FIG. 1

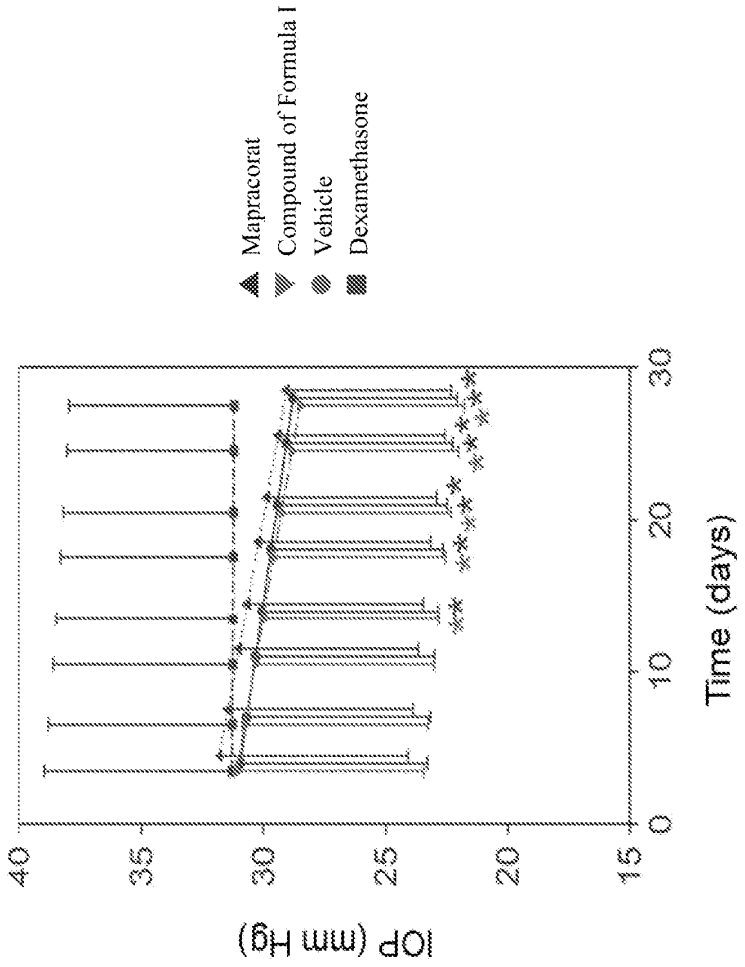

Mapracorat
Compound of Formula I
Vehicle
Dexamethasone
IOP (mm Hg)
40
35
30
25
20
15
0
10
20
30
Time (days)

