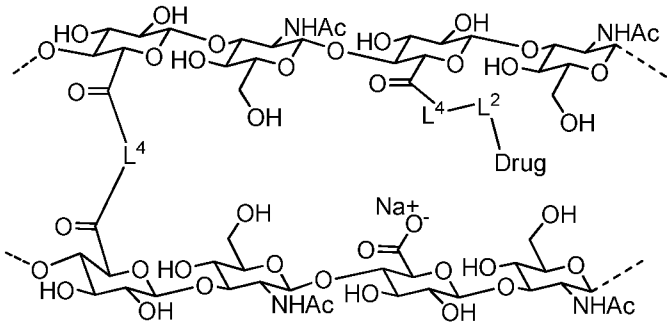


1

FIGURE 1

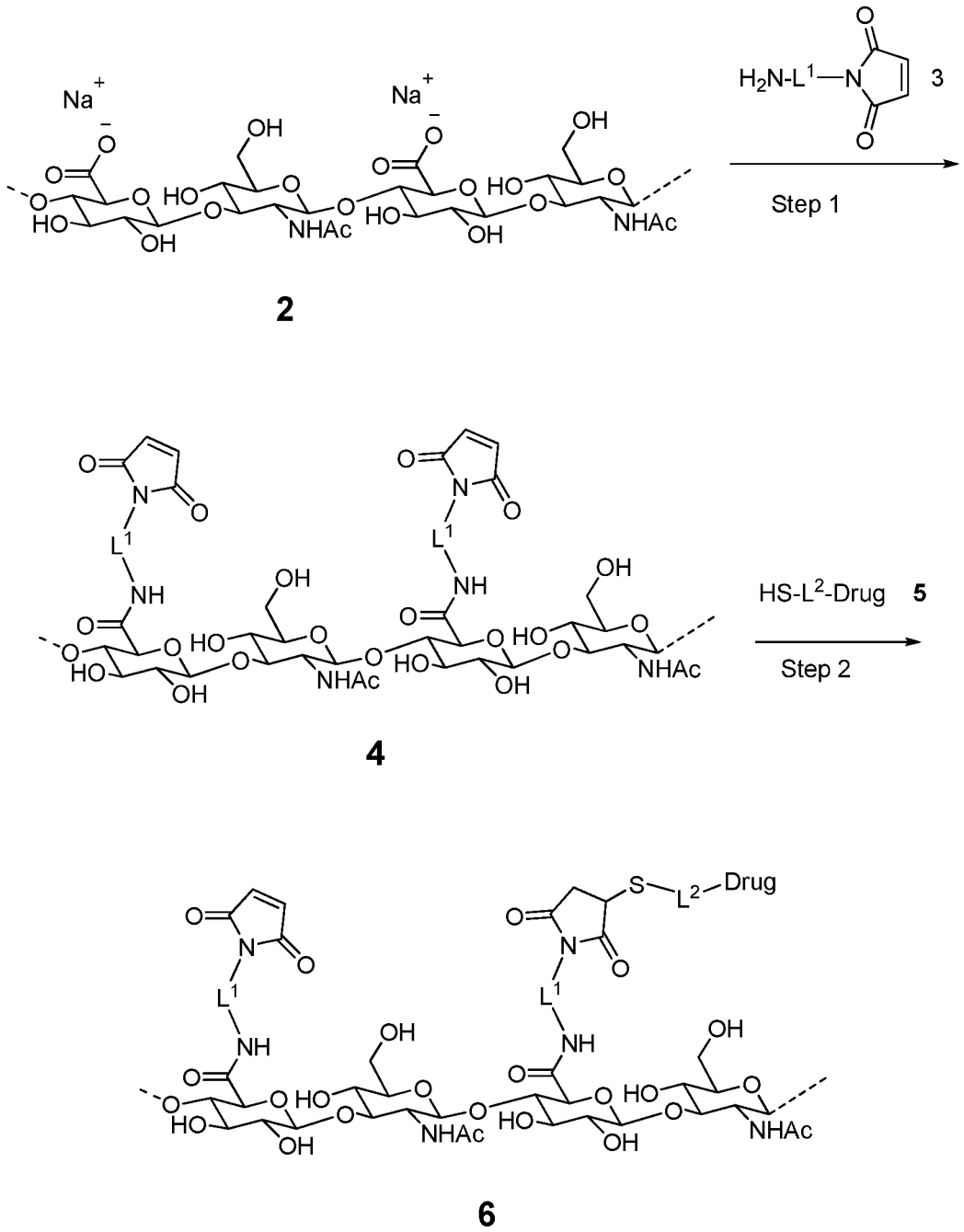
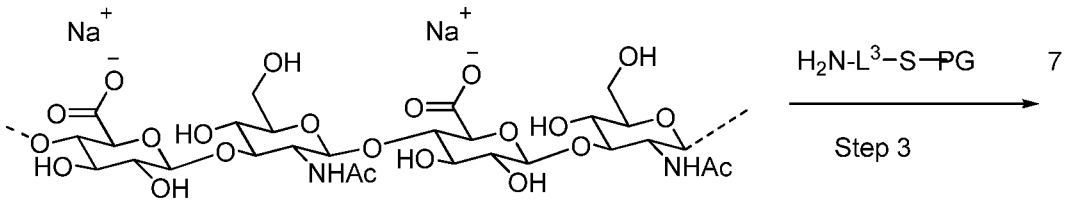
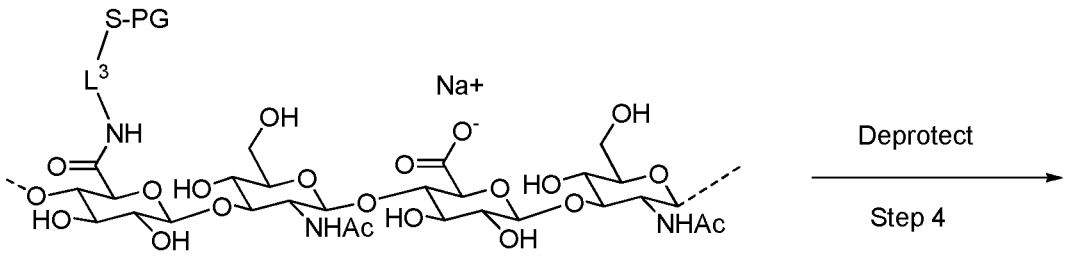


