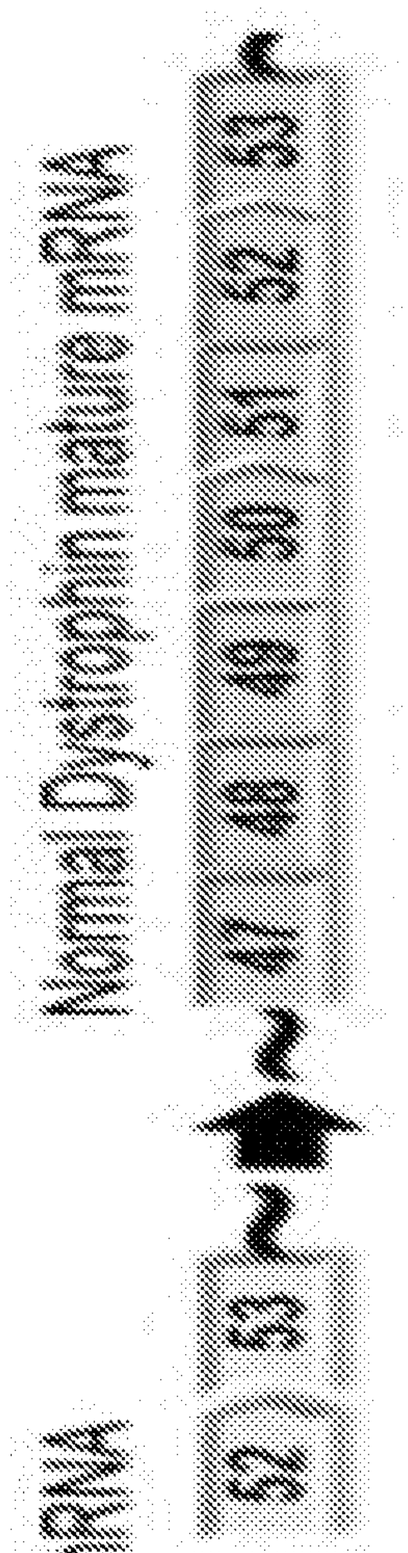

Normal Cystoid Matrix
Cystoid Matrix
Cystoid Pre-Mature
Mature
42
43
44
45
46
47
48
49
50
51
52
53

Figure 1

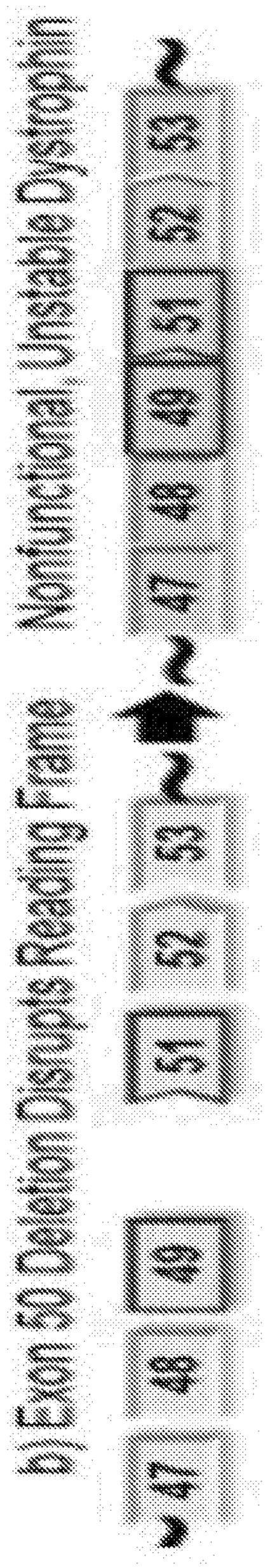

Non-functional Unstable Oligonucleotide
47
48
49
51
52
53
Oligonucleotide Reading Frame
51
53
49
47

Figure 2

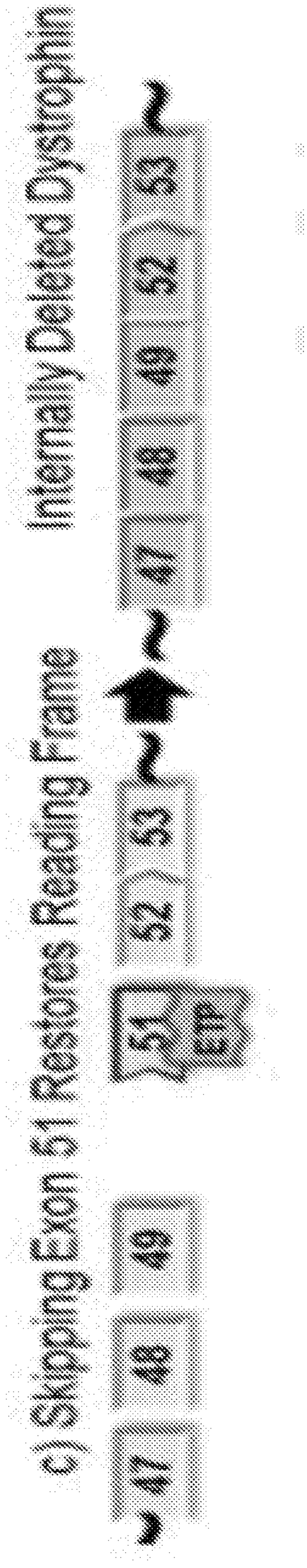


Figure 3

Source: Adapted from Kole 2012.

