Appendix Table 1: Differences in sleep and quality of life characteristics between genders

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Median (IQR)			
Sleep latency (minutes)	30 (10-50)	30 (20-60)	30 (15-60)	0.23 (0.03-0.44; p<0.01)
Sleep duration (hours)	6 (4.5-6.5)	6 (5-7)	6 (4.7-6.8)	0.10 (-0.10-0.30; p=0.99)
Time in bed (hours)	7 (6-8)	7.5 (6.5-8.5)	7 (6-8)	0.37 (0.17-0.57; p<0.01)
	Number of patients (%)			
Sleep latency				
Normative (≤30 minutes)	174 (63)	80 (52)	254 (59)	0.61 (0.40-0.94; p<0.05)
Prolonged (>31 minutes)	93 (34)	70 (45)	163 (38)	
Sleep duration				
Normative (6-9 hors)	144 (52)	74 (48)	218 (51)	1.20 (0.81-1.90; p=0.37)
Disturbed (≤5.9 or ≥9.1 hours)	124 (45)	78 (51)	202 (47)	
	Median (IQR)			
Quality of life (EQ-5, 0-10)	8 (7-10)	9 (8-10)	9 (7-10)	0.19 (-0.004-0.39; p=0.19)

IQR, Inter quartile range; CI, Confidence interval; NPS, Numerical pain scale; EQ-5, EuroQoL questionnaire.

Appendix Table 2: Sex differences in specific MC related AEs

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Central nervous system				
Confusion	21 (8)	24 (16)	45 (10)	2.20 (1.10-4.40; p<0.05)
Confusion frequency				
Rarely	17 (6)	12 (8)	29 (7)	0.16 (0.06-0.25; p<0.05)
Frequently	4 (2)	10 (7)	14 (3)	
Constantly	0	1 (<1)	1 (<1)	
Confusion severity				
Mild	16 (6)	11 (7)	27 (6)	0.16 (0.06-0.25; p<0.05)
Moderate	5 (2)	10 (7)	15 (4)	
Severe	0	2 (1)	0	
Disorientation	4 (2)	6 (4)	10 (2)	2.70 (0.64-13.0; p=0.20)
Disorientation frequency				
Rarely	1 (<1)	4 (3)	5 (1)	0.10 (0.009-0.2; p=0.09)
Frequently	2 (<1)	2 (1)	4 (<1)	
Constantly	0	0	0	
Disorientation severity				
Mild	1 (<1)	5 (3)	6 (1)	0.12 (0.2-0.21; p=0.051)
Moderate	2 (<1)	1 (<1)	3 (<1)	
Severe	0	0	0	
Impaired attention	28 (10)	32 (21)	60 (14)	2.30 (1.30-4.20; p<0.005)
Impaired attention frequency				
Rarely	8 (3)	2 (1)	10 (2)	0.19 (0.09-0.28; p<0.005)
Frequently	15 (6)	21 (14)	36 (8)	
Constantly	3 (1)	7 (5)	10 (2)	
Impaired attention severity				
Mild	11 (4)	6 (4)	17 (4)	0.19 (0.09-0.28; p<0.005)
Moderate	14 (5)	14 (9)	28 (7)	
Severe	2 (<1)	10 (7)	12 (3)	
Dizziness	12 (4)	26 (17)	38 (9)	4.40 (2.10-10.0; p<0.001)
Dizziness frequency				
Rarely	8 (3)	10 (7)	18 (4)	0.23 (0.14-0.32; p<0.001)
Frequently	4 (2)	13 (8)	17 (4)	
Constantly	0	3 (2)	3 (<1)	
Dizziness severity				
Mild	6 (2)	5 (3)	11 (3)	0.24 (0.15-0.32; p<0.001)
Moderate	6 (2)	16 (10)	22 (5)	
Severe	0	5 (3)	5 (1)	
Increased awareness	37 (13)	23 (15)	60 (14)	1.10 (0.61-2.00; p=0.78)

