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TREATING MAJOR DEPRESSION IN THE SETTING OF OUD

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SPEAKER DISCLOSURES

- ✓ Any conflicts of interest?
 - ✓ none

OBJECTIVES



Explore the relationship between mood sx and OUD.

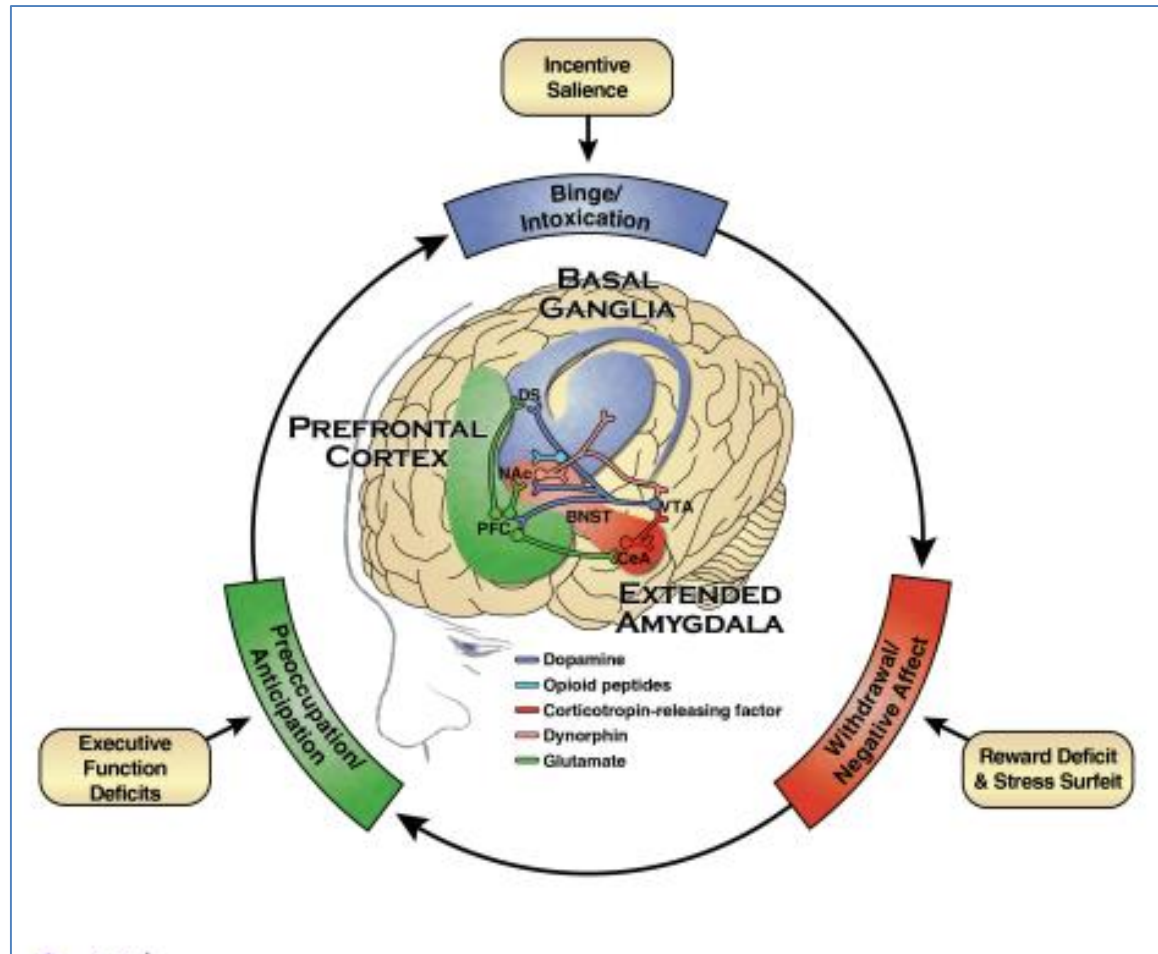


Review 6 papers examining tx MDD in OUD.



