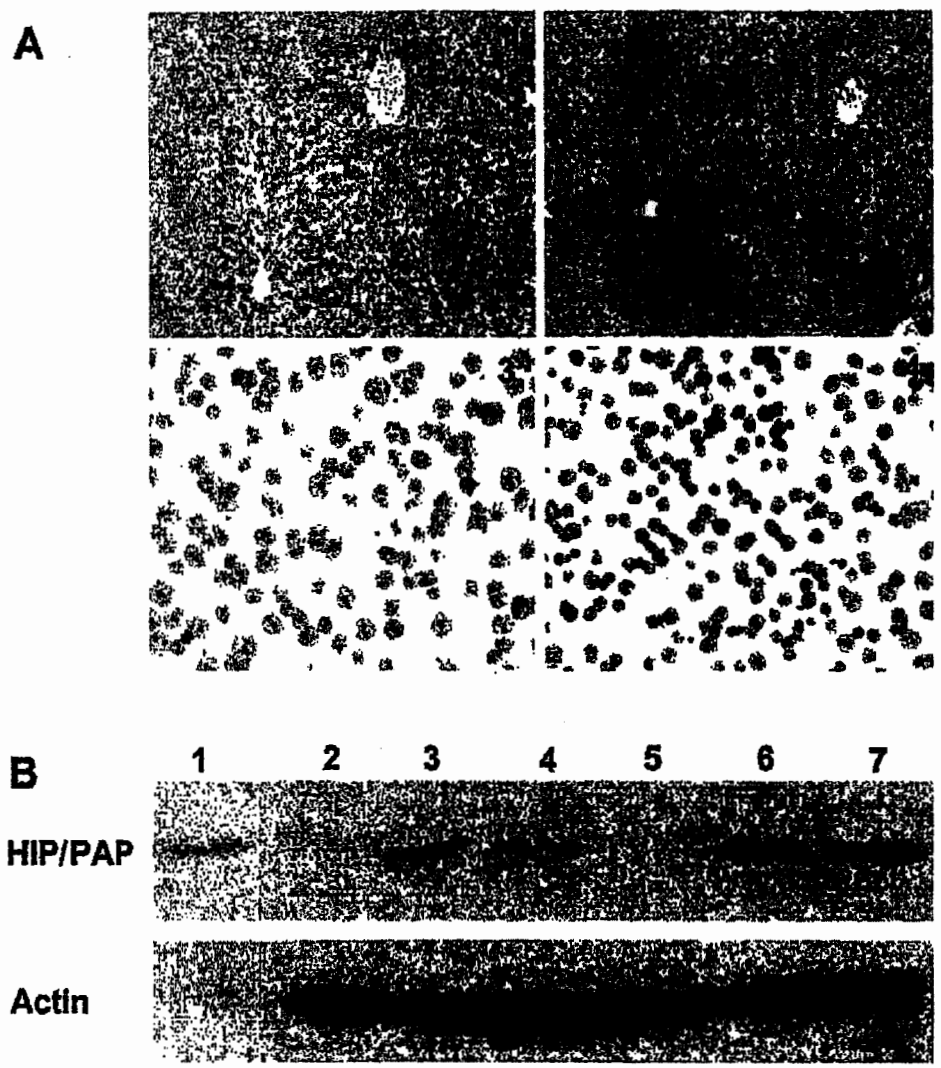


FIGURE 2

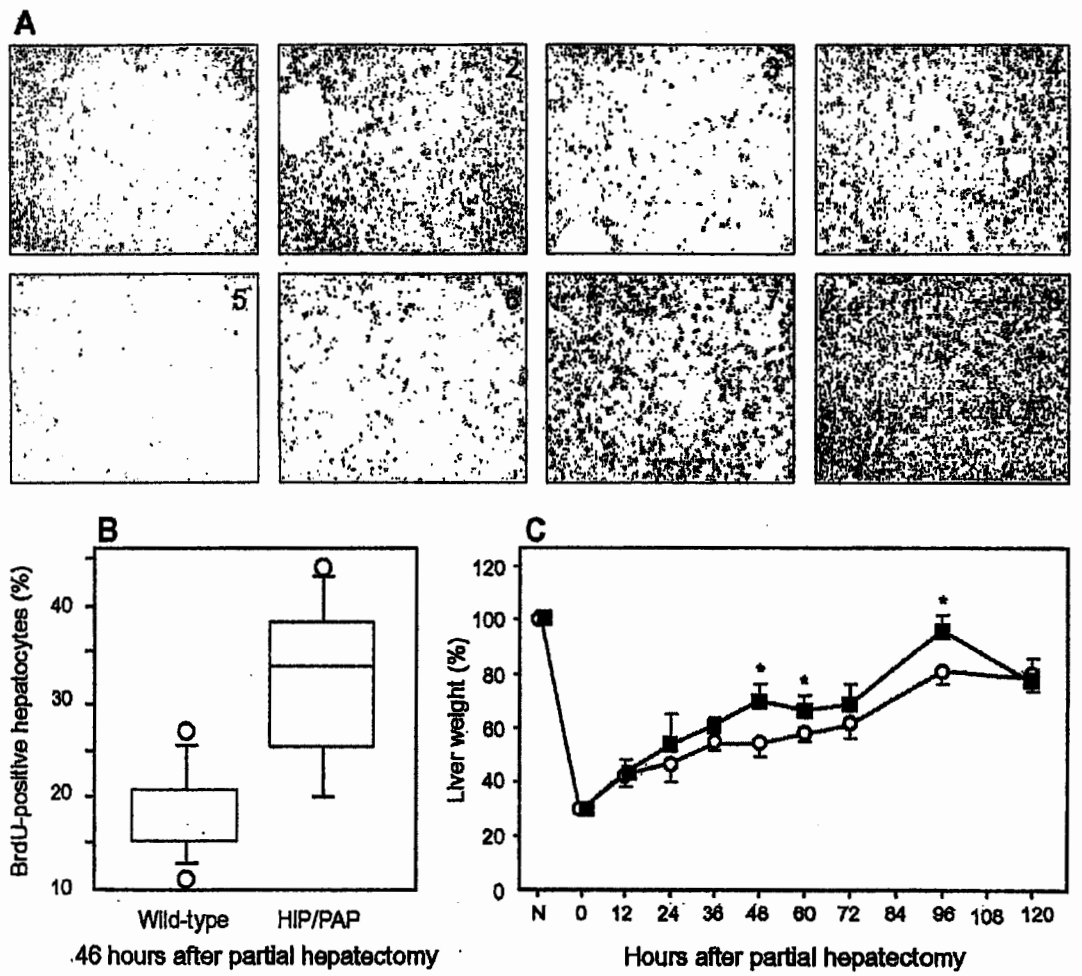

A
B
BrdU-positive hepatocytes (%)
40
30
20
10
Wild-type
HIP/PAP
46 hours after partial hepatectomy
C
Liver weight (%)
120
100
80
60
40
20
0
N
0
12
24
36
48
60
72
84
96
108
120
Hours after partial hepatectomy

