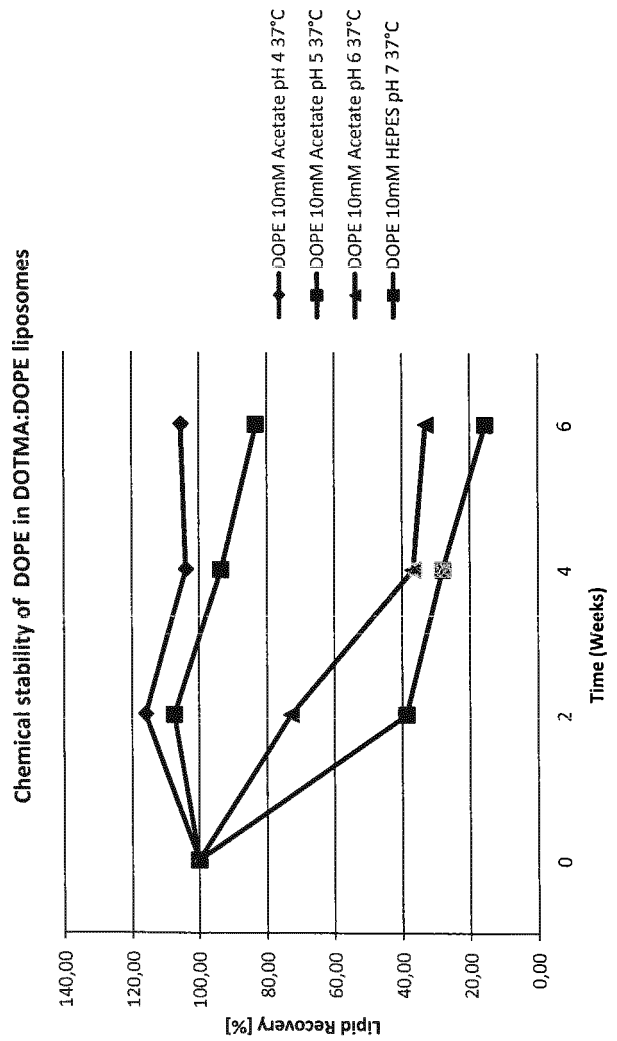
Figure 1


Chemical stability of DOPE in DOTMA:DOPE liposomes
DOPE 10mM Acetate pH 4 37°C
DOPE 10mM Acetate pH 5 37°C
DOPE 10mM Acetate pH 6 37°C
DOPE 10mM HEPES pH 7 37°C
Lipid Recovery [%]
140,00
120,00
100,00
80,00
60,00
40,00
20,00
0,00
0
2
4
6
Time (Weeks)