Summarize/Discuss

BACKGROUND



LITERATURE

All FDA treatments for depression excludes substance users

Studies examine:

- All Substance Use (alcohol, cocaine, opioids, nicotine) vs Any psychiatric dx;
- Specific SUD vs Any psychiatric dx;
- Specific SUD vs Specific Psychiatric dx (Mood, anxiety, PTSD, BiPD, SCZ);
- All vary by population demographics – gender, age, pregnancy status, SES, treatment type, diagnostic methods, SUD tx phase, SUD use pattern.
- Varied outcome of interest – Psychiatric sx, SU outcome

SIGNIFICANCE

Pathways between nonmedical opioid use/dependence and psychiatric disorders:
Results from the National Epidemiologic Survey on Alcohol and Related
Conditions

Silvia S. Martins^{a,*}, Katherine M. Keyes^{b,c}, Carla L. Storr^{a,d}, Hong Zhu^e, Howard D. Chilcoat^{a,f}

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- Bidirectional relationship
 - OUD increases risk of developing MDD
 - HR 4.6 (2.8, 7.6)
 - MDD increases risk of developing OUD:
 - HR 5.2(3.2, 8.2)

LITERATURE

Efficacy of antidepressants in substance use disorders with and without comorbid depression A systematic review and meta-analysis

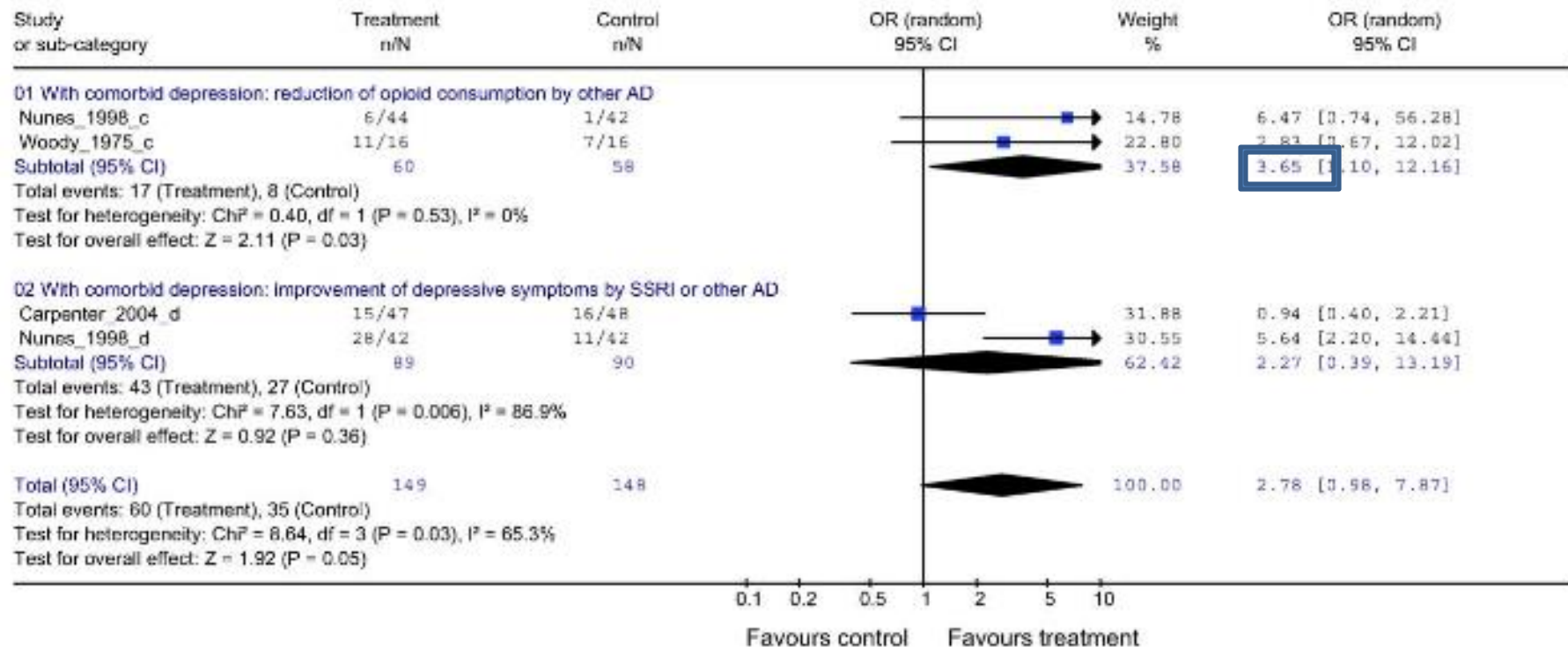
Marta Torrens^{a,*}, Francina Fonseca^a, Gerard Mateu^a, Magí Farré^b