FIGURE 3

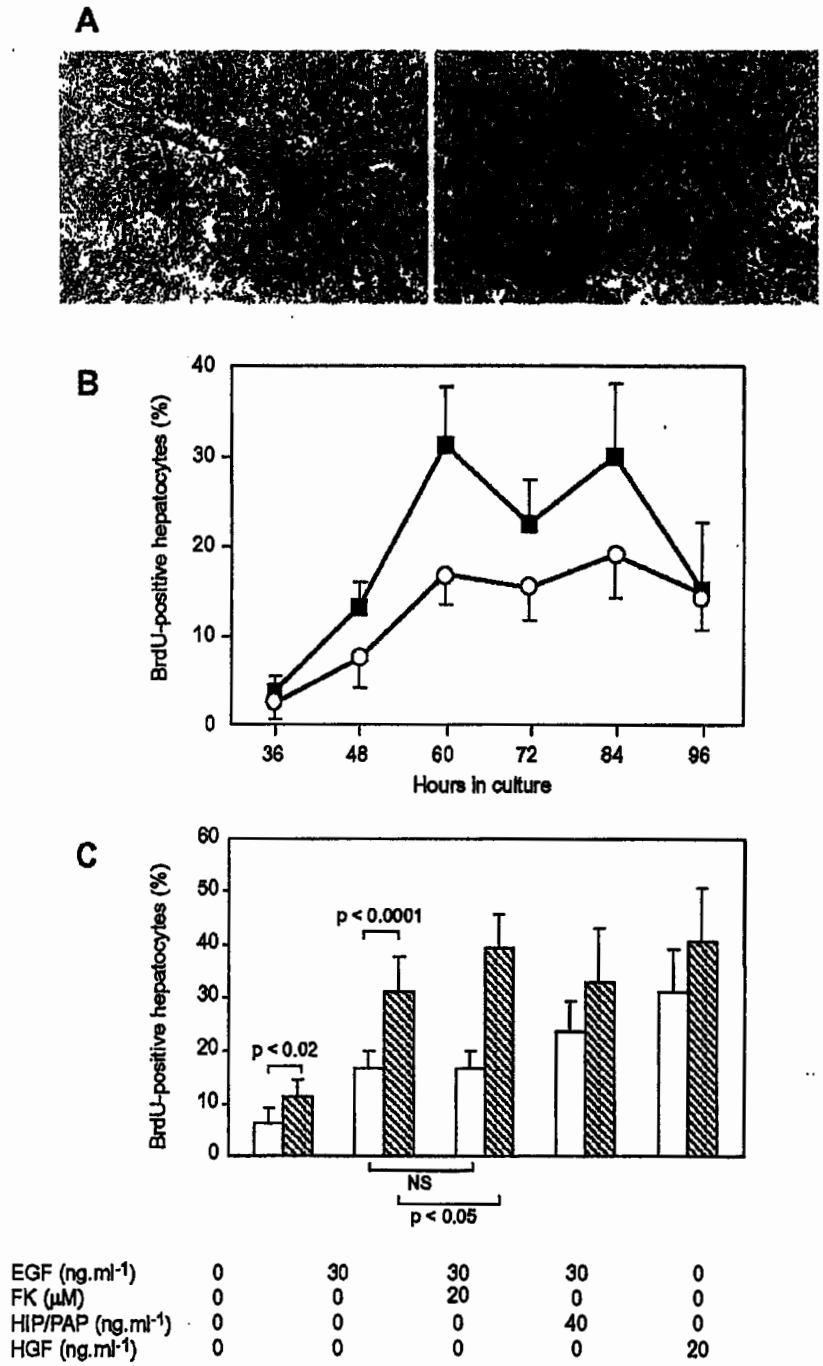

A
B
BrdU-positive hepatocytes (%)
40
30
20
10
0
36
48
60
72
84
96
Hours in culture
C
BrdU-positive hepatocytes (%)
60
50
40
30
20
10
0
p < 0.0001
p < 0.02
NS
p < 0.05
EGF (ng.ml-1) 0 30 30 30 0
FK (µM) 0 0 20 0 0
HIP/PAP (ng.ml-1) 0 0 0 40 0
HGF (ng.ml-1) 0 0 0 0 20