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Increased awareness frequency				
Rarely	10 (4)	2 (1)	12 (3)	0.23 (0-0.47; p=0.21)
Frequently	15 (6)	12 (8)	27 (6)	
Constantly	10 (4)	8 (5)	18 (4)	
Increased awareness severity				
Mild	13 (5)	8 (5)	21 (5)	0.10 (0.009-0.20; p=0.20)
Moderate	19 (7)	7 (5)	26 (6)	
Severe	4 (2)	7 (5)	11 (3)	
Decreased awareness	14 (5)	15 (10)	29 (7)	2.00 (0.88-4.60; p=0.10)
Decreased awareness frequency				
Rarely	7 (3)	4 (3)	11 (3)	0.14 (0.04-0.23; p<0.05)
Frequently	4 (2)	10 (7)	14 (3)	
Constantly	1 (<1)	0	1 (<1)	
Decreased awareness severity				
Mild	8 (3)	5 (3)	13 (3)	0.13 (0.03-0.22; p=0.07)
Moderate	4 (2)	9 (6)	13 (3)	
Severe	1 (<1)	0	1 (<1)	
Decreased physical sensation	11 (4)	6 (4)	17 (4)	0.97 (0.29-2.90; p=1.00)
Decreased physical sensation frequency				
Rarely	2 (<1)	2 (1)	4 (<1)	0.03 (0-0.13; p=0.92)
Frequently	7 (3)	3 (2)	10 (2)	
Constantly	2 (<1)	1 (<1)	3 (<1)	
Decreased physical sensation severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.03 (0-0.13; p=0.90)
Moderate	6 (2)	3 (2)	9 (2)	
Severe	3 (1)	1 (<1)	4 (<1)	
Intoxication feeling	8 (3)	5 (3)	13 (3)	1.10 (0.28-4.00; p=1.00)
Intoxication feeling frequency				
Rarely	6 (2)	3 (2)	9 (2)	0.02 (0-0.12; p=0.83)
Frequently	2 (<1)	2 (1)	4 (<1)	
Constantly	0	0	0	
Intoxication feeling severity				
Mild	5 (2)	2 (1)	7 (2)	0.04 (0-0.13; p=0.71)
Moderate	3 (1)	3 (2)	6 (1)	
Severe	0	0	0	
Impaired balance	13 (5)	24 (16)	37 (9)	3.70 (1.70-8.20; p<0.001)
Impaired balance frequency				
Rarely	6 (2)	7 (5)	13 (3)	0.21 (0.12-0.30; p<0.001)

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Frequently	5 (2)	17 (11)	22 (5)	
Constantly	0	0	0	
Impaired balance severity				
Mild	6 (2)	7 (5)	13 (3)	0.21 (0.12-0.30; p<0.001)
Moderate	6 (2)	13 (8)	19 (4)	
Severe	0	4 (3)	4 (<1)	
Fatigue	108 (39)	88 (57)	196 (45)	2.10 (1.40-3.10; p<0.001)
Fatigue frequency				
Rarely	20 (8)	6 (4)	26 (6)	0.26 (0.17-0.34; p<0.001)
Frequently	70 (25)	48 (31)	118 (28)	
Constantly	15 (6)	31 (20)	46 (11)	
Fatigue severity				
Mild	28 (10)	9 (6)	37 (9)	0.29 (0.20-0.38; p<0.001)
Moderate	67 (24)	43 (28)	110 (26)	
Severe	12 (4)	34 (22)	46 (11)	
Impaired memory	34 (12)	45 (29)	79 (18)	2.90 (1.70-5.00; p<0.001)
Impaired memory frequency				
Rarely	6 (2)	10 (7)	16 (4)	0.21 (0.12-0.30; p<0.001)
Frequently	19 (7)	23 (15)	42 (10)	
Constantly	7 (3)	11 (7)	18 (4)	
Impaired memory severity				
Mild	10 (4)	12 (8)	22 (5)	0.22 (0.13-0.31; p<0.001)
Moderate	17 (6)	18 (12)	35 (8)	
Severe	6 (4)	14 (9)	20 (5)	
Impaired coordination	6 (2)	13 (8)	19 (4)	4.10 (1.40-14.00; p<0.005)
Impaired coordination frequency				
Rarely	3 (1)	5 (3)	8 (2)	0.15 (0.05-0.24; p<0.05)
Frequently	2 (<1)	7 (5)	9 (2)	
Constantly	0	0	0	
Impaired coordination severity				
Mild	5 (2)	4 (3)	9 (2)	0.19 (0.09-0.28; p<0.005)
Moderate	0	7 (5)	7 (2)	
Severe	0	1 (<1)	1 (<1)	
Impaired speech	10 (4)	13 (8)	23 (5)	2.40 (0.96-6.40; p<0.01)
Impaired speech frequency				
Rarely	4 (2)	5 (3)	9 (2)	0.10 (0.006-0.19; p=0.23)
Frequently	3 (1)	4 (3)	7 (2)	
Constantly	2 (<1)	3 (2)	5 (1)	
Impaired speech severity				
Mild	6 (2)	4 (3)	10 (2)	0.13 (0.03-0.22; p=0.06)