FIG. 2

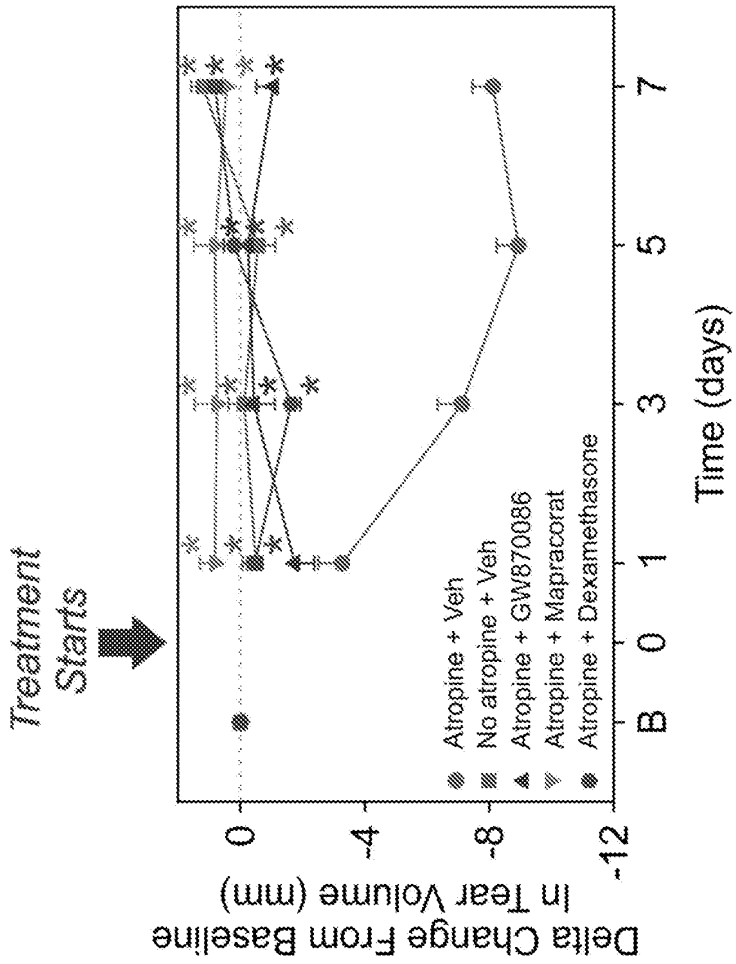

Treatment Starts
Delta Change From Baseline In Tear Volume (mm)
0
-4
-8
-12
B
0
1
3
5
7
Time (days)
Atropine + Veh
No atropine + Veh
Atropine + GW870086
Atropine + Mapracorat
Atropine + Dexamethasone