FIGURE 4

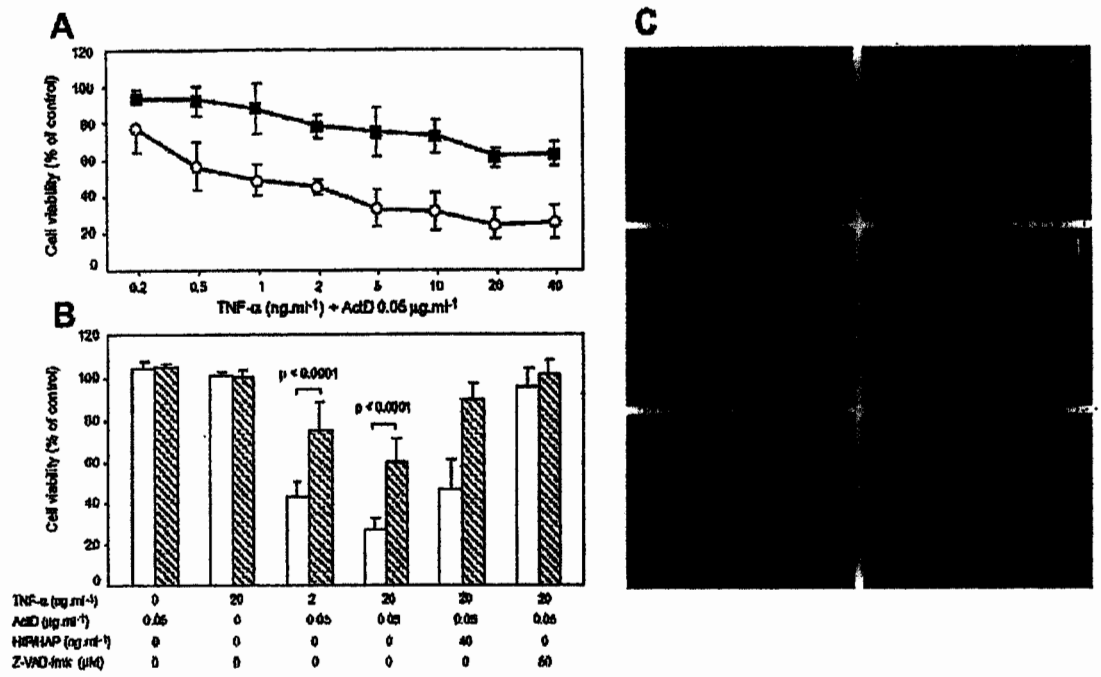

A
Cell viability (% of control)
TNF-α (ng.ml-1) + ActD 0.05 µg.ml-1
B
Cell viability (% of control)
p < 0.0001
p < 0.0001
TNF-α (ng.ml-1)
ActD (µg.ml-1)
HIRHAP (ng.ml-1)
Z-VAD-fmk (µM)
C

FIGURE 5

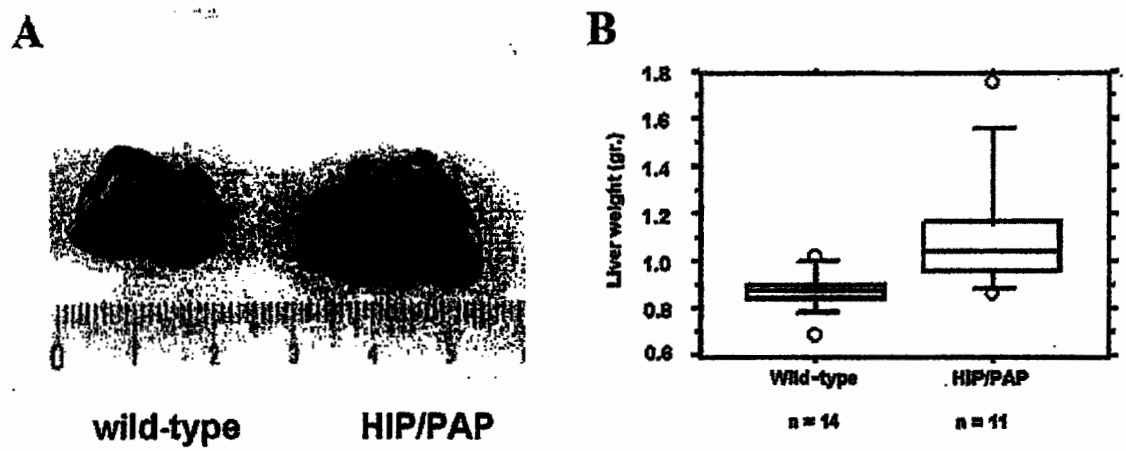

A
B
wild-type
HIP/PAP
Liver weight (gr.)
1.8
1.6
1.4
1.2
1.0
0.8
0.6
Wild-type
HIP/PAP
n = 14
n = 11

FIGURE 6

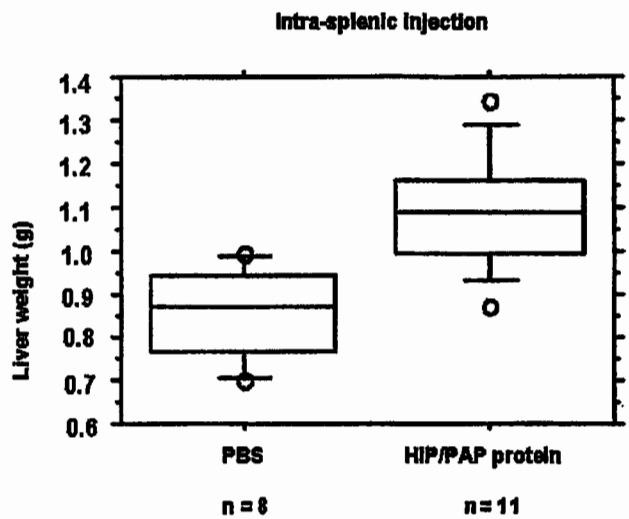

Intra-splenic injection
Liver weight (g)
1.4
1.3
1.2
1.1
1.0
0.9
0.8
0.7
0.6
PBS
HIP/PAP protein
n = 8
n = 11

