## Appendices

Appendix I

**Chemical synthesis** 

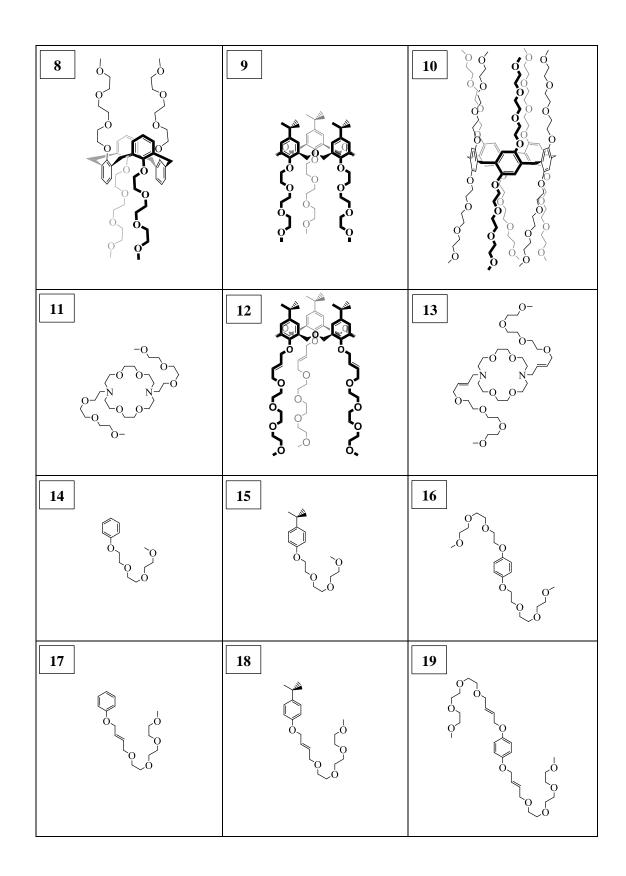
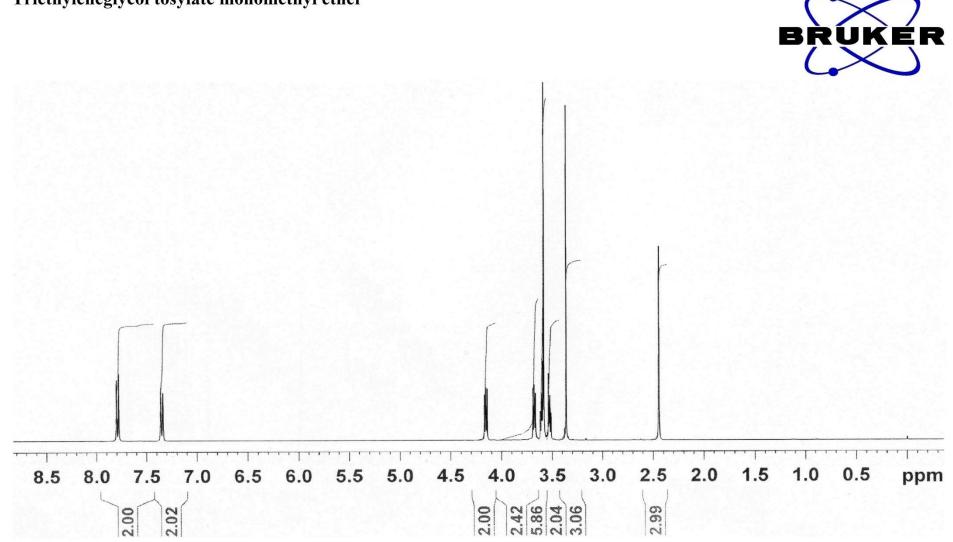


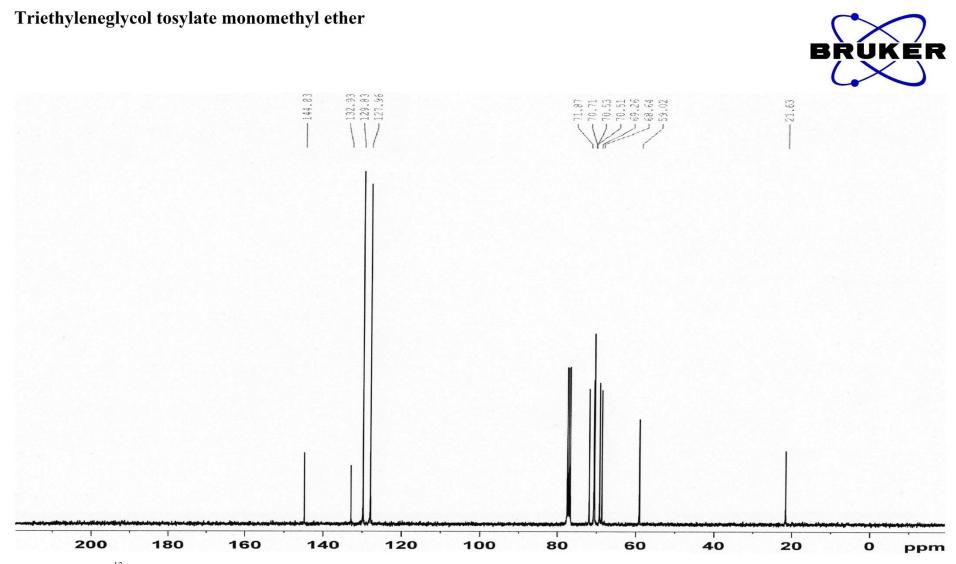
Table A1-1: Chemical structures of compounds 8-19, whose ion channel activities and antimicrobial properties were assessed.



Triethyleneglycol tosylate monomethyl ether

Figure A1 - 1(a): <sup>1</sup>H NMR spectra of 1 conducted in CDCl<sub>3</sub>.

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**Figure A1 - 1(b):** <sup>13</sup>C NMR spectra of **1** conducted in CDCl<sub>3</sub>.

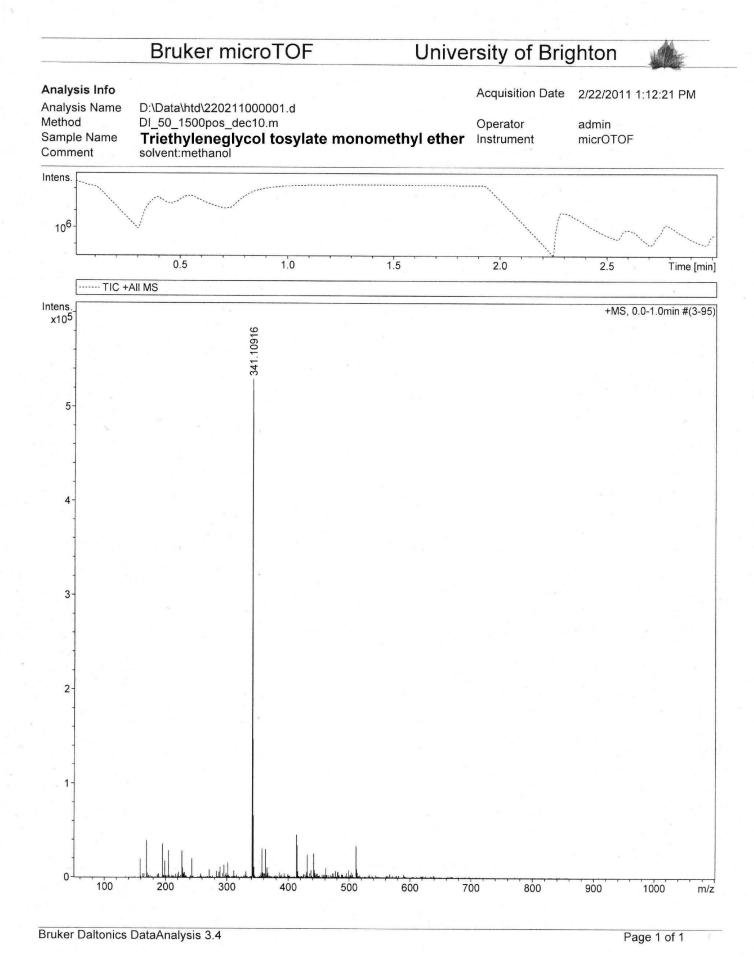
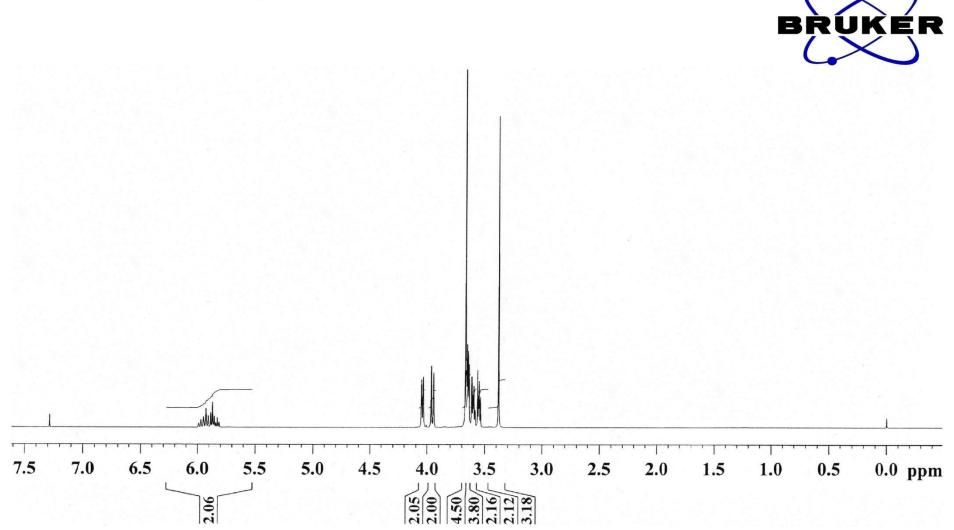
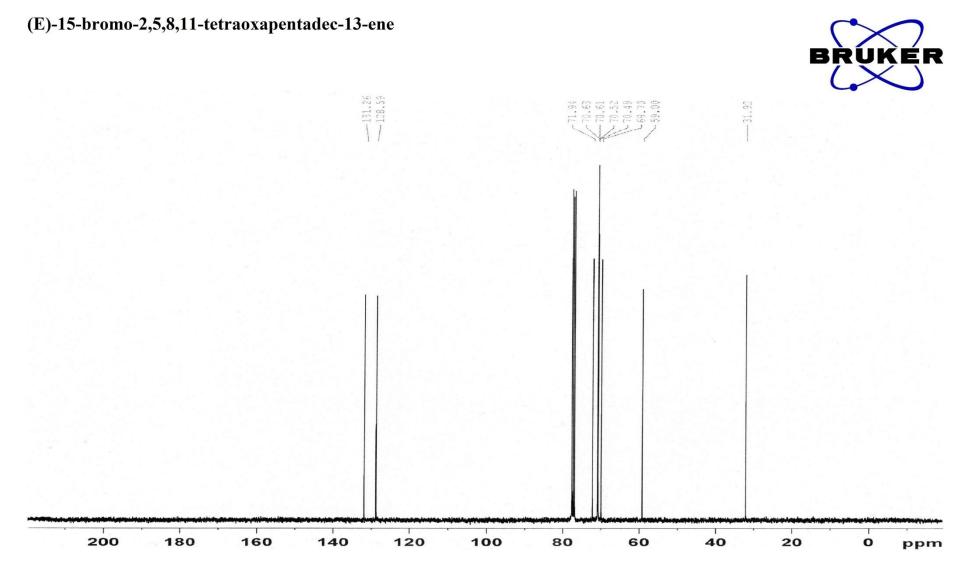


Figure A1 - 1(c): Mass spectra of 1 conducted in methanol.

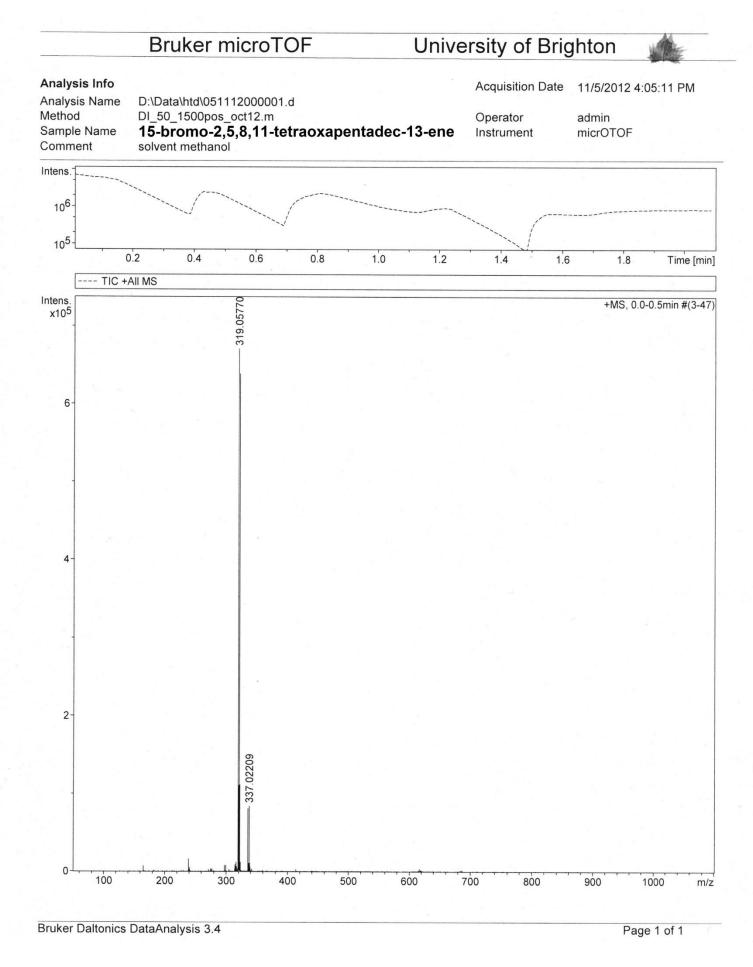


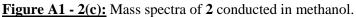
<sup>(</sup>E)-15-bromo-2,5,8,11-tetraoxapentadec-13-ene

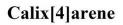
**Figure A1 - 2(a):** <sup>1</sup>H NMR spectra of **2** conducted in CDCl<sub>3</sub>.



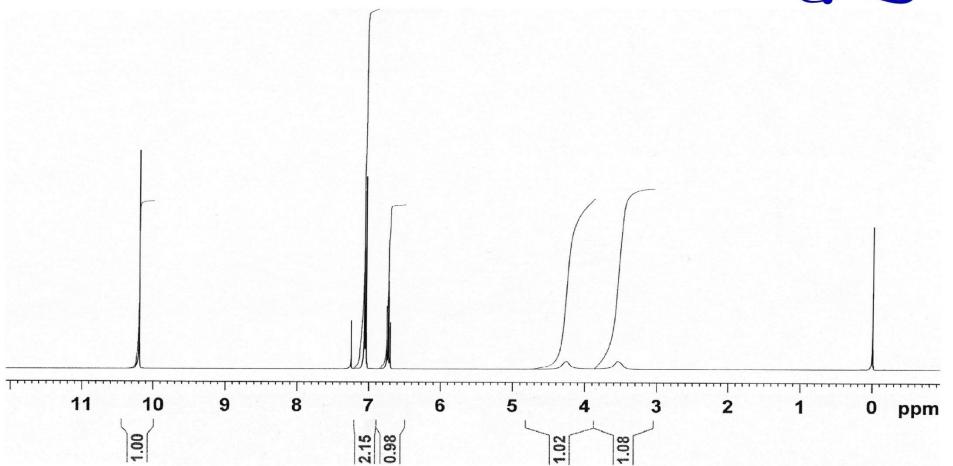
**Figure A1 - 2(b):** <sup>13</sup>C NMR spectra of **2** conducted in CDCl<sub>3</sub>.





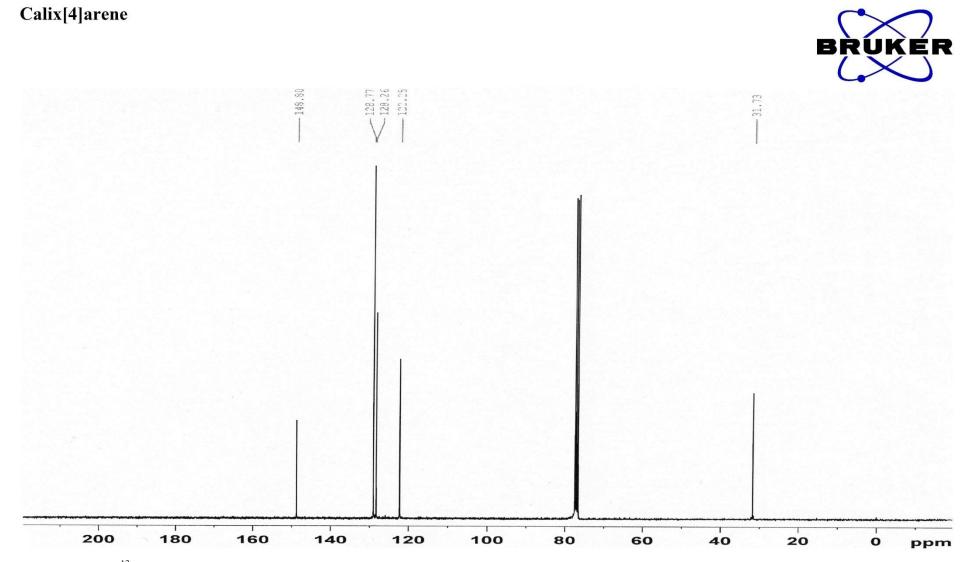






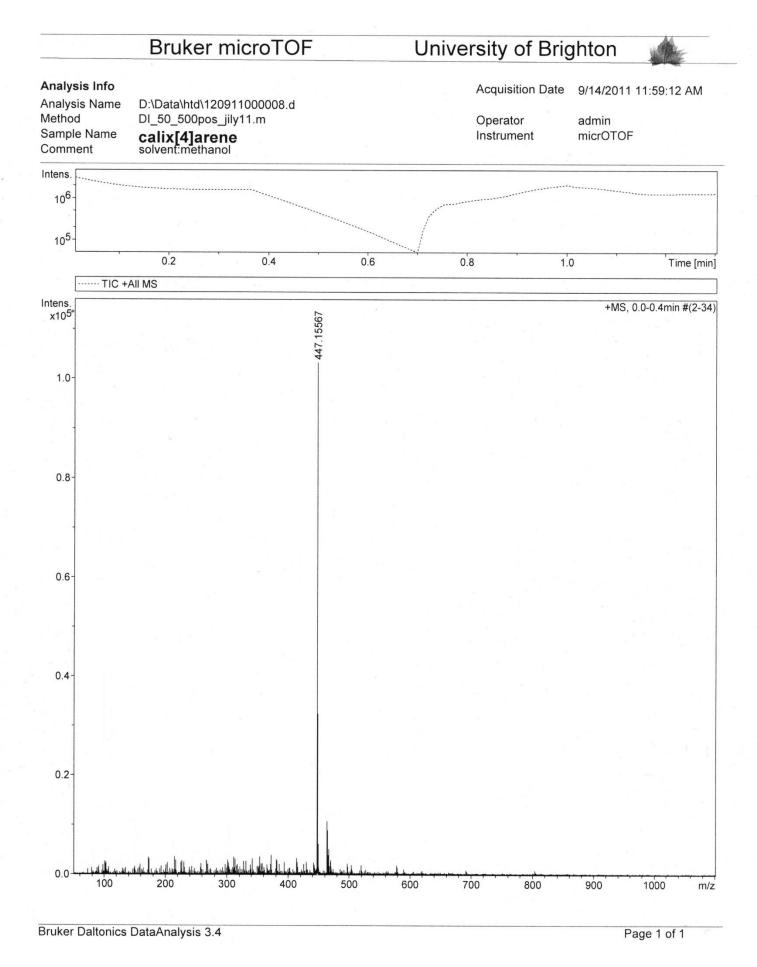
**Figure A1 - 3(a):** <sup>1</sup>H NMR spectra of **3** conducted in CDCl<sub>3</sub>.

 $\infty$ 

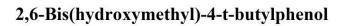


**<u>Figure A1 - 3(b)</u>**<sup>13</sup>C NMR spectra of **3** conducted in CDCl<sub>3</sub>.

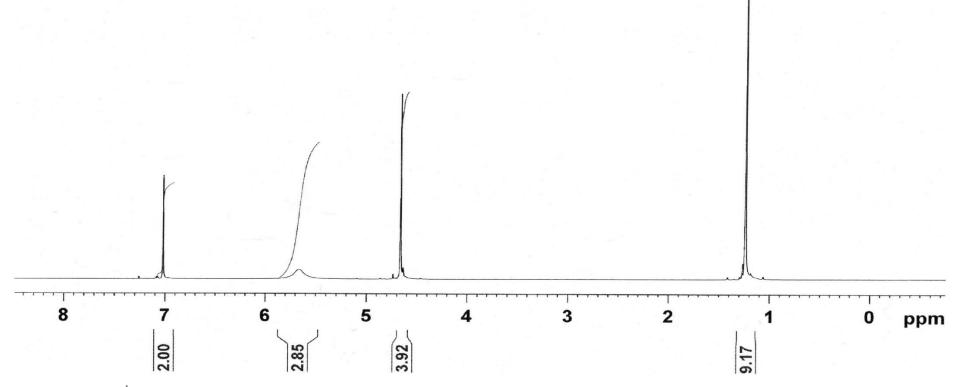
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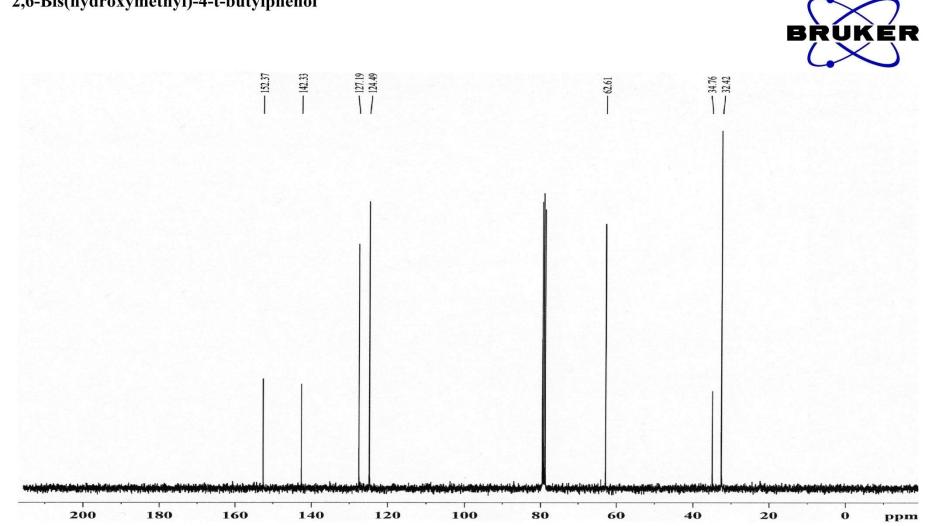








**Figure A1 - 4(a):** <sup>1</sup>H NMR spectra of **4** conducted in CDCl<sub>3</sub>.



2,6-Bis(hydroxymethyl)-4-t-butylphenol

**<u>Figure A1 - 4(b)</u>**<sup>13</sup>C NMR spectra of **4** conducted in CDCl<sub>3</sub>.

## Bruker microTOF

## University of Brighton

#### Analysis Info

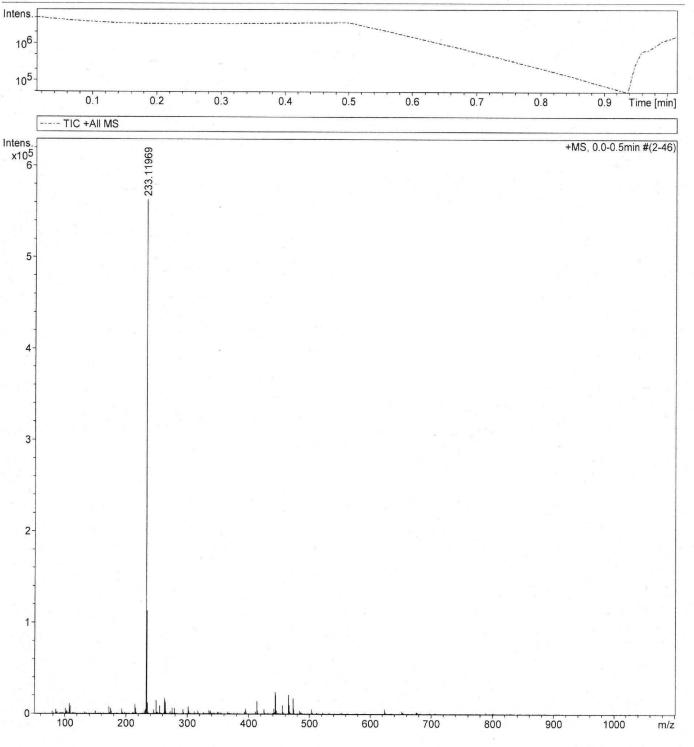


DI\_50\_500pos\_jily11.m 2,6-Bis(hydroxymethyl)-4-t-butylphenol Acquisition Date 9/16/2011 3:14:11 PM

Operator Instrument

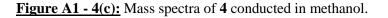
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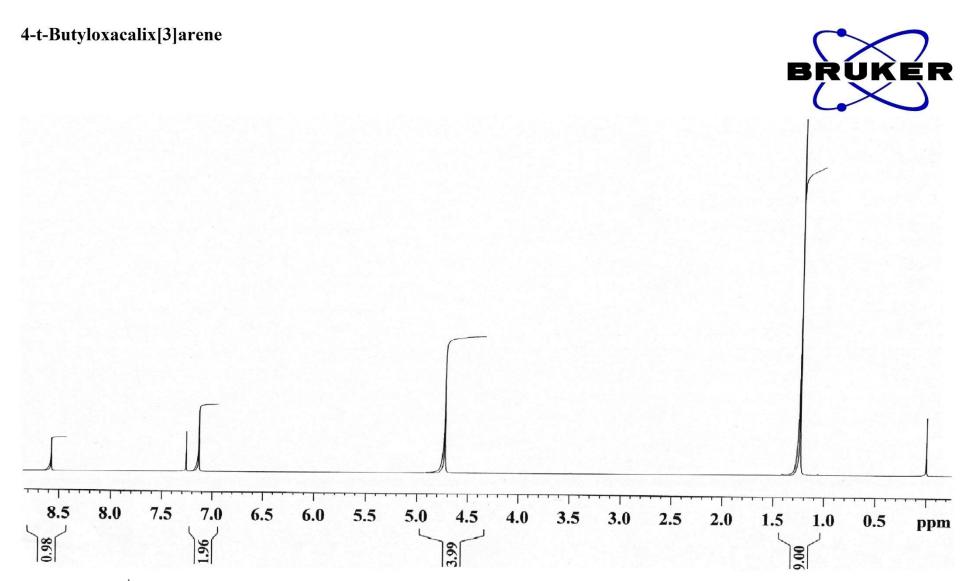
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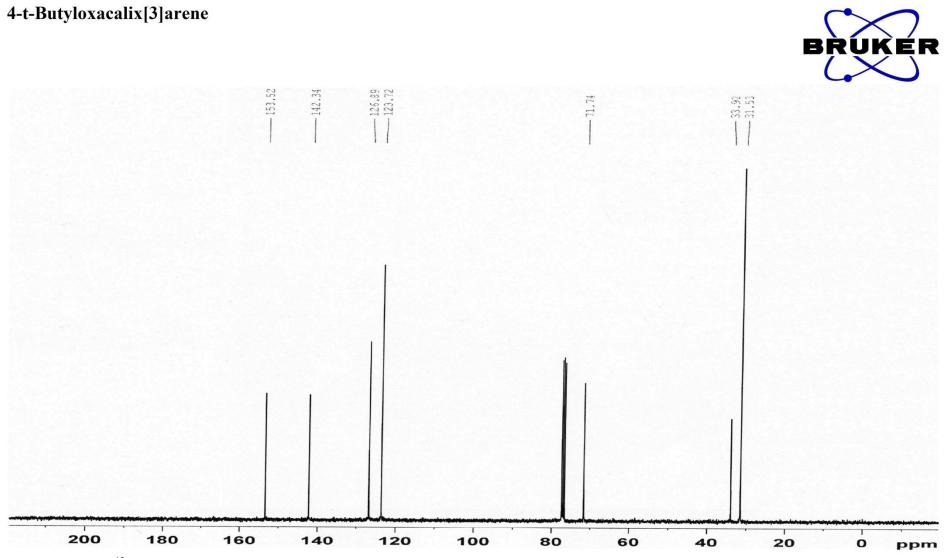
Bruker Daltonics DataAnalysis 3.4

Page 1 of 1



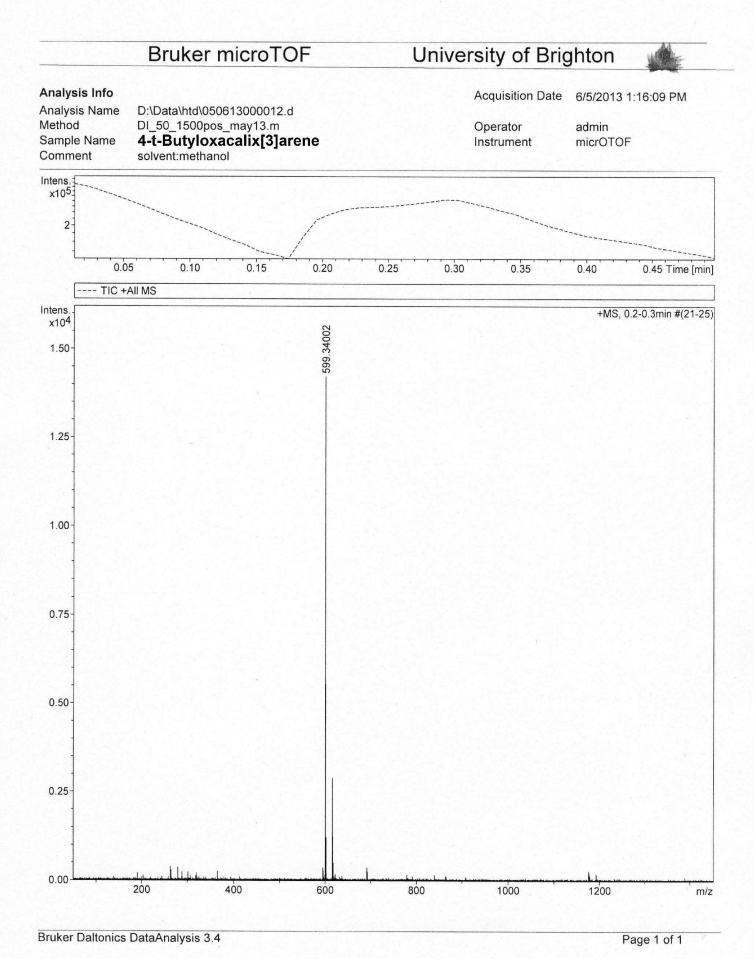


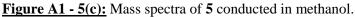
**Figure A1 - 5(a):** <sup>1</sup>H NMR spectra of **5** conducted in CDCl<sub>3</sub>.



**Figure A1 - 5(b):** <sup>13</sup>C NMR spectra of **5** conducted in CDCl<sub>3</sub>.

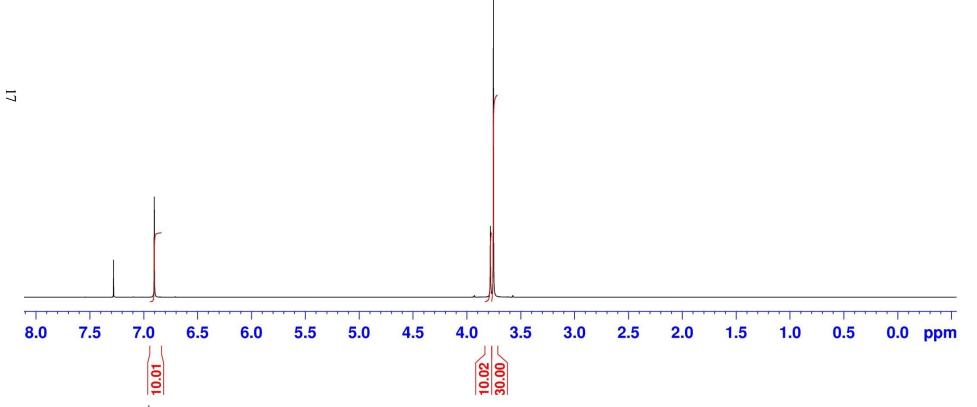
15



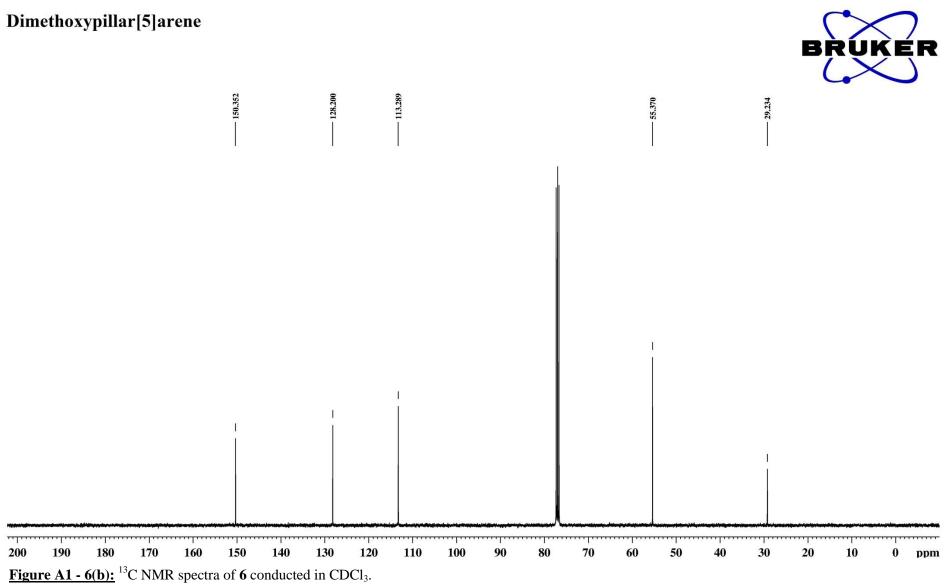


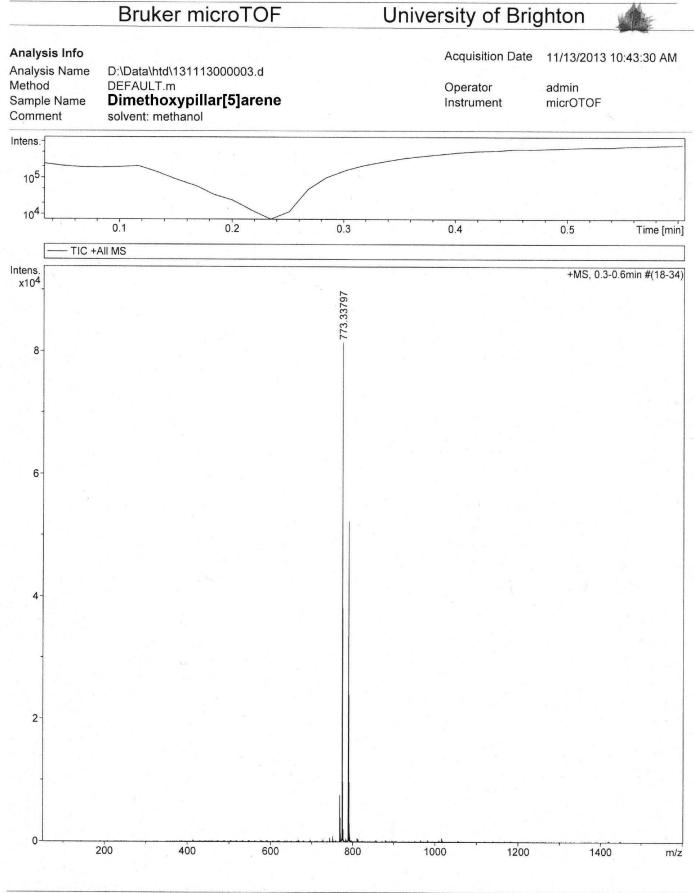
Dimethoxypillar[5]arene





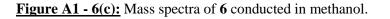
**Figure A1 - 6(a):** <sup>1</sup>H NMR spectra of **6** conducted in CDCl<sub>3</sub>.

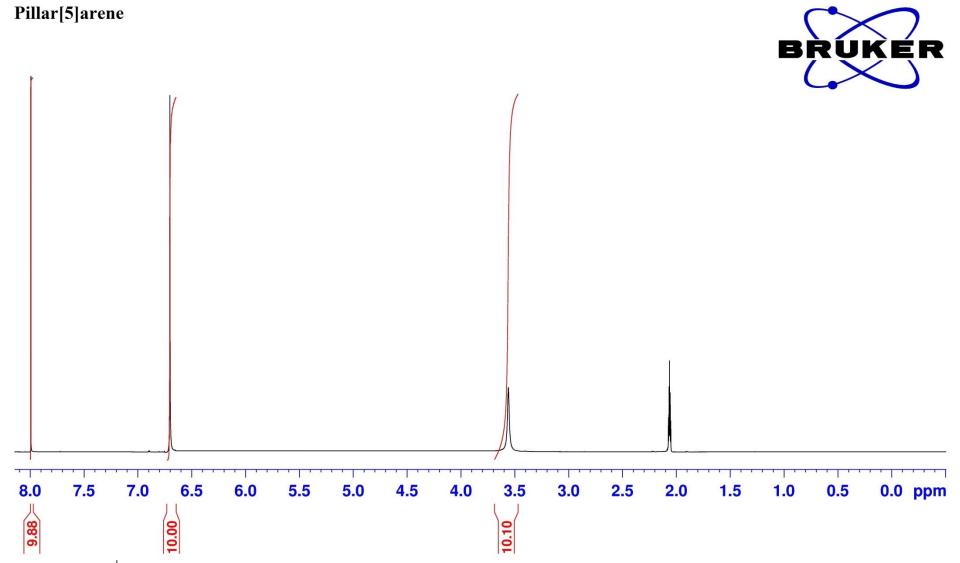




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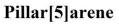
Page 1 of 1

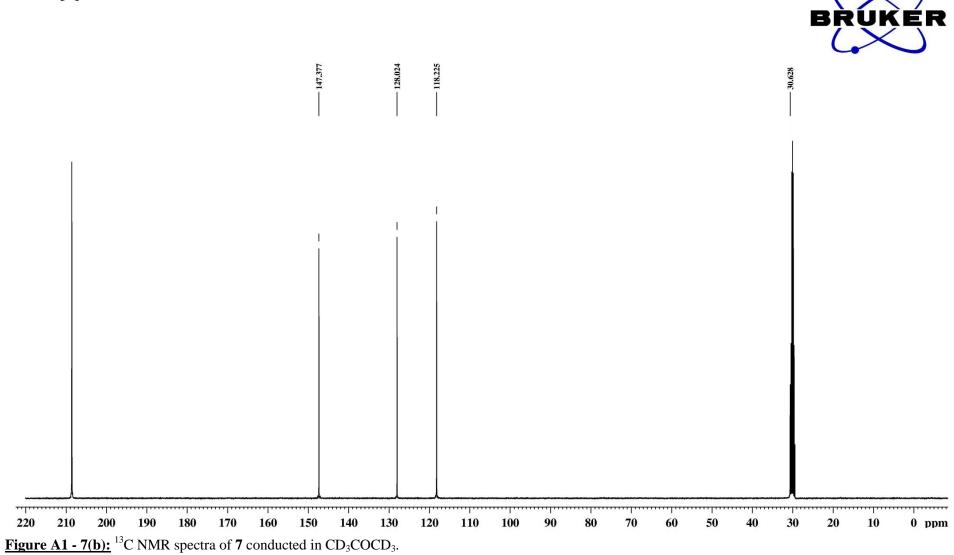




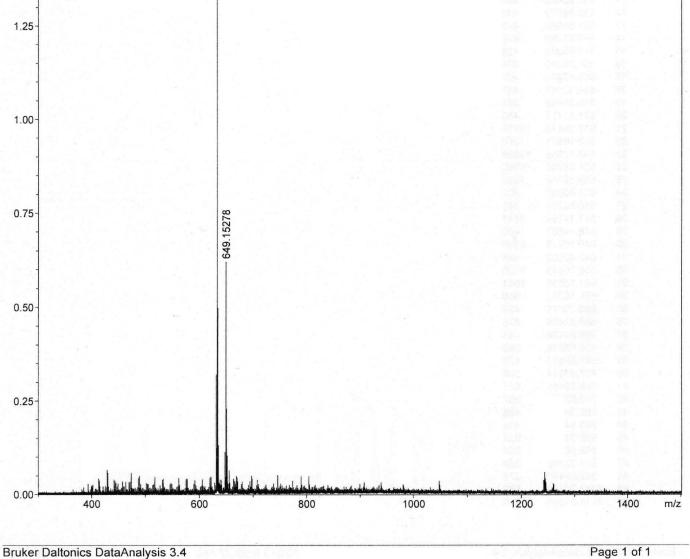
**<u>Figure A1 - 7(a)</u>** <sup>1</sup>H NMR spectra of **7** conducted in  $CD_3COCD_3$ .

20

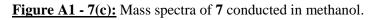


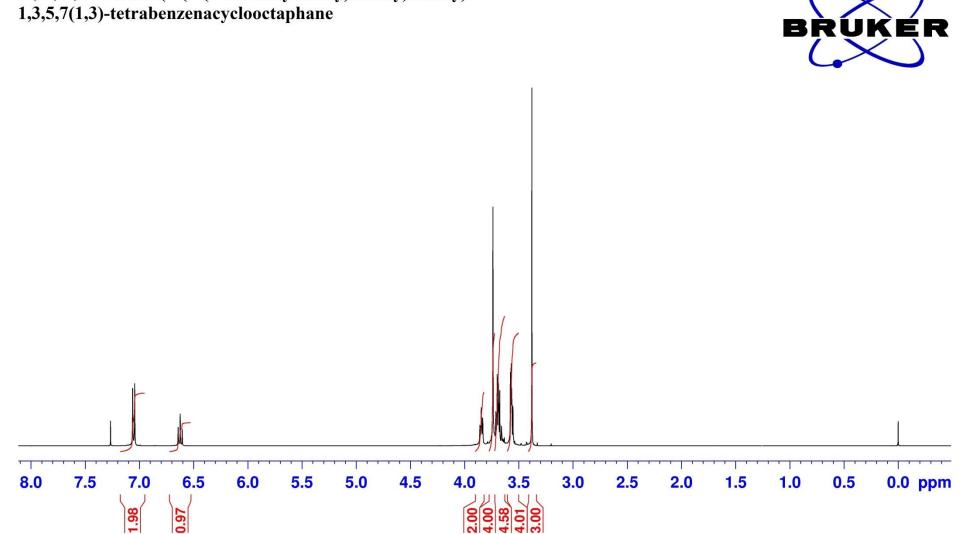


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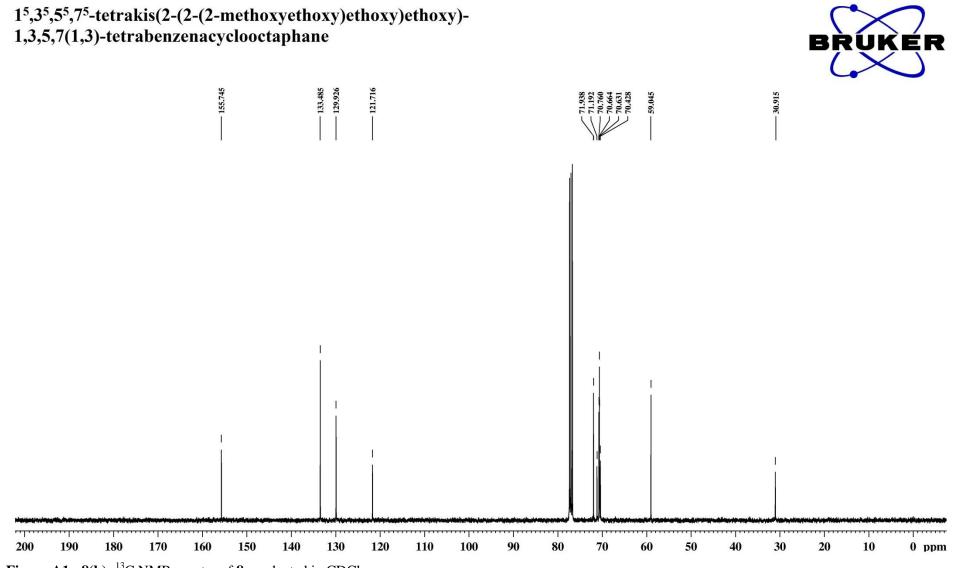
가까? 다 같은 같은 것 같아. 그는 것이 있는 것 같아?





1<sup>5</sup>,3<sup>5</sup>,5<sup>5</sup>,7<sup>5</sup>-tetrakis(2-(2-(2-methoxyethoxy)ethoxy)-

Figure A1 - 8(a): <sup>1</sup>H NMR spectra of 8 conducted in CDCl<sub>3</sub>.



**Figure A1 - 8(b):** <sup>13</sup>C NMR spectra of **8** conducted in CDCl<sub>3</sub>.

# Bruker microTOF University of Brighton

#### Analysis Info



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- TIC +All MS

0.04

Intens. x10<sup>6</sup> 2.50 2.25 2.00

Intens.

x105

4

3.