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Moderate	2 (<1)	7 (5)	9 (2)	
Severe	1 (<1)	1 (<1)	2 (<1)	
Psychological				
<u>Unusual thinking</u>	12 (5)	11 (7)	23 (5)	1.70 (0.65-4.30; p=0.32)
Unusual thinking frequency				
Rarely	9 (4)	5 (3)	14 (3)	0.05 (0-0.15; p=0.50)
Frequently	3 (1)	4 (3)	7 (2)	
Constantly	0	0	0	
Unusual thinking severity				
Mild	5 (2)	5 (3)	10 (2)	0.05 (0-0.15; p=0.51)
Moderate	6 (2)	5 (3)	11 (3)	
Severe	0	0	0	
<u>Anxiety</u>	19 (7)	22 (14)	41 (10)	2.20 (1.10-4.50; p<0.05)
Anxiety frequency				
Rarely	6 (2)	7 (5)	13 (3)	0.11 (0.01-0.20; p=0.15)
Frequently	9 (3)	9 (6)	18 (4)	
Constantly	4 (2)	5 (3)	9 (2)	
Anxiety severity				
Mild	4 (2)	3 (2)	7 (2)	0.12 (0.03-0.22; p=0.08)
Moderate	10 (4)	9 (6)	19 (4)	
Severe	5 (2)	9 (6)	14 (3)	
<u>Dysphoria</u>	39 (14)	45 (29)	84 (20)	2.50 (1.50-4.20; p<0.001)
Dysphoria frequency				
Rarely	12 (4)	13 (8)	25 (6)	0.19 (0.09-0.28; p<0.005)
Frequently	20 (7)	22 (14)	42 (10)	
Constantly	5 (2)	9 (6)	14 (3)	
Dysphoria severity				
Mild	12 (4)	11 (7)	23 (5)	0.18 (0.09-0.27; p<0.005)
Moderate	21 (8)	25 (16)	46 (11)	
Severe	5 (2)	8 (5)	13 (3)	
<u>Hyperactivity</u>	10 (4)	8 (5)	18 (4)	1.50 (0.49-4.20; p=0.60)
Hyperactivity frequency				
Rarely	1 (<1)	3 (2)	4 (<1)	0.08 (0-0.18; p=0.36)
Frequently	8 (3)	3 (2)	11 (3)	
Constantly	1 (<1)	1 (<1)	2 (<1)	
Hyperactivity severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.06 (0-0.16; p=0.63)
Moderate	7 (3)	3 (2)	10 (2)	
Severe	1 (<1)	2 (1)	3 (<1)	
<u>Euphoria</u>	15 (6)	3 (2)	18 (4)	0.35 (0.06-1.20; p=0.14)
Euphoria frequency				

