



Supplementary Material

Article Title: Is the Noradrenergic Symptom Cluster a Valid Construct in Adjunctive Treatment of Major Depressive Disorder?

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Supplementary Material

Supplementary eTable 1. Patient's Psychiatric History

Study	Tx	Age at MDD Onset, years, mean (SD)	Number of Previous MDD Episodes, mean (SD)	Duration of Current MDD Episode, weeks, mean (SD)
LNDK	EDX (N=111)	32 (13)	6 (9)	41 (40)
	PLB (N=116)	27 (13)	6 (7)	38 (50)
LNBM	EDX 12 mg (N=231)	36 (13)	3 (4)	47 (60)
	EDX 18 mg (N=230)	35 (13)	3 (4)	49 (74)
	PLB (N=240)	33 (12)	3 (7)	48 (64)
LNBQ	EDX 12-18 mg (N=232)	36 (14)	3 (7)	61 (146)
	EDX 6 mg (N=226)	35 (14)	3 (5)	58 (118)
	PLB (N=231)	35 (14)	3 (6)	54 (119)
LNBR	EDX (N=228)	34 (14)	5 (9)	72 (118)
	PLB (N=219)	34 (14)	6 (11)	87 (161)

Abbreviations: EDX = edivoxetine; MDD = major depressive disorder; N = number of patients; PLB = placebo; SD = standard deviation; Tx = treatment.

Supplementary eTable 2. SSRI Treatments

Study	Tx ^a	Sertraline		Paroxetine		Escitalopram		Fluoxetine		Citalopram		Fluvoxamine	
		n (%)	mean dose (SD)	n (%)	mean dose (SD)	n (%)	mean dose (SD)	n (%)	mean dose (SD)	n (%)	mean dose (SD)	n (%)	mean dose (SD)
LNDK	EDX	30 (27)	108 (40)	12 (11)	28 (13)	24 (22)	16 (6)	19 (17)	34 (13)	26 (23)	25 (9)	0	-
	PLB	21 (18)	100 (42)	18 (16)	28 (12)	21 (18)	18 (7)	23 (20)	32 (15)	32 (28)	31 (11)	1 (0.9)	200
LNBM	EDX 12 mg	70 (30)	106 (50)	34 (15)	29 (10)	40 (17)	13 (5)	39 (17)	39 (16)	39 (17)	26 (10)	9 (4)	103 (40)
	EDX 18 mg	66 (29)	93 (49)	41 (18)	29 (9)	40 (17)	15 (5)	40 (17)	35 (10)	31 (13)	27 (12)	12 (5)	108 (34)
	PLB	72 (30)	90 (47)	37 (15)	28 (9)	52 (22)	15 (5)	34 (14)	35 (17)	37 (15)	26 (13)	8 (3)	78 (36)
LNBQ	EDX 12-18 mg	61 (26)	95 (45)	18 (8)	27 (11)	59 (25)	14 (5)	28 (12)	32 (13)	58 (25)	26 (10)	8 (3)	122 (41)
	EDX 6 mg	73 (32)	97 (41)	18 (8)	29 (12)	63 (28)	13 (4)	21 (9)	36 (16)	43 (19)	27 (10)	8 (4)	81 (37)
	PLB	75 (32)	97 (47)	23 (10)	26 (9)	54 (23)	14 (4)	29 (13)	30 (17)	41 (18)	25 (10)	9 (4)	111 (33)
LNBR	EDX	60 (26)	98 (41)	17 (7)	23 (6)	36 (16)	16 (5)	45 (20)	30 (12)	71 (31)	25 (9)	1 (0.4)	300
	PLB	37 (17)	109 (48)	12 (5)	28 (10)	32 (15)	17 (7)	48 (22)	27 (10)	90 (41)	27 (11)	0	-

Abbreviations: EDX = edivoxetine; n = number of patients; PLB = placebo; SD = standard deviation; SSRI = selective serotonin reuptake inhibitor; Tx = treatment.

^aall treatment doses are presented in mg