Figure 1

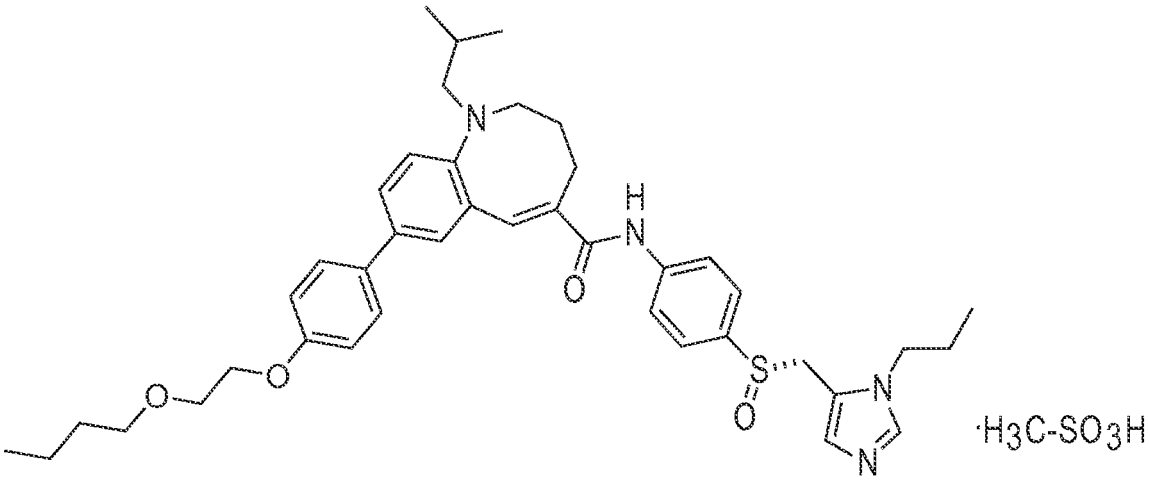


Figure 2

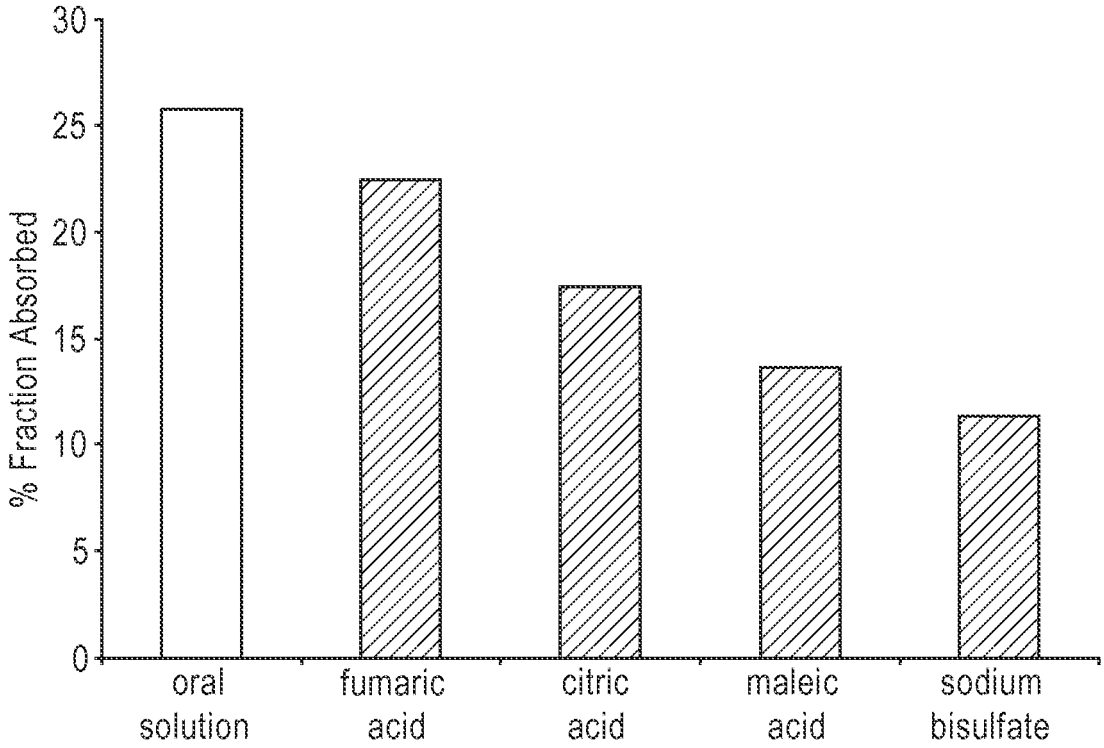

30
25
20
15
10
5
0
% Fraction Absorbed
oral solution
fumaric acid
citric acid
maleic acid
sodium bisulfate

Figure 3

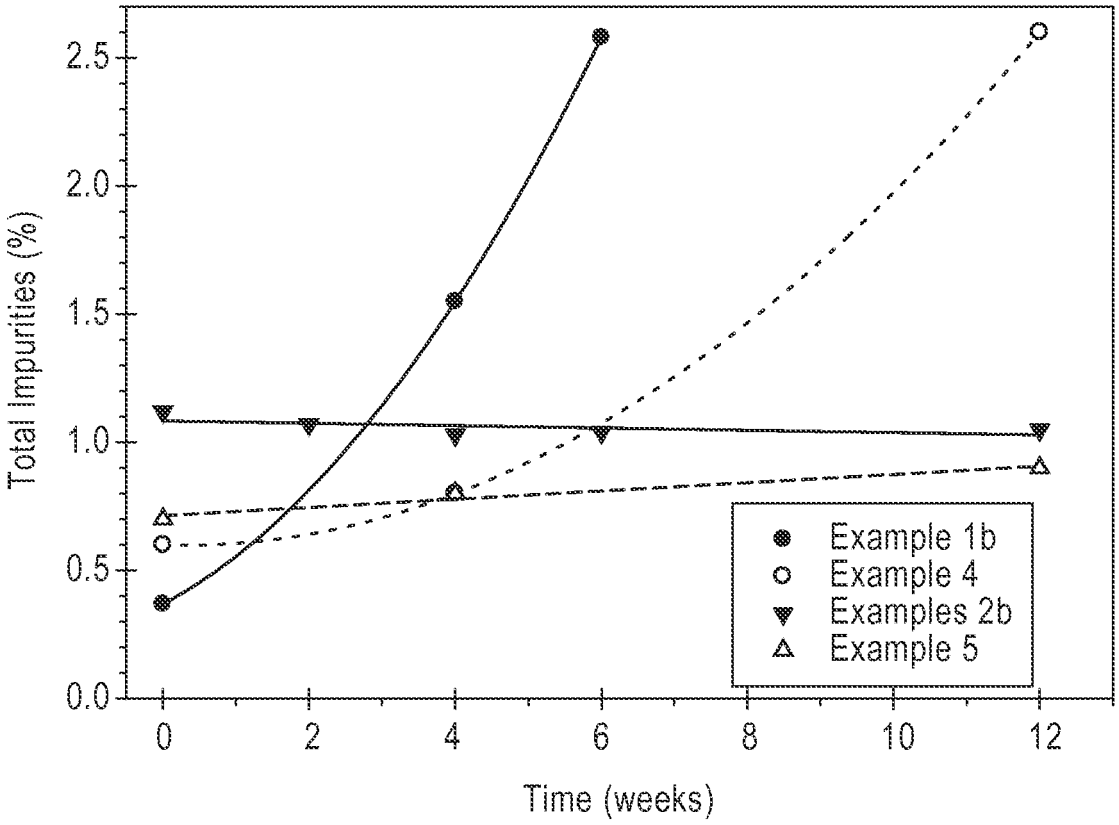

Total Impurities (%)
2.5
2.0
1.5
1.0
0.5
0.0
0
2
4
6
8
10
12
Time (weeks)
Example 1b
Example 4
Examples 2b
Example 5

Figure 4

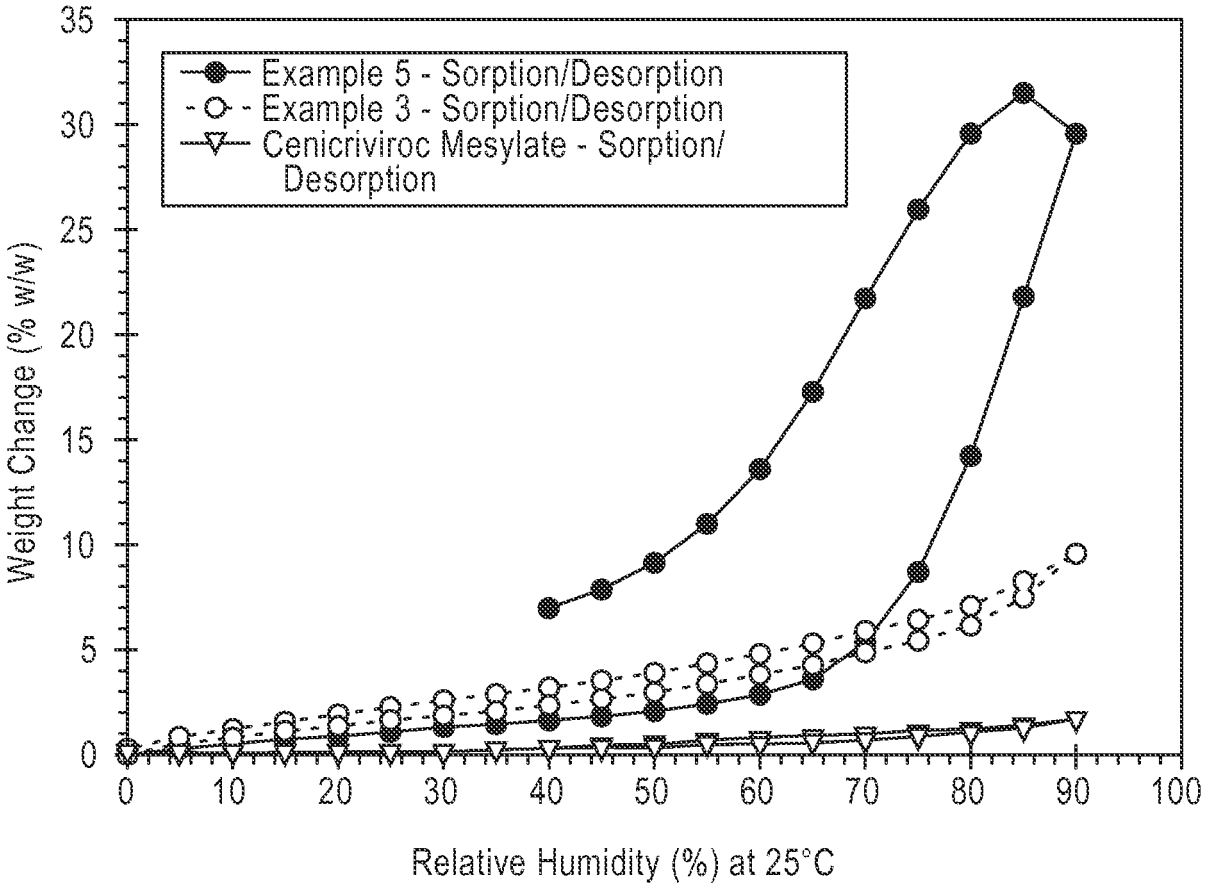

Example 5 - Sorption/Desorption
Example 3 - Sorption/Desorption
Cenicriviroc Mesylate - Sorption/ Desorption
Weight Change (% w/w)
Relative Humidity (%) at 25°C
0
5
10
15
20
25
30
35
0
10
20
30
40
50
60
70
80
90
100

Figure 5

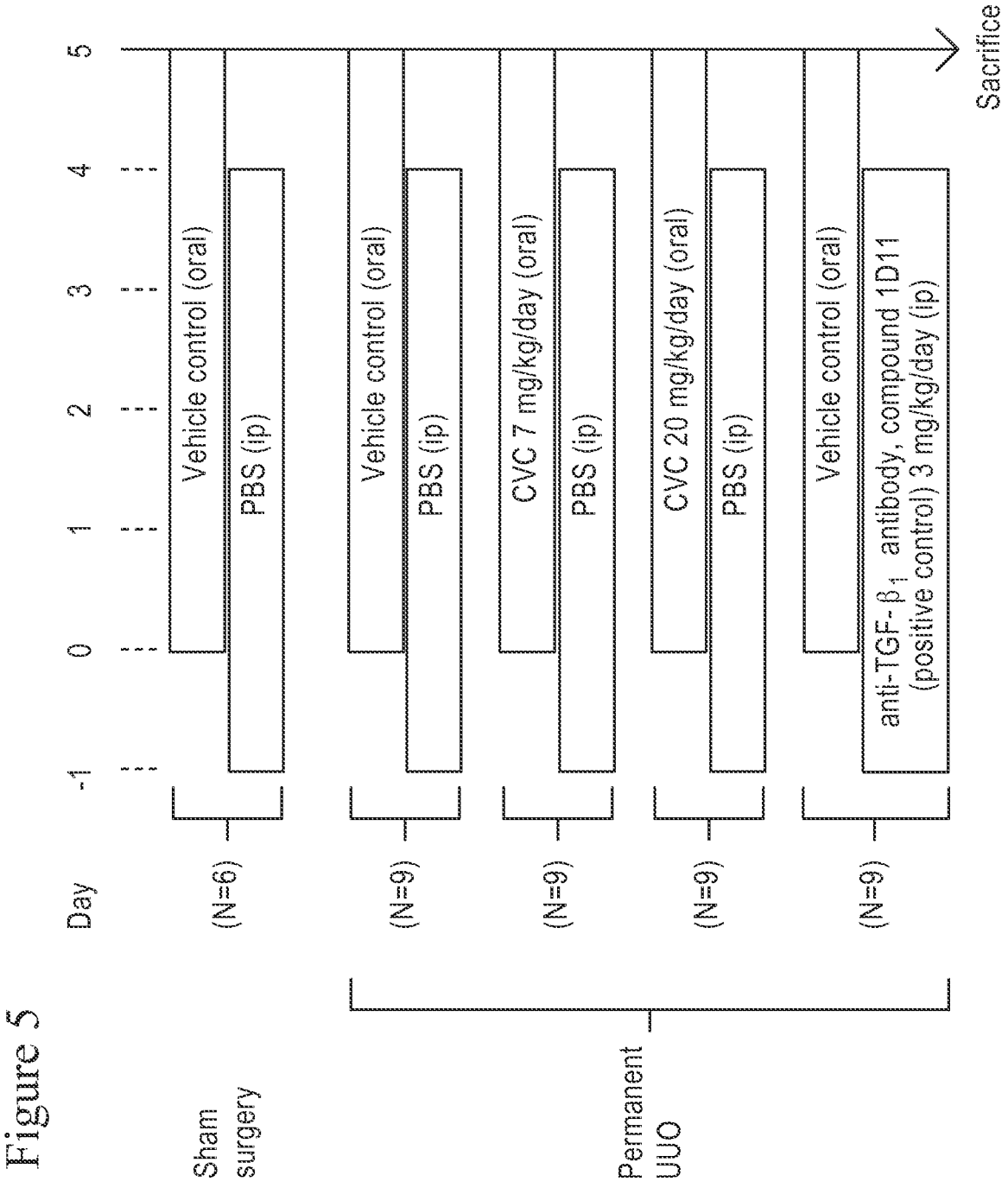

Day
-1
0
1
2
3
4
5
Sacrifice
Sham surgery
Permanent UUO
(N=6)
(N=9)
(N=9)
(N=9)
(N=9)
Vehicle control (oral)
PBS (ip)
Vehicle control (oral)
PBS (ip)
CVC 7 mg/kg/day (oral)
PBS (ip)
CVC 20 mg/kg/day (oral)
PBS (ip)
Vehicle control (oral)
anti-TGF-β1 antibody, compound 1D11
(positive control) 3 mg/kg/day (ip)

Figure 6

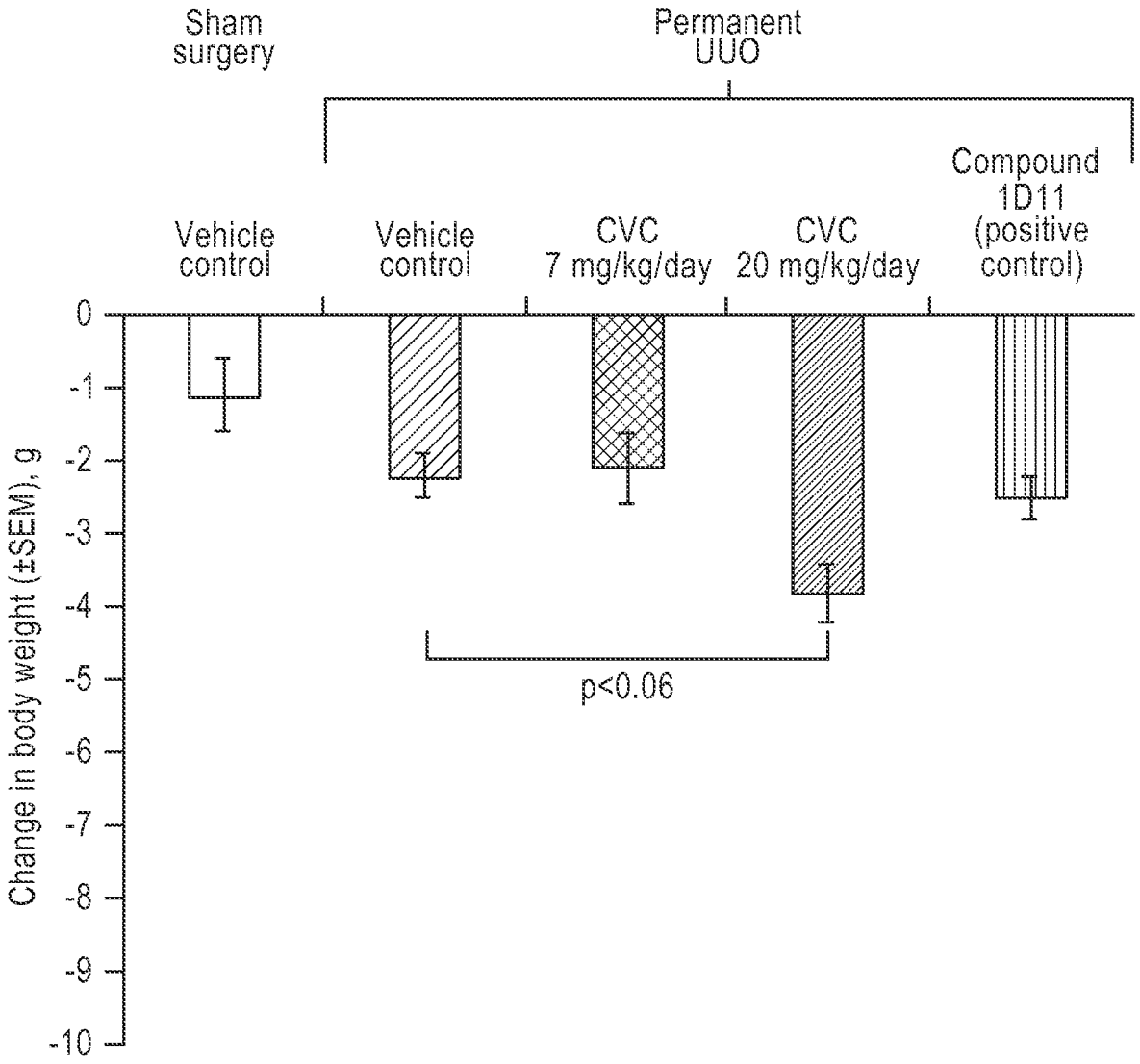

Sham surgery
Permanent UUO
Vehicle control
Vehicle control
CVC 7 mg/kg/day
CVC 20 mg/kg/day
Compound 1D11 (positive control)
Change in body weight (±SEM), g
0
-1
-2
-3
-4
-5
-6
-7
-8
-9
-10
p<0.06

Figure 7

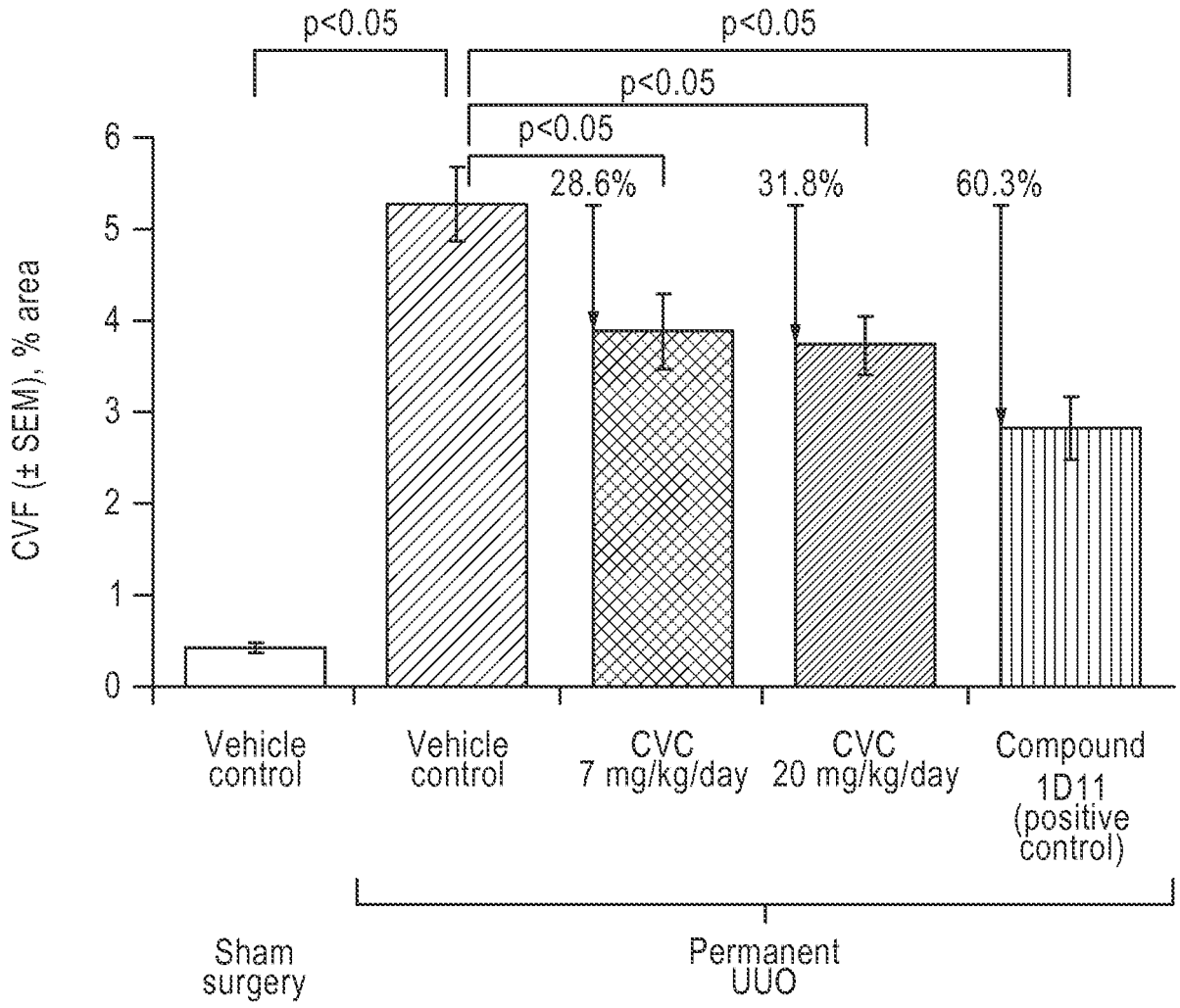

p<0.05
p<0.05
p<0.05
p<0.05
28.6%
31.8%
60.3%
CVF (± SEM), % area
0
1
2
3
4
5
6
Vehicle control
Vehicle control
CVC 7 mg/kg/day
CVC 20 mg/kg/day
Compound 1D11 (positive control)
Sham surgery
Permanent UUO