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Review: Effect of antidepressant medication in drug dependence (Version 02 (07_09_04))
Comparison: 04 Opioids
Outcome: 01 Opioid dependence



LITERATURE

Treatment of Depression in Patients with Opiate Dependence

Edward V. Nunes, Maria A. Sullivan, and Frances R. Levin

Table 1. Antidepressant trials in Methadone-Maintained, Opiate-Dependent Patients with Depressive Disorders or Symptoms: Methodology and Mood Outcome

Author, Year (Sample Size)	Method of Selecting Depression at Baseline	Medication Trial (Dose mg/day/ Duration Weeks)	Outcome Measures ^a	Outcome on Active Medication ^a			Outcome on Placebo ^a			Composite Effect Size ^b Cohen's d (95% CI)
				Baseline	End Study	% Reduction	Baseline	End Study	% Reduction	
Woody et al 1975 (N = 35)	Depressed by psychiatrist's judgement ^c	Doxepin (100-150/16)	HamD +: BDI**:	18.3 9.1	7.1 3.6	61 60	18.3 9.1	15.1 10.3	17 -13	.93 (.04 to 1.83)
Titievsky et al 1982 (N = 76)	HamD > 17 ^c	Doxepin (up to 100/4)	HamD*: POMS +:	27.5 103.6	19.5 89.5	29 14	27.0 111.0	22.6 105.6	16 5	.56 (-.05 to 1.17)
Kleber et al 1983 (N = 46)	Evaluation by psychiatrist, DSM- III MDD	Imipramine (150-225/8)	HamD: BDI:	20.1 15.1	10.1 10.2	50 32	19.5 13.3	11.2 10.4	43 22	.13 (-.48 to .74)
Ziedonis and Kosten 1991 (N = 15)	SCID for DSM-III-R MDD or DD (subgroup of cocaine dependent sample) ^d	Desipramine (150/12)	BDI:	9.6	9.0	7	8.5	17.3	-100	.0 (-1.14 to 1.14)
Arndt et al 1992 (N = 79)	None, depression outcome reported ^c	Desipramine (250-300/12)	BDI: ASI Psych Comp*:	15 .21	7 .12	53 43	13 .15	9 .20	31 -33	.27 (-.27 to .81)
Margolin et al 1995 (N = 149)	None, depression outcome reported; subgroup (N = 36) HamD > 12 also reported ^c	Bupropion (200-300/12)	HamD: ASI Psych Comp:	7.8 .09	4.6 .09	41 0	8.2 .10	6.2 .12	24 -20	.22 (-.14 to .57)
Nunes et al 1998 (N = 137)	SCID for DSM-III-R MDD, DD, or NOS (evaluation by psychiatrist) ^d	Imipramine (up to 300/12)	HamD***: %mood response*:	16.2	10.0 42	38	15.6	14.4 21	8	.68 (.33 to 1.00)
Petrakis et al 1998 (N = 144)	Evaluation by psychiatrist DSM- III MDD, DD, or NOS ^d	Fluoxetine (up to 60/12)	HamD: BDI:	14.0 17.6	8.0 9.6	43 45	14.9 12.6	7.2 7.9	52 37	-.20 (-.81 to .42)
Dean et al 2002 (N = 49)	BDI > 21	Fluoxetine (20/12)	MADRS:	28.6	19.9	30	27.8	17.8	36	-.18 (-.75 to .40)
Carpenter et al 2004 (N = 95)	SCID for DSM-III-R MDD or DD (evaluation by psychiatrist) ^d	Sertraline (up to 200/12)	HamD: %mood response:	21.1	14.5 32	31	21.1	14.9 33	29	.07 (-.24 to .48)
Pooled Estimate of Effect ^f										.26 (.04 to .47)