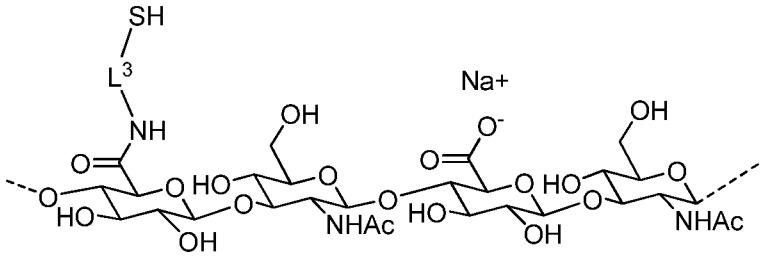
FIGURE 2



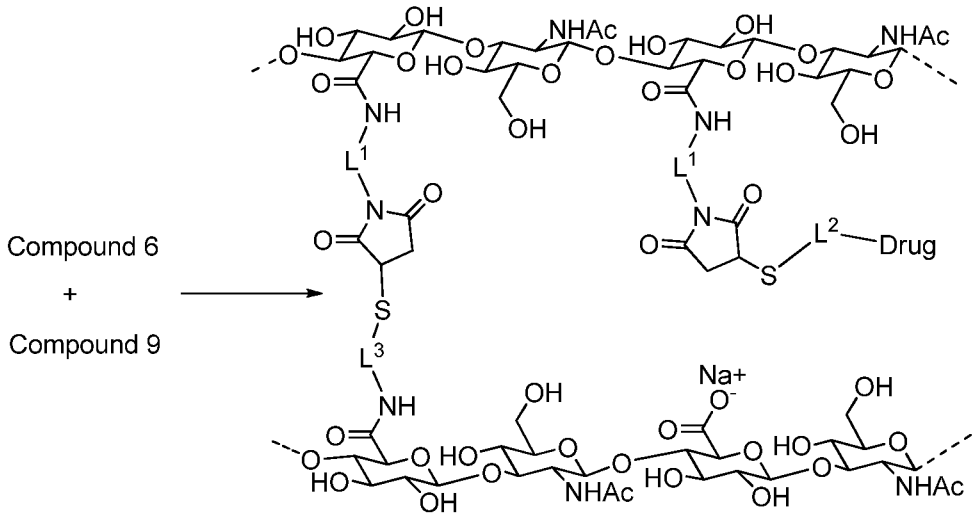
2



8

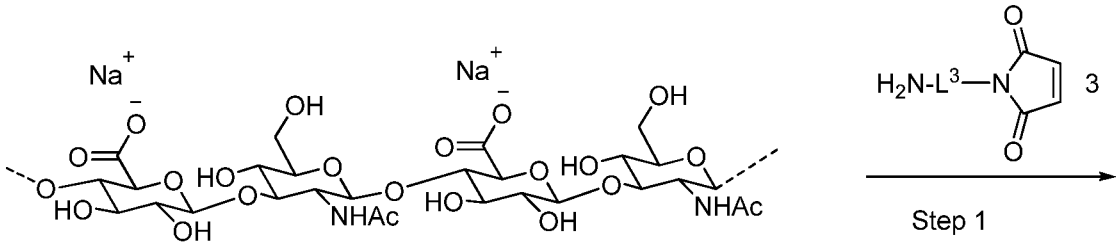


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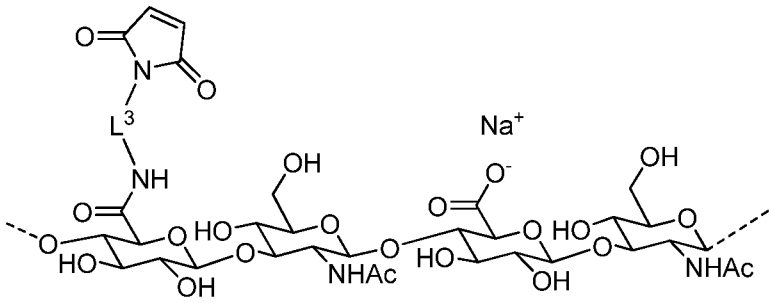