Figure 8A

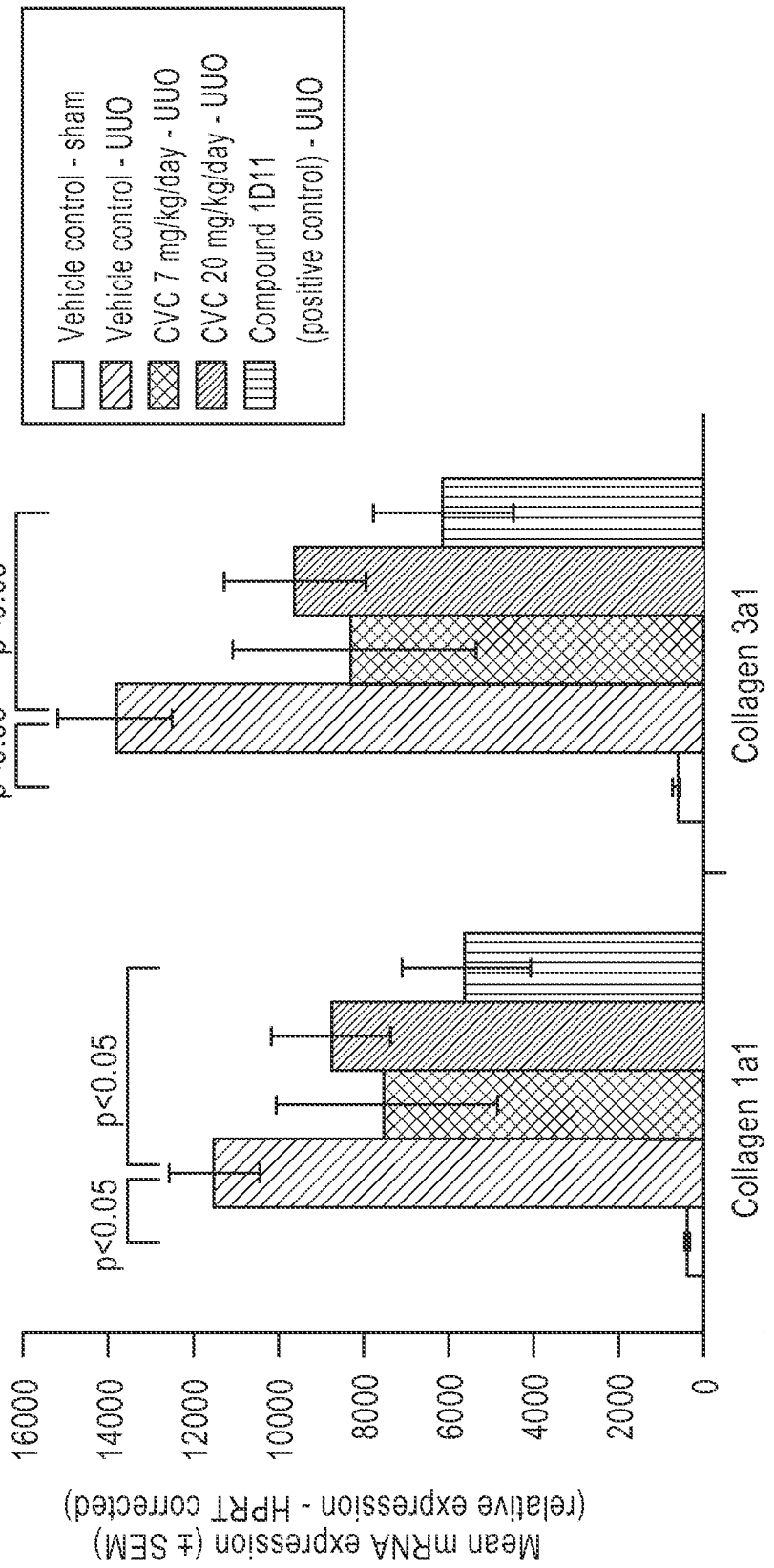

Mean mRNA expression (± SEM)
(relative expression - HPRT corrected)
16000
14000
12000
10000
8000
6000
4000
2000
0
p<0.05
p<0.05
p<0.05
p<0.05
Collagen 1a1
Collagen 3a1
Vehicle control - sham
Vehicle control - UUO
CVC 7 mg/kg/day - UUO
CVC 20 mg/kg/day - UUO
Compound 1D11
(positive control) - UUO

Figure 8B

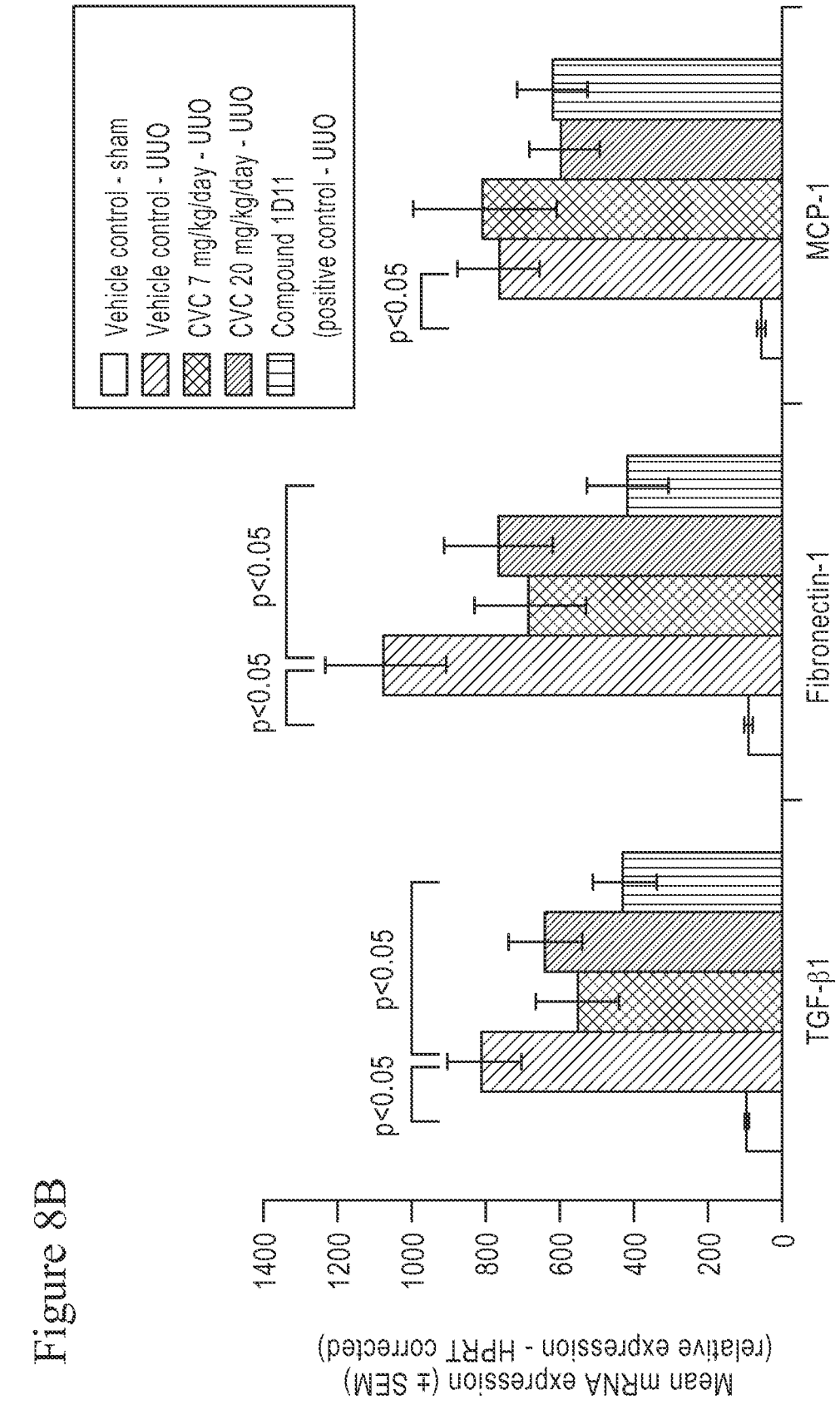

Vehicle control - sham
Vehicle control - UUO
CVC 7 mg/kg/day - UUO
CVC 20 mg/kg/day - UUO
Compound 1D11
(positive control - UUO
Mean mRNA expression (± SEM)
(relative expression - HPRT corrected)
1400
1200
1000
800
600
400
200
0
p<0.05
p<0.05
p<0.05
p<0.05
p<0.05
p<0.05
TGF-β1
Fibronectin-1
MCP-1

Figure 9

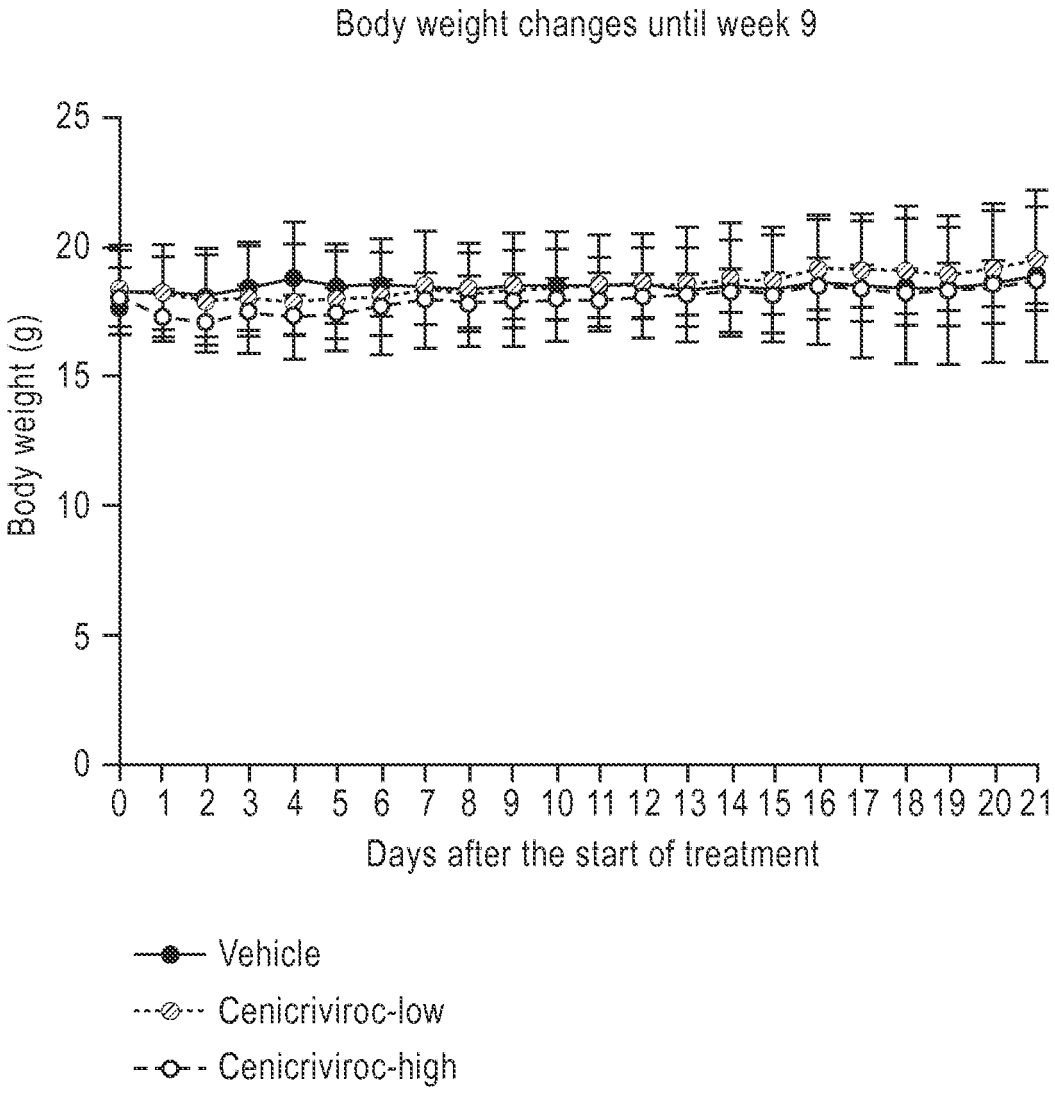

Body weight changes until week 9
Body weight (g)
25
20
15
10
5
0
0 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21
Days after the start of treatment
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 10A

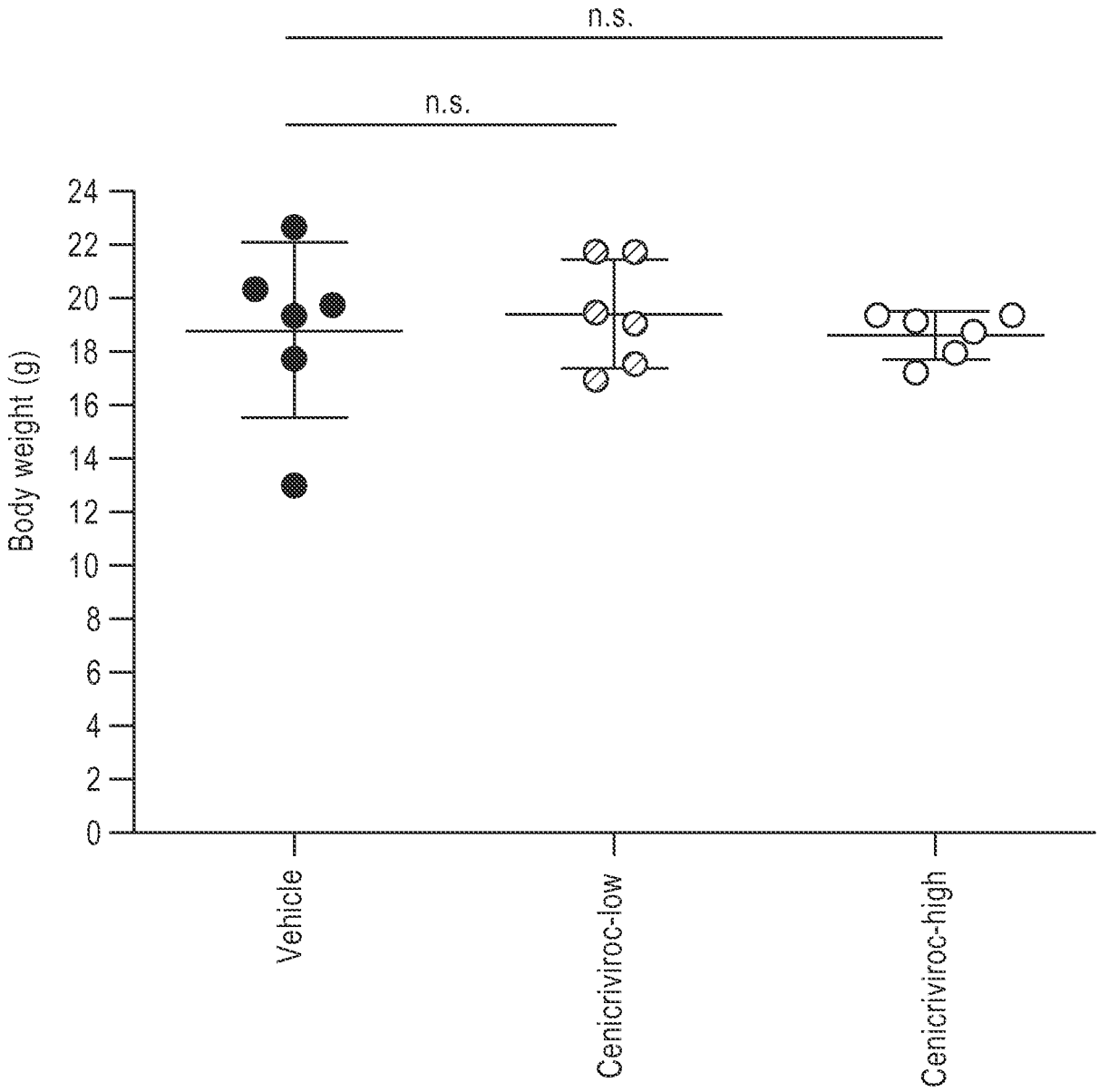

n.s.
n.s.
Body weight (g)
24
22
20
18
16
14
12
10
8
6
4
2
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 10B

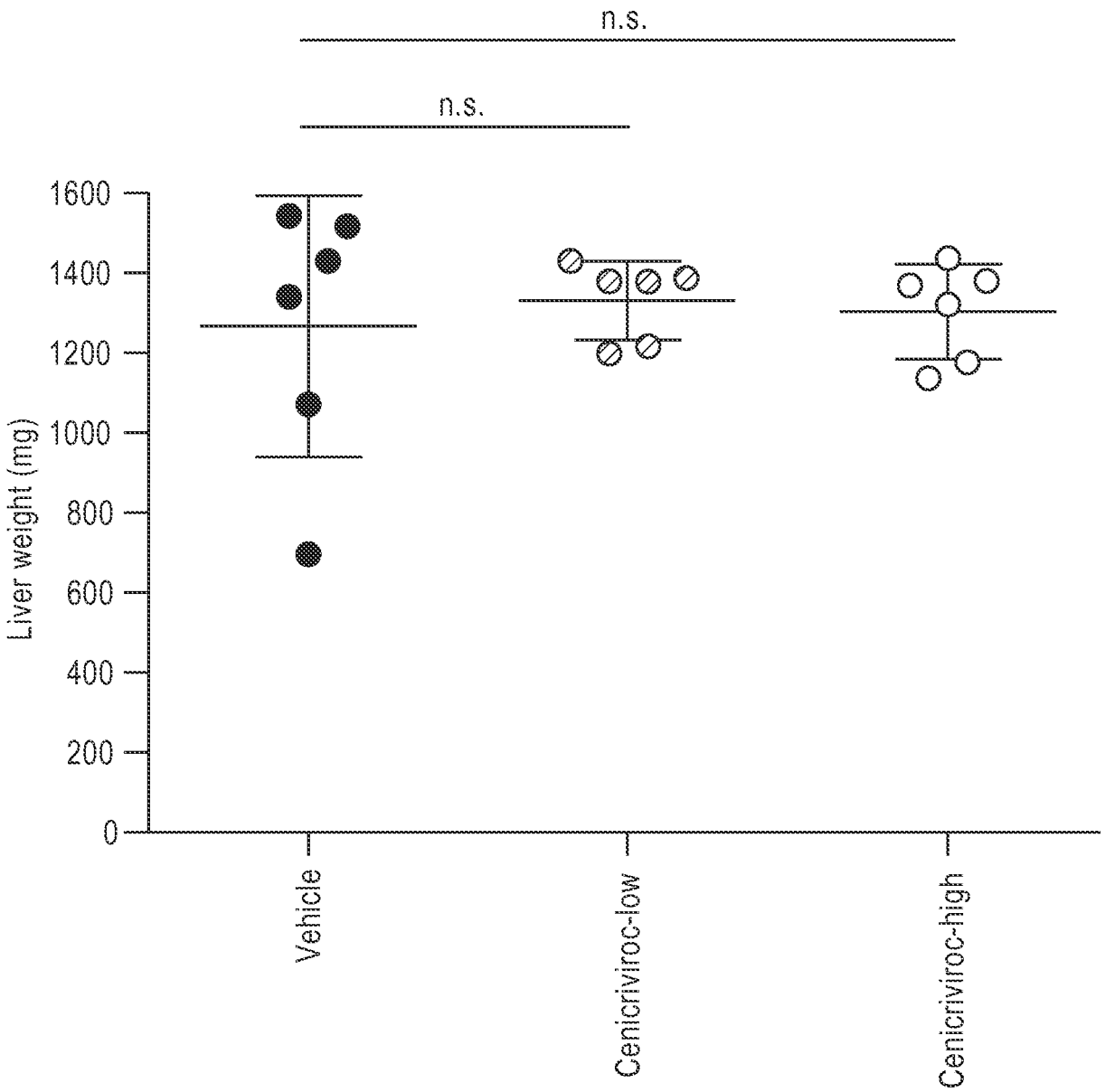

n.s.
n.s.
1600
1400
1200
1000
800
600
400
200
0
Liver weight (mg)
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 10C

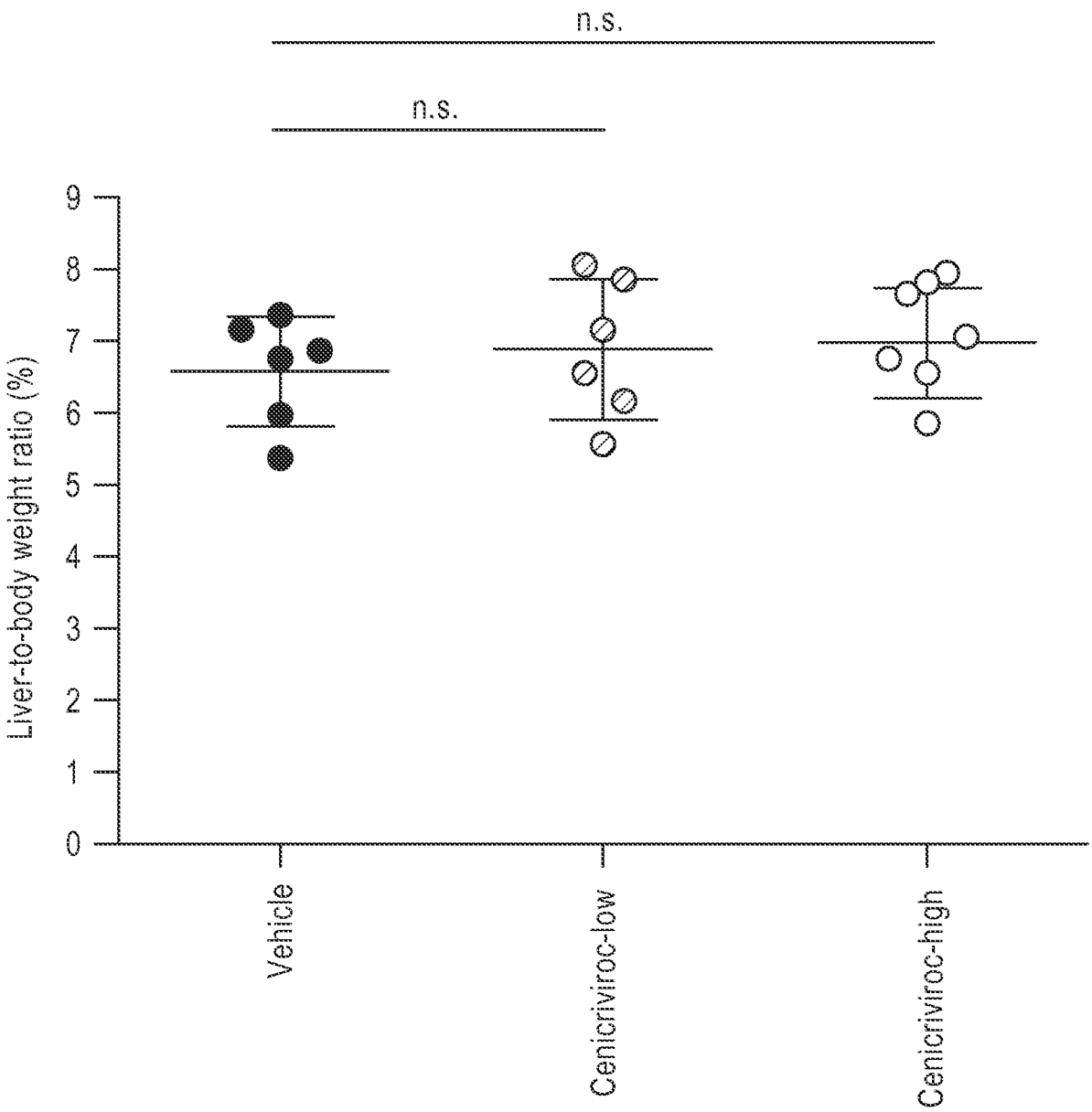

n.s.
n.s.
Liver-to-body weight ratio (%)
0
1
2
3
4
5
6
7
8
9
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 11

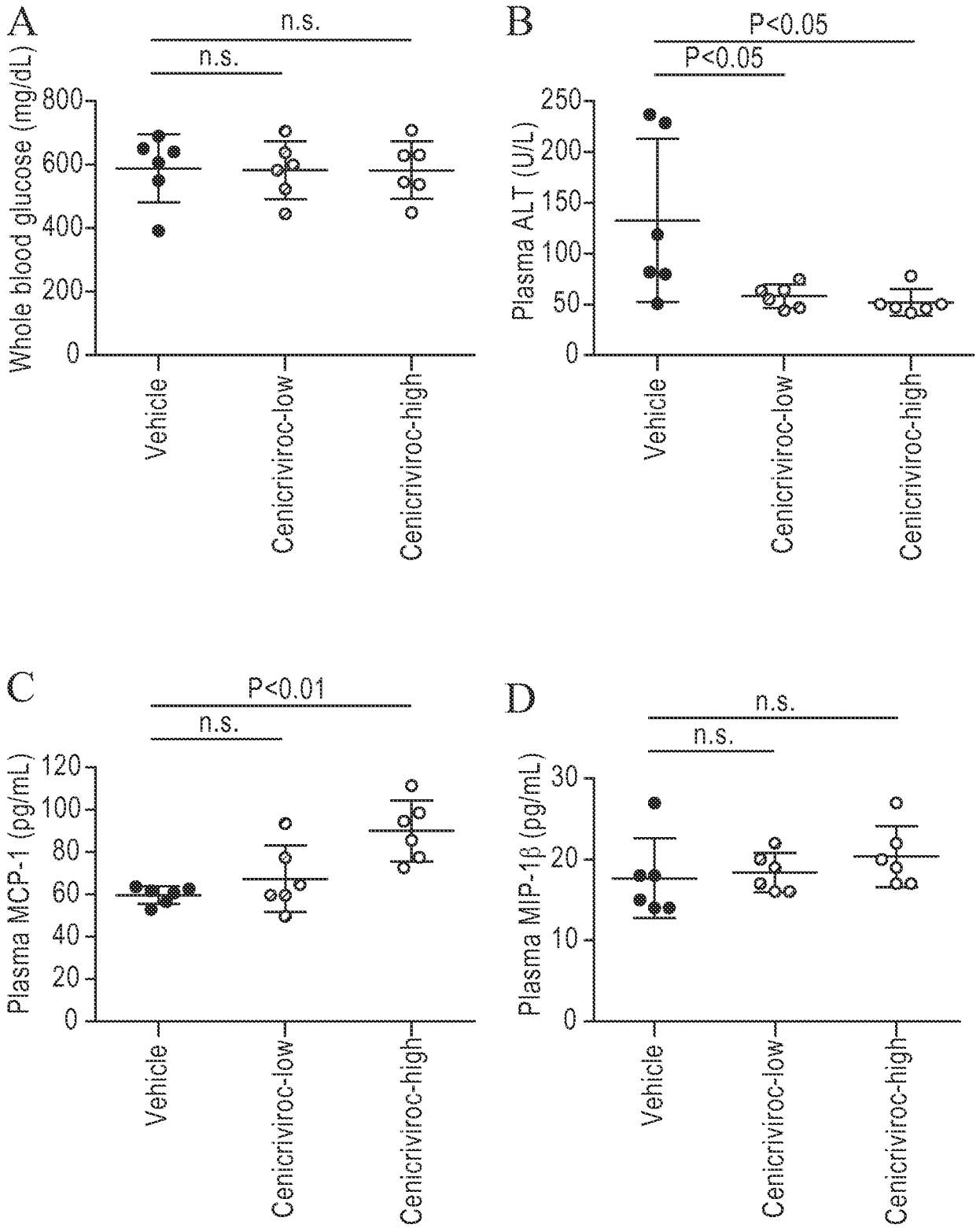

A
n.s.
n.s.
Whole blood glucose (mg/dL)
800
600
400
200
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high
B
P<0.05
P<0.05
Plasma ALT (U/L)
250
200
150
100
50
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high
C
P<0.01
n.s.
Plasma MCP-1 (pg/mL)
120
100
80
60
40
20
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high
D
n.s.
n.s.
Plasma MIP-1β (pg/mL)
30
20
10
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 11