Figure 4A

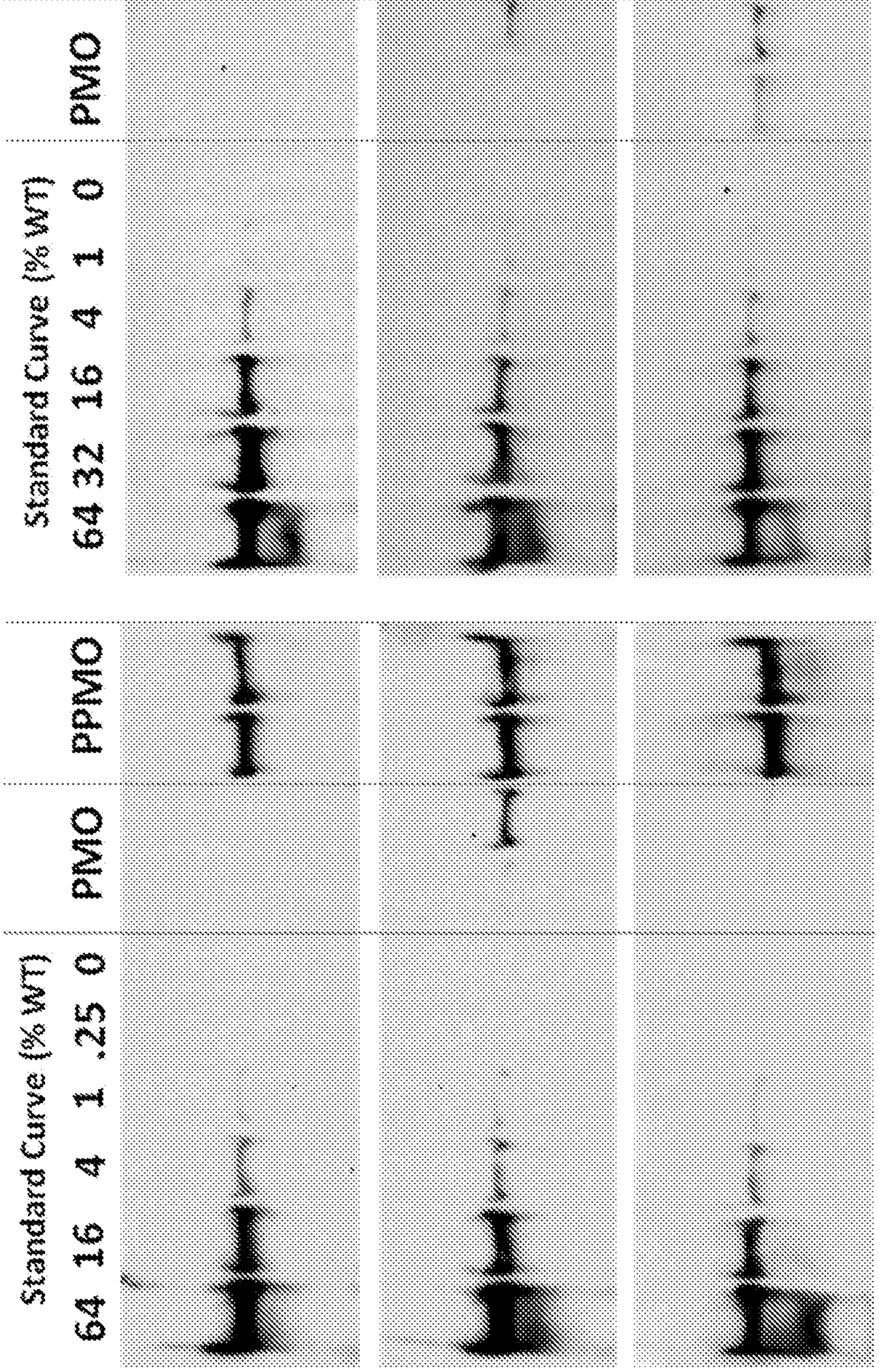


Figure 4B

Figure 4C

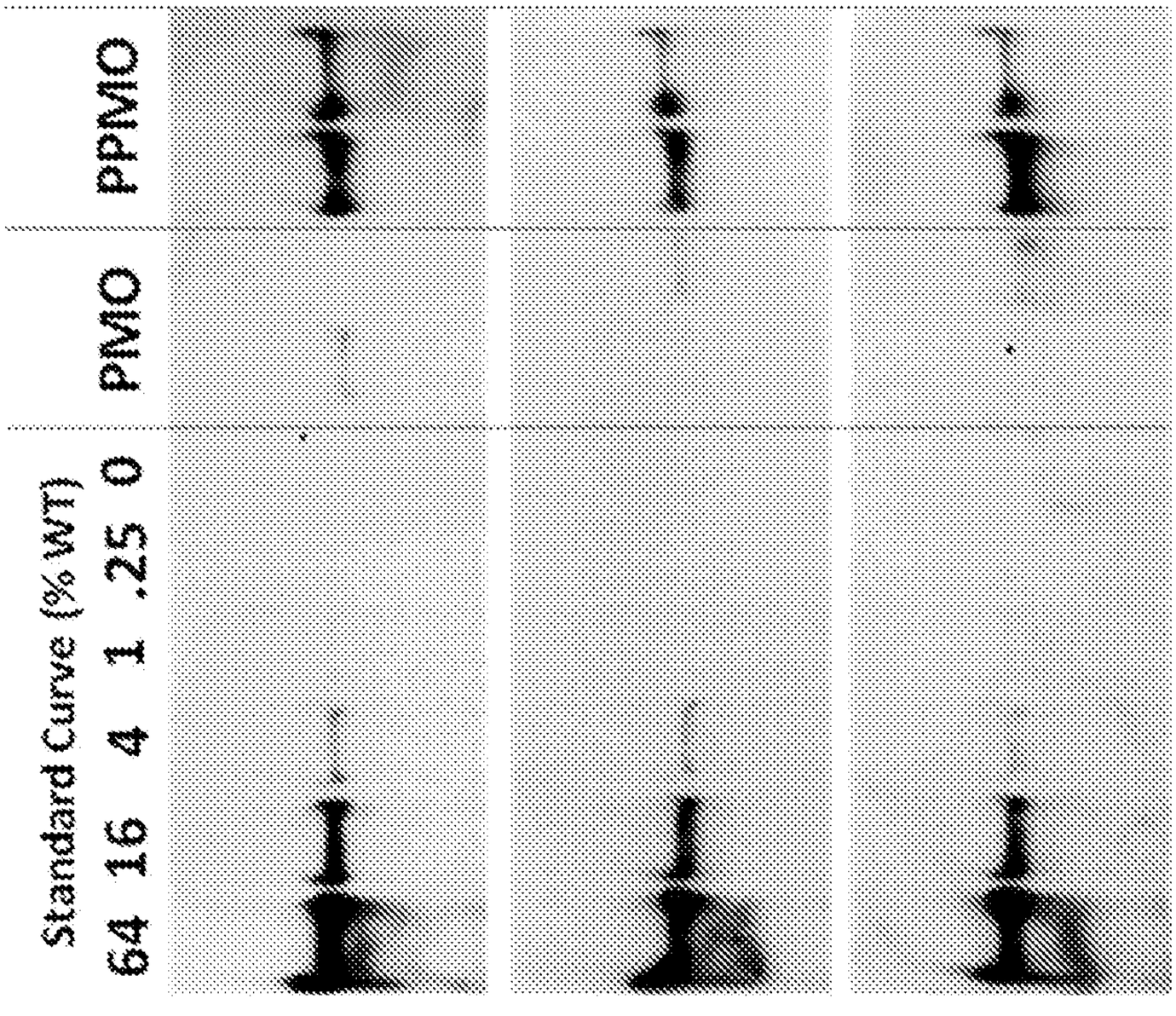


Figure 4D

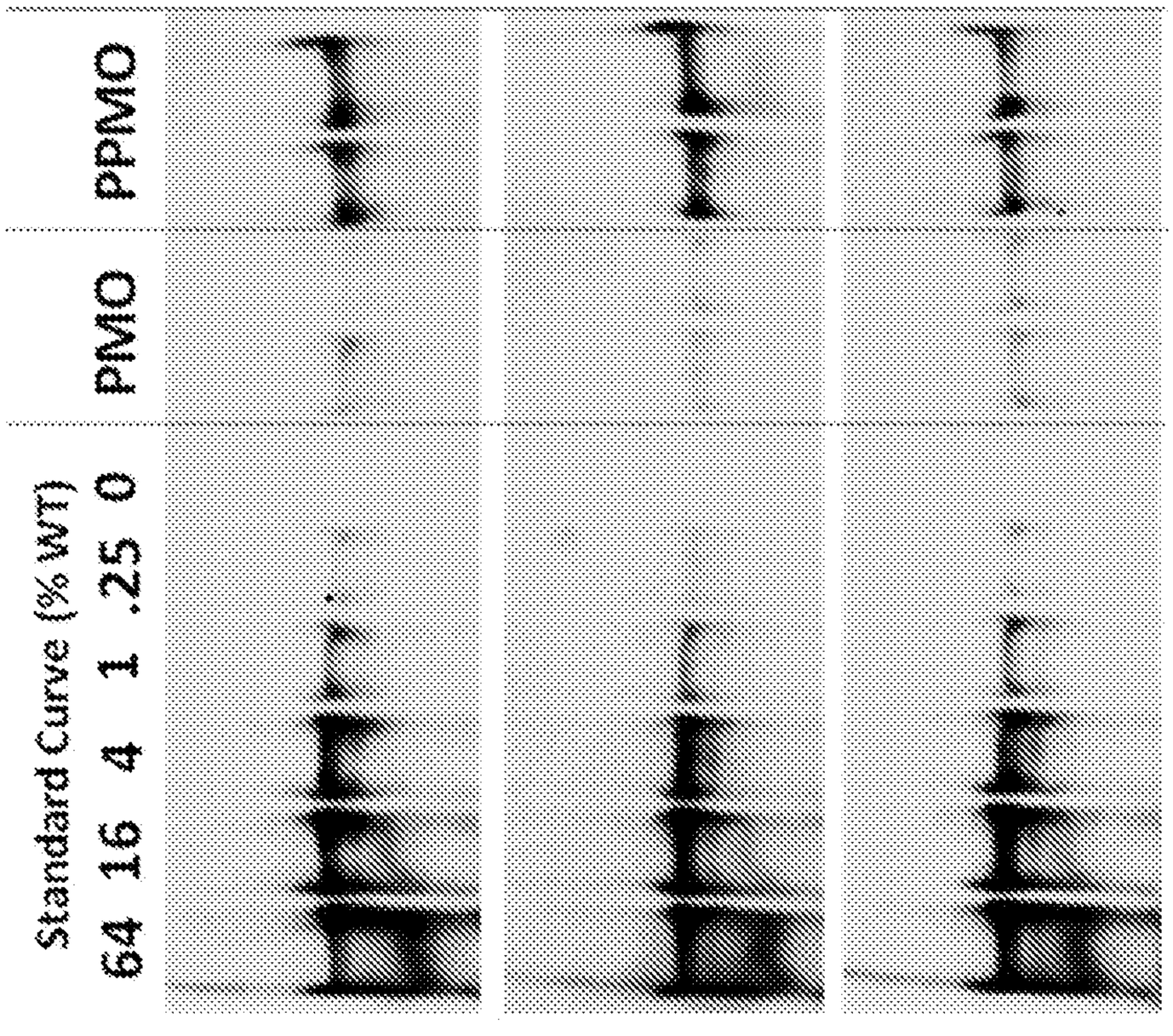


Figure 5B

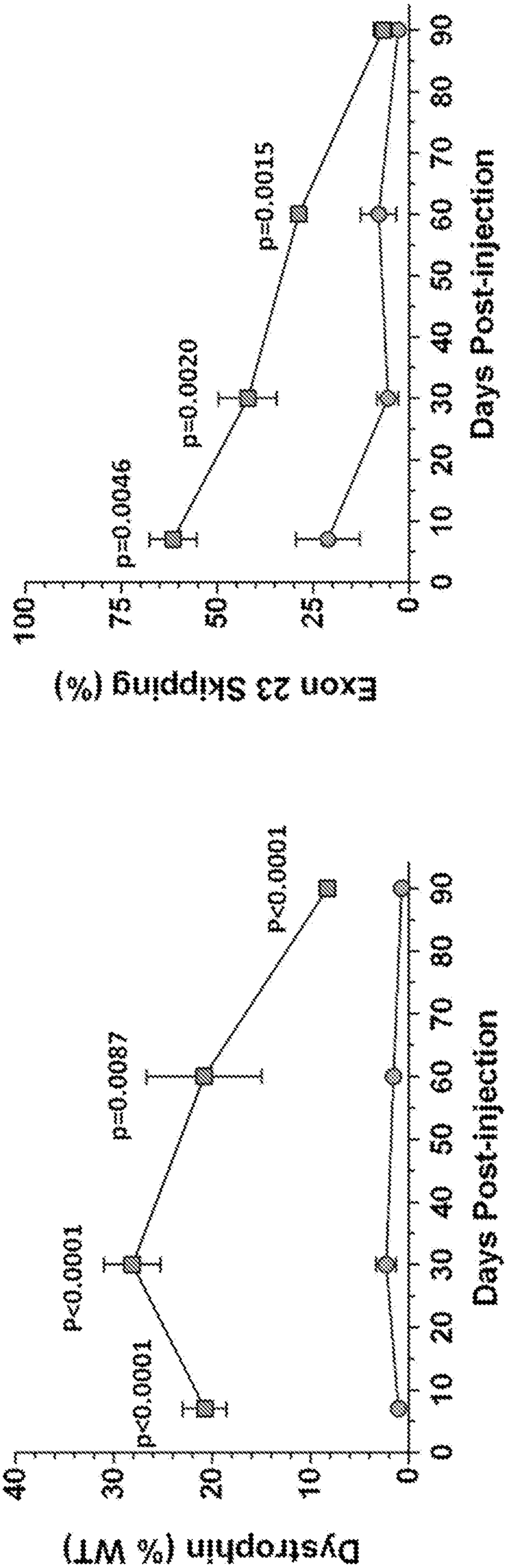


Figure 5A

Figure 6B