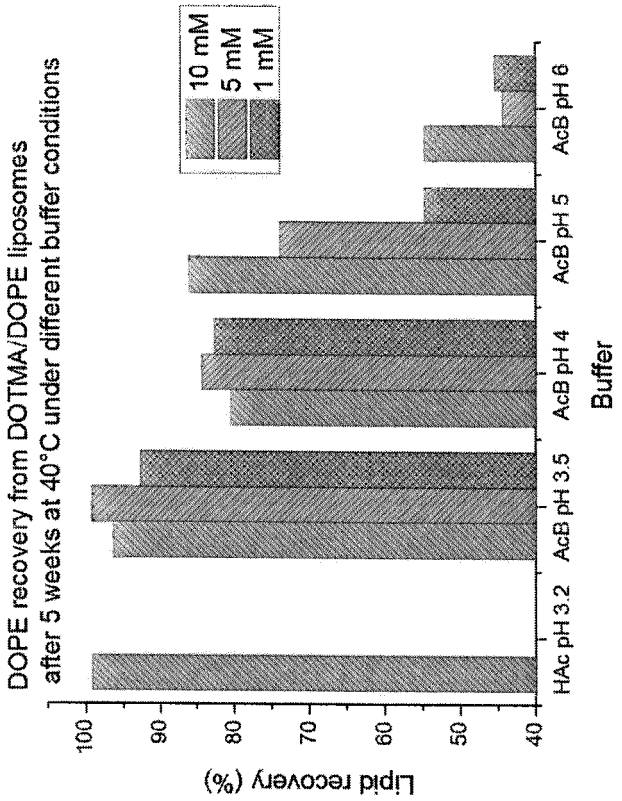

DOPE recovery from DOTMA/DOPE liposomes
after 5 weeks at 40°C under different buffer conditions
10 mM
5 mM
1 mM
Lipid recovery (%)
100
90
80
70
60
50
40
HAc pH 3.2
AcB pH 3.5
AcB pH 4
AcB pH 5
AcB pH 6
Buffer

Figure 2

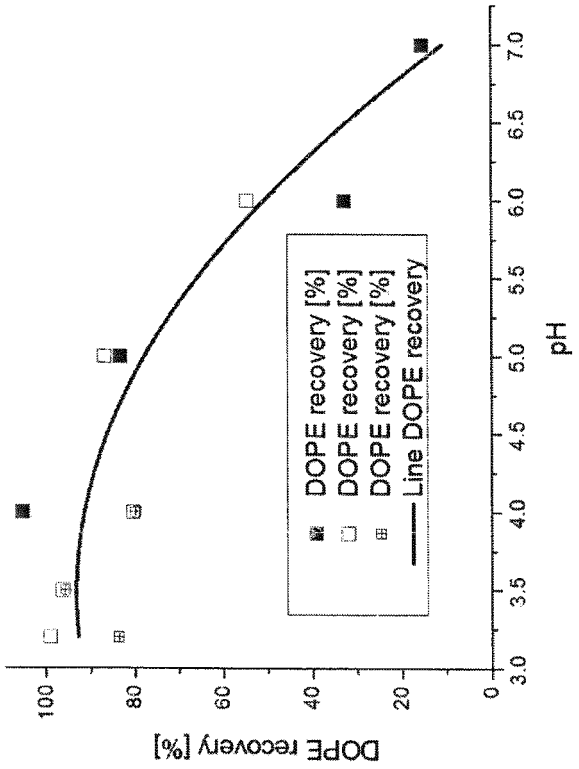

DOPE recovery [%]
DOPE recovery [%]
DOPE recovery [%]
Line DOPE recovery
DOPE recovery [%]
pH
0
20
40
60
80
100
3.0
3.5
4.0
4.5
5.0
5.5
6.0
6.5
7.0

Figure 3

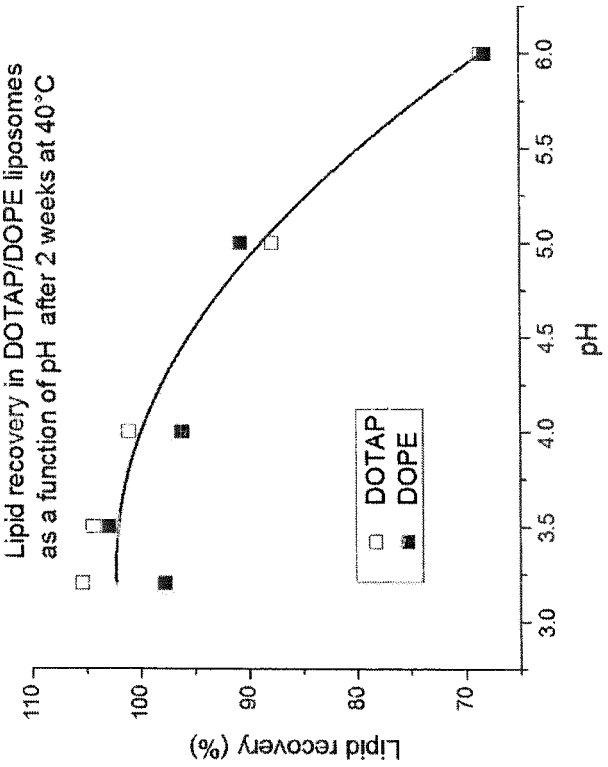

Lipid recovery in DOTAP/DOPE liposomes as a function of pH after 2 weeks at 40°C
Lipid recovery (%)
110
100
90
80
70
3.0
3.5
4.0
4.5
5.0
5.5
6.0
pH
DOTAP
DOPE