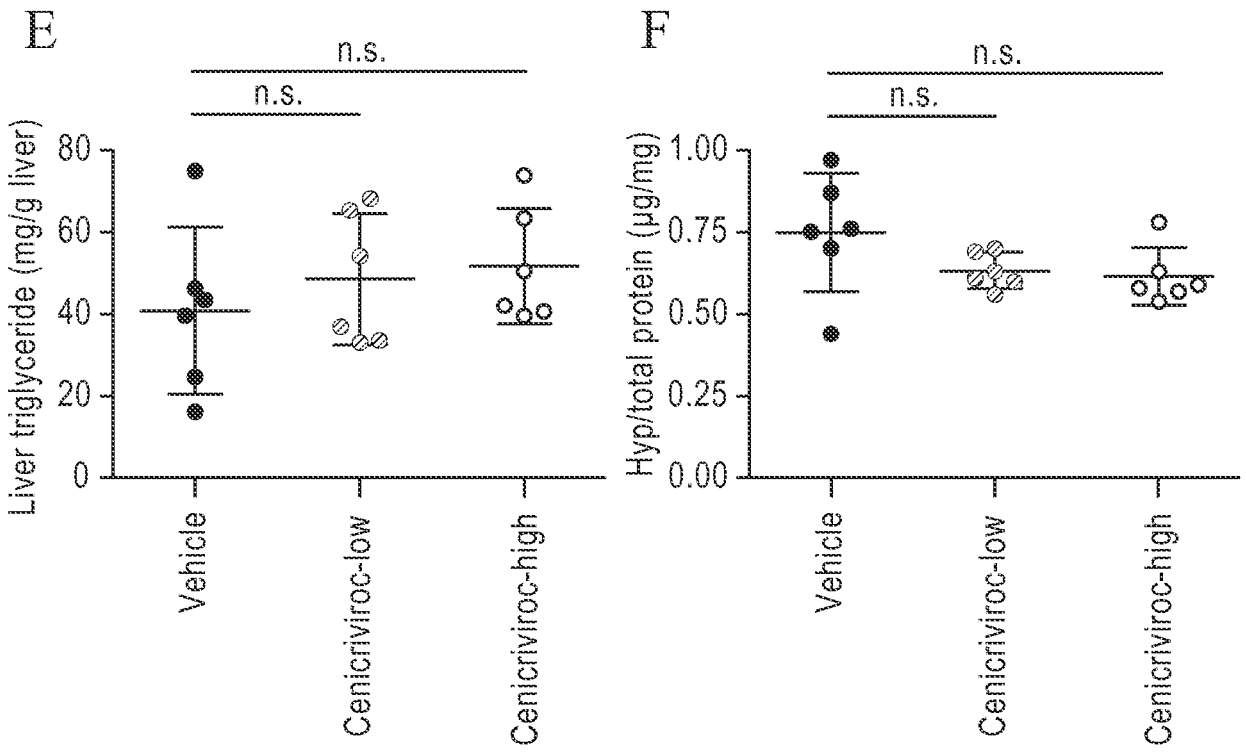

E
n.s.
n.s.
Liver triglyceride (mg/g liver)
80
60
40
20
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high
F
n.s.
n.s.
Hyp/total protein (µg/mg)
1.00
0.75
0.50
0.25
0.00
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 12

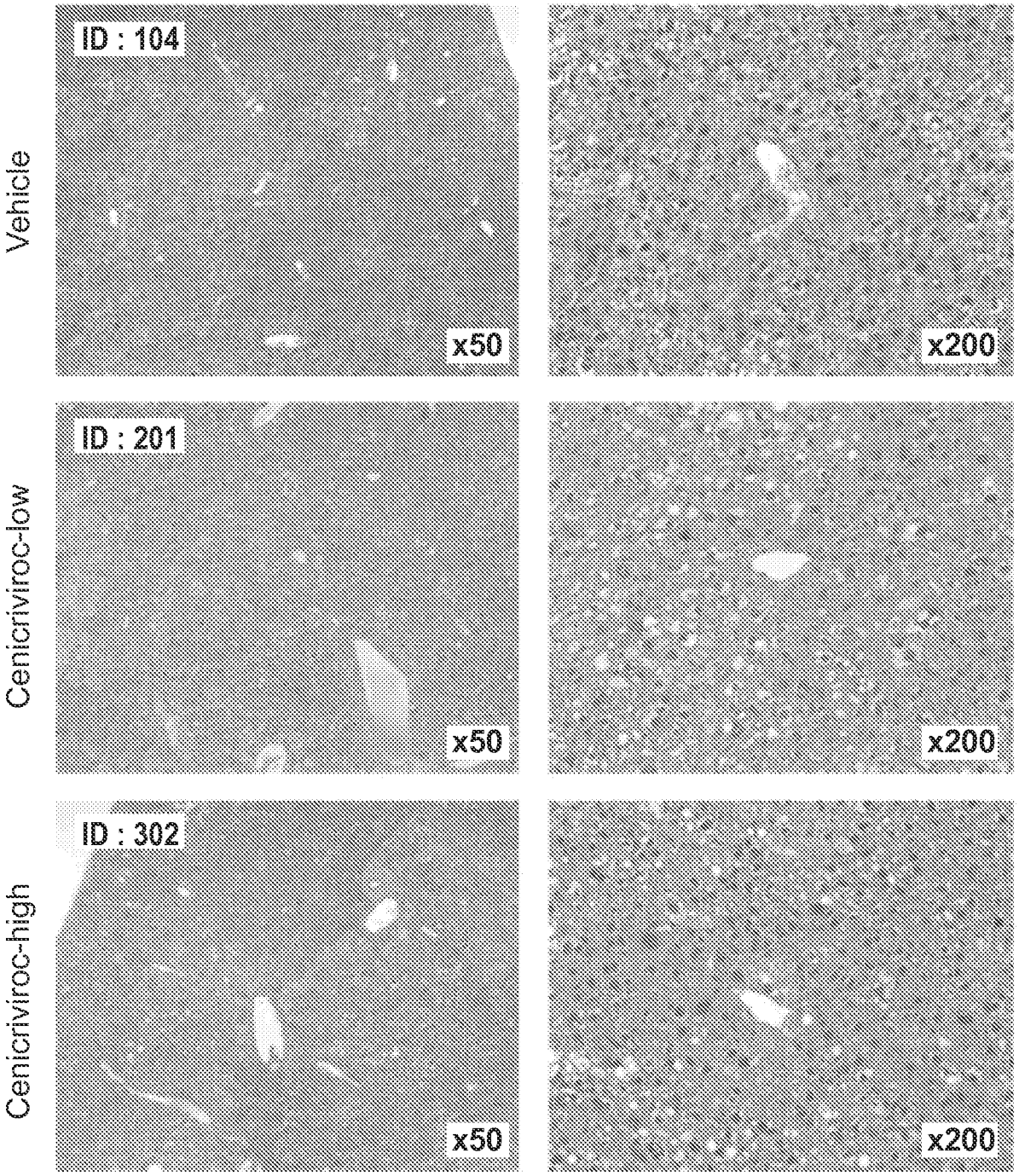

Vehicle
ID : 104
x50
x200
Cenicriviroc-low
ID : 201
x50
x200
Cenicriviroc-high
ID : 302
x50
x200

Figure 13

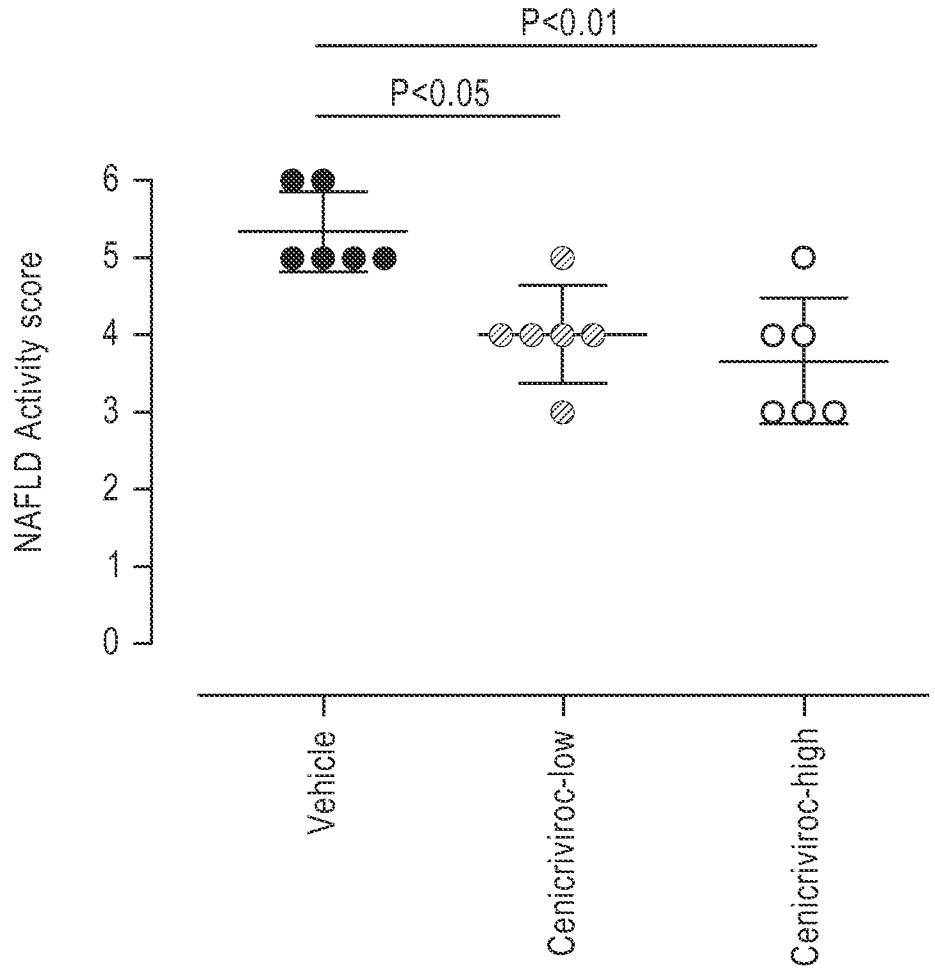

P<0.01
P<0.05
NAFLD Activity score
0
1
2
3
4
5
6
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 14

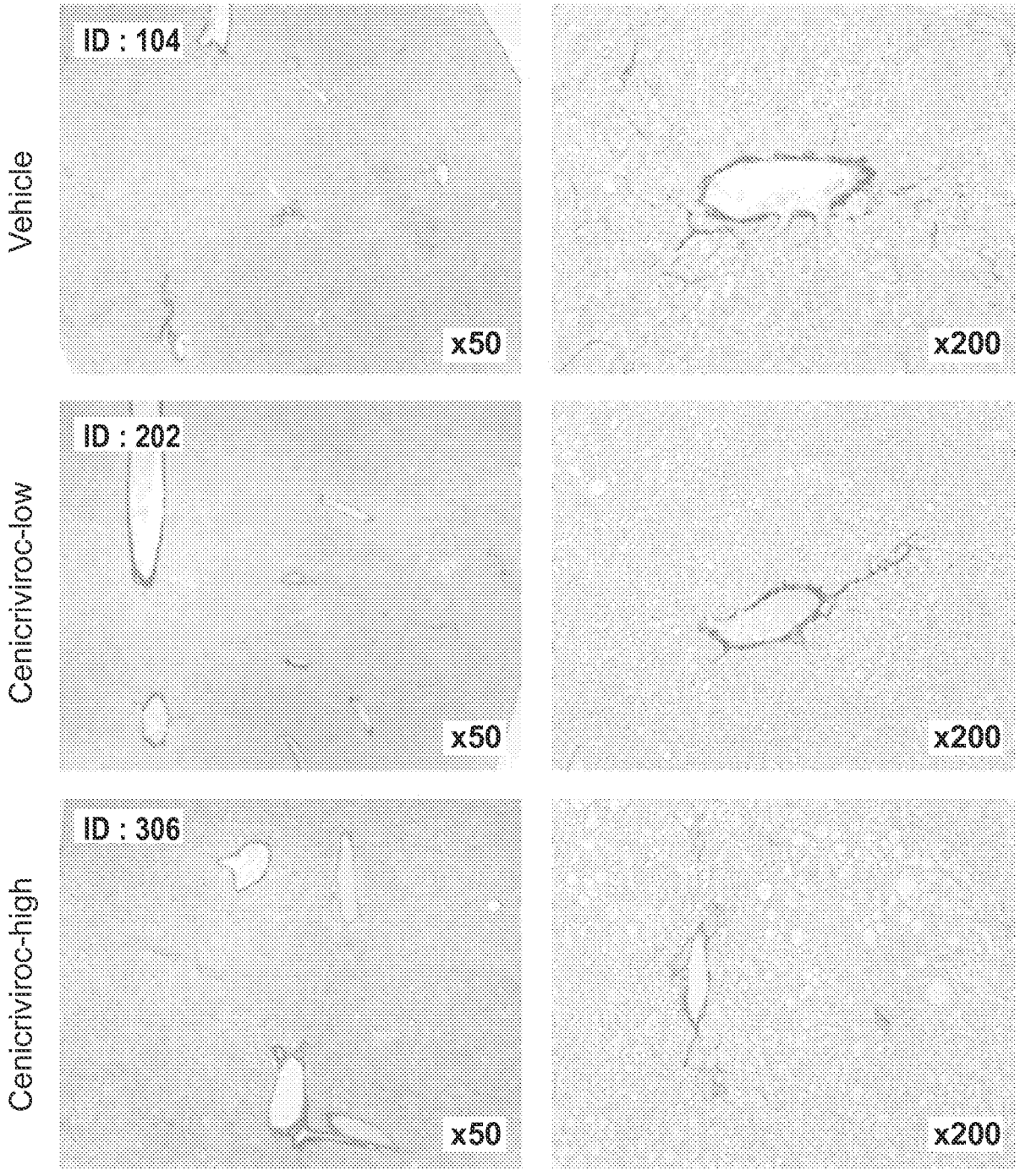

Vehicle
ID : 104
x50
x200
Cenicriviroc-low
ID : 202
x50
x200
Cenicriviroc-high
ID : 306
x50
x200

Figure 15

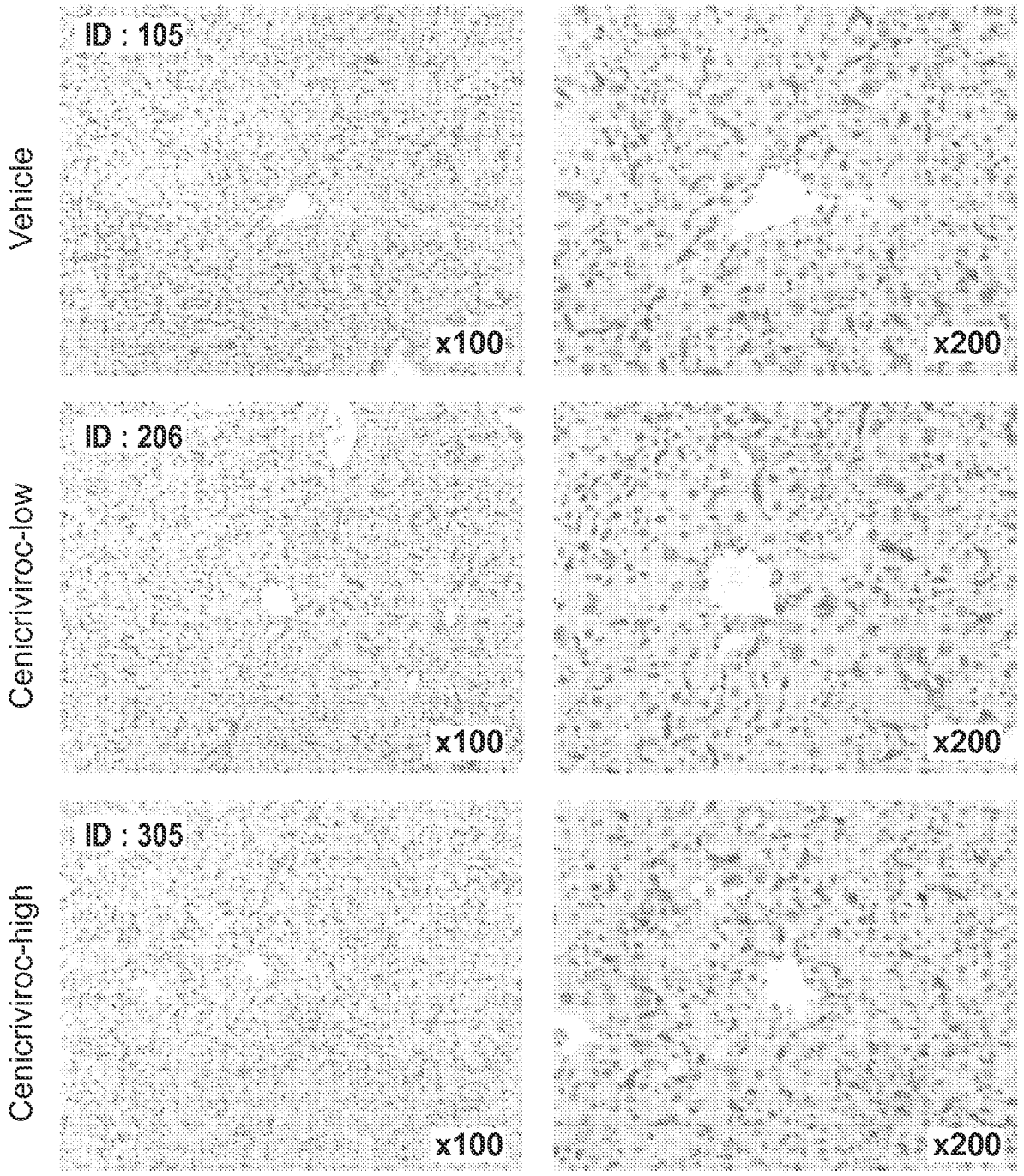

Vehicle
ID : 105
x100
x200
Cenicriviroc-low
ID : 206
x100
x200
Cenicriviroc-high
ID : 305
x100
x200

Figure 16

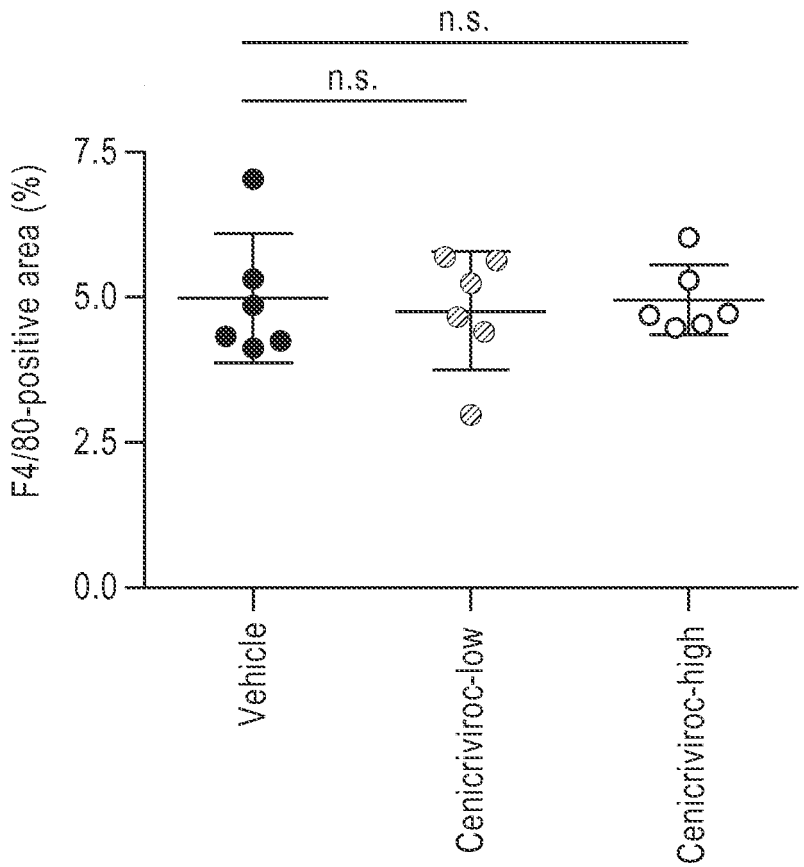

n.s.
n.s.
F4/80-positive area (%)
7.5
5.0
2.5
0.0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 17

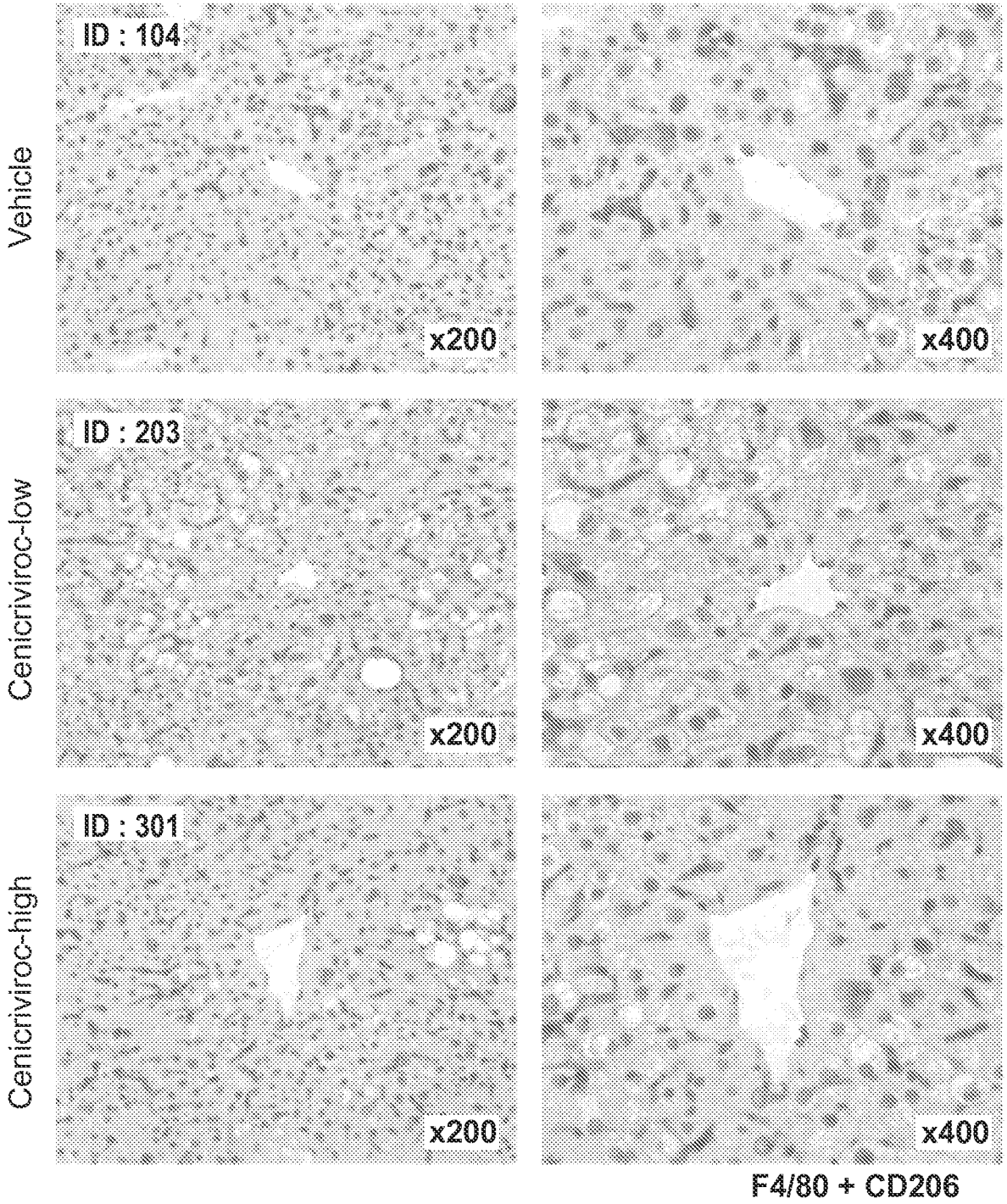

Vehicle
ID : 104
x200
x400
Cenicriviroc-low
ID : 203
x200
x400
Cenicriviroc-high
ID : 301
x200
x400
F4/80 + CD206

Figure 18

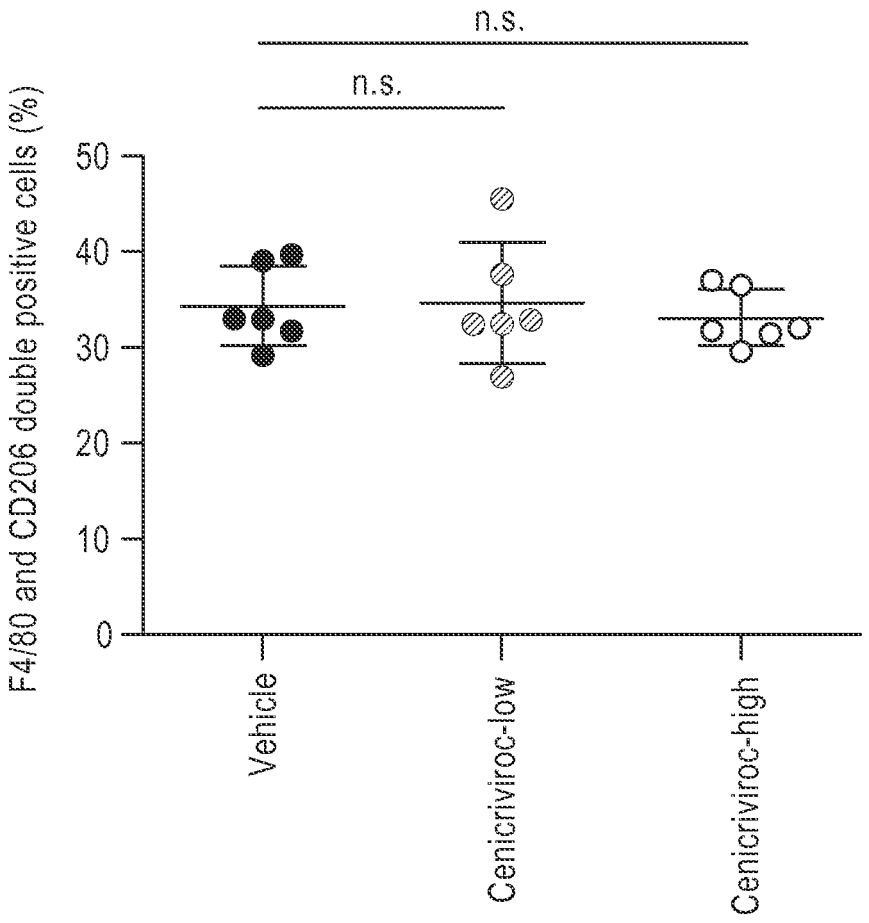

n.s.
n.s.
F4/80 and CD206 double positive cells (%)
50
40
30
20
10
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 19