30 Days

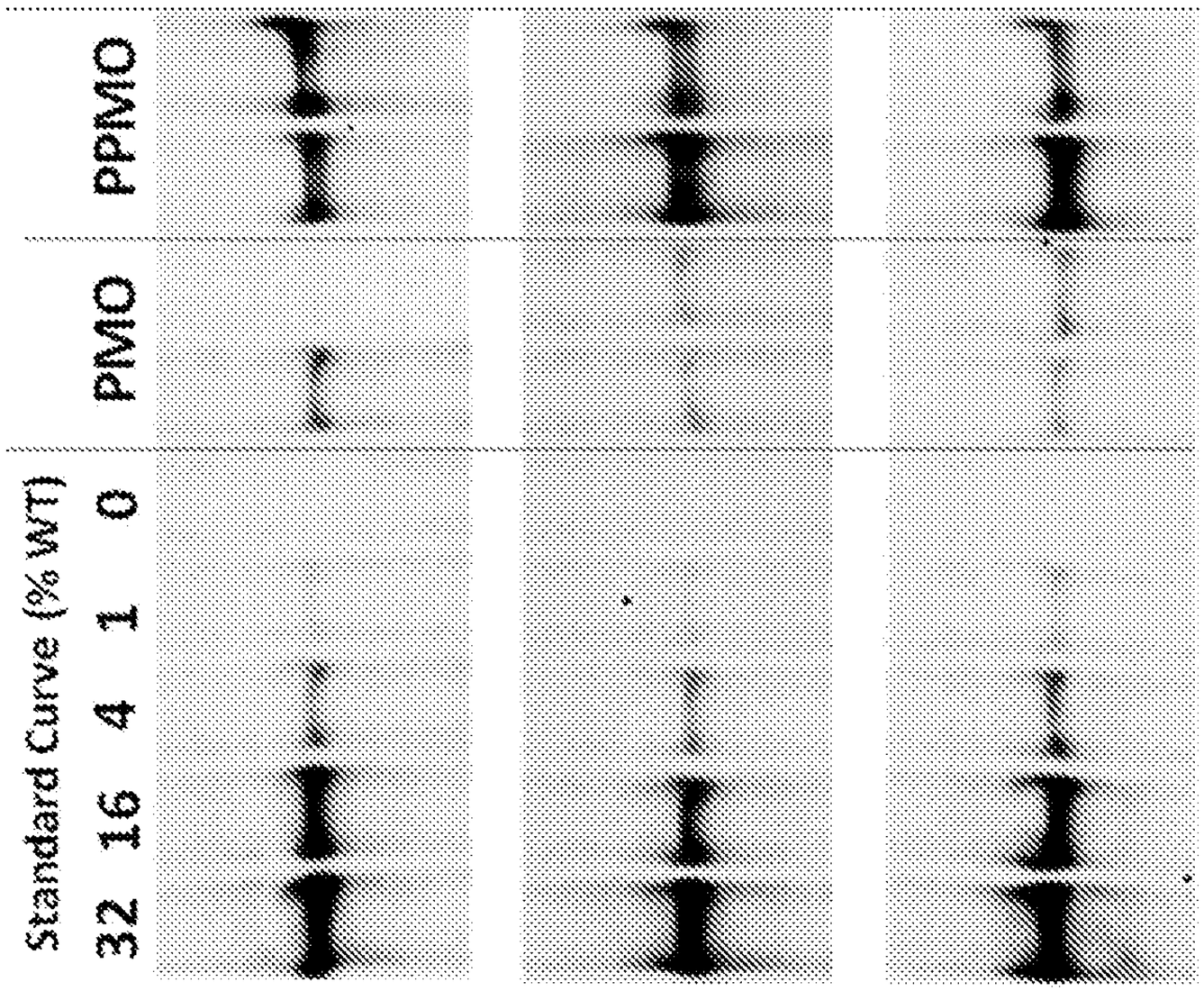


Figure 6A
7 Days

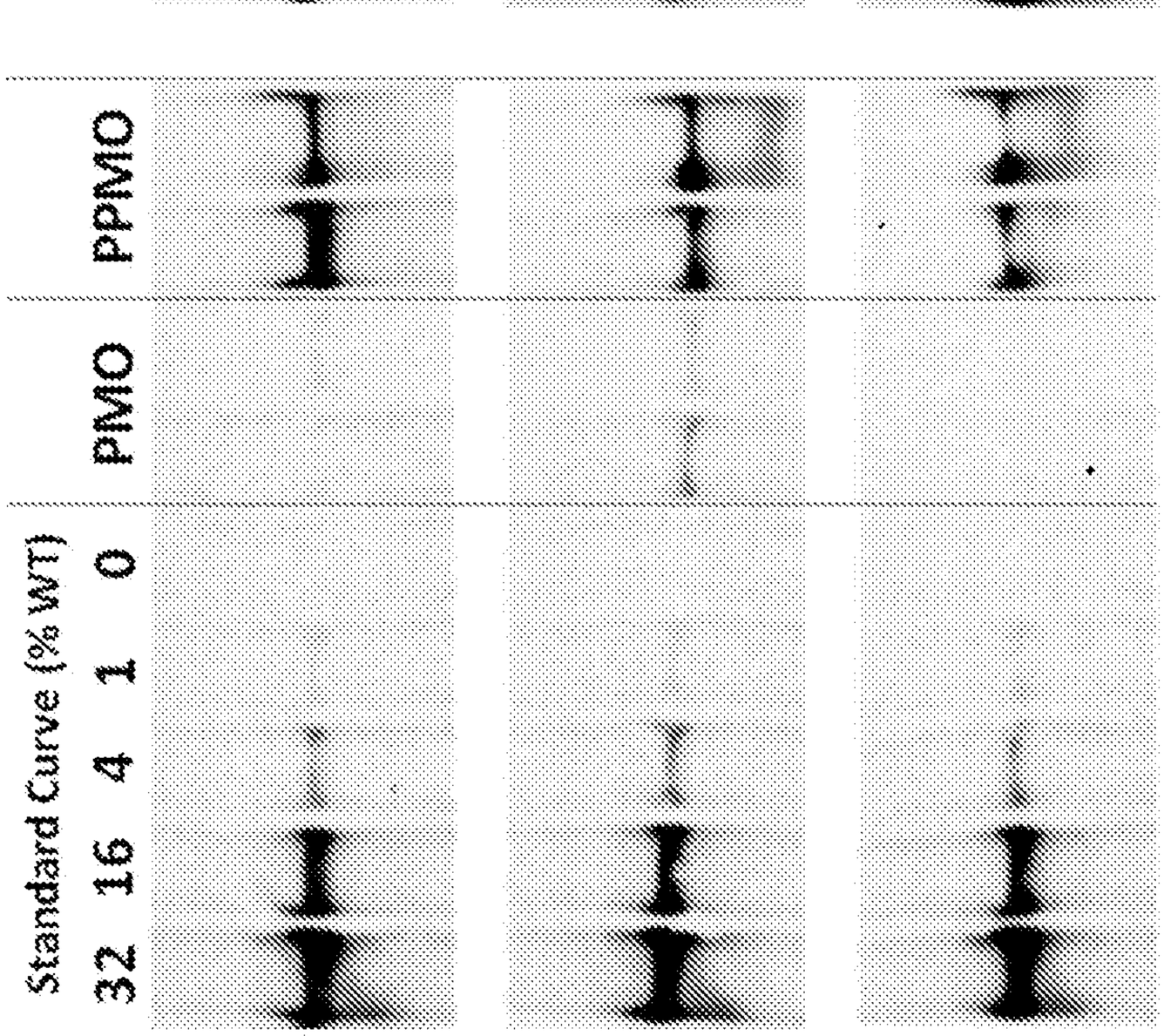


Figure 6C

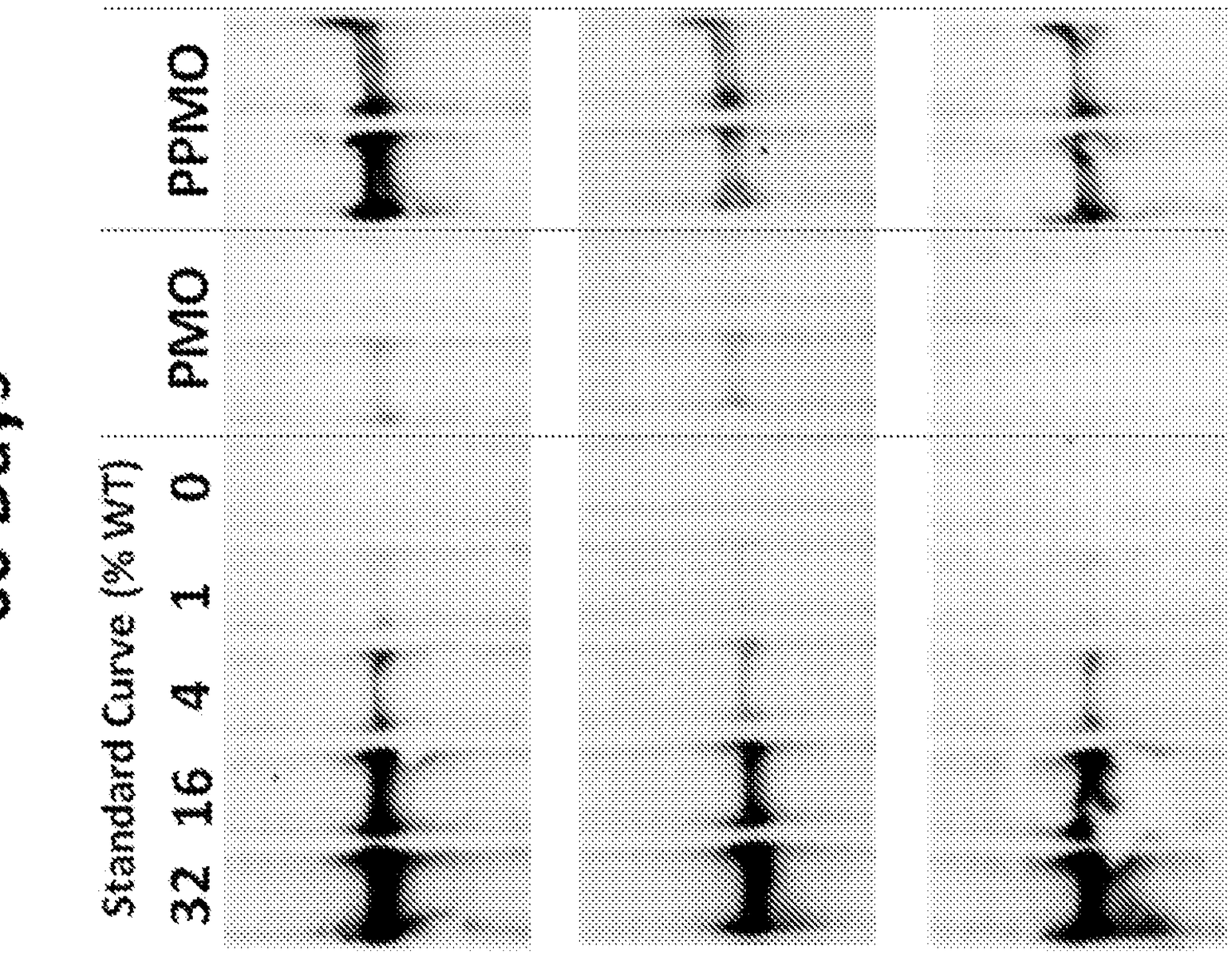


Figure 6D

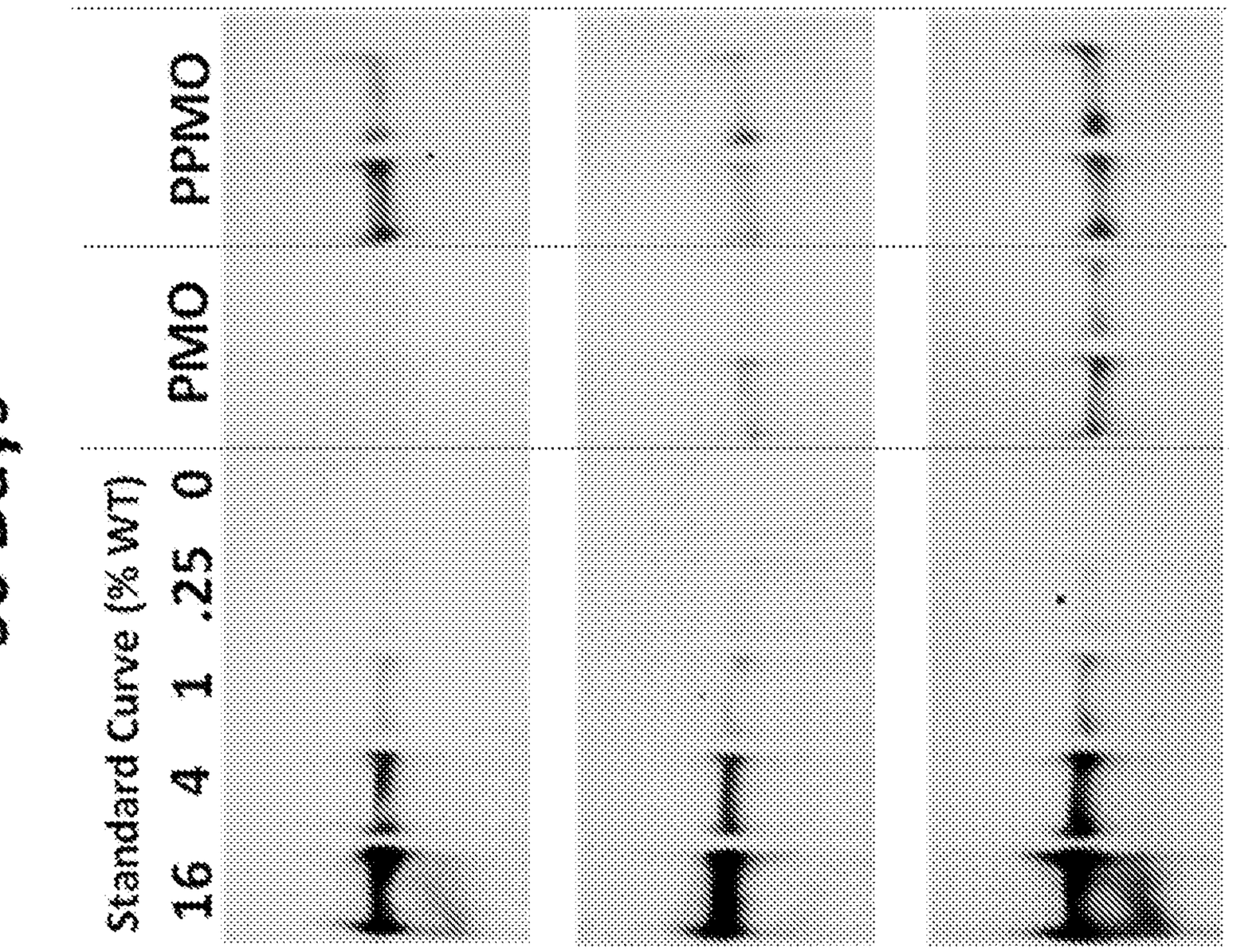


Figure 7A

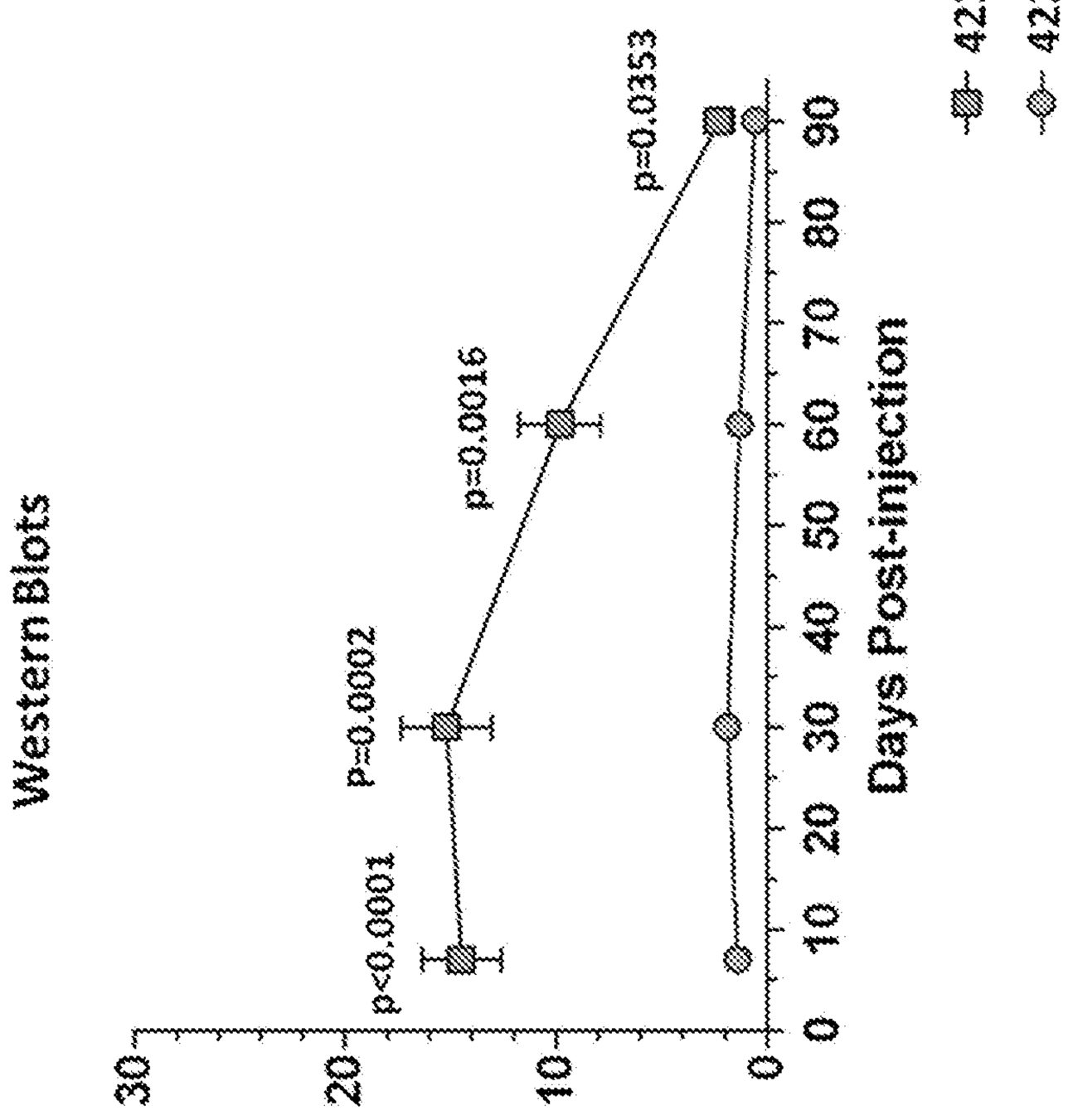