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Rarely	5 (2)	0	5 (1)	0.09 (0.003-0.19; p=0.25)
Frequently	9 (3)	3 (2)	12 (3)	
Constantly	1 (<1)	0	1 (<1)	
Euphoria severity				
Mild	8 (3)	0	8 (2)	0.11 (0.01-0.20; p=0.08)
Moderate	6 (2)	2 (1)	8 (2)	
Severe	0	0	0	
Loss of time sensation	14 (5)	13 (8)	27 (6)	1.70 (0.72-4.10; p=0.24)
Loss of time sensation frequency				
Rarely	11 (4)	4 (3)	15 (4)	0.13 (0.035-0.22; p<0.05)
Frequently	3 (1)	8 (5)	11 (3)	
Constantly	0	0	0	
Loss of time sensation severity				
Mild	10 (4)	7 (5)	17 (4)	0.08 (0-0.18; p=0.35)
Moderate	3 (1)	5 (3)	8 (2)	
Severe	1 (<1)	0	1 (<1)	
Forgetfulness	35 (13)	34 (22)	69 (16)	1.90 (1.10-3.40; p<0.05)
Forgetfulness frequency				
Rarely	13 (5)	10 (7)	23 (5)	0.15 (0.05-0.24; p<0.05)
Frequently	14 (5)	15 (10)	29 (7)	
Constantly	4 (2)	8 (5)	12 (3)	
Forgetfulness severity				
Mild	15 (6)	12 (8)	27 (6)	0.16 (0.06-0.25; p<0.05)
Moderate	15 (6)	10 (7)	25 (6)	
Severe	4 (2)	11 (7)	15 (4)	
Nervousness	36 (13)	29 (19)	65 (15)	1.50 (0.87-2.70; p=0.15)
Nervousness frequency				
Rarely	14 (5)	7 (5)	21 (5)	0.08 (0-0.18; p=0.41)
Frequently	18 (7)	17 (11)	35 (8)	
Constantly	4 (2)	3 (2)	7 (2)	
Nervousness severity				
Mild	9 (4)	5 (3)	14 (3)	0.07 (0-0.16; p=0.54)
Moderate	18 (7)	16 (10)	34 (8)	
Severe	9 (3)	6 (4)	14 (3)	
Stress	38 (14)	33 (21)	71 (17)	1.70 (0.98-2.90; p=0.058)
Stress frequency				
Rarely	16 (6)	9 (6)	25 (6)	0.13 (0.03-0.22; p=0.056)
Frequently	19 (7)	14 (9)	33 (8)	
Constantly	3 (1)	8 (5)	11 (3)	
Stress severity				

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Mild	14 (5)	5 (3)	19 (5)	0.14 (0.04-0.23; p<0.05)
Moderate	19 (7)	17 (11)	36 (8)	
Severe	5 (2)	9 (6)	14 (3)	
Tantrums	17 (6)	12 (8)	29 (7)	1.30 (0.54-2.90; p=0.66)
Tantrums frequency				
Rarely	7 (3)	3 (2)	10 (2)	0.09 (0-0.19; p=0.28)
Frequently	7 (3)	8 (5)	15 (4)	
Constantly	3 (1)	0	3 (<1)	
Tantrums severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.04 (0-0.14; p=0.85)
Moderate	8 (3)	6 (4)	14 (3)	
Severe	7 (3)	3 (2)	10 (2)	
Gastrointestinal				
Abdominal discomfort	35 (13)	29 (19)	64 (15)	1.60 (0.89-2.80; p=0.12)
Abdominal discomfort frequency				
Rarely	9 (3)	4 (3)	13 (3)	0.12 (0.02-0.21; p=0.12)
Frequently	18 (7)	12 (8)	30 (7)	
Constantly	8 (3)	12 (8)	20 (5)	
Abdominal discomfort severity				
Mild	8 (3)	6 (4)	14 (3)	0.11 (0.01-0.20; p=0.19)
Moderate	18 (7)	10 (7)	28 (7)	
Severe	9 (3)	12 (8)	21 (5)	
Abdominal pain	23 (8)	32 (21)	55 (13)	2.90 (1.60-5.40; p<0.001)
Abdominal pain frequency				
Rarely	5 (2)	3 (2)	8 (2)	0.18 (0.08-0.27; p<0.005)
Frequently	10 (4)	17 (11)	27 (6)	
Constantly	7 (3)	10 (7)	17 (4)	
Abdominal pain severity				
Mild	4 (3)	2 (1)	6 (1)	0.19 (0.09-0.28; p<0.005)
Moderate	10 (4)	13 (8)	23 (5)	
Severe	8 (3)	16 (10)	24 (6)	
Heartburn	20 (7)	10 (7)	30 (7)	0.89 (0.36-2.00; p=0.92)
Heartburn frequency				
Rarely	6 (2)	3 (2)	9 (2)	0.02 (0-0.12; p=0.98)
Frequently	11 (4)	5 (3)	16 (4)	
Constantly	2 (<1)	1 (<1)	3 (<1)	
Heartburn severity				
Mild	6 (2)	5 (3)	11 (3)	0.06 (0-0.16; p=0.58)
Moderate	9 (3)	2 (1)	11 (3)	
Severe	3 (1)	2 (1)	5 (1)	