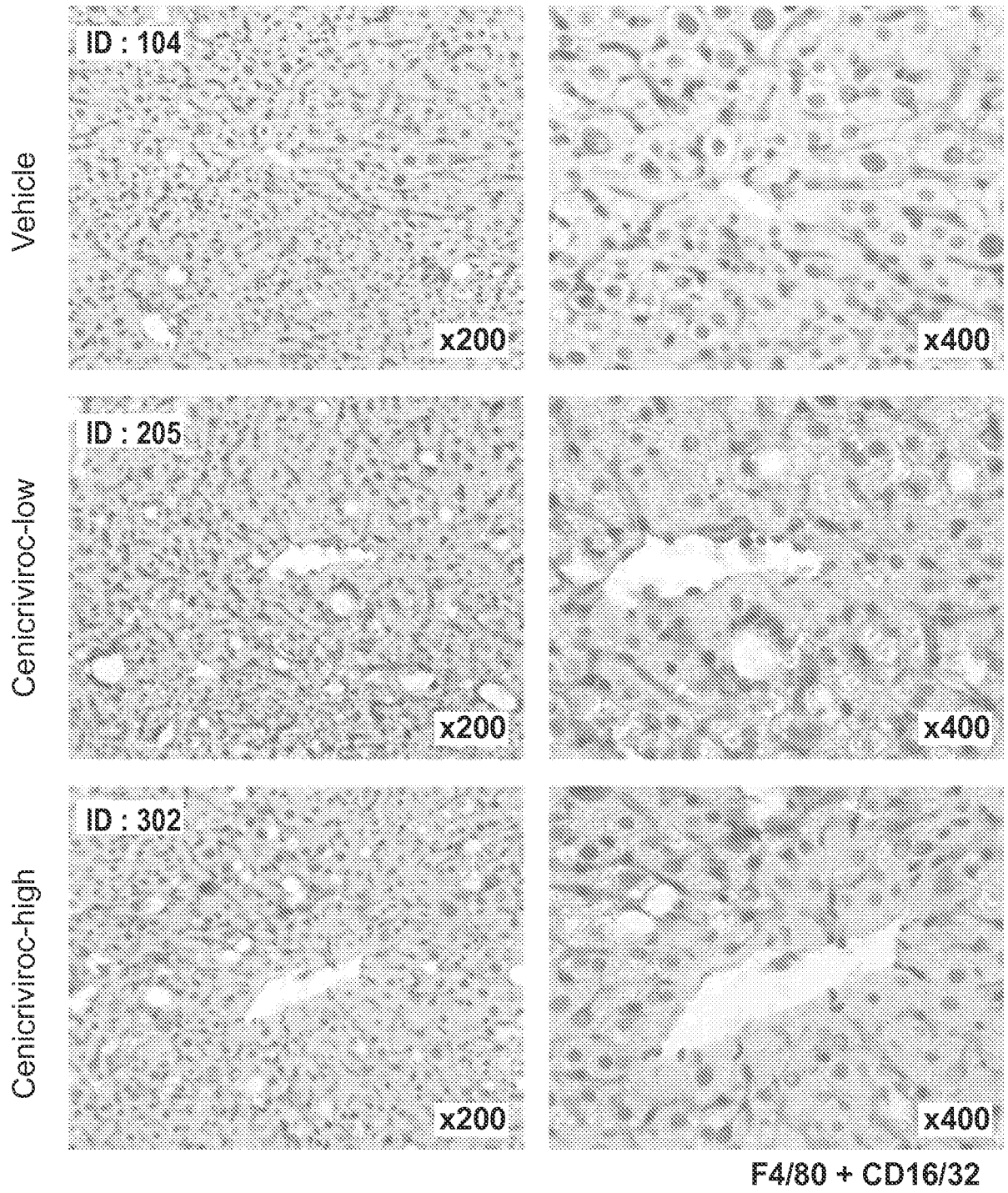

Vehicle
Cenicriviroc-low
Cenicriviroc-high
ID : 104
ID : 205
ID : 302
x200
x400
x200
x400
x200
x400
F4/80 + CD16/32

Figure 20

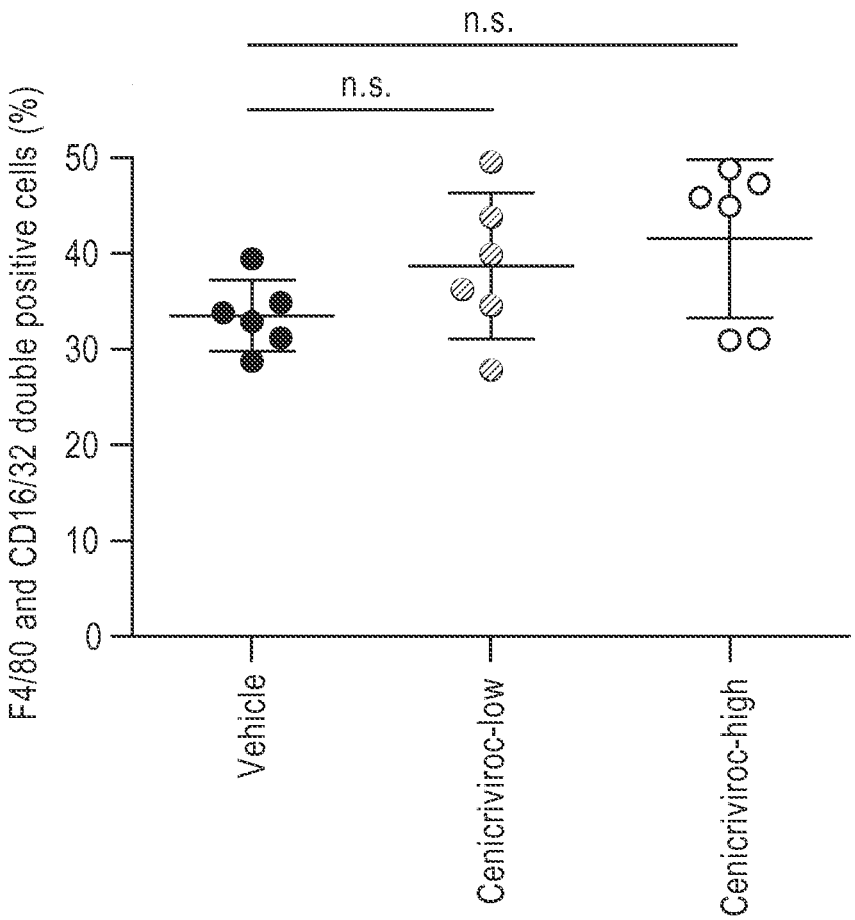

n.s.
n.s.
F4/80 and CD16/32 double positive cells (%)
50
40
30
20
10
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 21

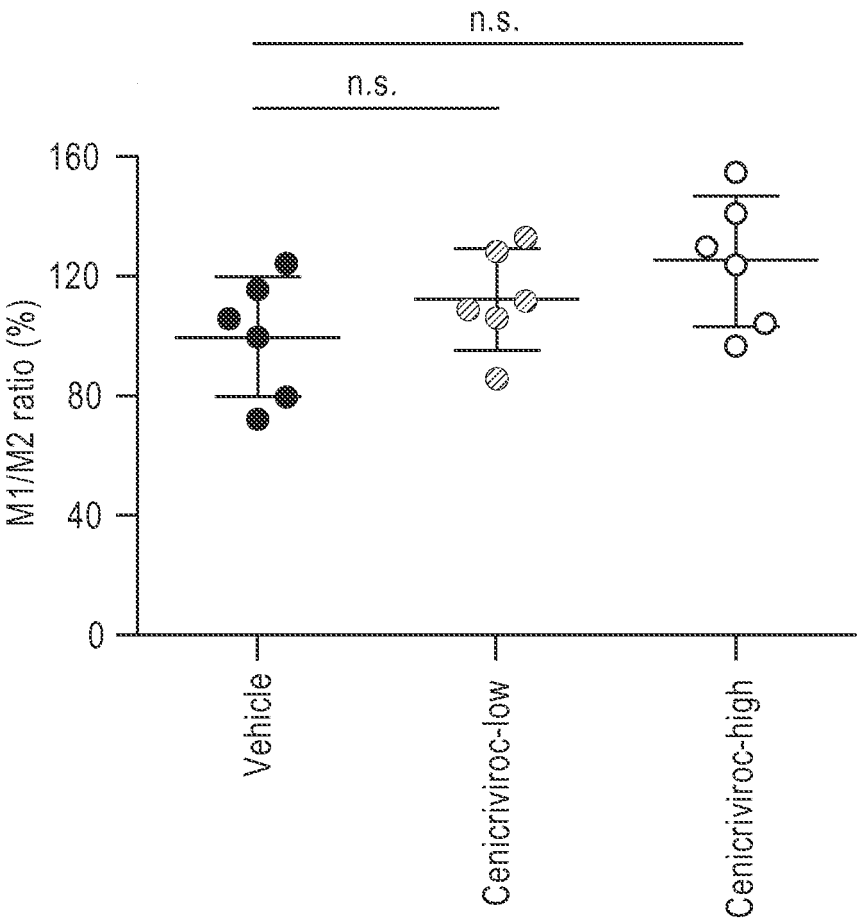

n.s.
n.s.
160
120
80
40
0
M1/M2 ratio (%)
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 22

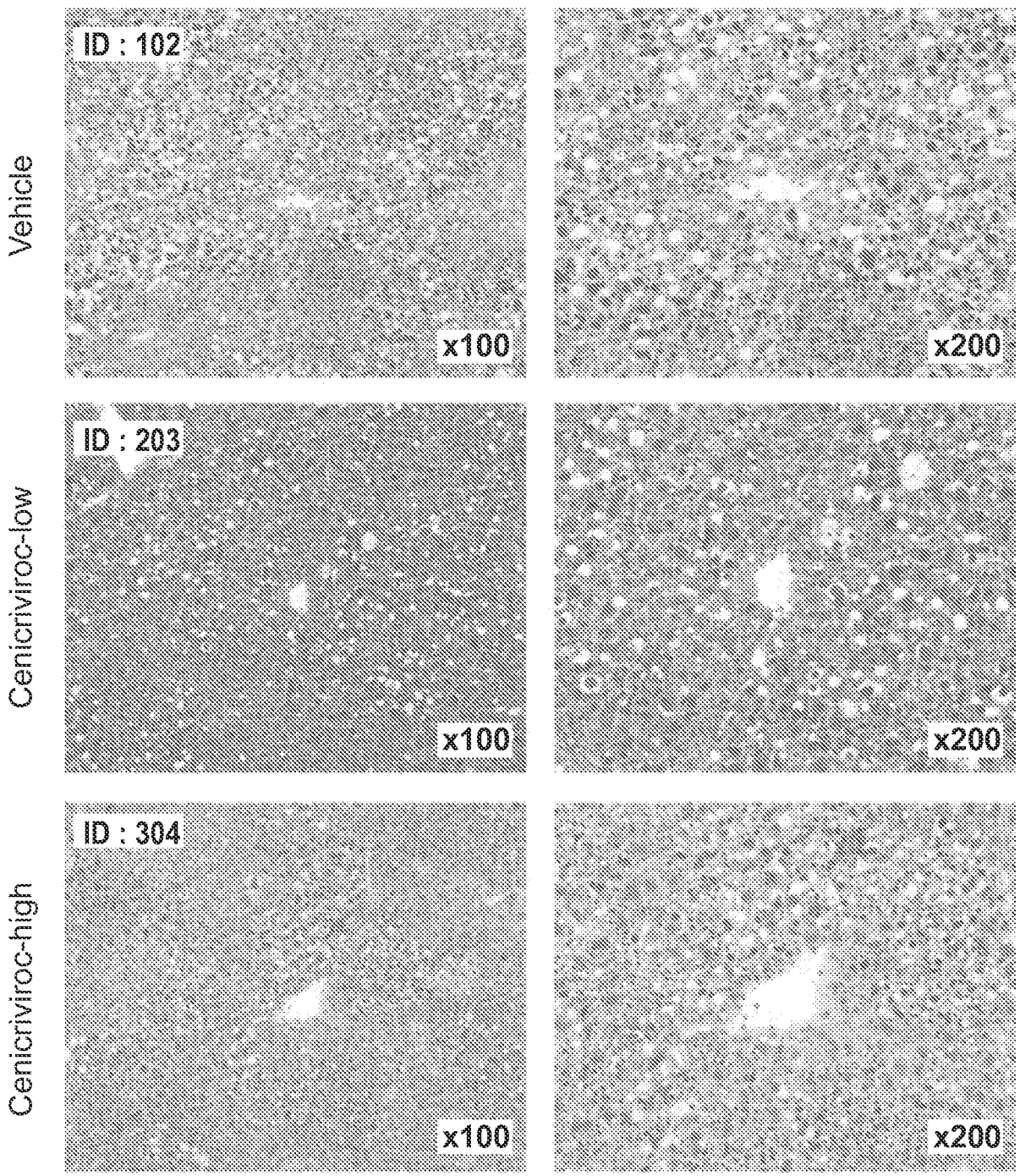

Vehicle
ID : 102
x100
x200
Cenicriviroc-low
ID : 203
x100
x200
Cenicriviroc-high
ID : 304
x100
x200

Figure 23

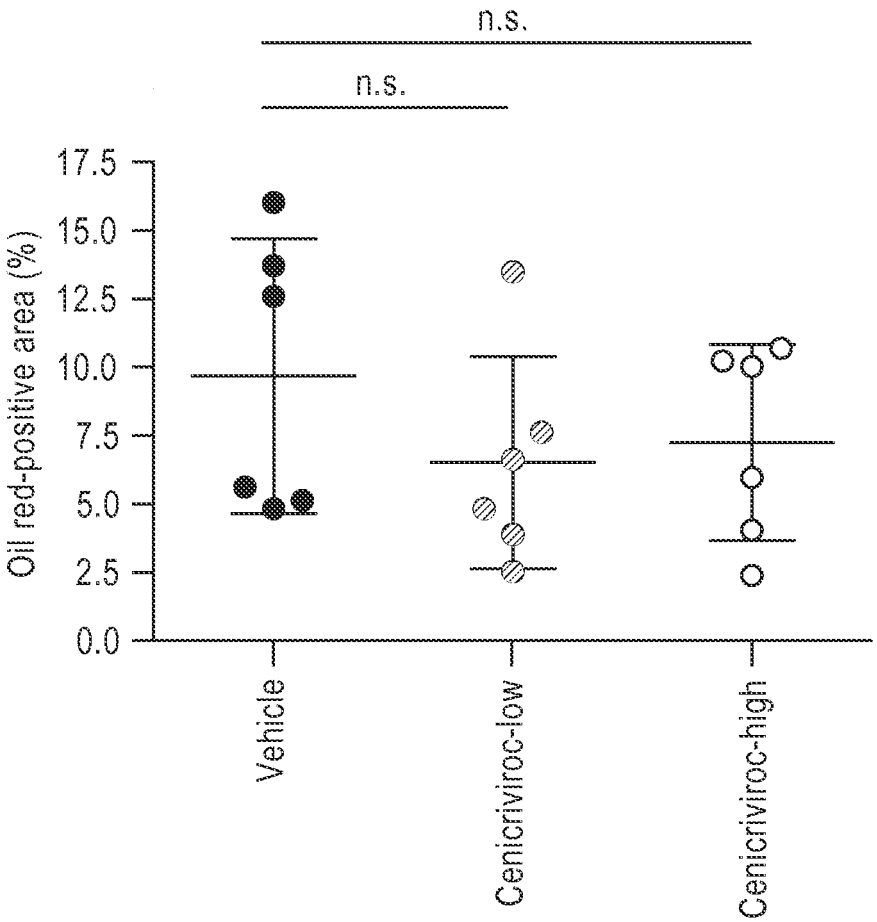

n.s.
n.s.
Oil red-positive area (%)
17.5
15.0
12.5
10.0
7.5
5.0
2.5
0.0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 24

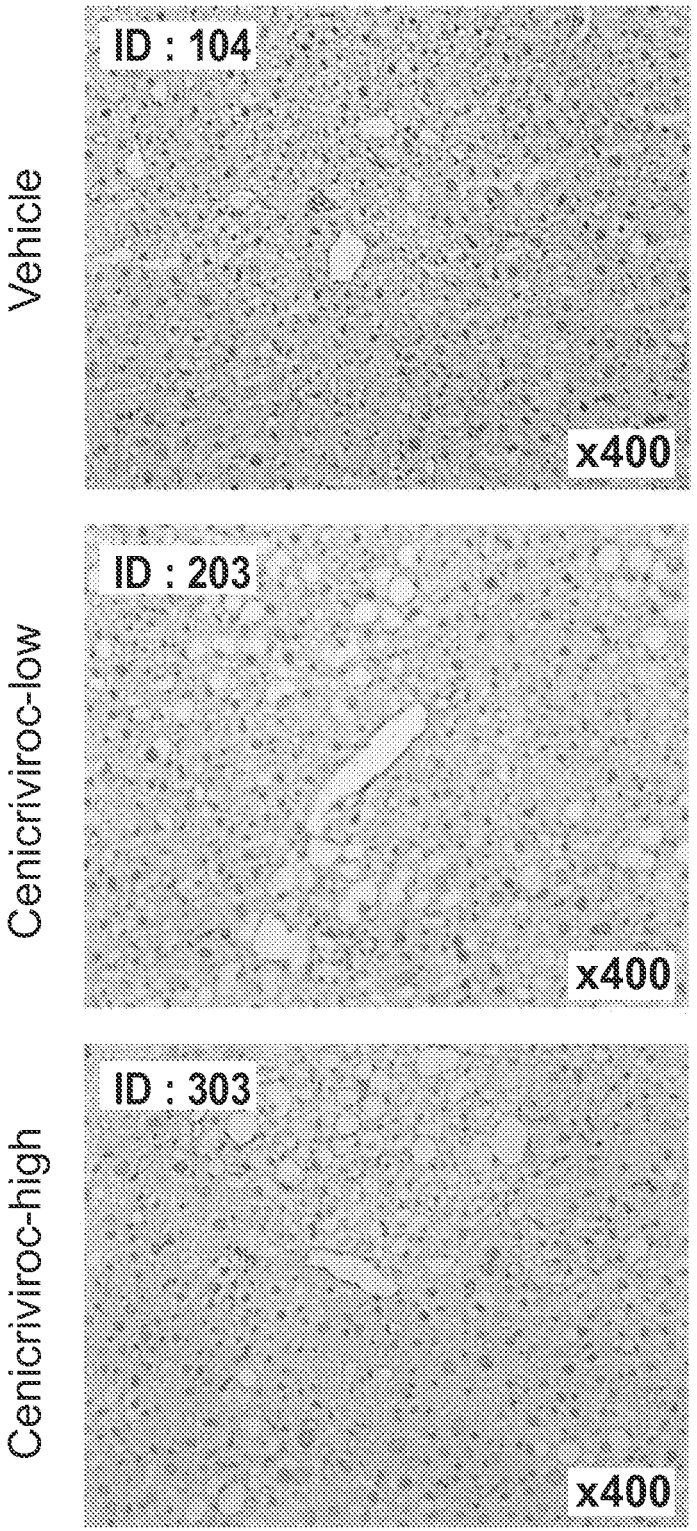

Vehicle
ID : 104
x400
Cenicriviroc-low
ID : 203
x400
Cenicriviroc-high
ID : 303
x400

Figure 25

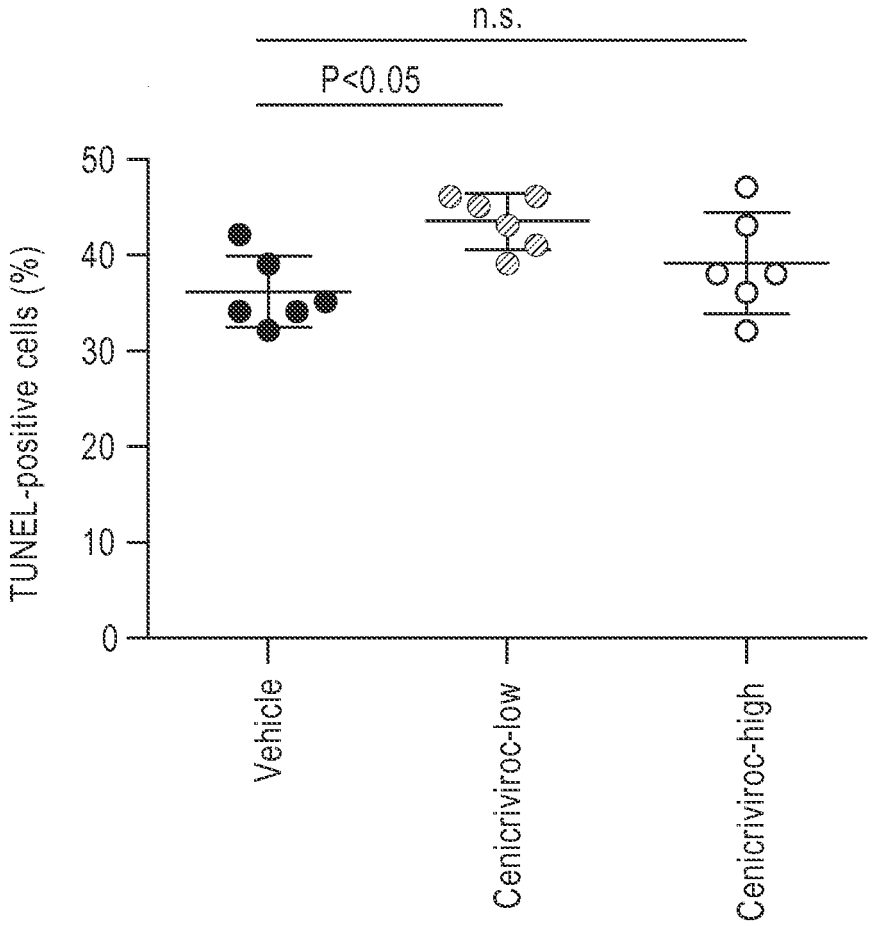

n.s.
P<0.05
TUNEL-positive cells (%)
50
40
30
20
10
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 26

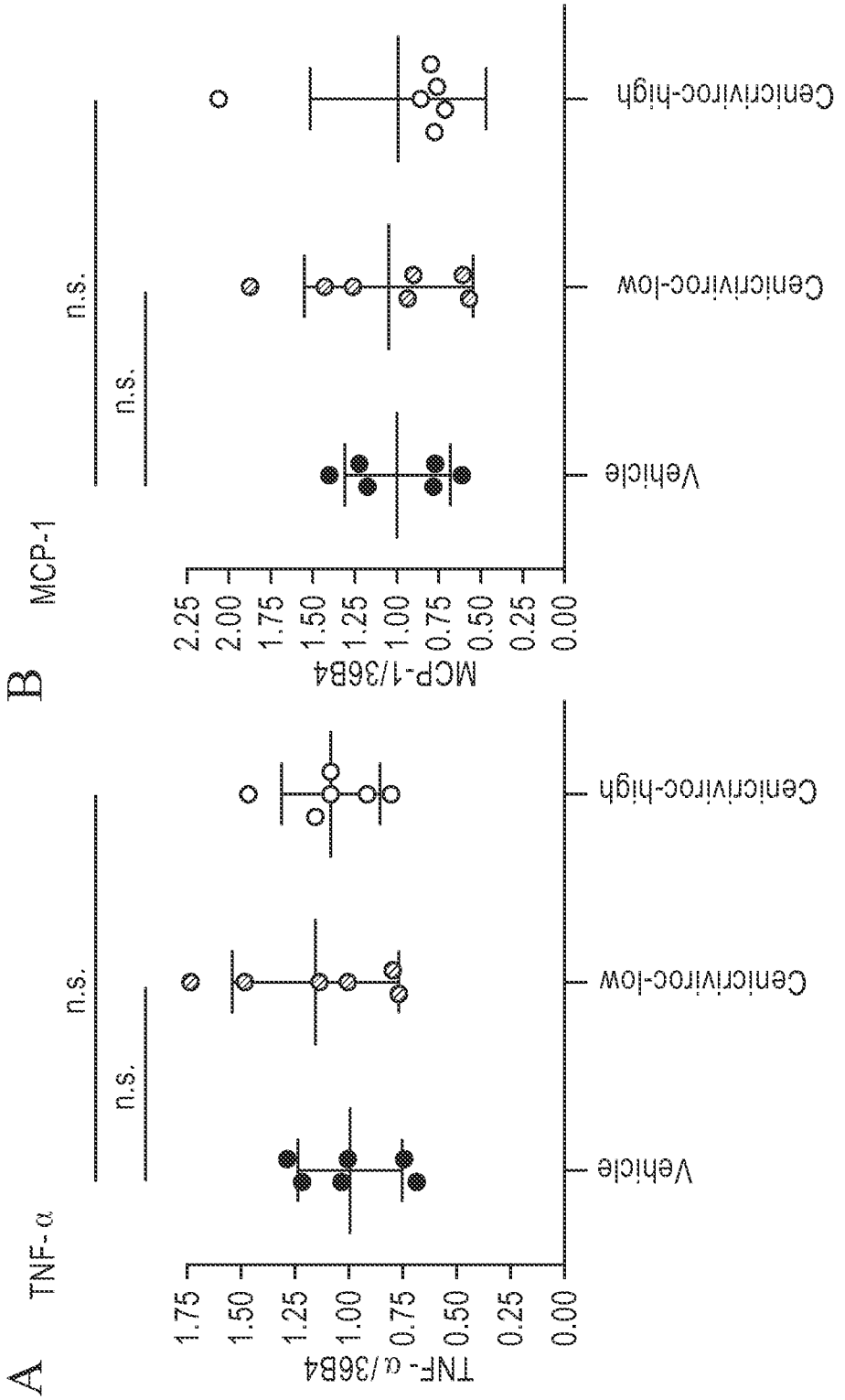

A
TNF-α
TNF-α/36B4
1.75
1.50
1.25
1.00
0.75
0.50
0.25
0.00
n.s.
n.s.
Vehicle
Cenicriviroc-low
Cenicriviroc-high
B
MCP-1
MCP-1/36B4
2.25
2.00
1.75
1.50
1.25
1.00
0.75
0.50
0.25
0.00
n.s.
n.s.
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 26