Figure 7B

Figure 8B

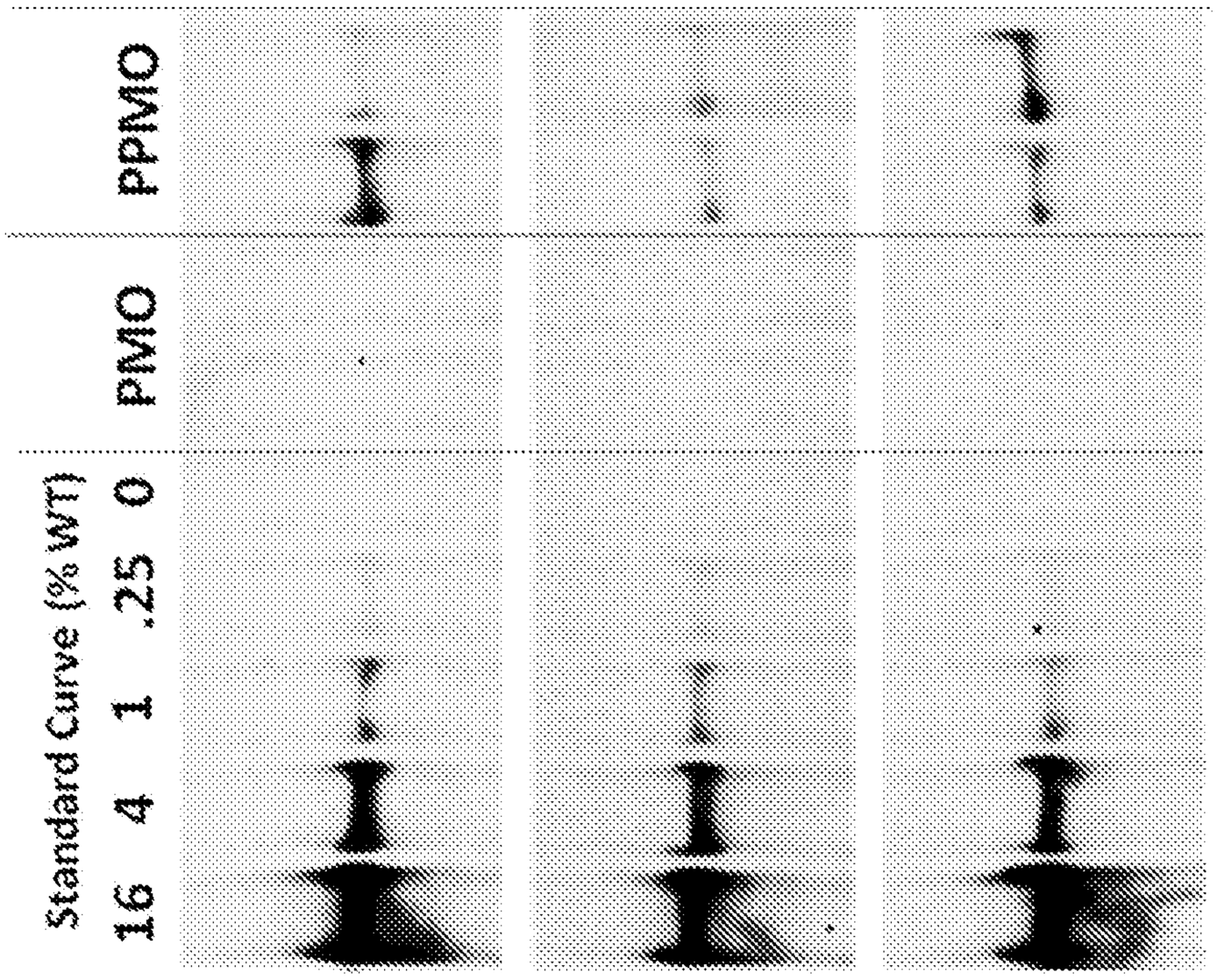


Figure 8A

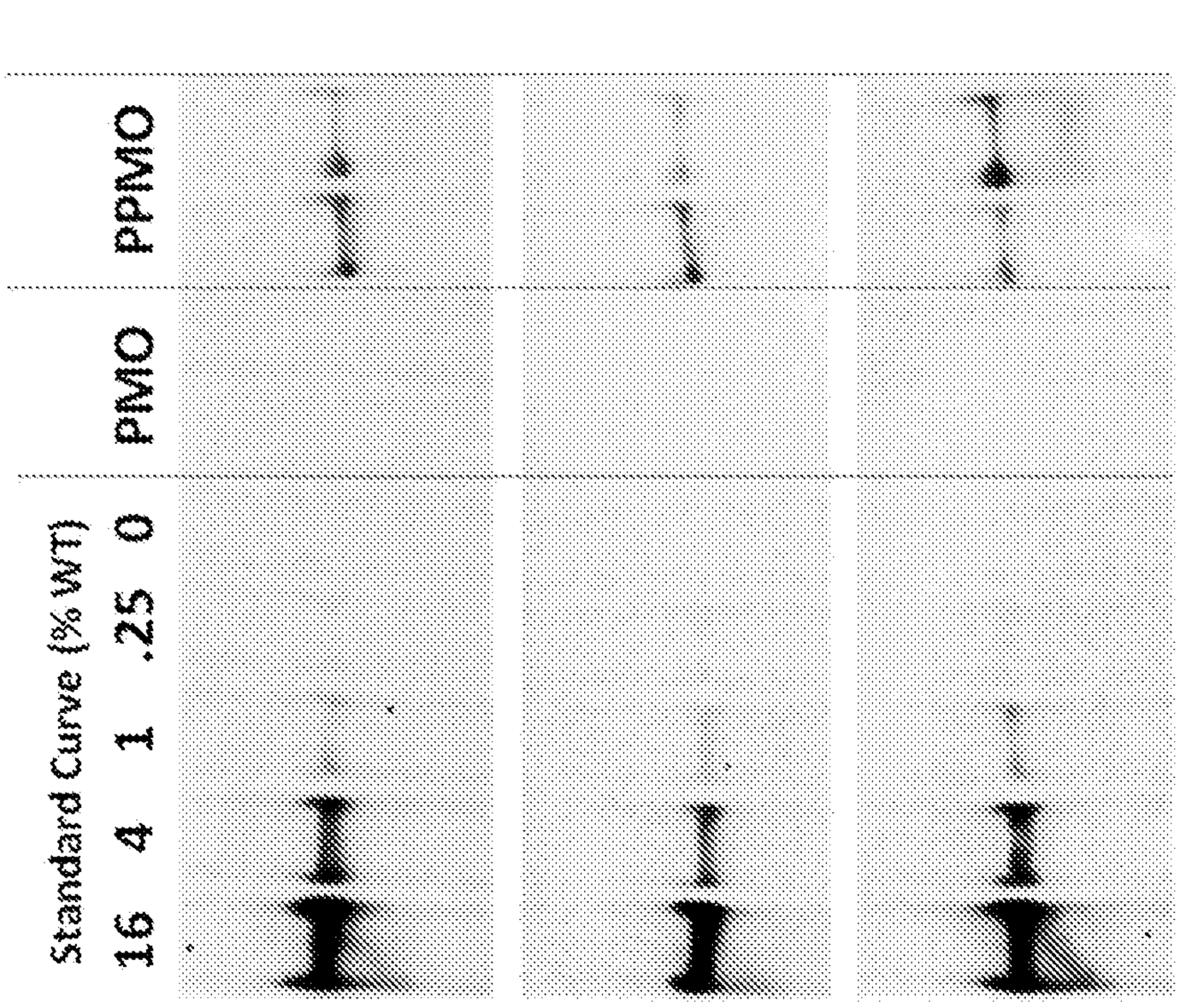


Figure 8C

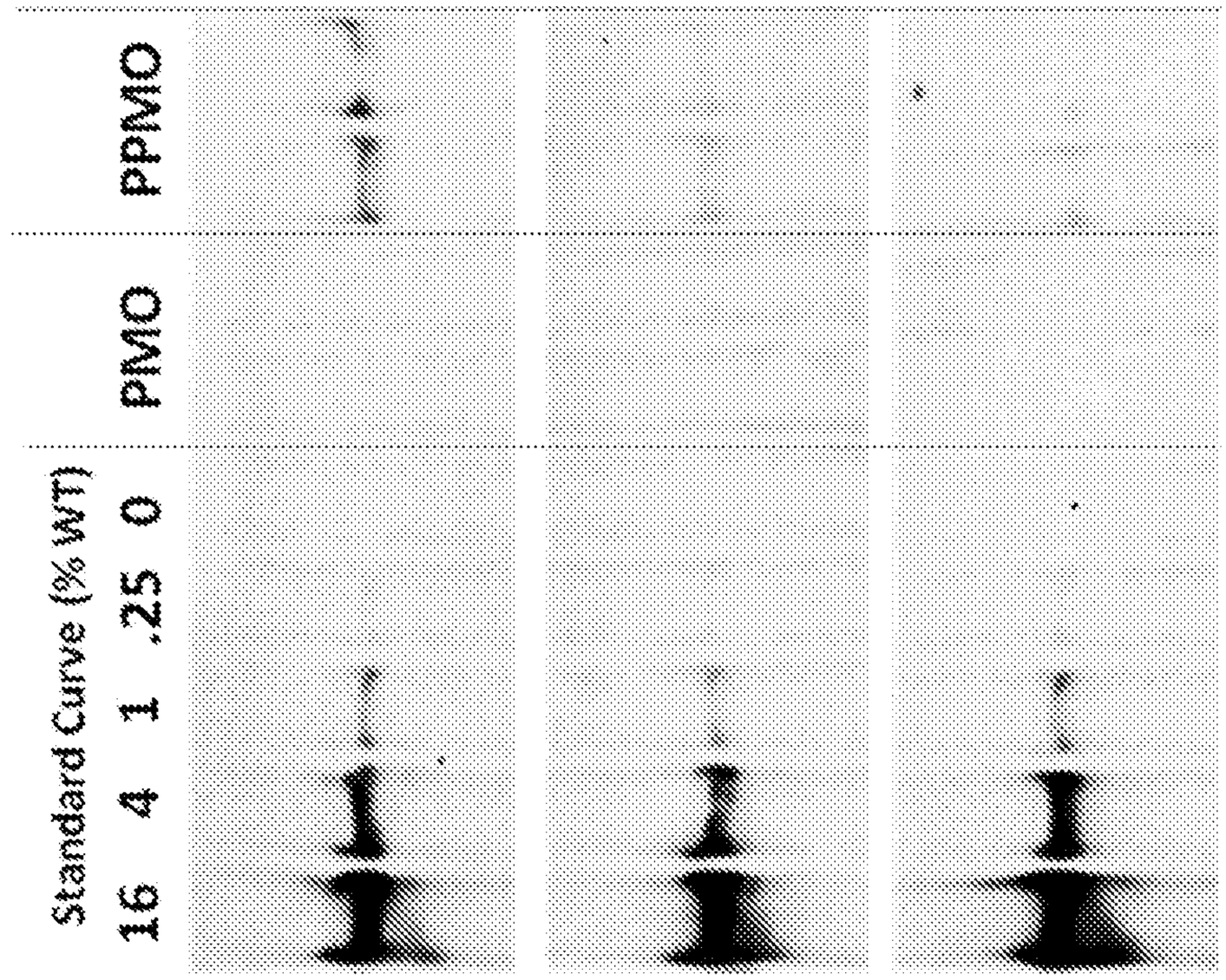


Figure 8D

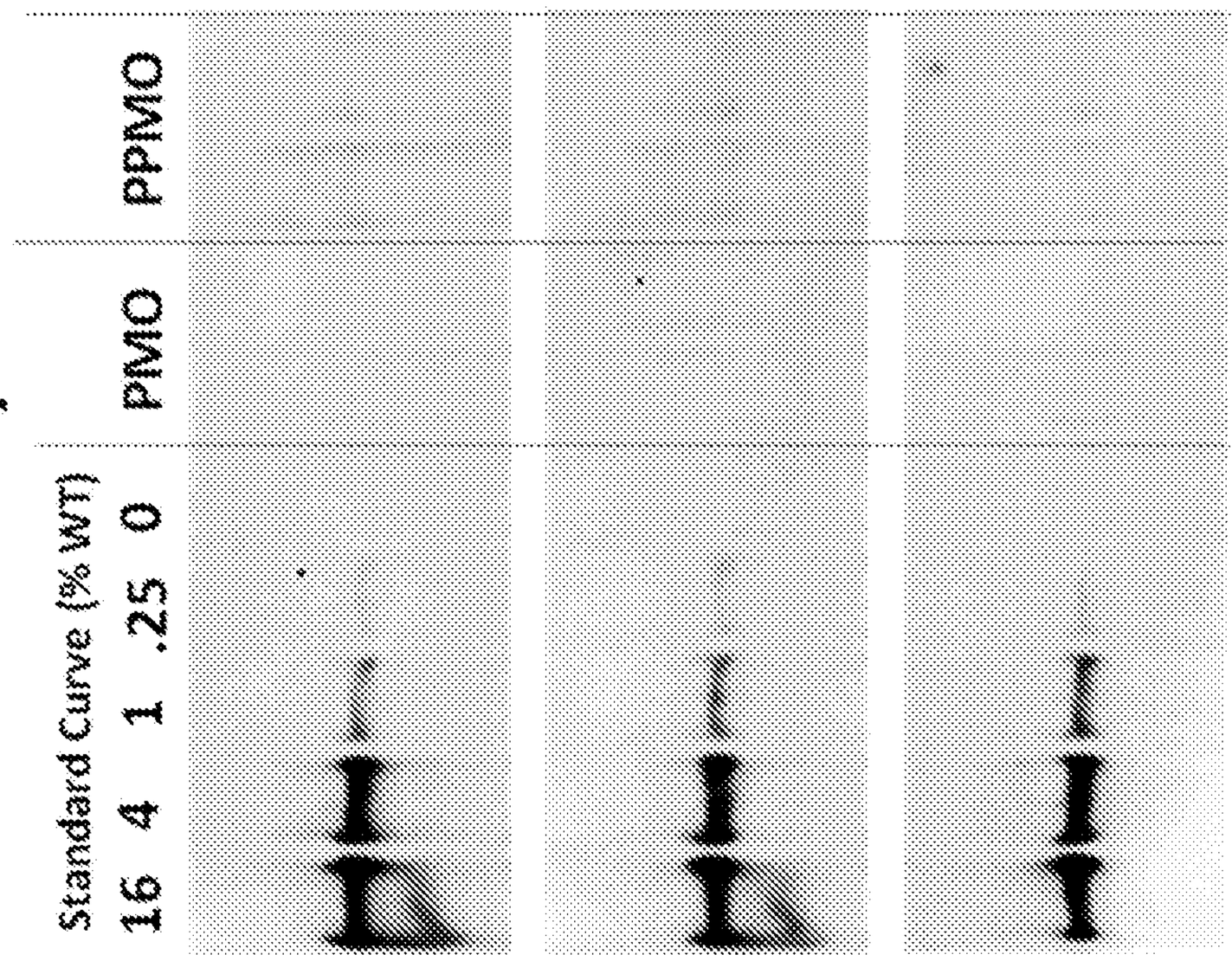


Figure 9A