FIGURE 7

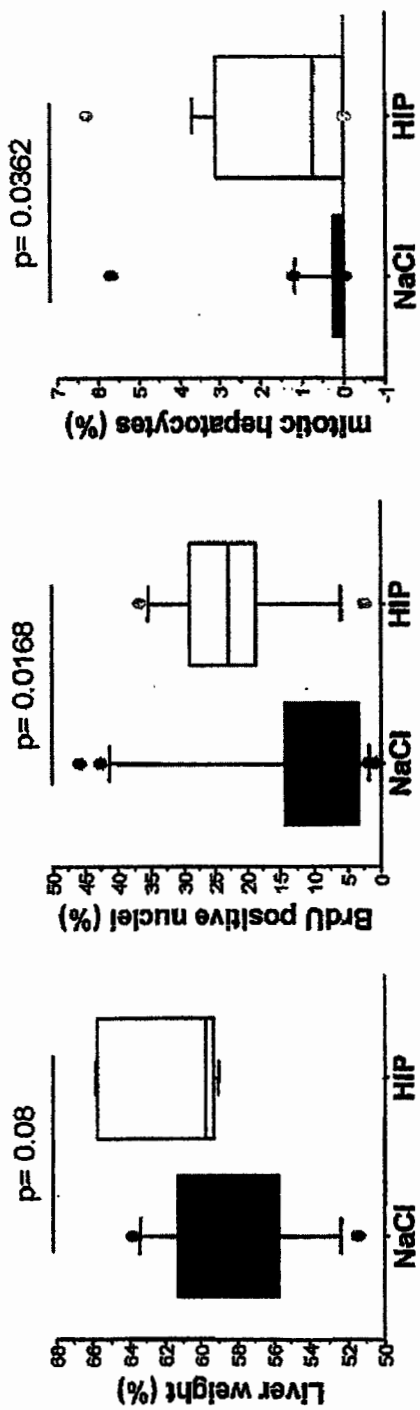

p= 0.08
NaCl
HIP
Liver weight (%)
68
66
64
62
60
58
56
54
52
50
p= 0.0168
NaCl
HIP
BrdU positive nuclei (%)
50
45
40
35
30
25
20
15
10
5
0
p= 0.0362
NaCl
HIP
mitotic hepatocytes (%)
7
6
5
4
3
2
1
0
-1

FIGURE 8

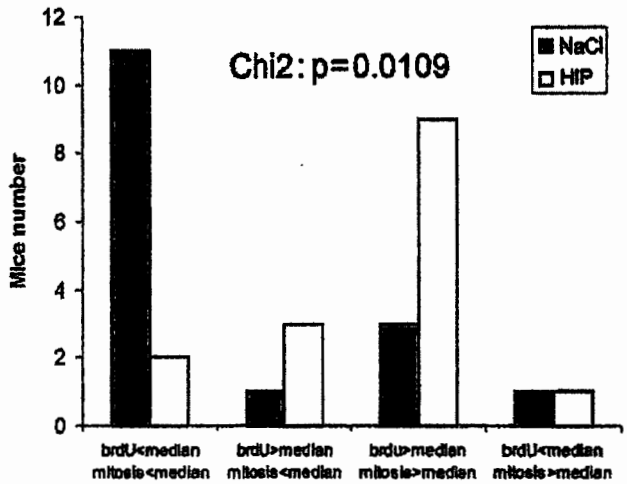

Chi2: p=0.0109
NaCl
HIP
Mice number
12
10
8
6
4
2
0
brdU<median mitosis<median
brdU>median mitosis<median
brdU>median mitosis>median
brdU<median mitosis>median