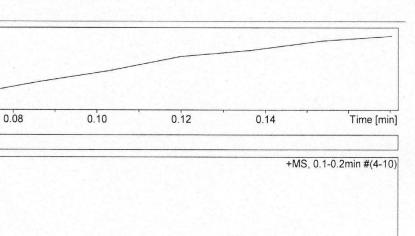
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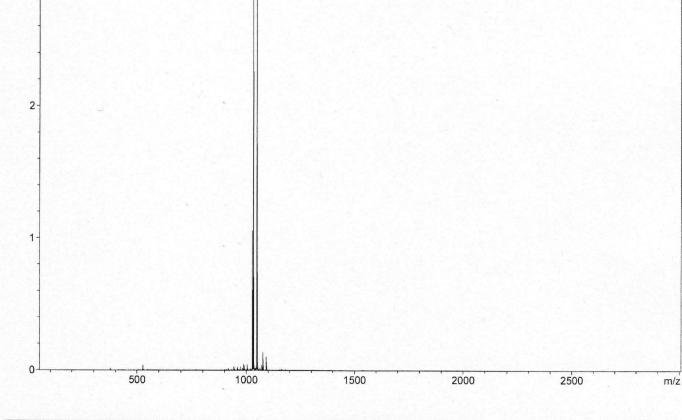
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Operator Instrument admin micrOTOF





Bruker Daltonics DataAnalysis 3.4

Page 1 of 1



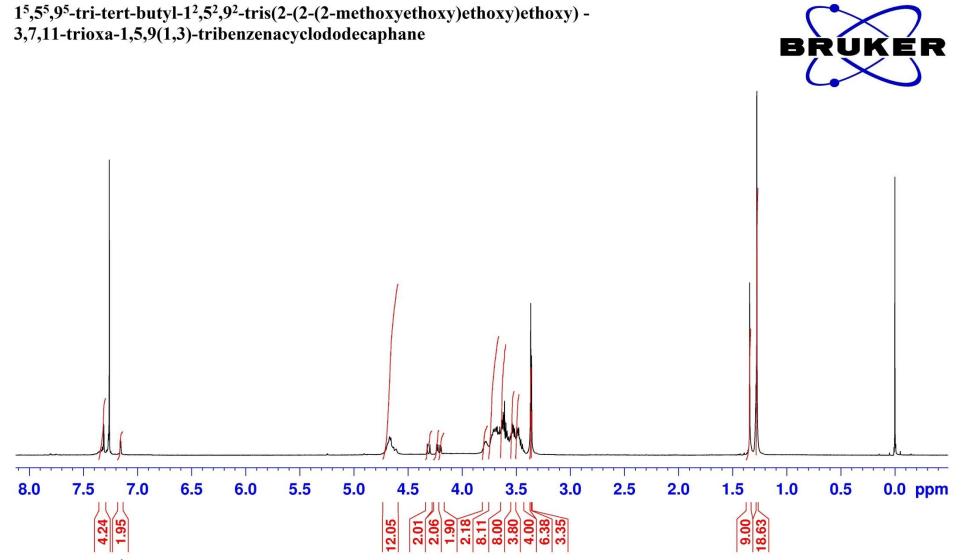
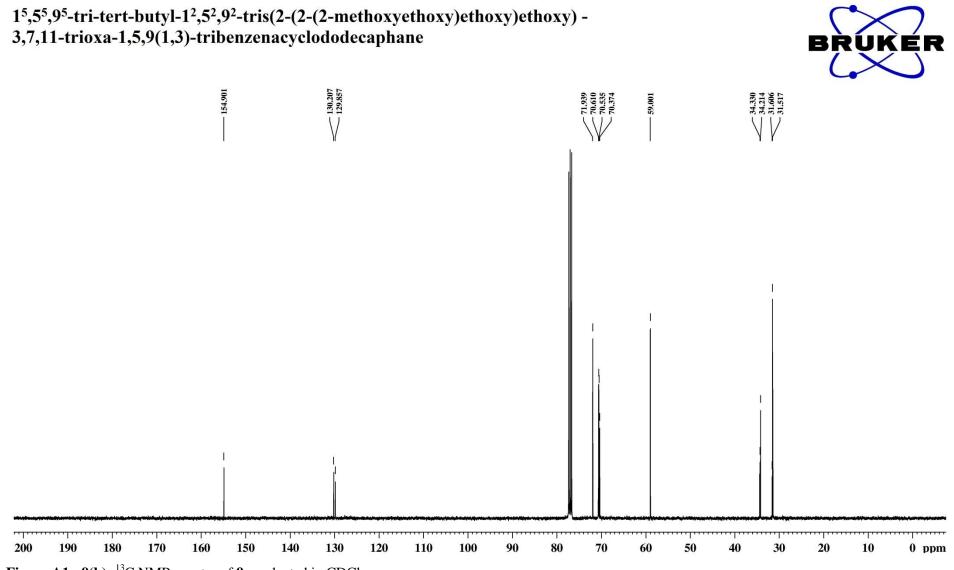


Figure A1 - 9(a): <sup>1</sup>H NMR spectra of 9 conducted in CDCl<sub>3</sub>.



**Figure A1 - 9(b):** <sup>13</sup>C NMR spectra of **9** conducted in CDCl<sub>3</sub>.

### Bruker microTOF

## University of Brighton

#### Analysis Info

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Operator Instrument admin micrOTOF

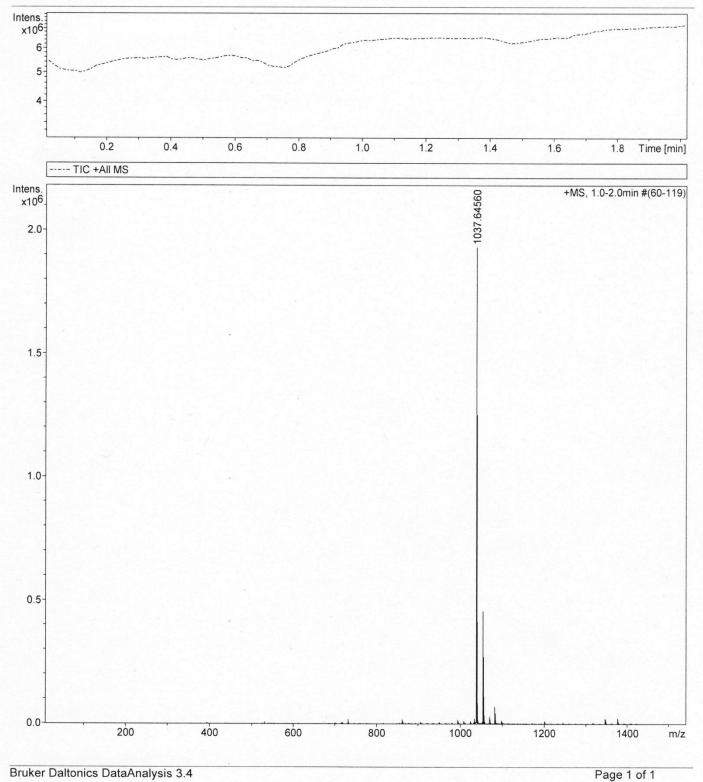


Figure A1 - 9(c): Mass spectra of 9 conducted in methanol.

1<sup>2</sup>,1<sup>5</sup>,3<sup>2</sup>,3<sup>5</sup>,5<sup>2</sup>,5<sup>5</sup>,7<sup>2</sup>,7<sup>5</sup>,9<sup>2</sup>,9<sup>5</sup>-decakis(2-(2-(2-methoxyethoxy)ethoxy) ethoxy)-1,3,5,7,9(1,4)-pentabenzenacyclodecaphane



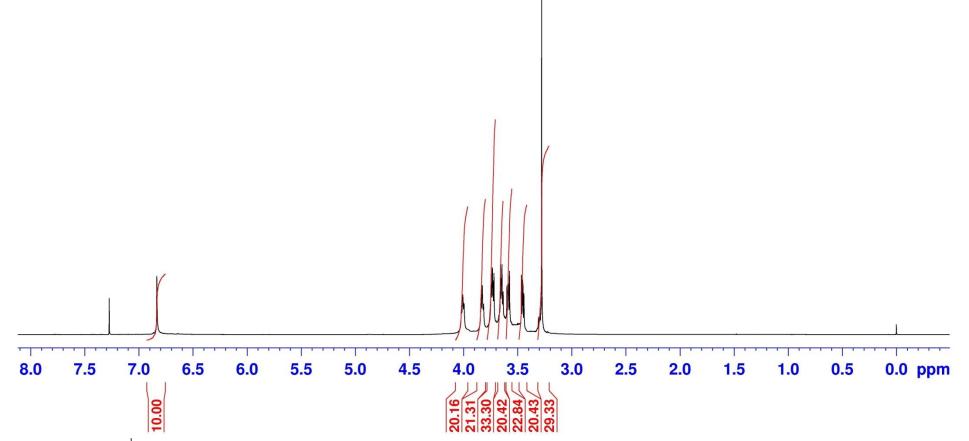
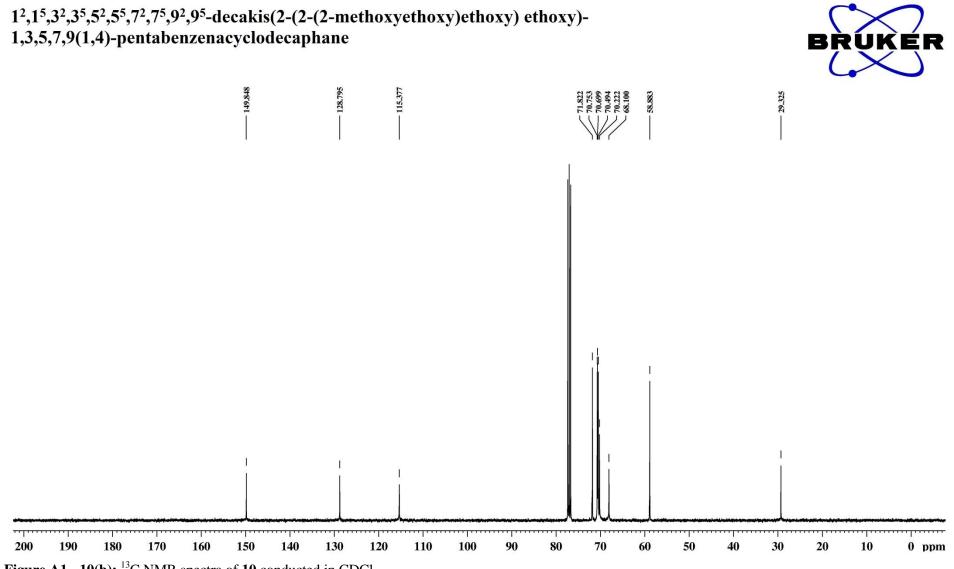


Figure A1 - 10(a): <sup>1</sup>H NMR spectra of 10 conducted in CDCl<sub>3</sub>.



**Figure A1 - 10(b):** <sup>13</sup>C NMR spectra of **10** conducted in CDCl<sub>3</sub>.

## Bruker microTOF

## University of Brighton

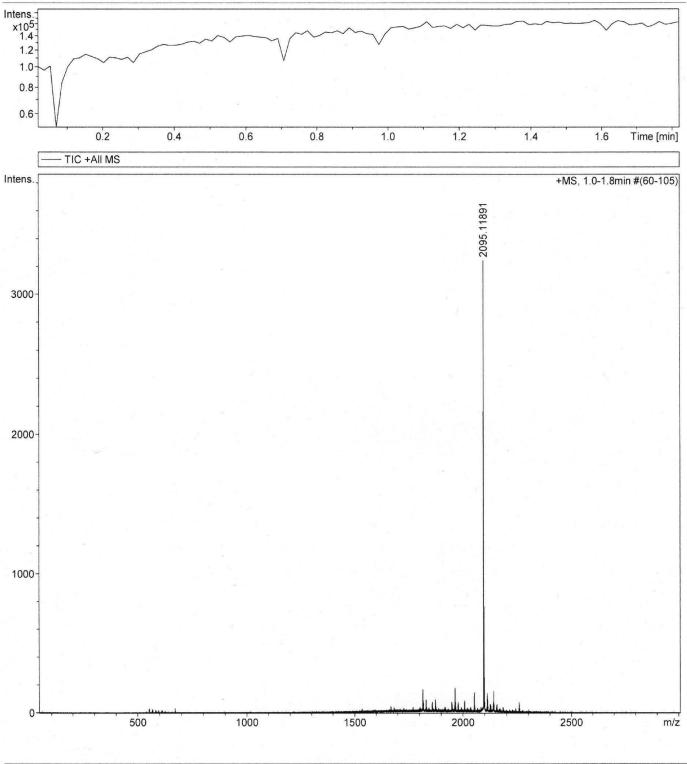


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Operator Instrument 8 11

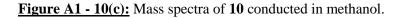
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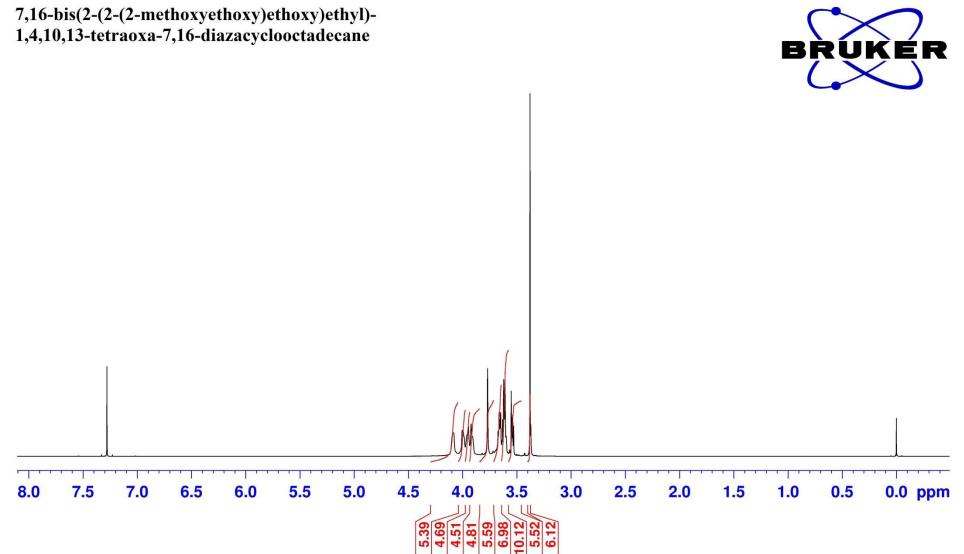
admin micrOTOF



Bruker Daltonics DataAnalysis 3.4

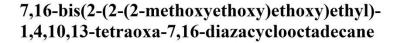
Page 1 of 1



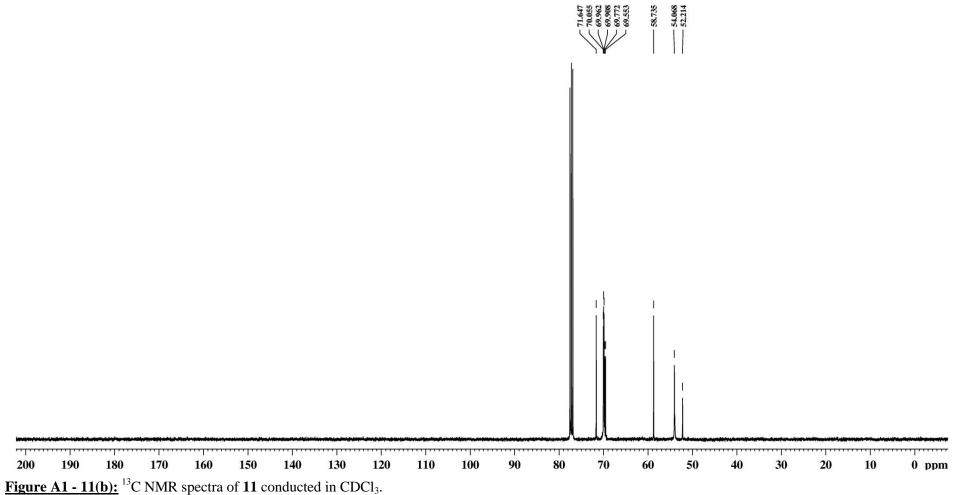


**Figure A1 - 11(a):** <sup>1</sup>H NMR spectra of **11** conducted in CDCl<sub>3</sub>.

32







#### Bruker microTOF

## University of Brighton

#### Analysis Info

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#### Acquisition Date 3/4/2014 6:43:02 PM

Operator Instrument admin micrOTOF

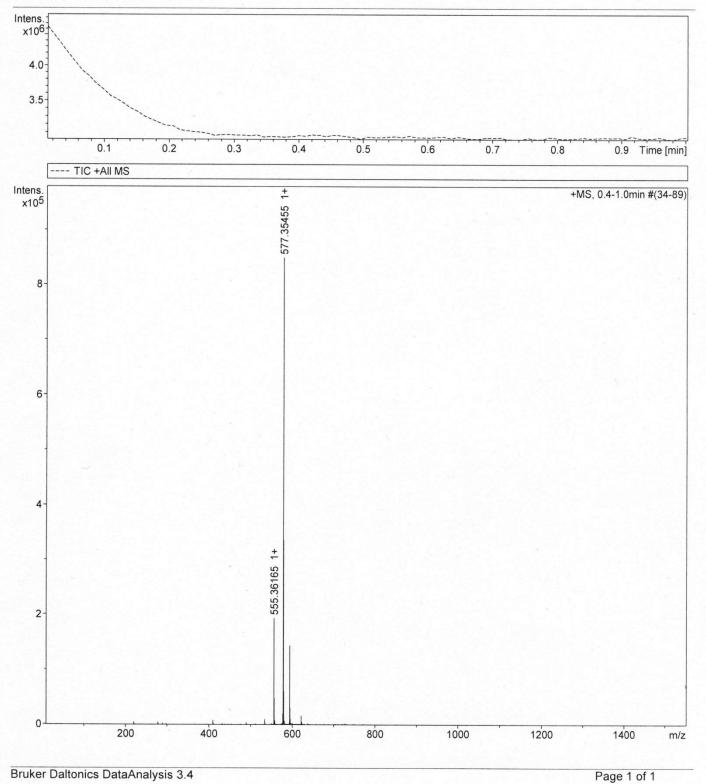


Figure A1 - 11(c): Mass spectra of 11 conducted in methanol.

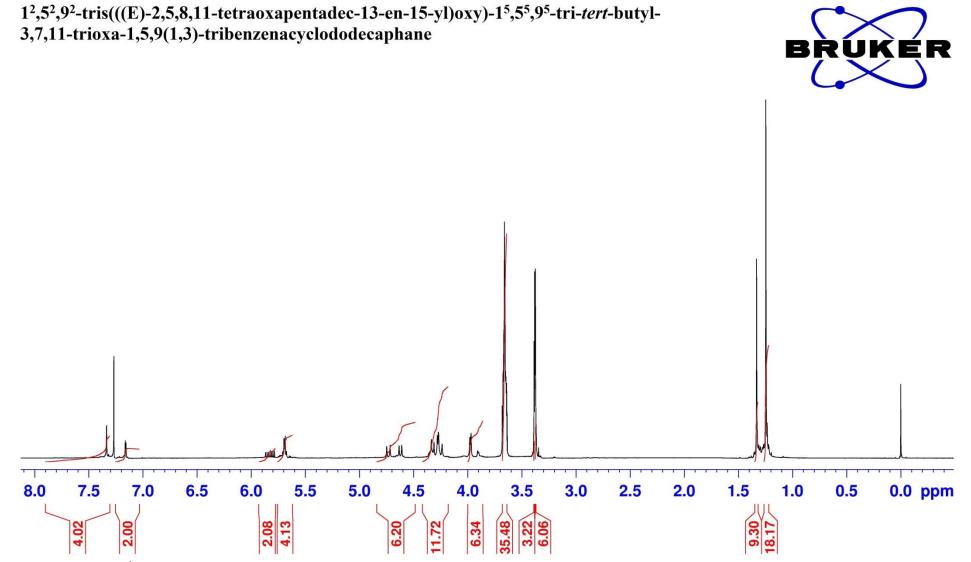
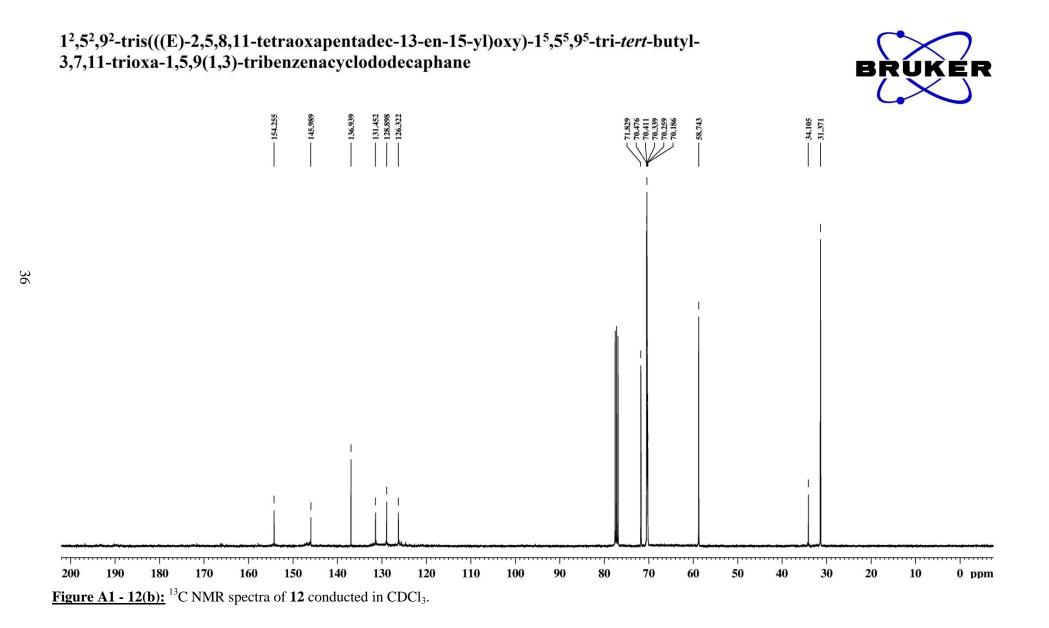
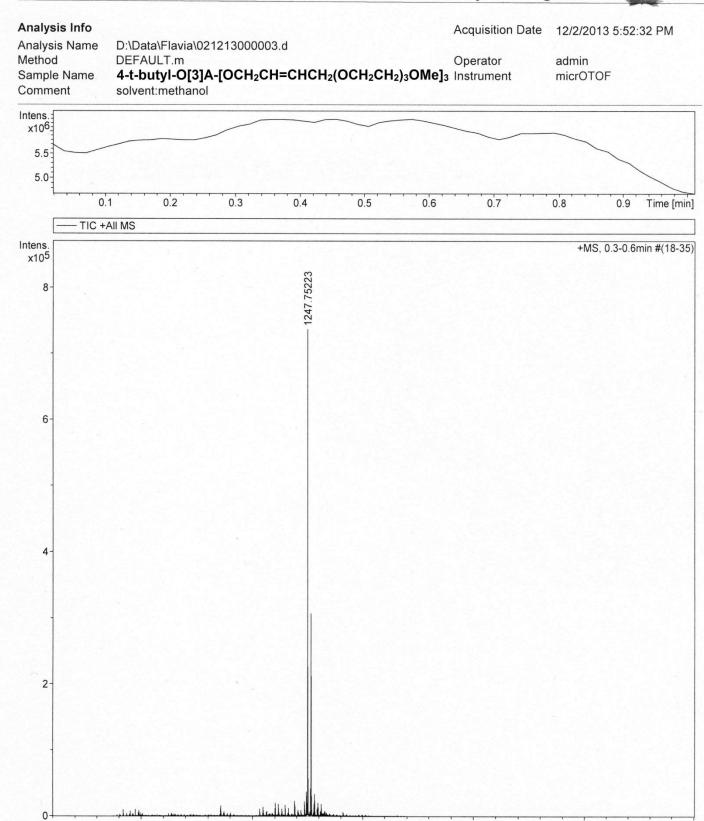


Figure A1 - 12(a): <sup>1</sup>H NMR spectra of 12 conducted in CDCl<sub>3</sub>.



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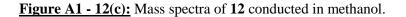
#### Bruker Daltonics DataAnalysis 3.4

500

Page 1 of 1

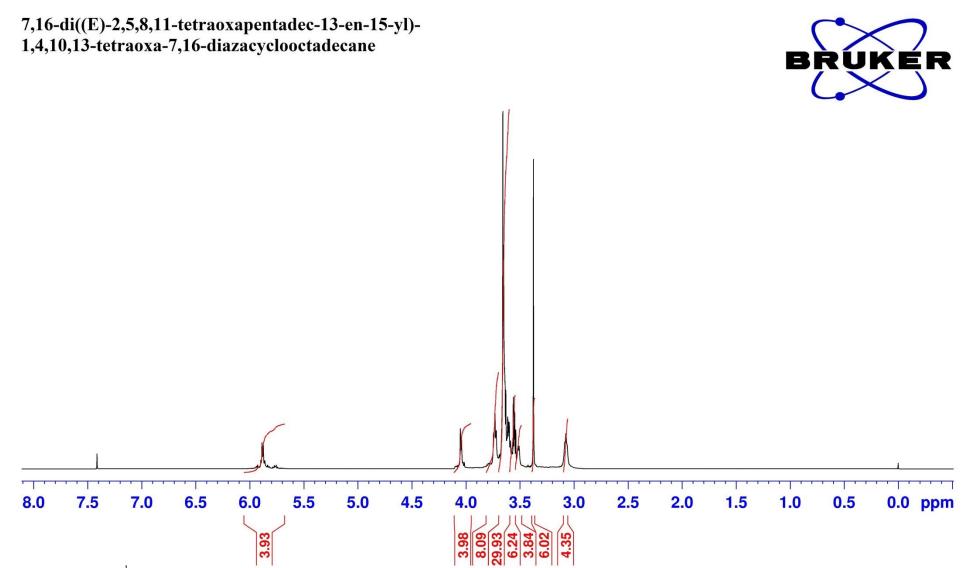
m/z

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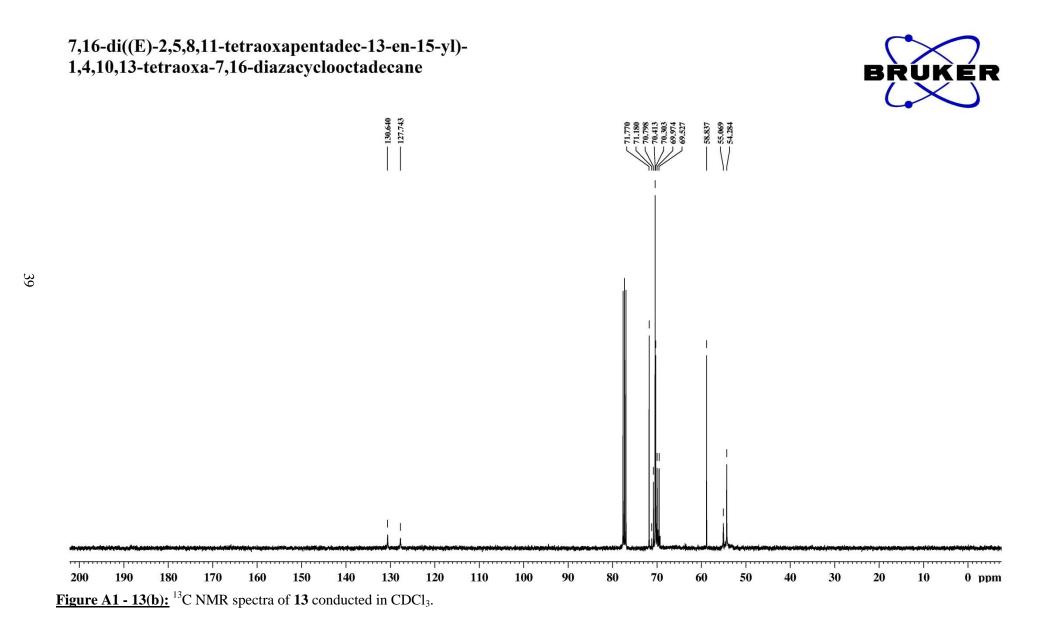


1000

1500



**<u>Figure A1 - 13(a)</u>** <sup>1</sup>H NMR spectra of **13** conducted in CDCl<sub>3</sub>.



# University of Brighton

### Analysis Info

Analysis Name Method Sample Name Comment

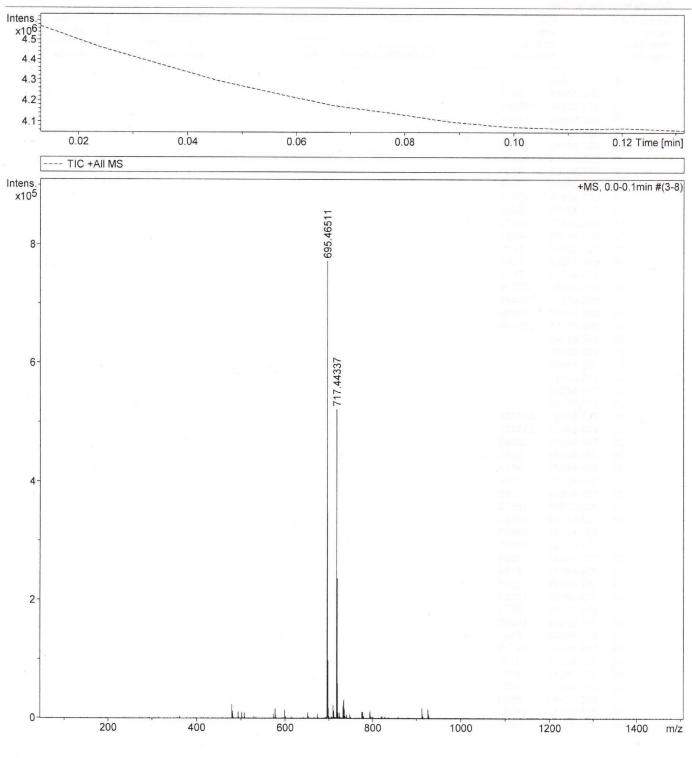
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Acquisition Date 12/11/2013 3:57:36 PM

Operator

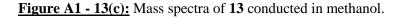
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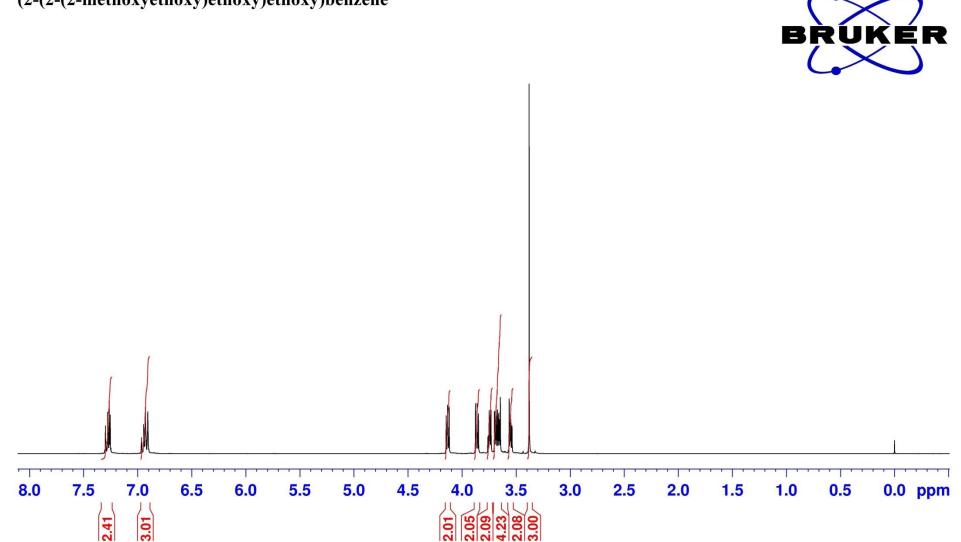
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Bruker Daltonics DataAnalysis 3.4

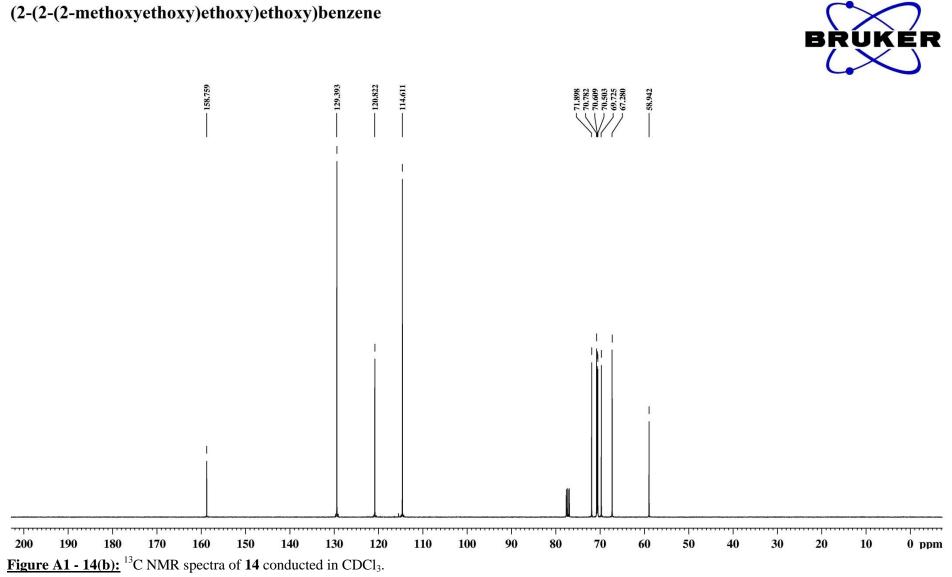
Page 1 of 1

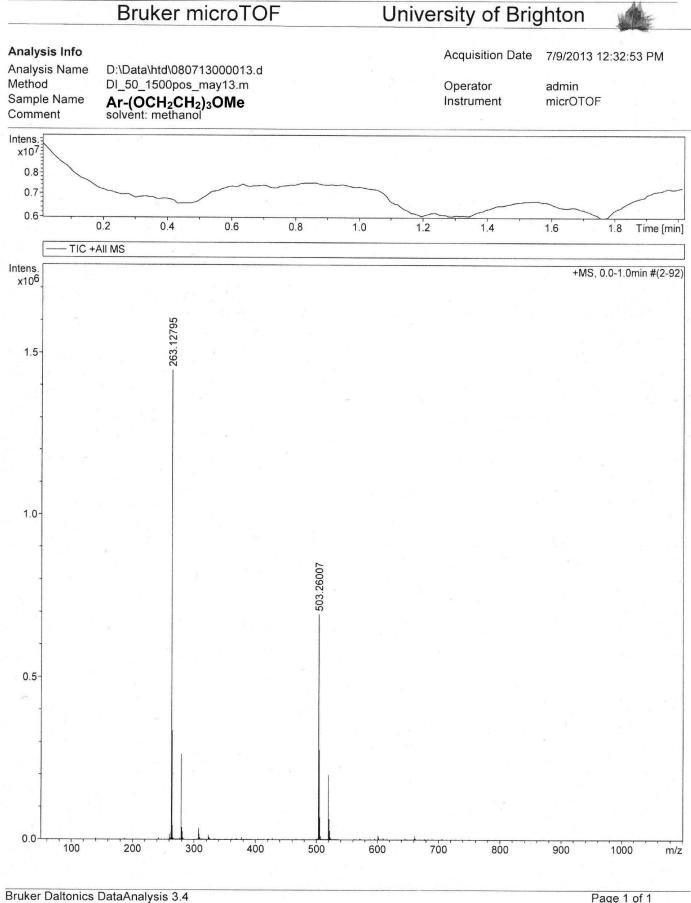




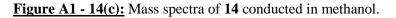
(2-(2-(2-methoxy)ethoxy)ethoxy)benzene

Figure A1 - 14(a): <sup>1</sup>H NMR spectra of 14 conducted in CDCl<sub>3</sub>.





Page 1 of 1



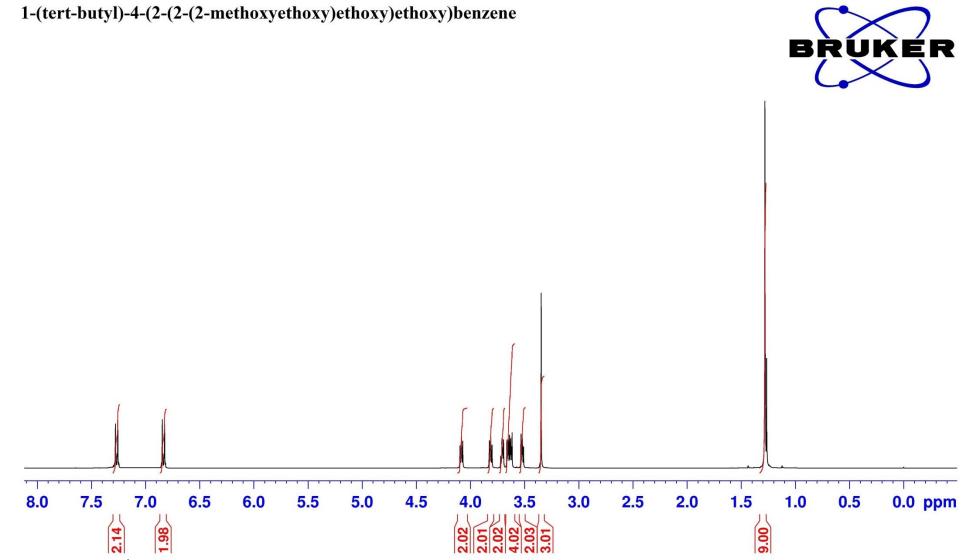
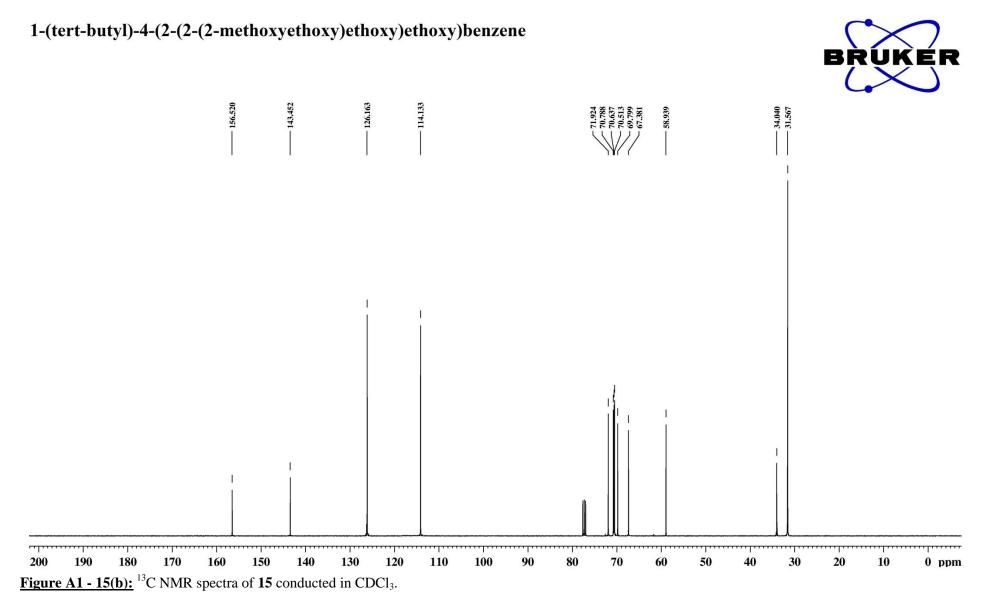


Figure A1 - 15(a): <sup>1</sup>H NMR spectra of 15 conducted in CDCl<sub>3</sub>.



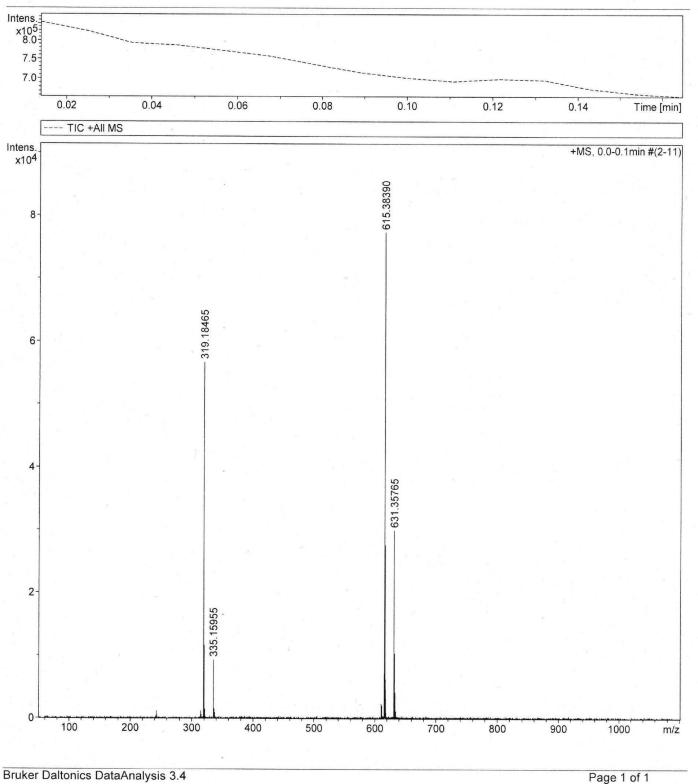
# University of Brighton

### Analysis Info

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#### Acquisition Date 7/24/2013 3:22:35 PM

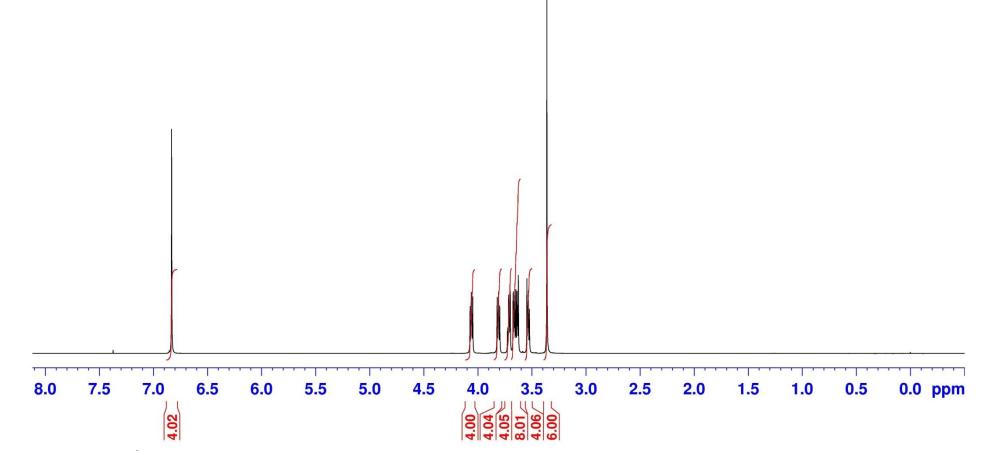
Operator Instrument admin micrOTOF



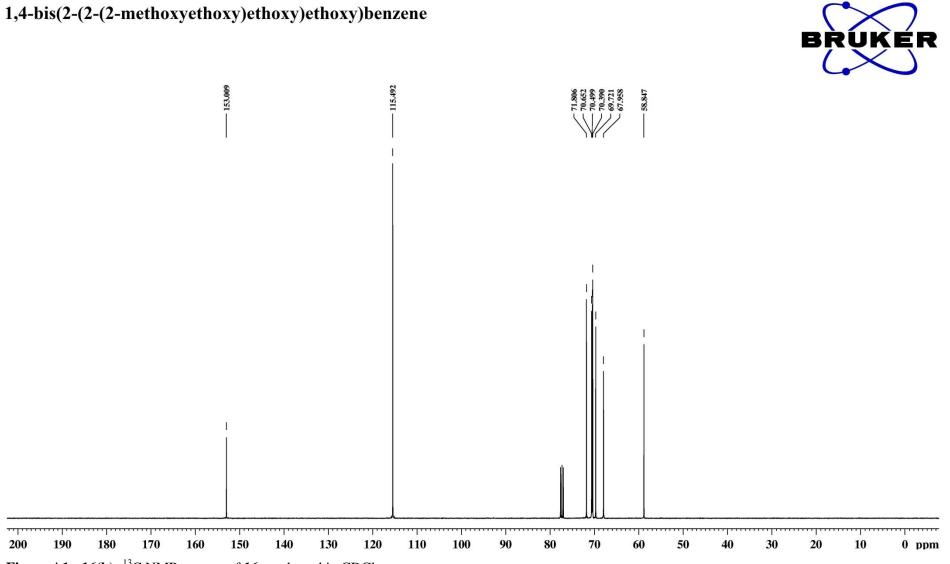




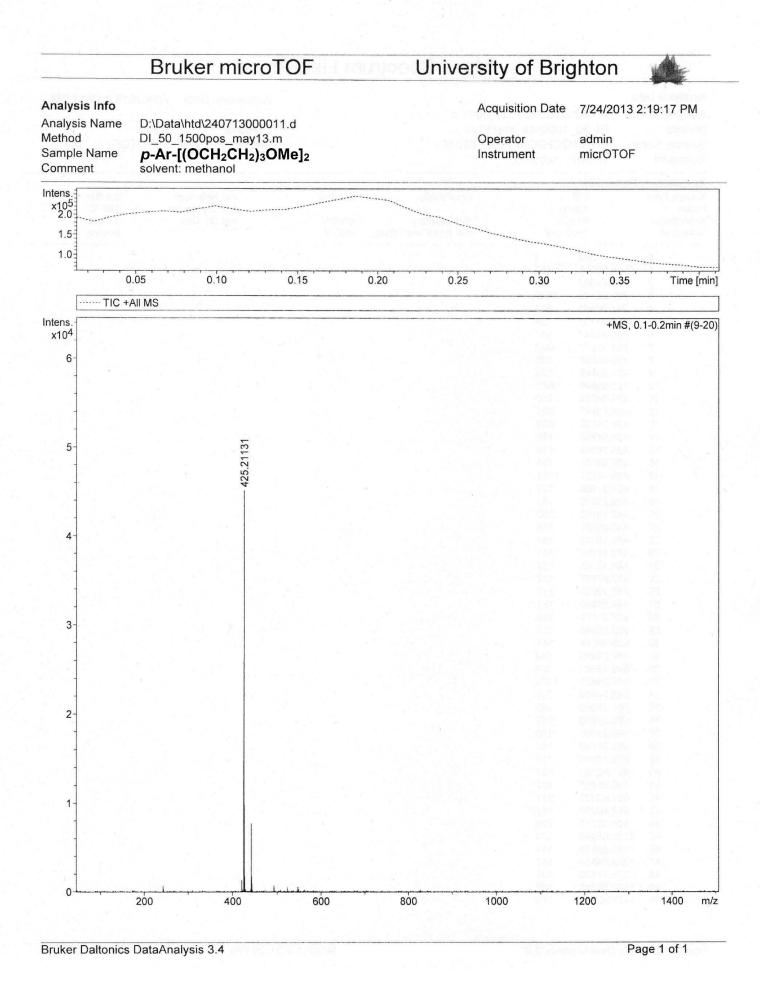
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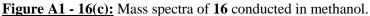


**Figure A1 - 16(a):** <sup>1</sup>H NMR spectra of **16** conducted in CDCl<sub>3</sub>.



**Figure A1 - 16(b):** <sup>13</sup>C NMR spectra of **16** conducted in CDCl<sub>3</sub>.





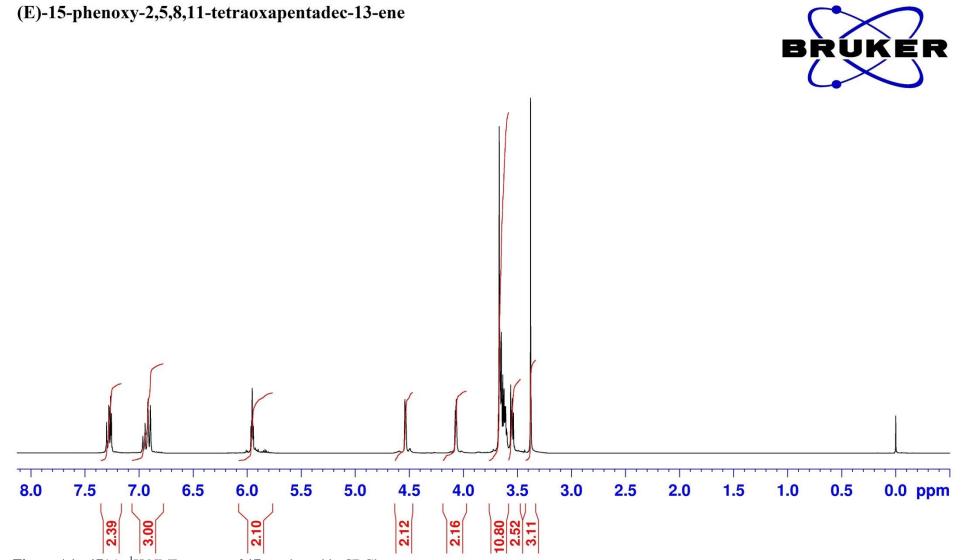


Figure A1 - 17(a): <sup>1</sup>H NMR spectra of 17 conducted in CDCl<sub>3</sub>.