LITERATURE

The effect of sertraline and environmental context on treating depression and illicit substance use among methadone maintained opiate dependent patients: a controlled clinical trial

Kenneth M. Carpenter*, Adam C. Brooks, Suzanne K. Vosburg, Edward V. Nunes

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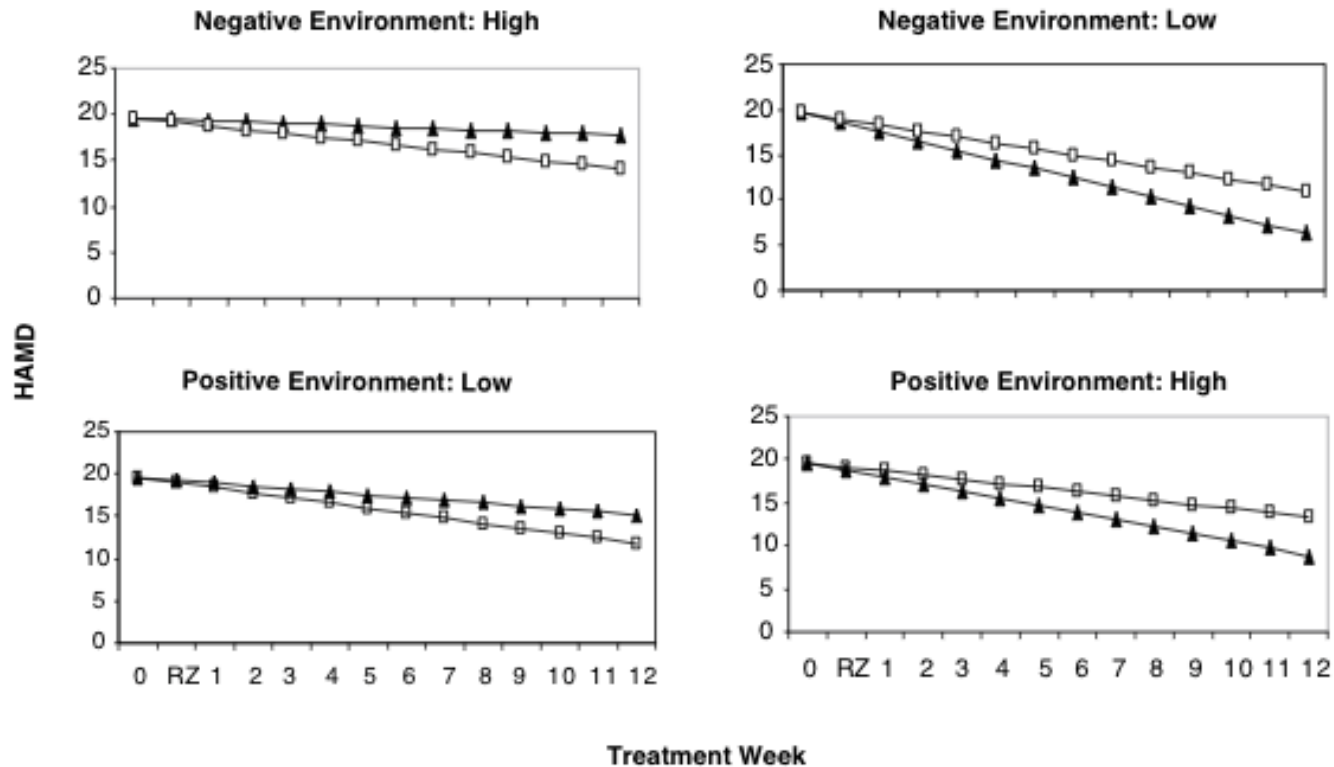


Fig. 1. Depression Scores (HAMD) by treatment group (sertraline (▲); placebo (□)) under high and low positive and negative environment conditions.

LITERATURE

Depression history as a predictor of outcomes during buprenorphine-naloxone treatment of prescription opioid use disorder

Andrew D. Peckham*, Margaret L. Griffin, R. Kathryn McHugh, Roger D. Weiss

Department of Psychiatry, McLean Hospital and Harvard Medical School, Belmont, MA, USA

Table 1

Models to predict successful opioid outcomes during a 12-week treatment study.