FIG. 3B

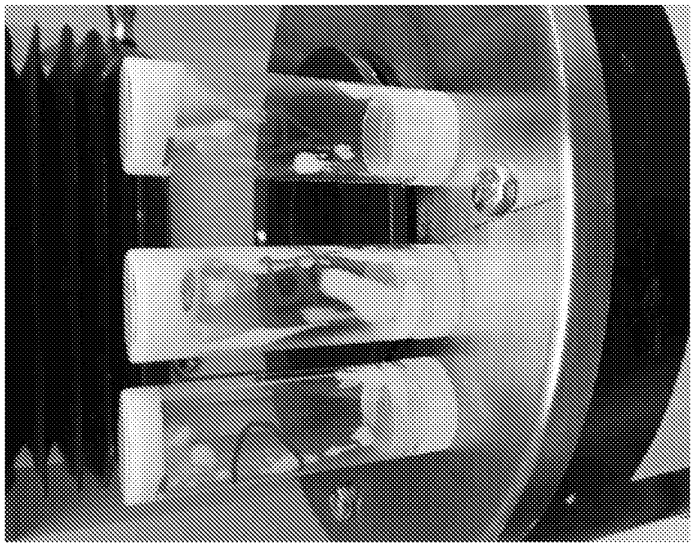


FIG. 3A

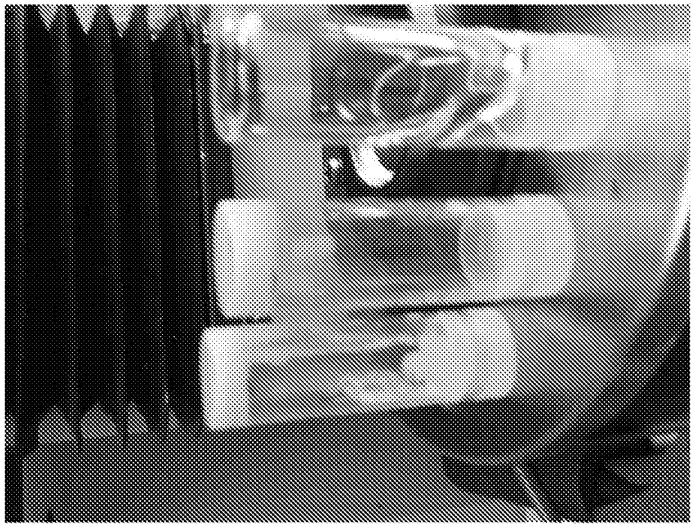


FIG. 4A

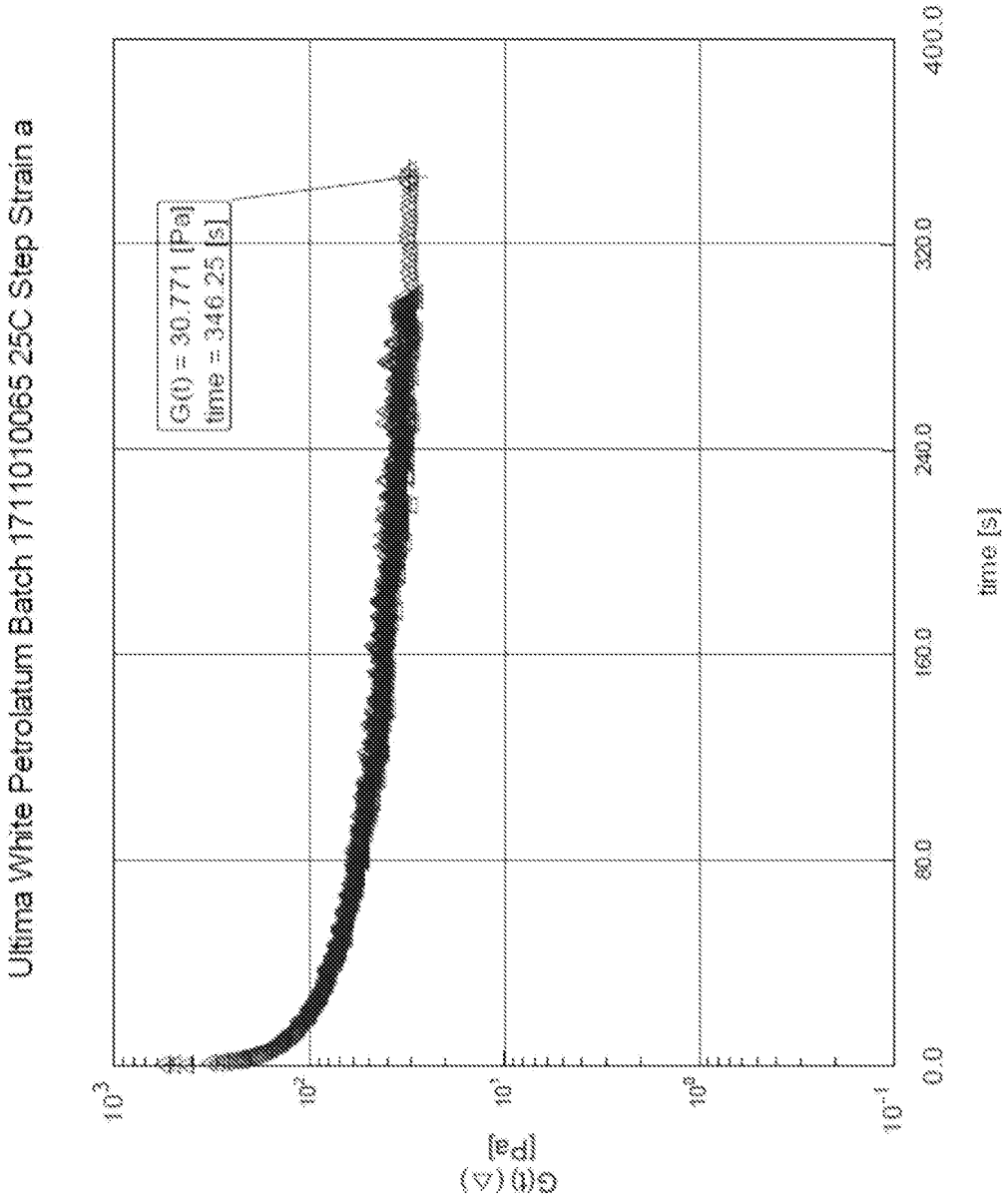

Ultima White Petrolatum Batch 171101065 25C Step Strain a
G(0) = 30.771 [Pa]
time = 346.25 [s]
G(t) (△) [Pa]
time [s]
0.0
80.0
160.0
240.0
320.0
400.0