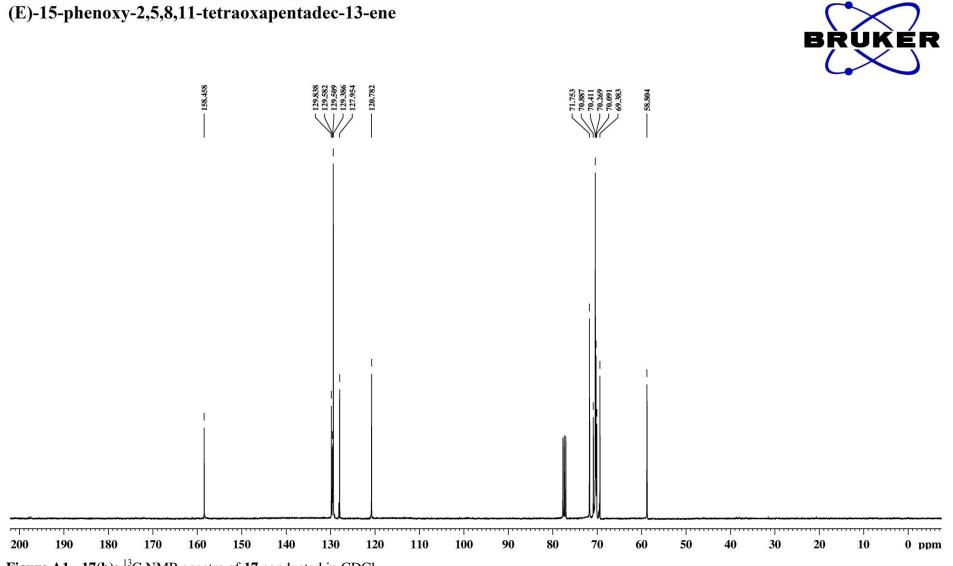


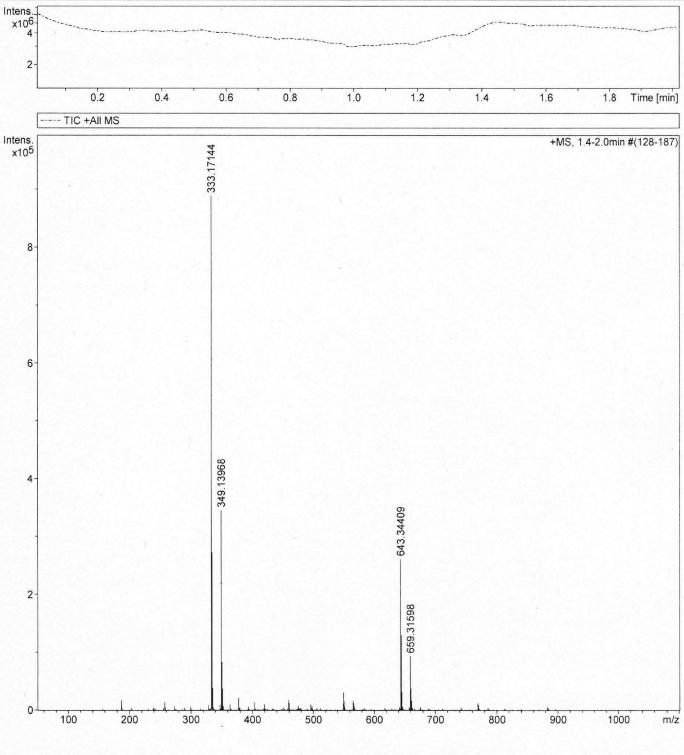
Figure A1 - 17(b): <sup>13</sup>C NMR spectra of 17 conducted in CDCl<sub>3</sub>.

# University of Brighton

#### Analysis Info

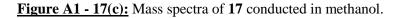
Analysis Name Method Sample Name Comment D:\Data\htd\050713000008.d DI\_50\_1500pos\_may13.m **Ar-OCH<sub>2</sub>CH=CHCH<sub>2</sub>(OCH<sub>2</sub>CH<sub>2</sub>)<sub>3</sub>OMe** solvent:methanol Acquisition Date 7/5/2013 4:01:55 PM

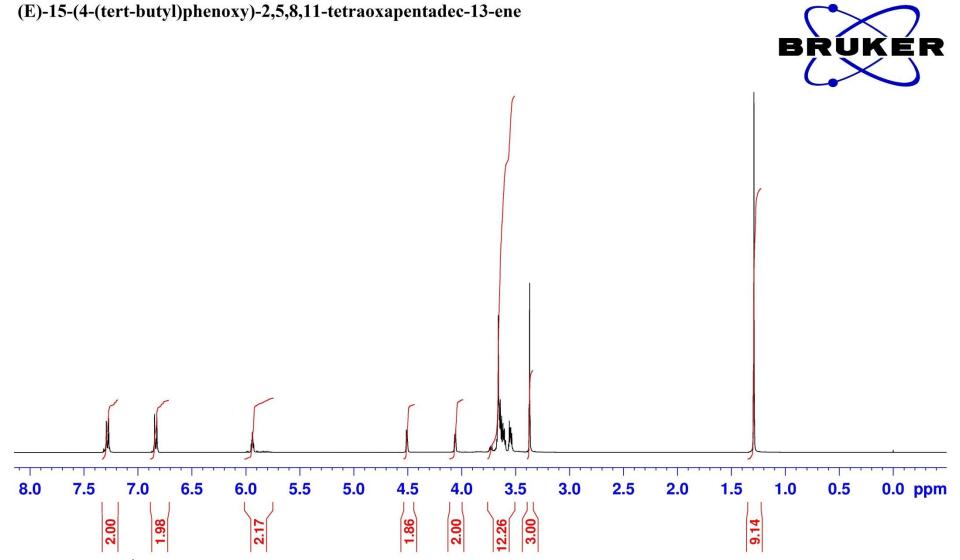
Operator Instrument admin micrOTOF



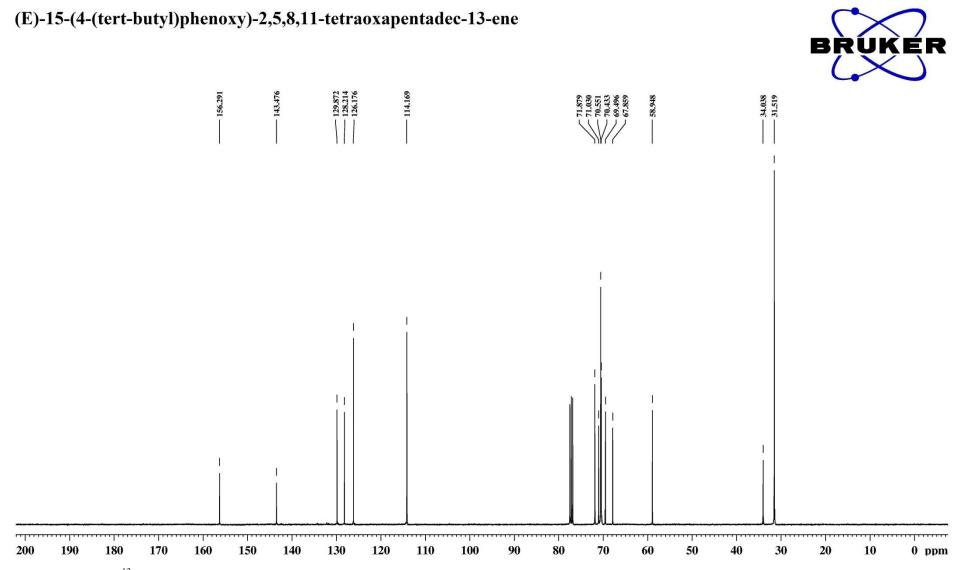
Bruker Daltonics DataAnalysis 3.4

Page 1 of 1



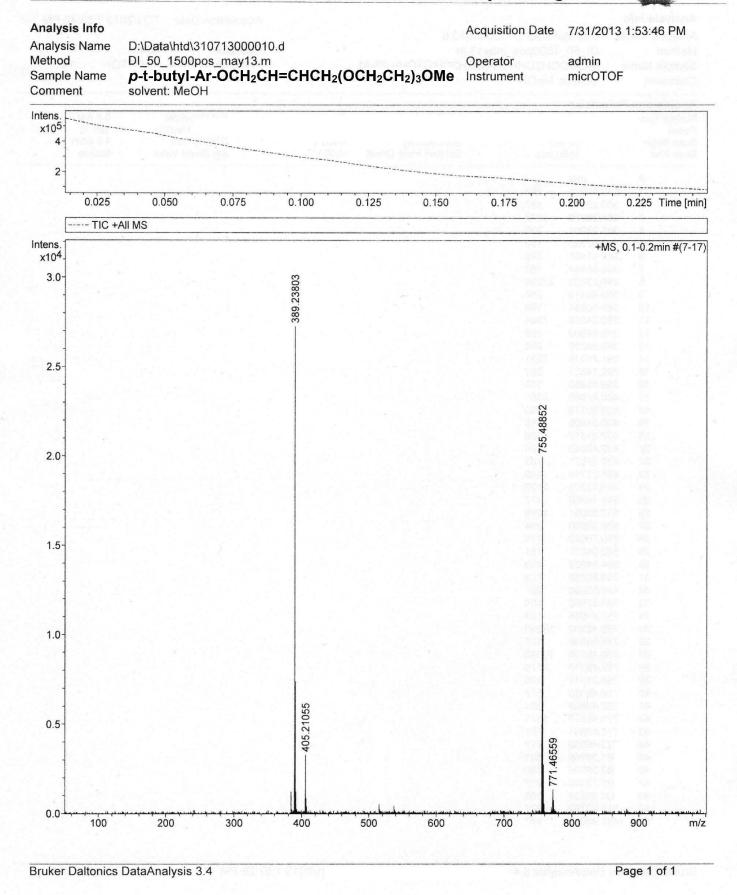


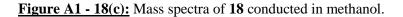
**Figure A1 - 18(a):** <sup>1</sup>H NMR spectra of **18** conducted in CDCl<sub>3</sub>.

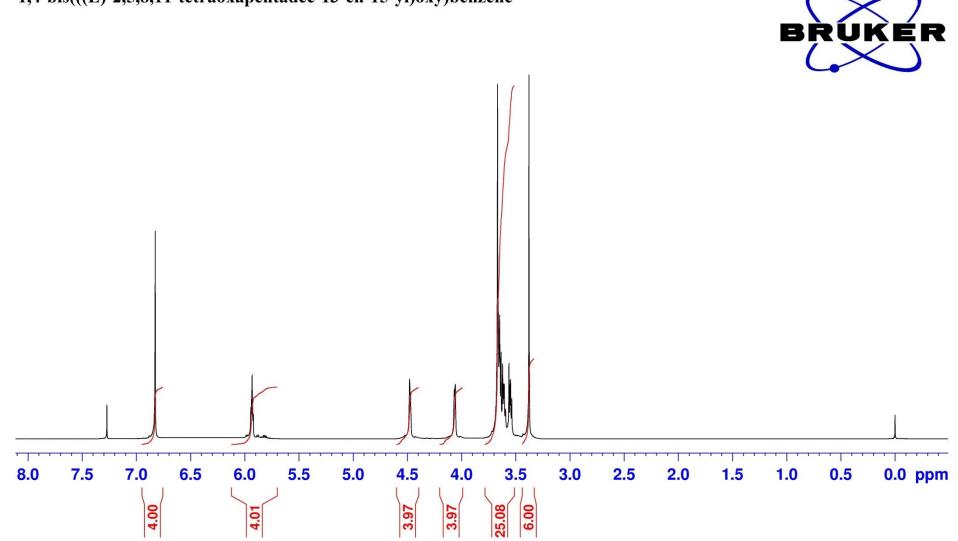


**Figure A1 - 18(b):** <sup>13</sup>C NMR spectra of **18** conducted in CDCl<sub>3</sub>.

# University of Brighton

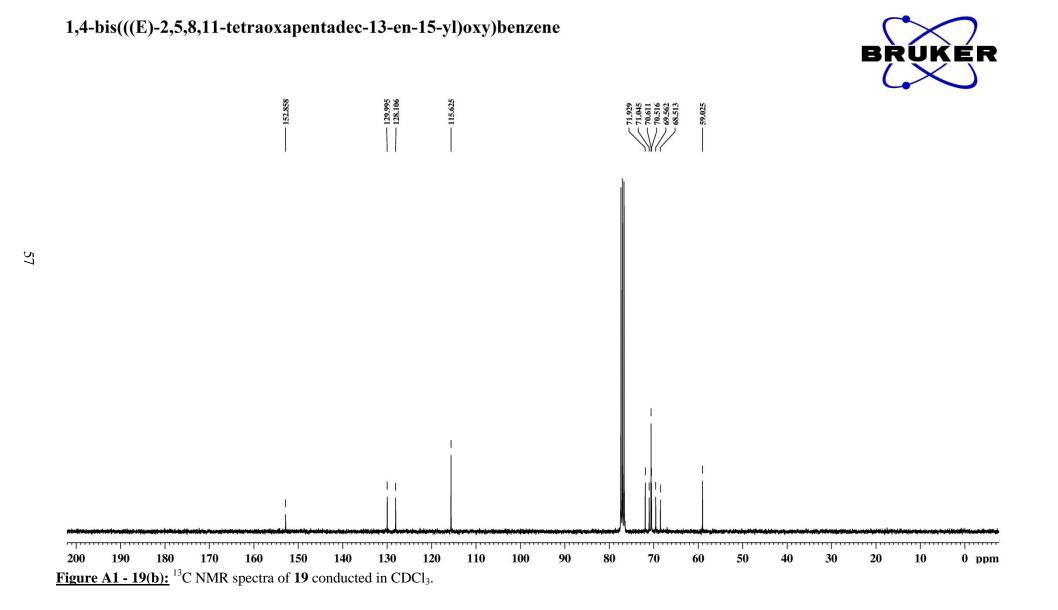






### 1,4-bis(((E)-2,5,8,11-tetraoxapentadec-13-en-15-yl)oxy)benzene

Figure A1 - 19(a): <sup>1</sup>H NMR spectra of 19 conducted in CDCl<sub>3</sub>.



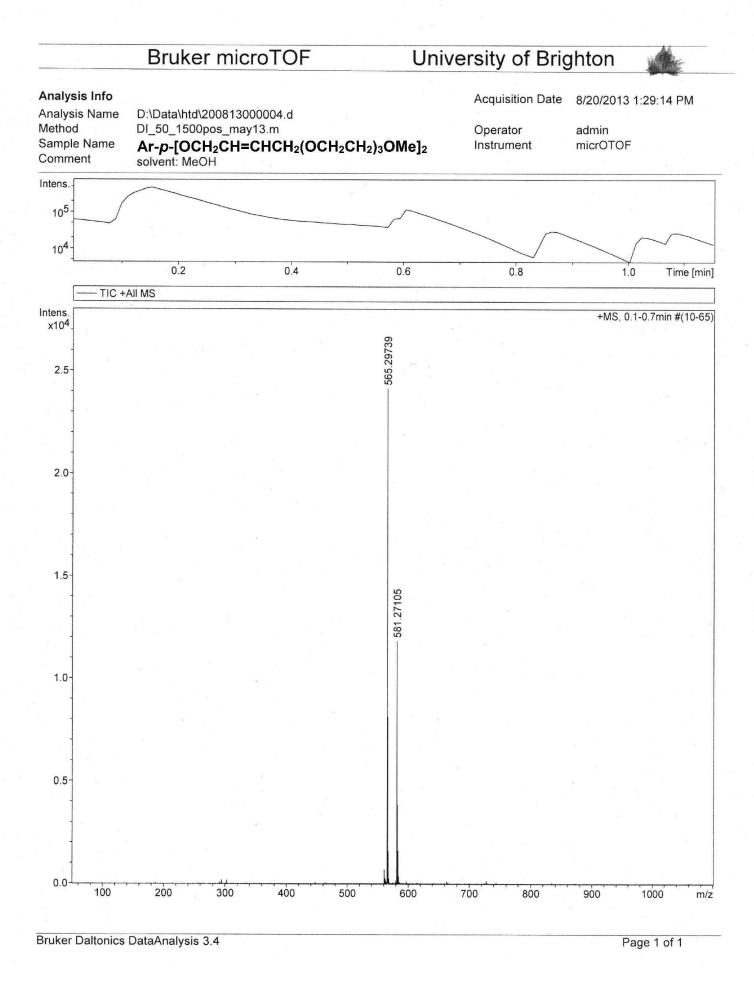
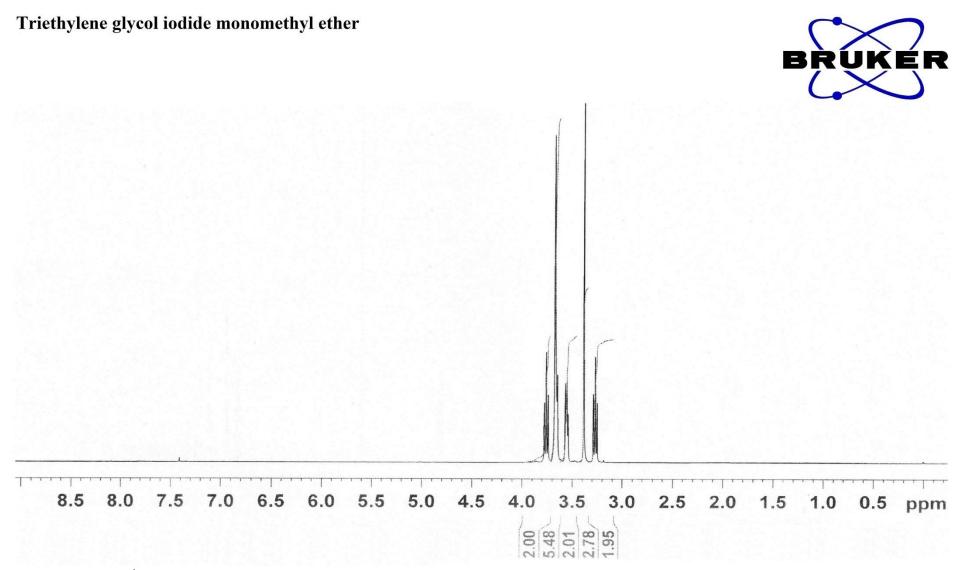
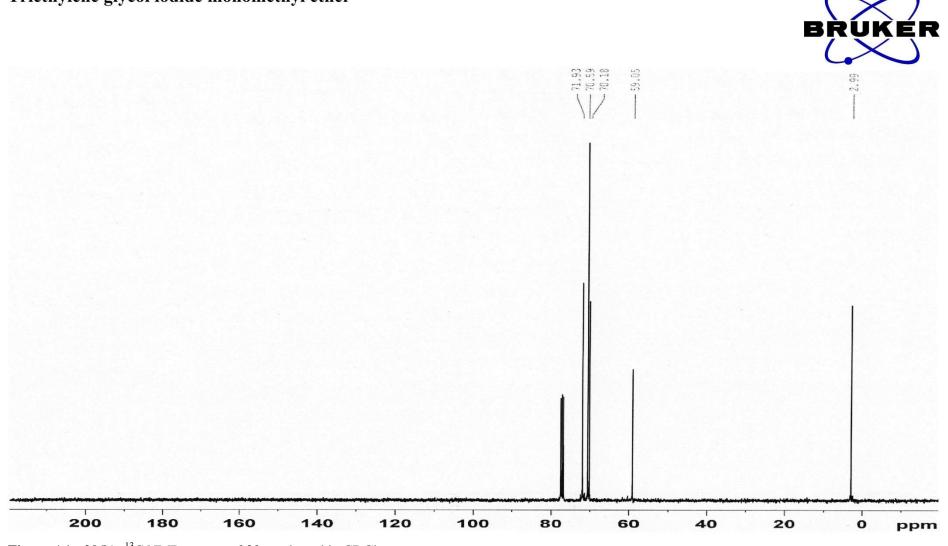


Figure A1 - 19(c): Mass spectra of 19 conducted in methanol.



**Figure A1 - 20(a):** <sup>1</sup>H NMR spectra of **20** conducted in CDCl<sub>3</sub>.



Triethylene glycol iodide monomethyl ether

**Figure A1 - 20(b):** <sup>13</sup>C NMR spectra of **20** conducted in CDCl<sub>3</sub>.

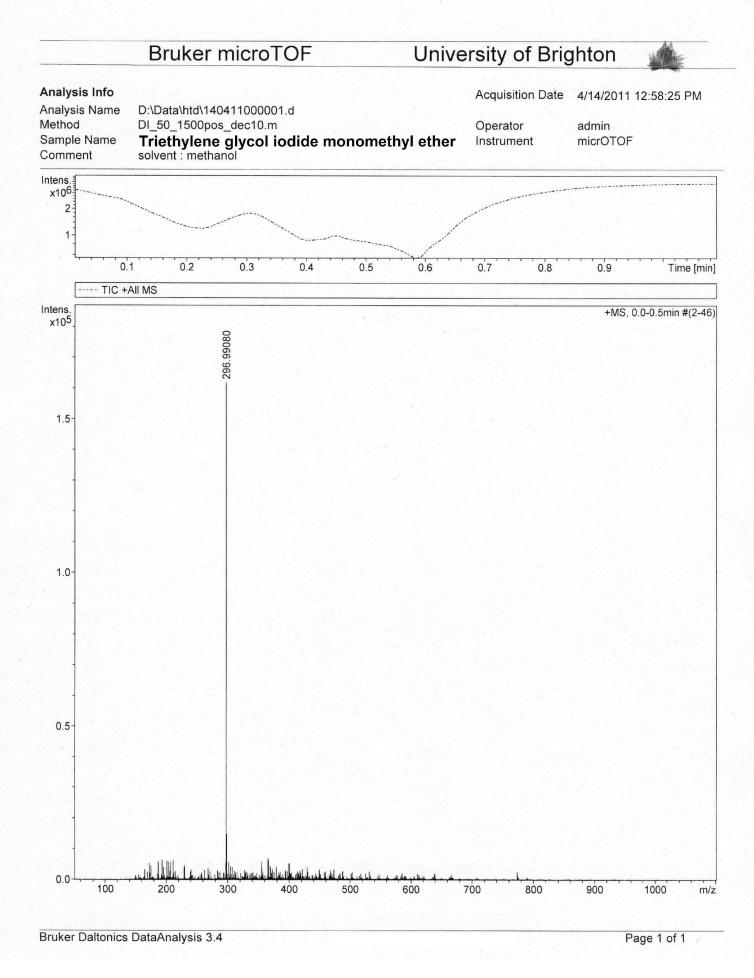
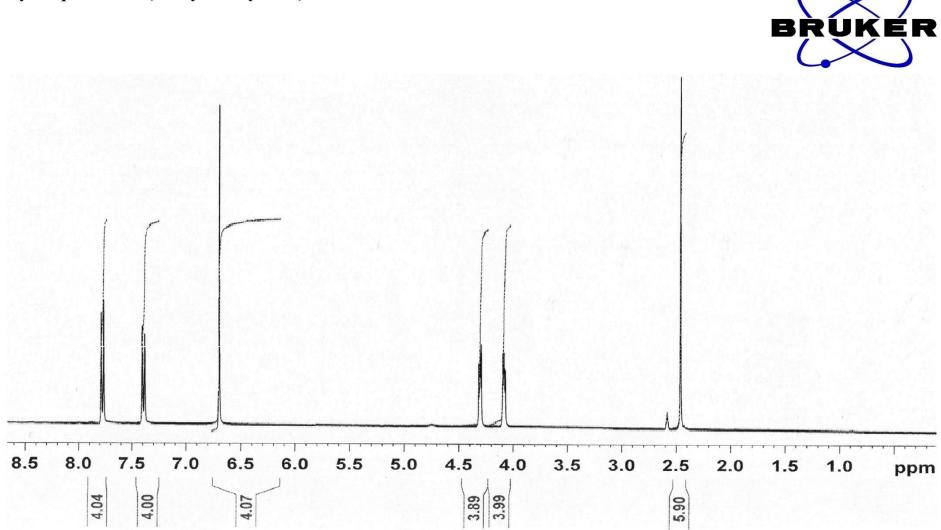
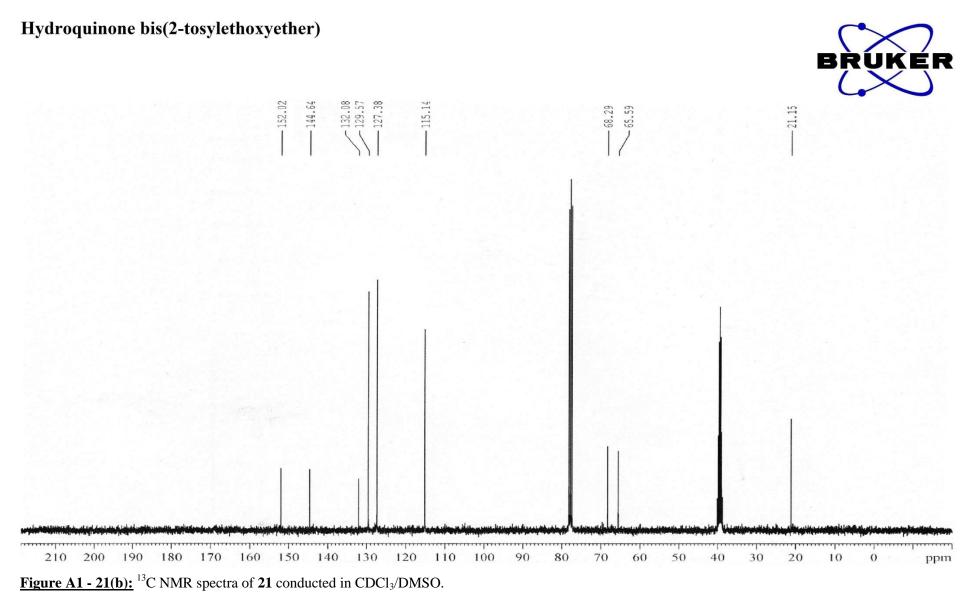


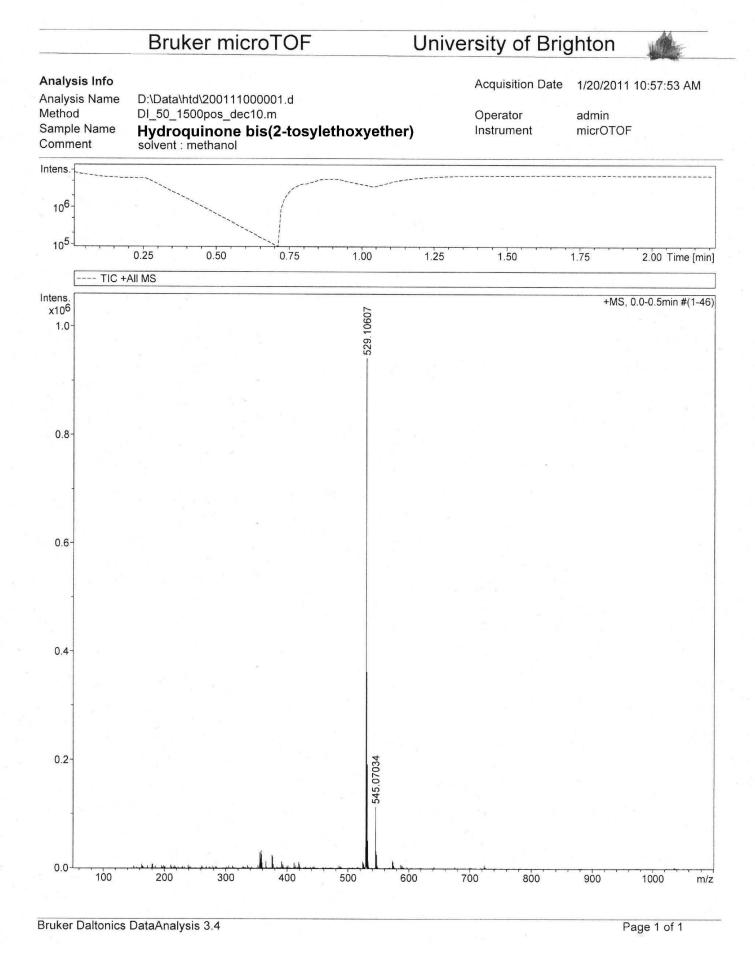
Figure A1 - 20(c): Mass spectra of 20 conducted in methanol.

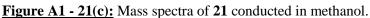


Hydroquinone bis(2-tosylethoxyether)

**Figure A1 - 21(a):** <sup>13</sup>C NMR spectra of **21** conducted in CDCl<sub>3</sub>/DMSO.

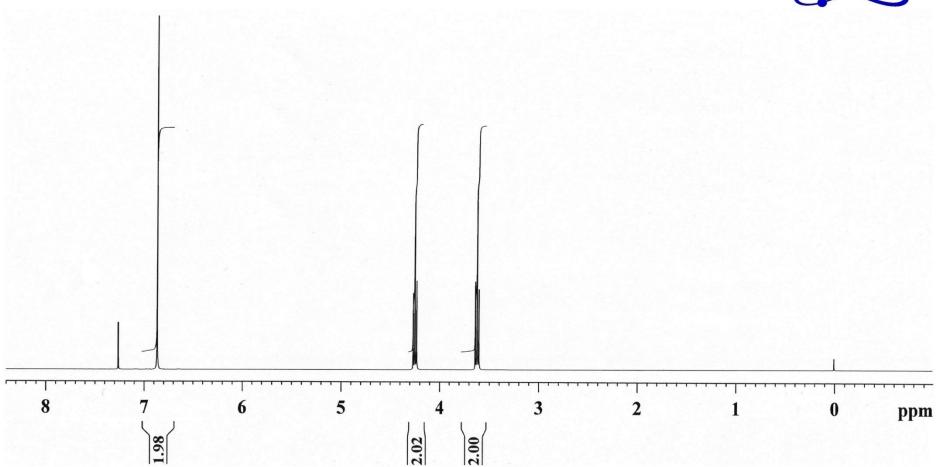






1,4-Bis(2-bromoethoxy)benzene

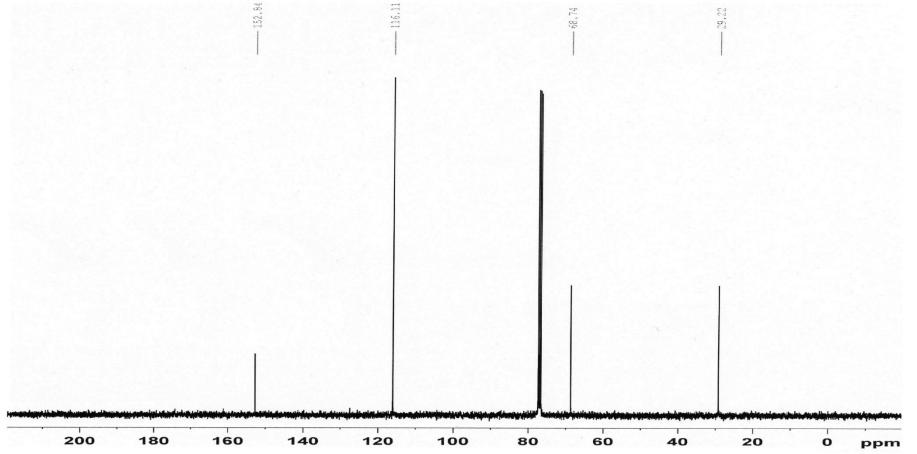




**Figure A1 - 22(a):** <sup>1</sup>H NMR spectra of **22** conducted in CDCl<sub>3</sub>.

1,4-Bis(2-bromoethoxy)benzene







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Analysis Name Method Sample Name Comment D:\Data\htd\061011000001.d DI\_50\_500pos\_jily11.m **1,4-Bis(2-bromoethoxy)benzene** solvent: methanol Acquisition Date

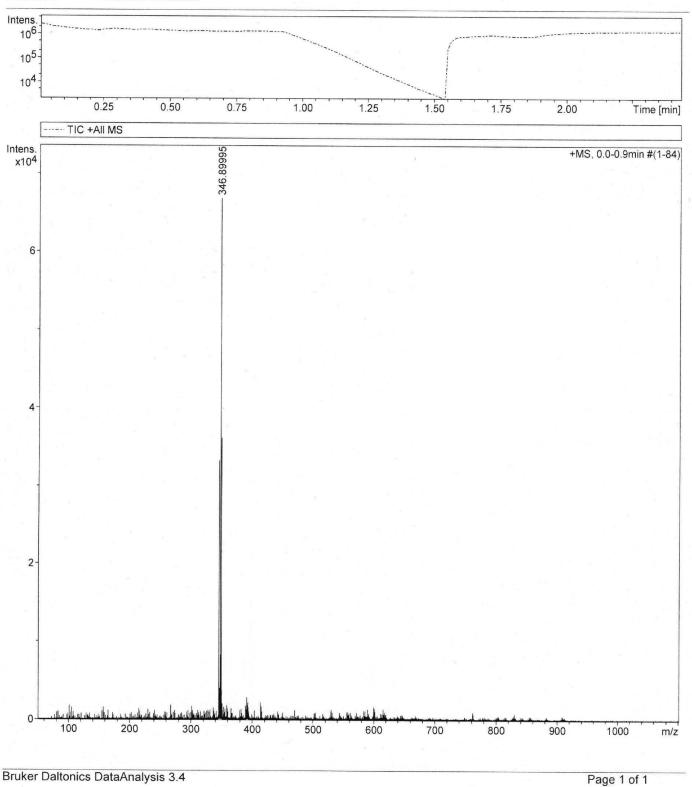
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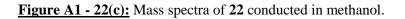
Instrument

admin

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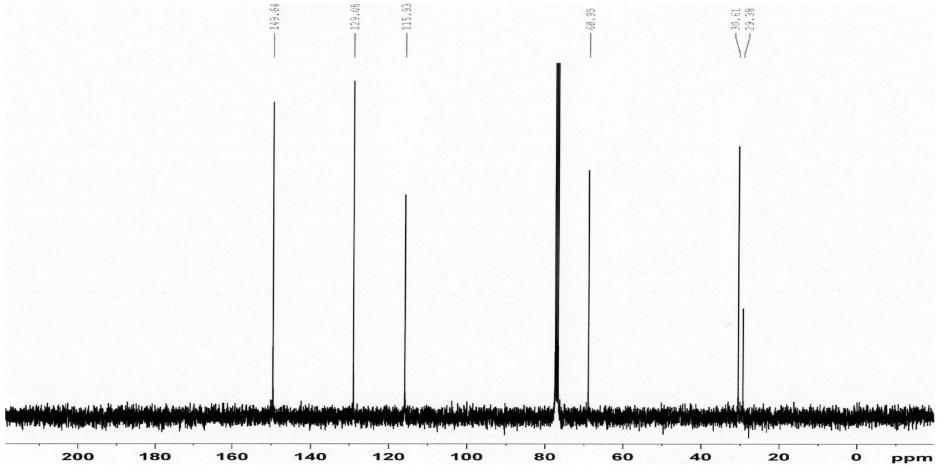
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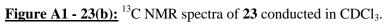
<sup>1,4-</sup>Bis(2-bromoethoxy)pillar[5]arene

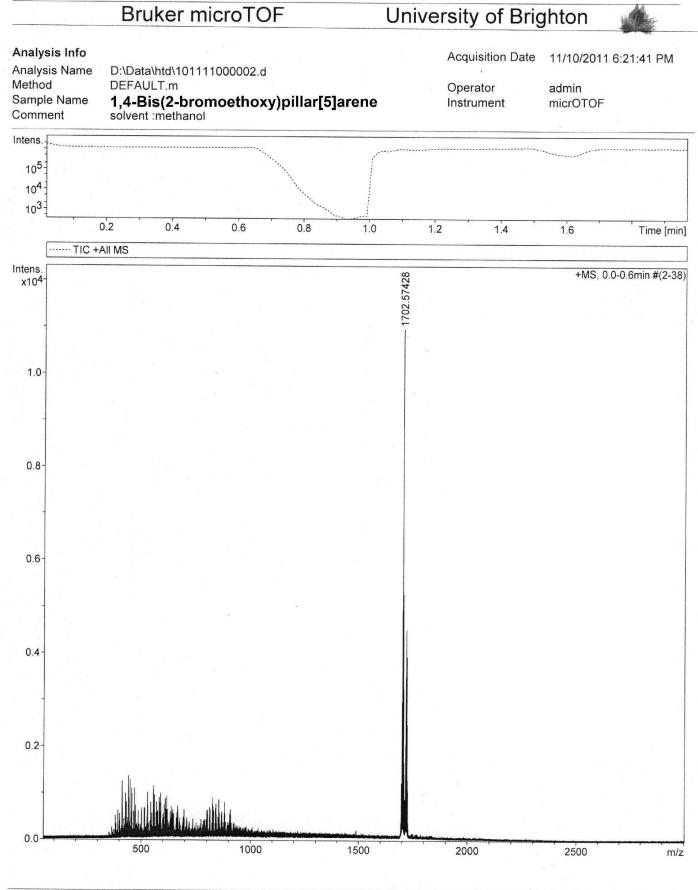
Figure A1 - 23(a): <sup>1</sup>H NMR spectra of 23 conducted in CDCl<sub>3</sub>.

1,4-Bis(2-bromoethoxy)pillar[5]arene



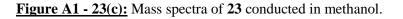


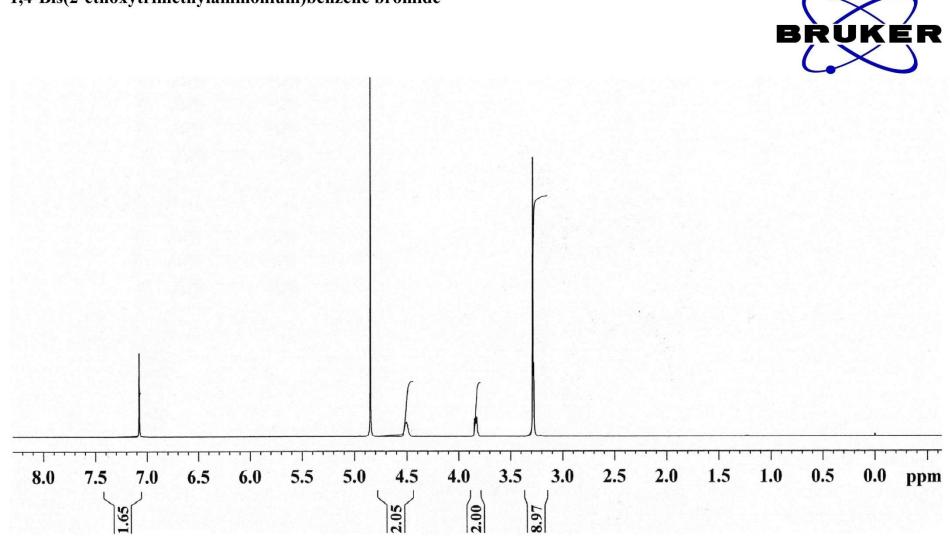




Bruker Daltonics DataAnalysis 3.4

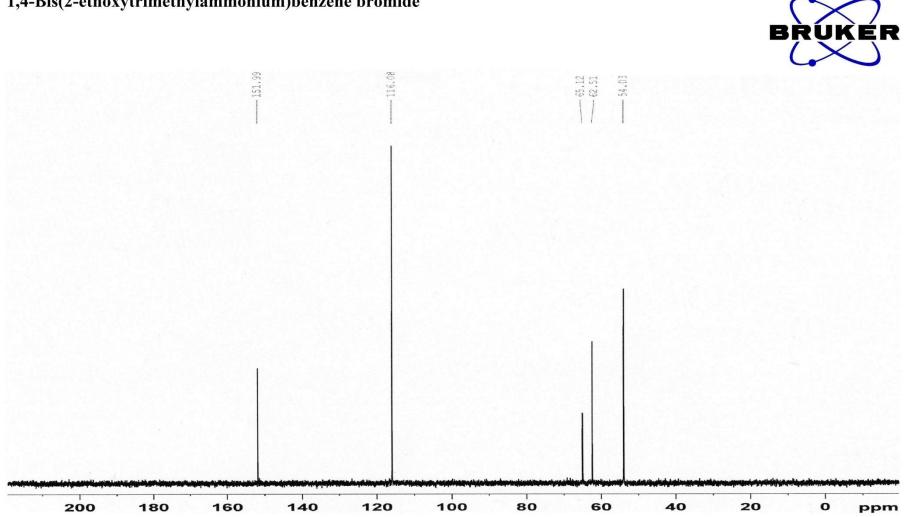
Page 1 of 1





<sup>1,4-</sup>Bis(2-ethoxytrimethylammonium)benzene bromide

**Figure A1 - 24(a):** <sup>1</sup>H NMR spectra of **24** conducted in  $D_2O$ .



1,4-Bis(2-ethoxytrimethylammonium)benzene bromide

**<u>Figure A1 - 24(b)</u>**: <sup>13</sup>C NMR spectra of **24** conducted in  $D_2O$ .

### Bruker microTOF

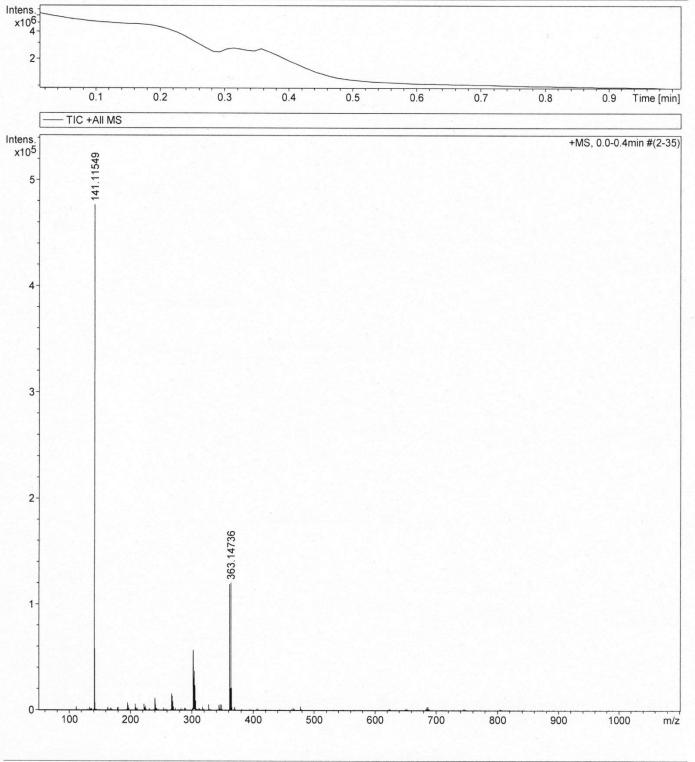
## University of Brighton

#### Analysis Info

Analysis Name Method Sample Name Comment D:\Data\htd\072111000012.d DI\_50\_1500pos\_jily11.m **Ar-p-[OCH<sub>2</sub>CH<sub>2</sub>N<sup>+</sup>(CH<sub>3</sub>)<sub>3</sub>Br<sup>-</sup>]**<sub>2</sub> solvent:Methanol Acquisition Date

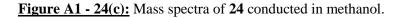
Operator Instrument admin micrOTOF

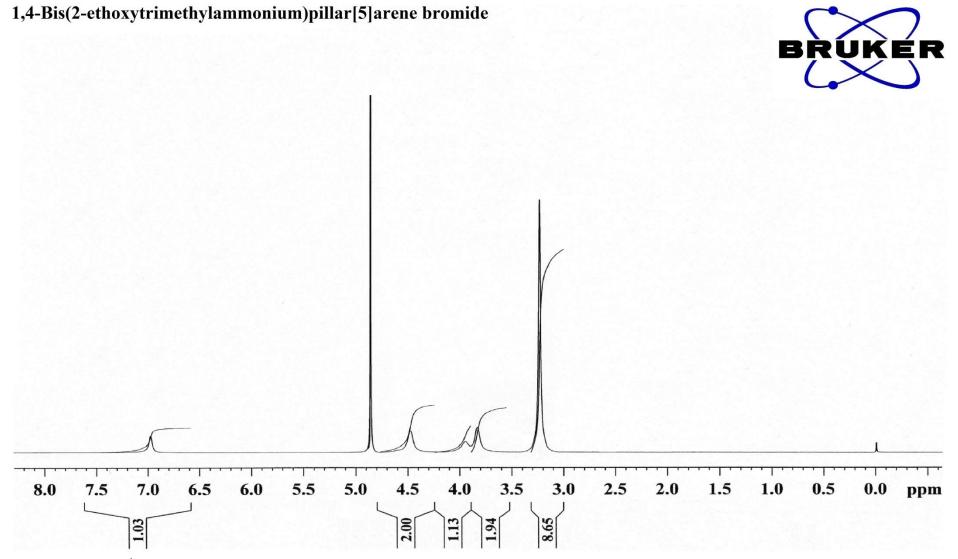
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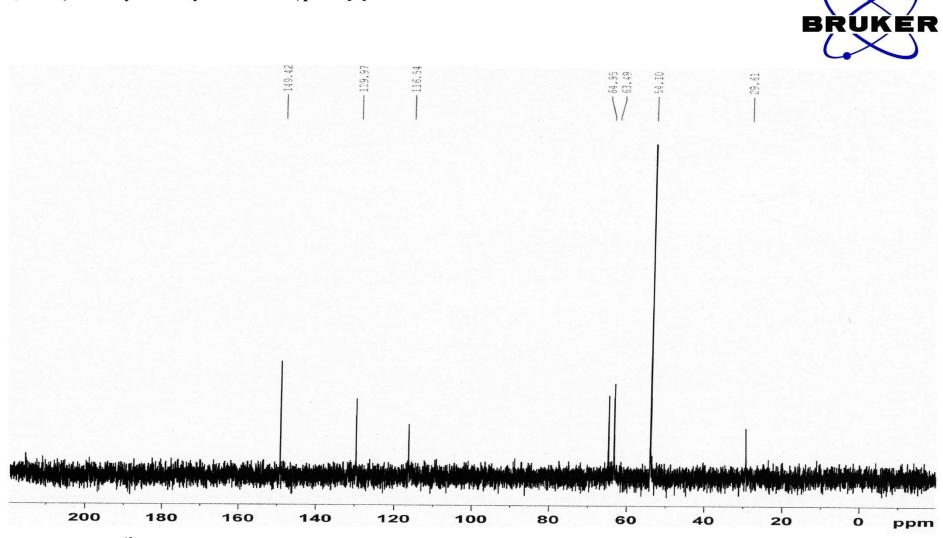
Bruker Daltonics DataAnalysis 3.4

Page 1 of 1



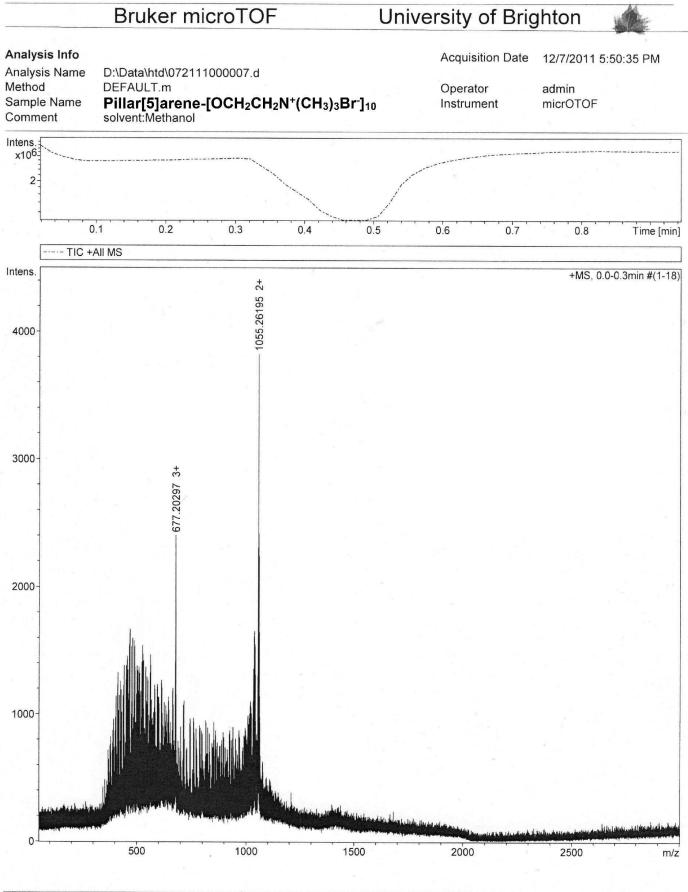


**Figure A1 - 25(a):** <sup>1</sup>H NMR spectra of **25** conducted in  $D_2O$ .

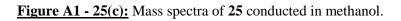


## 1,4-Bis(2-ethoxytrimethylammonium)pillar[5]arene bromide

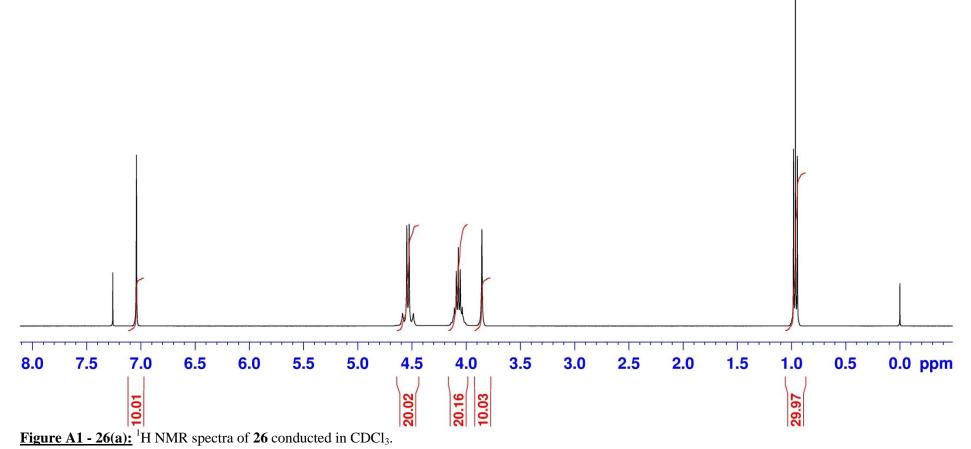
**Figure A1 - 25(b):** <sup>13</sup>C NMR spectra of **25** conducted in  $D_2O$ .