Figure 4

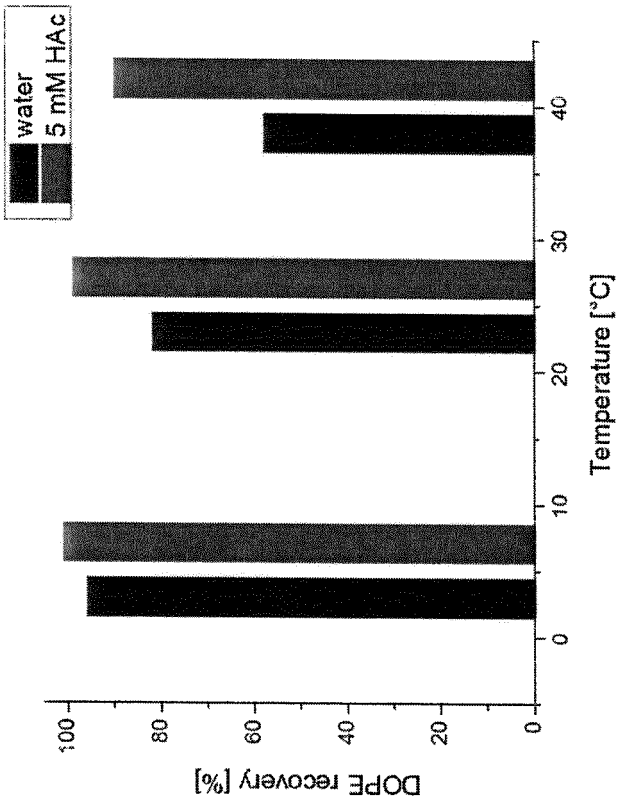

water
5 mM HAc
DOPE recovery [%]
0
20
40
60
80
100
0
10
20
30
40
Temperature [°C]