FIGURE 9

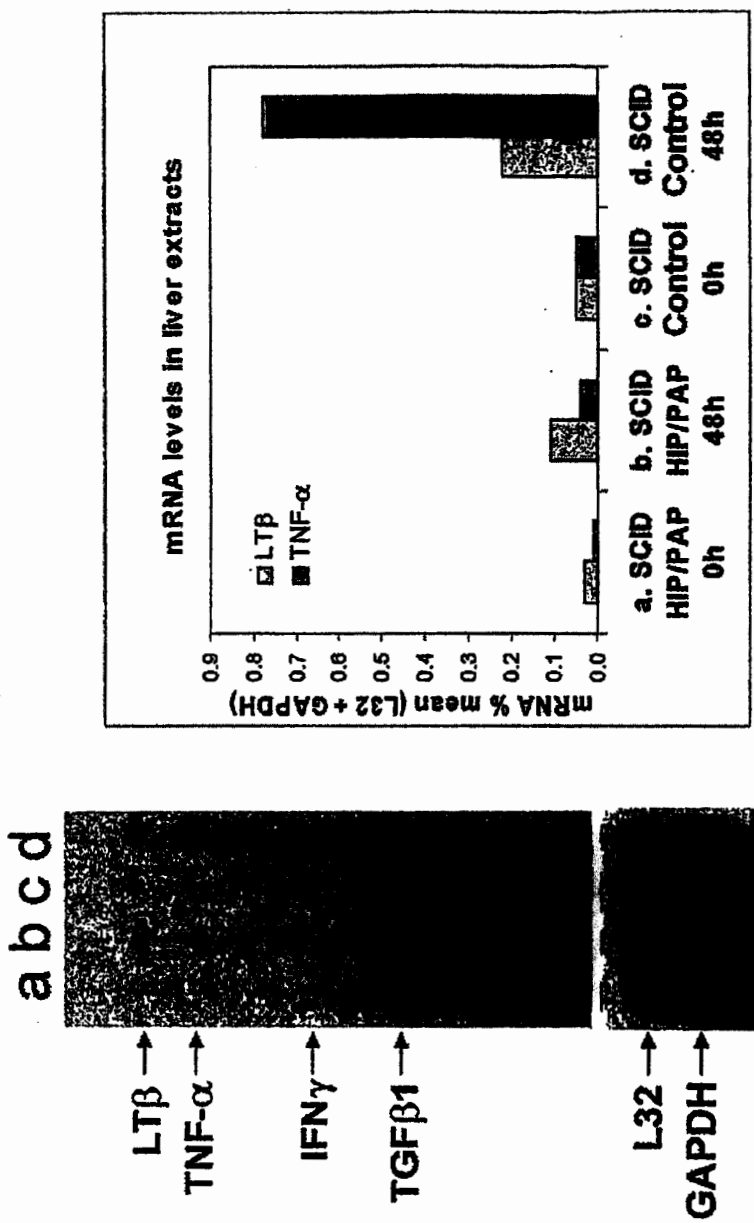
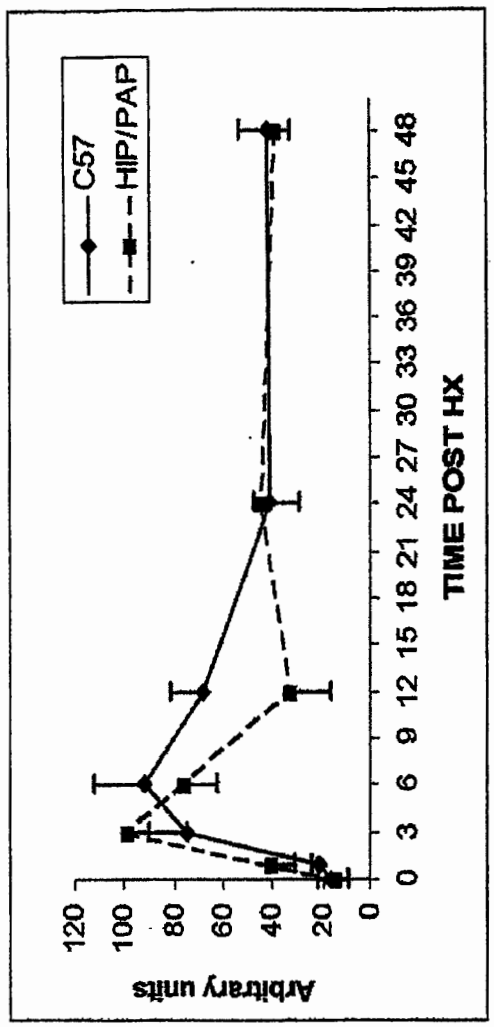

mRNA levels in liver extracts
LTβ
TNF-α
mRNA % mean (L32 + GAPDH)
0.9
0.8
0.7
0.6
0.5
0.4
0.3
0.2
0.1
0.0
a. SCID HIP/PAP 0h
b. SCID HIP/PAP 48h
c. SCID Control 0h
d. SCID Control 48h
a b c d
LTβ
TNF-α
IFNγ
TGFβ1
L32
GAPDH

FIGURE 10


C57
HIP/PAP
Arbitrary units
120
100
80
60
40
20
0
0 3 6 9 12 15 18 21 24 27 30 33 36 39 42 45 48
TIME POST HX