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Nausea	26 (10)	28 (18)	54 (13)	2.10 (1.10-3.90; p<0.05)
Nausea frequency				
Rarely	9 (3)	6 (4)	15 (4)	0.14 (0.05-0.24; p<0.05)
Frequently	11 (4)	17 (11)	28 (7)	
Constantly	3 (1)	3 (2)	6 (1)	
Nausea severity				
Mild	6 (2)	6 (4)	12 (3)	0.13 (0.04-0.23; p=0.054)
Moderate	11 (4)	15 (10)	26 (6)	
Severe	6 (2)	5 (3)	11 (3)	
Vomiting	8 (3)	7 (5)	15 (3)	1.60 (0.48-5.10; p=0.54)
Vomiting frequency				
Rarely	7 (3)	5 (3)	12 (3)	0.05 (0-0.15; p=0.49)
Frequently	1 (<1)	2 (1)	3 (<1)	
Constantly	0	0	0	
Vomiting severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.06 (0-0.16; p=0.66)
Moderate	4 (2)	2 (1)	6 (1)	
Severe	1 (<1)	2 (1)	3 (<1)	
Diarrhea	14 (5)	17 (11)	31 (7)	2.30 (1.00-5.20; p<0.05)
Diarrhea frequency				
Rarely	3 (1)	3 (2)	6 (1)	0.10 (0.006-0.19; p=0.22)
Frequently	7 (3)	9 (6)	16 (4)	
Constantly	4 (2)	4 (3)	8 (2)	
Diarrhea severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.11 (0.01-0.21; p=0.14)
Moderate	6 (2)	6 (4)	12 (3)	
Severe	5 (2)	8 (5)	13 (3)	
Decreased appetite	28 (10)	36 (23)	64 (15)	2.70 (1.50-4.80; p<0.001)
Decreased appetite frequency				
Rarely	1 (<1)	8 (5)	9 (2)	0.21 (0.12-0.30; p<0.001)
Frequently	17 (7)	15 (10)	32 (8)	
Constantly	7 (3)	11 (7)	18 (4)	
Decreased appetite severity				
Mild	4 (2)	9 (6)	13 (3)	
Moderate	13 (5)	14 (9)	27 (6)	
Severe	9 (3)	10 (7)	19 (4)	
Increased appetite	59 (21)	43 (28)	102 (24)	1.40 (0.87-2.30; p=0.16)
Increased appetite frequency				
Rarely	3 (1)	6 (4)	9 (2)	0.10 (0.009-0.20; p=0.20)
Frequently	38 (14)	25 (16)	63 (15)	
Constantly	16 (6)	10 (7)	26 (6)	