Figure 5

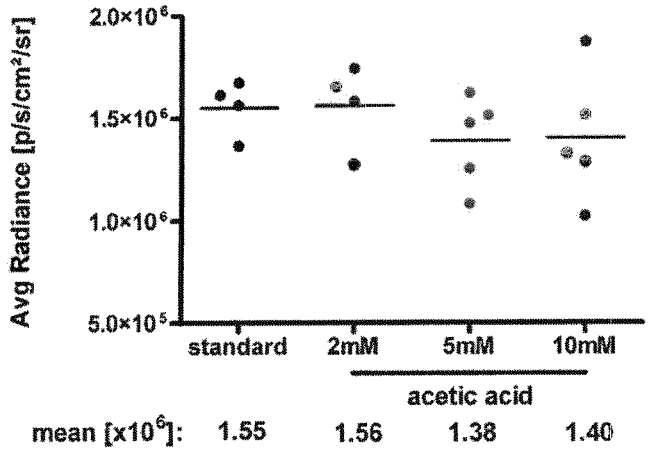


Figure 6

WO 2016/046060 PCT/EP2015/071344

A

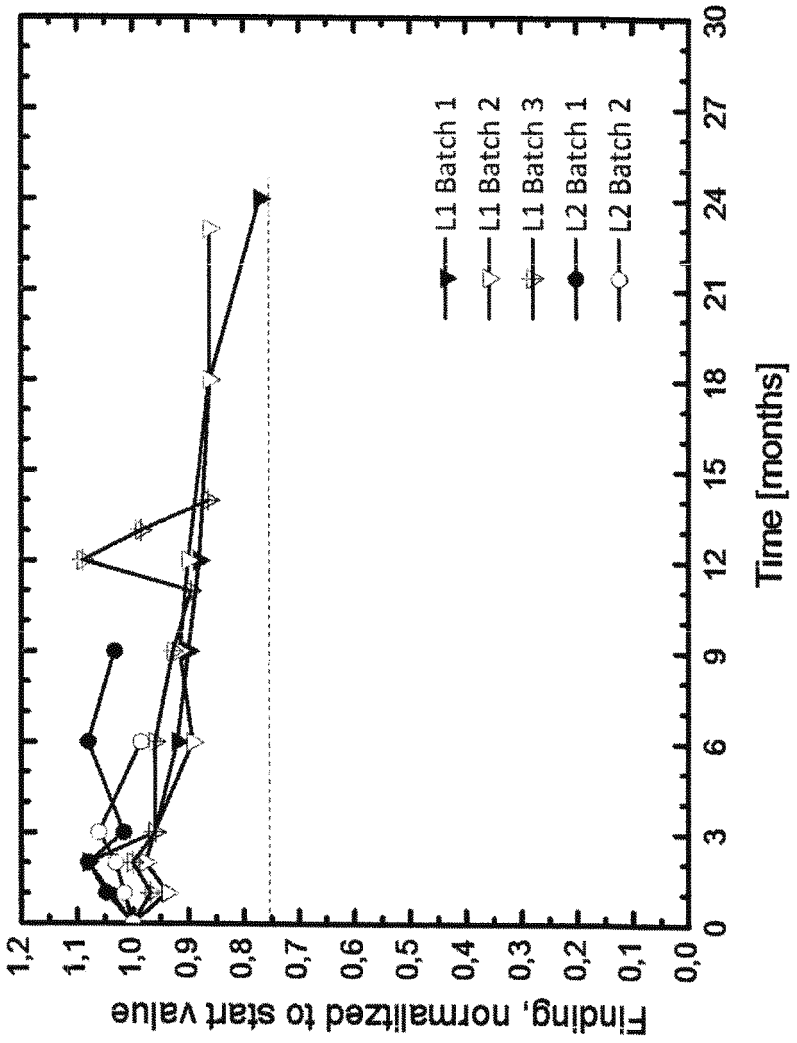

Finding, normalized to start value
1,2
1,1
1,0
0,9
0,8
0,7
0,6
0,5
0,4
0,3
0,2
0,1
0,0
0
3
6
9
12
15
18
21
24
27
30
Time [months]
L1 Batch 1
L1 Batch 2
L1 Batch 3
L2 Batch 1
L2 Batch 2