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Page 1 of 1



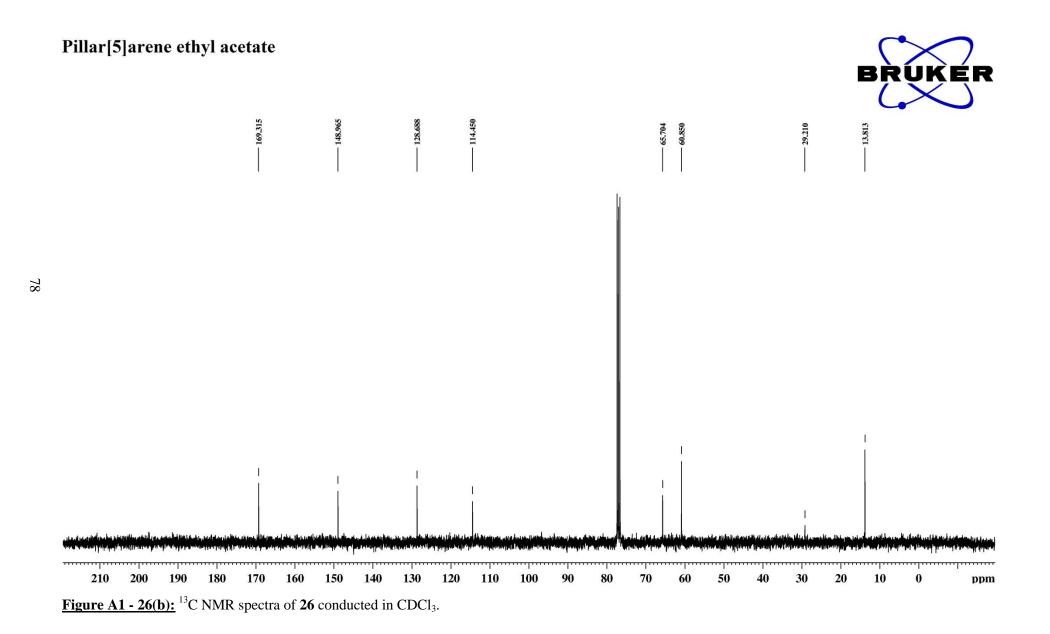
RUK

B

ÉR

Pillar[5]arene ethyl acetate

ΓΓ



## Bruker microTOF

## University of Brighton



 Analysis Name
 D:\Data\htd\030913000018.d

 Method
 DEFAULT.m

 Sample Name
 Pillar[5]arene ethyl acetate

 Comment
 solvent:Methanol

Acquisition Date 9/5/2013 3:11:35 PM

Operator Instrument admin micrOTOF

Page 1 of 1

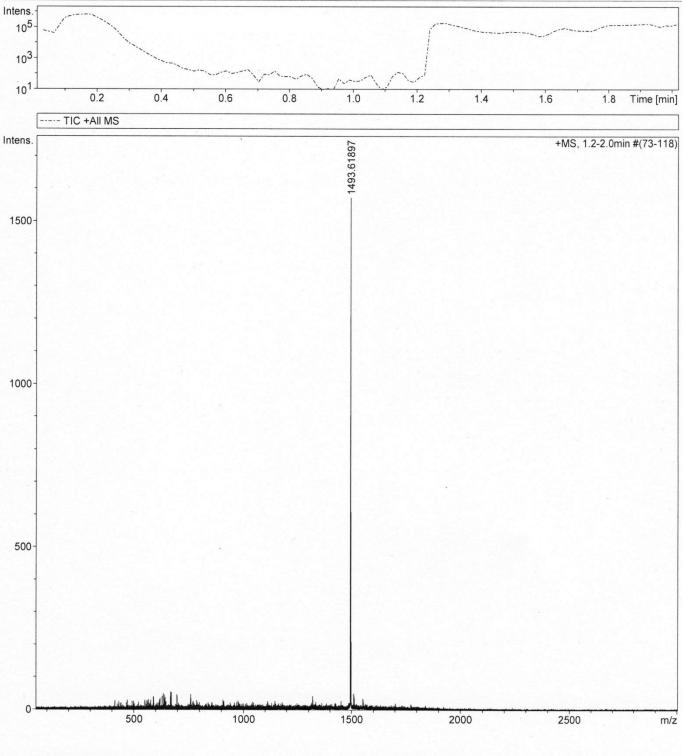


Figure A1 - 26(c): Mass spectra of 26 conducted in methanol.

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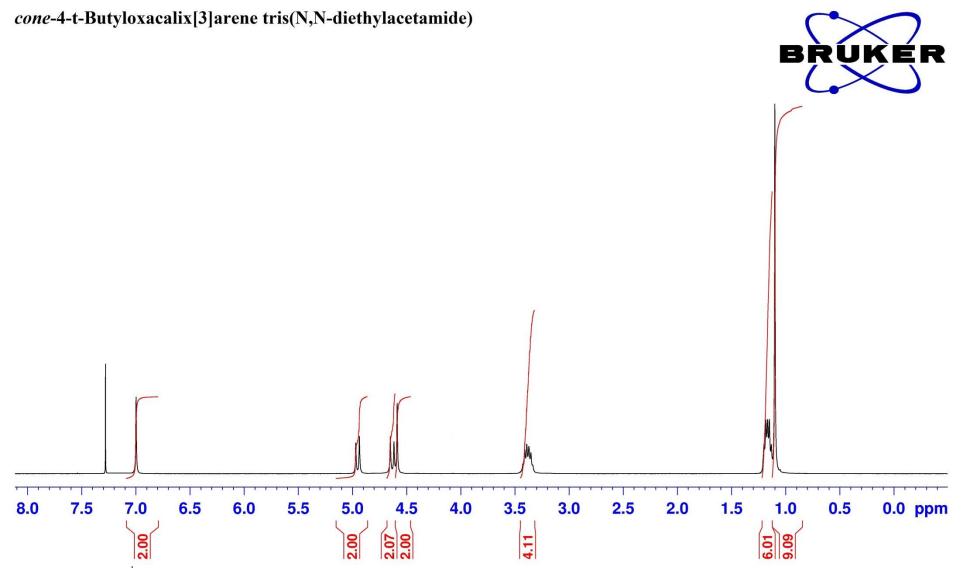


Figure A1 - 27(a): <sup>1</sup>H NMR spectra of 27 conducted in CDCl<sub>3</sub>.

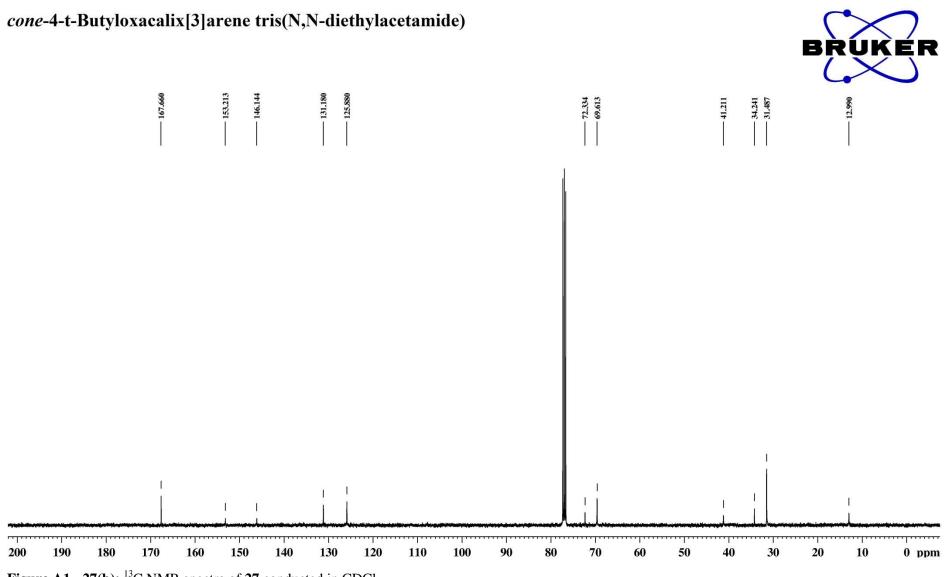


Figure A1 - 27(b): <sup>13</sup>C NMR spectra of 27 conducted in CDCl<sub>3</sub>.

## Bruker microTOF

## University of Brighton

Acquisition Date

#### Analysis Info

Analysis Name Method Sample Name Comment

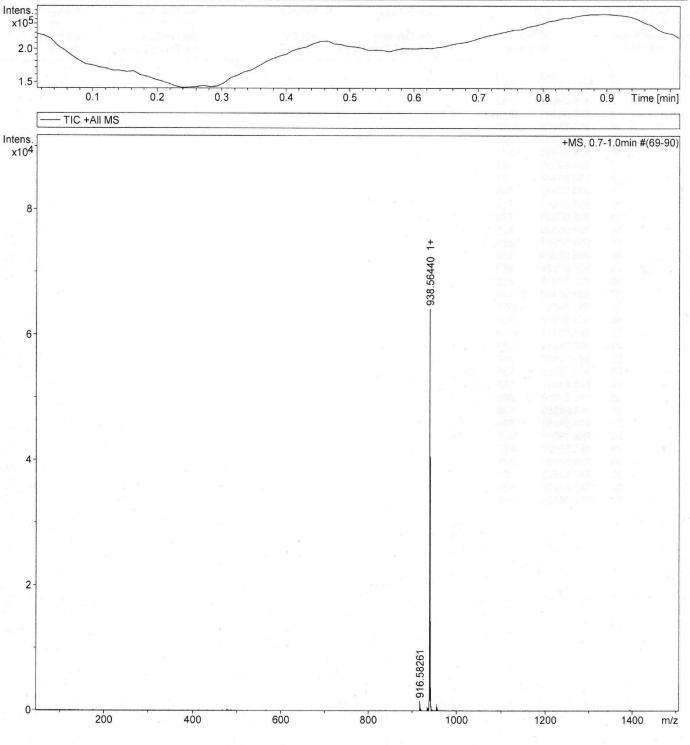
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Operator

Instrument

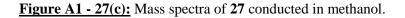
admin micrOTOF

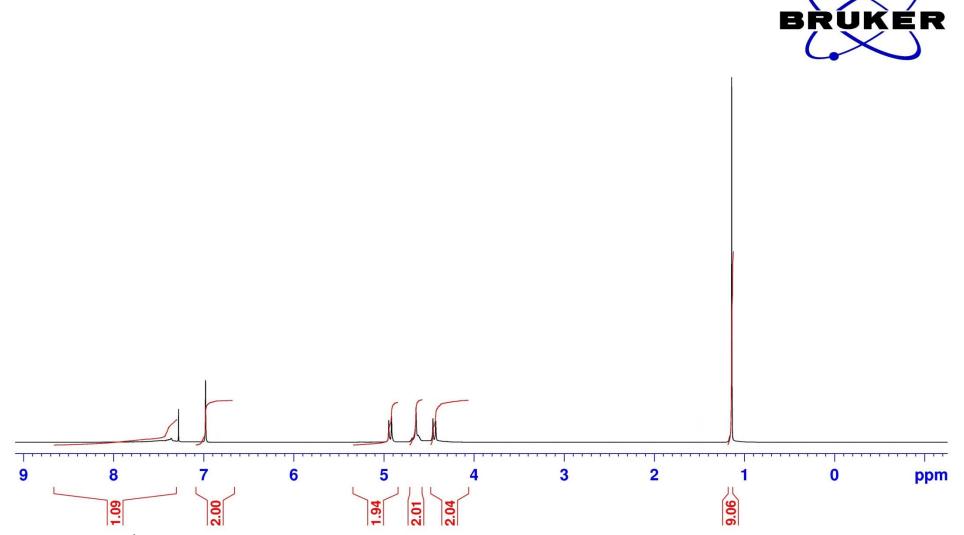
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Bruker Daltonics DataAnalysis 3.4

Page 1 of 1





cone-p-tert-butylhexahomotrioxacalix[3]arenetris(acetic acid)

Figure A1 - 28(a): <sup>1</sup>H NMR spectra of 28 conducted in CDCl<sub>3</sub>.

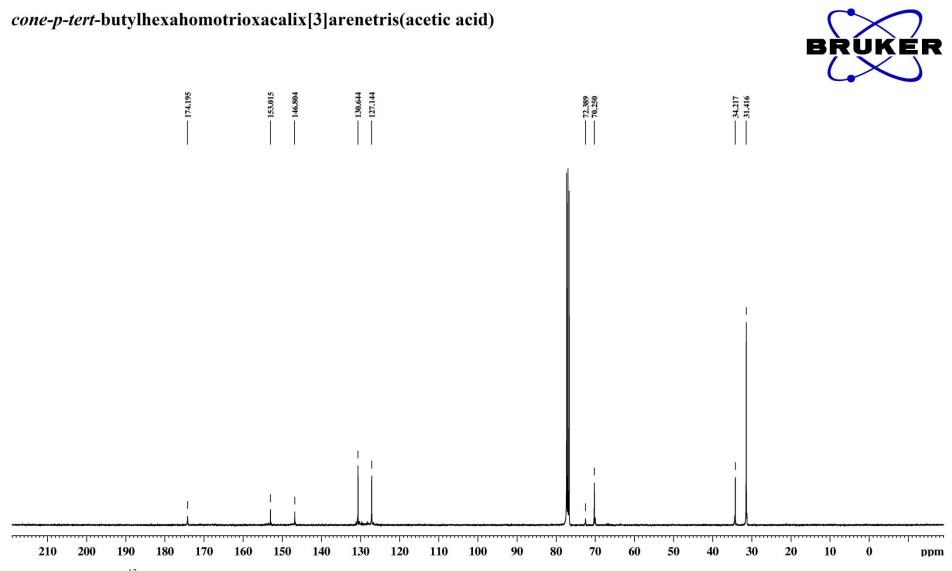


Figure A1 - 28(b): <sup>13</sup>C NMR spectra of 28 conducted in CDCl<sub>3</sub>.

## Bruker microTOF

## University of Brighton

Operator

Instrument

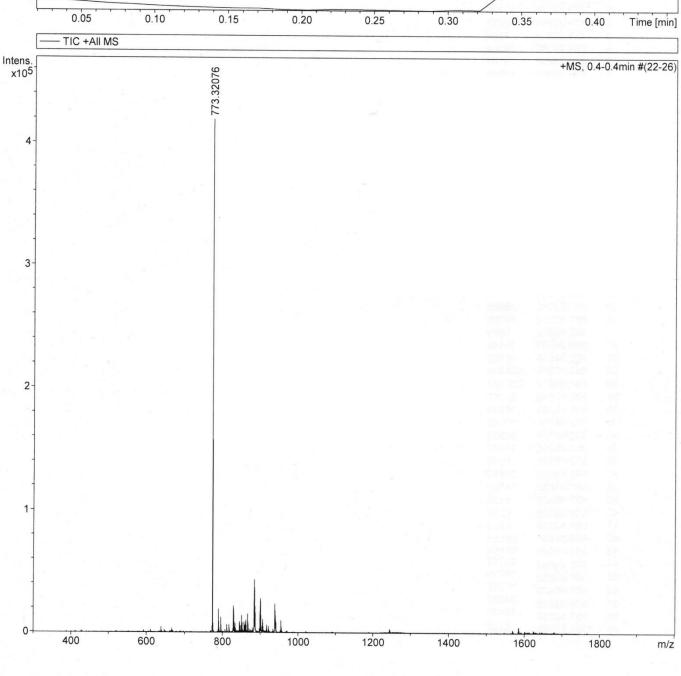
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admin

micrOTOF

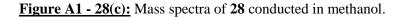
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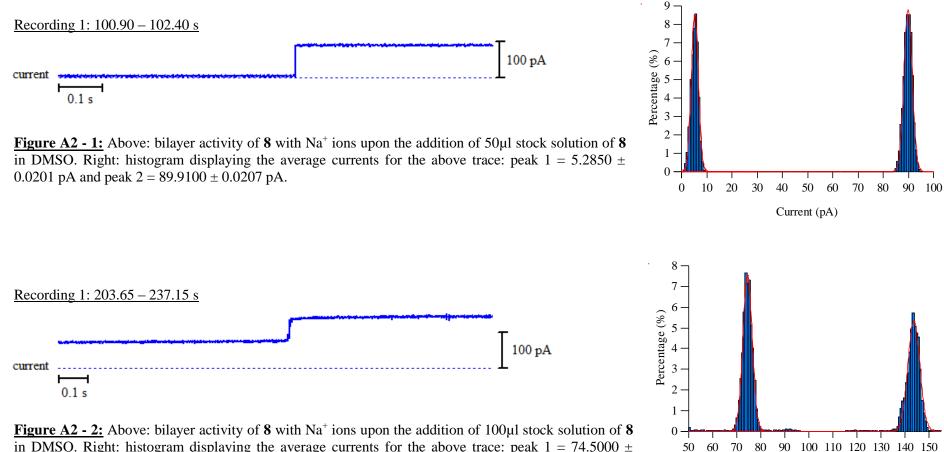
Page 1 of 1

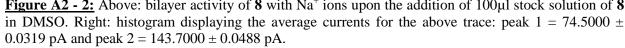


# **Appendix II**

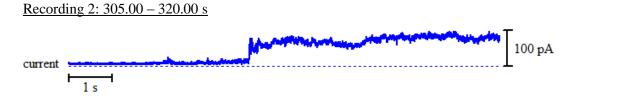
**Planar lipid bilayer experiments** 

#### Planar phospholipid bilayer activity of 8 towards Na<sup>+</sup> across the POPE and POPS bilayer system

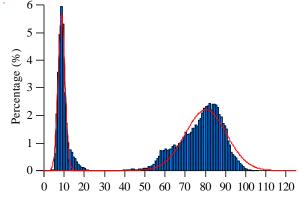






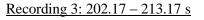


**Figure A2 - 3:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150 $\mu$ l stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 8.7040 \pm$ 0.0589 pA and peak  $2 = 80.1100 \pm 0.3392 \text{ pA}$ .



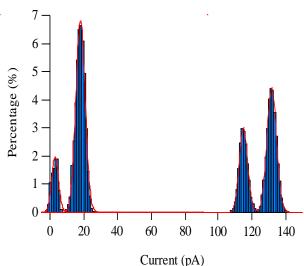
Current (pA)

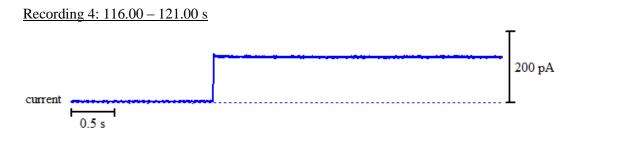
pA.





1 s **Figure A2 - 4:** Above: bilayer activity of **8** with  $Na^+$  ions upon the addition of 100µl stock solution of 8 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.9080 \pm$ 0.1642 pA, peak 2 = 17.9500 ± 0.0548 pA, peak 3 = 115 ± 0.0396 pA, and peak 4 = 131.8 ± 0.02815





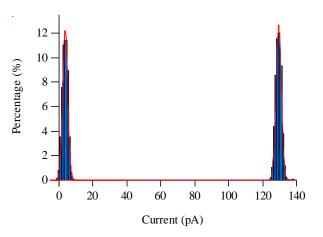
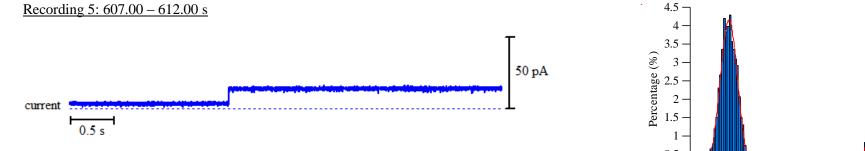
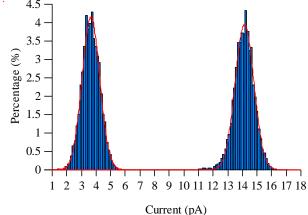


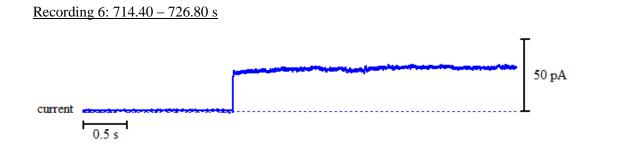
Figure A2 - 5: Above: bilayer activity of 8 with Na<sup>+</sup> ions upon the addition of 50µl stock solution of 8 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.787 \pm$ 0.01876 pA and peak  $2 = 129.4 \pm 0.01811 \text{ pA}$ .

88

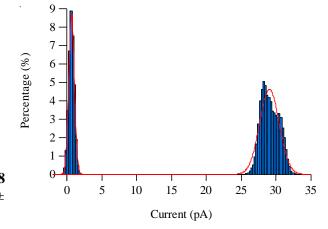


**Figure A2 - 6:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 200 $\mu$ l stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.642 \pm$ 0.008 pA and peak  $2 = 14.14 \pm 0.0084$  pA.





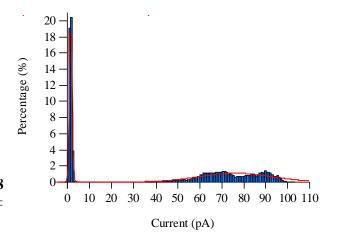
**Figure A2 - 7:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.6366 \pm 0.0122$  pA and peak  $2 = 29.01 \pm 0.0397$  pA.

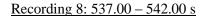


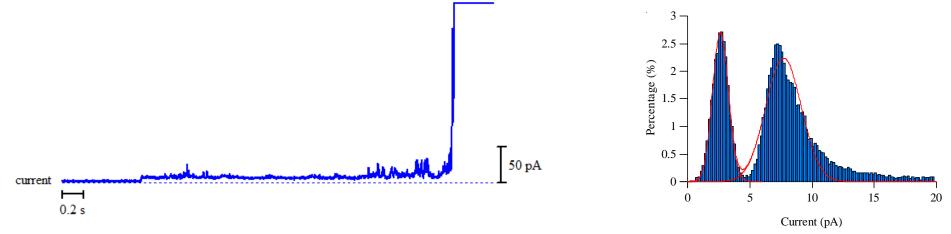




**Figure A2 - 8:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.427 \pm 0.0047$  pA and peak  $2 = 76.14 \pm 0.8700$  pA.







**Figure A2 - 9:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 225µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.632 \pm 0.0247$  pA and peak  $2 = 7.657 \pm 0.0436$  pA.

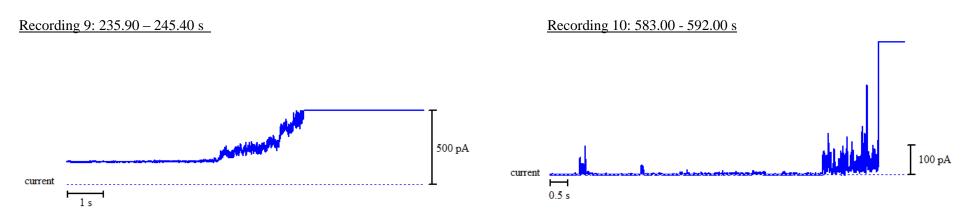
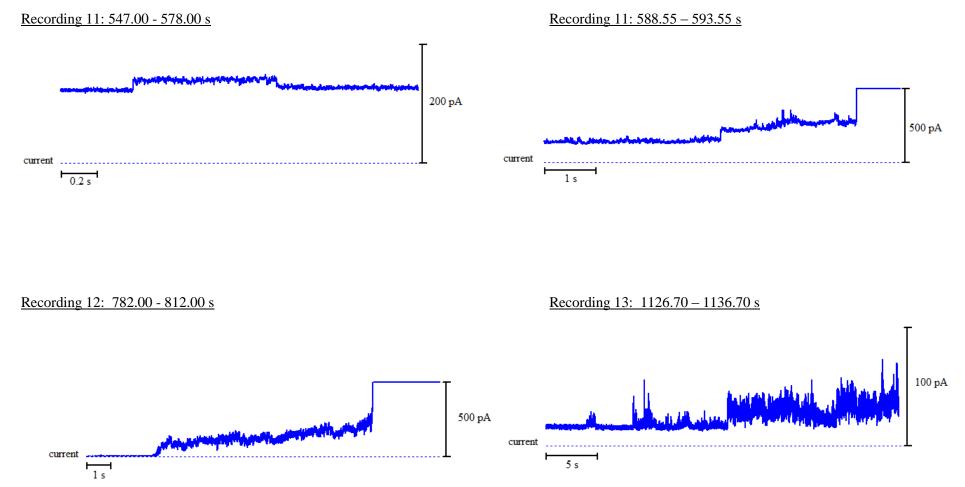
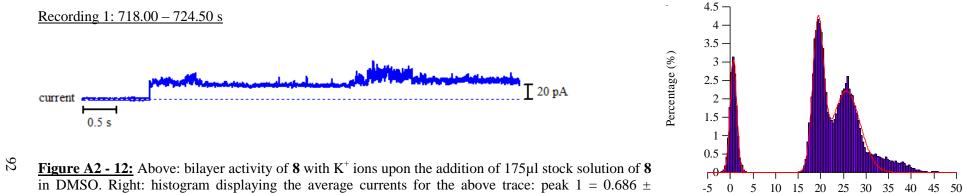


Figure A2 - 10: Bilayer activity of 8 with Na<sup>+</sup> ions upon the addition of 100µl (recording 9), 125µl (recording 10) stock solution of 8 in DMSO.



**Figure A2 - 11:** Bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150 $\mu$ l (recordings 11), 225 $\mu$ l (recording 12) and 250 $\mu$ l (recording 13) stock solution of **8** in DMSO.

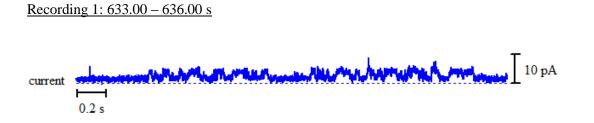
### Planar phospholipid bilayer activity of 8 towards K<sup>+</sup> across the POPE and POPS bilayer system



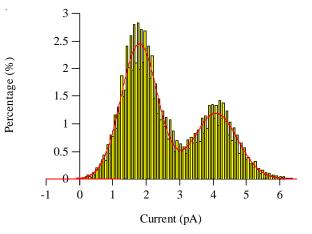
in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.686 \pm$ 0.0274 pA, peak 2 = 19.4 ± 0.0317 pA and peak 3 = 25.6 ± 0.1118 pA.



### Planar phospholipid bilayer activity of 8 towards Li<sup>+</sup> across the POPE and POPS bilayer system



**Figure A2 - 13:** Above: bilayer activity of **8** with Li<sup>+</sup> ions upon the addition of 150µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.787 \pm 0.0139$  pA and peak  $2 = 4.071 \pm 0.0309$  pA.



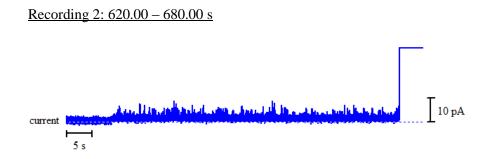


Figure A2 - 14: Bilayer activity of 8 with K<sup>+</sup> ions upon the addition of 150µl (recording 2) stock solution of 8 in DMSO

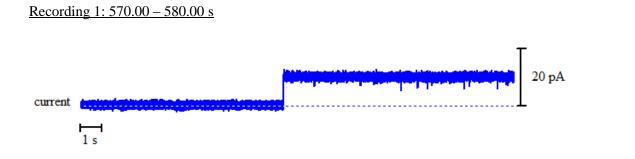
### Planar phospholipid bilayer activity of 8 towards Rb<sup>+</sup> across the POPE and POPS bilayer system

Due to experimental limitations, the ion channel activity of 8 towards  $Rb^+$  was unable to be investigated across this bilayer system.

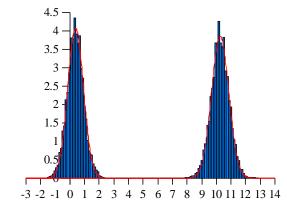
## Planar phospholipid bilayer activity of 8 towards Cs<sup>+</sup> across the POPE and POPS bilayer system

No activity was produced with  $\mathbf{8}$  towards  $Cs^+$  across this bilayer system

### Planar phospholipid bilayer activity of 8 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

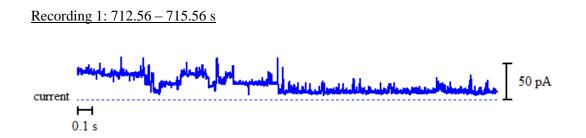


**Figure A2 - 15:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.3814 \pm 0.0069$  pA and peak  $2 = 10.25 \pm 0.0077$  pA.

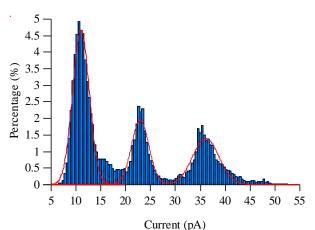


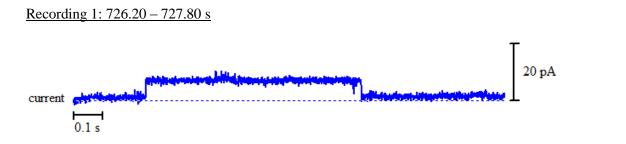
Percentage (%)

Current (pA)

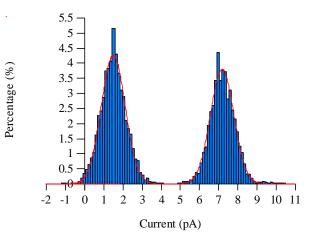


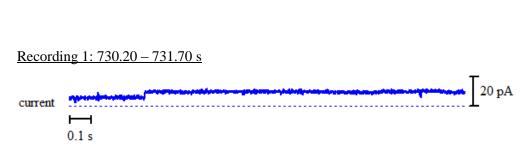
**Figure A2 - 16:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 10.98 \pm 0.0510$  pA, peak  $2 = 22.88 \pm 0.1265$  pA and peak  $3 = 36.05 \pm 0.2159$  pA.



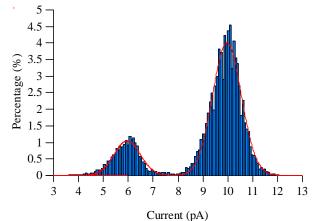


**Figure A2 - 17:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.489 \pm 0.0120$  pA and peak  $2 = 7.173 \pm 0.0134$  pA.



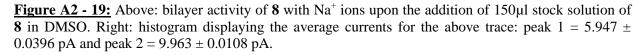


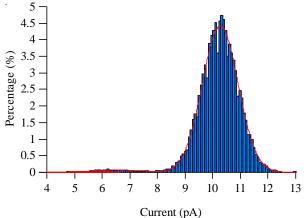
**Figure A2 - 18:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 5.947 \pm 0.0396$  pA and peak  $2 = 9.963 \pm 0.0108$  pA.

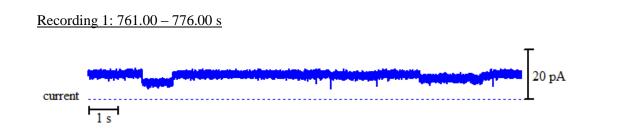


Recording 1: 739.70 -746.70 s

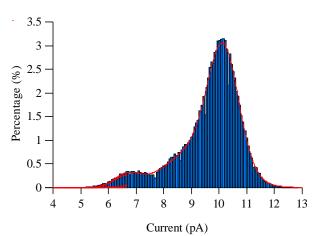
current I = 20 pA

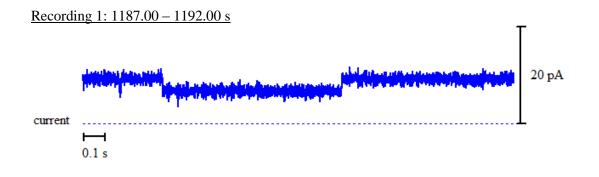


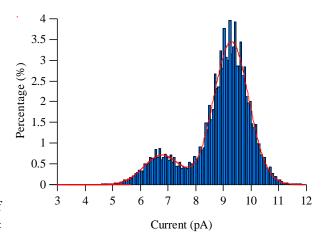




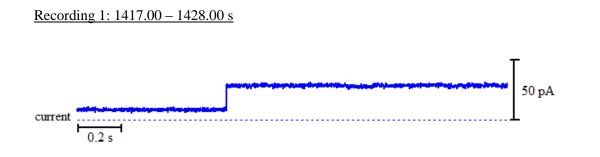
**Figure A2 - 20:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.67 \pm 0.0173$  pA and peak  $2 = 10.16 \pm 0.0101$  pA.

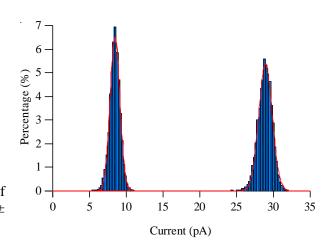




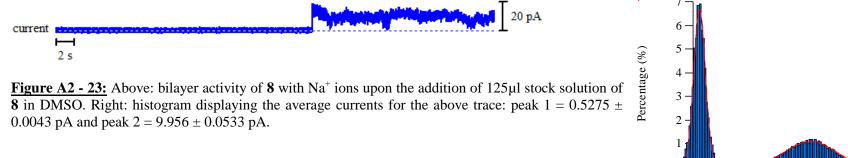


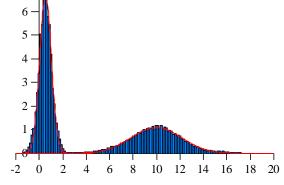
**Figure A2 - 21:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 225µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.772 \pm 0.0545$  pA and peak  $2 = 9.279 \pm 0.0112$  pA.





**Figure A2 - 22:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 250µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 8.463 \pm 0.0061$  pA and peak  $2 = 28.92 \pm 0.0085$  pA. Recording 2: 500.00 - 550.00 s









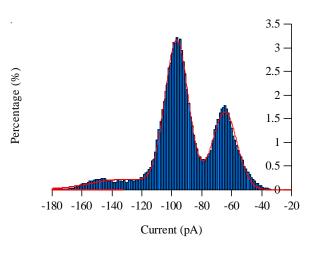
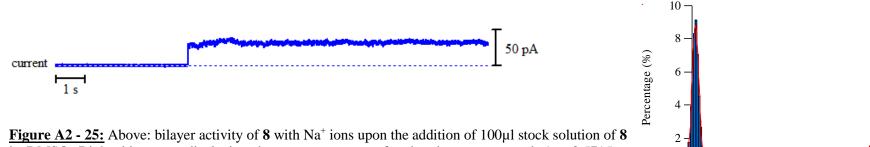
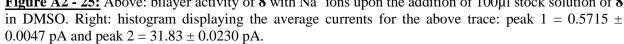
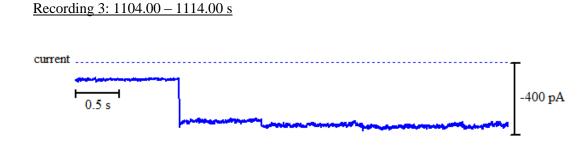
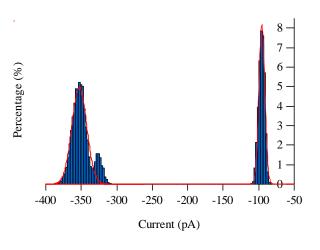


Figure A2 - 24: Above: bilayer activity of 8 with Na<sup>+</sup> ions upon the addition of 150µl stock solution of 8 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -131.3 \pm 0.9404$ pA, peak  $2 = -96.84 \pm 0.0781$  pA and peak  $3 = -64.66 \pm 0.1414$  pA. Recording 3: 254.00 – 264.00 s

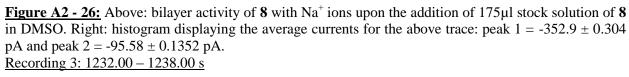




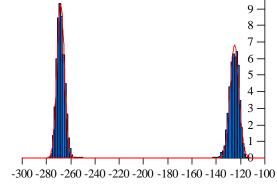




Current (pA)







10 -

**Figure A2 - 27:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -268.4 \pm 0.0413$  pA and peak  $2 = -125.0 \pm 0.0591$  pA.

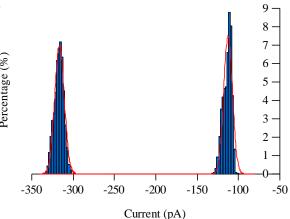
Currnet (pA)

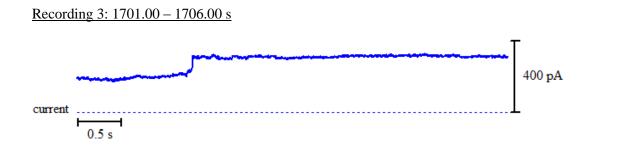


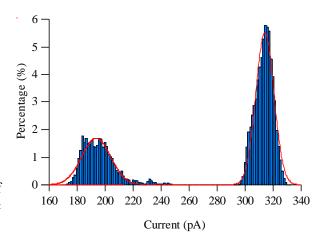


Recording 3: 1544.00 – 1552.00 s current -400 pA (%) effetting and a second s

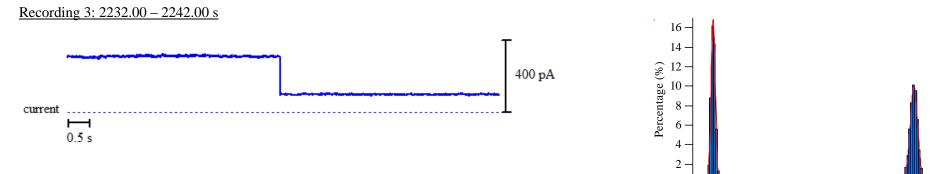
**Figure A2 - 28:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -316.6 \pm 0.2497$  pA and peak  $2 = -112.7 \pm 0.2241$  pA.







**Figure A2 - 29:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 193.4 \pm 0.5221$  pA and peak  $2 = 314.1 \pm 0.1313$  pA.

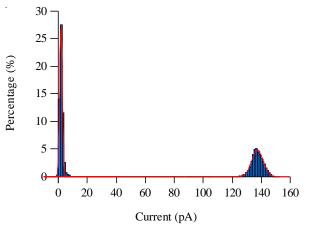


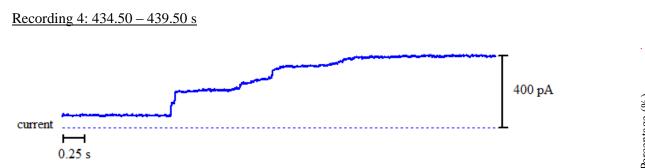
**Figure A2 - 30:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 250µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 101.7 \pm 0.0195$  pA and peak  $2 = 313.2 \pm 0.0346$  pA.

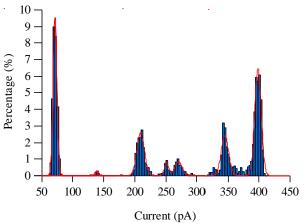
80 100 120 140 160 180 200 220 240 260 280 300 320 340



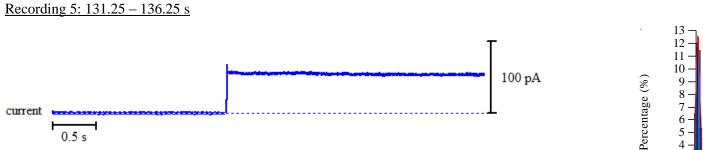
**Figure A2 - 31:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.87 \pm 0.0150$  pA and peak  $2 = 137.3 \pm 0.1119$  pA.



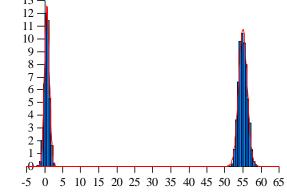




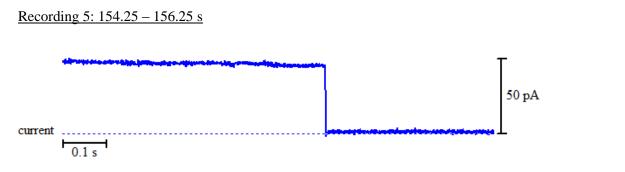
**Figure A2 - 32:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 71.21 \pm 0.0281$  pA, peak  $2 = 138.7 \pm 1.0160$  pA, peak  $3 = 208.5 \pm 0.2716$  pA, peak  $4 = 250.4 \pm 0.8182$  pA, peak  $5 = 269.5 \pm 0.9469$  pA, peak  $6 = 344.3 \pm 0.7990$  pA and peak  $7 = 398 \pm 0.2874$  pA.



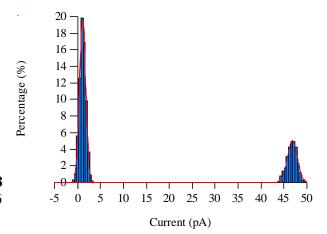
**Figure A2 - 33:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.6164 \pm 0.0077$  pA and peak  $2 = 54.92 \pm 0.0107$  pA.



Current (pA)

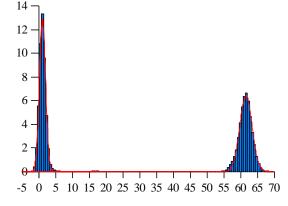


**Figure A2 - 34:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 46.77 \pm 0.0176$  pA and peak  $2 = 1.063 \pm 0.0038$  pA.





**Figure A2 - 35:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.041 \pm 0.0073$  pA and peak  $2 = 61.49 \pm 0.0190$  pA.



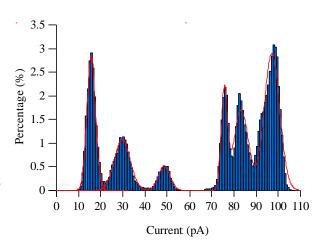
Percentage (%)

Current (pA)

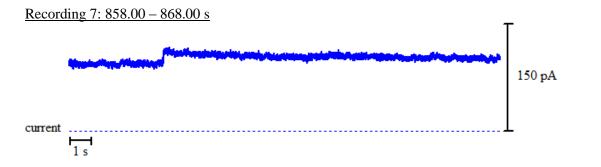
105

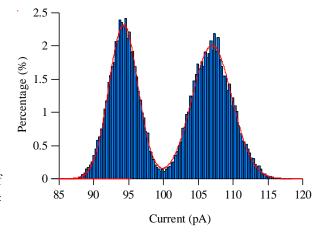
Recording 7: 807.00 – 837.00 s

current I s

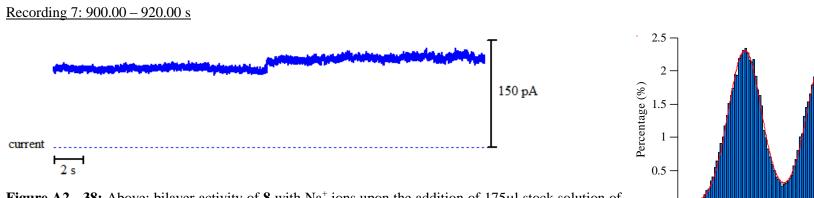


**Figure A2 - 36:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 15.73 \pm 0.0267$  pA, peak  $2 = 29.74 \pm 0.0967$  pA, peak  $3 = 48.69 \pm 0.1719$  pA, peak  $4 = 75.59 \pm 0.1255$  pA, peak  $5 = 83.15 \pm 0.2860$  pA and peak  $6 = 97.09 \pm 0.1591$  pA.