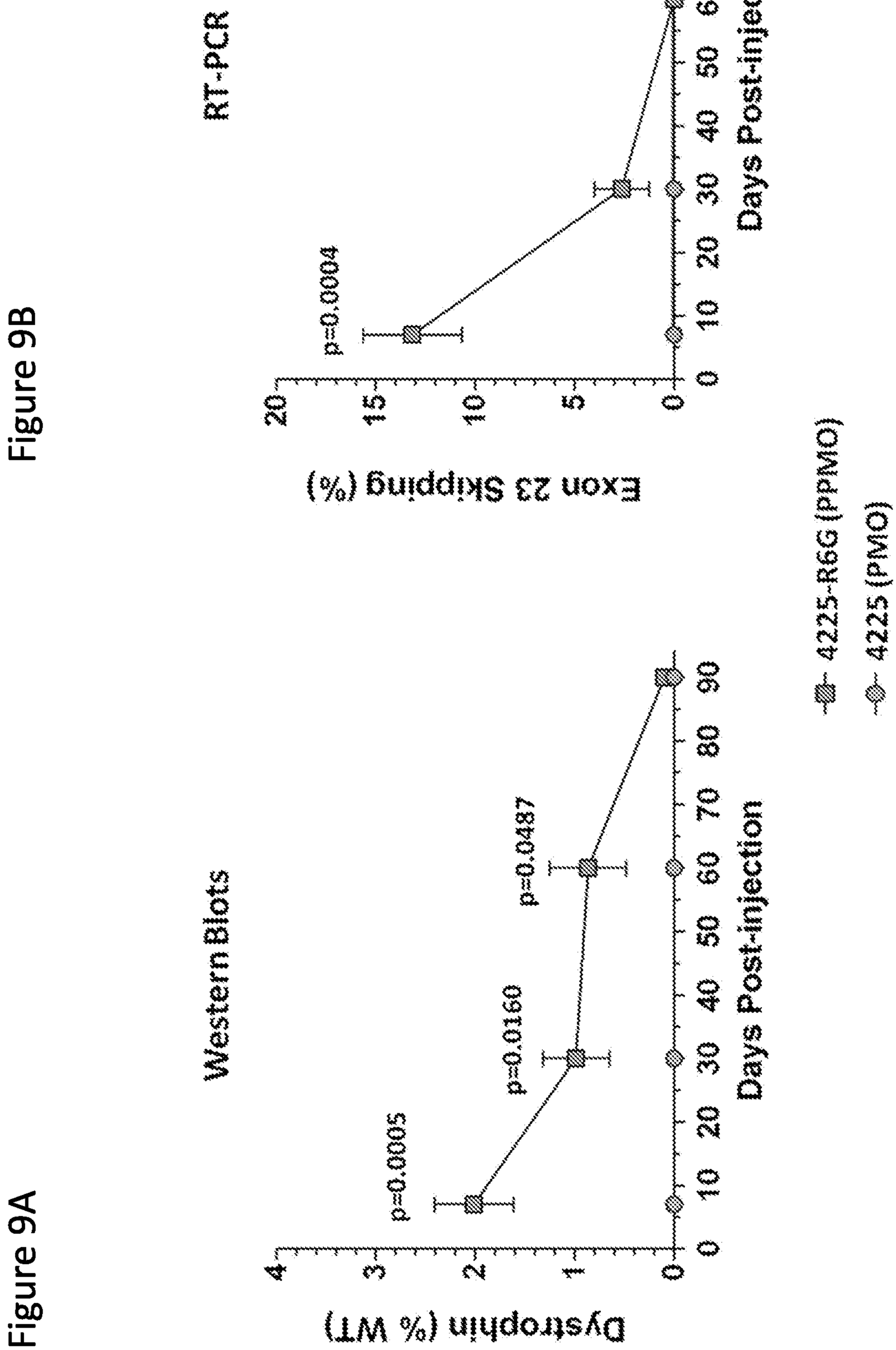


Figure 9B

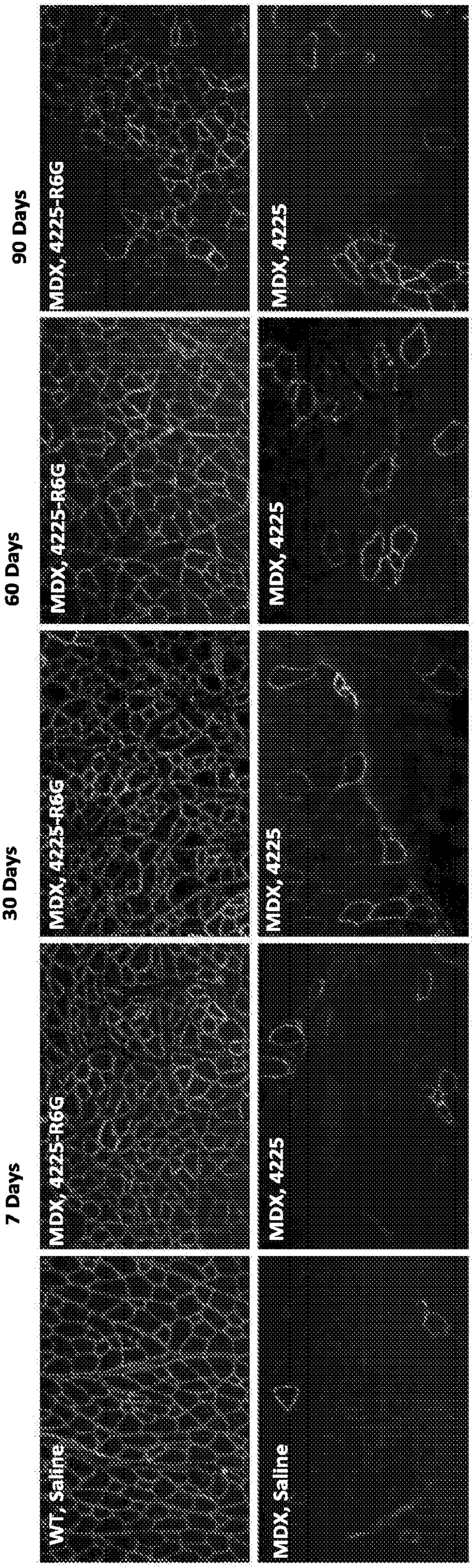

90 Days
MDX, 4225-R6G
MDX, 4225
60 Days
MDX, 4225-R6G
MDX, 4225
30 Days
MDX, 4225-R6G
MDX, 4225
7 Days
MDX, 4225-R6G
MDX, 4225
WT, Saline
MDX, Saline