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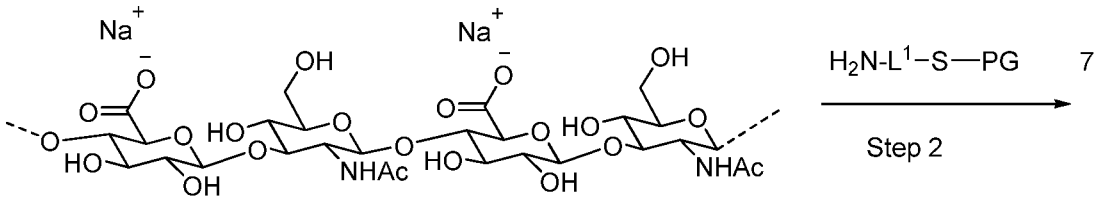
FIGURE 3



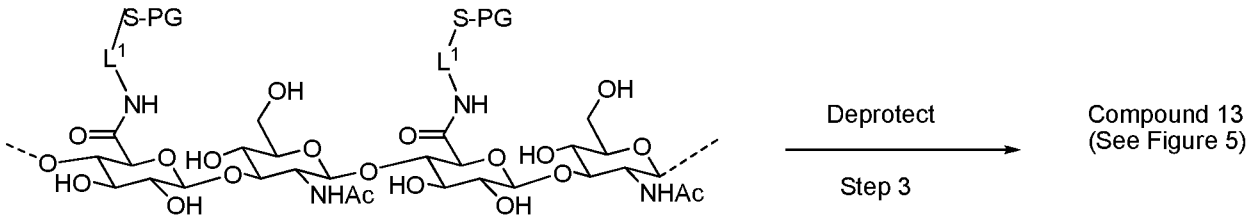
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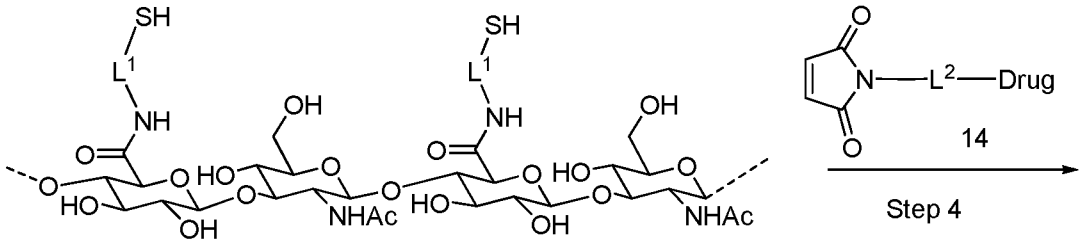


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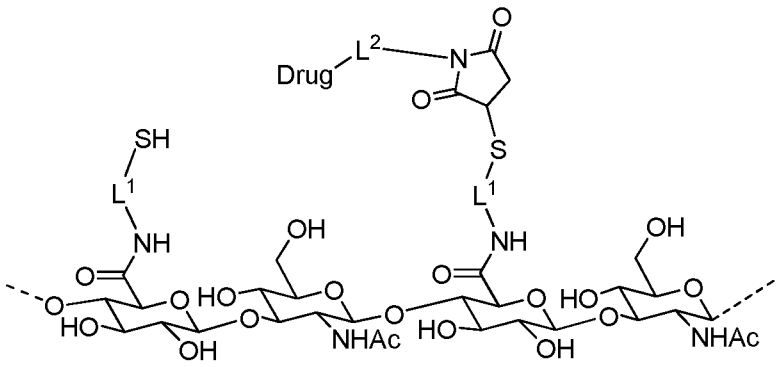


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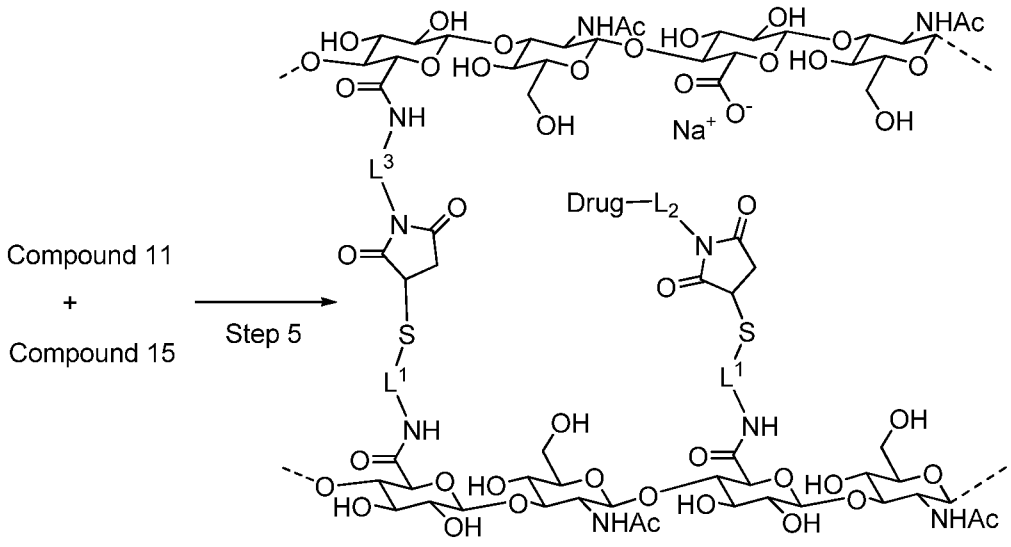
FIGURE 4