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Increased appetite severity				
Mild	14 (5)	9 (6)	23 (5)	0.10 (0.007-0.19; p=0.22)
Moderate	39 (14)	24 (16)	63 (15)	
Severe	6 (2)	9 (6)	15 (4)	
Sweet cravings	58 (21)	39 (25)	97 (23)	1.30 (0.77-2.10; p=0.38)
Sweet cravings frequency				
Rarely	8 (3)	4 (3)	12 (3)	0.09 (0.001-0.19; p=0.27)
Frequently	36 (13)	18 (12)	54 (13)	
Constantly	12 (4)	14 (9)	26 (6)	
Sweet cravings severity				
Mild	19 (7)	7 (5)	26 (6)	0.11 (0.01-0.20; p=0.17)
Moderate	31 (11)	19 (12)	50 (12)	
Severe	7 (3)	10 (7)	17 (4)	
Bad taste	24 (9)	22 (14)	43 (11)	1.70 (0.89-3.40; p=0.10)
Bad taste frequency				
Rarely	8 (3)	4 (3)	12 (3)	0.11 (0.01-0.20; p=0.15)
Frequently	11 (4)	10 (7)	21 (5)	
Constantly	4 (2)	7 (5)	11 (3)	
Bad taste severity				
Mild	9 (3)	4 (3)	13 (3)	0.12 (0.03-0.22; p=0.08)
Moderate	11 (4)	7 (5)	18 (4)	
Severe	4 (2)	9 (6)	13 (3)	
Thirst	76 (28)	59 (38)	135 (31)	1.60 (1.00-2.50; p<0.05)
Thirst frequency				
Rarely	7 (3)	5 (3)	12 (3)	0.13 (0.03-0.22; p=0.059)
Frequently	49 (18)	27 (18)	76 (18)	
Constantly	18 (7)	22 (14)	40 (9)	
Thirst severity				
Mild	22 (8)	14 (9)	36 (8)	0.17 (0.08-0.26; p<0.01)
Moderate	48 (17)	26 (17)	74 (17)	
Severe	5 (2)	14 (9)	19 (4)	
Dry mouth	97 (35)	66 (43)	163 (38)	1.40 (0.90-2.10; p=0.15)
Dry mouth frequency				
Rarely	14 (5)	8 (5)	22 (5)	0.17 (0.07-0.26; p<0.01)
Frequently	62 (23)	28 (18)	90 (21)	
Constantly	18 (7)	26 (17)	44 (10)	
Dry mouth severity				
Mild	37 (13)	14 (9)	51 (12)	0.22 (0.13-0.31; p<0.001)
Moderate	53 (19)	27 (18)	80 (19)	
Severe	7 (3)	21 (14)	28 (7)	
Musculoskeletal				

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
<u>Joint pain</u>	37 (13)	39 (25)	76 (18)	2.20 (1.30-3.70; p<0.005)
Joint pain frequency				
Rarely	3 (1)	3 (2)	6 (1)	0.14 (0.04-0.23; p<0.05)
Frequently	16 (6)	15 (10)	31 (7)	
Constantly	16 (6)	18 (12)	34 (8)	
Joint pain severity				
Mild	2 (<1)	2 (1)	4 (<1)	0.15 (0.05-0.24; p<0.05)
Moderate	17 (6)	16 (10)	33 (8)	
Severe	16 (6)	19 (12)	35 (8)	
<u>Weak limbs</u>	29 (11)	29 (19)	58 (14)	2.00 (1.10-3.60; p<0.05)
Weak limbs frequency				
Rarely	3 (1)	6 (4)	9 (2)	0.14 (0.04-0.23; p<0.05)
Frequently	14 (5)	9 (6)	23 (5)	
Constantly	11 (4)	13 (8)	24 (6)	
Weak limbs severity				
Mild	6 (2)	5 (3)	11 (3)	0.12 (0.02-0.21; p=0.09)
Moderate	10 (4)	11 (7)	21 (5)	
Severe	11 (4)	12 (8)	23 (5)	
<u>Tremor</u>	16 (6)	9 (6)	25 (6)	1.00 (0.38-2.50; p=1.00)
Tremor frequency				
Rarely	4 (2)	7 (5)	11 (7)	0.14 (0.04-0.23; p<0.05)
Frequently	7 (3)	1 (<1)	8 (2)	
Constantly	5 (2)	0	5 (1)	
Tremor severity				
Mild	6 (2)	4 (3)	10 (2)	0.07 (0-0.17; p=0.50)
Moderate	6 (2)	4 (3)	10 (2)	
Severe	4 (2)	0	4 (<1)	
<u>Spasms</u>	21 (8)	20 (13)	41 (10)	1.80 (0.89-3.60; p=0.10)
Spasms frequency				
Rarely	2 (<1)	5 (3)	7 (2)	0.12 (0.02-0.21; p=0.11)
Frequently	12 (4)	11 (7)	23 (5)	
Constantly	5 (2)	4 (3)	9 (2)	
Spasms severity				
Mild	0	2 (1)	2 (<1)	0.12 (0.03-0.22; p=0.08)
Moderate	11 (4)	11 (7)	22 (5)	
Severe	8 (3)	7 (5)	15 (4)	
Cardiovascular				
<u>Palpitations</u>	21 (8)	21 (14)	42 (10)	1.90 (0.95-3.80; p=0.06)
Palpitations frequency				
Rarely	10 (4)	11 (7)	21 (5)	0.09 (0-0.18; p=0.31)