FIG. 4B

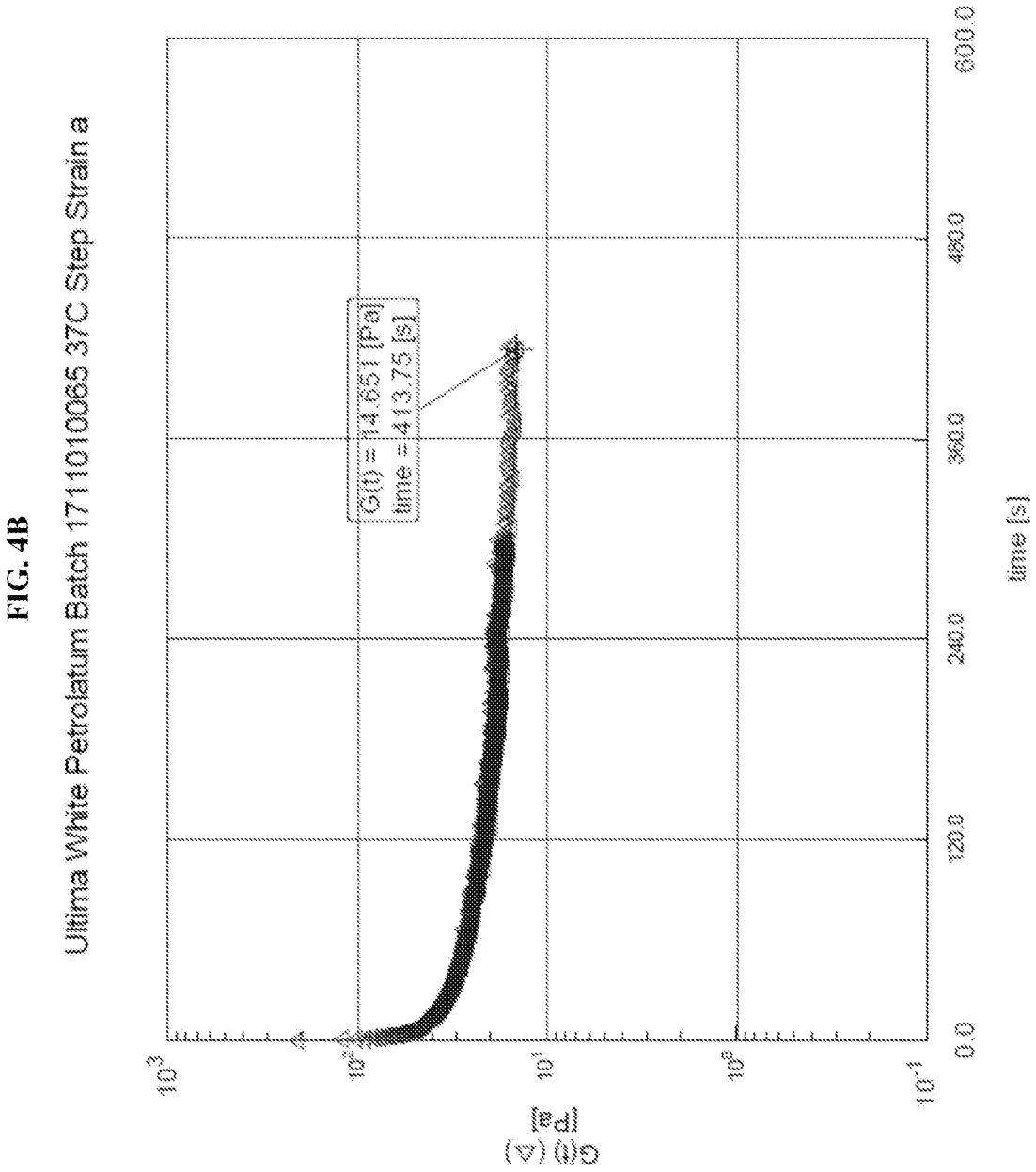

Ultima White Petrolatum Batch 1711010065 37C Step Strain a
G(t) = 14.651 [Pa]
time = 413.75 [s]
G(t) (Δ) [Pa]
time [s]
10^-3
10^-2
10^-1
10^0
10^1
0.0
120.0
240.0
360.0
480.0
600.0