Figure 10

Figure 11A

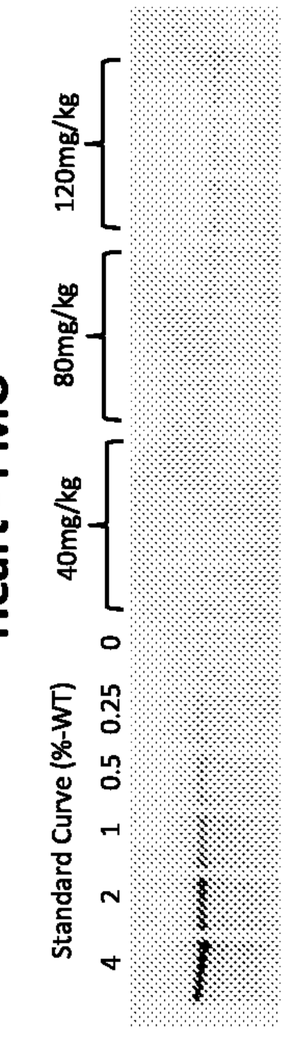
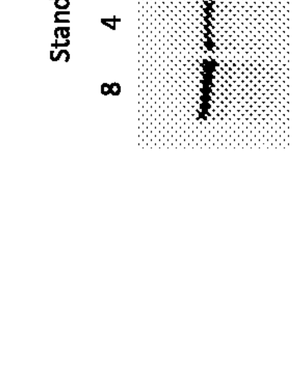


Figure 11B



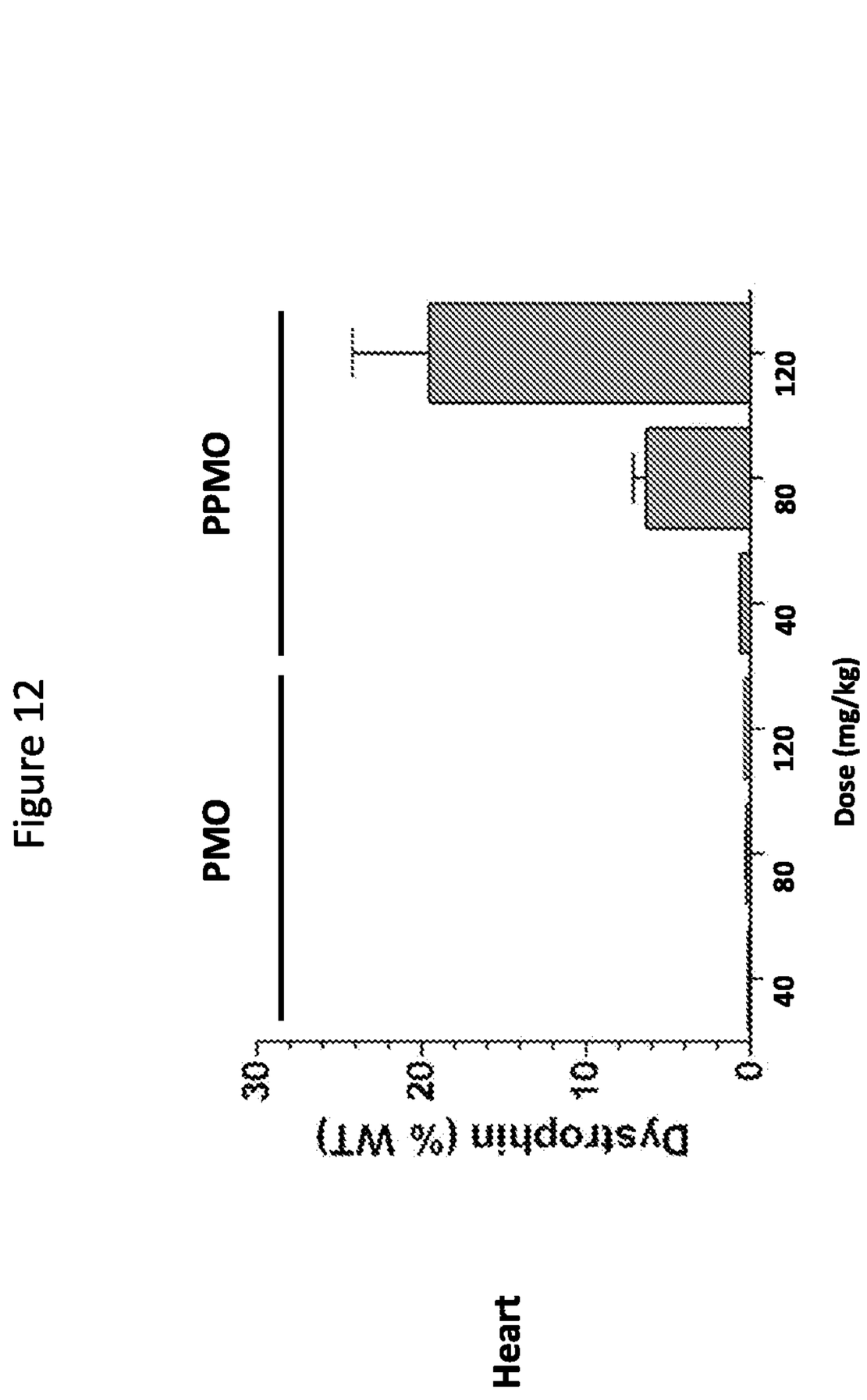


Figure 12

Figure 13A

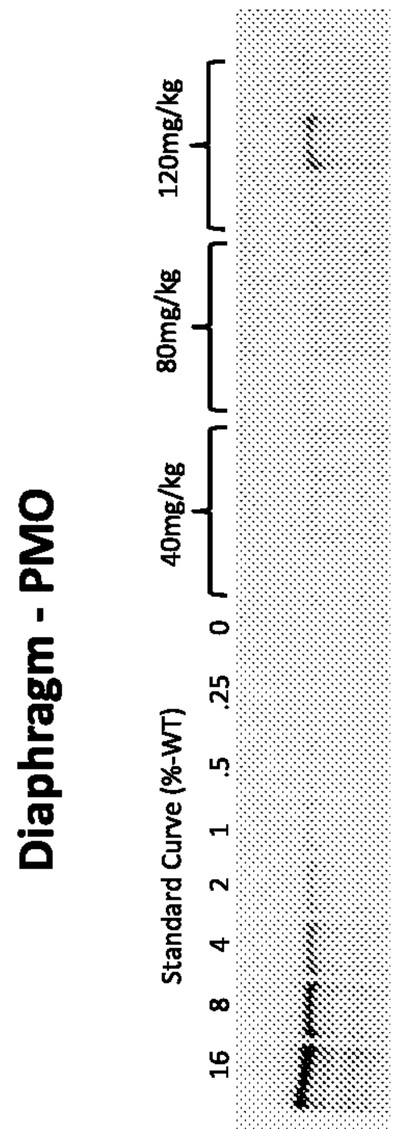
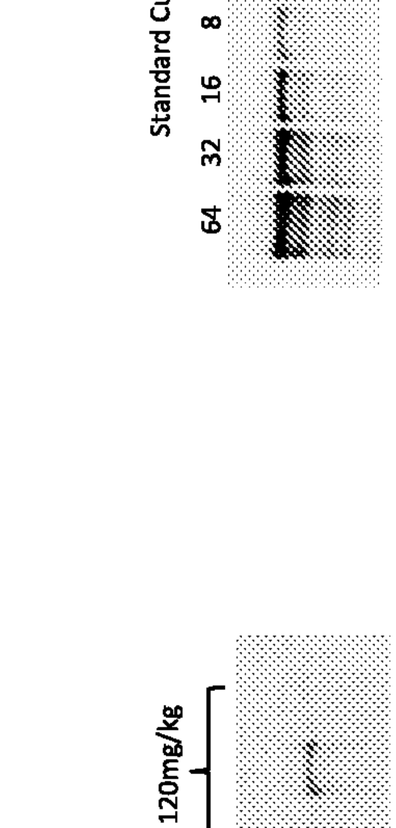


Figure 13B



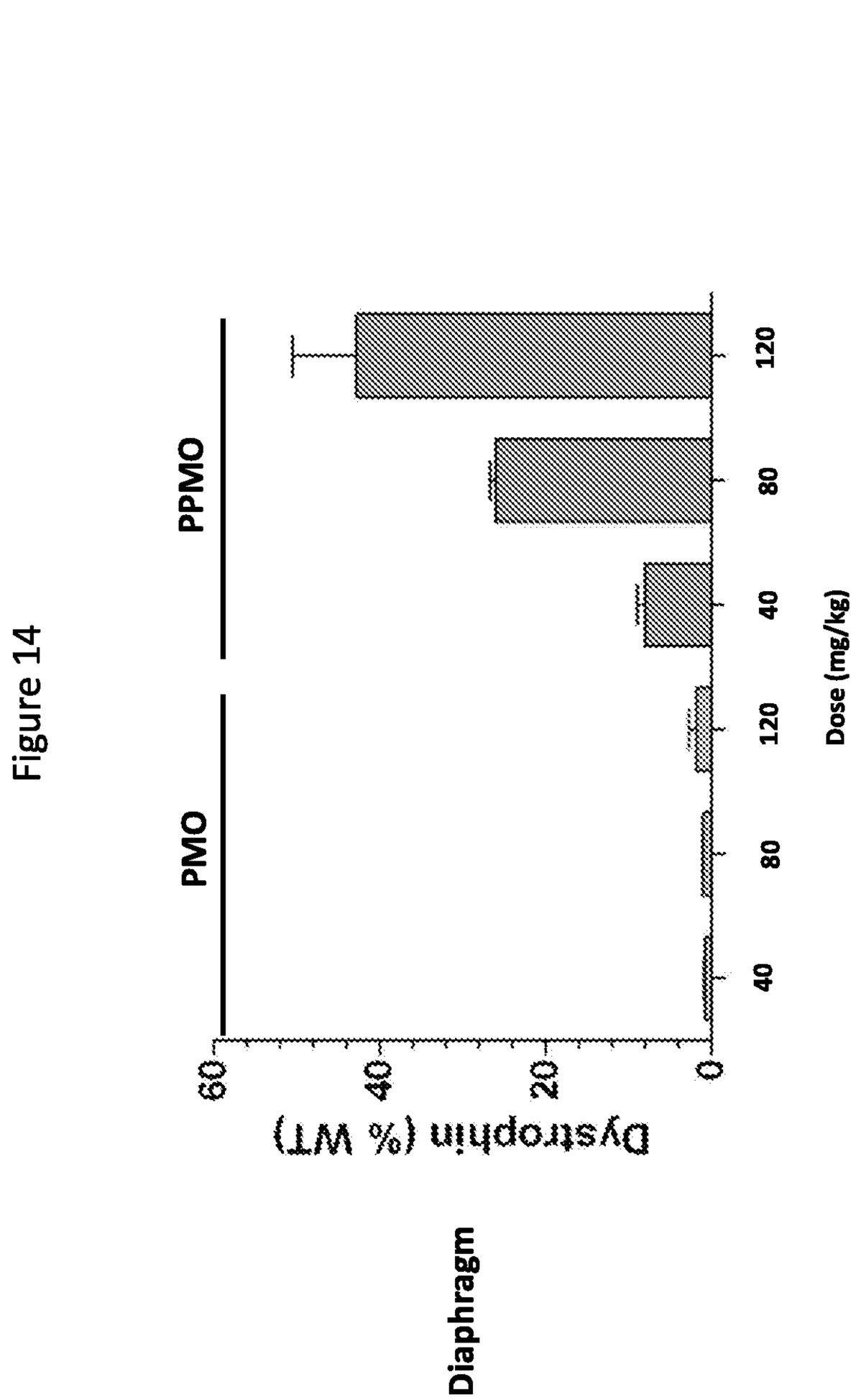

Diaphragm
Dystrophin (% WT)
0
20
40
60
PMO
PPMO
40
80
120
Dose (mg/kg)