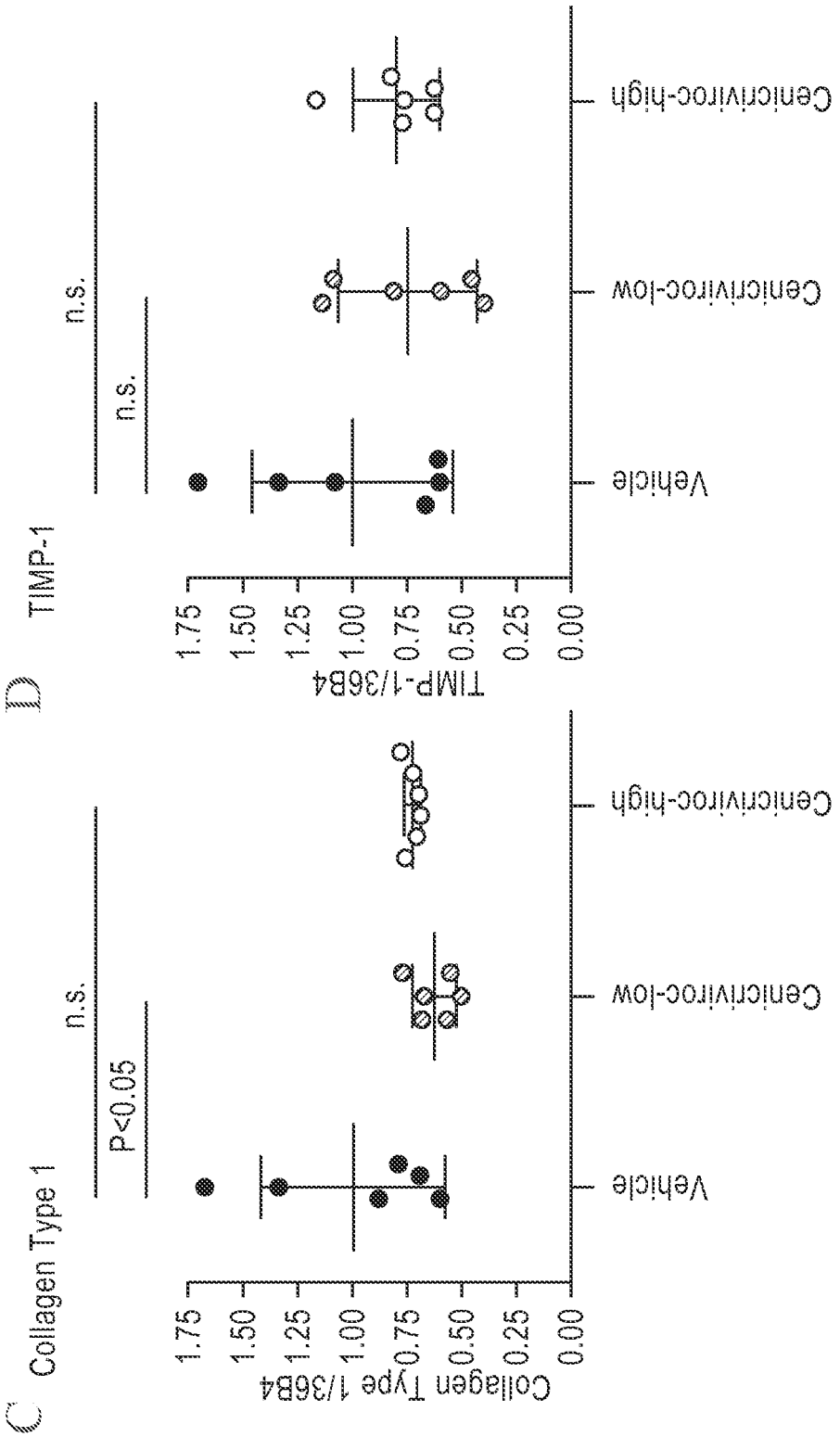

C
Collagen Type 1
n.s.
P<0.05
Collagen Type 1/36B4
1.75
1.50
1.25
1.00
0.75
0.50
0.25
0.00
Vehicle
Cenicriviroc-low
Cenicriviroc-high
D
TIMP-1
n.s.
n.s.
TIMP-1/36B4
1.75
1.50
1.25
1.00
0.75
0.50
0.25
0.00
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 27A Plate 1-1 : 36B4

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	21.3	-	4.00E+00	3.67E+00
A3	UNKN	A	101	21.49	-	3.54E+00	3.67E+00
A4	UNKN	A	101	21.52	-	3.47E+00	3.67E+00
B2	UNKN	A	102	21.37	-	3.82E+00	4.09E+00
B3	UNKN	A	102	21.28	-	4.05E+00	4.09E+00
B4	UNKN	A	102	21.15	-	4.40E+00	4.09E+00
C2	UNKN	A	103	21.1	-	4.54E+00	4.21E+00
C3	UNKN	A	103	21.31	-	3.97E+00	4.21E+00
C4	UNKN	A	103	21.26	-	4.10E+00	4.21E+00
D2	UNKN	A	104	21.06	-	4.66E+00	4.28E+00
D3	UNKN	A	104	21.32	-	3.95E+00	4.28E+00
D4	UNKN	A	104	21.21	-	4.23E+00	4.28E+00
E2	UNKN	A	105	20.97	-	4.94E+00	4.86E+00
E3	UNKN	A	105	21.1	-	4.54E+00	4.86E+00
E4	UNKN	A	105	20.92	-	5.10E+00	4.86E+00
F2	UNKN	A	106	20.9	-	5.16E+00	5.16E+00
F3	UNKN	A	106	20.91	-	5.13E+00	5.16E+00
F4	UNKN	A	106	20.89	-	5.19E+00	5.16E+00
A5	UNKN	A	201	21.26	-	4.10E+00	4.30E+00
A6	UNKN	A	201	21.11	-	4.51E+00	4.30E+00
A7	UNKN	A	201	21.19	-	4.29E+00	4.30E+00
B5	UNKN	A	202	20.96	-	4.97E+00	5.03E+00
B6	UNKN	A	202	20.86	-	5.29E+00	5.03E+00
B7	UNKN	A	202	21	-	4.84E+00	5.03E+00
C5	UNKN	A	203	21.31	-	3.97E+00	4.20E+00
C6	UNKN	A	203	21.02	-	4.78E+00	4.20E+00
C7	UNKN	A	203	21.36	-	3.85E+00	4.20E+00
D5	UNKN	A	204	21.17	-	4.34E+00	4.45E+00
D6	UNKN	A	204	21.12	-	4.48E+00	4.45E+00
D7	UNKN	A	204	21.11	-	4.51E+00	4.45E+00
E5	UNKN	A	205	20.98	-	4.90E+00	4.77E+00
E6	UNKN	A	205	21.1	-	4.54E+00	4.77E+00
E7	UNKN	A	205	20.99	-	4.87E+00	4.77E+00
F5	UNKN	A	206	21.11	-	4.51E+00	4.53E+00
F6	UNKN	A	206	21.01	-	4.81E+00	4.53E+00
F7	UNKN	A	206	21.2	-	4.26E+00	4.53E+00
A8	UNKN	A	301	21.1	-	4.54E+00	4.49E+00
A9	UNKN	A	301	21.1	-	4.54E+00	4.49E+00
A10	UNKN	A	301	21.16	-	4.37E+00	4.49E+00
B8	UNKN	A	302	21.34	-	3.90E+00	3.97E+00
B9	UNKN	A	302	21.25	-	4.13E+00	3.97E+00
B10	UNKN	A	302	21.34	-	3.90E+00	3.97E+00
C8	UNKN	A	303	21.03	-	4.75E+00	4.79E+00
C9	UNKN	A	303	20.97	-	4.94E+00	4.79E+00
C10	UNKN	A	303	21.05	-	4.69E+00	4.79E+00
D8	UNKN	A	304	20.87	-	5.26E+00	5.10E+00
D9	UNKN	A	304	20.94	-	5.03E+00	5.10E+00
D10	UNKN	A	304	20.95	-	5.00E+00	5.10E+00
E8	UNKN	A	305	21.02	-	4.78E+00	4.82E+00
E9	UNKN	A	305	20.98	-	4.90E+00	4.82E+00
E10	UNKN	A	305	21.02	-	4.78E+00	4.82E+00
F8	UNKN	A	306	21.17	-	4.34E+00	4.51E+00
F9	UNKN	A	306	21.04	-	4.72E+00	4.51E+00
F10	UNKN	A	306	21.13	-	4.46E+00	4.51E+00
B11	STD	A		19.8	1.00E+01	1.04E+01	-
C11	STD	A		19.92	1.00E+01	9.65E+00	-
D11	STD	A		19.8	1.00E+01	1.04E+01	-
E11	STD	A		22.06	2.50E+00	2.46E+00	-
F11	STD	A		22.1	2.50E+00	2.40E+00	-
G11	STD	A		22.02	2.50E+00	2.52E+00	-
B12	STD	A		24.11	6.25E-01	6.65E-01	-
C12	STD	A		24.29	6.25E-01	5.92E-01	-
D12	STD	A		24.33	6.25E-01	5.77E-01	-
E12	STD	A		26.3	1.56E-01	1.64E-01	-
F12	STD	A		26.11	1.56E-01	1.85E-01	-
G12	STD	A		26.63	1.56E-01	1.33E-01	-

Figure 27B Plate 1-2 : TNF-α

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	30.72	-	4.40E+00	4.47E+00
A3	UNKN	A	101	30.65	-	4.59E+00	4.47E+00
A4	UNKN	A	101	30.71	-	4.42E+00	4.47E+00
B2	UNKN	A	102	30.44	-	5.21E+00	5.26E+00
B3	UNKN	A	102	30.44	-	5.21E+00	5.26E+00
B4	UNKN	A	102	30.39	-	5.37E+00	5.26E+00
C2	UNKN	A	103	31.19	-	3.31E+00	2.88E+00
C3	UNKN	A	103	31.53	-	2.69E+00	2.88E+00
C4	UNKN	A	103	31.56	-	2.65E+00	2.88E+00
D2	UNKN	A	104	30.9	-	3.94E+00	4.40E+00
D3	UNKN	A	104	30.54	-	4.90E+00	4.40E+00
D4	UNKN	A	104	30.73	-	4.37E+00	4.40E+00
E2	UNKN	A	105	30.91	-	3.92E+00	3.62E+00
E3	UNKN	A	105	31.05	-	3.60E+00	3.62E+00
E4	UNKN	A	105	31.18	-	3.33E+00	3.62E+00
F2	UNKN	A	106	30.49	-	5.05E+00	5.16E+00
F3	UNKN	A	106	30.59	-	4.76E+00	5.16E+00
F4	UNKN	A	106	30.3	-	5.67E+00	5.16E+00
A5	UNKN	A	201	30.51	-	4.99E+00	4.86E+00
A6	UNKN	A	201	30.69	-	4.48E+00	4.86E+00
A7	UNKN	A	201	30.47	-	5.11E+00	4.86E+00
B5	UNKN	A	202	30.31	-	5.63E+00	5.06E+00
B6	UNKN	A	202	30.62	-	4.67E+00	5.06E+00
B7	UNKN	A	202	30.55	-	4.87E+00	5.06E+00
C5	UNKN	A	203	31.05	-	3.60E+00	3.23E+00
C6	UNKN	A	203	31.18	-	3.33E+00	3.23E+00
C7	UNKN	A	203	31.5	-	2.74E+00	3.23E+00
D5	UNKN	A	204	31.17	-	3.35E+00	3.55E+00
D6	UNKN	A	204	31.01	-	3.69E+00	3.55E+00
D7	UNKN	A	204	31.04	-	3.62E+00	3.55E+00
E5	UNKN	A	205	30.15	-	6.20E+00	7.06E+00
E6	UNKN	A	205	29.78	-	7.76E+00	7.06E+00
E7	UNKN	A	205	29.9	-	7.22E+00	7.06E+00
F5	UNKN	A	206	29.9	-	7.22E+00	7.82E+00
F6	UNKN	A	206	29.79	-	7.71E+00	7.82E+00
F7	UNKN	A	206	29.62	-	8.55E+00	7.82E+00
A8	UNKN	A	301	30.59	-	4.76E+00	5.19E+00
A9	UNKN	A	301	30.47	-	5.11E+00	5.19E+00
A10	UNKN	A	301	30.29	-	5.70E+00	5.19E+00
B8	UNKN	A	302	30.72	-	4.40E+00	4.30E+00
B9	UNKN	A	302	30.6	-	4.73E+00	4.30E+00
B10	UNKN	A	302	30.97	-	3.78E+00	4.30E+00
C8	UNKN	A	303	29.8	-	7.67E+00	7.02E+00
C9	UNKN	A	303	30.04	-	6.63E+00	7.02E+00
C10	UNKN	A	303	30.01	-	6.75E+00	7.02E+00
D8	UNKN	A	304	30.93	-	3.87E+00	4.10E+00
D9	UNKN	A	304	30.86	-	4.04E+00	4.10E+00
D10	UNKN	A	304	30.72	-	4.40E+00	4.10E+00
E8	UNKN	A	305	30.58	-	4.78E+00	4.40E+00
E9	UNKN	A	305	30.75	-	4.32E+00	4.40E+00
E10	UNKN	A	305	30.84	-	4.09E+00	4.40E+00
F8	UNKN	A	306	30.59	-	4.76E+00	4.89E+00
F9	UNKN	A	306	30.46	-	5.14E+00	4.89E+00
F10	UNKN	A	306	30.59	-	4.76E+00	4.89E+00
B11	STD	A		29.33	1.00E+01	1.02E+01	-
C11	STD	A		29.36	1.00E+01	1.00E+01	-
D11	STD	A		29.44	1.00E+01	9.53E+00	-
E11	STD	A		31.59	2.50E+00	2.60E+00	-
F11	STD	A		31.8	2.50E+00	2.29E+00	-
G11	STD	A		31.75	2.50E+00	2.36E+00	-
B12	STD	A		33.59	6.25E-01	7.76E-01	-
C12	STD	A		34.07	6.25E-01	5.81E-01	-
D12	STD	A		33.71	6.25E-01	7.22E-01	-
F12	STD	A		36.38	1.56E-01	1.44E-01	-
H11	STD	A		36.37	1.56E-01	1.45E-01	-

Figure 27C Plate 1-3 : TIMP-1

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	29.74	-	5.21E+00	5.14E+00
A3	UNKN	A	101	29.77	-	5.12E+00	5.14E+00
A4	UNKN	A	101	29.78	-	5.09E+00	5.14E+00
B2	UNKN	A	102	29.14	-	7.52E+00	7.34E+00
B3	UNKN	A	102	29.25	-	7.03E+00	7.34E+00
B4	UNKN	A	102	29.15	-	7.48E+00	7.34E+00
C2	UNKN	A	103	30.88	-	2.60E+00	2.65E+00
C3	UNKN	A	103	30.78	-	2.76E+00	2.65E+00
C4	UNKN	A	103	30.89	-	2.58E+00	2.65E+00
D2	UNKN	A	104	30.58	-	3.12E+00	2.73E+00
D3	UNKN	A	104	30.8	-	2.73E+00	2.73E+00
D4	UNKN	A	104	31.05	-	2.34E+00	2.73E+00
E2	UNKN	A	105	30.56	-	3.16E+00	3.38E+00
E3	UNKN	A	105	30.52	-	3.24E+00	3.38E+00
E4	UNKN	A	105	30.28	-	3.75E+00	3.38E+00
F2	UNKN	A	106	29.36	-	6.57E+00	5.82E+00
F3	UNKN	A	106	29.7	-	5.34E+00	5.82E+00
F4	UNKN	A	106	29.64	-	5.54E+00	5.82E+00
A5	UNKN	A	201	30.43	-	3.42E+00	3.66E+00
A6	UNKN	A	201	30.17	-	4.01E+00	3.66E+00
A7	UNKN	A	201	30.36	-	3.57E+00	3.66E+00
B5	UNKN	A	202	30.38	-	3.52E+00	3.14E+00
B6	UNKN	A	202	30.88	-	2.60E+00	3.14E+00
B7	UNKN	A	202	30.49	-	3.30E+00	3.14E+00
C5	UNKN	A	203	31.4	-	1.89E+00	1.75E+00
C6	UNKN	A	203	31.63	-	1.64E+00	1.75E+00
C7	UNKN	A	203	31.54	-	1.73E+00	1.75E+00
D5	UNKN	A	204	31.22	-	2.11E+00	2.12E+00
D6	UNKN	A	204	31.35	-	1.95E+00	2.12E+00
D7	UNKN	A	204	31.08	-	2.30E+00	2.12E+00
E5	UNKN	A	205	29.75	-	5.15E+00	5.44E+00
E6	UNKN	A	205	29.66	-	5.47E+00	5.44E+00
E7	UNKN	A	205	29.59	-	5.71E+00	5.44E+00
F5	UNKN	A	206	29.65	-	5.51E+00	5.40E+00
F6	UNKN	A	206	29.68	-	5.41E+00	5.40E+00
F7	UNKN	A	206	29.72	-	5.28E+00	5.40E+00
A8	UNKN	A	301	30.34	-	3.61E+00	3.89E+00
A9	UNKN	A	301	30.05	-	4.31E+00	3.89E+00
A10	UNKN	A	301	30.28	-	3.75E+00	3.89E+00
B8	UNKN	A	302	30.81	-	2.71E+00	2.62E+00
B9	UNKN	A	302	30.85	-	2.64E+00	2.62E+00
B10	UNKN	A	302	30.94	-	2.50E+00	2.62E+00
C8	UNKN	A	303	29.47	-	6.15E+00	5.89E+00
C9	UNKN	A	303	29.6	-	5.68E+00	5.89E+00
C10	UNKN	A	303	29.55	-	5.85E+00	5.89E+00
D8	UNKN	A	304	30.05	-	4.31E+00	4.14E+00
D9	UNKN	A	304	30.32	-	3.66E+00	4.14E+00
D10	UNKN	A	304	30	-	4.45E+00	4.14E+00
E8	UNKN	A	305	30.86	-	2.63E+00	3.18E+00
E9	UNKN	A	305	30.47	-	3.34E+00	3.18E+00
E10	UNKN	A	305	30.36	-	3.57E+00	3.18E+00
F8	UNKN	A	306	30.31	-	3.68E+00	3.60E+00
F9	UNKN	A	306	30.51	-	3.25E+00	3.60E+00
F10	UNKN	A	306	30.23	-	3.86E+00	3.60E+00
B11	STD	A		28.72	1.00E+01	9.72E+00	-
C11	STD	A		28.81	1.00E+01	9.20E+00	-
D11	STD	A		28.8	1.00E+01	9.26E+00	-
E11	STD	A		31.05	2.50E+00	2.34E+00	-
F11	STD	A		30.82	2.50E+00	2.69E+00	-
G11	STD	A		30.75	2.50E+00	2.81E+00	-
B12	STD	A		33.04	6.25E-01	6.93E-01	-
C12	STD	A		32.89	6.25E-01	7.60E-01	-
F12	STD	A		36.03	1.56E-01	1.11E-01	-
H11	STD	A		35.34	1.56E-01	1.70E-01	-

Figure 27D Plate 1-4 : Collagen Type 1

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	27.62	-	5.75E+00	5.44E+00
A3	UNKN	A	101	27.83	-	5.07E+00	5.44E+00
A4	UNKN	A	101	27.7	-	5.48E+00	5.44E+00
B2	UNKN	A	102	27.15	-	7.64E+00	7.59E+00
B3	UNKN	A	102	27.25	-	7.19E+00	7.59E+00
B4	UNKN	A	102	27.09	-	7.92E+00	7.59E+00
C2	UNKN	A	103	28.18	-	4.10E+00	4.08E+00
C3	UNKN	A	103	28.24	-	3.96E+00	4.08E+00
C4	UNKN	A	103	28.15	-	4.18E+00	4.08E+00
D2	UNKN	A	104	28.48	-	3.42E+00	2.84E+00
D3	UNKN	A	104	29	-	2.50E+00	2.84E+00
D4	UNKN	A	104	28.94	-	2.59E+00	2.84E+00
E2	UNKN	A	105	28	-	4.58E+00	4.24E+00
E3	UNKN	A	105	28.34	-	3.73E+00	4.24E+00
E4	UNKN	A	105	28.06	-	4.41E+00	4.24E+00
F2	UNKN	A	106	28.28	-	3.86E+00	3.95E+00
F3	UNKN	A	106	28.24	-	3.96E+00	3.95E+00
F4	UNKN	A	106	28.21	-	4.03E+00	3.95E+00
A5	UNKN	A	201	28.54	-	3.30E+00	3.23E+00
A6	UNKN	A	201	28.45	-	3.49E+00	3.23E+00
A7	UNKN	A	201	28.75	-	2.91E+00	3.23E+00
B5	UNKN	A	202	28.59	-	3.21E+00	3.10E+00
B6	UNKN	A	202	28.51	-	3.36E+00	3.10E+00
B7	UNKN	A	202	28.85	-	2.74E+00	3.10E+00
C5	UNKN	A	203	28.87	-	2.71E+00	2.65E+00
C6	UNKN	A	203	28.78	-	2.86E+00	2.65E+00
C7	UNKN	A	203	29.07	-	2.40E+00	2.65E+00
D5	UNKN	A	204	28.99	-	2.52E+00	2.46E+00
D6	UNKN	A	204	28.9	-	2.66E+00	2.46E+00
D7	UNKN	A	204	29.21	-	2.20E+00	2.46E+00
E5	UNKN	A	205	28.29	-	3.84E+00	3.55E+00
E6	UNKN	A	205	28.42	-	3.55E+00	3.55E+00
E7	UNKN	A	205	28.56	-	3.26E+00	3.55E+00
F5	UNKN	A	206	28.37	-	3.66E+00	3.86E+00
F6	UNKN	A	206	28.06	-	4.41E+00	3.86E+00
F7	UNKN	A	206	28.44	-	3.51E+00	3.86E+00
A8	UNKN	A	301	28.41	-	3.57E+00	3.41E+00
A9	UNKN	A	301	28.64	-	3.11E+00	3.41E+00
A10	UNKN	A	301	28.42	-	3.55E+00	3.41E+00
B8	UNKN	A	302	28.41	-	3.57E+00	3.43E+00
B9	UNKN	A	302	28.46	-	3.47E+00	3.43E+00
B10	UNKN	A	302	28.57	-	3.24E+00	3.43E+00
C8	UNKN	A	303	28.33	-	3.75E+00	3.83E+00
C9	UNKN	A	303	28.25	-	3.93E+00	3.83E+00
C10	UNKN	A	303	28.31	-	3.79E+00	3.83E+00
D8	UNKN	A	304	28.13	-	4.23E+00	3.92E+00
D9	UNKN	A	304	28.54	-	3.30E+00	3.92E+00
D10	UNKN	A	304	28.13	-	4.23E+00	3.92E+00
E8	UNKN	A	305	28.4	-	3.59E+00	4.04E+00
E9	UNKN	A	305	28.17	-	4.13E+00	4.04E+00
E10	UNKN	A	305	28.07	-	4.39E+00	4.04E+00
F8	UNKN	A	306	28.47	-	3.45E+00	3.52E+00
F9	UNKN	A	306	28.44	-	3.51E+00	3.52E+00
F10	UNKN	A	306	28.4	-	3.59E+00	3.52E+00
B11	STD	A		26.55	1.00E+01	1.10E+01	-
C11	STD	A		26.75	1.00E+01	9.73E+00	-
D11	STD	A		26.87	1.00E+01	9.05E+01	-
E11	STD	A		29.08	2.50E+00	2.38E+00	-
F11	STD	A		29.14	2.50E+00	2.30E+00	-
G11	STD	A		28.88	2.50E+00	2.69E+00	-
B12	STD	A		31.01	6.25E-01	7.44E-01	-
C12	STD	A		31.26	6.25E-01	6.40E-01	-
D12	STD	A		31.31	6.25E-01	6.21E-01	-
E12	STD	A		33.4	1.56E-01	1.76E-01	-
F12	STD	A		33.54	1.56E-01	1.62E-01	-
G12	STD	A		34.05	1.56E-01	1.19E-01	-