0 1 3 6 12 24 48
Control
HIP/PAP
STAT 3 P
STAT 3 P

FIGURE 11

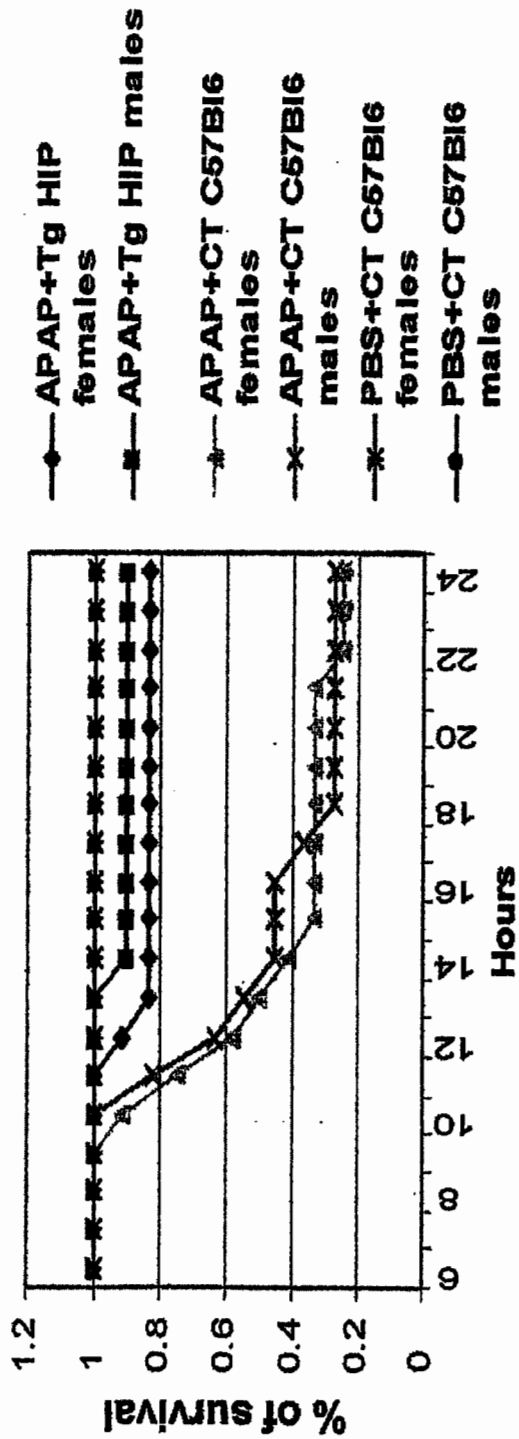

APAP+Tg HIP females
APAP+Tg HIP males
APAP+CT C57Bl6 females
APAP+CT C57Bl6 males
PBS+CT C57Bl6 females
PBS+CT C57Bl6 males
% of survival
1.2
1
0.8
0.6
0.4
0.2
0
6
8
10
12
14
16
18
20
22
24
Hours

FIGURE 12

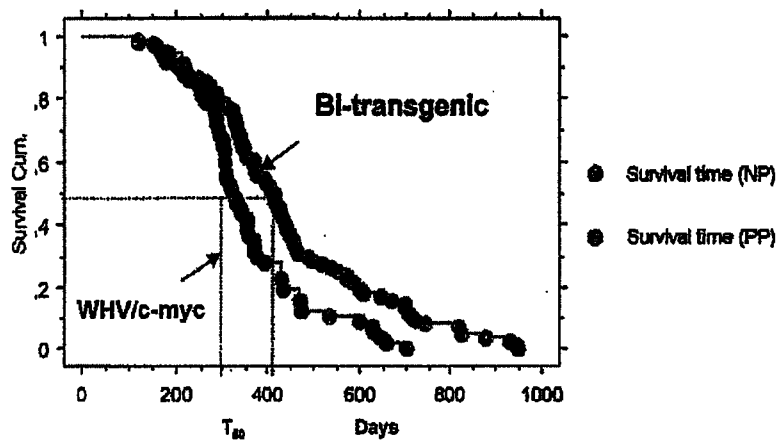

Survival Cum.
1
,8
,6
,4
,2
0
0
200
400
600
800
1000
Days
T_{50}
Bi-transgenic
WHV/c-myc
Survival time (NP)
Survival time (PP)

FIGURE 13