Figure 14

Figure 15A

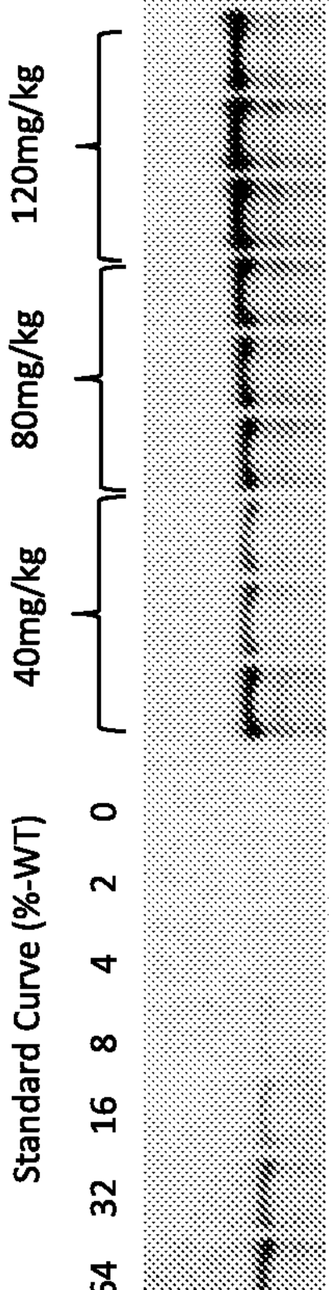
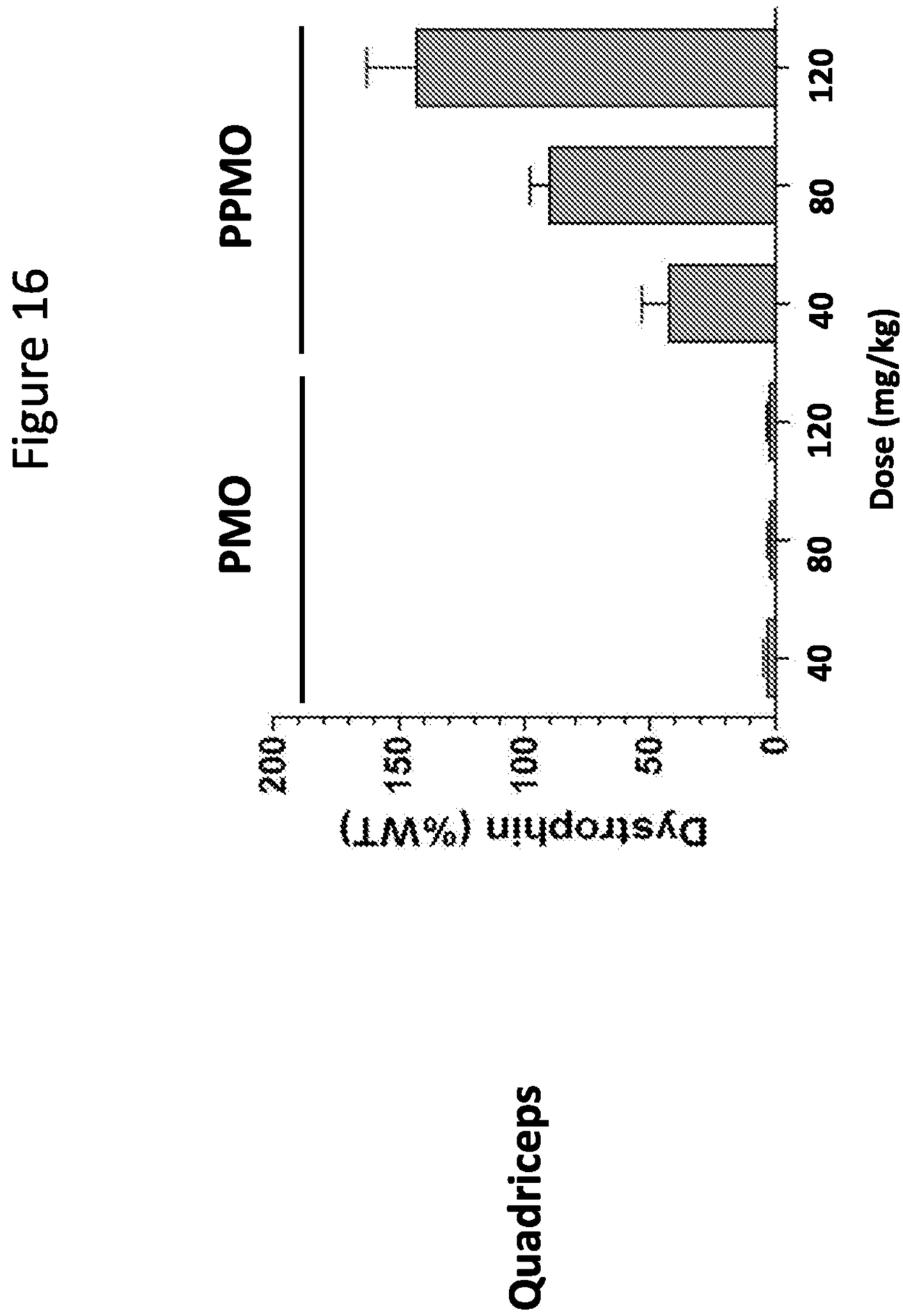


Figure 15B

Figure 16


PPMO
PMO
Dystrophin (%WT)
200
150
100
50
0
120
80
40
120
80
40
Dose (mg/kg)
Quadriceps

Cycle No.: Subunit (SU)	Pre-coupling Treatment				Coupling Cycle		Post-Coupling Treatment	
	1	2	3	4			1	2
	30% TFE/DCM Wash	CYTFA Solution[1]	Neutralization Solution	DCM Wash	Quantity SU (g) NEM (L) DMI (L)	RT Coupling Time (Hrs.)	DCM Wash	30% TFE/DCM Wash
1:C	5.5L	a) 5,5L b) 5.5L, 122ml	3x5.5L	5.5L	584g; 195 ml NEM; 3.2L DMI	5	5.5L	2x5.5L
2:A	7.0L	a) 7L b) 7L, 158ml	3x7L	2x7L	592.2g and 195ml NEM 3.2L DMI	4.25	7L	2x7L
3:A	8L	a) 8L b) 8L, 182ml	3x8L	2x8L	592.2g; 195ml NEM; 3.4L DMI	4.25	8L	2x8L
4:T	9L	a) 9L b) 9L, 206ml	3x9L	2x9L	514.2g; 195ml NEM; 3.6L DMI	4.25	9L	2x9L
5:G	9.5L	a) 9.5L b) 9.5L, 220ml	3x9.5L	2x9.5L	755.5g; 195ml NEM; 3.4L DMI	4.25	9.5L	2x9.5L
6:C	10L	a) 10L b) 10L, 232ml	3x10L	2x10L	584.4g; 195ml NEM; 3.45L DMI	4.25	10L	2x10L
7:C	11L	a) 11L b) 11L, 256ml	3x11L	2x11L	584.5g; 195ml NEM; 3.57L DMI	4.25	11L	2x11L
8:A	11L	a) 11L b) 11L, 256ml	3x11L	2x11L	592.5g; 195ml NEM; 3.64L DMI	4.25	11L	2x11L
9:T	11.5L	a) 11.5L b) 11.5L 268ml	3x11.5L	2x 11.5L	514.5g; 195ml NEM;	4.25	11.5L	2x11.5L

Figure 17

[1] ml indicates the amount of 1:1 NEM/DCM

Cycle No.: Subunit (SU)	Pre-coupling Treatment				Coupling Cycle		Post-Coupling Treatment	
	1	2	3	4			1	2
	30% TFE/DCM Wash	CYTFA Solution[1]	Neutralization Solution	DCM Wash	Quantity SU (g) NEM (L) DMI (L)	RT Coupling Time (Hrs.)	DCM Wash	30% TFE/DCM Wash
					3.72L DMI			
10:C	12L	a) 12L b) 12L, 280ml	3x12L	2x12L	338.5g; 195ml NEM; 3.96L DMI	4.25	12L	2x12L
11:C	13.5L	a) 13.5L b) 13.5L, 204ml	3x13.5L	2x 13.5L	770.4g; 253ml NEM; 4.02L DMI	4.25	13.5L	2x13.5L
12:T	13.5L	a) 13.5L b) 13.5L, 204ml	3x13.5L	2x 13.5L	668.7g; 253ml NEM; 4.02L DMI	4.25	13.5L	2x13.5L
13:DPG	14L	a) 14L b) 14L, 216ml	3x14L	2x14L	982.3g; 253ml NEM; 4.02L DMI	4.25	14L	2x14L
14:DPG	14.5L	a) 14.5L b) 14.5L, 228ml	3x14.5L	2x 14.5L	982.3g; 253ml NEM; 4.1L DMI	4.25	14.5L	2x14.5L
15:A	15.5L	a) 15.5L b) 15.5L, 254ml	3x15.5L	2x 15.5L	770.1g; 253ml NEM; 4.26L DMI	4.25	15.5L	2x15.5L
16:DPG	15.5L	a) 15.5L b) 15.5L, 254ml	3x15.5L	2x 15.5L	982.4g; 253ml NEM; 4.26L DMI	4.25	15.5L	2x15.5L
17:T	16L	a) 16L b) 16L, 366ml	3x16L	2x16L	549.6g; 253ml NEM;	4.75	16L	2x16L

Figure 17 (continued)

Cycle No.: Subunit (SU)	Pre-coupling Treatment				Coupling Cycle		Post-Coupling Treatment	
	1	2	3	4			1	2
	30% TFE/DCM Wash	CYTFA Solution[1]	Neutralization Solution	DCM Wash	Quantity SU (g) NEM (L) DMI (L)	RT Coupling Time (Hrs.)	DCM Wash	30% TFE/DCM Wash
					4.4L DMI			
18:T	16.5L	a) 16.5L b) 16.5L, 378ml	3x16.5L	2x 16.5L	630.7g; 253ml NEM; 4.4L DMI	4.25	16.5L	2x16.5L
19:C	16.5L	a) 16.5L b) 16.5L, 378ml	3x16.5L	2x 16.5L	770.4g; 253ml NEM; 4.57L DMI	4.25	16.5L	2x16.5L
20:C	17L	a) 17L b) 17L, 390ml	3x17L	2x17L	770.4g; 253ml NEM; 4.57L DMI	4.75	17L	2x17L
21:T	17L	a) 17L b) 17L, 390ml	3x17L	2x17L	776.4g; 311ml NEM; 4.72L DMI	4.25	17L	2x17L
22:DPG	17.5L	a) 17.5L b) 17.5L, 402ml	3x17.5L	2x 17.5L	1208.7g; 311ml NEM; 4.72L DMI	4.75	17.5L	2x17.5L