13



15



16

FIGURE 5

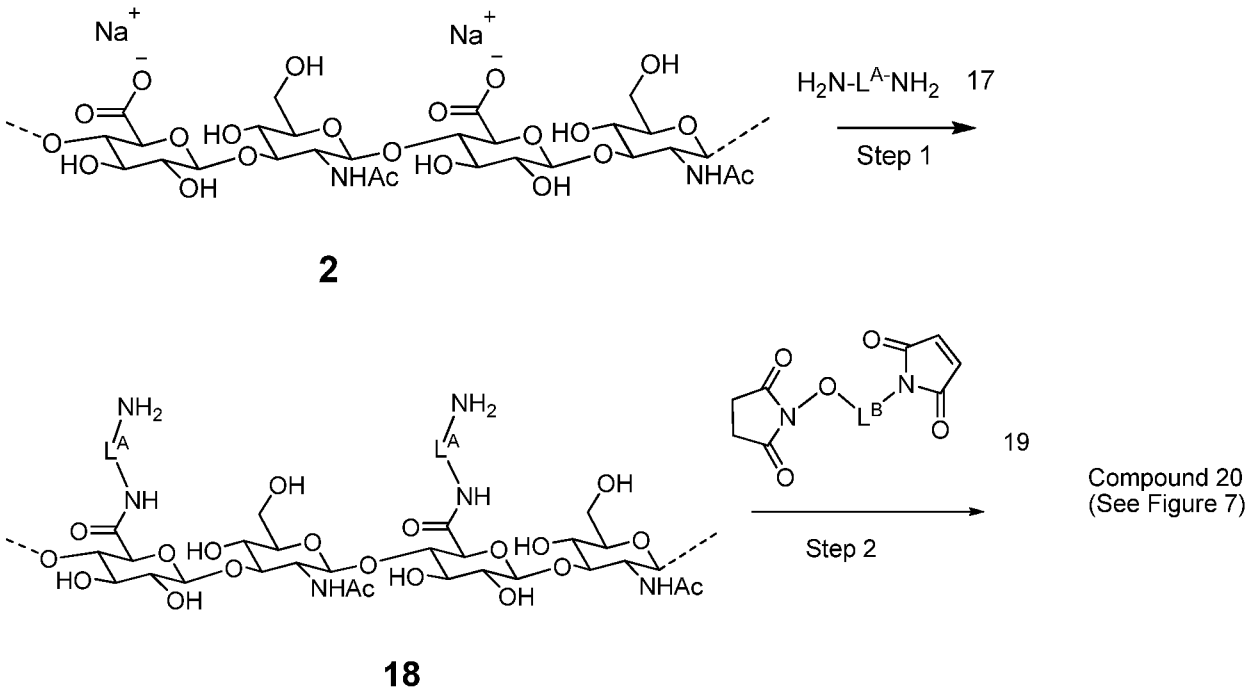
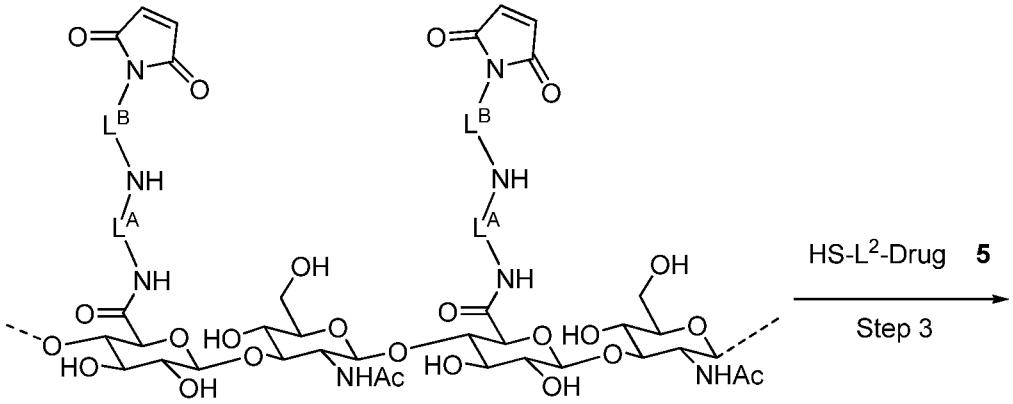
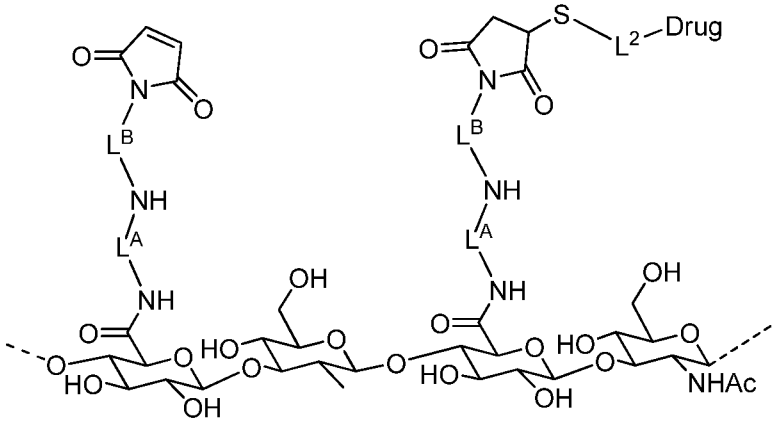

Na+
OH
HO
NHAc
OH
H2N-LA-NH2 17
Step 1
2
NH2
LA
NH
19
Step 2
Compound 20
(See Figure 7)
18