FIG. 4C

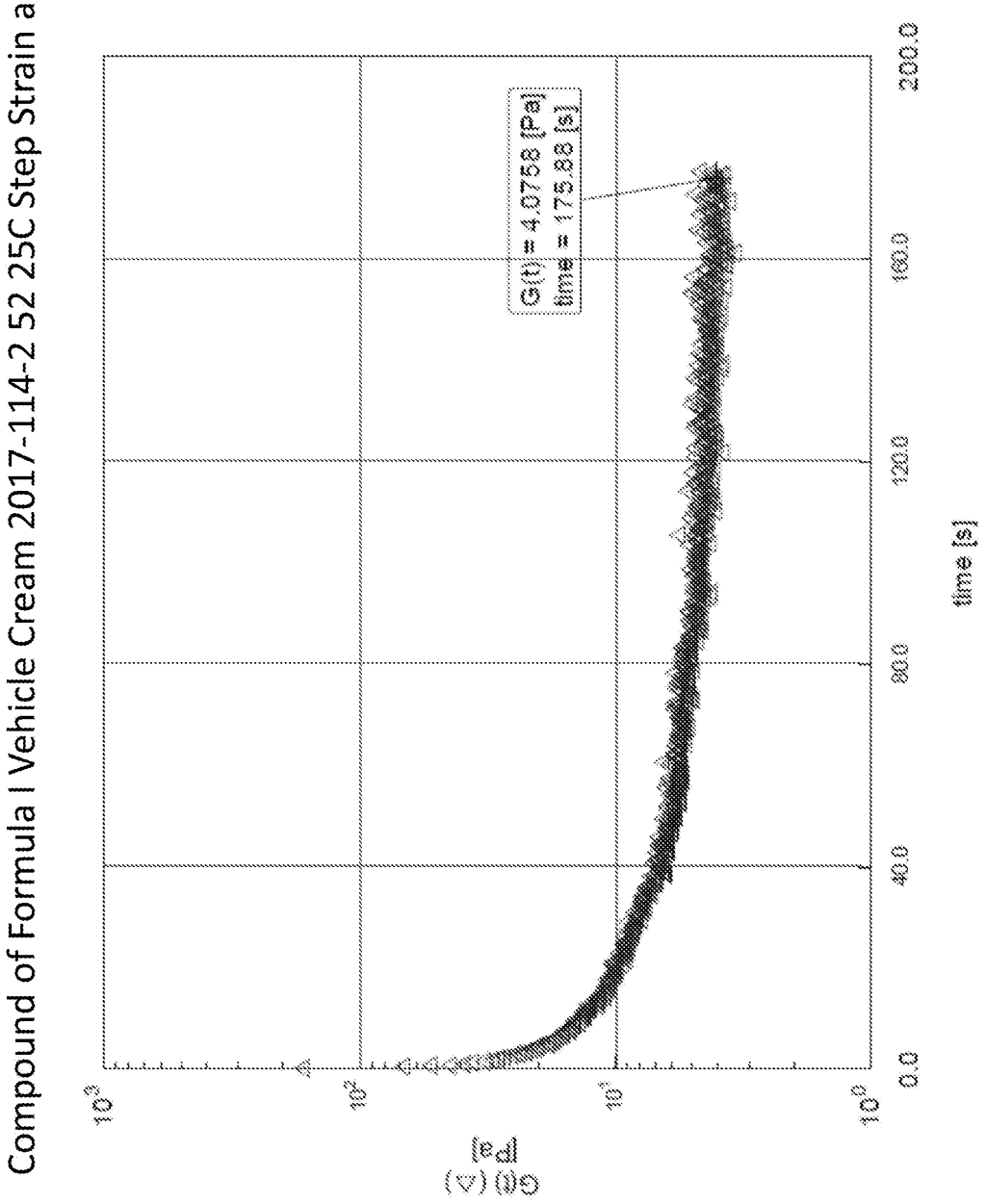

Compound of Formula I Vehicle Cream 2017-114-2 52 25C Step Strain a
G(t) = 4.0758 [Pa]
time = 175.88 [s]
G(t) (△) [Pa]
time [s]
10^3
10^2
10^1
10^0
0.0
40.0
80.0
120.0
160.0
200.0