Figure 17 (continued)

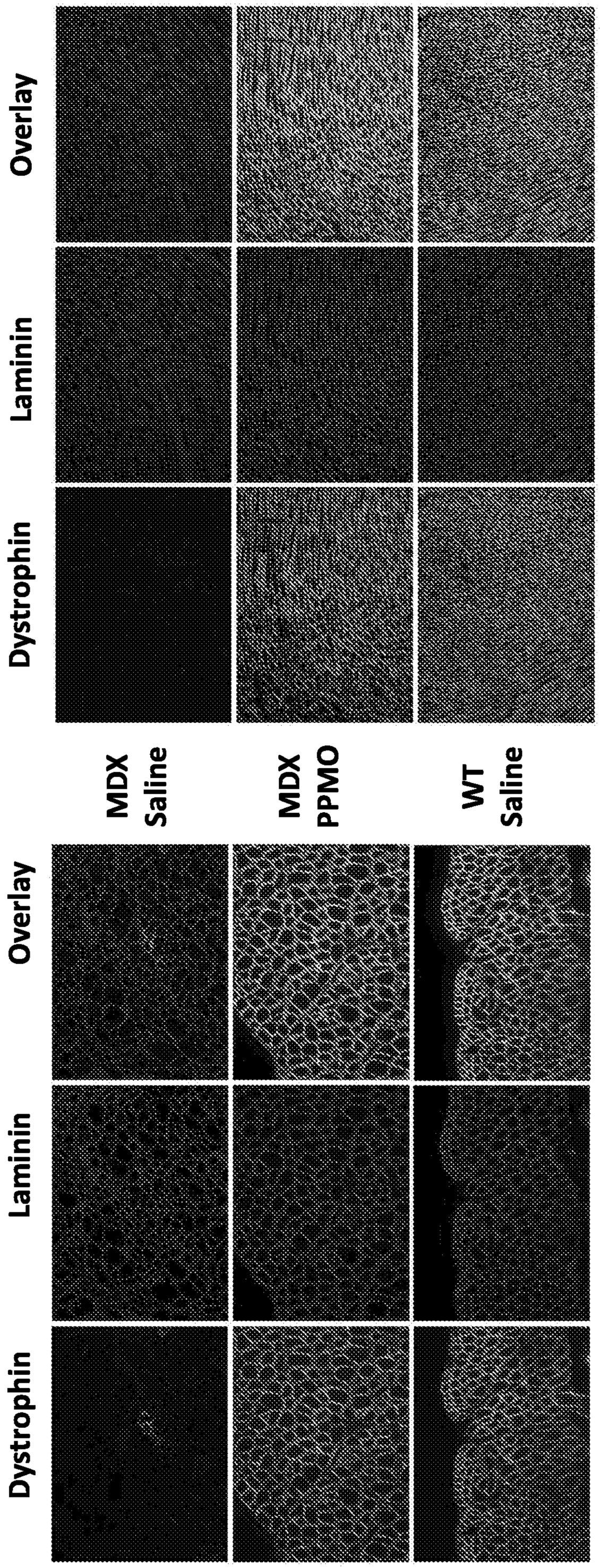

Heart
Dystrophin
Laminin
Overlay
MDX Saline
MDX PPMO
WT Saline
Diaphragm
Laminin
Overlay
Dystrophin

Figure 18

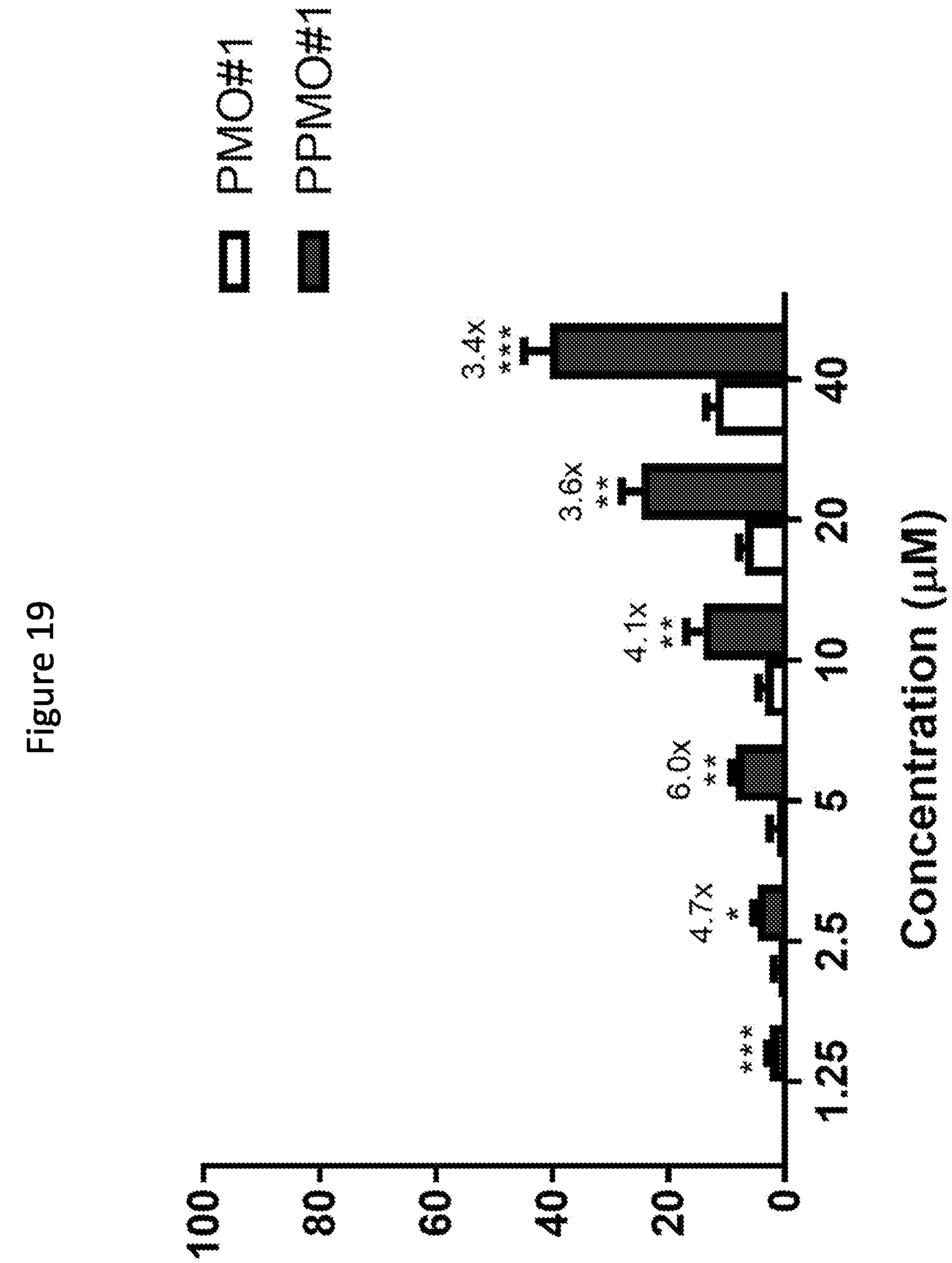

PPMO#1
PPMO#1
3.4x
***
3.6x
**
4.1x
**
6.0x
**
4.7x
*
***
40
20
10
5
2.5
1.25
Concentration (μM)
100
80
60
40
20
0
% skipping, DMD exon 45

Figure 19

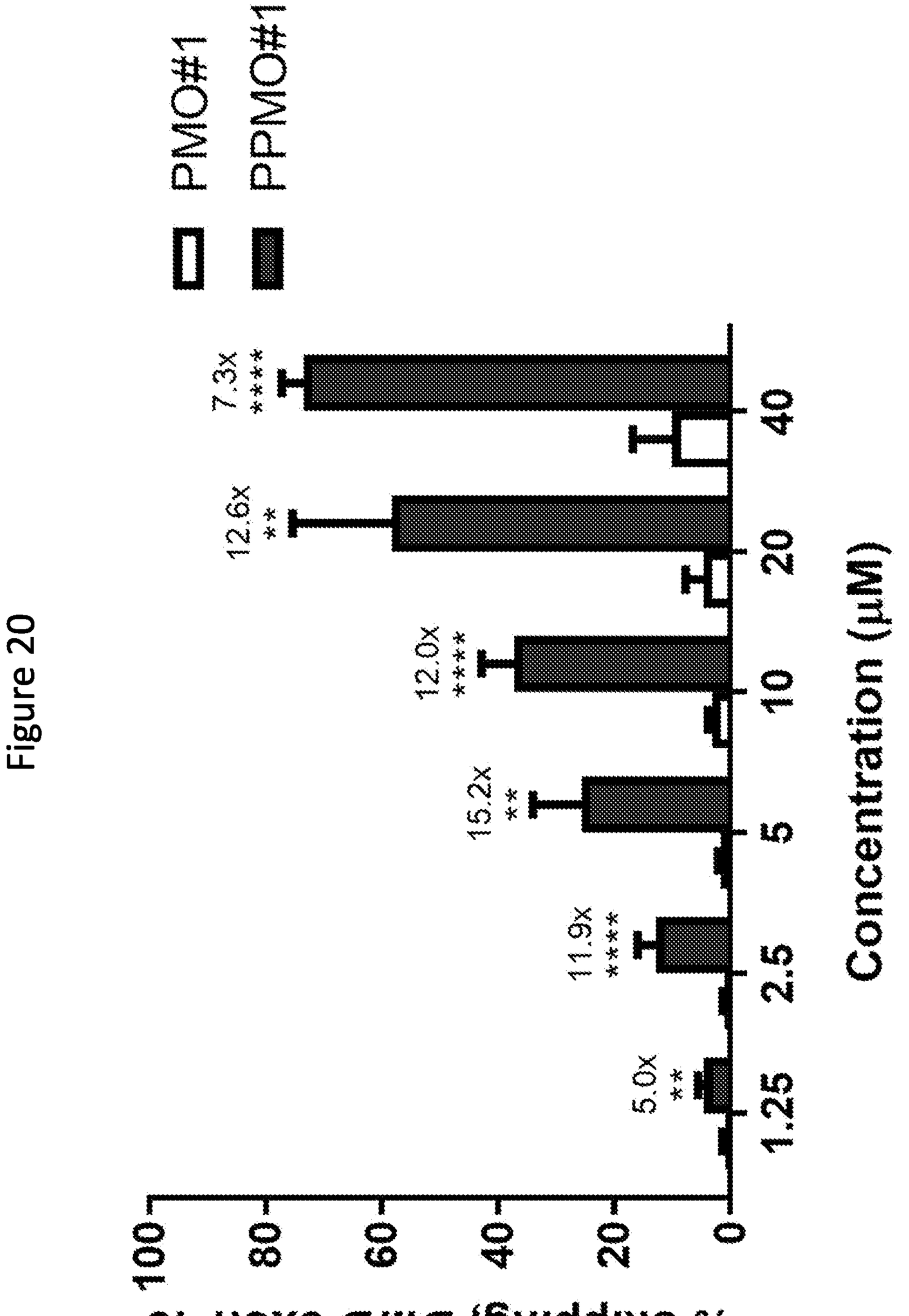

PPMO#1
PPMO#1
7.3x
****
12.6x
**
12.0x
****
15.2x
**
11.9x
****
5.0x
**
40
20
10
5
2.5
1.25
Concentration (μM)
100
80
60
40
20
0
% skipping, DMD exon 45

Figure 20