FIGURE 6



20



21

FIGURE 7

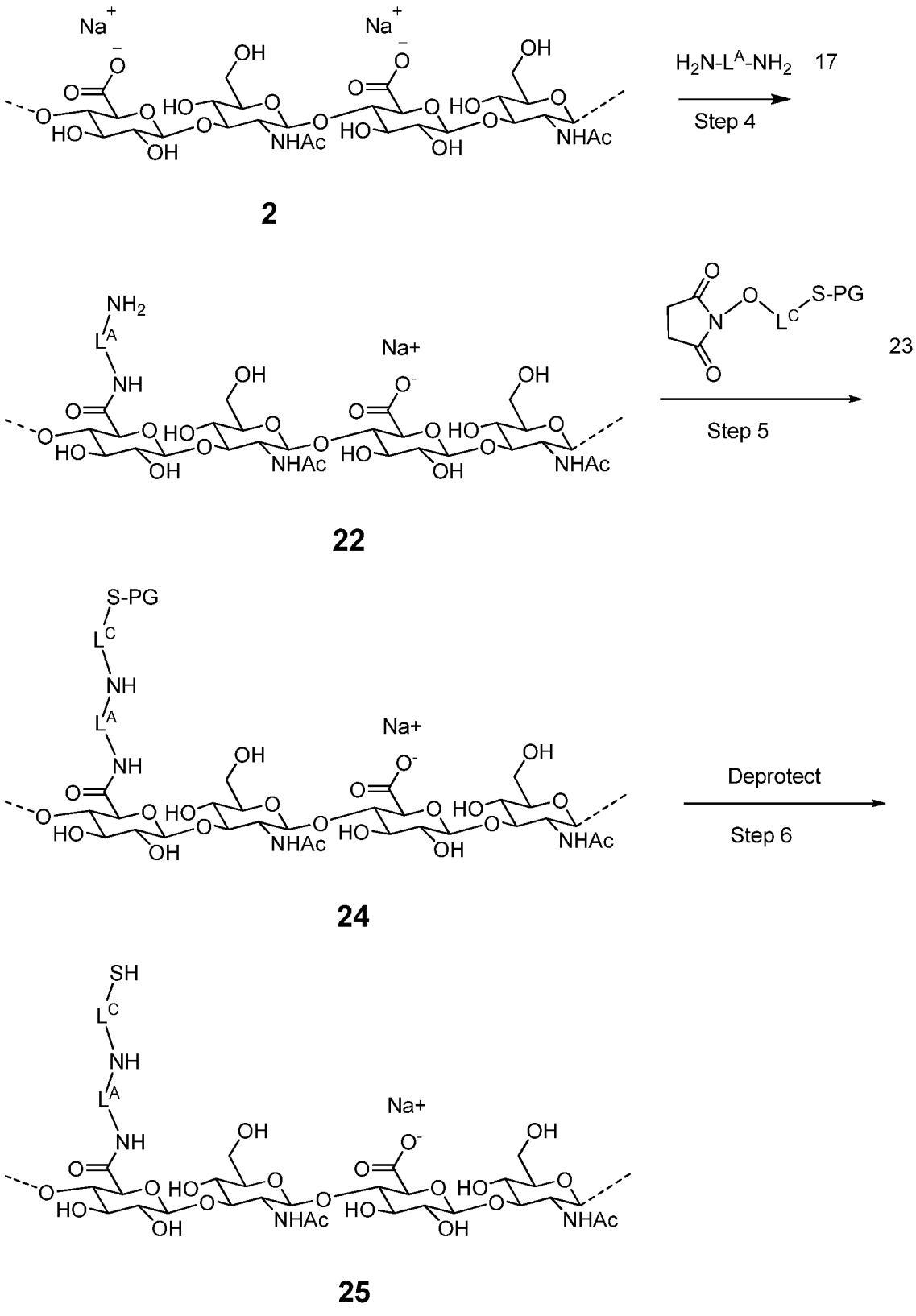
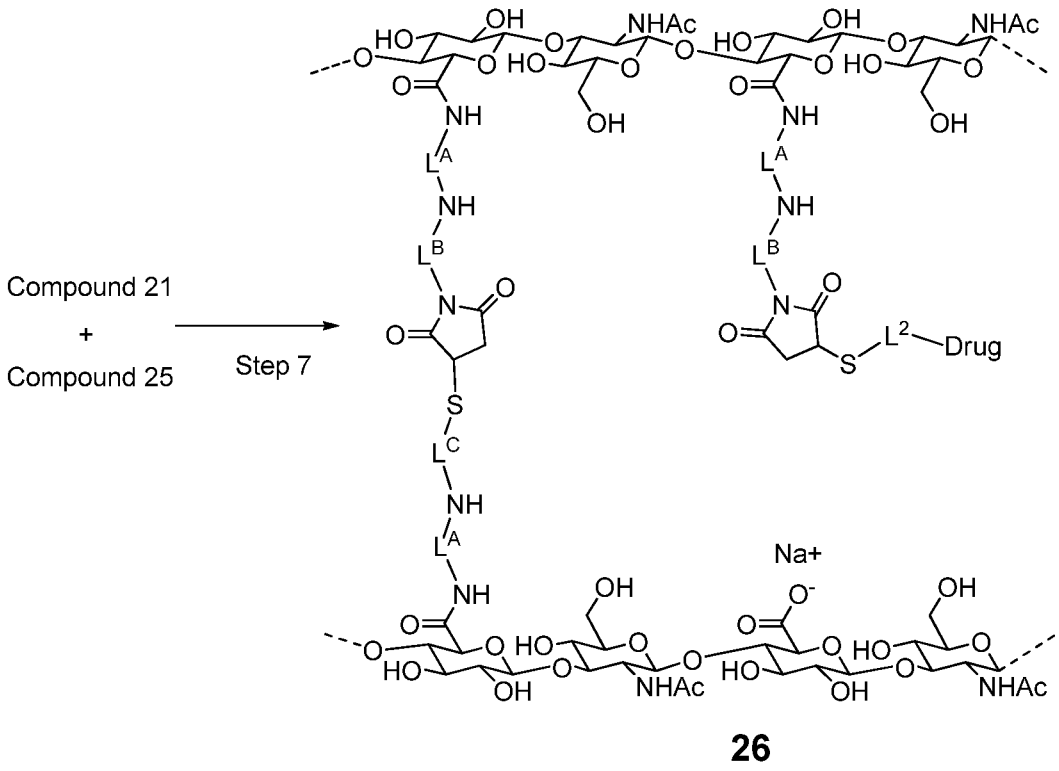