FIG. 4D

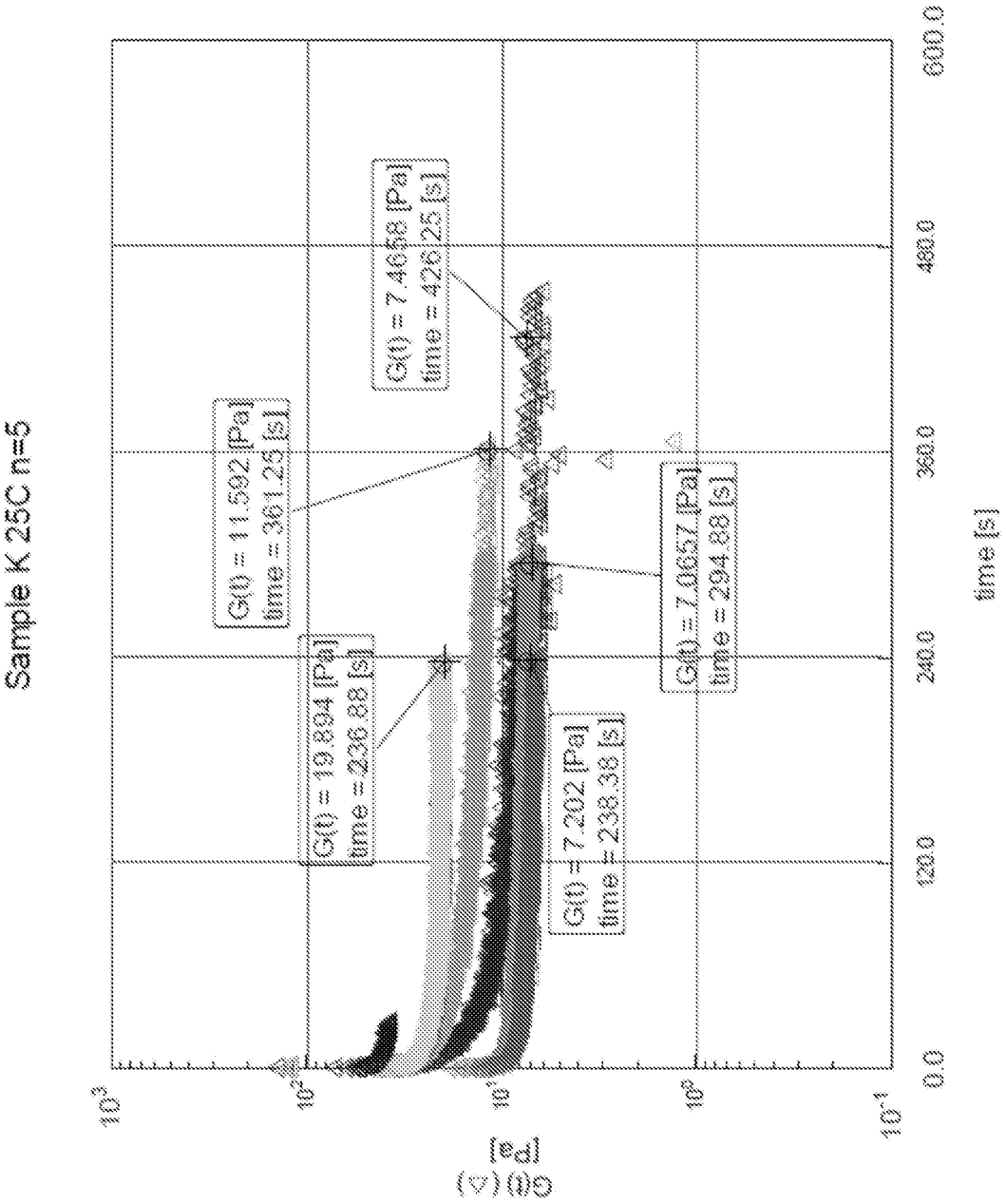

Sample K 25C n=5
G(t) = 19.894 [Pa]
time = 236.88 [s]
G(t) = 11.592 [Pa]
time = 361.23 [s]
G(t) = 7.4658 [Pa]
time = 426.25 [s]
G(t) = 7.202 [Pa]
time = 238.38 [s]
G(t) = 7.0657 [Pa]
time = 294.88 [s]
G(t) (△) (▽) [Pa]
time [s]
0.0
120.0
240.0
360.0
480.0
600.0