Figure 27E Plate 2-1 : 36B4

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	19.42	-	2.63E+01	2.53E+01
A3	UNKN	A	101	19.47	-	2.54E+01	2.53E+01
A4	UNKN	A	101	19.54	-	2.41E+01	2.53E+01
B2	UNKN	A	102	19.3	-	2.86E+01	2.79E+01
B3	UNKN	A	102	19.32	-	2.82E+01	2.79E+01
B4	UNKN	A	102	19.38	-	2.70E+01	2.79E+01
C2	UNKN	A	103	19.33	-	2.80E+01	2.83E+01
C3	UNKN	A	103	19.37	-	2.72E+01	2.83E+01
C4	UNKN	A	103	19.24	-	2.98E+01	2.83E+01
D2	UNKN	A	104	19.26	-	2.94E+01	2.70E+01
D3	UNKN	A	104	19.49	-	2.50E+01	2.70E+01
D4	UNKN	A	104	19.41	-	2.65E+01	2.70E+01
E2	UNKN	A	105	19.17	-	3.13E+01	2.94E+01
E3	UNKN	A	105	19.3	-	2.86E+01	2.94E+01
E4	UNKN	A	105	19.32	-	2.82E+01	2.94E+01
F2	UNKN	A	106	19.03	-	3.46E+01	3.32E+01
F3	UNKN	A	106	19.14	-	3.20E+01	3.32E+01
F4	UNKN	A	106	19.1	-	3.29E+01	3.32E+01
A5	UNKN	A	201	19.21	-	3.05E+01	3.20E+01
A6	UNKN	A	201	19.03	-	3.46E+01	3.20E+01
A7	UNKN	A	201	19.19	-	3.09E+01	3.20E+01
B5	UNKN	A	202	19.11	-	3.27E+01	3.08E+01
B6	UNKN	A	202	19.14	-	3.20E+01	3.08E+01
B7	UNKN	A	202	19.35	-	2.76E+01	3.08E+01
C5	UNKN	A	203	19.55	-	2.40E+01	2.26E+01
C6	UNKN	A	203	19.53	-	2.43E+01	2.26E+01
C7	UNKN	A	203	19.84	-	1.95E+01	2.26E+01
D5	UNKN	A	204	19.45	-	2.57E+01	2.52E+01
D6	UNKN	A	204	19.38	-	2.70E+01	2.52E+01
D7	UNKN	A	204	19.62	-	2.28E+01	2.52E+01
E5	UNKN	A	205	19.23	-	3.00E+01	3.03E+01
E6	UNKN	A	205	19.15	-	3.15E+01	3.03E+01
E7	UNKN	A	205	19.28	-	2.90E+01	3.03E+01
F5	UNKN	A	206	19.37	-	2.72E+01	2.75E+01
F6	UNKN	A	206	19.36	-	2.74E+01	2.75E+01
F7	UNKN	A	206	19.34	-	2.78E+01	2.75E+01
A8	UNKN	A	301	19.14	-	3.20E+01	3.00E+01
A9	UNKN	A	301	19.23	-	3.00E+01	3.00E+01
A10	UNKN	A	301	19.33	-	2.80E+01	3.00E+01
B8	UNKN	A	302	19.46	-	2.55E+01	2.65E+01
B9	UNKN	A	302	19.36	-	2.74E+01	2.65E+01
B10	UNKN	A	302	19.41	-	2.65E+01	2.65E+01
C8	UNKN	A	303	19.19	-	3.09E+01	3.18E+01
C9	UNKN	A	303	19.1	-	3.29E+01	3.18E+01
C10	UNKN	A	303	19.16	-	3.16E+01	3.18E+01
D8	UNKN	A	304	19.27	-	2.92E+01	3.16E+01
D9	UNKN	A	304	19.07	-	3.36E+01	3.16E+01
D10	UNKN	A	304	19.15	-	3.18E+01	3.16E+01
E8	UNKN	A	305	19.37	-	2.72E+01	2.84E+01
E9	UNKN	A	305	19.22	-	3.03E+01	2.84E+01
E10	UNKN	A	305	19.35	-	2.76E+01	2.84E+01
F8	UNKN	A	306	19.46	-	2.55E+01	2.80E+01
F9	UNKN	A	306	19.33	-	2.80E+01	2.80E+01
F10	UNKN	A	306	19.21	-	3.05E+01	2.80E+01
B11	STD	A		18.7	4.00E+01	4.37E+01	-
C11	STD	A		18.83	4.00E+01	3.99E+01	-
D11	STD	A		18.86	4.00E+01	3.90E+01	-
E11	STD	A		20.9	1.00E+01	9.22E+00	-
F11	STD	A		20.81	1.00E+01	9.83E+00	-
G11	STD	A		20.81	1.00E+01	9.83E+00	-
B12	STD	A		22.69	2.50E+00	2.60E+00	-
C12	STD	A		22.71	2.50E+00	2.56E+00	-
D12	STD	A		22.8	2.50E+00	2.40E+00	-
E12	STD	A		24.77	6.25E-01	5.97E-01	-
F12	STD	A		24.68	6.25E-01	6.36E-01	-
G12	STD	A		24.63	6.25E-01	6.59E-01	-

Figure 27F Plate 2-2 : MCP-1

Well	Sample Type	Target Mark	Association Mark	Ct(SDM)	Init Qty	Qty(SDM)	Rel.Qty(SDM)
A2	UNKN	A	101	28.15	-	4.51E+01	4.43E+01
A3	UNKN	A	101	29.21	-	4.31E+01	4.43E+01
A4	UNKN	A	101	28.16	-	4.48E+01	4.43E+01
B2	UNKN	A	102	28.26	-	4.16E+01	4.12E+01
B3	UNKN	A	102	28.18	-	4.41E+01	4.12E+01
B4	UNKN	A	102	28.38	-	3.80E+01	4.12E+01
C2	UNKN	A	103	29.26	-	1.98E+01	2.19E+01
C3	UNKN	A	103	28.94	-	2.51E+01	2.19E+01
C4	UNKN	A	103	29.19	-	2.08E+01	2.19E+01
D2	UNKN	A	104	28.19	-	4.38E+01	4.12E+01
D3	UNKN	A	104	28.44	-	3.64E+01	4.12E+01
D4	UNKN	A	104	28.2	-	4.35E+01	4.12E+01
E2	UNKN	A	105	28.71	-	2.97E+01	2.84E+01
E3	UNKN	A	105	28.74	-	2.91E+01	2.84E+01
E4	UNKN	A	105	28.87	-	2.64E+01	2.84E+01
F2	UNKN	A	106	28.57	-	3.30E+01	3.25E+01
F3	UNKN	A	106	28.65	-	3.11E+01	3.25E+01
F4	UNKN	A	106	28.55	-	3.35E+01	3.25E+01
A5	UNKN	A	201	28.34	-	3.92E+01	3.62E+01
A6	UNKN	A	201	28.45	-	3.61E+01	3.62E+01
A7	UNKN	A	201	28.56	-	3.33E+01	3.62E+01
B5	UNKN	A	202	28.42	-	3.69E+01	3.60E+01
B6	UNKN	A	202	28.5	-	3.48E+01	3.60E+01
B7	UNKN	A	202	28.44	-	3.64E+01	3.60E+01
C5	UNKN	A	203	29.55	-	1.59E+01	1.61E+01
C6	UNKN	A	203	29.46	-	1.70E+01	1.61E+01
C7	UNKN	A	203	29.59	-	1.55E+01	1.61E+01
D5	UNKN	A	204	29.28	-	1.95E+01	1.93E+01
D6	UNKN	A	204	29.4	-	1.78E+01	1.93E+01
D7	UNKN	A	204	29.21	-	2.05E+01	1.93E+01
E5	UNKN	A	205	28.08	-	4.75E+01	5.32E+01
E6	UNKN	A	205	27.92	-	5.35E+01	5.32E+01
E7	UNKN	A	205	27.8	-	5.85E+01	5.32E+01
F5	UNKN	A	206	27.6	-	6.79E+01	6.45E+01
F6	UNKN	A	206	27.72	-	6.21E+01	6.45E+01
F7	UNKN	A	206	27.69	-	6.35E+01	6.45E+01
A8	UNKN	A	301	28.75	-	2.89E+01	3.00E+01
A9	UNKN	A	301	28.68	-	3.04E+01	3.00E+01
A10	UNKN	A	301	28.67	-	3.06E+01	3.00E+01
B8	UNKN	A	302	28.92	-	2.54E+01	2.84E+01
B9	UNKN	A	302	28.66	-	3.09E+01	2.84E+01
B10	UNKN	A	302	28.75	-	2.89E+01	2.84E+01
C8	UNKN	A	303	27.44	-	7.65E+01	8.22E+01
C9	UNKN	A	303	27.24	-	8.88E+01	8.22E+01
C10	UNKN	A	303	27.36	-	8.12E+01	8.22E+01
D8	UNKN	A	304	28.78	-	2.82E+01	2.82E+01
D9	UNKN	A	304	28.6	-	3.23E+01	2.82E+01
D10	UNKN	A	304	28.99	-	2.42E+01	2.82E+01
E8	UNKN	A	305	28.61	-	3.20E+01	2.70E+01
E9	UNKN	A	305	29	-	2.40E+01	2.70E+01
E10	UNKN	A	305	28.94	-	2.51E+01	2.70E+01
F8	UNKN	A	306	28.81	-	2.76E+01	2.73E+01
F9	UNKN	A	306	28.78	-	2.82E+01	2.73E+01
F10	UNKN	A	306	28.89	-	2.60E+01	2.73E+01
B11	STD	A		28.29	4.00E+01	4.07E+01	-
C11	STD	A		28.14	4.00E+01	4.55E+01	-
D11	STD	A		28.12	4.00E+01	4.61E+01	-
E11	STD	A		30.33	1.00E+01	8.91E+00	-
F11	STD	A		30.18	1.00E+01	9.96E+00	-
G11	STD	A		30.37	1.00E+01	8.65E+00	-
B12	STD	A		31.93	2.50E+00	2.71E+00	-
C12	STD	A		32.36	2.50E+00	1.97E+00	-
D12	STD	A		32.3	2.50E+00	2.06E+00	-
F12	STD	A		33.76	6.25E-01	6.95E-01	-
H11	STD	A		33.61	6.25E-01	7.77E-01	-

Figure 28

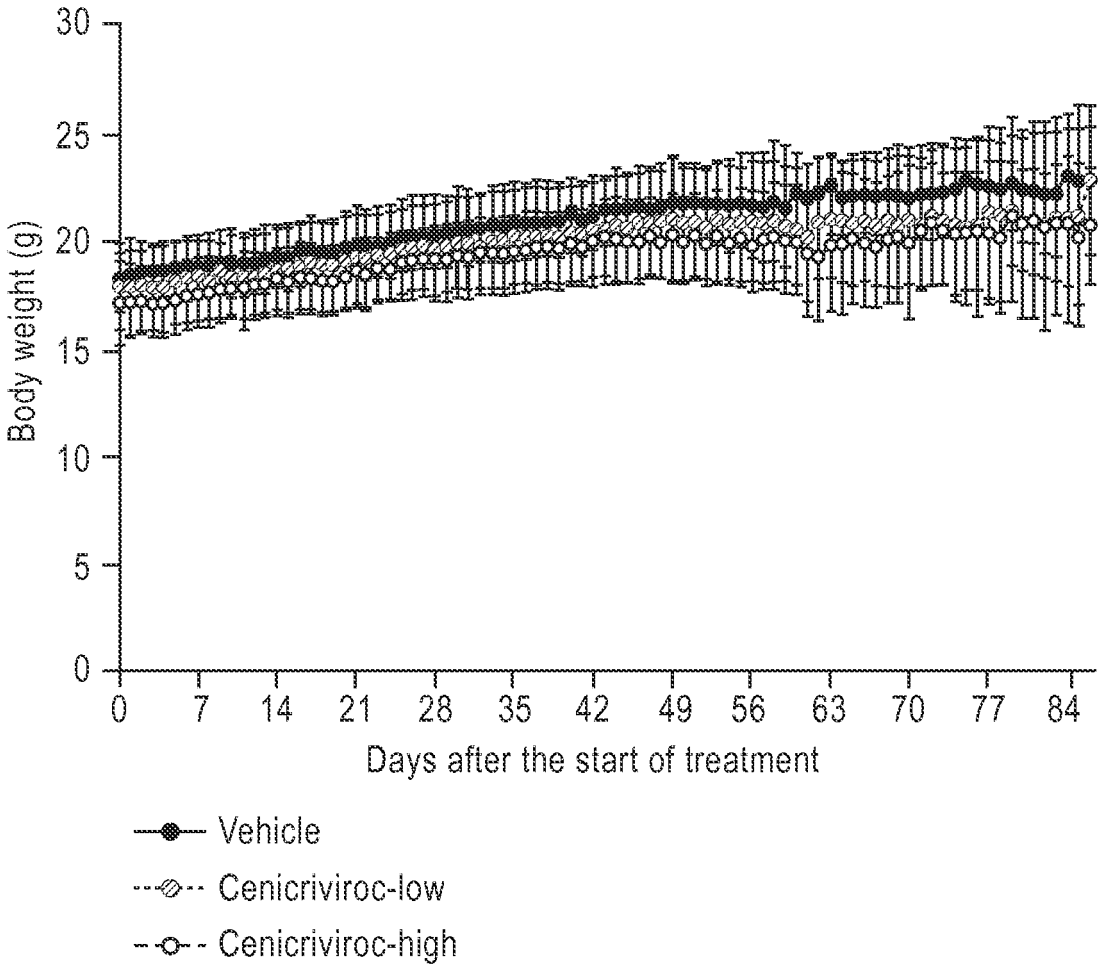

30
25
20
15
10
5
0
Body weight (g)
0
7
14
21
28
35
42
49
56
63
70
77
84
Days after the start of treatment
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 29

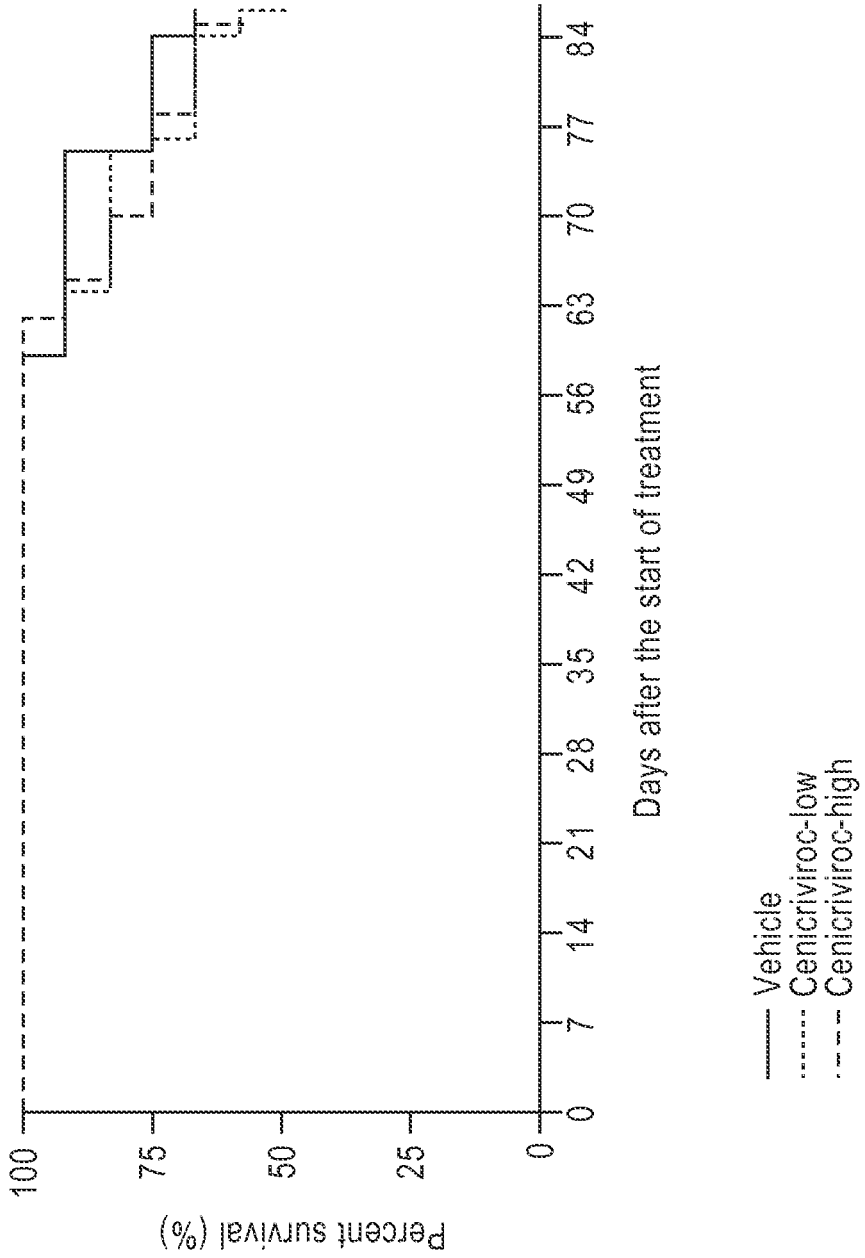

Percent survival (%)
100
75
50
25
0
0
7
14
21
28
35
42
49
56
63
70
77
84
Days after the start of treatment
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 30

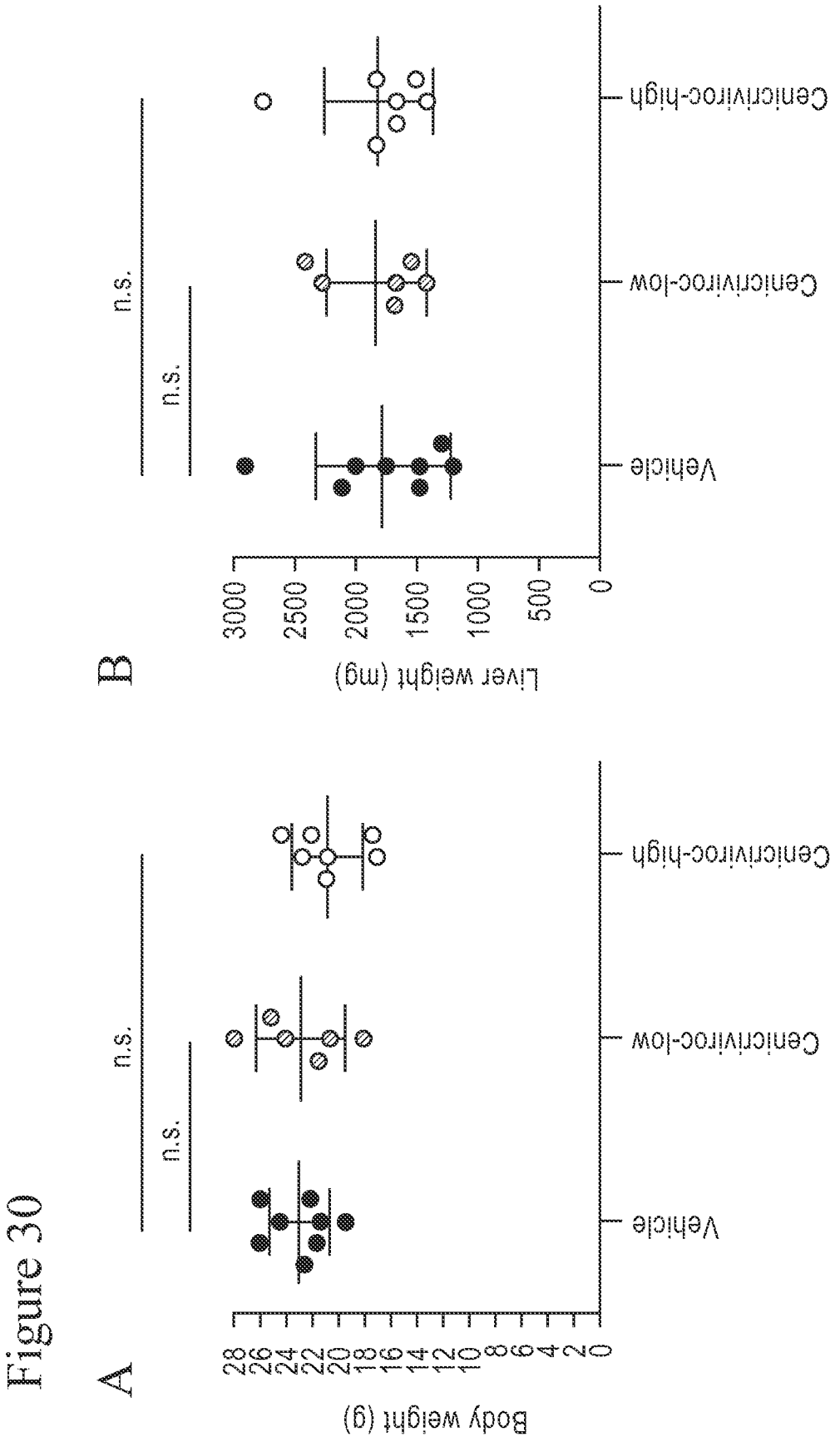

A
Body weight (g)
28
26
24
22
20
18
16
14
12
10
8
6
4
2
0
n.s.
n.s.
Vehicle
Cenicriviroc-low
Cenicriviroc-high
B
Liver weight (mg)
3000
2500
2000
1500
1000
500
0
n.s.
n.s.
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 30

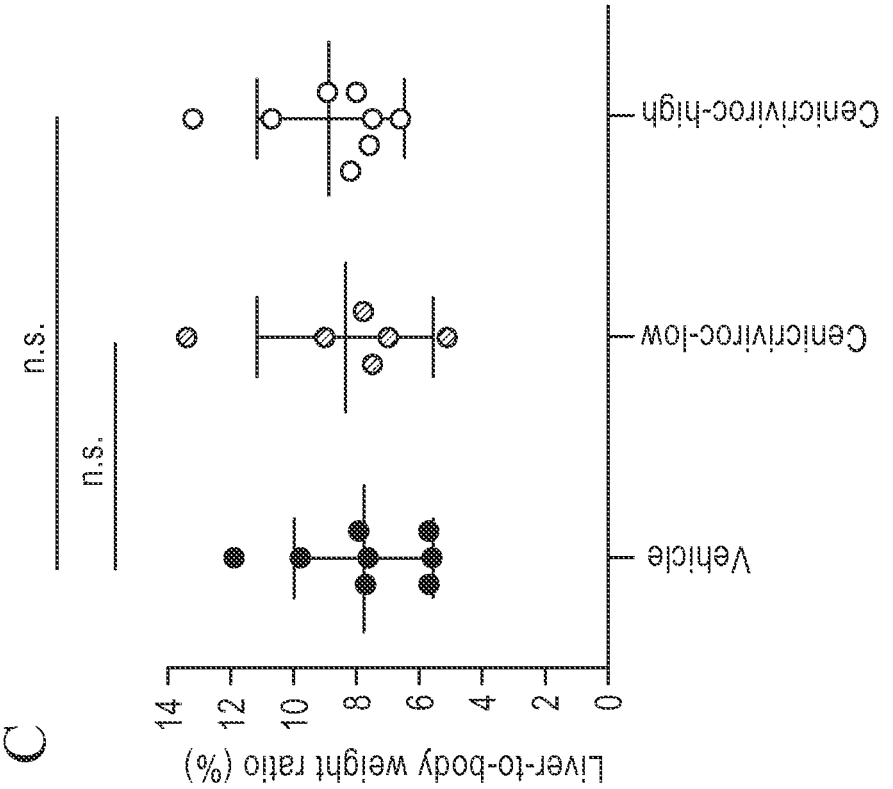

C
n.s.
n.s.
Liver-to-body weight ratio (%)
0
2
4
6
8
10
12
14
Vehicle
Cenicriviroc-low
Cenicriviroc-high

Figure 31A

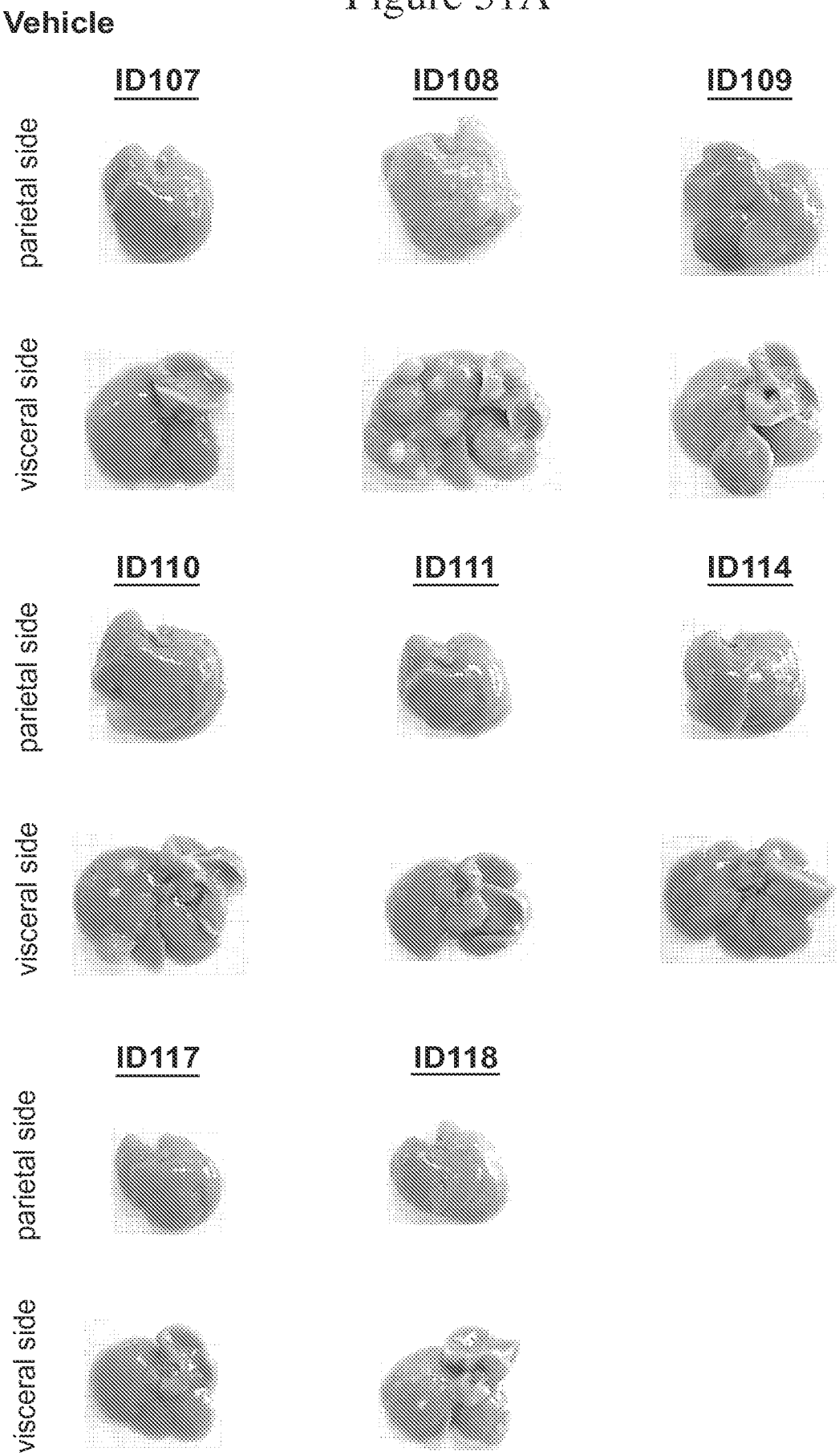

Vehicle
ID107
ID108
ID109
parietal side
visceral side
ID110
ID111
ID114
parietal side
visceral side
ID117
ID118
parietal side
visceral side