Na+
OH
HO
O
NHAc
H_2N-L^A-NH_2 17
Step 4
2
NH_2
L^A
NH
Na+
S-PG
L^C
23
Step 5
22
Deprotect
Step 6
24
SH
25

FIGURE 8



26

FIGURE 9

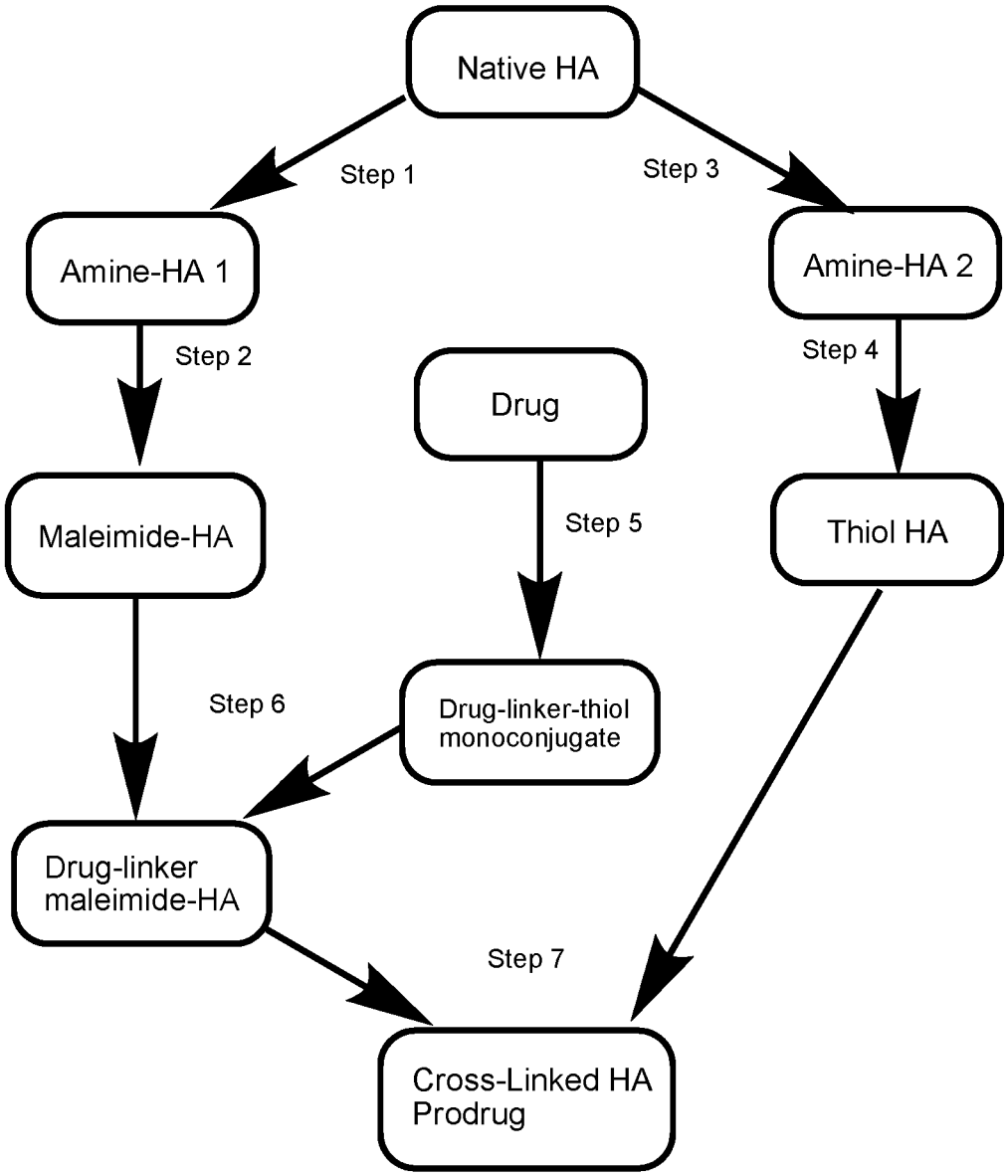

Native HA
Step 1
Step 3
Amine-HA 1
Amine-HA 2
Step 2
Step 4
Drug
Maleimide-HA
Step 5
Thiol HA
Step 6
Drug-linker-thiol monoconjugate
Drug-linker maleimide-HA
Step 7
Cross-Linked HA Prodrug