FIG. 5

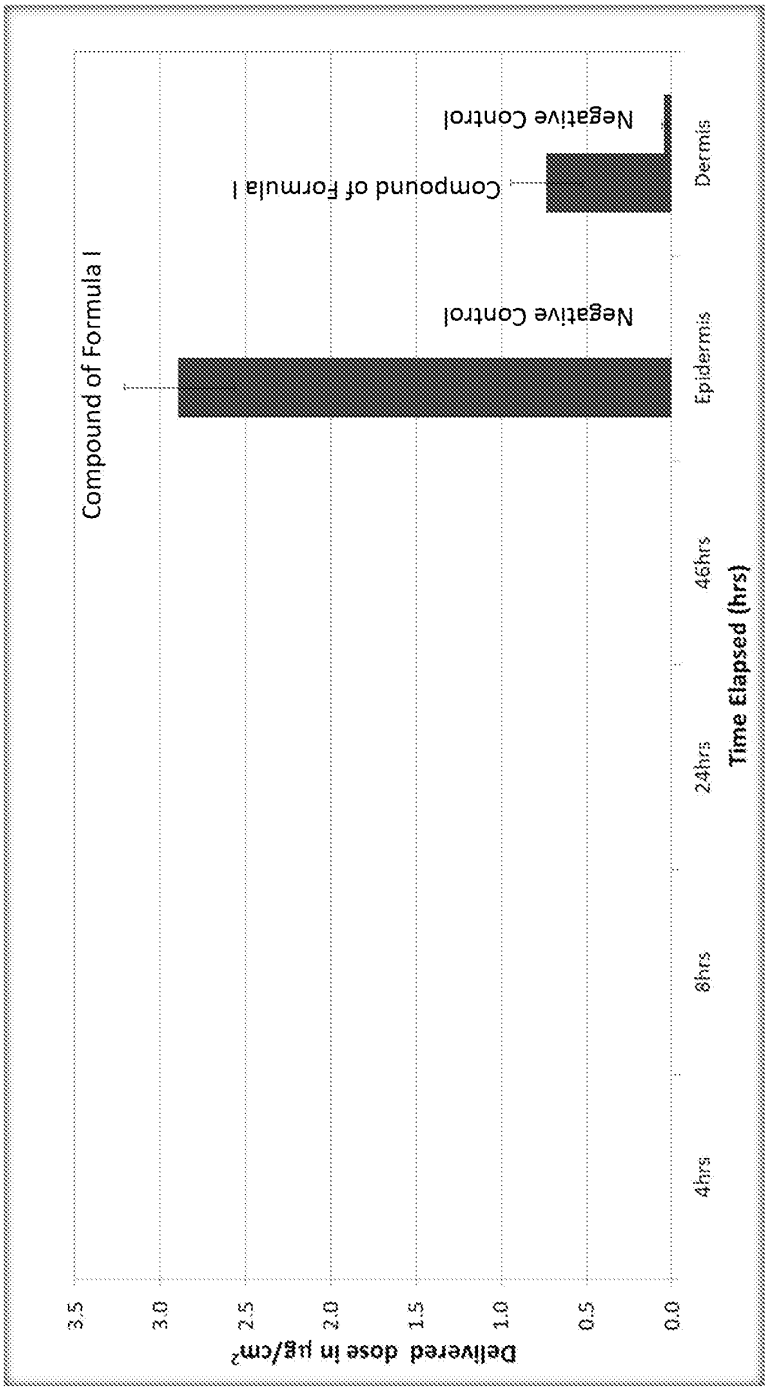

Delivered dose in µg/cm²
3.5
3.0
2.5
2.0
1.5
1.0
0.5
0.0
4hrs
8hrs
24hrs
46hrs
Time Elapsed (hrs)
Epidermis
Dermis
Compound of Formula I
Negative Control
Compound of Formula I
Negative Control