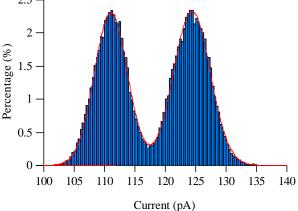


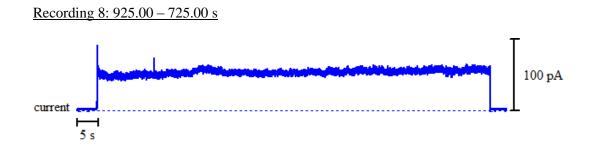


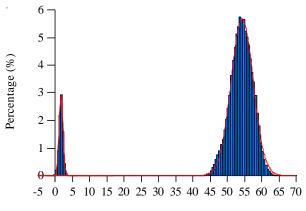
**Figure A2 - 37:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 94.23 \pm 0.0255$  pA and peak  $2 = 106.9 \pm 0.0334$  pA.



**Figure A2 - 38:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 111 \pm 0.0275$  pA and peak  $2 = 124.4 \pm 0.0286$  pA.



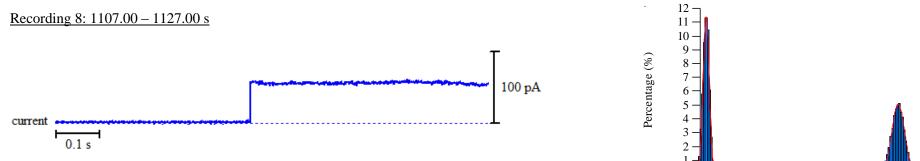




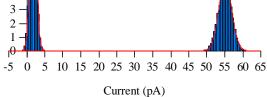
**Figure A2 - 39:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.637 \pm 0.0184$  pA and peak  $2 = 54.29 \pm 0.0167$  pA.







**Figure A2 - 40:** Above: bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 300µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.774 \pm 0.0060$  pA and peak  $2 = 55.1 \pm 0.0178$  pA.



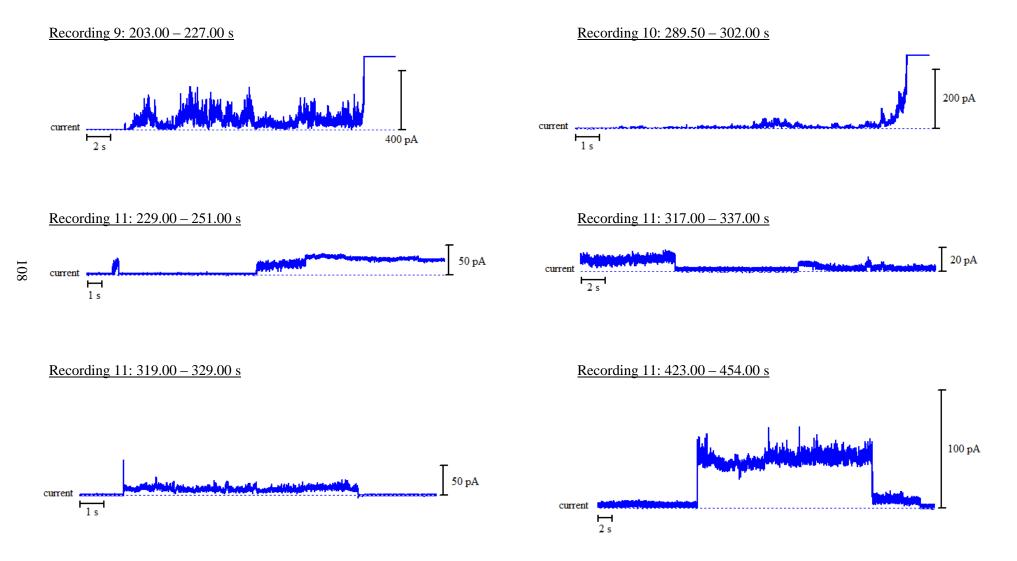
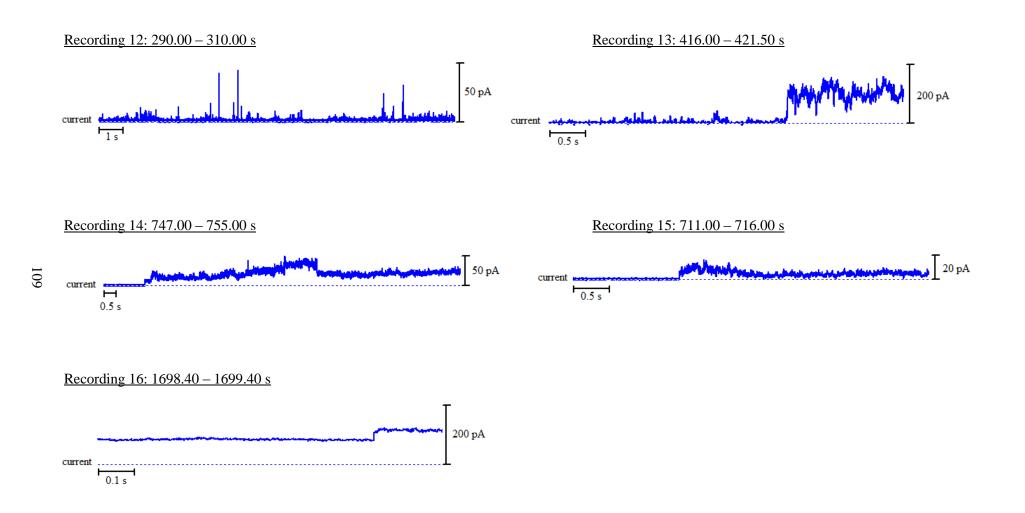
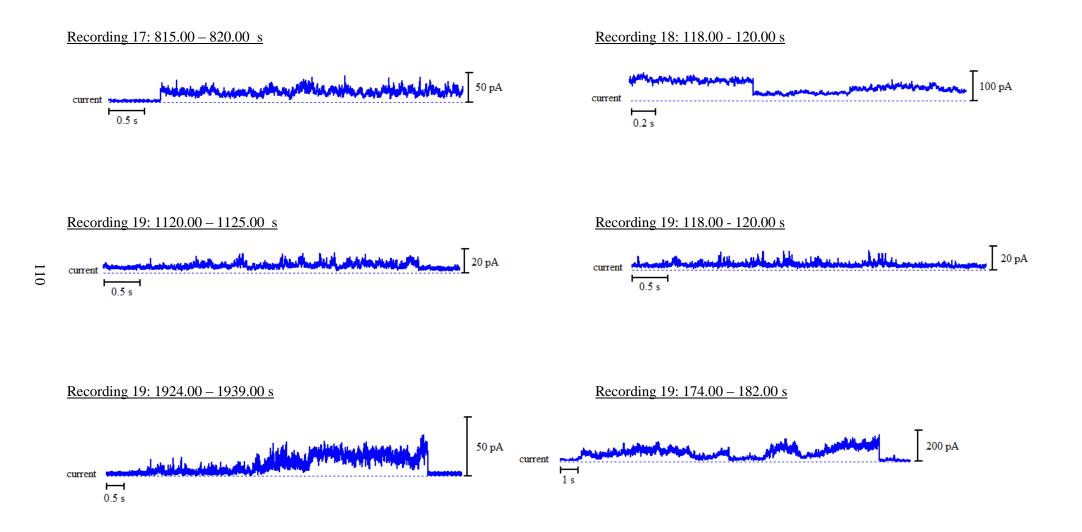


Figure A2 - 41: Bilayer activity of 8 with Na<sup>+</sup> ions upon the addition of 50µl (recordings 9-11) and 75µl (recordings 11- lower) stock solution of 8 in DMSO.

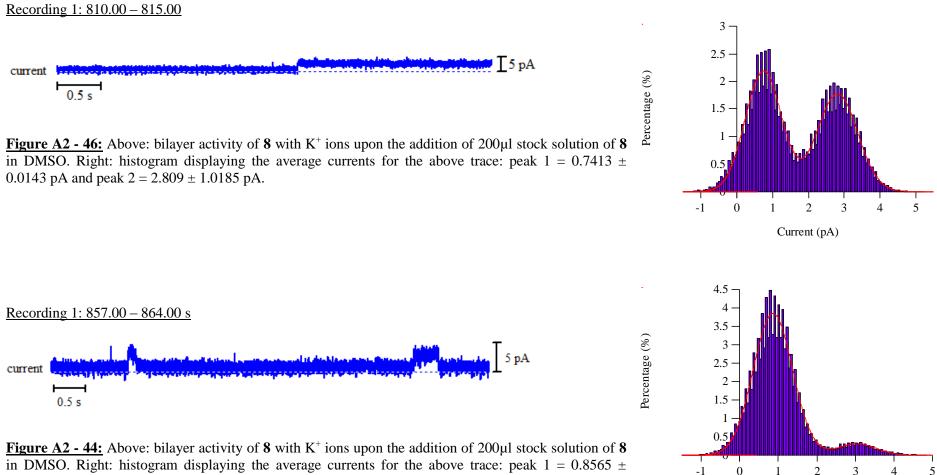


**Figure A2 - 42:** Bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 100 $\mu$ l (recording 12), 150 $\mu$ l (recording 13) and 175 $\mu$ l (recordings 14-16) stock solution of **8** in DMSO.



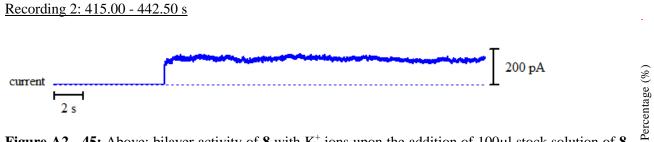
**Figure A2 - 43:** Bilayer activity of **8** with Na<sup>+</sup> ions upon the addition of 200µl (recordings 17 and 18), 225µl (recording 19- centre) and 250µl (recordings 19 - lower) stock solution of **8** in DMSO.

#### Planar phospholipid bilayer activity of 8 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

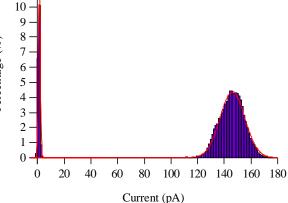


in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.8565 \pm$ 0.0107 pA and peak  $2 = 3.035 \pm 0.129$  pA.



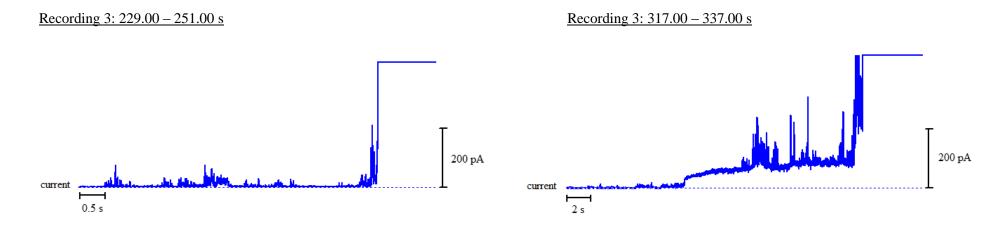


**Figure A2 - 45:** Above: bilayer activity of **8** with  $K^+$  ions upon the addition of 100µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.343 \pm 0.0174$  pA and peak  $2 = 146.00 \pm 0.797$  pA.

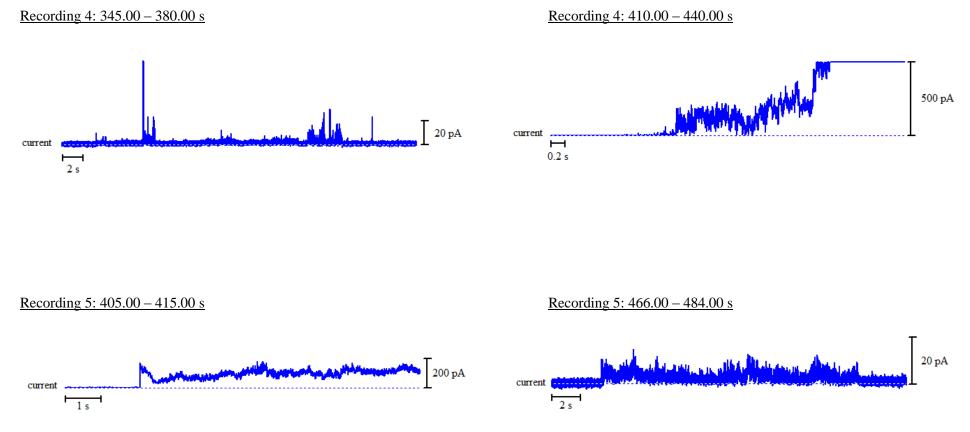


11 -

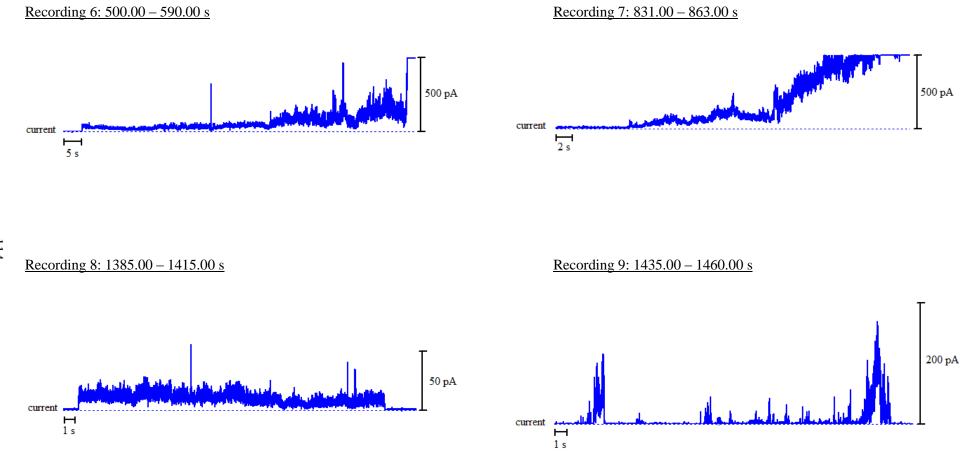




**Figure A2 - 47:** Bilayer activities of **8** with  $K^+$  ions upon the addition of 50µl stock solution of **8** in DMSO.

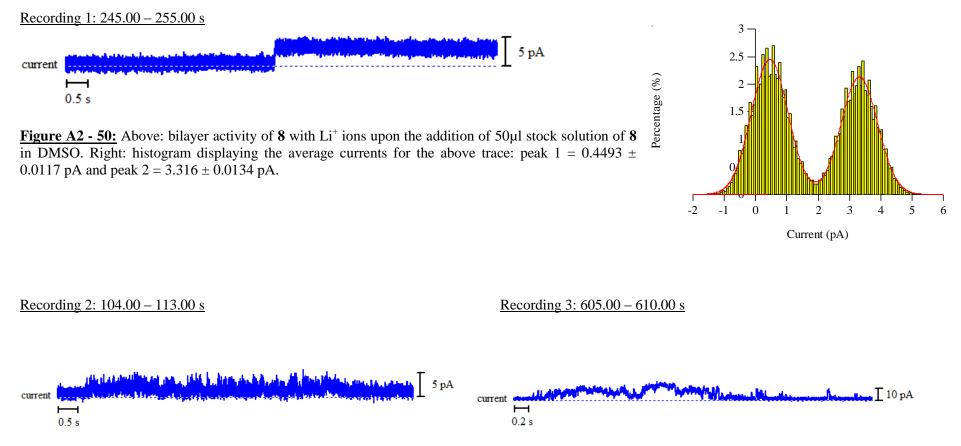


**Figure A2 - 48:** Bilayer activity of **8** with  $K^+$  ions upon the addition of 75µl (recording 4 – left), 100µl (recording 4 – right) and 100µl (recordings 5) stock solution of **8** in DMSO.



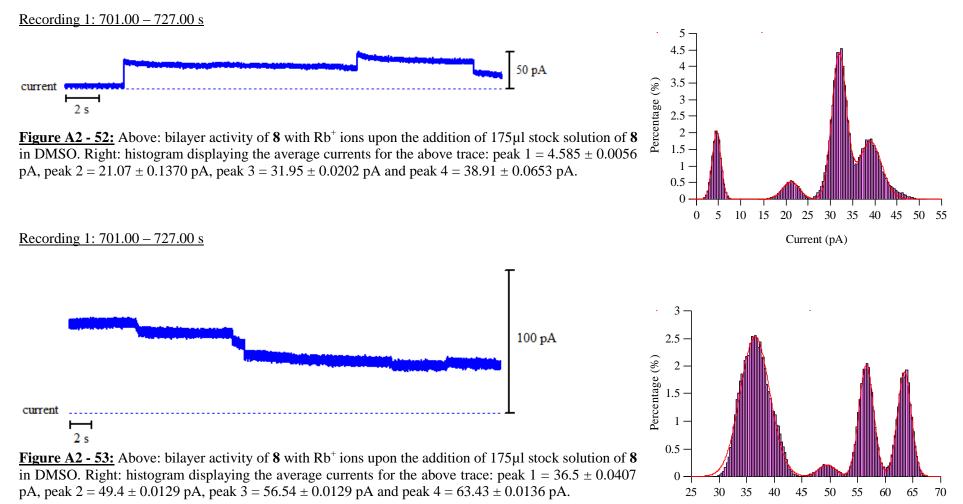
**Figure A2 - 49:** Bilayer activity of **8** with  $K^+$  ions upon the addition of 125µl (recording 6), 175µl (recording 7) and 300µl (recordings 8 and 9) stock solution of **8** in DMSO.

### Planar phospholipid bilayer activity of 8 towards Li<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



**Figure A2 - 51:** Bilayer activity of **8** with  $K^+$  ions upon the addition of 25µl (recording 2) and 150µl (recording 3) stock solution of **8** in DMSO.

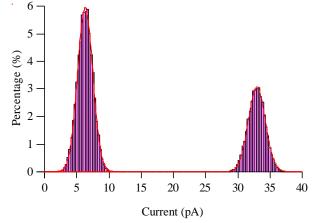
Planar phospholipid bilayer activity of 8 towards Rb<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



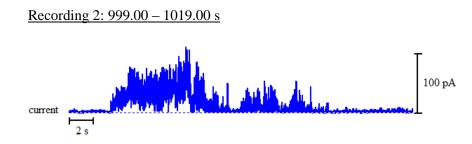
Current (pA)



**Figure A2 - 54:** Above: bilayer activity of **8** with Rb<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.311 \pm 0.0050$  pA and peak  $2 = 32.94 \pm 0.0102$  pA.

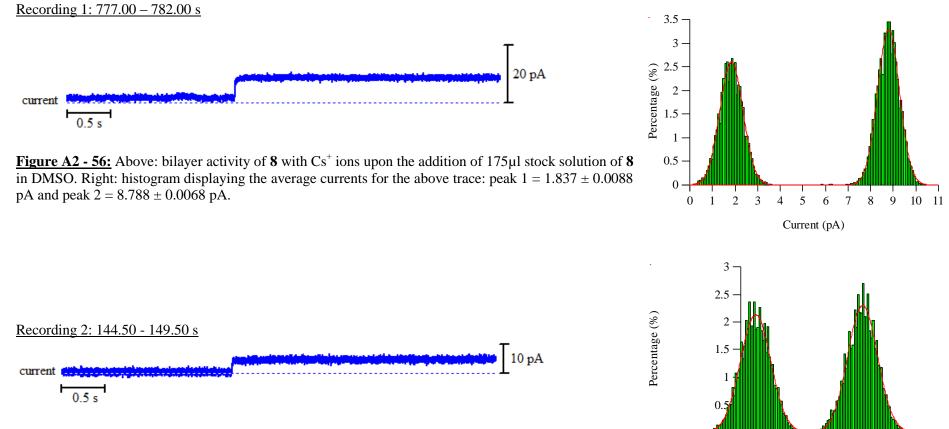






**Figure A2 - 55:** Bilayer activity of **8** with  $Rb^+$  ions upon the addition of 200µl (recording 2) stock solution of **8** in DMSO.

## Planar phospholipid bilayer activity of 8 towards Cs<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



**Figure A2 - 57:** Above: bilayer activity of **8** with Cs<sup>+</sup> ions upon the addition of  $25\mu$ l stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.601 \pm 0.0135$  pA and peak  $2 = 4.875 \pm 0.0124$  pA.

118



3

-2 -1

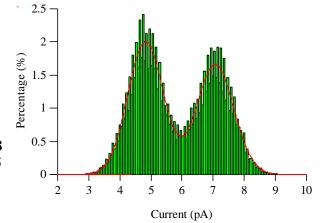
0 1 2

5

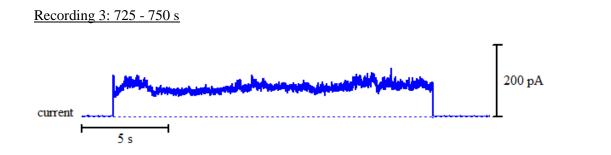
6

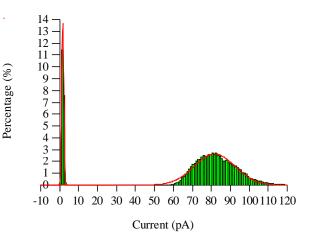
4





**Figure A2 - 58:** Above: bilayer activity of **8** with Cs<sup>+</sup> ions upon the addition of 75µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.823 \pm 0.0165$  pA and peak  $2 = 7.062 \pm 0.0203$  pA.





**Figure A2 - 59:** Above: bilayer activity of **8** with Cs<sup>+</sup> ions upon the addition of 175µl stock solution of **8** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.659 \pm 0.0125$  pA and peak  $2 = 81.59 \pm 0.1376$  pA.

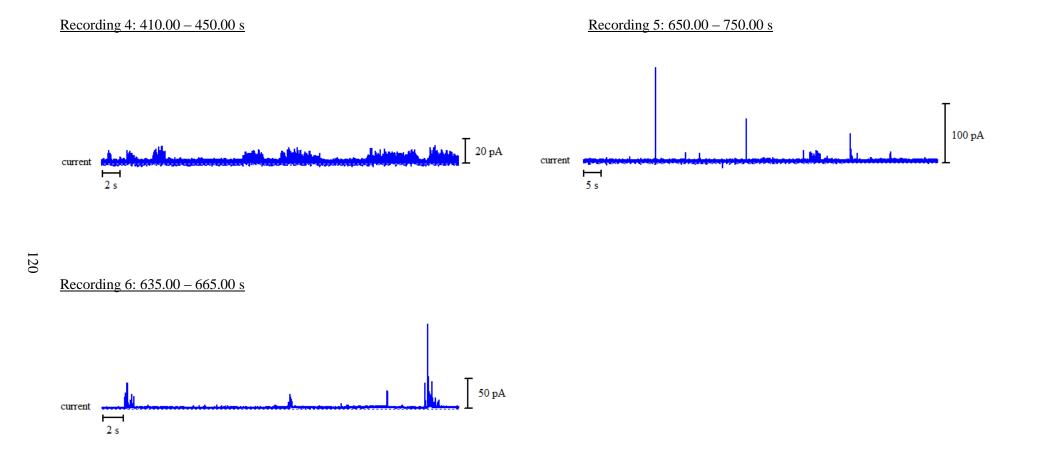
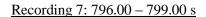
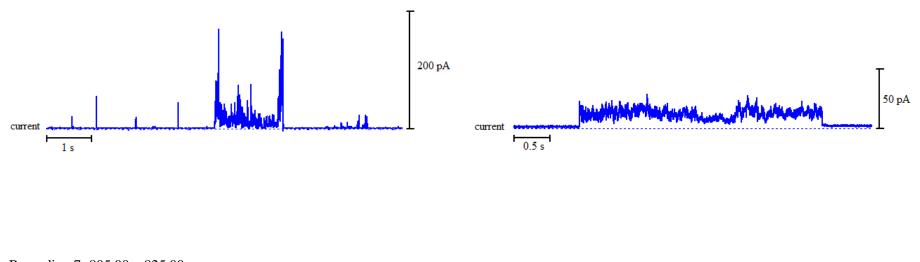


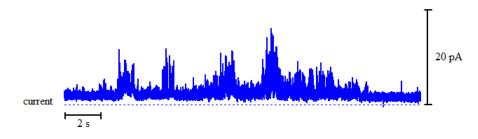
Figure A2 - 60: Bilayer activity of 8 with Cs<sup>+</sup> ions upon the addition of 100µl (recording 4) and 150µl (recordings 5 and 6) stock solution of 8 in DMSO.

<u>Recording 7: 150.00 – 155.00 s</u>

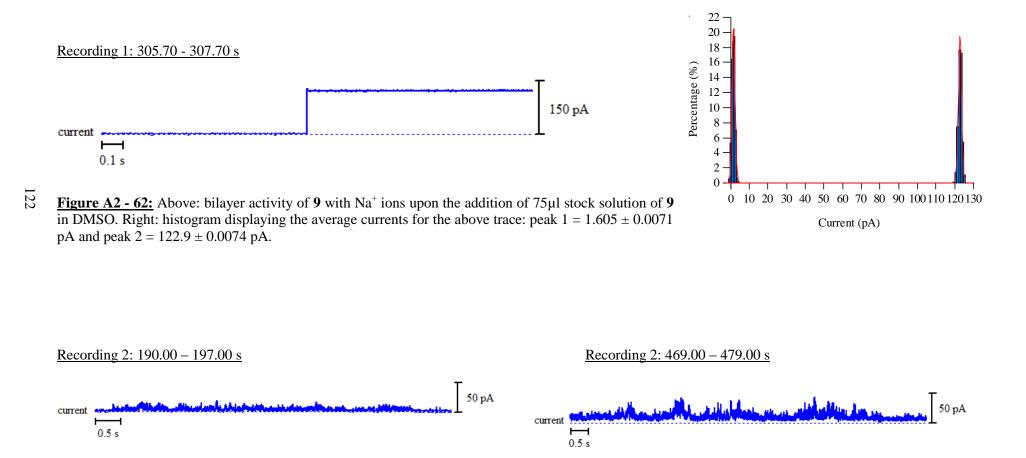




Recording 7: 805.00 - 825.00 s

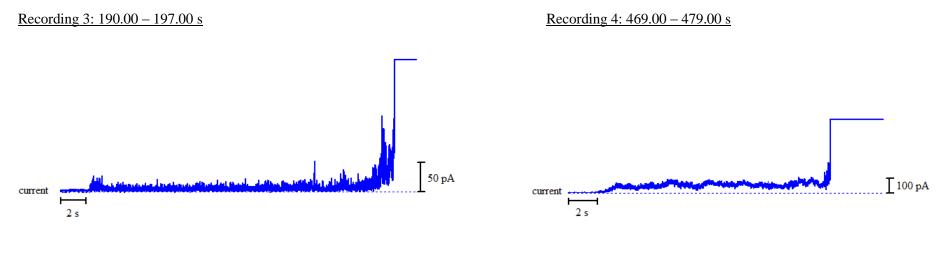


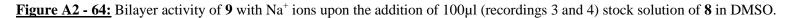
**Figure A2 - 61:** Bilayer activity of **8** with  $Cs^+$  ions upon the addition of 25µl (recording 7 – upper), 175µl (recording 7) and 200µl (recording 7 – lower) stock solution of **8** in DMSO.



## Planar phospholipid bilayer activity of 9 towards Na<sup>+</sup> across the POPE and POPS bilayer system

**Figure A2 - 63:** Bilayer activity of **9** with Na<sup>+</sup> ions upon the addition of  $25\mu l$  (recording 2 – left) and  $100\mu l$  (recording 2 – right) stock solution of **8** in DMSO.

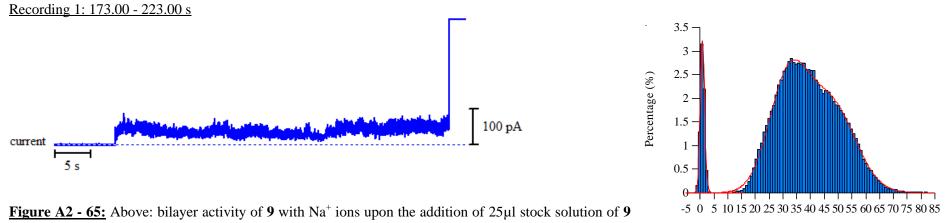




## Planar phospholipid bilayer activity of 9 towards $\mathbf{K}^{\scriptscriptstyle +}$ across the POPE and POPS bilayer system

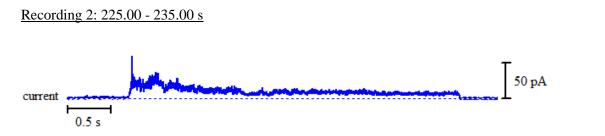
Due to experimental limitations, the ion channel activity of **9** towards  $K^+$  was unable to be investigated across this bilayer system.

### Planar phospholipid bilayer activity of 9 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

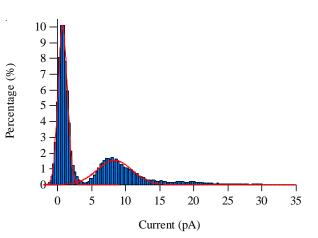


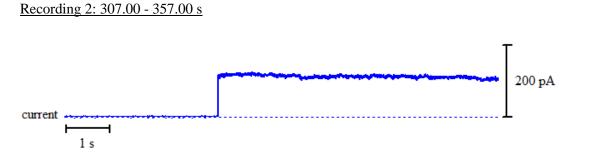
**Figure A2 - 65:** Above: bilayer activity of **9** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **9** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.8466 \pm 0.0138$  pA, peak  $2 = 31.92 \pm 0.2449$  pA and peak  $3 = 47.53 \pm 0.456$  pA.

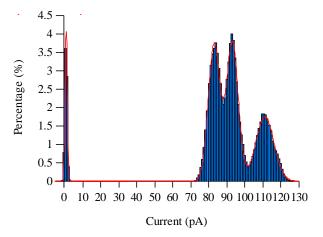




**Figure A2 - 66** Above: bilayer activity of **9** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **9** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.7661 \pm 0.0100$  pA and peak  $2 = 8.237 \pm 0.1265$  pA.

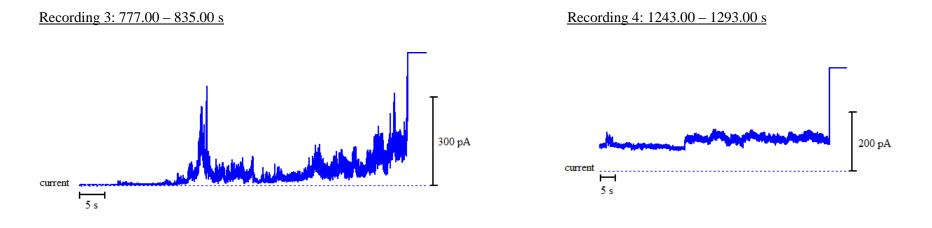




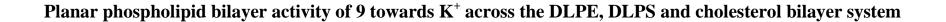


**Figure A2 - 67:** Above: bilayer activity of **9** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **9** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.18 \pm 0.0004$  pA, peak  $2 = 111 \pm 0.1246$  pA, peak  $3 = 92.95 \pm 0.0537$  pA and peak  $4 = 82.85 \pm 0.0454$  pA.





**Figure A2 - 68:** Bilayer activity of **9** with Na<sup>+</sup> ions upon the addition of 150 $\mu$ l (recordings 7 and 8) stock solution of **8** in DMSO.



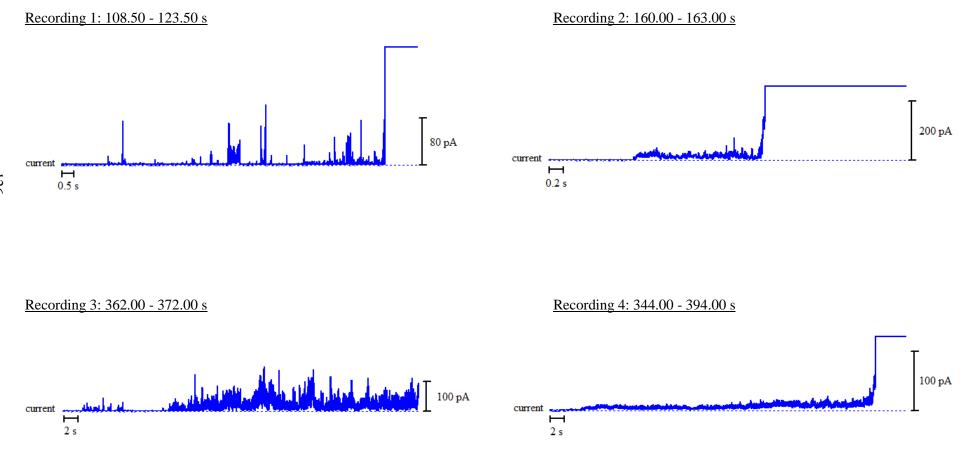
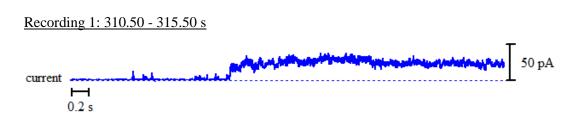
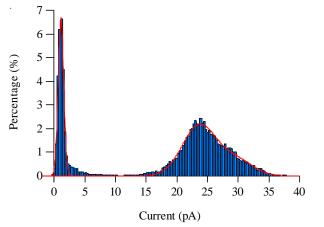


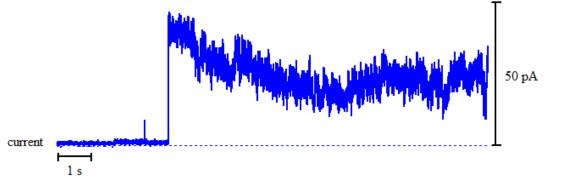
Figure A2 - 69: Bilayer activity of 9 with K<sup>+</sup> ions upon the addition of 25µl (recording 1 and 2) and 75µl (recording 3 and 4) stock solution of 9 in DMSO.

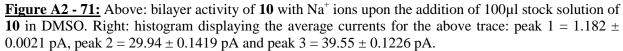
#### Planar phospholipid bilayer activity of 10 towards Na<sup>+</sup> across the POPE and POPS bilayer system

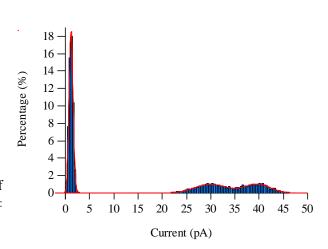


**Figure A2 - 70:** Above: bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.108 \pm 0.0072$  pA and peak  $2 = 23.63 \pm 0.2292$  pA.





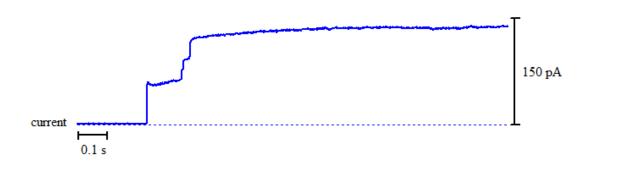


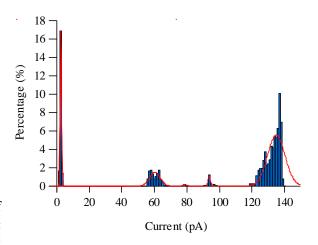


# 127

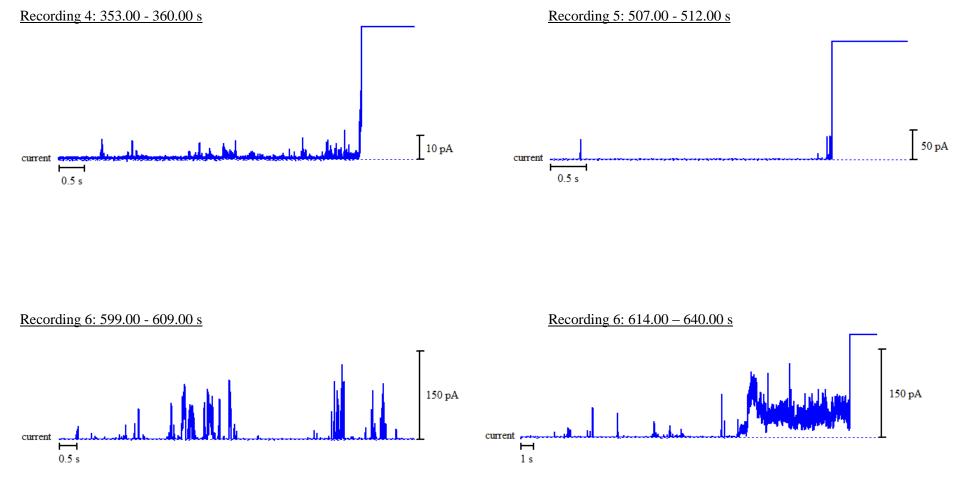
Recording 2: 507.30 - 311.30 s



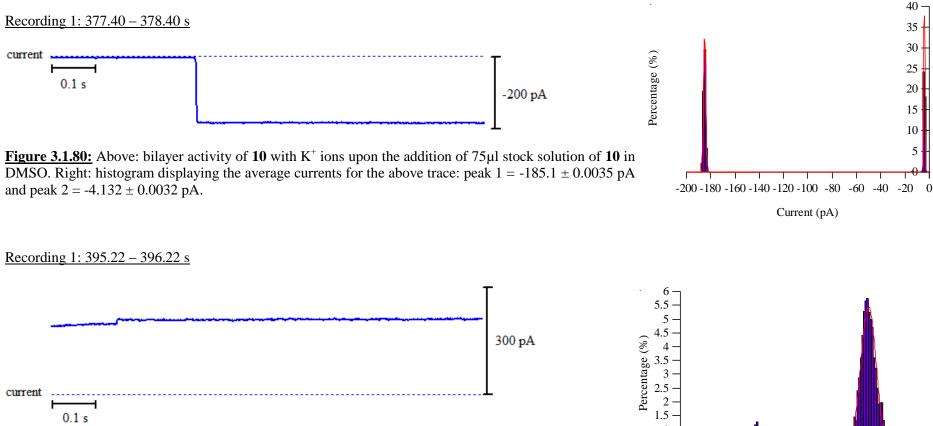




**Figure A2 - 72:** Above: bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.538 \pm 0.0144$  pA, peak  $2 = 60.33 \pm 0.2579$  pA, peak  $3 = 78.63 \pm 9.275$  pA, peak  $4 = 93.6 \pm 1.502$  pA and peak  $5 = 135 \pm 0.6027$  pA.

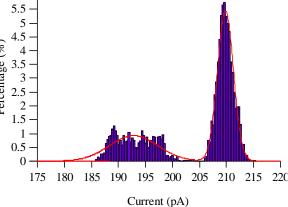


**Figure A2 - 73:** Bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of  $75\mu$ l (recording 4),  $125\mu$ l (recording 5 and 6 - left) and  $150\mu$ l (recording 6 - right) stock solution of **10** in DMSO.

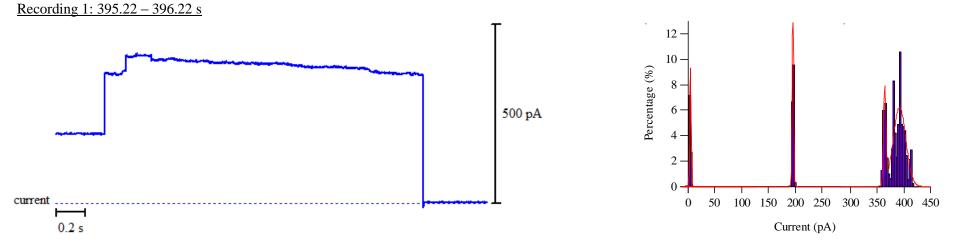


### Planar phospholipid bilayer activity of 10 towards K<sup>+</sup> across the POPE and POPS bilayer system

Figure A2 - 74: Above: bilayer activity of 10 with K<sup>+</sup> ions upon the addition of 75µl stock solution of 10 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 192.8 \pm 0.3911$ pA and peak  $2 = 209.8 \pm 0.0512$  pA.







**Figure A2 - 75:** Above: bilayer activity of **10** with K<sup>+</sup> ions upon the addition of 75µl stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.765 \pm 0.0104$  pA, peak  $2 = 195.8 \pm 0.0094$  pA, peak  $3 = 364.5 \pm 1.2290$  pA and peak  $4 = 391.8 \pm 0.4682$  pA.

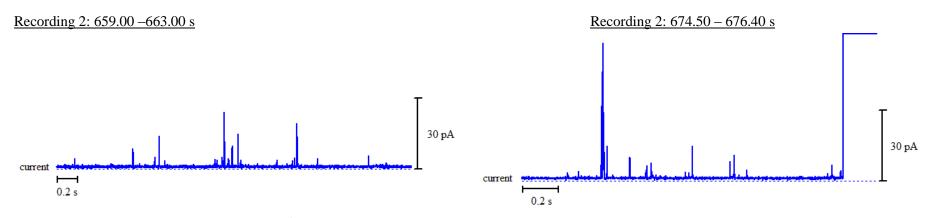
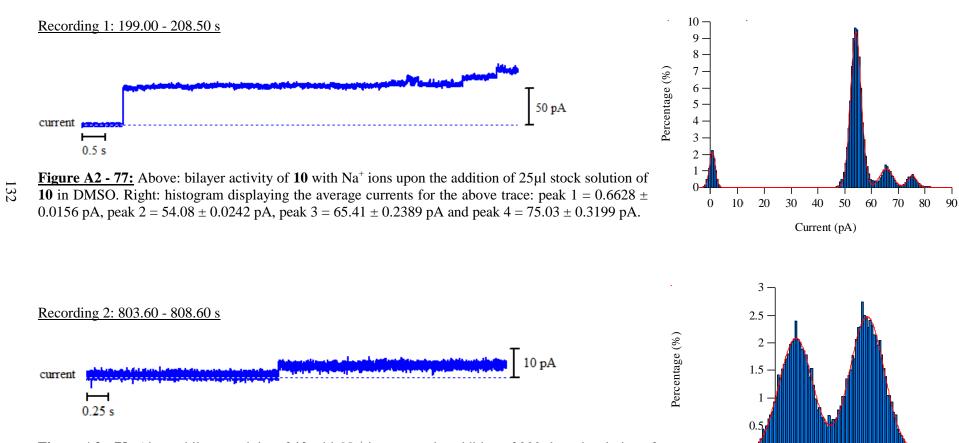


Figure A2 - 76: Bilayer activity of 10 with Na<sup>+</sup> ions upon the addition of 125µl (recordings 2), stock solution of 10 in DMSO.

### Planar phospholipid bilayer activity of 10 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



**Figure A2 - 78:** Above: bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.9736 \pm 0.0106$  pA and peak  $2 = 4.382 \pm 0.0090$  pA.

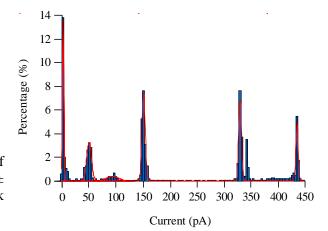
-3 -2 -1 0 1 2 3 4 5 6 Current (pA)

78

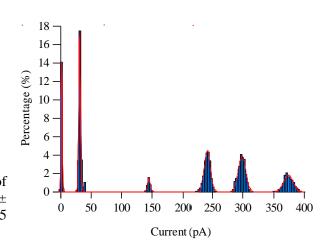
Recording 3: 100.02 – 100.12 s



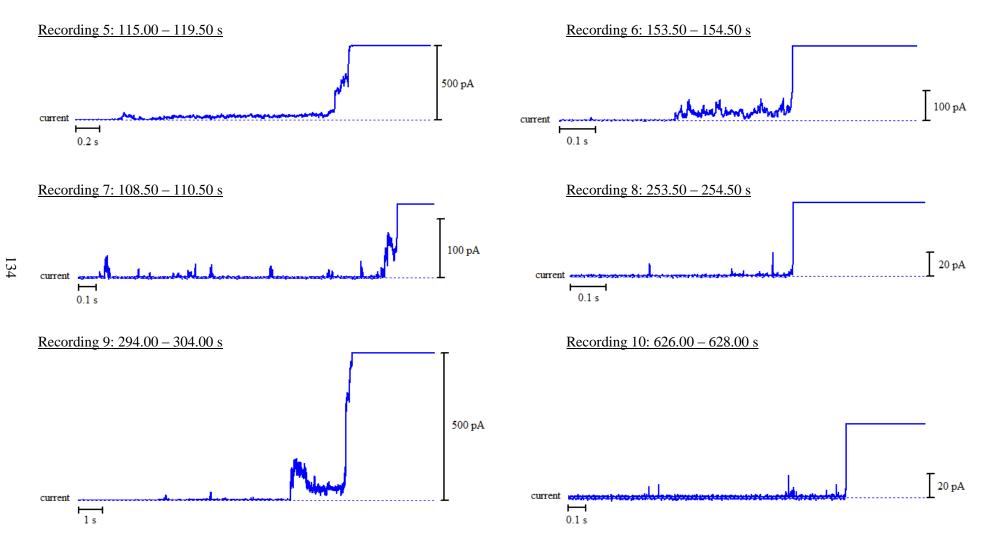
**Figure A2 - 79:** Above: bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of  $25\mu$ l stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.228 \pm 0.0500$  pA, peak  $2 = 50.7 \pm 0.4058$  pA, peak  $3 = 94.28 \pm 4.5320$  pA, peak  $4 = 149.9 \pm 0.2852$  pA, peak  $5 = 330.1 \pm 0.3153$  pA and peak  $6 = 435 \pm 0.4353$  pA.





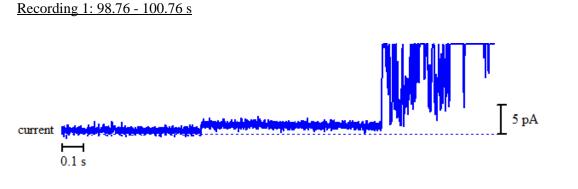


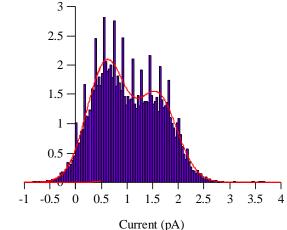
**Figure A2 - 80:** Above: bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **10** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.071 \pm 0.2221$  pA, peak  $2 = 31.5 \pm 0.0511$  pA, peak  $3 = 144.2 \pm 0.0162$  pA, peak  $4 = 241.7 \pm 0.1588$  pA, peak  $5 = 298.1 \pm 0.2119$  pA and peak  $6 = 373.7 \pm 0.4968$  pA.



**Figure A2 - 81:** Bilayer activity of **10** with Na<sup>+</sup> ions upon the addition of  $25\mu$ l (recordings 5, 6 and 7),  $50\mu$ l (recording 8) and  $75\mu$ l (recordings 9 and 10) stock solution of **10** in DMSO.

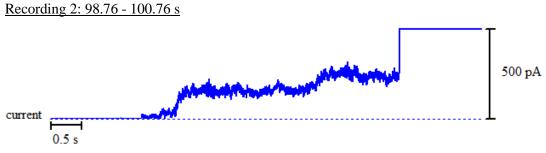
## Planar phospholipid bilayer activity of 10 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system





Percentage (%)

Figure A2 - 82: Above: bilayer activity of 10 with  $K^+$  ions upon the addition of 25µl stock solution of 10 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.5903 \pm$ 0.0295 pA and peak  $2 = 1.584 \pm 0.0417$  pA.