Figure 7

B

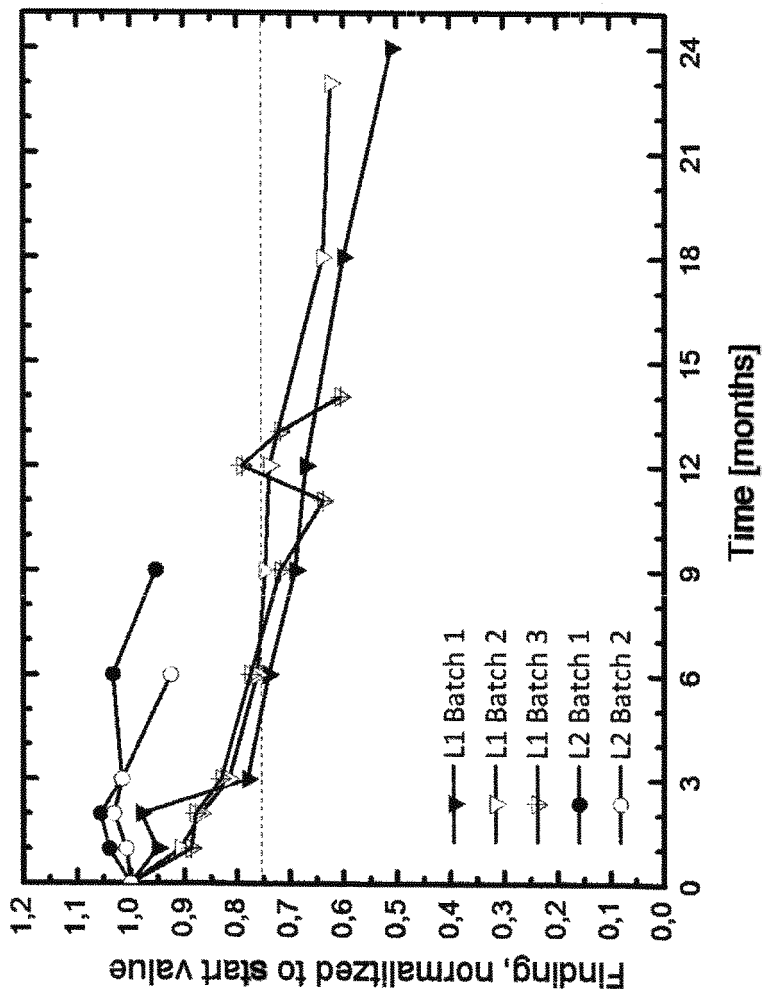


Figure 7 (cont.)

C

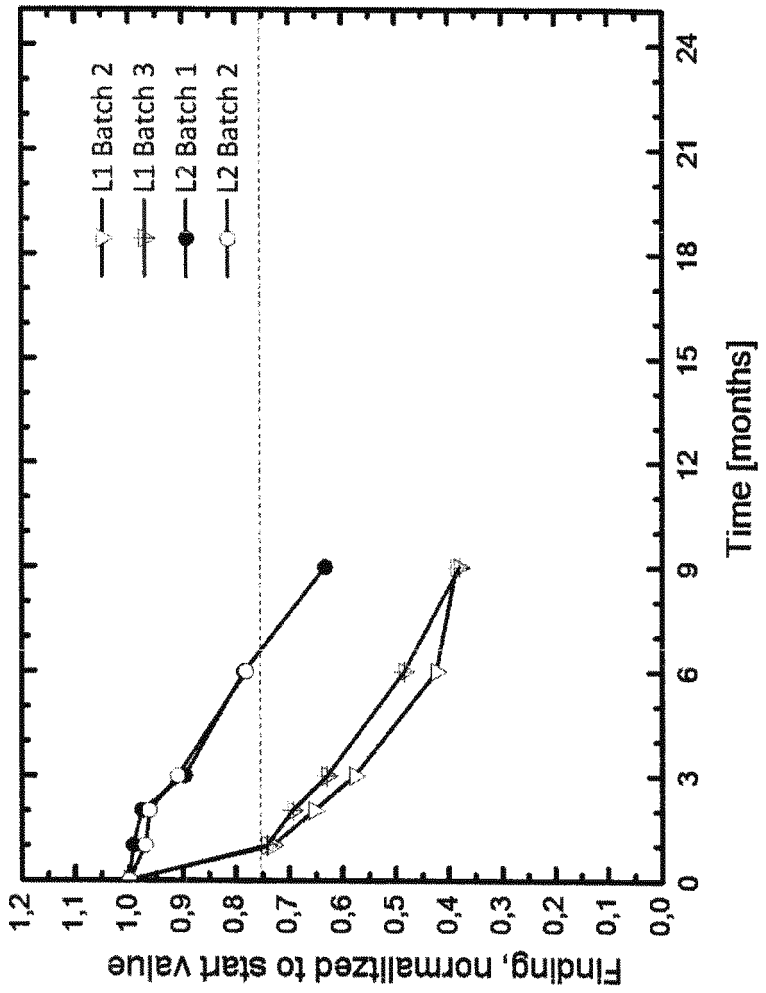


Figure 7 (cont.)