FIG. 6

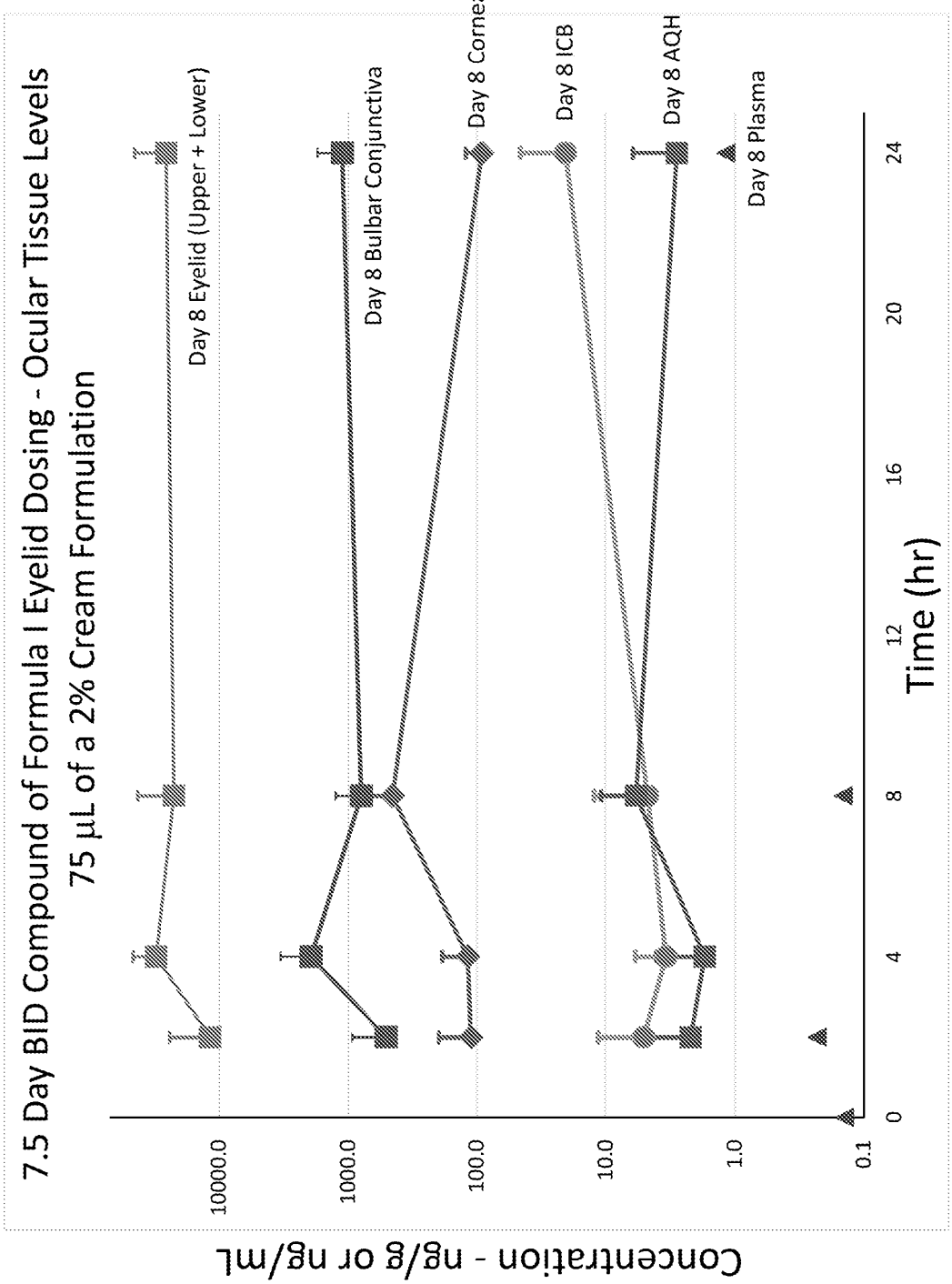

7.5 Day BID Compound of Formula I Eyelid Dosing - Ocular Tissue Levels
75 µL of a 2% Cream Formulation
Day 8 Eyelid (Upper + Lower)
Day 8 Bulbar Conjunctiva
Day 8 Cornea
Day 8 ICB
Day 8 AQH
Day 8 Plasma
Concentration - ng/g or ng/mL
10000.0
1000.0
100.0
10.0
1.0
0.1
0
4
8
12
16
20
24
Time (hr)

FIG. 7 Compound of Formula I Cream Manufacturing Process - Flowchart