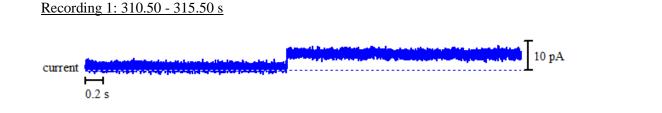
**Figure A2 - 83:** Bilayer activity of **10** with  $K^+$  ions upon the addition of 25µl stock solution of **10** in DMSO.

## Planar phospholipid bilayer activity of 11 towards Na<sup>+</sup> across the POPE and POPS bilayer system

Due to experimental limitations, the ion channel activity of **11** towards Na<sup>+</sup> was unable to be investigated across this bilayer system.

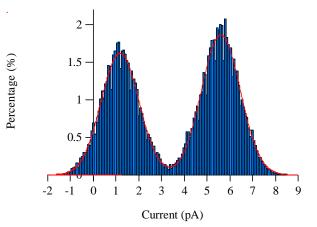
## Planar phospholipid bilayer activity of 11 towards K<sup>+</sup> across the POPE and POPS bilayer system

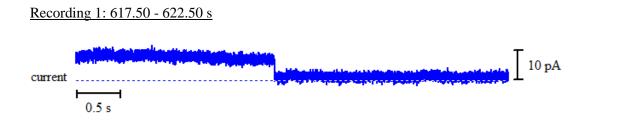
Due to experimental limitations, the ion channel activity of 11 towards  $K^+$  was unable to be investigated across this bilayer system.

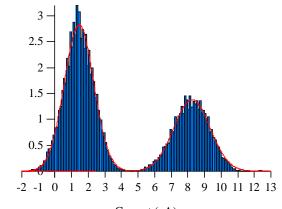


### Planar phospholipid bilayer activity of 11 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

Figure A2 - 84: Above: bilayer activity of 11 with Na<sup>+</sup> ions upon the addition of 75µl stock solution of 11 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.173 \pm 0.0156$  pA and peak  $2 = 5.601 \pm 0.0137$  pA.







Percentage (%)

**Figure A2 - 85:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.439 \pm 0.0102$  pA and peak  $2 = 8.233 \pm 0.0230$  pA.

Current (pA)



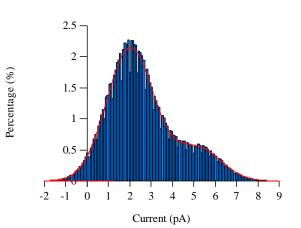
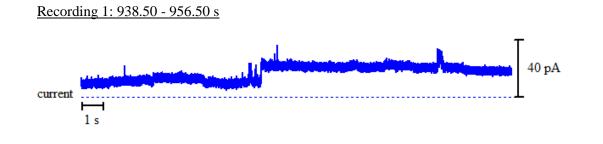


Figure A2 - 86: Above: bilayer activity of 11 with Na<sup>+</sup> ions upon the addition of 150µl stock solution of 11 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.018 \pm$ 0.0229 pA and peak  $2 = 5.162 \pm 0.0919 \text{ pA}$ .





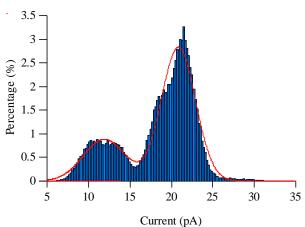
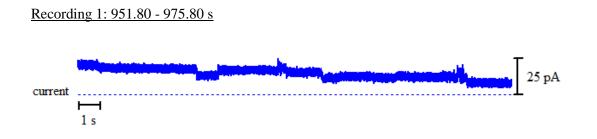
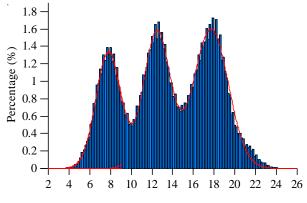


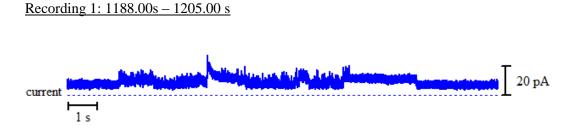
Figure A2 - 87: Above: bilayer activity of 11 with Na<sup>+</sup> ions upon the addition of 175µl stock solution of 11 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 11.89 \pm$ 0.1455 pA and peak  $2 = 20.83 \pm 0.0413$  pA.

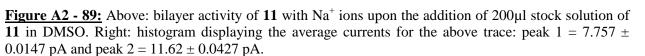


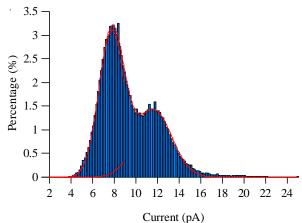


**Figure A2 - 88:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 7.807 \pm 0.0219$  pA, peak  $2 = 12.47 \pm 0.0240$  and peak  $3 = 17.68 \pm 0.0271$  pA.

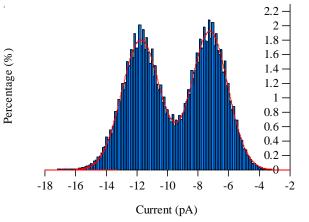






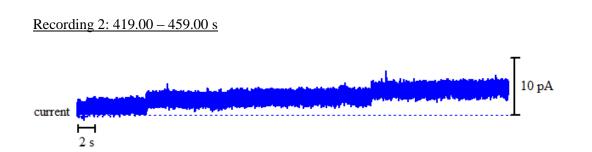


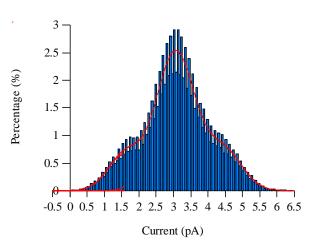




**Figure A2 - 90:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -11.75 \pm 0.0183$  pA and peak  $2 = -7.244 \pm 0.0168$  pA.







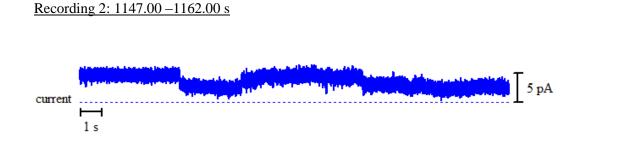
**Figure A2 - 91:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.611 \pm 0.0740$  pA, peak  $2 = 3.054 \pm 0.0347$  pA and peak  $3 = 4.439 \pm 0.2179$  pA.

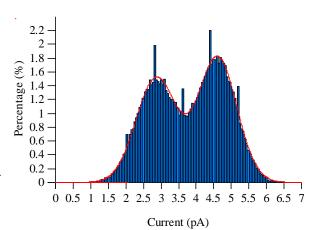


4.5 -4 3.5 2 -1 -0.5 -0. 4 7 8 9 10 11 12 13 14 5 6 Current (pA)

**Figure A2 - 92:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.784 \pm 0.0581$  pA and peak  $2 = 9.733 \pm 0.0083$  pA.

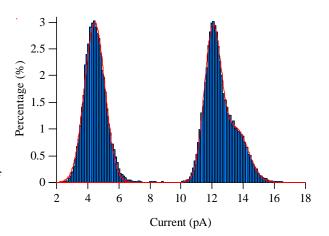
141





**Figure A2 - 93:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.862 \pm 0.0144$  pA and peak  $2 = 4.603 \pm 0.0118$  pA.

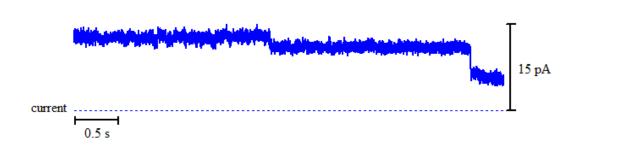


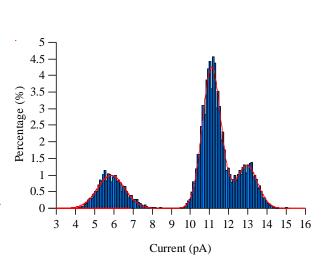


**Figure A2 – 94:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.445 \pm 0.0048$  pA, peak  $2 = 12.03 \pm 0.0109$  pA and peak  $3 = 13.58 \pm 0.0474$  pA.

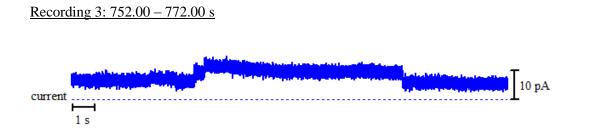
142

Recording 2: 1403.00 - 1407.00 s





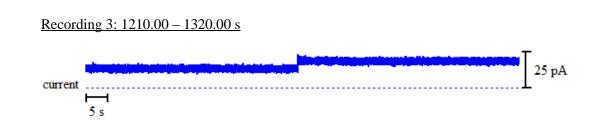
**Figure A2 – 95:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 5.845 \pm 0.0396$  pA, peak  $2 = 11.1 \pm 0.0097$  pA and peak  $3 = 12.94 \pm 0.0337$  pA.



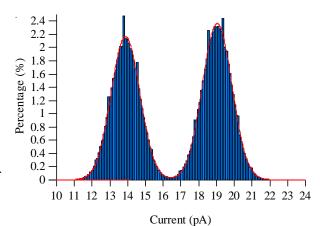
2 - $1.8 \cdot$ 1.6 - 6.1 - 2.1 - 1.4 - 1.5 - 1.6 0.4 -0.2 -0 2 3 4 5 8 9 10 11 12 13 14 15 6 7 Current (pA)

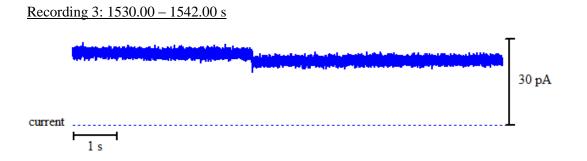
**Figure A2 - 96:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.342 \pm 0.0204$  pA and peak  $2 = 10.06 \pm 0.0265$  pA.

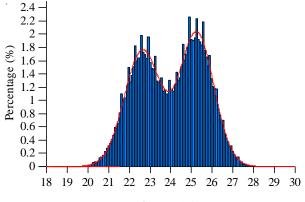




**Figure A2 - 97:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 13.87 \pm 0.0077$  pA and peak  $2 = 19.03 \pm 0.0071$  pA.



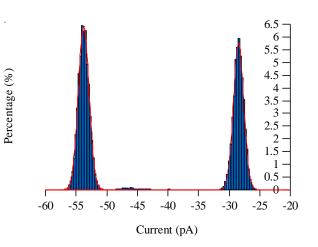




**Figure A2 - 98:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 22.62 \pm 0.0231$  pA and peak  $2 = 25.24 \pm 0.0195$  pA.



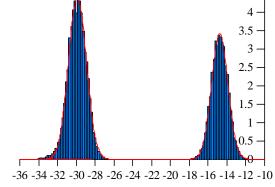




**Figure A2 - 99:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -53.77 \pm 0.0061$  pA and peak  $2 = -28.45 \pm 0.0066$  pA.



**Figure A2 - 100:** Above: bilayer activity of **11** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **11** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -29.88 \pm$ 



Percentage (%)

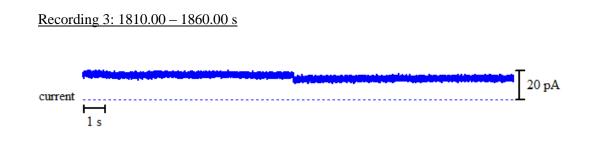
4.5

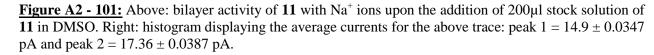
Current (pA)

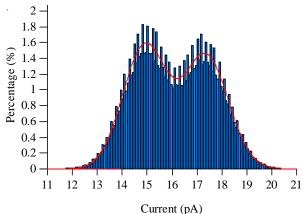


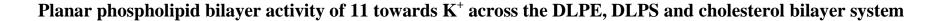


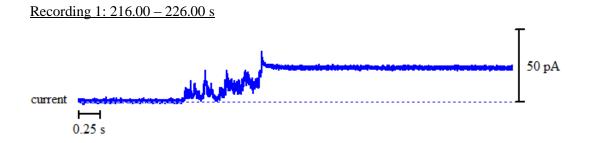
0.0068 pA and peak  $2 = -14.77 \pm 0.0085$  pA.



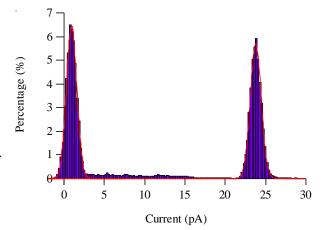


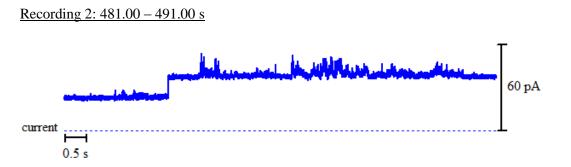


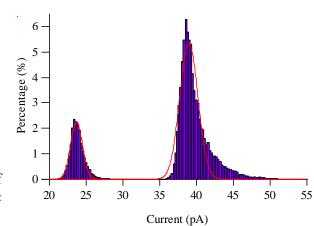




**Figure A2 - 102:** Above: bilayer activity of 11 with  $K^+$  ions upon the addition of 75µl stock solution of 11 in DMSO. Right: histogram displaying the average currents for the above trace: peak 1 = 0.923 ± 0.0067 pA and peak 2 = 23.77 ± 0.0077 pA.







**Figure A2 - 103:** Above: bilayer activity of 11 with K<sup>+</sup> ions upon the addition of 75µl stock solution of 11 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 23.86 \pm 0.0798$  pA and peak  $2 = 38.95 \pm 0.0397$  pA.



<u>Recording 3: 1006.10 – 1058.60 s</u>

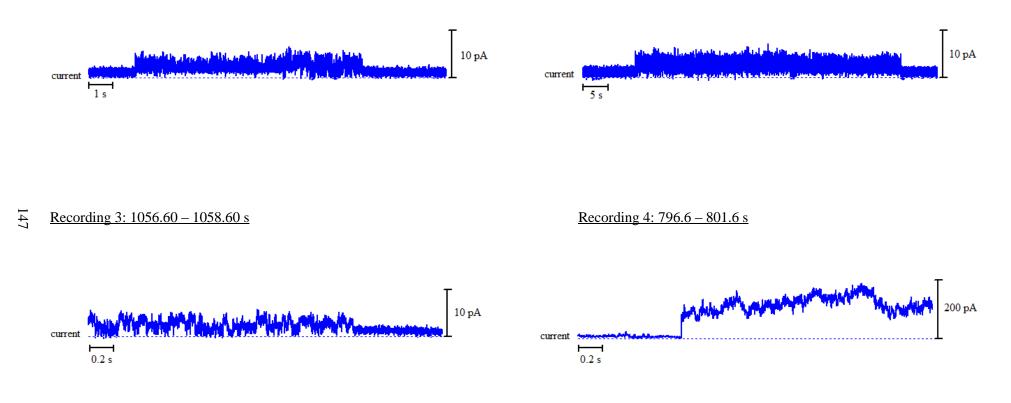
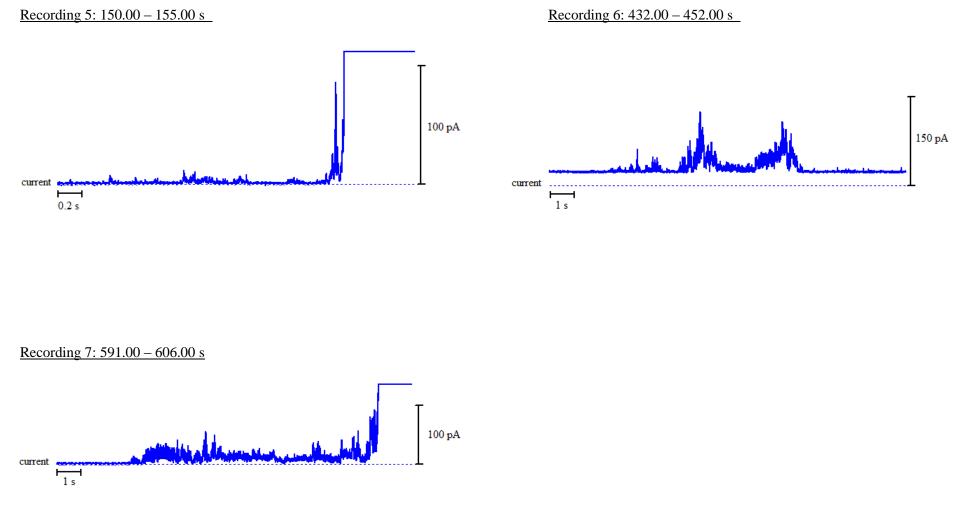


Figure A2 - 104: Bilayer activity of 11 with K<sup>+</sup> ions upon the addition of 200µl (recordings 3) and 150µl (recording 4) stock solution of 11 in DMSO.

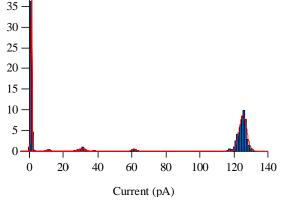


<u>Figure A2 - 105</u>: Bilayer activity of 11 with  $K^+$  ions upon the addition of 25µl (recording 5), 75µl (recording 6) and 100µl (recording 7) stock solution of 11 in DMSO.

### Planar phospholipid bilayer activity of 12 towards Na<sup>+</sup> across the POPE and POPS bilayer system

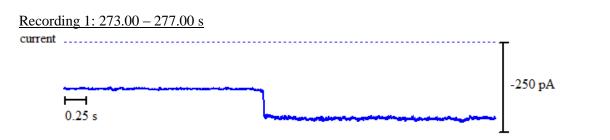
# 150 pA current 0.05 s

5 -0 Figure A2 - 106: Above: bilayer activity of 12 with Na<sup>+</sup> ions upon the addition of 100µl stock solution 0 of 12 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.414 \pm$ 0.0025 pA, peak 2 =  $10.92 \pm 0.2131$  pA, peak 3 =  $30.73 \pm 0.6173$  pA, peak 4 =  $61.16 \pm 0.9055$  pA and



40

Percentage (%)



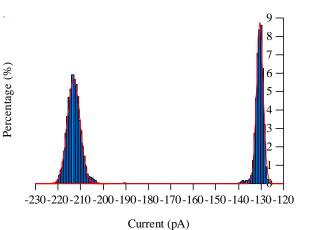


Figure A2 - 107: Above: bilayer activity of 12 with Na<sup>+</sup> ions upon the addition of 100µl stock solution of 12 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -213.1 \pm$ 0.0399 pA and peak  $2 = -130.5 \pm 0.0250$  pA.

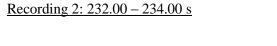
peak  $5 = 125.2 \pm 0.0628$  pA.

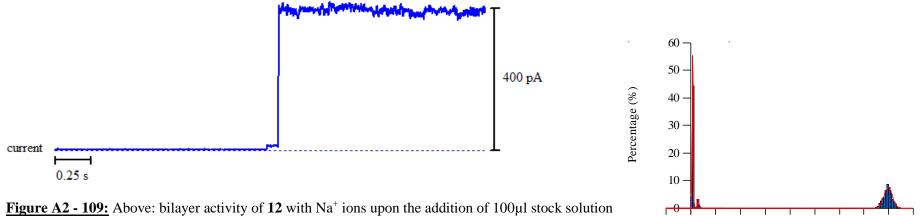
Recording 1: 258.00 – 259.00 s



**Figure A2 - 108:** Bilayer activity of **12** with Na<sup>+</sup> ions upon the addition of 100 $\mu$ l stock solution of **12** in DMSO.





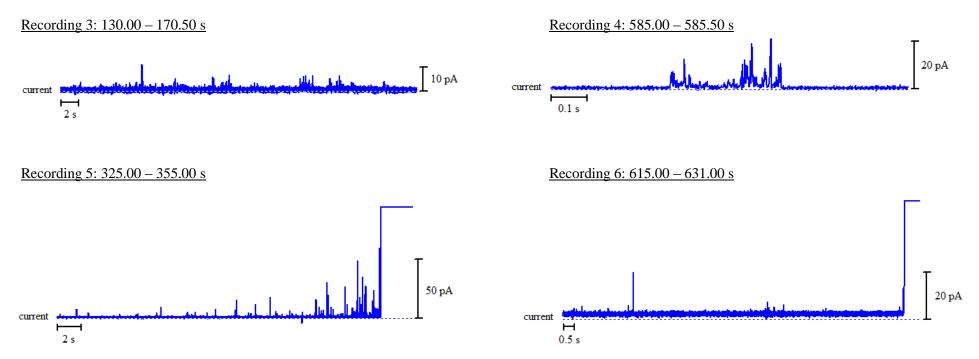


**Figure A2 - 109:** Above: bilayer activity of **12** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **12** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.207 \pm 0.1242$  pA, peak  $2 = 15.17 \pm 0.2261$  pA and peak  $3 = 398.7 \pm 0.1815$  pA.

Current (pA)

-50 0

50 100 150 200 250 300 350 400 450



**Figure A2 - 110:** Bilayer activity of **12** with Na<sup>+</sup> ions upon the addition of  $50\mu$ l (recording 3),  $125\mu$ l (recordings 4 and 5) and  $150\mu$ l (recording 6) stock solution of **12** in DMSO.

## Planar phospholipid bilayer activity of 12 towards K<sup>+</sup> across the POPE and POPS bilayer system

Due to experimental limitations, the ion channel activity of 12 towards  $K^+$  was unable to be investigated across this bilayer system.

## Planar phospholipid bilayer activity of 12 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

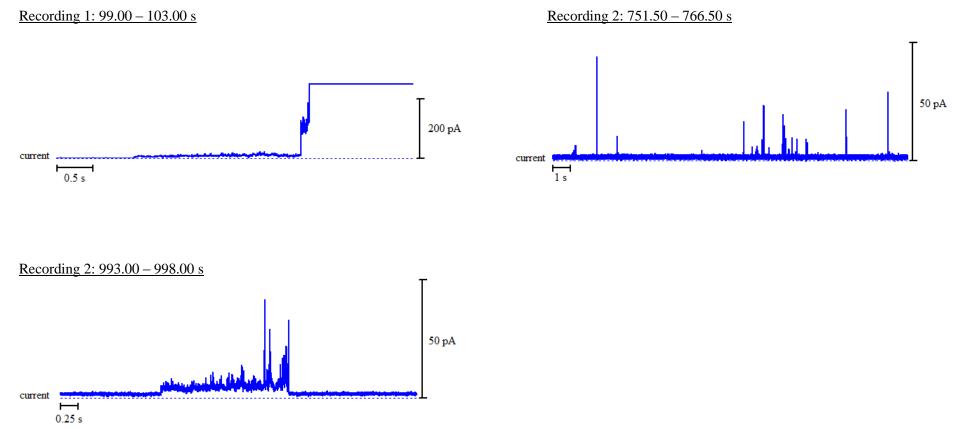
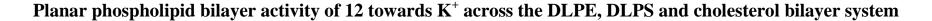
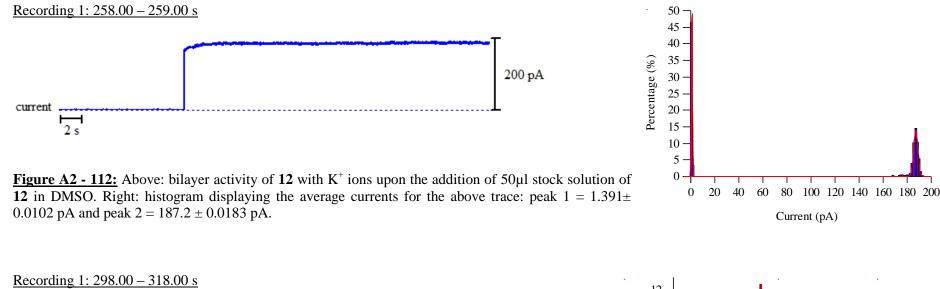
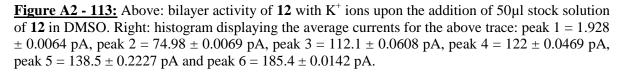


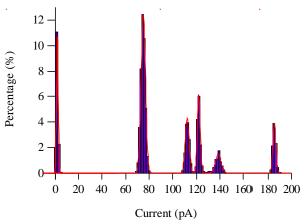
Figure A2 - 111: Bilayer activity of 12 with Na<sup>+</sup> ions upon the addition of 25µl (recording 1) and 200µl (recordings 2) stock solution of 12 in DMSO.



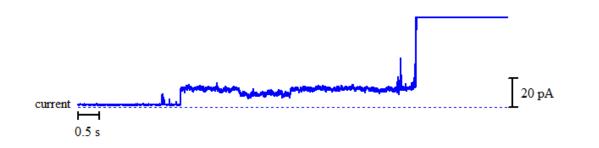




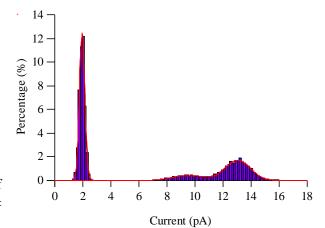




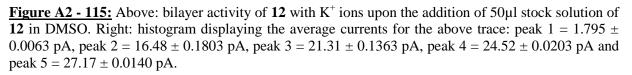
Recording 2: 320.00 - 335.00 s

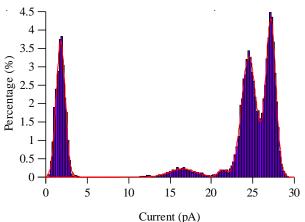


**Figure A2 - 114:** Above: bilayer activity of **12** with K<sup>+</sup> ions upon the addition of 75µl stock solution of **12** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.951 \pm 0.0017$  pA, peak  $2 = 9.542 \pm 0.1378$  pA and peak  $3 = 13.04 \pm 0.0312$  pA.



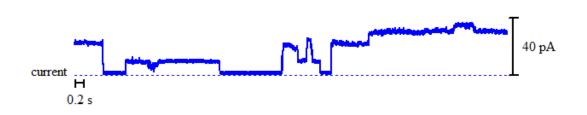


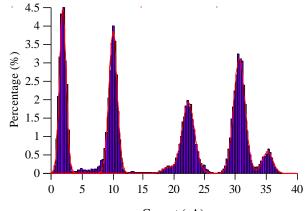






Recording 3: 315.50 - 340.50 s



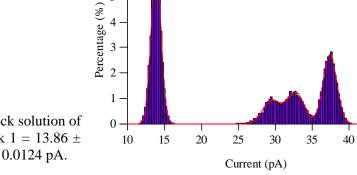


**Figure A2 - 116:** Above: bilayer activity of **12** with K<sup>+</sup> ions upon the addition of 50µl stock solution of **12** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.886 \pm 0.0109$  pA, peak  $2 = 9.962 \pm 0.0139$  pA, peak  $3 = 19.06 \pm 0.2169$  pA, peak  $4 = 22.33 \pm 0.0192$  pA, peak  $5 = 30.66 \pm 0.0151$  pA and peak  $6 = 35.16 \pm 0.0774$  pA.

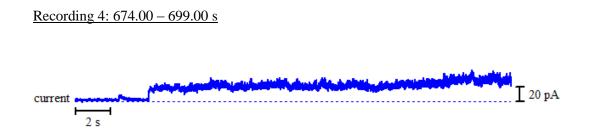


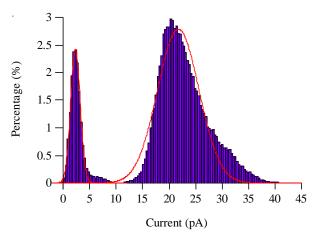
45





**Figure A2 - 117:** Above: bilayer activity of **12** with K<sup>+</sup> ions upon the addition of 50µl stock solution of **12** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 13.86 \pm 0.0052$  pA, peak  $2 = 29.47 \pm 0.0388$  pA, peak  $3 = 32.72 \pm 0.0304$  pA and peak  $4 = 37.4 \pm 0.0124$  pA.





**Figure A2 - 118:** Above: bilayer activity of **12** with K<sup>+</sup> ions upon the addition of 150µl stock solution of **12** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.284 \pm 0.0599$  pA and peak  $2 = 21.7 \pm 0.1054$  pA.



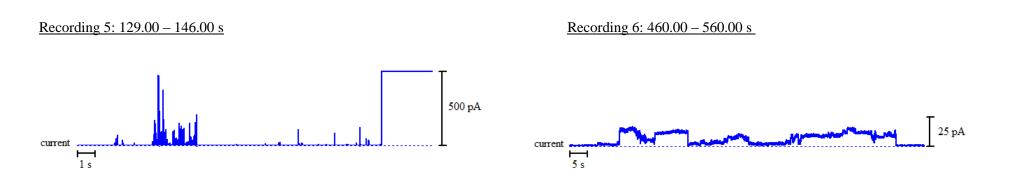


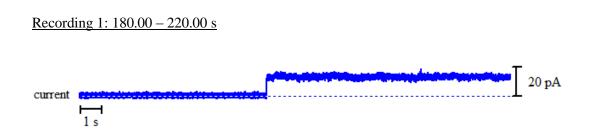
Figure A2 - 119: Bilayer activity of 12 with K<sup>+</sup> ions upon the addition of 25µl (recording 5) and 50µl (recording 6) stock solution of 12 in DMSO.

## Planar phospholipid bilayer activity of 13 towards Na<sup>+</sup> across the POPE and POPS bilayer system

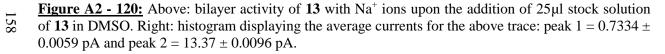
Due to experimental limitations, the ion channel activity of 13 towards Na<sup>+</sup> was unable to be investigated across this bilayer system.

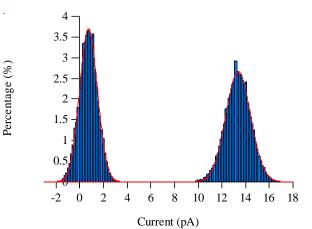
## Planar phospholipid bilayer activity of 13 towards K<sup>+</sup> across the POPE and POPS bilayer system

Due to experimental limitations, the ion channel activity of 13 towards  $K^+$  was unable to be investigated across this bilayer system.

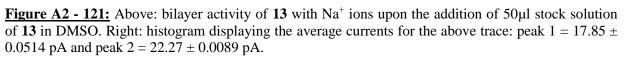


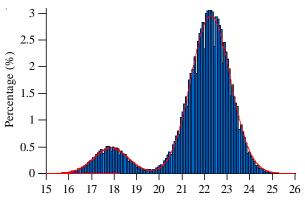
## Planar phospholipid bilayer activity of 13 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



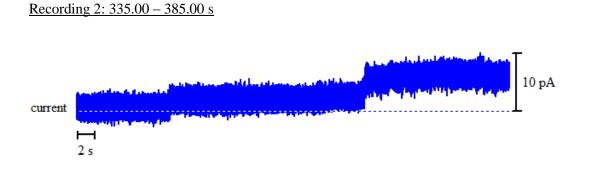


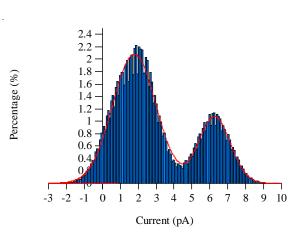
Recording 1: 235.00 – 335.00 s 30 pA5 s





Current (pA)

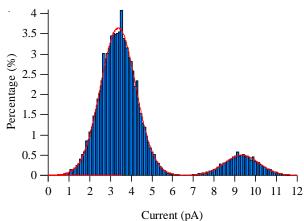


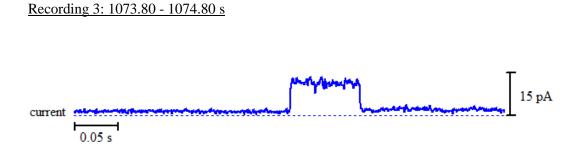


**Figure A2 - 122:** Above: bilayer activity of **13** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **13** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.776 \pm 0.0160$  pA and peak  $2 = 6.249 \pm 0.0267$  pA.



**Figure A2 - 123:** Above: bilayer activity of **13** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **13** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.37 \pm 0.0059$  pA and peak  $2 = 9.311 \pm 0.0427$  pA.



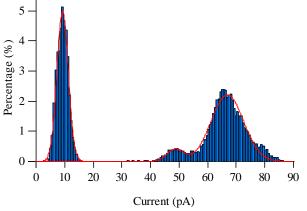


 $\begin{array}{c} 8 \\ 7 \\ 6 \\ 6 \\ 9 \\ 5 \\ - \\ 0 \\ 1 \\ 2 \\ - \\ 1 \\ 0 \\ 0 \\ 1 \\ 2 \\ 3 \\ 4 \\ 5 \\ 6 \\ 7 \\ 8 \\ 9 \\ 10 \\ 11 \\ 12 \\ 13 \\ 14 \\ \end{array}$ 

Current (pA)

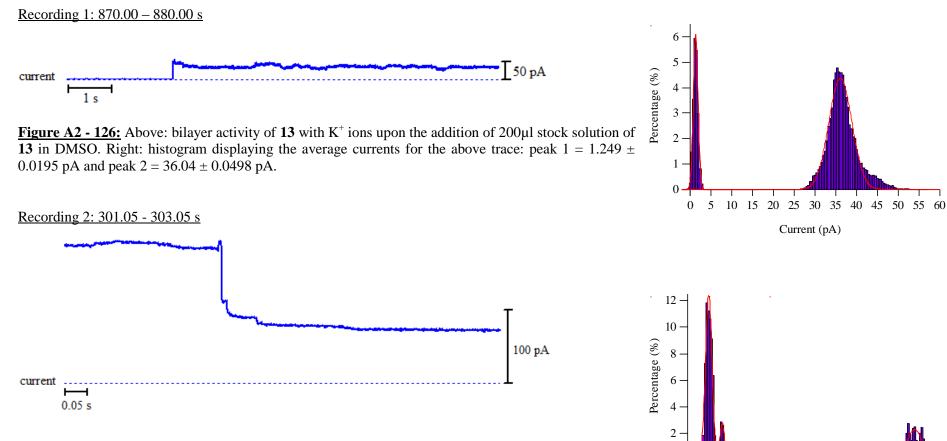
**Figure A2 - 124:** Above: bilayer activity of **13** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **13** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.729 \pm 0.0118$  pA and peak  $2 = 11.3 \pm 0.2743$  pA.

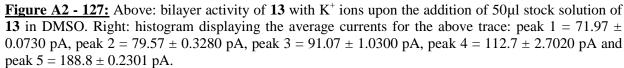




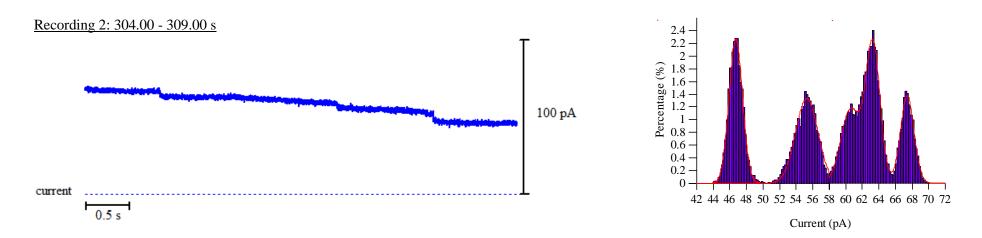
**Figure A2 - 125:** Above: bilayer activity of **13** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **13** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 9.247 \pm 0.0034$  pA, peak  $2 = 48.57 \pm 0.5281$  pA and peak  $3 = 66.81 \pm 0.1317$  pA.

Planar phospholipid bilayer activity of 13 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

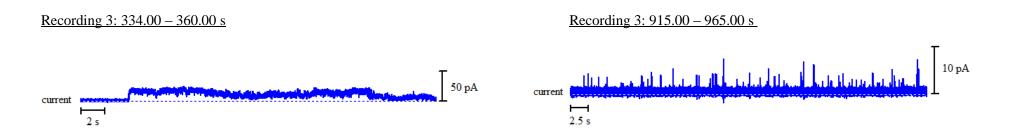




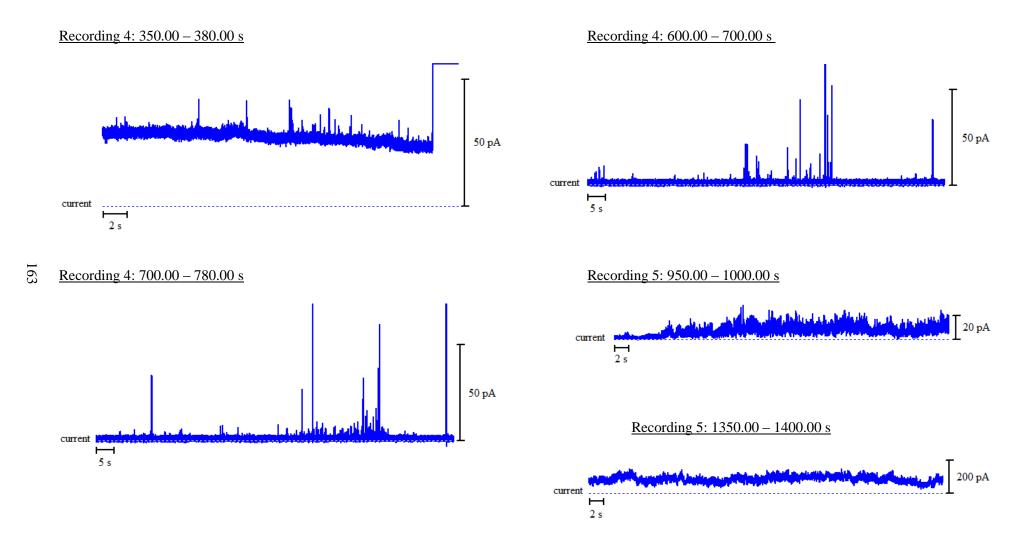
Current (pA)



 $\overline{5}$  Figure A2 - 128: Above: bilayer activity of 13 with K<sup>+</sup> ions upon the addition of 50µl stock solution of 13 in DMSO. Right: histogram displaying the average currents for the above trace: peak 1 = 46.7 ± 0.0149 pA, peak 2 = 55.26 ± 0.0307 pA, peak 3 = 60.58 ± 0.0380 pA, peak 4 = 63.27 ± 0.0184 pA and peak 5 = 67.42 ± 0.0287 pA.



**Figure A2 - 129:** Bilayer activity of **13** with  $K^+$  ions upon the addition of 50µl (recording 3- left) and 225µl (recording 3- right) stock solution of **13** in DMSO.

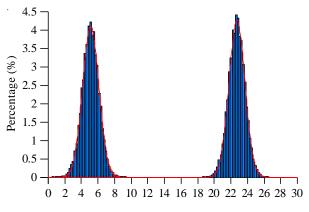


**Figure A2 - 130:** Bilayer activity of **13** with  $K^+$  ions upon the addition of 150ul (recordings 4) and 175µl and 200µl (recordings 5) stock solution of **13** in DMSO.

### Planar phospholipid bilayer activity of 14 towards Na<sup>+</sup> across the POPE and POPS bilayer system

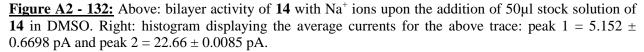


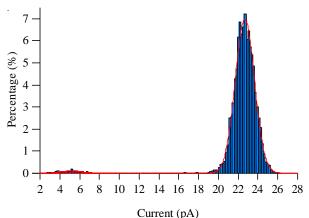
**Figure A2 - 131:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 5.113 \pm 0.0071$  pA and peak  $2 = 22.72 \pm 0.0068$  pA.

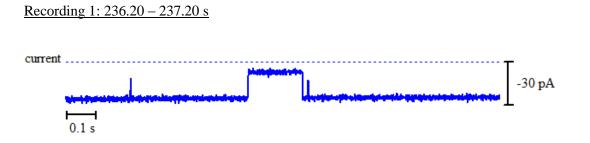




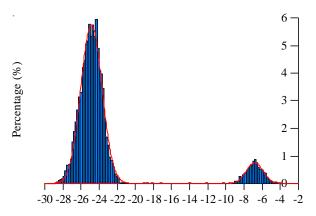




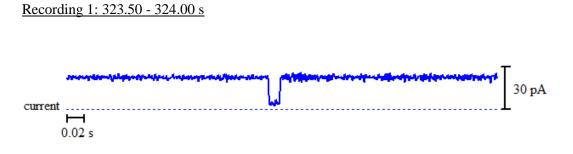


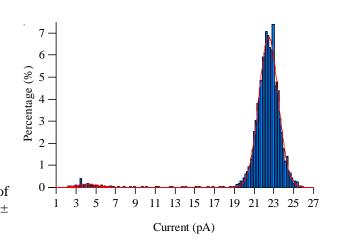


**Figure A2 - 133:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -24.86 \pm 0.0110$  pA and peak  $2 = -6.856 \pm 0.0760$  pA.

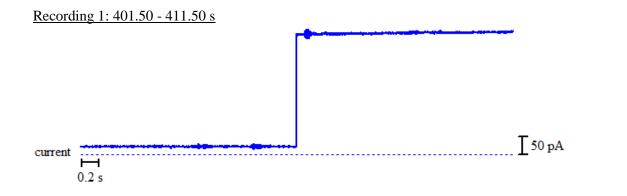


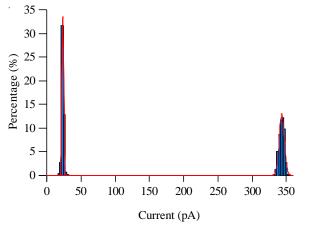
Current (pA)



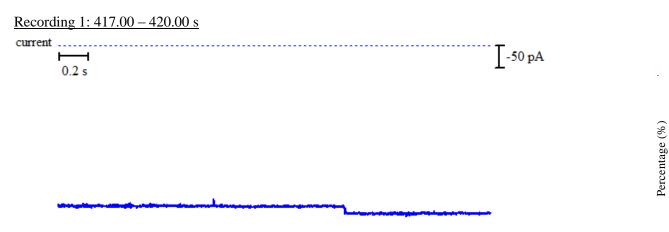


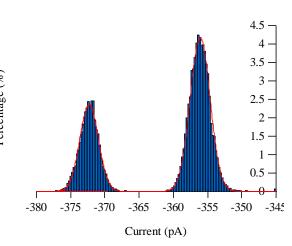
**Figure A2 - 134:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.223 \pm 1.13$  pA and peak  $2 = 22.44 \pm 0.0115$  pA.



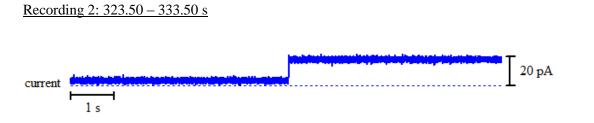


**Figure A2 - 135:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 23.39 \pm 0.0305$  pA and peak  $2 = 343.1 \pm 0.0665$  pA.

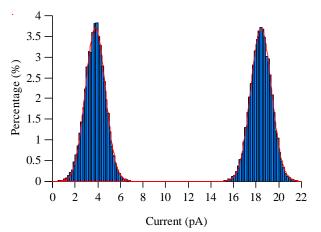




**Figure A2 - 136:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -372.3 \pm 0.0492$  pA and peak  $2 = -356.1 \pm 0.0268$  pA.

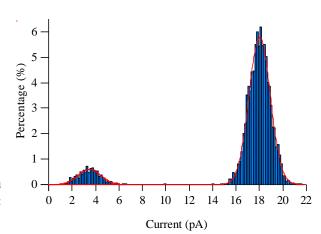


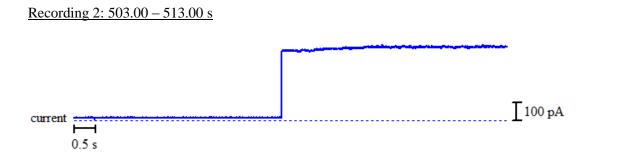
**Figure A2 - 137:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.782 \pm 0.0051$  pA and peak  $2 = 18.43 \pm 0.0052$  pA.



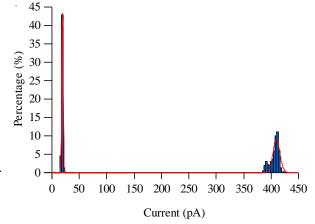


**Figure A2 - 138:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.464 \pm 0.0883$  pA and peak  $2 = 18.01 \pm 0.0089$  pA.

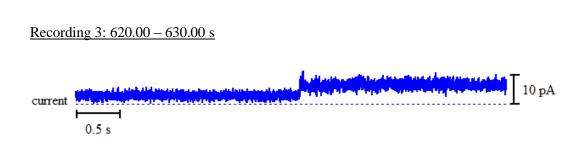


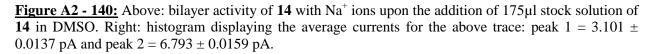


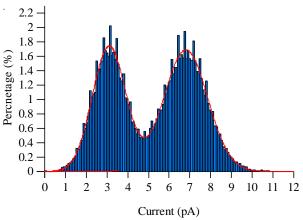
**Figure A2 - 139:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 19.18 \pm 0.0857$  pA and peak  $2 = 409.7 \pm 0.3002$  pA.

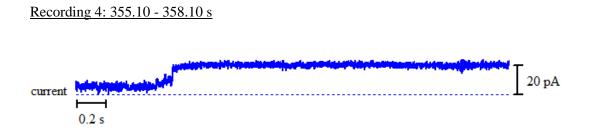


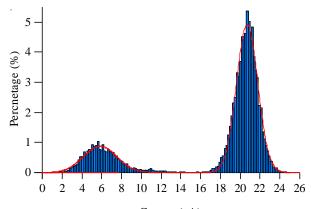








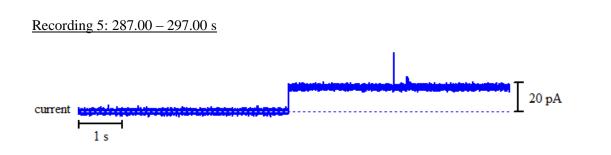




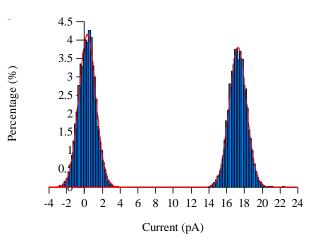
Current (pA)

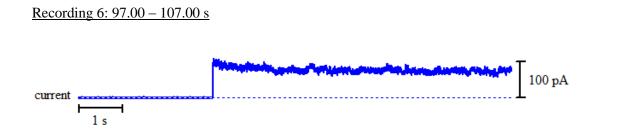
**Figure A2 - 141:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 5.813 \pm 0.0756$  pA and peak  $2 = 20.68 \pm 0.0109$  pA.

169

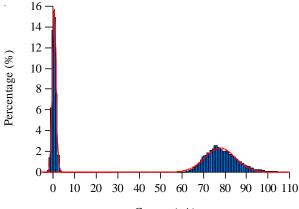


**Figure A2 - 142:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.3026 \pm 0.0086$  pA and peak  $2 = 17.25 \pm 0.0095$  pA.