Figure 31B

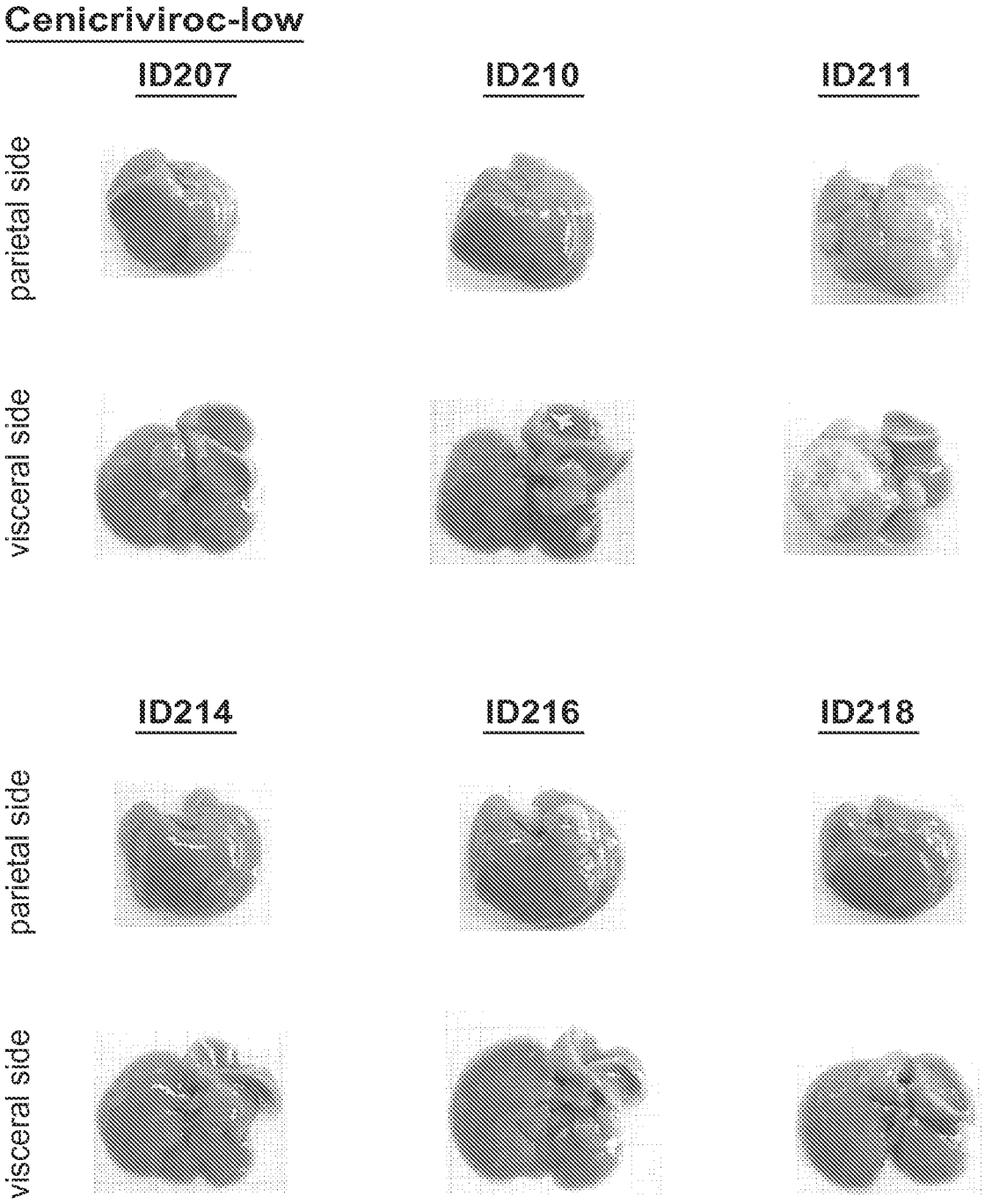
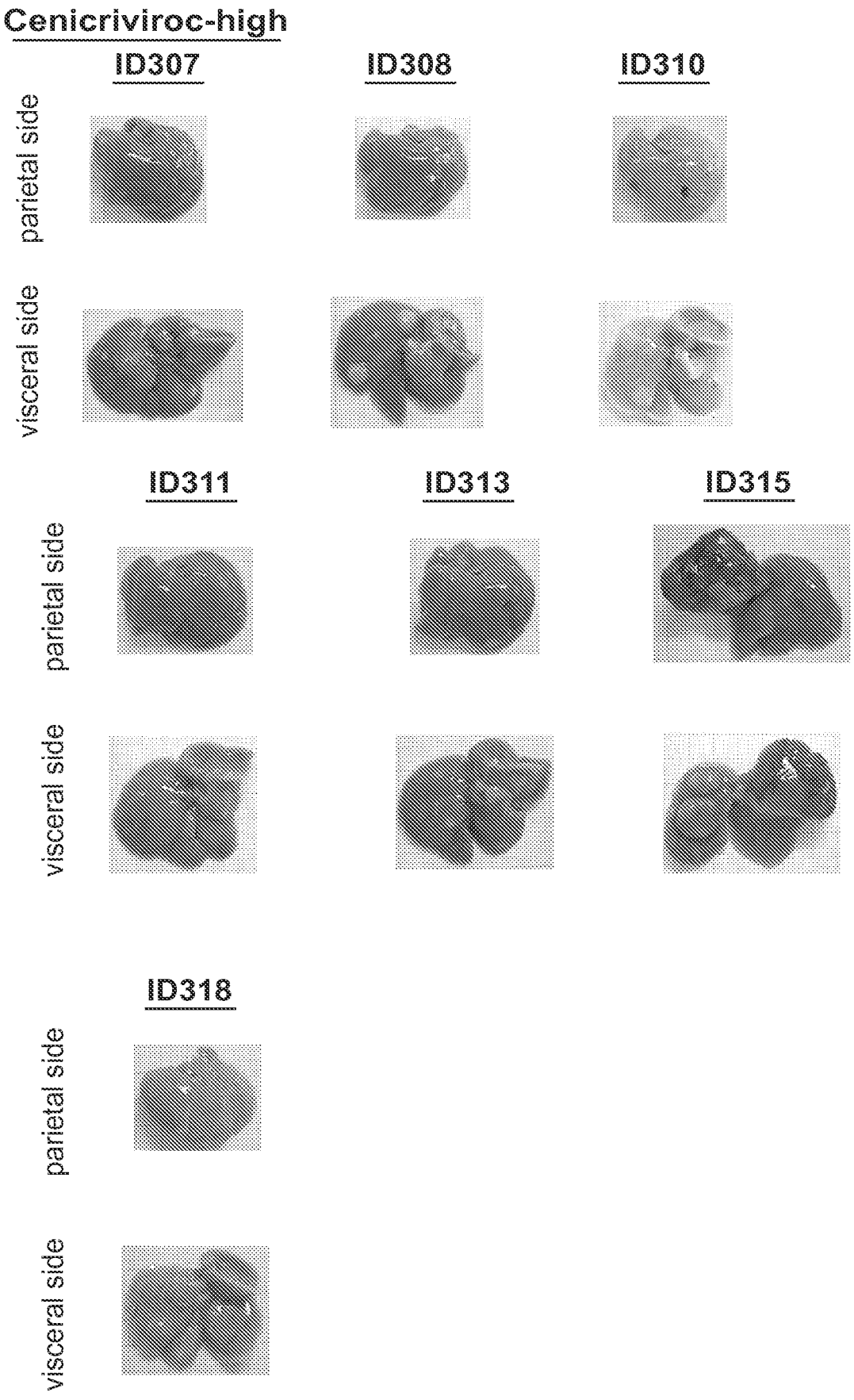

Cenicriviroc-low
ID207
ID210
ID211
parietal side
visceral side
ID214
ID216
ID218
parietal side
visceral side

Figure 31C


Cenicriviroc-high
ID307
ID308
ID310
parietal side
visceral side
ID311
ID313
ID315
parietal side
visceral side
ID318
parietal side
visceral side

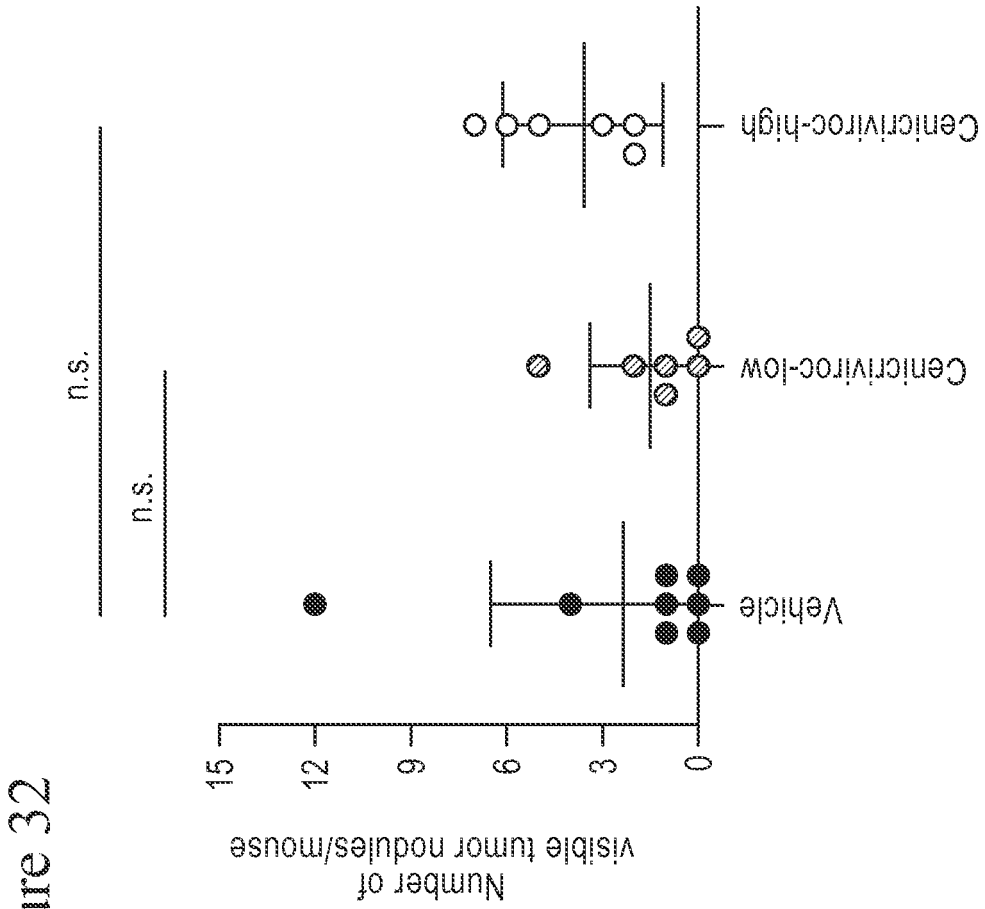

Number of
visible tumor nodules/mouse
15
12
9
6
3
0
Vehicle
Cenicriviroc-low
Cenicriviroc-high
n.s.
n.s.

Figure 32

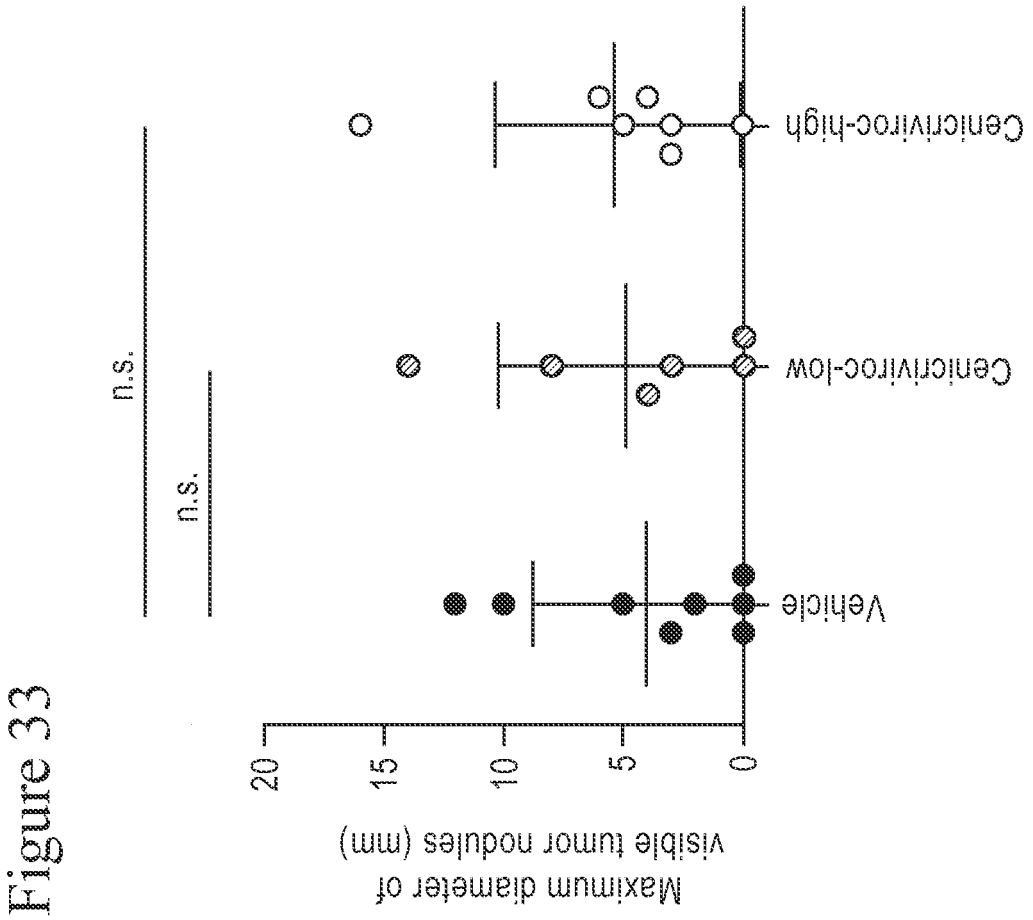

Maximum diameter of
visible tumor nodules (mm)
0
5
10
15
20
Vehicle
Cenicriviroc-low
Cenicriviroc-high
n.s.
n.s.

Figure 33

Figure 34

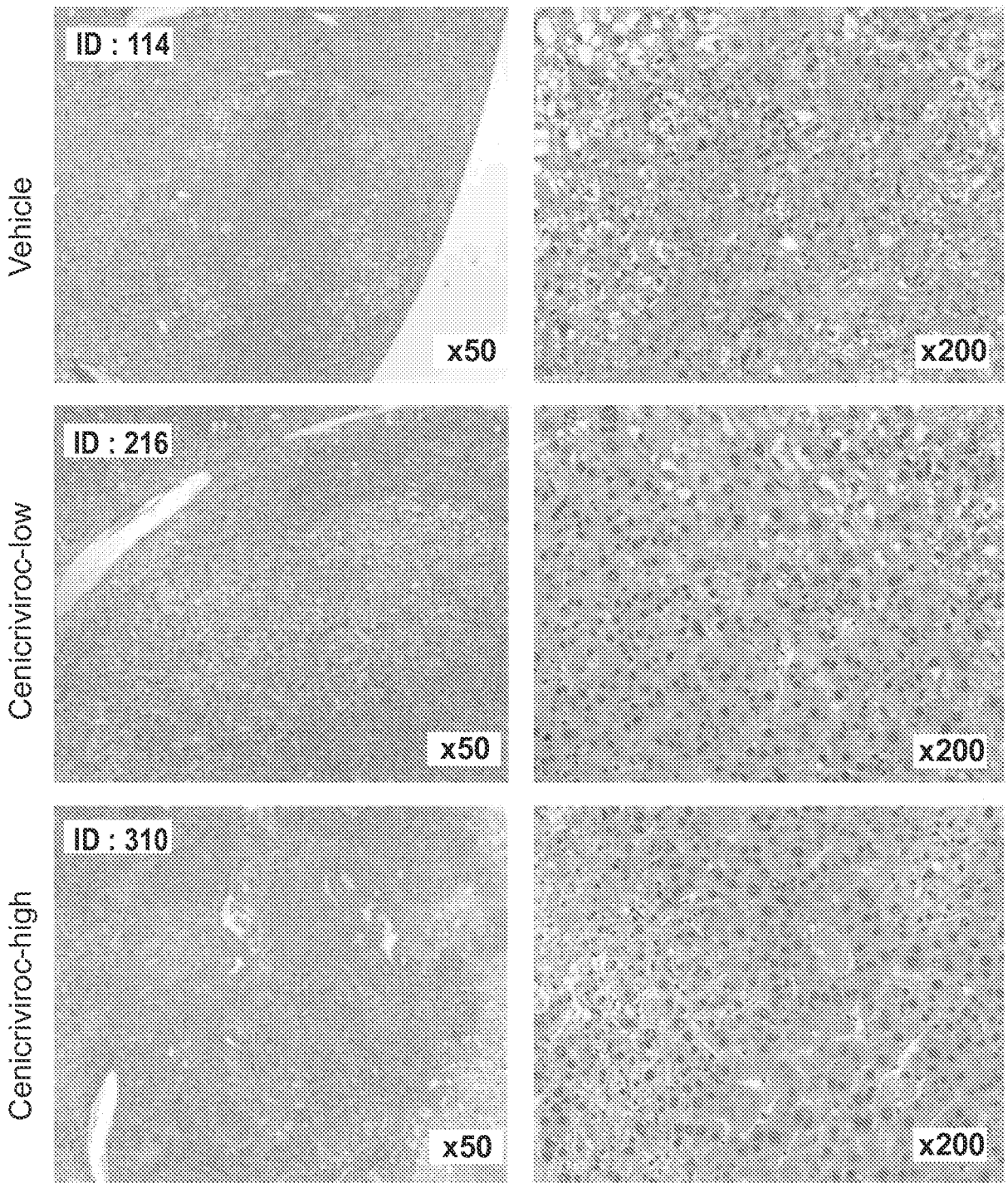

Vehicle
ID : 114
x50
x200
Cenicriviroc-low
ID : 216
x50
x200
Cenicriviroc-high
ID : 310
x50
x200

Figure 35

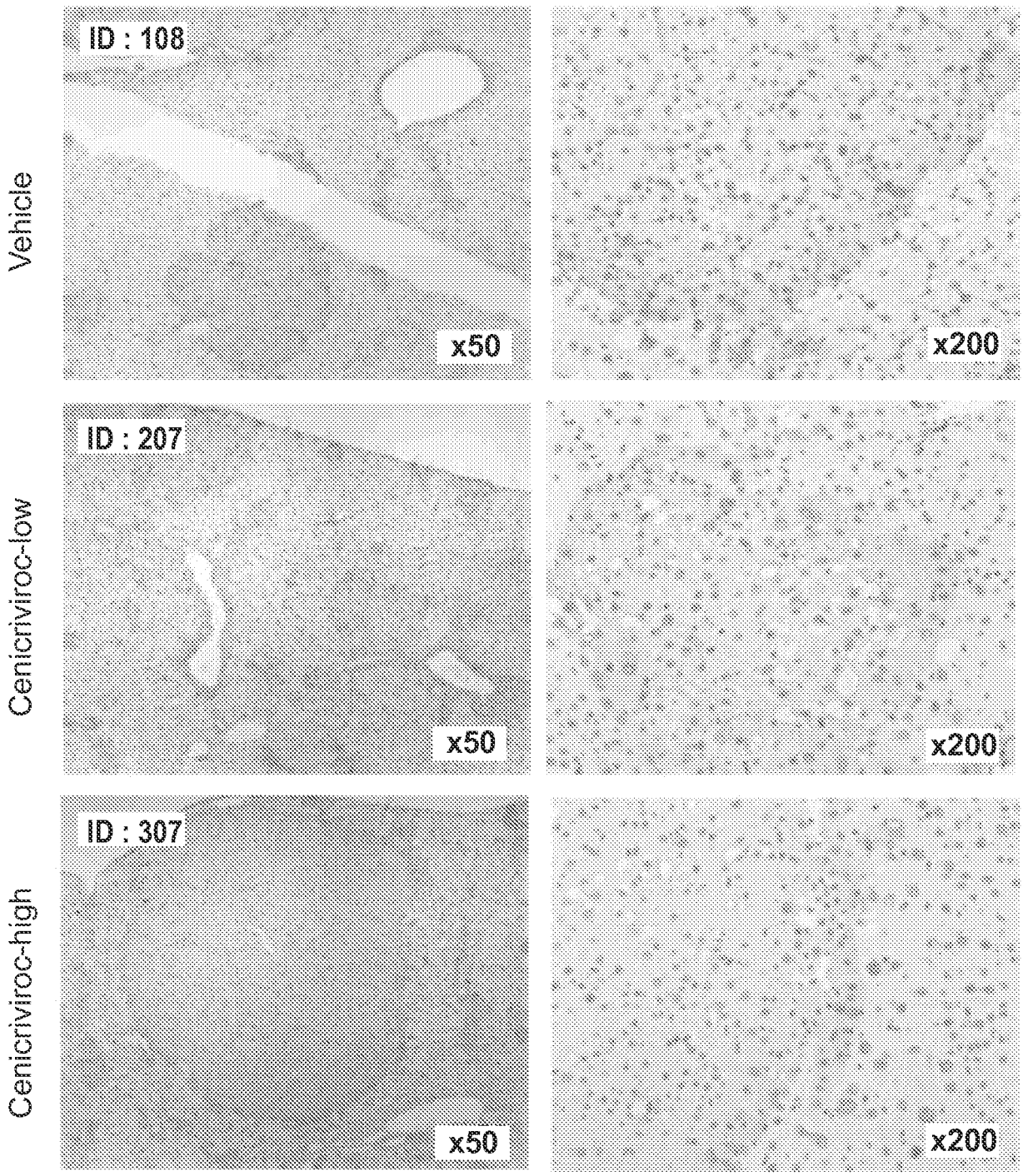
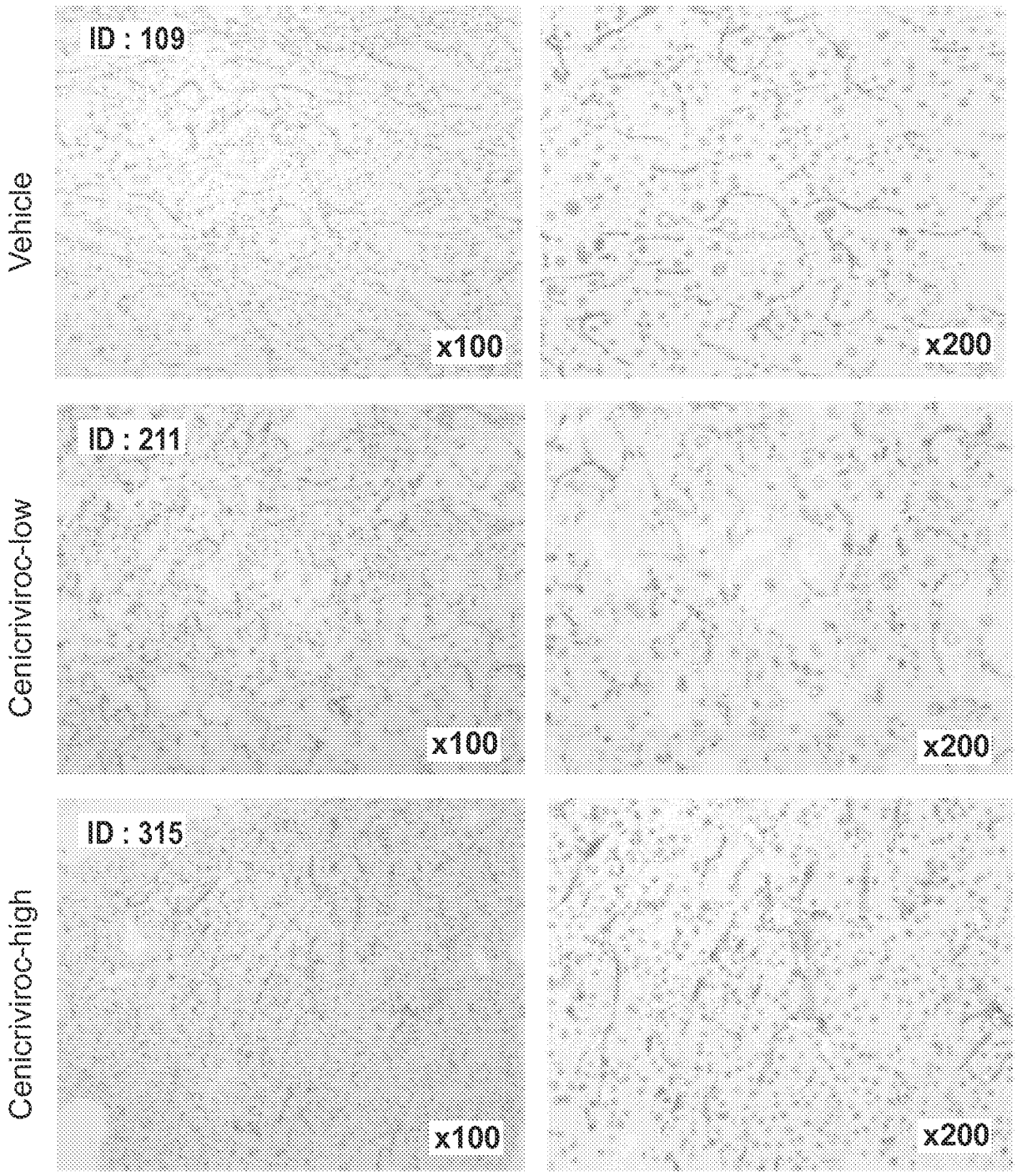