FIGURE 10

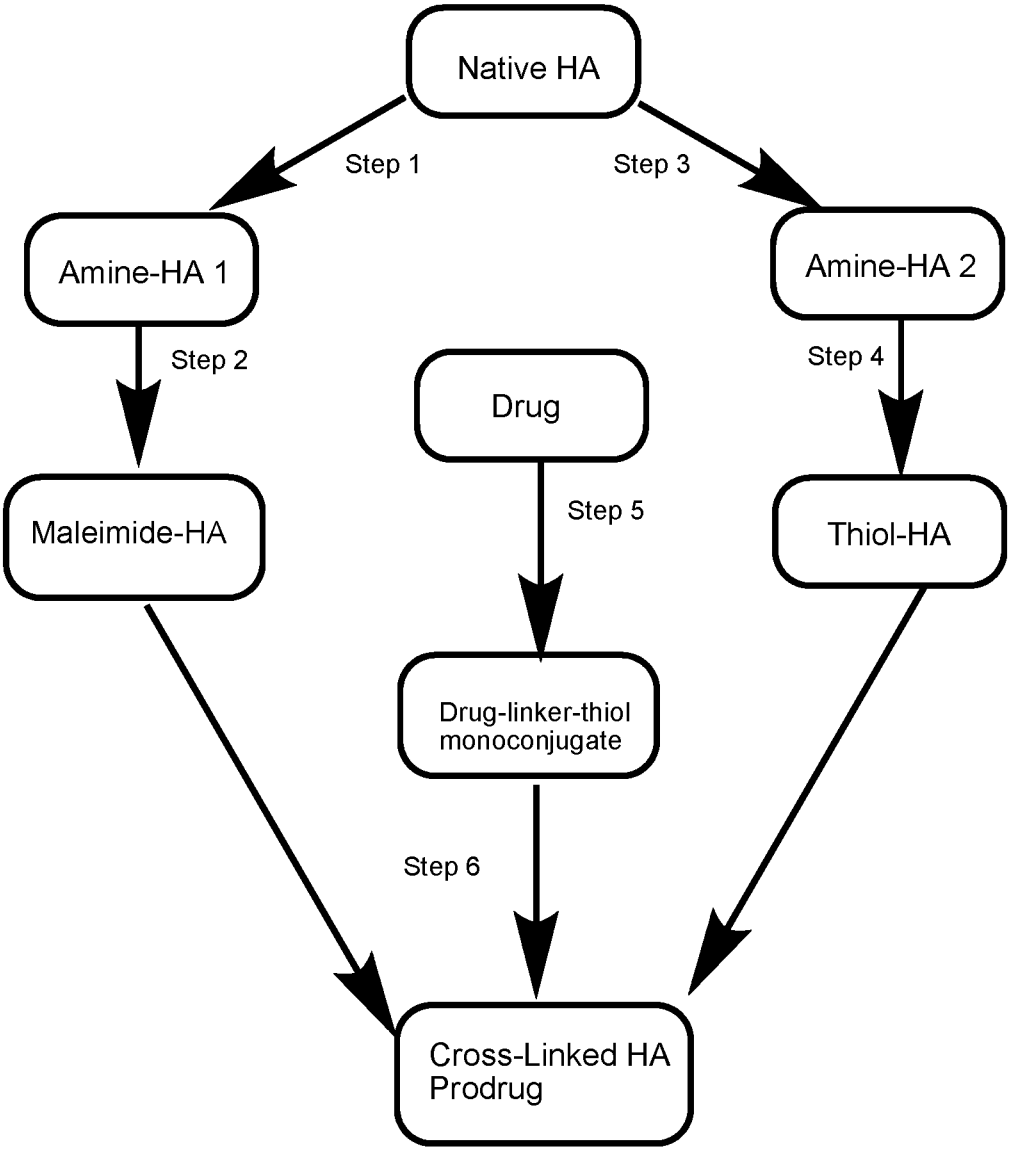

Native HA
Step 1
Step 3
Amine-HA 1
Amine-HA 2
Step 2
Step 4
Drug
Maleimide-HA
Step 5
Thiol-HA
Drug-linker-thiol monoconjugate
Step 6
Cross-Linked HA Prodrug

FIGURE 11

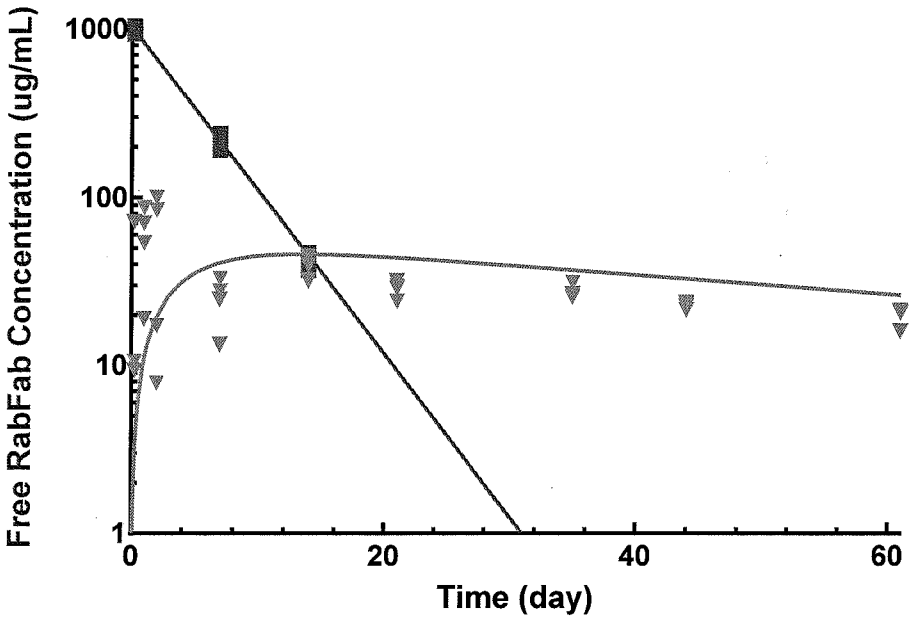

Free RabFab Concentration (ug/mL)
1000
100
10
1
0
20
40
60
Time (day)