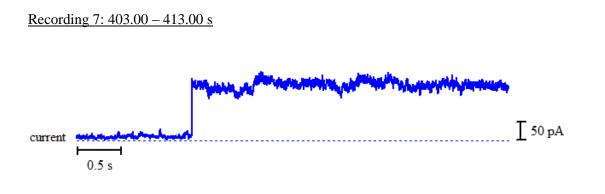




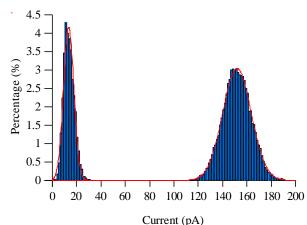
**Figure A2 - 143:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.4345 \pm 0.0070$  pA and peak  $2 = 77.52 \pm 0.0996$  pA.

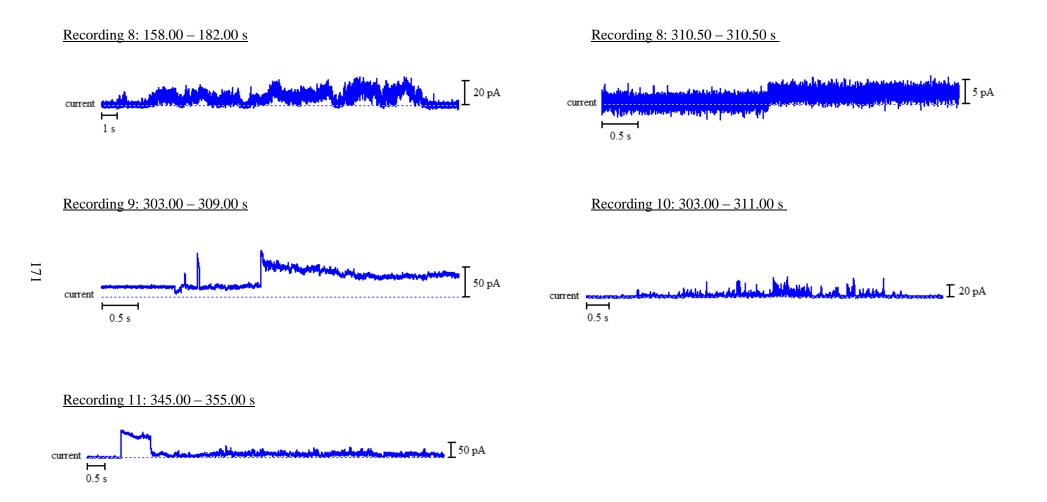






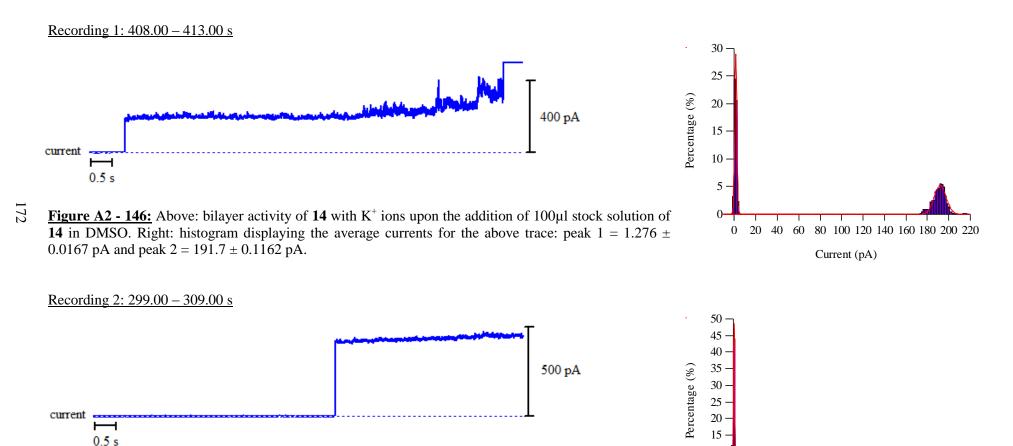
**Figure A2 - 144:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 13.4 \pm 0.0695$  pA and peak  $2 = 151.7 \pm 0.1517$  pA.

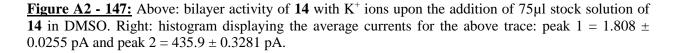


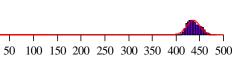


**Figure A2 - 145:** Bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of  $25\mu$ l (recording 8- left), 75ul (recording 8- right) and 100µl (recordings 9, 10 and 11) stock solution of **13** in DMSO.

#### Planar phospholipid bilayer activity of 14 towards K<sup>+</sup> across the POPE and POPS bilayer system





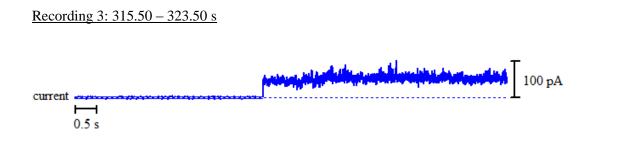


Current (pA)

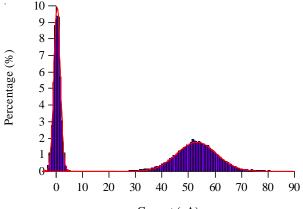
10 -

5 -

0-

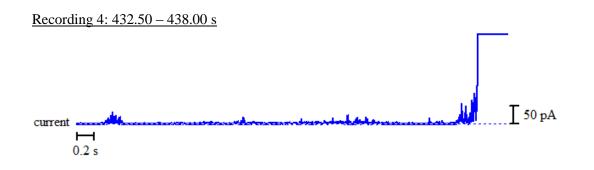


**Figure A2 - 148:** Above: bilayer activity of **14** with K<sup>+</sup> ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.4798 \pm 0.0088$  pA and peak  $2 = 53.0 \pm 0.1112$  pA.



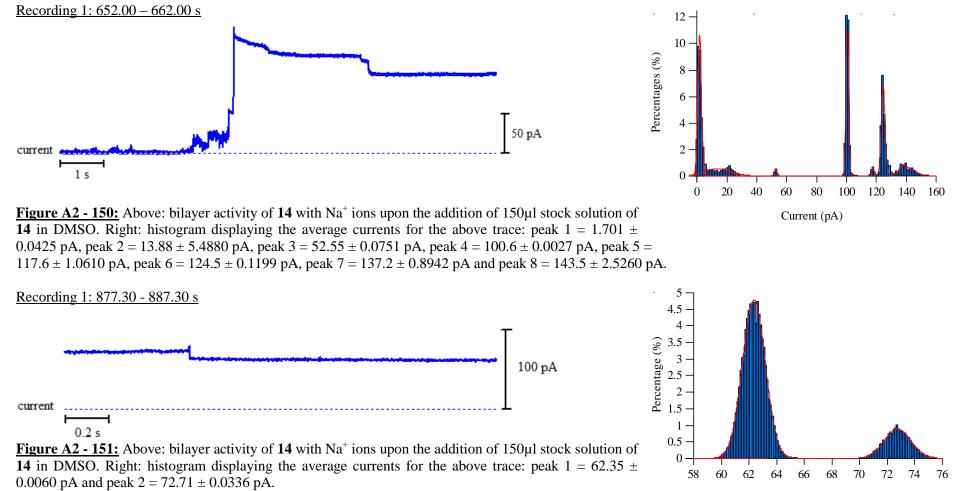
Current (pA)

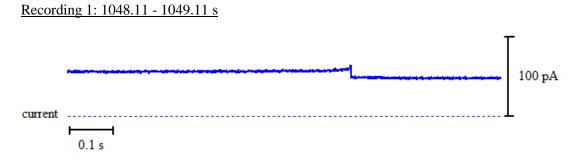


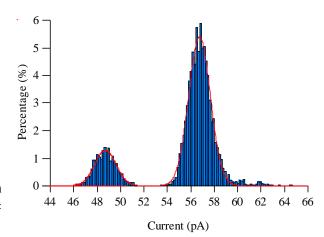


**Figure A2 - 149:** Bilayer activity of **14** with  $K^+$  ions upon the addition of 100µl stock solution of **13** in DMSO.

#### Planar phospholipid bilayer activity of 14 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system







**Figure A2 - 152:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 48.69 \pm 0.0494$  pA and peak  $2 = 56.74 \pm 0.0119$  pA.

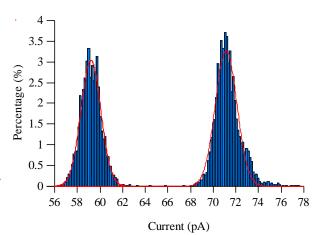


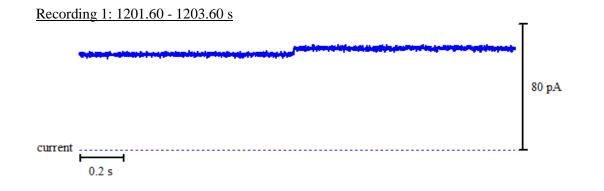


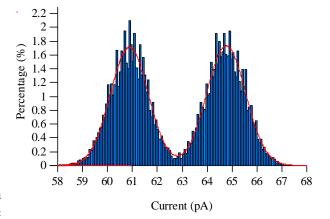
Recording 1: 1161.10 - 1162.10 s



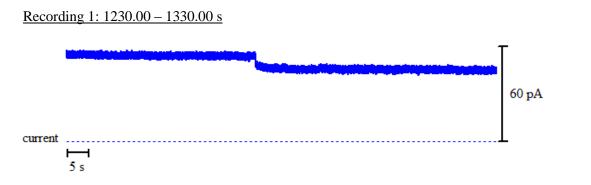
**Figure A2 - 153:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 59.19 \pm 0.0268$  pA and peak  $2 = 71.13 \pm 0.0252$  pA.

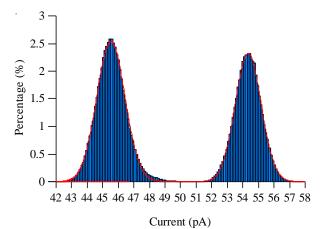






**Figure A2 - 154:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 60.87 \pm 0.0218$  pA and peak  $2 = 64.76 \pm 0.0224$  pA.

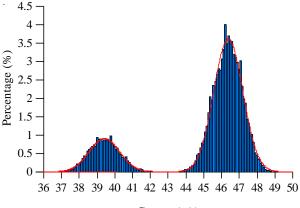




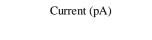
**Figure A2 - 155:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 60.87 \pm 0.0218$  pA and peak  $2 = 64.76 \pm 0.0224$  pA.

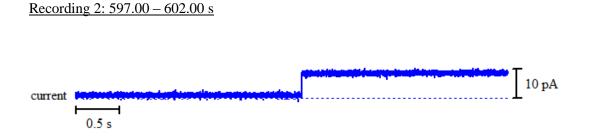
Recording 1: 1398.5 - 1403.5 s



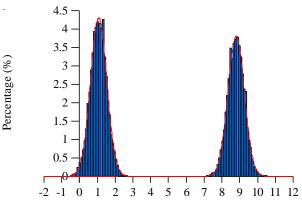


**Figure A2 - 156:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 200µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 39.39 \pm 0.0319$  pA and peak  $2 = 46.37 \pm 0.0079$  pA.

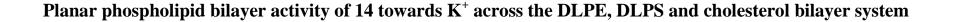


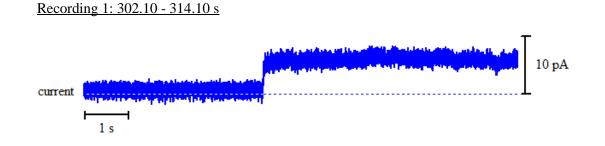


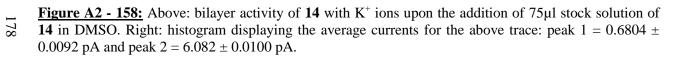
**Figure A2 - 157:** Above: bilayer activity of **14** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.069 \pm 0.0044$  pA and peak  $2 = 8.796 \pm 0.0050$  pA.

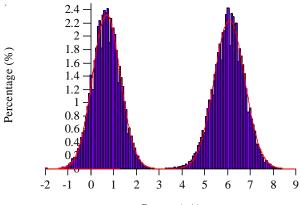


Current (pA)

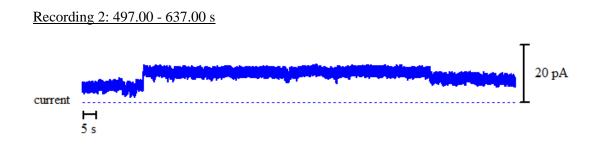


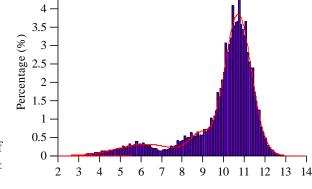










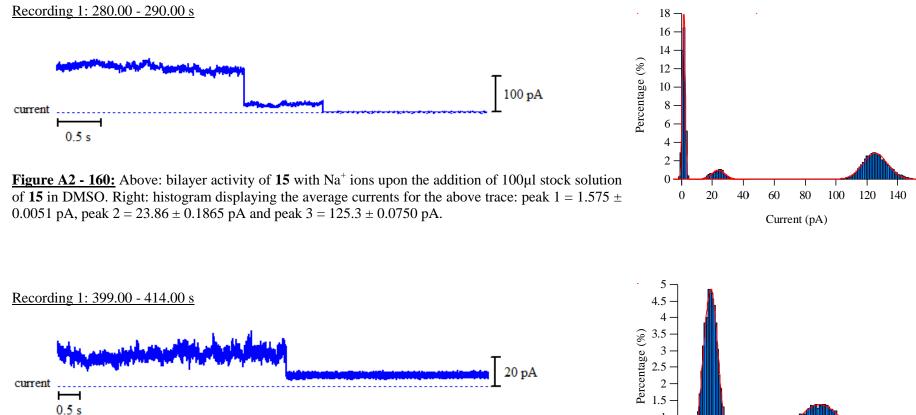


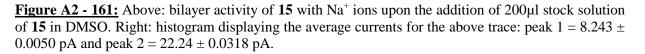
4.5

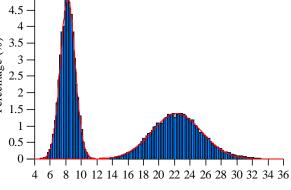
**Figure A2 - 159:** Above: bilayer activity of **14** with  $K^+$  ions upon the addition of 75µl stock solution of **14** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 6.236 \pm 0.0597$  pA and peak  $2 = 8.833 \pm 0.1513$  pA.

Current (pA)

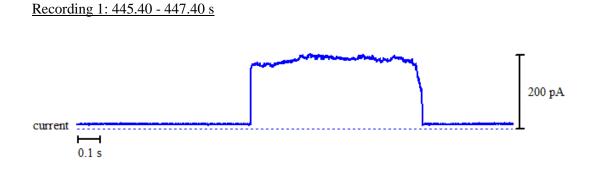
## Planar phospholipid bilayer activity of 15 towards Na<sup>+</sup> across the POPE and POPS bilayer system







Current (pA)



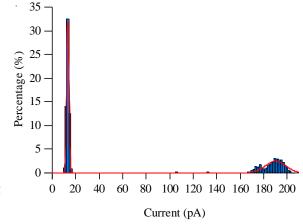


Figure 3.2.31: Above: bilayer activity of 15 with Na<sup>+</sup> ions upon the addition of 200µl stock solution of 15 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 13.26 \pm 0.0170$ pA and peak  $2 = 189.9 \pm 0.3542$  pA.

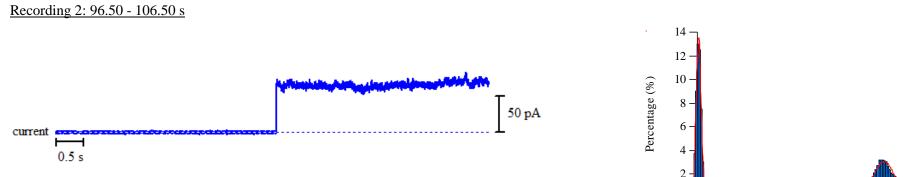
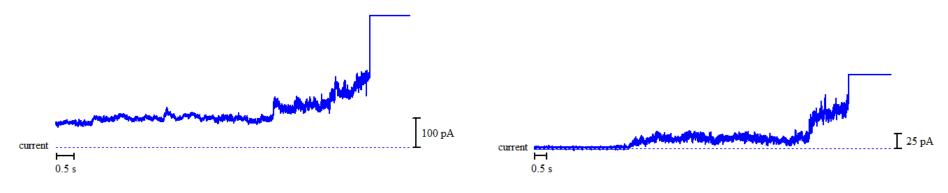


Figure A2 - 162: Above: bilayer activity of 15 with Na<sup>+</sup> ions upon the addition of 50µl stock solution of 15 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.3315 \pm$ 0.0049 pA and peak  $2 = 66.41 \pm 0.0376$  pA.

 $-5 \ 0 \ 5 \ 10 \ 15 \ 20 \ 25 \ 30 \ 35 \ 40 \ 45 \ 50 \ 55 \ 60 \ 65 \ 70 \ 75 \ 80 \ 85$ Current (pA)

<u>Recording 3: 120.00 – 130.00 s</u>

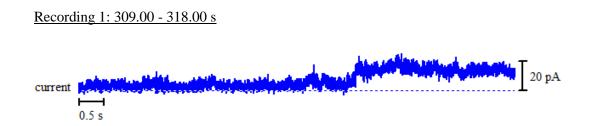
<u>Recording 4: 493.00 – 507.00 s</u>



181

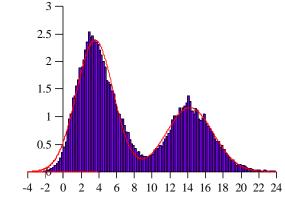
Figure A2 - 163: Bilayer activity of 15 with Na<sup>+</sup> ions upon the addition of 50µl (recording 3) and 100µl (recording 4) stock solution of 15 in DMSO.

### Planar phospholipid bilayer activity of 15 towards K<sup>+</sup> across the POPE and POPS bilayer system



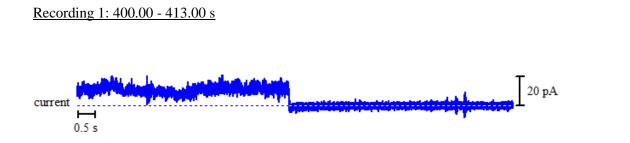
182

**Figure A2 - 164:** Above: bilayer activity of **15** with  $K^+$  ions upon the addition of 75µl stock solution of **15** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.536 \pm 0.0260$  pA and peak  $2 = 14.21 \pm 0.0598$  pA.

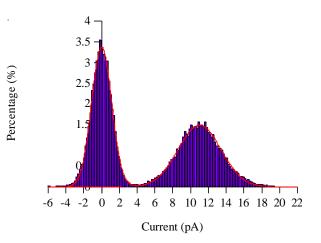


Percentage (%)

Current (pA)



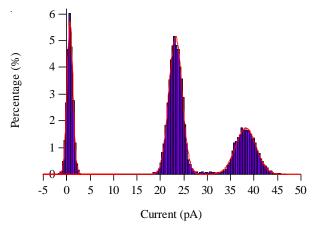
**Figure A2 - 165:** Above: bilayer activity of **15** with  $K^+$  ions upon the addition of 75µl stock solution of **15** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -0.0017 \pm 0.0069$  pA and peak  $2 = 10.98 \pm 0.0227$  pA.



Recording 2: 913.00 - 918.00 s



**Figure A2 - 166:** Above: bilayer activity of **15** with K<sup>+</sup> ions upon the addition of 250µl stock solution of **15** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.6839 \pm 0.0068$  pA, peak  $2 = 23.22 \pm 0.0114$  pA and peak  $3 = 38.18 \pm 0.04161$  pA.





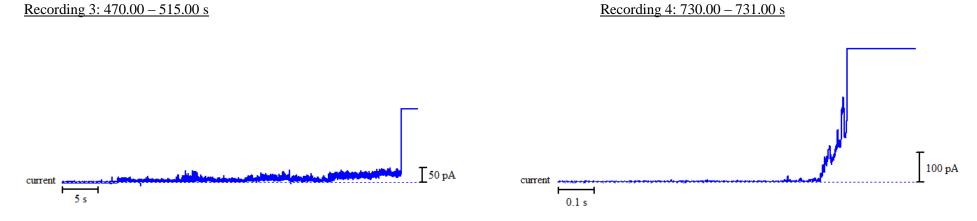


Figure A2 - 167: Bilayer activity of 15 with K<sup>+</sup> ions upon the addition of 100µl (recording 3) and 175µl (recording 4) stock solution of 15 in DMSO.

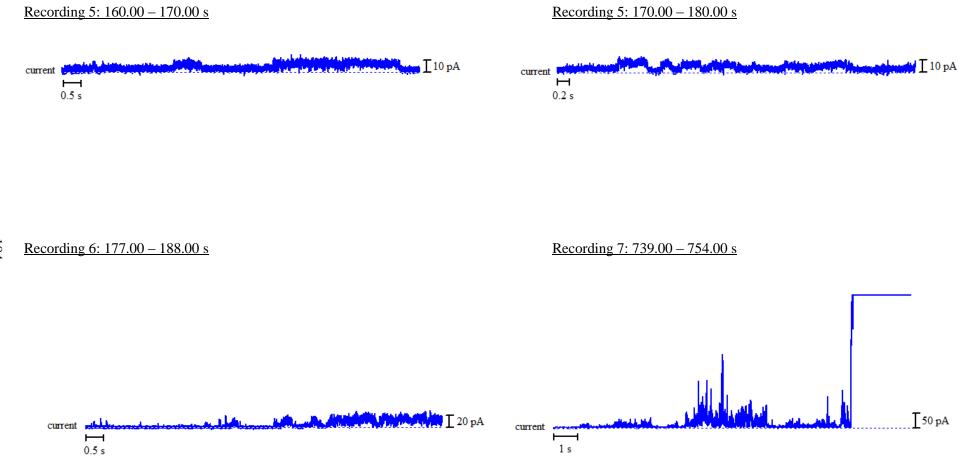


Figure A2 - 168: Bilayer activity of 15 with K<sup>+</sup> ions upon the addition of 25µl (recordings 5 and 6) and 175µl (recording 7) stock solution of 15 in DMSO.

# Planar phospholipid bilayer activity of 15 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

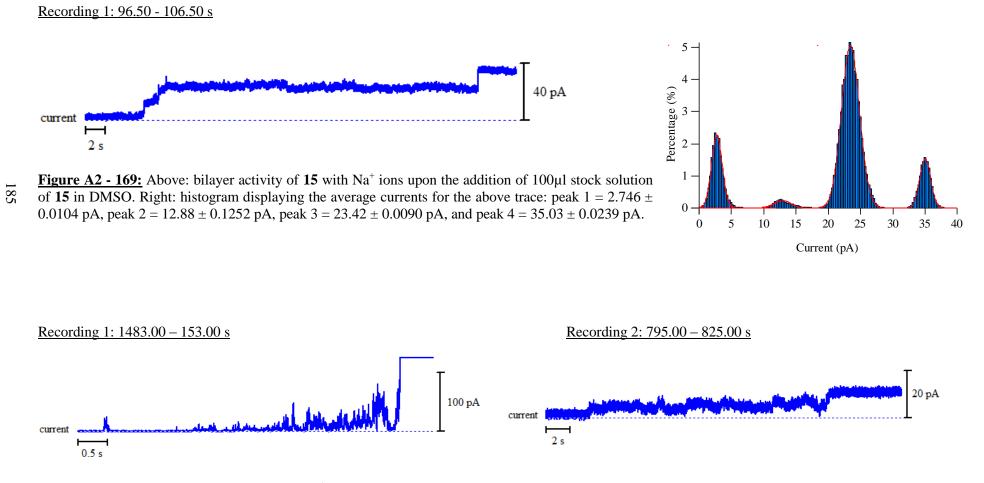
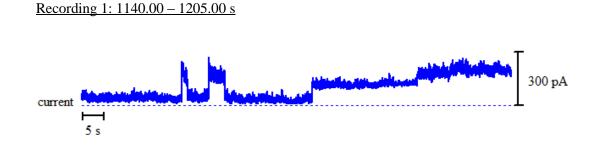


Figure A2 - 170: Bilayer activity of 15 with Na<sup>+</sup> ions upon the addition of 25µl (recording 1) and 175µl (recording 2) stock solution of 15 in DMSO.

Planar phospholipid bilayer activity of 15 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



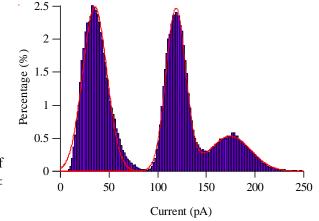


Figure A2 - 171: Above: bilayer activity of 15 with K<sup>+</sup> ions upon the addition of 175µl stock solution of 15 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 35.34 \pm 0.1766$  pA, peak  $2 = 118.6 \pm 0.182$  pA and peak  $3 = 174.2 \pm 1.214$  pA.

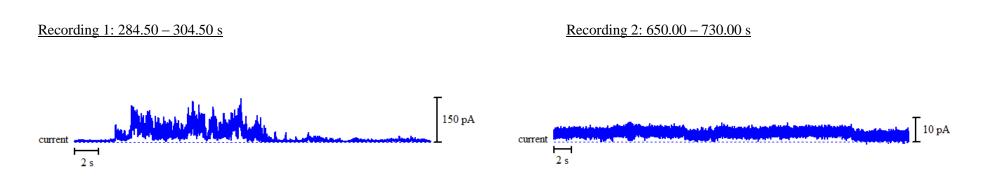
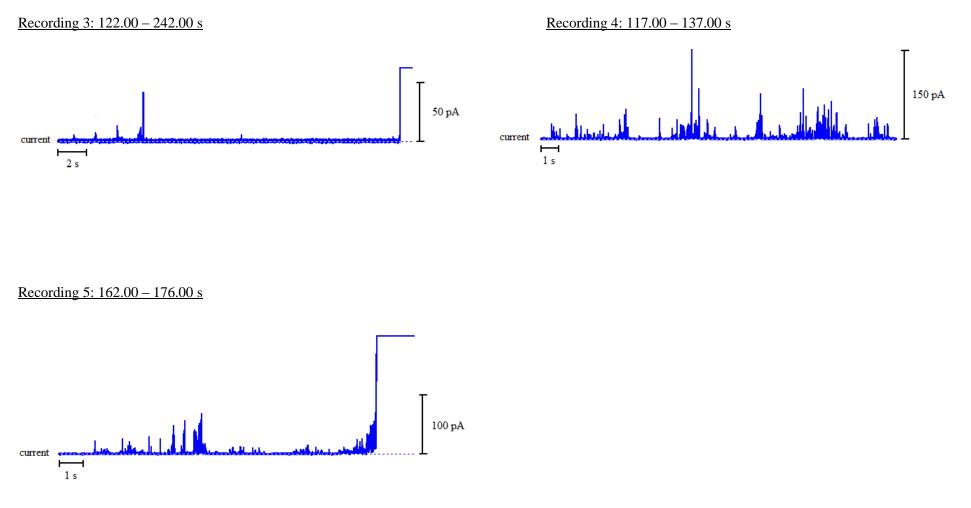


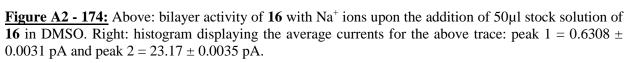
Figure A2 - 172: Bilayer activity of 15 with K<sup>+</sup> ions upon the addition of 50µl (recording 1) and 150µl (recording 2) stock solution of 15 in DMSO.

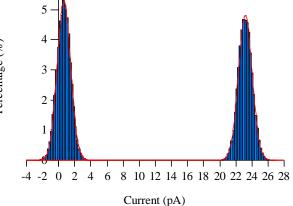


**<u>Figure A2 - 173</u>**: Bilayer activity of **15** with  $K^+$  ions upon the addition of 25µl (recordings 3, 4 and 5) stock solution of **15** in DMSO.

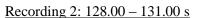












Recording 3: 177.50 – 193.50 s

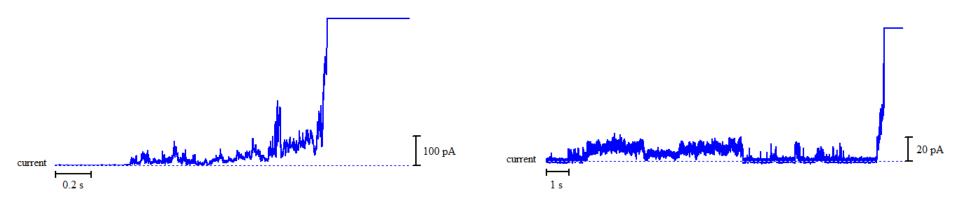
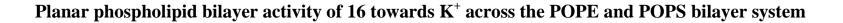
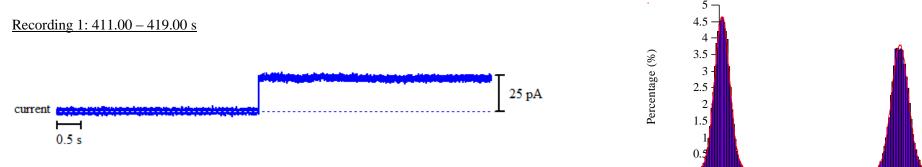
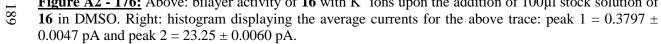
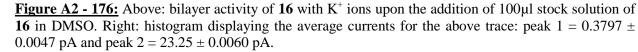


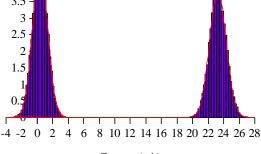
Figure A2 - 175: Bilayer activity of 16 with Na<sup>+</sup> ions upon the addition of 25µl (recording 2) and 50µl (recording 3) stock solution of 16 in DMSO.



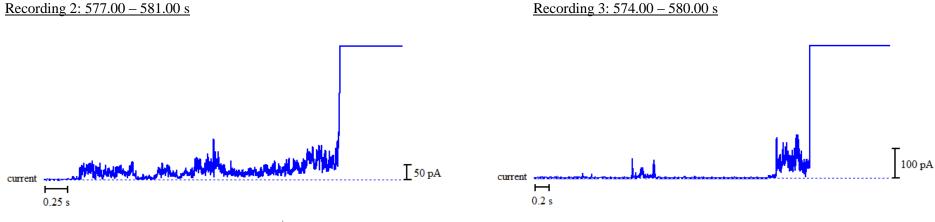




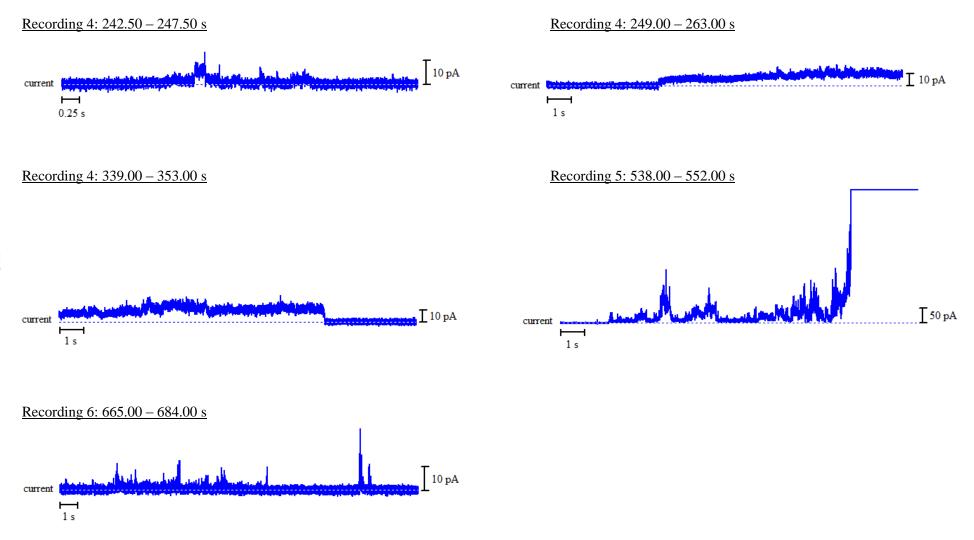






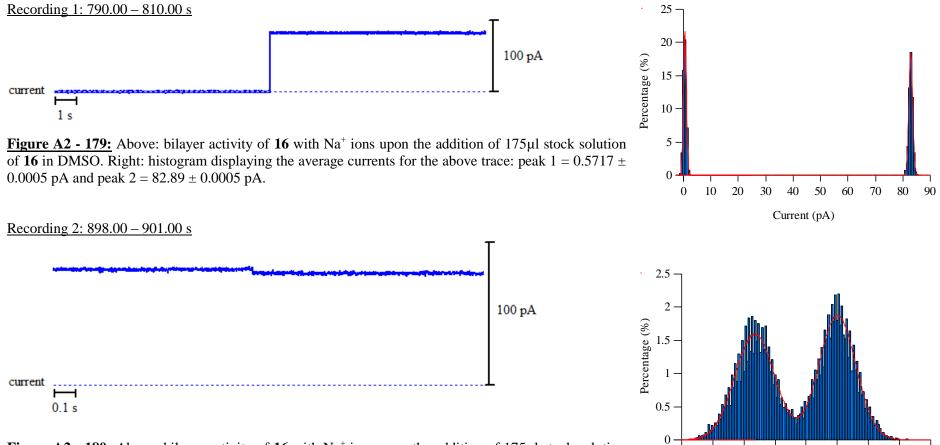


**Figure A2 - 177:** Bilayer activity of **16** with  $K^+$  ions upon the addition of 125µl (recordings 2 and 3) stock solution of **16** in DMSO.



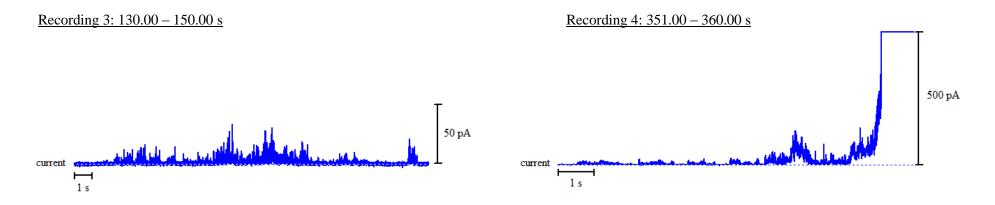
**Figure A2 - 178:** Bilayer activity of **16** with  $K^+$  ions upon the addition of 50µl (recordings 4- top left and right), 75µl (recording 4- centre left), 125µl (recording 5) and 150µl (recording 6) stock solution of **16** in DMSO.

### Planar phospholipid bilayer activity of 16 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system



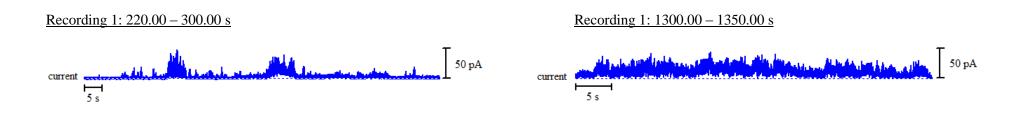
**Figure A2 - 180:** Above: bilayer activity of **16** with Na<sup>+</sup> ions upon the addition of 175µl stock solution of **16** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 78.34 \pm 0.0352$  pA and peak  $2 = 81.0 \pm 0.0303$  pA.

Current (pA)

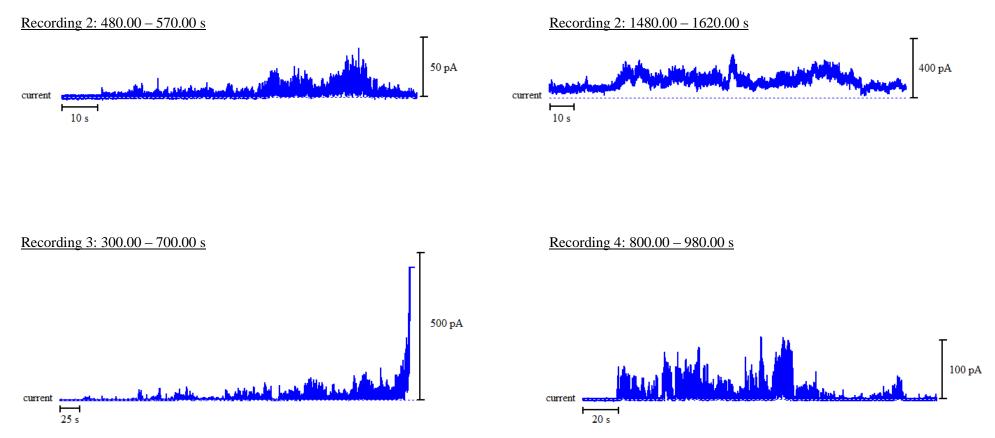


**Figure A2 - 181:** Bilayer activity of **16** with Na<sup>+</sup> ions upon the addition of  $25\mu$ l (recording 3) and  $75\mu$ l (recording 4) stock solution of **16** in DMSO.

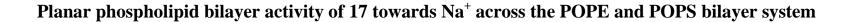
# Planar phospholipid bilayer activity of 16 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

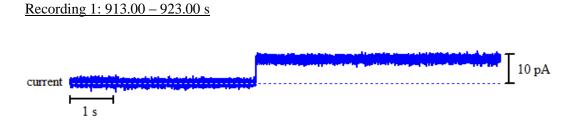


**Figure A2 - 182:** Bilayer activity of **16** with  $K^+$  ions upon the addition of 50µl (recording 1- left) and 175µl (recording 1- right) stock solution of **16** in DMSO.

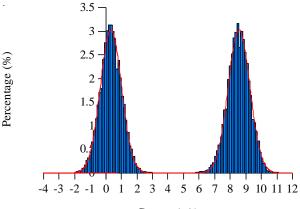


**Figure A2 - 183:** Bilayer activity of **16** with  $K^+$  ions upon the addition of 100µl (Recording 2- left), 200µl (Recording 2- right), 125µl (Recording 3) and 175µl (Recording 4) stock solution of **16** in DMSO.





**Figure A2 - 184:** Above: bilayer activity of **17** with Na<sup>+</sup> ions upon the addition of 250µl stock solution of **17** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.3271 \pm 0.0072$  pA and peak  $2 = 8.563 \pm 0.0073$  pA.





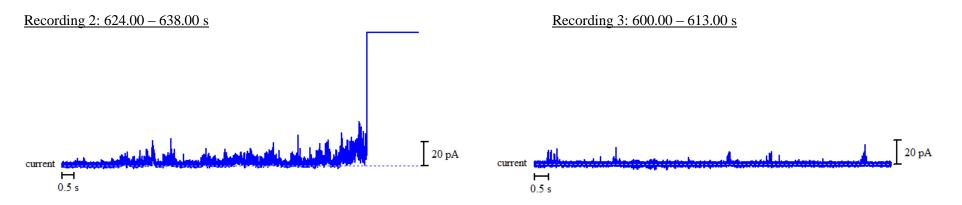
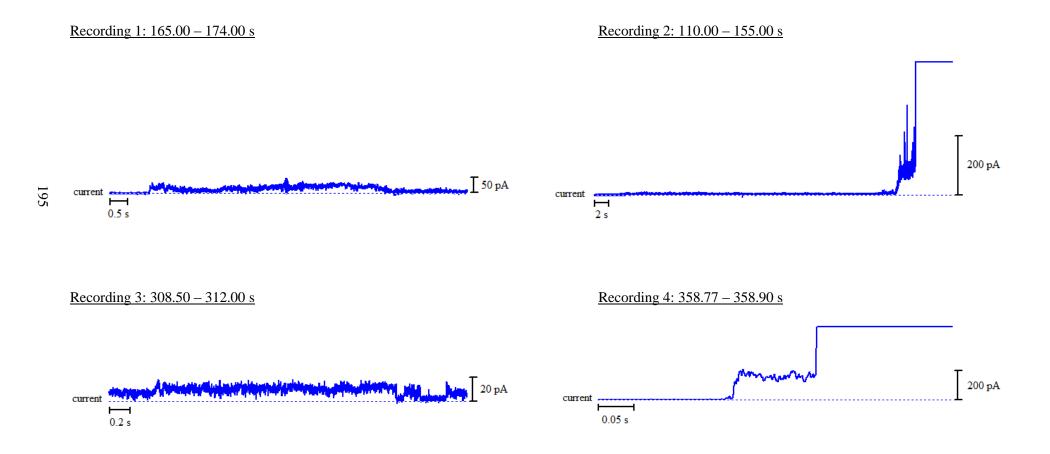
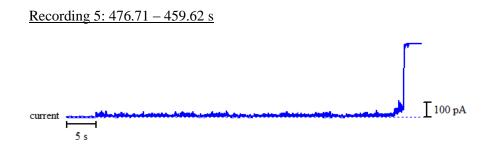


Figure A2 - 185: Bilayer activity of 17 with Na<sup>+</sup> ions upon the addition of 150µl (recordings 2 and 3) stock solution of 17 in DMSO.

# Planar phospholipid bilayer activity of 17 towards K<sup>+</sup> across the POPE and POPS bilayer system



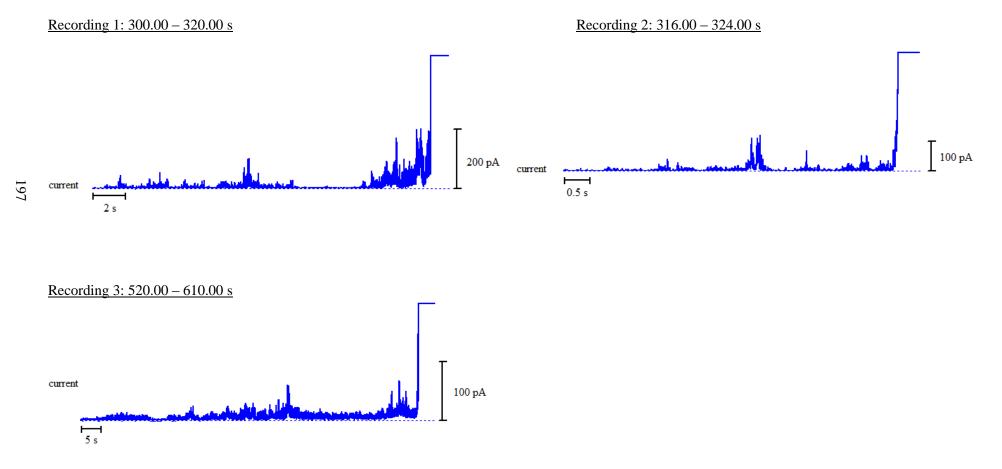
**Figure A2 - 186:** Bilayer activity of **17** with  $K^+$  ions upon the addition of 25µl (recordings 1 and 2) and 75µl (recordings 3 and 4) stock solution of **17** in DMSO.



**Figure A2 - 187:** Bilayer activity of **17** with  $K^+$  ions upon the addition of 100µl (recording 5), stock solution of **17** in DMSO.

# Planar phospholipid bilayer activity of 17 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

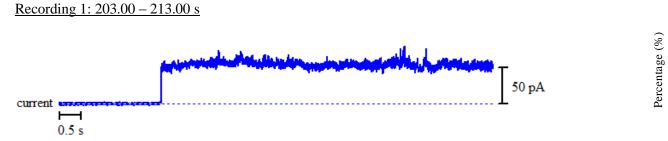
No activity was produced with 17 towards Na<sup>+</sup> across this bilayer system



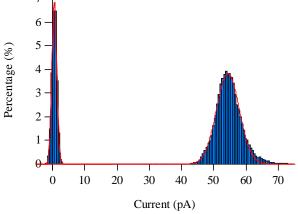
# Planar phospholipid bilayer activity of 17 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

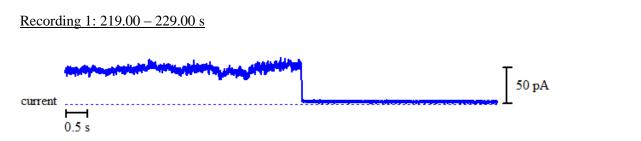
**Figure A2 - 188:** Bilayer activity of **17** with  $K^+$  ions upon the addition of 50µl (recording 1), 75µl (recording 2) and 100µl (recording 3), stock solution of **17** in DMSO.

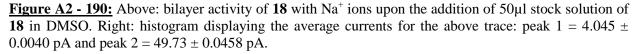
## Planar phospholipid bilayer activity of 18 towards Na<sup>+</sup> across the POPE and POPS bilayer system

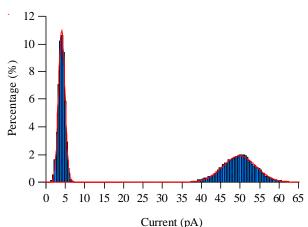


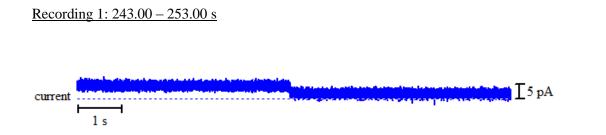
**Figure A2 - 189:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.5316 \pm 0.0099$  pA and peak  $2 = 54.36 \pm 0.0317$  pA.

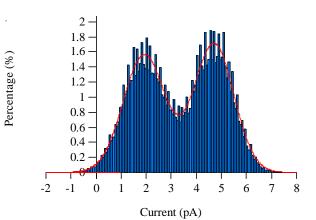








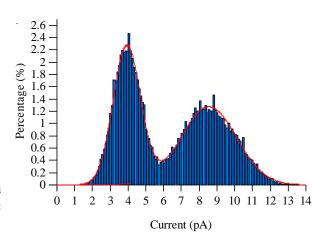




**Figure A2 - 191:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 50µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 1.915 \pm 0.0226$  pA and peak  $2 = 4.695 \pm 0.0202$  pA.



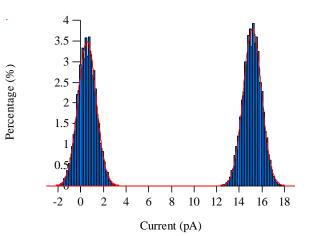




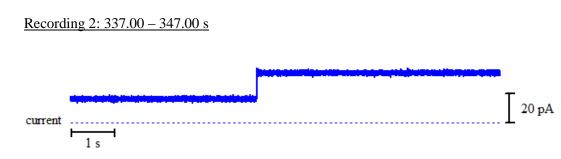
**Figure A2 - 192:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 3.927 \pm 0.0079$  pA and peak  $2 = 8.524 \pm 0.0196$  pA.



Figure A2 - 193: Above: bilayer activity of 18 with Na<sup>+</sup> ions upon the addition of 50µl stock solution of 18 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.5521 \pm$ 0.0097 pA and peak  $2 = 15.16 \pm 0.0086$  pA.