Vehicle
ID : 108
x50
x200
Cenicriviroc-low
ID : 207
x50
x200
Cenicriviroc-high
ID : 307
x50
x200

Figure 36


Vehicle
ID : 109
x100
x200
Cenicriviroc-low
ID : 211
x100
x200
Cenicriviroc-high
ID : 315
x100
x200

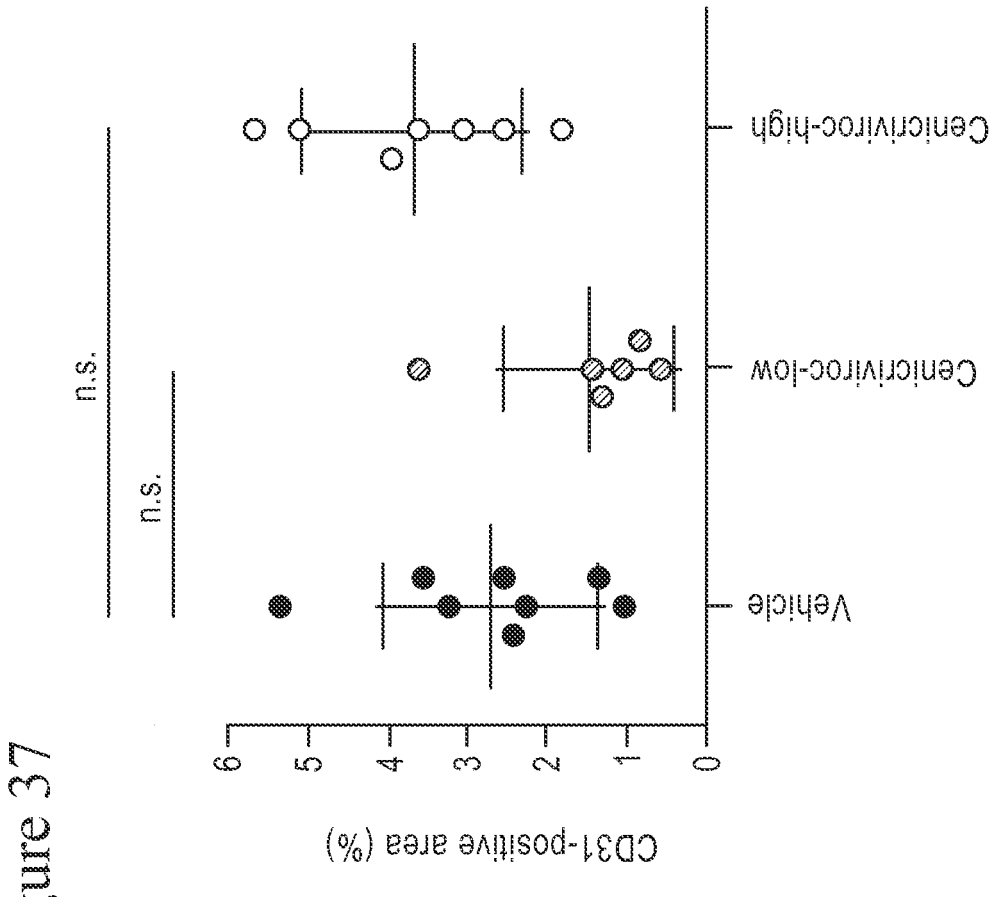
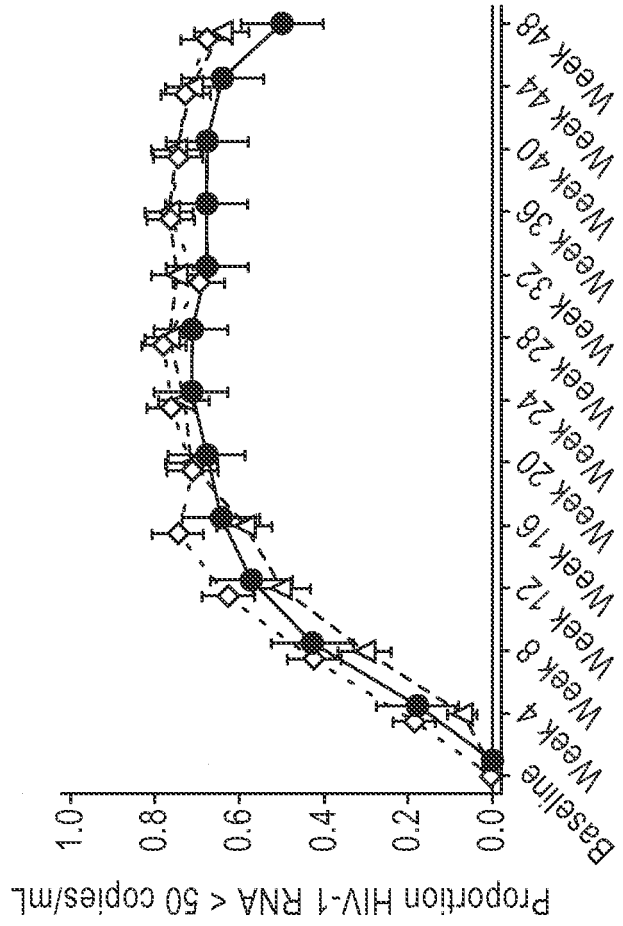

n.s.
n.s.
Vehicle
Cenicriviroc-low
Cenicriviroc-high
0
1
2
3
4
5
6
CD31-positive area (%)

Figure 37

Figure 38



No. of subjects													
CVC 100 + FTC/TDF	0/59	11/59	25/59	37/59	44/59	42/49	45/59	46/59	41/59	45/59	44/59	43/59	40/59
CVC 200 + FTC/TDF	0/56	4/56	17/56	28/56	33/56	40/56	41/56	43/56	42/56	43/56	42/56	40/56	36/56
EFV + FTC/TDF	0/28	5/28	12/28	16/28	18/28	19/28	20/28	20/28	19/28	19/28	19/28	18/28	14/28

Treatment group
- CVC 100 mg + EFV placebo + FTC/TDF
- CVC 200 mg + EFV placebo + FTC/TDF
- CVC placebo + EFV 600 mg + FTC/TDF

n/N: number of subjects with HIV-1 RNA <50 copies/mL/number of subjects with data.
Source: Week 48 final CSR 652-2-202 [22].

Figure 39

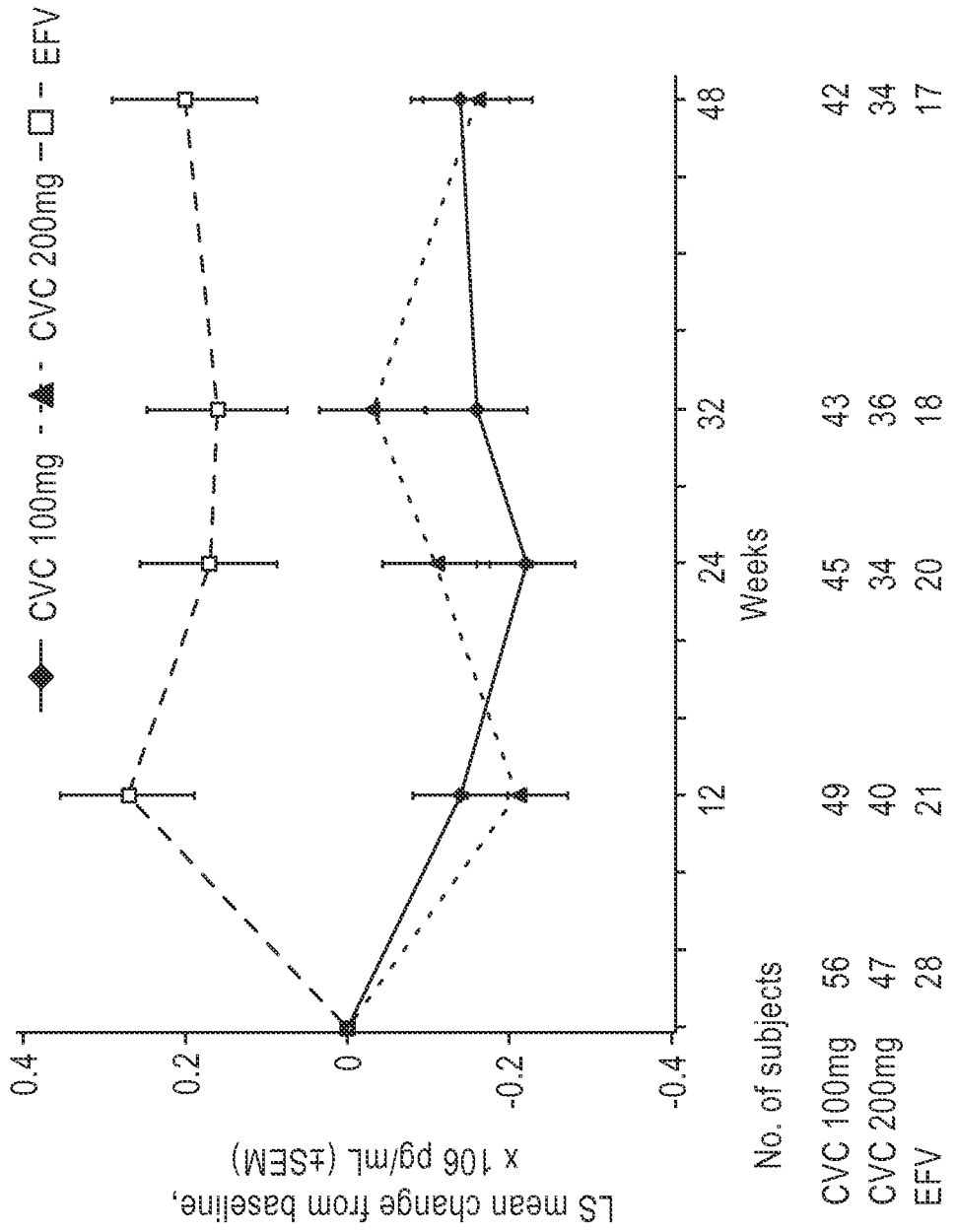
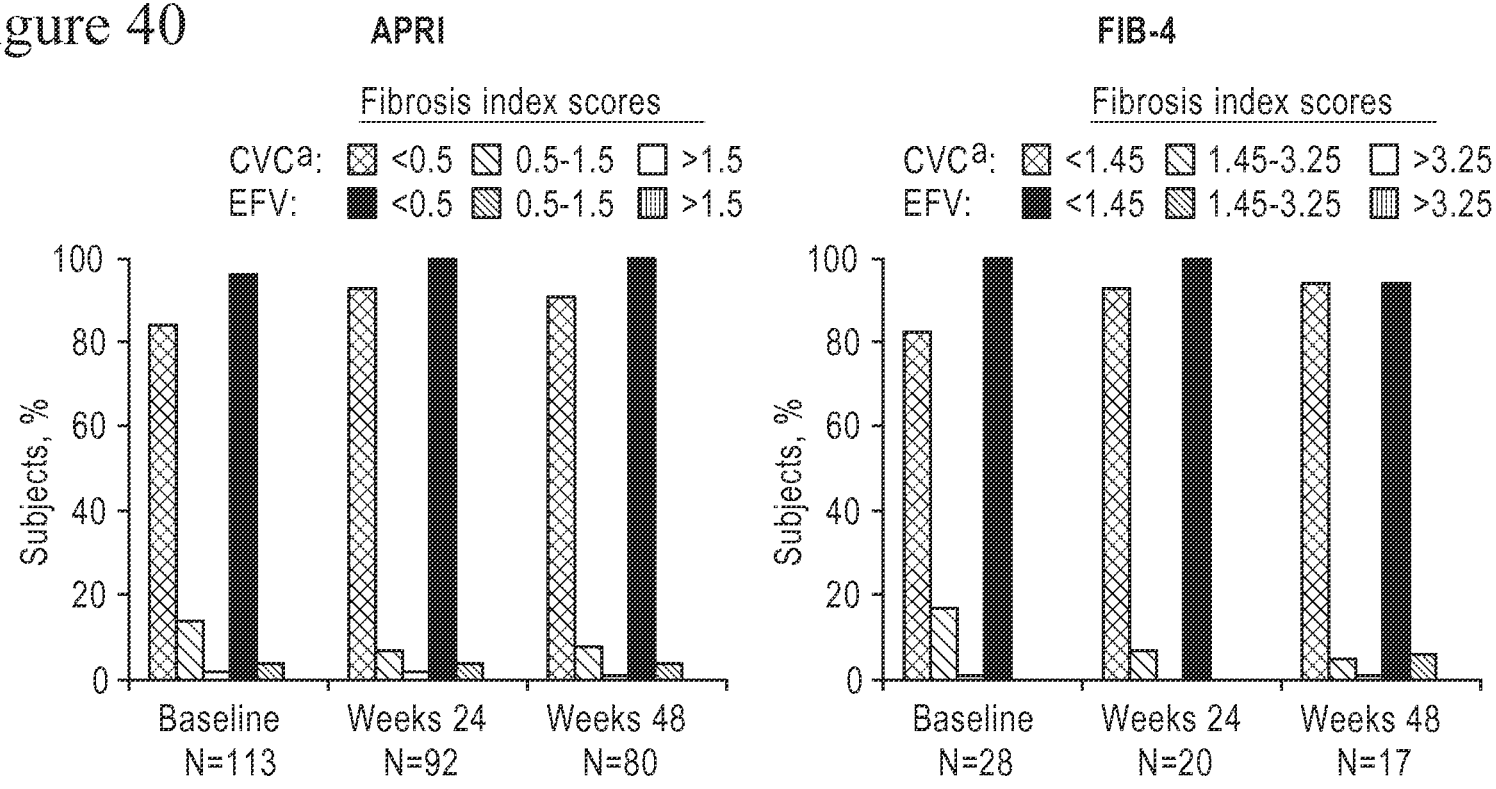

CVC 100mg
CVC 200mg
EFV
LS mean change from baseline, x 10⁶ pg/mL (±SEM)
0.4
0.2
0
-0.2
-0.4
12
24
32
48
Weeks
No. of subjects
CVC 100mg 56 49 45 43 42
CVC 200mg 47 40 34 36 34
EFV 28 21 20 18 17

Figure 40



	Decreased by one category from baseline		
	Baseline	Week 24	Week 48
CVC	N/A	14%	10%
EFV	N/A	5%	6%

	Decreased by one category from baseline		
	Baseline	Week 24	Week 48
CVC	N/A	13%	14%
EFV	N/A	-	-

Note: The proportion of subjects decreasing by one category is shown below the figures.

a CVC 100-mg and 200-mg arms pooled

Figure 41

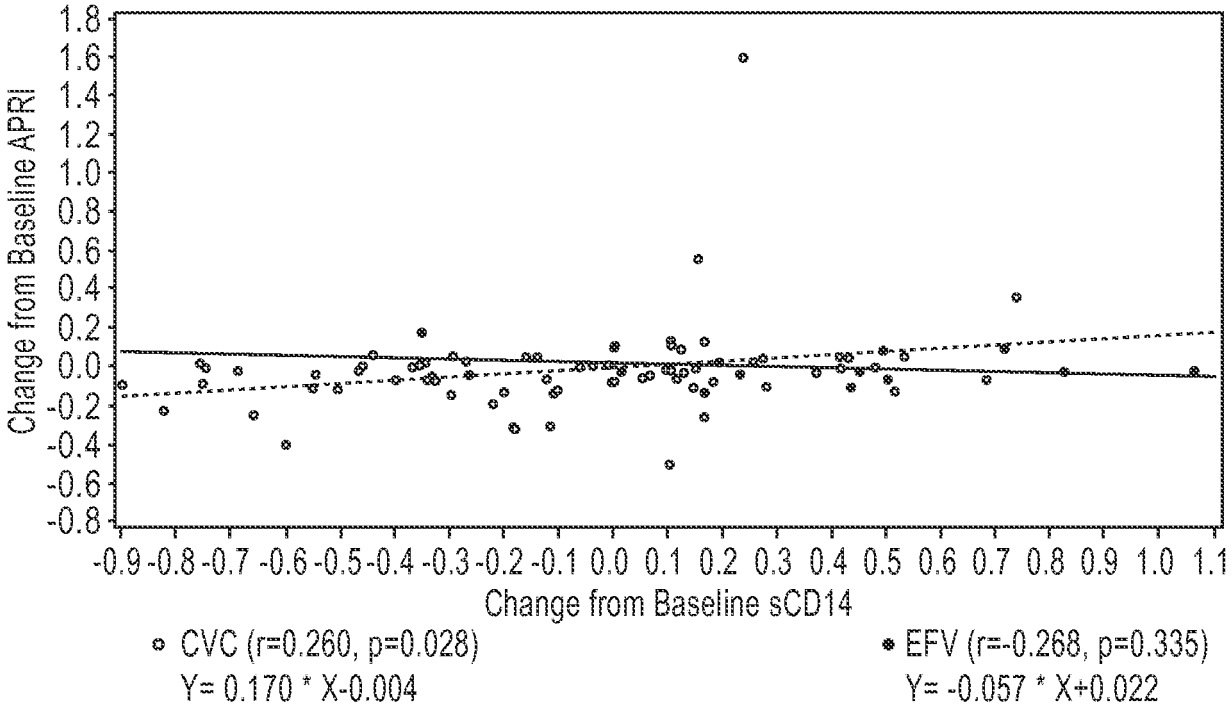


Figure 42

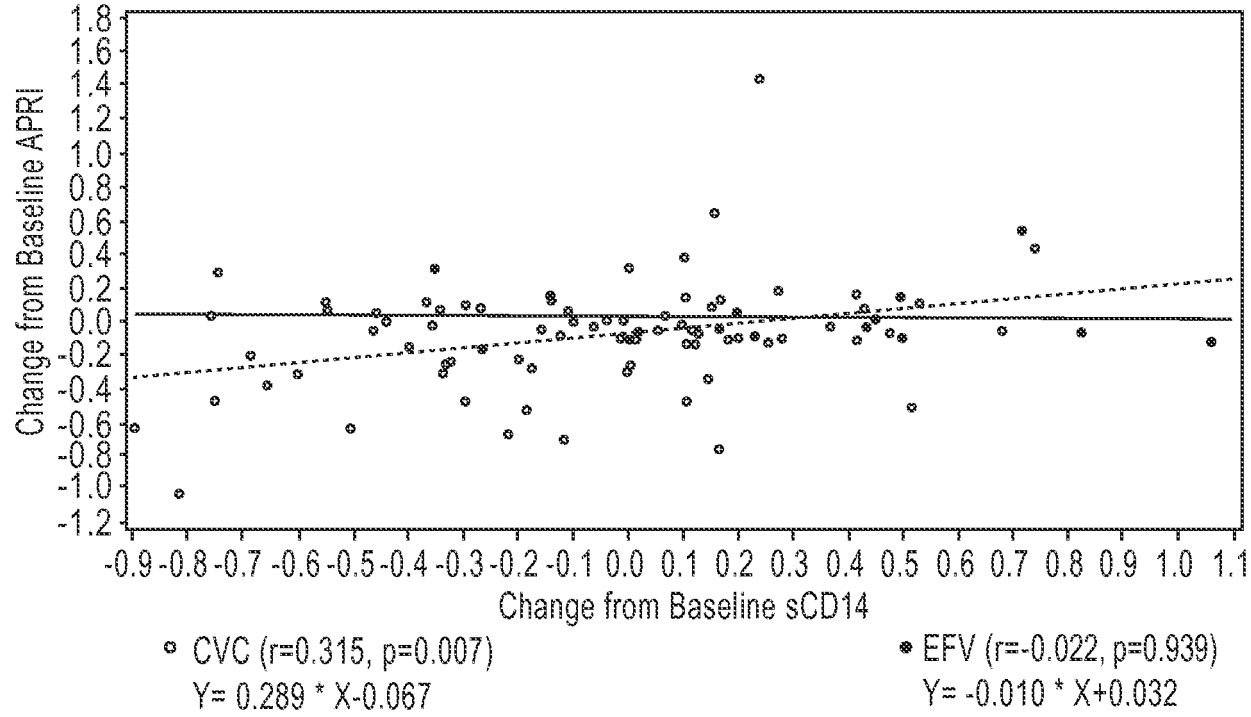
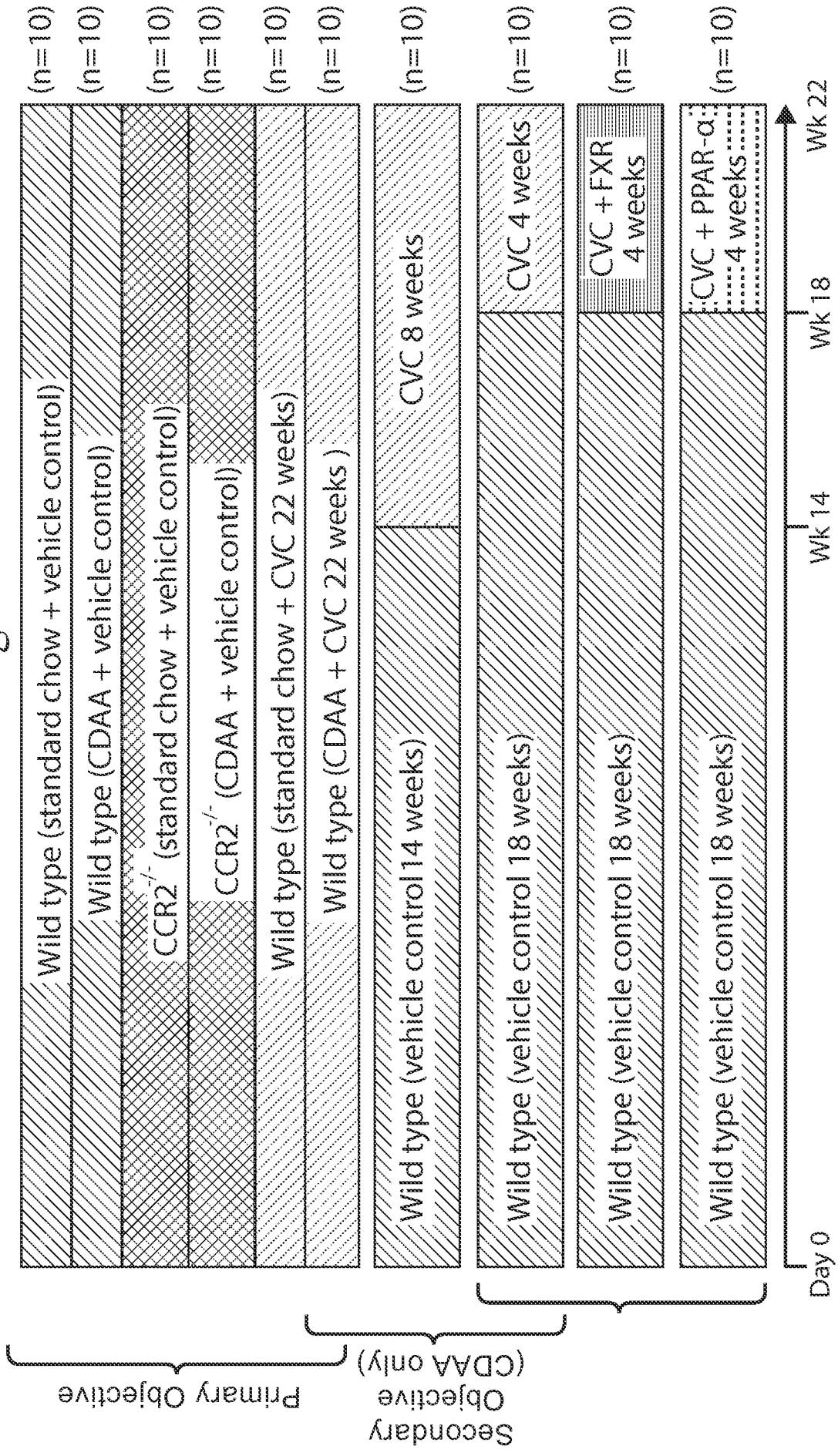


Figure 43


(n=10)
Wild type (standard chow + vehicle control)
(n=10)
Wild type (CDAA + vehicle control)
(n=10)
CCR2-/- (standard chow + vehicle control)
(n=10)
CCR2-/- (CDAA + vehicle control)
(n=10)
Wild type (standard chow + CVC 22 weeks)
(n=10)
Wild type (CDAA + CVC 22 weeks)
(n=10)
Wild type (vehicle control 14 weeks)
CVC 8 weeks
(n=10)
Wild type (vehicle control 18 weeks)
CVC 4 weeks
(n=10)
Wild type (vehicle control 18 weeks)
CVC + FXR 4 weeks
(n=10)
Wild type (vehicle control 18 weeks)
CVC + PPAR-α: 4 weeks
Day 0
Wk 14
Wk 18
Wk 22
Secondary Objective (CDAA only)
Primary Objective