FIGURE 12

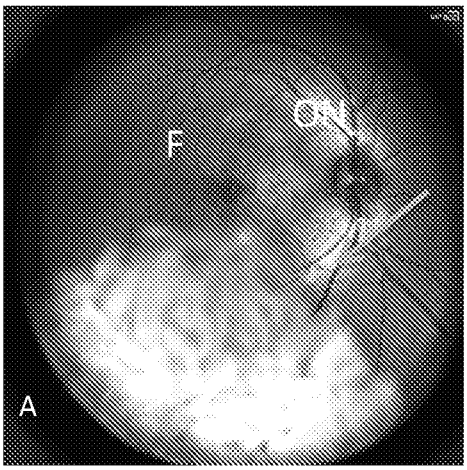
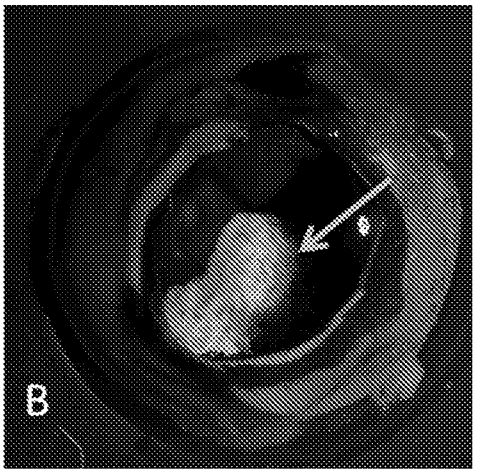


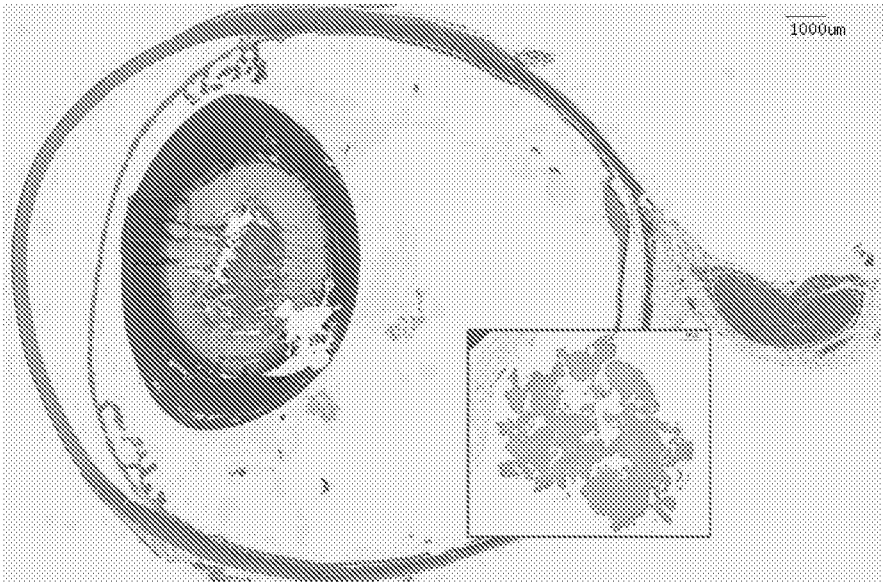
FIGURE 13A



Test article

FIGURE 13B

FIGURE 14



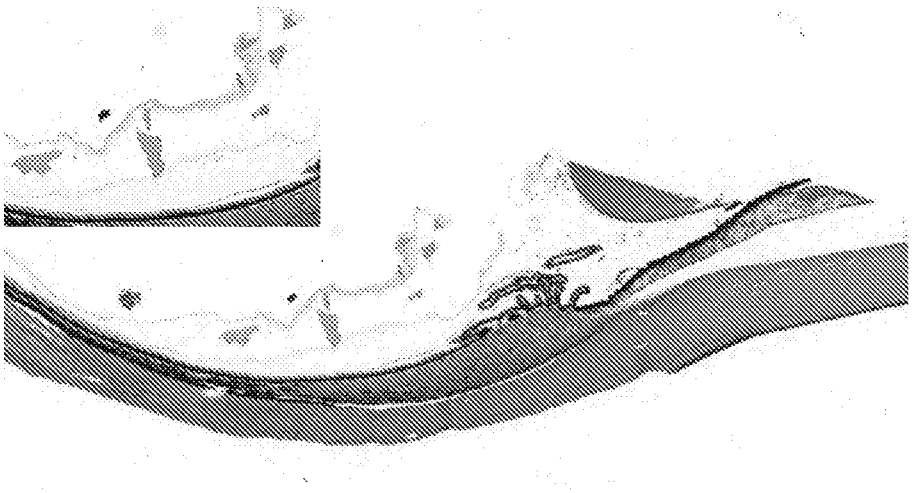


FIGURE 15