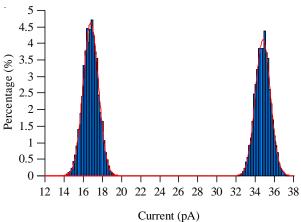
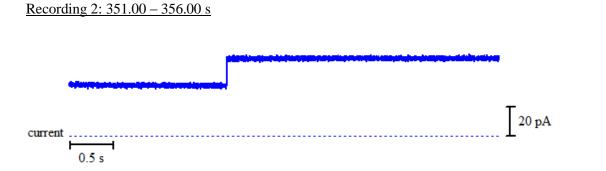
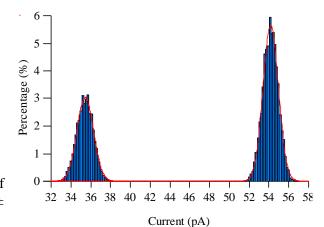


Figure A2 - 194: Above: bilayer activity of 18 with Na<sup>+</sup> ions upon the addition of 75µl stock solution of 18 in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 16.78 \pm$ 0.0080 pA and peak  $2 = 34.79 \pm 0.0089 \text{ pA}$ .



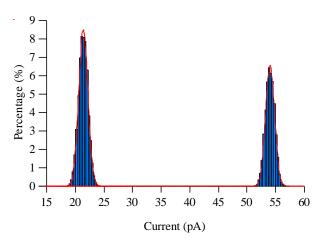
**Figure A2 - 195:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 75µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 35.43 \pm 0.0134$  pA and peak  $2 = 54.17 \pm 0.0071$  pA.



Recording 2: 402.00 – 412.00 s

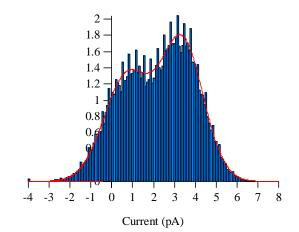


**Figure A2 - 196:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 100µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 21.39 \pm 0.0050$  pA and peak  $2 = 54.01 \pm 0.0064$  pA.



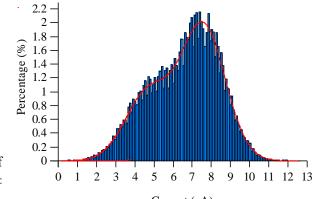


**Figure A2 - 197:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.7199 \pm 0.0624$  pA and peak  $2 = 3.36 \pm 0.0437$  pA.

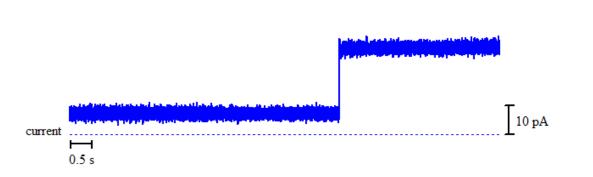


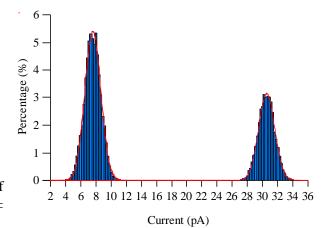
Percentage (%)



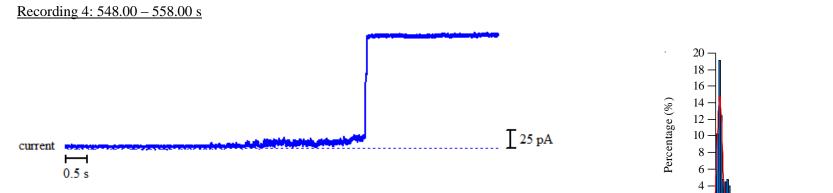


**Figure A2 - 198:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.70 \pm 0.1066$  pA and peak  $2 = 7.558 \pm 0.0527$  pA.

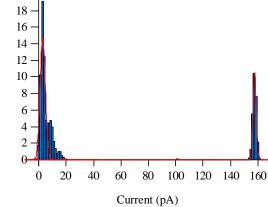




**Figure A2 - 199:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 25µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 7.59 \pm 0.0073$  pA and peak  $2 = 30.53 \pm 0.0125$  pA.

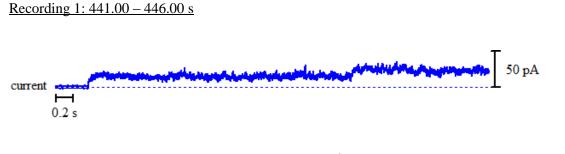


**Figure A2 - 200:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 2.911 \pm 0.115$  pA and peak  $2 = 157.6 \pm 0.1542$  pA.

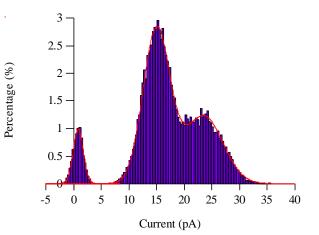


Recording 3: 145.50 - 155.50 s

## Planar phospholipid bilayer activity of 18 towards K<sup>+</sup> across the POPE and POPS bilayer system



**Figure A2 - 201:** Above: bilayer activity of **18** with K<sup>+</sup> ions upon the addition of 100µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 0.8334 \pm 0.0347$  pA, peak  $2 = 14.97 \pm 0.0332$  pA and peak  $3 = 23.45 \pm 0.0962$  pA.



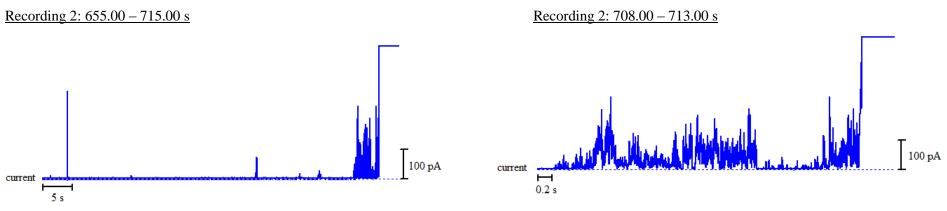
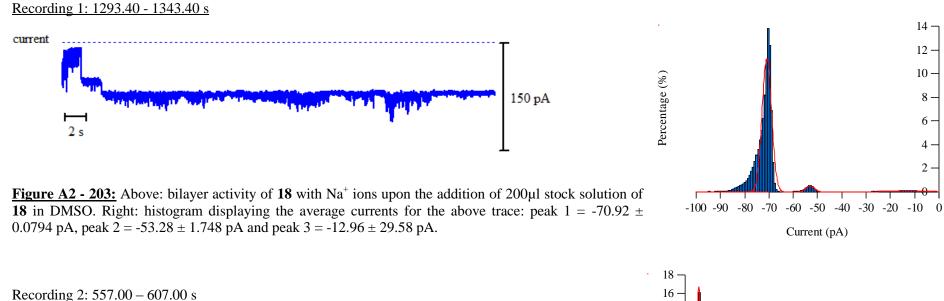


Figure A2 - 202: Bilayer activity of 18 with K<sup>+</sup> ions upon the addition of 150µl (recordings 2) stock solution of 18 in DMSO.

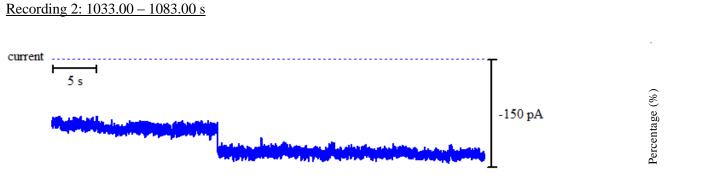
# Planar phospholipid bilayer activity of 18 towards Na<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

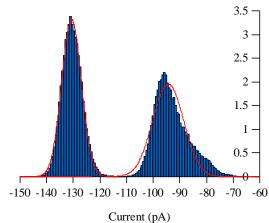




Percentage (%) Current (pA)

**Figure A2 - 204:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 125µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = 4.324 \pm 0.0039$  pA and peak  $2 = 58.01 \pm 0.0625$  pA.





**Figure A2 - 205:** Above: bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 150µl stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak  $1 = -130.7 \pm 0.0701$  pA and peak  $2 = -94.43 \pm 0.1459$  pA.

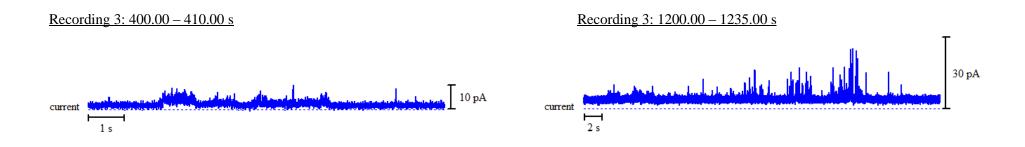
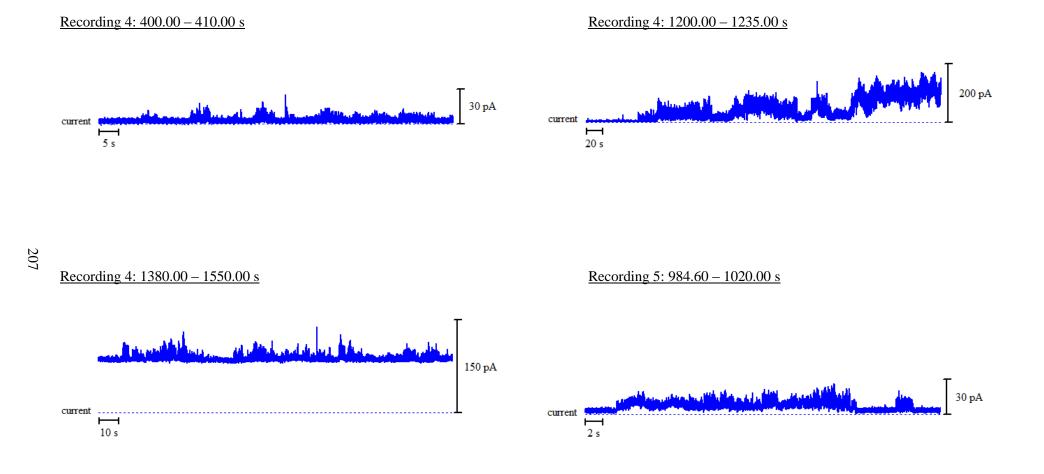
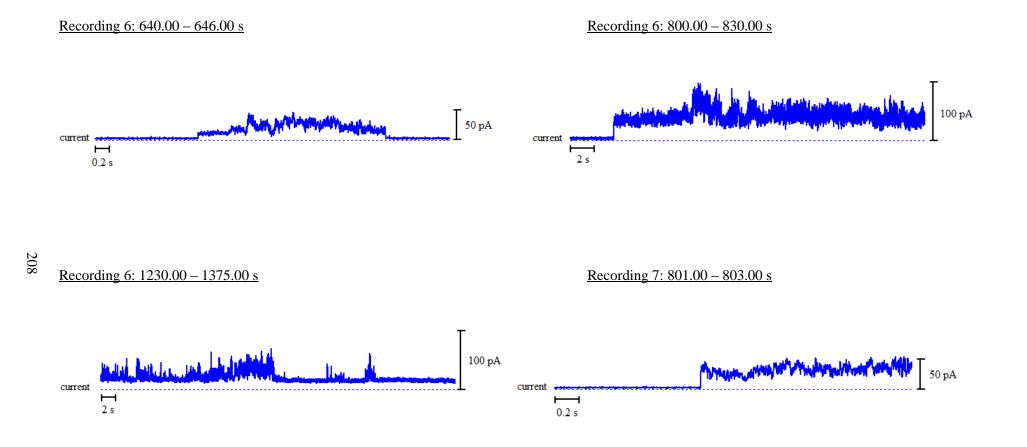


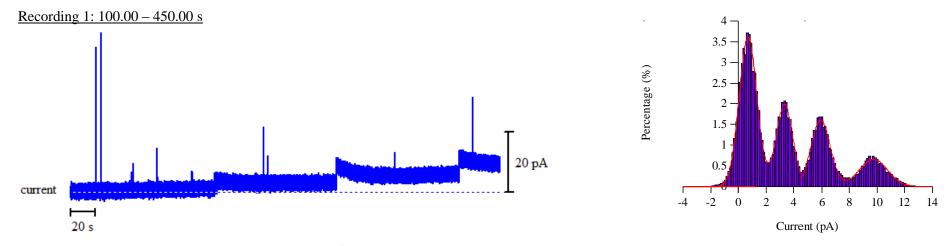
Figure A2 - 206: Bilayer activity of 18 with Na<sup>+</sup> ions upon the addition of 75µl and 200µl (recordings 3) stock solution of 18 in DMSO.



**Figure A2 - 207:** Bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of  $125\mu$ l (recording 4- top left),  $175\mu$ l (recording 4- top right),  $200\mu$ l (recording 4- bottom left) and  $200\mu$ l (recording 5) stock solution of **18** in DMSO.



**Figure A2 - 208:** Bilayer activity of **18** with Na<sup>+</sup> ions upon the addition of 150 $\mu$ l (recording 6- top left), 175 $\mu$ l (recording 6- top right), 200 $\mu$ l (recording 6- top left), 175 $\mu$ l (recording 7) stock solution of **18** in DMSO.



209

Planar phospholipid bilayer activity of 18 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system

**Figure A2 - 209:** Above: bilayer activity of **18** with K<sup>+</sup> ions upon the addition of  $25\mu$ l,  $50\mu$ l,  $75\mu$ l and  $100\mu$ l stock solution of **18** in DMSO. Right: histogram displaying the average currents for the above trace: peak 1 = 0.7004 ± 0.0089 pA, peak 2 =  $3.335 \pm 0.0156$  pA, peak 3 =  $5.908 \pm 0.0110$  pA and peak 4 =  $9.708 \pm 0.0316$  pA.

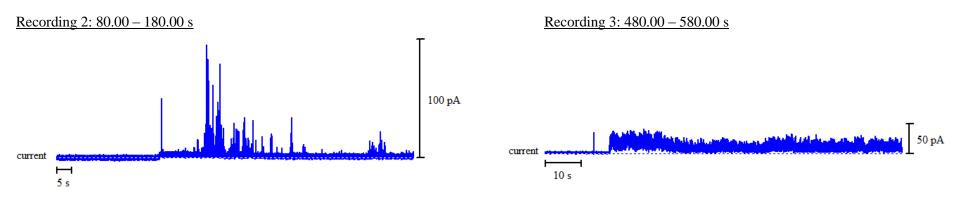
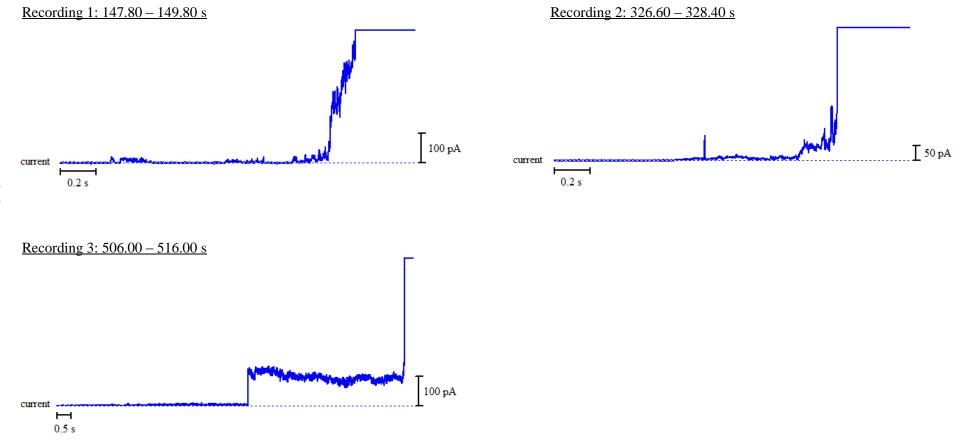
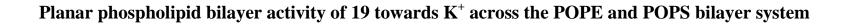


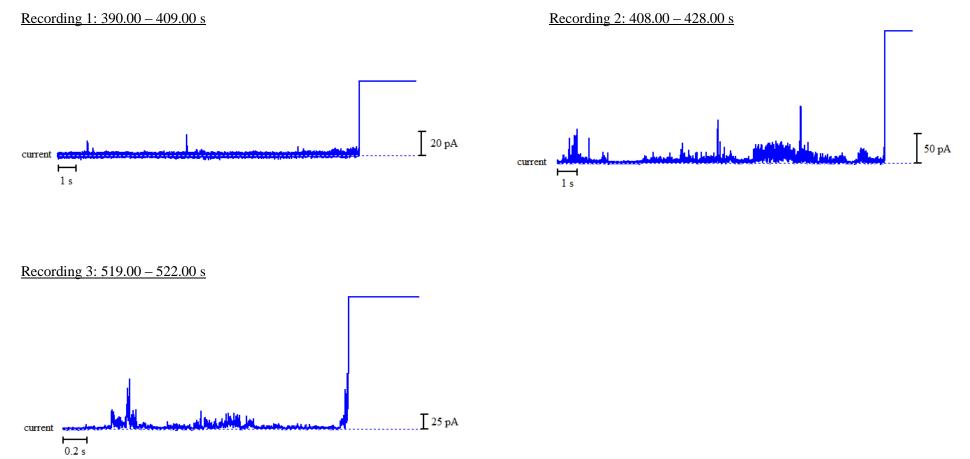
Figure A2 - 210: Bilayer activity of 18 with K<sup>+</sup> ions upon the addition of 25µl (recording 2) and 125µl (recording 3) stock solution of 18 in DMSO.



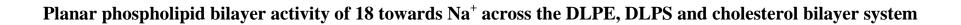
Planar phospholipid bilayer activity of 19 towards Na<sup>+</sup> across the POPE and POPS bilayer system

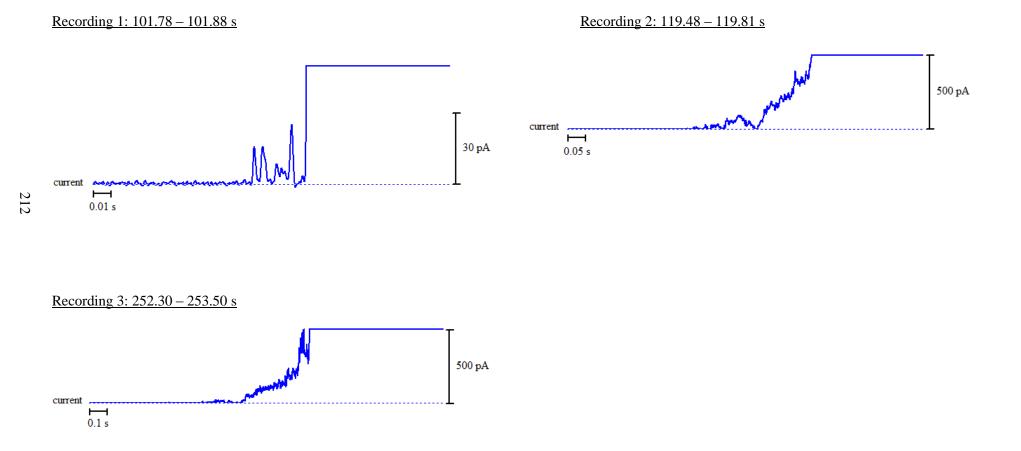
**Figure A2 - 211:** Bilayer activity of **19** with Na<sup>+</sup> ions upon the addition of  $25\mu$ l (recording 1), 75ul (recording 2) and 125ul (recording 3) stock solution of **19** in DMSO.

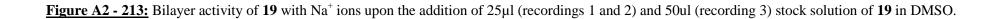




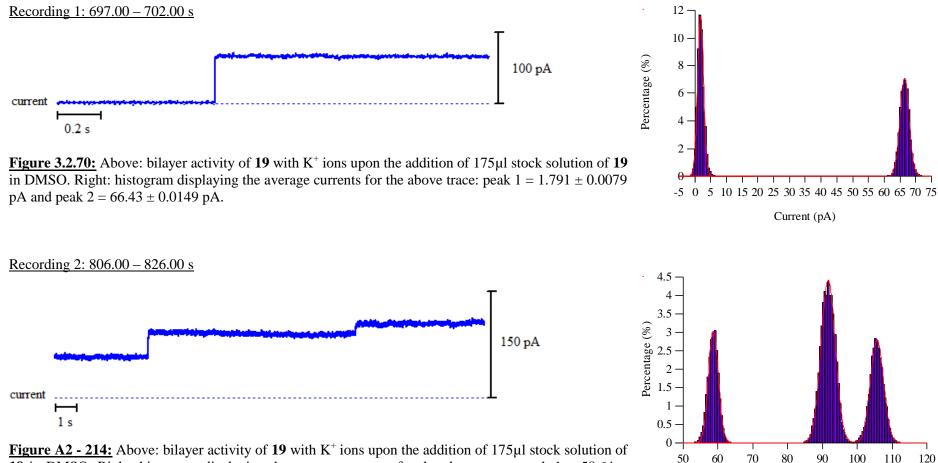
**Figure A2 - 212:** Bilayer activity of **19** with  $K^+$  ions upon the addition of 75µl (recording 1), 100µl (recording 2) and 125µl (recording 3) stock solution of **19** in DMSO.

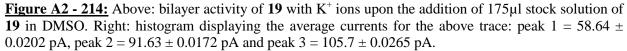




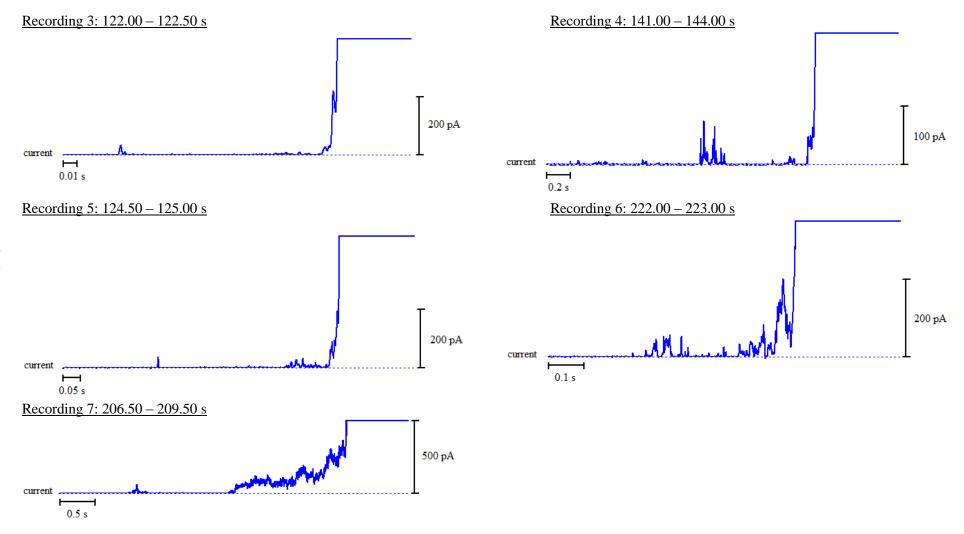


### Planar phospholipid bilayer activity of 18 towards K<sup>+</sup> across the DLPE, DLPS and cholesterol bilayer system









**Figure A2 - 215:** Bilayer activity of **19** with  $K^+$  ions upon the addition of 25µl (recordings 3, 4 and 5) and 50ul (recordings 6 and 7) stock solution of **19** in DMSO.

## **Rationale**

1 amp = 1 coulomb charge/sec 1 mole of Na<sup>+</sup> ions = 96,458 coulombs (Faraday's constant) 1 mole of Na<sup>+</sup> ions =  $6.022 \times 10^{23}$  ions (Avogadro's constant)

Channel current = 96,458 amps = 96,458 coulombs = 1 mole =  $6.022 \times 10^{23}$  Na<sup>+</sup> ions flowing through the channel per second.

<u>**Calculation**</u> – worked example (Figure A2 - 1)

Reading from experiment: 89.910 pA - 5.285 pA = 84.625 pA = **8.4625 x 10^{-11} amps 8.4625 x 10^{-11} amps = 8.4625 x 10^{-11} coulombs/second** 

Ratio for the number of moles of Na<sup>+</sup> ions:

Since 1 mole of Na<sup>+</sup> ions = 96,458 coulombs  $x = 8.4625 \times 10^{-11}$  coulombs/second

Where x is equal to the number of moles of  $Na^+$  ions to be found

 $\frac{1}{96,458} = 1.03672 \times 10^{-5}$ 

Number of moles of Na<sup>+</sup> ions =  $1.03672 \times 10^{-5} \times 8.4625 \times 10^{-11}$  coulombs/second Number of moles of Na<sup>+</sup> ions =  $8.7732 \times 10^{-16}$ 

Ratio for the number of Na<sup>+</sup> ions flowing per second:

Since 1 mole of Na<sup>+</sup> ions =  $6.022 \times 10^{23}$  ions (Avogadro's constant) x moles of Na<sup>+</sup> ions =  $8.7732 \times 10^{-16}$  coulombs/sec where x is equal to the number of ions flowing per second

 $\frac{6.022 \text{ x } 10^{23} \text{ ions}}{1 \text{ mole}} = 6.022 \text{ x } 10^{23}$ 

x Na<sup>+</sup> ions/sec =  $6.022 \times 10^{23} \times 8.7732 \times 10^{-16}$  coulombs/sec

 $Na^+$  ions/sec = 528325022.3 ions flowing per second

 $Na^+$  ions/sec = 5.28 x 10<sup>8</sup> ions flowing per second

<u>Calculation A2 - 1:</u> Calculation to determine the number of ions flowing across the bilayer membrane per second based from the current produced.

	Compound 8					Compound 8			
	С	onductance (pS	5)		Current (pA)				
Li <sup>+</sup>	$Na^+$	$\mathbf{K}^+$	$Rb^+$	$Cs^+$	Li <sup>+</sup>	Na <sup>+</sup>	$\mathbf{K}^+$	$Rb^+$	$Cs^+$
45.68	1692.50	374.28		0.00	2.28	84.63	18.71		0.00
0.00	1384.00	124.00		0.00	0.00	69.20	6.20		0.00
0.00	300.84	0.00		0.00	0.00	15.04	0.00		0.00
0.00	1941.00	0.00		0.00	0.00	97.05	0.00		0.00
	336.00	0.00		0.00		16.80	0.00		0.00
	2512.26	0.00		0.00		125.61	0.00		0.00
	209.96	0.00				10.50	0.00		0.00
	567.47	0.00				28.37	0.00		
	0.00	0.00				0.00	0.00		
	0.00	0.00				0.00	0.00		
	0.00	0.00				0.00	0.00		
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			
	0.00					0.00			

<u>**Table A2 - 1:**</u> A table showing the raw data for Conductance (pS) and Current (pA) obtained for **8** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

0.00		0.0	0	
0.00		0.0	0	
0.00		0.0	0	
0.00		0.0	0	
0.00		0.0	0	
0.00		0.0	0	

	$Li^+$	Na <sup>+</sup>	$\mathbf{K}^{+}$	<b>Rb</b> <sup>+</sup>	$\mathbf{Cs}^+$
Average Conductance (pS)	45.68	1118.00	249.14	N/a	0.00
Average Current (pA)	2.28	55.90	12.46	N/a	0.00
Average ion flow rate (ions/second)	14259303.99	348991938.80	77770643.50	N/a	0.000

	Compound 8					Compound 8			
	С	onductance (p	<b>S</b> )		Current (pA)				
Li <sup>+</sup>	Na <sup>+</sup>	$\mathbf{K}^+$	$Rb^+$	$Cs^+$	Li <sup>+</sup>	Na <sup>+</sup>	$\mathbf{K}^+$	$Rb^+$	Cs <sup>+</sup>
57.33	197.37	41.35	547.30	139.02	2.87	9.87	2.07	27.37	6.95
0.00	62.63	43.57	139.20	85.48	0.00	11.90	2.18	6.96	4.27
0.00	263.40	2893.14	356.80	44.78	0.00	13.17	144.66	17.84	2.24
0.00	71.05	0.00	137.80	1598.62	0.00	5.68	0.00	6.89	79.93
0.00	80.32	0.00	142.80	0.00	0.00	4.02	0.00	7.14	0.00
0.00	80.32	0.00	258.00	0.00	0.00	4.02	0.00	12.90	0.00
0.00	69.80	0.00	532.58	0.00	0.00	3.49	0.00	26.63	0.00
	50.14	0.00	0.00			2.51	0.00	0.00	
	409.14	0.00	0.00			20.46	0.00	0.00	
	188.57	0.00	0.00			9.43	0.00	0.00	
	643.60	0.00	0.00			32.18	0.00	0.00	
	689.20	0.00	0.00			34.46	0.00	0.00	
	625.17	0.00	0.00			31.26	0.00	0.00	
	3216.50		0.00			257.32		0.00	
	3585.00					143.40			
	4078.00					203.90			
	2414.00					120.70			
	4230.00					211.50			
	2708.60					135.43			
	1349.80					67.49			

Table A2 - 2: A table showing the raw data for Conductance (pS) and Current (pA) obtained for 8 across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

1396.00	69.80	
838.00	41.90	
382.00	19.10	
1496.00	74.80	
926.00	46.30	
1086.07	54.30	
914.14	45.71	
1208.98	60.45	
280.20	14.01	
379.00	18.95	
538.00	26.90	
151.20	7.56	
278.80	13.94	
253.40	12.67	
268.00	13.40	
1053.06	52.65	
1066.52	53.33	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	
0.00	0.00	

0.00		0.00		
0.00		0.00		
0.00		0.00		
0.00		0.00		
0.00		0.00		
0.00		0.00		

	$Li^+$	$\mathbf{Na}^+$	$\mathbf{K}^+$	<b>Rb</b> <sup>+</sup>	$\mathbf{Cs}^+$
Average Conductance (pS)	57.33	1014.27	992.69	302.07	466.98
Average Current (pA)	2.87	52.65	49.63	15.10	23.35
Average ion flow rate (ions/second)	17897174.58	328682828.80	309873904.50	94292635.39	145769231.20

**Table A2 - 3:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **9** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 9				
Conductance (pS)				
Na <sup>+</sup>	$\mathbf{K}^+$			
2425.90				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				

Compound 9				
Current (pA)				
Na <sup>+</sup>	$\mathbf{K}^+$			
121.30				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				
0.00				

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	2425.90	N/a
Average Current (pA)	121.30	N/a
Average ion flow rate (ions/second)	757260191.35	0.00

**Table A2 - 4:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **9** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 9			
Conductan	ice (pS)		
Na <sup>+</sup>	$\mathbf{K}^+$		
2196.40	0.00		
361.00	0.00		
202.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		

Compound 9				
Current (pA)				
$Na^+$	$\mathbf{K}^+$			
109.82	0.00			
18.05	0.00			
10.10	0.00			
0.00	0.00			
0.00	0.00			
0.00	0.00			
0.00	0.00			

	$Na^+$	$\mathbf{K}^+$
Average Conductance (pS)	919.80	0.00
Average Current (pA)	45.99	0.00
Average ion flow rate (ions/second)	287121449.40	0.00

**Table A2 - 5:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **10** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 10		
tance (pS)		
K <sup>+</sup>		
3619.36		
3374.00		
546.00		
546.00		
7194.70		
0.00		
0.00		
0.00		
0.00		
0.00		
0.00		

Compound 10			
Current (pA)			
$Na^+$	$\mathbf{K}^+$		
57.79	180.97		
18.30	168.70		
14.97	27.30		
41.40	27.30		
0.00	359.74		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00			
0.00			

	$\mathbf{Na}^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	662.31	3056.01
Average Current (pA)	33.12	152.80
Average ion flow rate (ions/second)	206744300.00	953953679.80

**Table A2 - 5:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **10** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compour	nd <b>10</b>	Compo	und 10
Conductan	ce (pS)	Currer	nt (pA)
Na <sup>+</sup>	K+	Na <sup>+</sup>	K <sup>+</sup>
1068.34	0.00	53.42	0.00
226.60	0.00	11.33	0.00
192.40	0.00	9.62	0.00
68.17	0.00	3.41	0.00
969.44	0.00	48.47	0.00
871.60	0.00	43.58	0.00
1112.40	0.00	55.62	0.00
3604.00	0.00	180.20	0.00
2098.00		104.90	
588.58		29.43	
2254.00		112.70	
1950.00		97.50	
1128.00		56.40	
1512.00		75.60	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.000		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	

	$Na^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	1260.25	0.00
Average Current (pA)	63.01	0.00
Average ion flow rate (ions/second)	393395806.50	0.00

**Table A2 - 6:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **11** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	und <b>11</b>	Compound 1	
Conducta	unce (pS)	Current (pA)	
Na <sup>+</sup>	$K^+$	Na <sup>+</sup> K <sup>+</sup>	

	Na <sup>+</sup>	<b>K</b> <sup>+</sup>
Average Conductance (pS)	N/a	N/a
Average Current (pA)	N/a	N/a
Average ion flow rate (ions/second)	N/a	N/a

**<u>Table A2 - 7</u>**: A table showing the raw data for Conductance (pS) and Current (pA) obtained for **11** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 11		Compo	und <b>11</b>
Conducta	nce (pS)	Current (pA)	
Na <sup>+</sup>	K <sup>+</sup>	Na <sup>+</sup>	$\mathbf{K}^+$
88.56	301.80	4.43	15.09
135.88	0.00	6.79	0.00
62.88	0.00	3.14	0.00
178.80	0.00	8.94	0.00
104.20	0.00	5.21	0.00
93.26	0.00	4.66	0.00
77.26	0.00	3.86	0.00
90.12	0.00	4.51	0.00
28.86	0.00	1.44	0.00
27.70		1.39	
58.98		2.95	
34.82		1.74	
151.70		7.59	
31.00		1.55	
36.80		1.84	
105.10		5.26	
74.36		3.72	
103.20		5.16	
32.75		2.62	
281.33		25.32	
302.20		15.11	
49.20		2.46	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	

	$\mathbf{Na}^{+}$	$\mathbf{K}^{+}$
Average Conductance (pS)	108.80	301.80
Average Current (pA)	5.44	15.09
Average ion flow rate (ions/second)	33963750.56	94208799.11

**Table A2 - 8:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **12** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 12		] [	Compound 12	
Conductance (pS)			Current (pA)	
$Na^+$	$\mathbf{K}^+$	] [	$Na^+$	$\mathbf{K}^+$
190.12		] [	9.51	
396.20			19.81	
608.60			30.43	
1280.80			64.04	
1652.00			82.60	
219.26			10.96	
7670.60		] [	383.53	
0.00			0.00	
0.00		] [	0.00	
0.00		] [	0.00	
0.00			0.00	
0.00		] [	0.00	
0.00		] [	0.00	
0.00			0.00	

	$\mathbf{Na}^{+}$	$\mathbf{K}^{+}$
Average Conductance (pS)	1716.80	N/a
Average Current (pA)	85.84	N/a
Average ion flow rate (ions/second)	535909201.90	N/a

**Table A2 - 9:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **12** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Com	pound 12	] [	Compound 12		
Condu	ctance (pS)		Curre	ent (pA)	
Na <sup>+</sup>	K <sup>+</sup>	1	Na <sup>+</sup> K <sup>+</sup>		
0.00	3716.18		0.00	185.81	
0.00	938.00		0.00	46.90	
0.00	330.00		0.00	16.50	
0.00	198.00		0.00	9.90	
0.00	742.40		0.00	37.12	
0.00	1461.04		0.00	73.05	
0.00	221.78		0.00	11.09	
0.00	69.96		0.00	3.50	
	69.96			3.50	
	408.88			20.44	
	161.52			8.08	
	161.52			8.08	
	343.48			17.17	
	181.96			9.10	
	247.36			12.37	
	247.36			12.37	
	161.52			8.08	
	408.88			20.44	
	166.60			8.33	
	90.00			4.50	
	90.00			4.50	
	377.20			18.86	
	312.20	] [		15.61	
	158.60	] [		7.93	
	0.00			0.00	
	0.00	] [		0.00	
	0.00	] [		0.00	
	0.00	] [		0.00	
	0.00	] [		0.00	
	0.00			0.00	

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	0.00	469.35
Average Current (pA)	0.00	23.47
Average ion flow rate (ions/second)	0.00	146510602.60

**Table A2 - 10:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **13** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	und 13		Compound 13	
Conducta	ince (pS)		Current (pA)	
Na <sup>+</sup>	K <sup>+</sup>	I	Na <sup>+</sup>	K <sup>+</sup>
		-		

	Na <sup>+</sup>	$\mathbf{K}^+$
Average Conductance (pS)	N/a	N/a
Average Current (pA)	N/a	N/a
Average ion flow rate (ions/second)	N/a	N/a

**Table A2 - 11:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **13** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Comp	bound 13		
Conduc	Conductance (pS)		
Na <sup>+</sup>	K <sup>+</sup>		
252.73	695.82		
88.40	1522.00		
89.46	432.60		
118.82	230.00		
191.42	152.00		
786.46	83.00		
364.80	53.80		
0.00	106.40		
0.00	171.20		
0.00	0.00		
0.00	0.00		
0.00	0.00		
	0.00		
	0.00		
	0.00		
	0.00	1	

Compound 13		
Current (pA)		
Na <sup>+</sup>	$K^+$	
12.64	34.79	
4.42	76.10	
4.47	21.63	
5.94	11.50	
9.57	7.60	
39.32	4.15	
18.24	2.69	
0.00	5.32	
0.00	8.56	
0.00	0.00	
0.00	0.00	
0.00	0.00	
	0.00	
	0.00	
	0.00	
	0.00	

	$Na^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	270.30	382.98
Average Current (pA)	13.51	19.15
Average ion flow rate (ions/second)	84375516.01	119549655.01

**Table A2 - 12:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **14** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	ound <b>14</b>	Compound 14	
Conducta	ance (pS)	Current (pA)	
$Na^+$	$\mathbf{K}^+$	Na <sup>+</sup> K <sup>+</sup>	
352.14	3808.48	17.61 190.42	2
350.16	8681.84	17.51 434.09	)
360.08	1050.40	18.00 52.52	
364.34	0.00	18.22 0.00	
6394.20	0.00	319.71 0.00	
324.00		16.20	
292.96		14.65	
290.92		14.55	
7810.40		390.52	
73.84		3.69	
297.34		14.87	
338.95		16.95	
1541.71		77.09	
2766.00		138.30	
0.00		0.00	
0.00		0.00	
0.00		0.00	
0.00		0.00	

	$Na^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	1539.79	4513.57
Average Current (pA)	76.99	225.68
Average ion flow rate (ions/second)	480654800.30	1408941182.97

**Table A2 - 13:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **14** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

		1 1		
	Compound 14		Compound 14	
Conducta			Current (pA)	
$Na^+$	$\mathbf{K}^+$		$Na^+$	$\mathbf{K}^+$
243.58	108.03		12.18	5.40
773.40	51.94		38.67	2.60
1819.00	0.00		90.95	0.00
126.00	0.00		6.30	0.00
254.00	0.00		12.70	0.00
138.00	0.00		6.90	0.00
340.00	0.00		17.00	0.00
207.20	0.00		10.36	0.00
201.25			8.05	
238.80			11.94	
77.80			3.89	
77.80			3.89	
139.60			6.98	
154.54			7.73	
0.00			0.00	
0.00			0.00	
0.00			0.00	
0.00			0.00	
0.00			0.00	

	$\mathbf{Na}^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	342.21	79.99
Average Current (pA)	16.97	4.00
Average ion flow rate (ions/second)	105926258.19	24968141.17

**Table A2 - 14:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **15** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 15		Compound 15	
Conducta	nce (pS)	Curre	ent (pA)
$Na^+$	$\mathbf{K}^+$	Na <sup>+</sup>	$\mathbf{K}^+$
2028.80	213.48	101.44	10.67
445.70	219.63	22.29	10.98
279.94	450.72	14.00	22.54
3532.80	299.20	176.64	14.96
1321.57	0.00	66.08	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
	0.00		0.00

	Na⁺	$\mathbf{K}^{+}$
Average Conductance (pS)	1521.76	295.76
Average Current (pA)	76.09	14.79
Average ion flow rate (ions/second)	475027735.40	92323062.34

**Table A2 - 15:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **15** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Comp	ound 15	1 [	Compo	ound 15
^	tance (pS)		-	nt (pA)
Na <sup>+</sup>	K <sup>+</sup>		$Na^+$	K <sup>+</sup>
202.68	2777.20		10.13	138.86
210.80	2777.20		10.54	138.86
232.20	2777.20		11.61	138.86
0.00	2777.20	1 [	0.00	138.86
0.00	1665.20		0.00	83.26
0.00	1112.00	1 [	0.00	55.60
0.00	0.00		0.00	0.00
0.00	0.00	1 [	0.00	0.00
0.00	0.00	1 [	0.00	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
	0.00	1 [		0.00
	0.00	] [		0.00
	0.00			0.00

	$\mathbf{Na}^{+}$	$\mathbf{K}^{+}$
Average Conductance (pS)	215.23	2314.33
Average Current (pA)	10.76	115.72
Average ion flow rate (ions/second)	67184379.73	722433943.22

**Table A2 - 16:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **16** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	Compound 16		Compound 16	
Conduct	Conductance (pS)		Curren	nt (pA)
Na <sup>+</sup>	$\mathbf{K}^+$		$Na^+$	$\mathbf{K}^+$
450.78	457.41		22.54	22.87
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
0.00			0.00	
0.00			0.00	
0.00			0.00	

	Na⁺	$\mathbf{K}^{+}$
Average Conductance (pS)	450.78	457.41
Average Current (pA)	22.54	22.87
Average ion flow rate (ions/second)	140715107.01	142782206.64

**Table A2 - 17:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **16** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 16		Com	pound 16
Conductan	ce (pS)	Curr	ent (pA)
$Na^+$	$\mathbf{K}^+$	Na <sup>+</sup>	K <sup>+</sup>
1646.37	0.00	82.32	0.00
53.20	0.00	2.66	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00		0.00	
0.00		0.00	

 $\mathbf{K}^+$ 0.00 0.00 0.00 0.00 0.00 0.00 0.00 0.00

	$\mathbf{Na}^{+}$	$\mathbf{K}^+$
Average Conductance (pS)	849.78	0.00
Average Current (pA)	42.49	0.00
Average ion flow rate (ions/second)	265265195.26	0.00

**Table A2 - 18:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **17** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	Compound 17 Compound 17		ound <b>17</b>	
Conducta	Conductance (pS)		Current (pA)	
$Na^+$	$\mathbf{K}^+$		$Na^+$	$\mathbf{K}^+$
164.72	0.00		8.24	0.00
0.00	0.00		0.00	0.00
0.00	0.00		0.00	0.00
	0.00			0.00
	0.00			0.00
	0.00			0.00
	0.00			0.00
	0.00			0.00
	0.00			0.00
	0.00	]		0.00
	0.00			0.00

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	164.72	0.00
Average Current (pA)	8.24	0.00
Average ion flow rate (ions/second)	51417776.58	0

**Table A2 - 19:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **17** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Comp	ound 17	Compound 17	
Conduc	ctance (pS)	Current (pA)	
Na <sup>+</sup>	$\mathbf{K}^+$	$Na^+$	$K^+$
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00	0.00	0.00	0.00
0.00		0.00	
0.00		0.00	

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	0.00	0.00
Average Current (pA)	0.00	0.00
Average ion flow rate (ions/second)	0.00	0.00

**<u>Table A2 - 20</u>**: A table showing the raw data for Conductance (pS) and Current (pA) obtained for **18** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compo	C	
Conduct	Conductance (pS)	
$Na^+$	$K^+$	N
1076.57	282.73	53
913.70	169.60	45
55.60	0.00	2.
91.94	0.00	4.
292.16	0.00	14
360.20	0.00	18
374.80	0.00	18
652.40	0.00	32
52.80	0.00	2.
458.80	0.00	22
3093.78		154
0.00		0.
0.00		0.
0.00		0.
0.00		0.

Compound 18			
Current (pA)			
Na <sup>+</sup> K <sup>+</sup>			
53.83	14.14		
45.69	8.48		
2.78	0.00		
4.60	0.00		
14.61	0.00		
18.01	0.00		
18.74	0.00		
32.62	0.00		
2.64	0.00		
22.94	0.00		
154.69			
0.00			
0.00			
0.00			
0.00			

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	674.80	226.17
Average Current (pA)	33.74	11.31
Average ion flow rate (ions/second)	210641657.67	70599162.55

**Table A2 - 21:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **18** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 18		Compo
Conductance (pS)		Curren
Na <sup>+</sup>	$\mathbf{K}^+$	Na <sup>+</sup>
806.40	52.69	40.32
352.80	51.46	17.64
1073.72	76.00	53.69
725.40	0.00	36.27
0.00	0.00	0.00
0.00	0.00	0.00
0.00	0.00	0.00
0.00	0.00	0.00
0.00	0.00	0.00
0.00	0.00	0.00
0.00	0.00	0.00
	0.00	

Compound 18			
Current (pA)			
Na <sup>+</sup> K <sup>+</sup>			
40.32 2.64			
17.64	2.57		
53.69	3.80		
36.27 0.00			
0.00 0.00			
0.00	0.00		
0.00 0.00			
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00			

	$Na^+$	$\mathbf{K}^{+}$
Average Conductance (pS)	739.58	60.05
Average Current (pA)	36.98	3.00
Average ion flow rate (ions/second)	230864624.40	18745199.44

**Table A2 - 22:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **19** across the POPE and POPS bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

$\begin{tabular}{ c c c c c c c c c c c c c c c c c c c$		
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	Compound 19	
0.00         0.00 <th< td=""><td>pA)</td></th<>	pA)	
0.00         0.00         0.00         0.00         0           0.00         0.00         0.00         0         0         0	$\mathbf{K}^+$	
0.00 0.00 0.00 0	0.00	
	0.00	
	0.00	
0.00 0.00 0.00 0	0.00	
0.00 0.00 0.00 0	0.00	
0.00 0.00 0.00 0	0.00	
0.00 0.00 0.00 0	0.00	
0.00	0.00	

	Na <sup>+</sup>	$\mathbf{K}^{+}$
Average Conductance (pS)	0.00	0.00
Average Current (pA)	0.00	0.00
Average ion flow rate (ions/second)	0.00	0.00

**Table A2 - 23:** A table showing the raw data for Conductance (pS) and Current (pA) obtained for **19** across the DLPE, DLPS and cholesterol bilayer system and the calculated average rate of ions per second (coulombs/second) of each ion respectively.

Compound 19			
Conductance (pS)			
Na <sup>+</sup> K <sup>+</sup>			
0.00	1292.78		
0.00	659.80		
0.00 281.40			
0.00 0.00			
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
	0.00		

Compound 19			
Current (pA)			
Na <sup>+</sup>	$\mathbf{K}^+$		
0.00	64.64		
0.00	32.99		
0.00	14.07		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00	0.00		
0.00			

	Na <sup>+</sup>	K <sup>+</sup>
Average Conductance (pS)	0.00	744.66
Average Current (pA)	0.00	37.23
Average ion flow rate (ions/second)	0.00	232450378.87

## **Appendix III**

Antimicrobial susceptibility testing

	Microorganism			
Compound	E. coli	S. aureus	P. aeruginosa	S. pyogenes
8	-	-	-	-
9	-	-	-	-
10	-	-	-	-
11	9.33	8.50	8.00	22.16
12	9.60	-	-	-
13	-	-	-	8.33
14	16.16	-	15.67	-
15	18.50	17.30	-	-
16	-	-	-	-
17	16.50	12.16	17.00	25.00
18	11.67	22.33	10.00	54.33
19	9.00	7.67	10.83	16.00

<u>**Table A3 – 1:**</u> The average diameter (mm) of microbial inhibition zones produced upon contact with compounds 8-19.