Measure	Males (N = 275, 64%)	Females (N = 154, 36%)	Total (N = 429)	Effect size (CI; P value)
	Number of patients (%)			
Frequently	9 (3)	7 (5)	16 (4)	
Constantly	2 (<1)	2 (1)	4 (<1)	
Palpitations severity				
Mild	10 (4)	9 (6)	19 (4)	0.12 (0.02-0.21; p=0.13)
Moderate	9 (3)	6 (4)	15 (4)	
Severe	1 (<1)	4 (3)	5 (1)	
Visual				
<u>Blurred vision</u>	14 (5)	19 (12)	33 (8)	2.60 (1.20-5.80; p<0.05)
Blurred vision frequency				
Rarely	7 (3)	4 (3)	11 (3)	0.16 (0.07-0.25; p<0.01)
Frequently	6 (2)	12 (8)	18 (4)	
Constantly	0	2 (1)	2 (<1)	
Blurred vision severity				
Mild	5 (2)	3 (2)	8 (2)	0.16 (0.06-0.25; p<0.05)
Moderate	7 (3)	14 (9)	21 (5)	
Severe	0	1 (<1)	1 (<1)	
<u>Red eyes</u>	59 (21)	20 (13)	79 (18)	0.55 (0.30-0.97; p<0.05)
Red eyes frequency				
Rarely	9 (3)	4 (3)	13 (3)	0.10 (0.01-0.20; p=0.20)
Frequently	34 (12)	12 (8)	46 (11)	
Constantly	15 (6)	4 (3)	19 (4)	
Red eyes severity				
Mild	26 (10)	8 (5)	34 (8)	0.11 (0.01-0.20; p=0.15)
Moderate	26 (10)	8 (5)	34 (8)	
Severe	6 (2)	4 (3)	10 (2)	
<u>Dry eyes</u>	35 (13)	34 (22)	69 (16)	1.90 (1.10-3.40; p<0.05)
Dry eyes frequency				
Rarely	10 (4)	8 (5)	18 (4)	0.11 (0.01-0.20; p=0.18)
Frequently	19 (7)	15 (10)	34 (8)	
Constantly	6 (2)	8 (5)	14 (3)	
Dry eyes severity				
Mild	14 (5)	13 (8)	27 (6)	0.12 (0.02-0.21; p=0.09)
Moderate	18 (8)	12 (8)	30 (7)	
Severe	3 (1)	6 (4)	10 (2)	
Other				
Headaches	1 (<1)	1(<1)	2 (<1)	NA
Fatigue	0	1 (<1)	1 (<1)	NA
Migraine	0	1 (<1)	1 (<1)	NA
Weakness	0	1 (<1)	1 (<1)	NA
Hypoglycemia	1 (<1)	0	1 (<1)	NA
Enhanced vision	1 (<1)	0	1 (<1)	NA
Tranquility	1 (<1)	0	1 (<1)	NA

IQR, Inter quartile range; OR, Odds ratio; CI, Confidence interval; NA, Non-applicable; percentages under 1 are displayed as <1; Percentages are rounded without decimal point.