	Odds ratios, adjusted		
	Model 1	Model 2	Model 3
Predictors of good opioid outcomes	(N = 360)	(N = 317)	(N = 317)
Gender	0.93	0.82	0.97
Treatment condition ^a	0.78	0.71	0.76
Heroin ever	0.56*	0.57*	0.53*
Major depression lifetime	1.82**	1.63 [†]	1.81*
Depression score at baseline		1.01	
Depression score at week 4 of treatment		0.98	
Abstinence goal			1.39
Mutual-help groups during treatment			1.67*

Note. All variables were assessed at baseline (N = 360), with the exception of week 4 depression symptoms and mutual-help group attendance (N = 317).

[†] $p < .055$.

* $p < .05$.

** $p < .01$.

^a Standard medical management alone or combined with opioid counseling.

TREATMENT GUIDELINES

Treatment for Substance Use Disorder With Co-Occurring Mental Illness

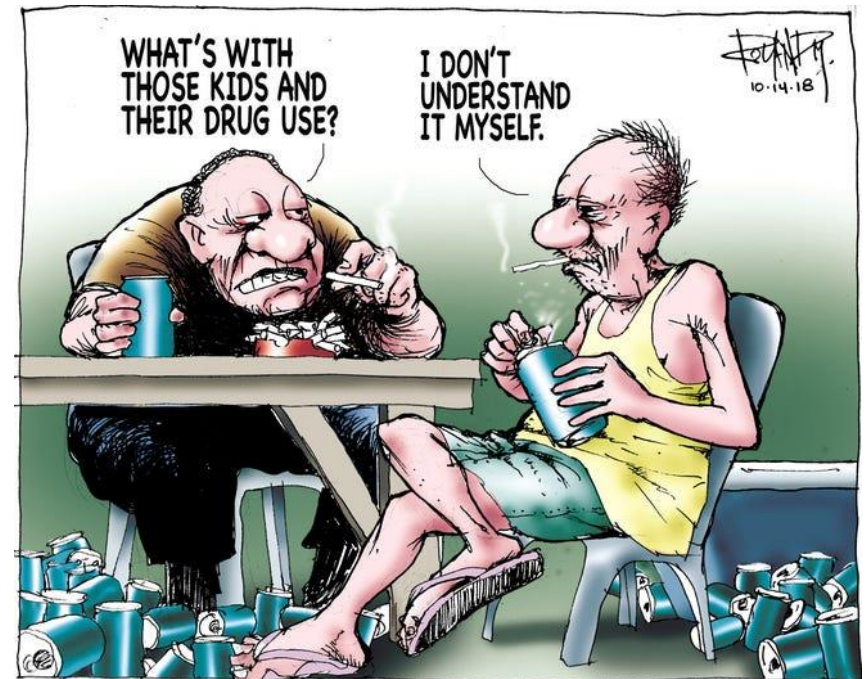
Muhammad N. Iqbal, M.D., Charles J. Levin, B.A., Frances R. Levin, M.D.

1. SSRI's are first line given safety profile for those with OUD and MDD;
2. TCA's have more data, but carry SE risk; cardiac, QTc, hypotension.
3. All should include Behavioral interventions; 12-step, MI, CBT.



SUMMARY

- Martin et al - Epi study demonstrate 5x risk of MDD in OUD and 5x risk of OUD among those with MDD;
- Torrens et al - meta-analysis (2004); very small effect of antidepressants on opioid use;
- Nunes et al - meta-analysis (2010); very small effect of antidepressant on depressive sx, Cohen's D – 0.26.
- Carpenter et al - RTC (2004); sertraline and social context have small effect on mood sx.
- Peckman et al- secondary analysis demonstrated self-help engagement as a predictor of improved Opioid use outcomes
- Take home: tx MDD with SSRI's, despite limited data.
- More data for TCA's, but SSRI's safer;
- **Context matters – more engagement improves all outcomes.**



BIBLIOGRAPHY

- Martins et al, 2009
- Torrens et al, 2004
- Nunes et al, 2010
- Carpenter et al, 2004
- Nunes et al, 2004
- Peckman et al, 2020
- Pettinati et al 2013
- Iqbal et al, 2019

THANK YOU!



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