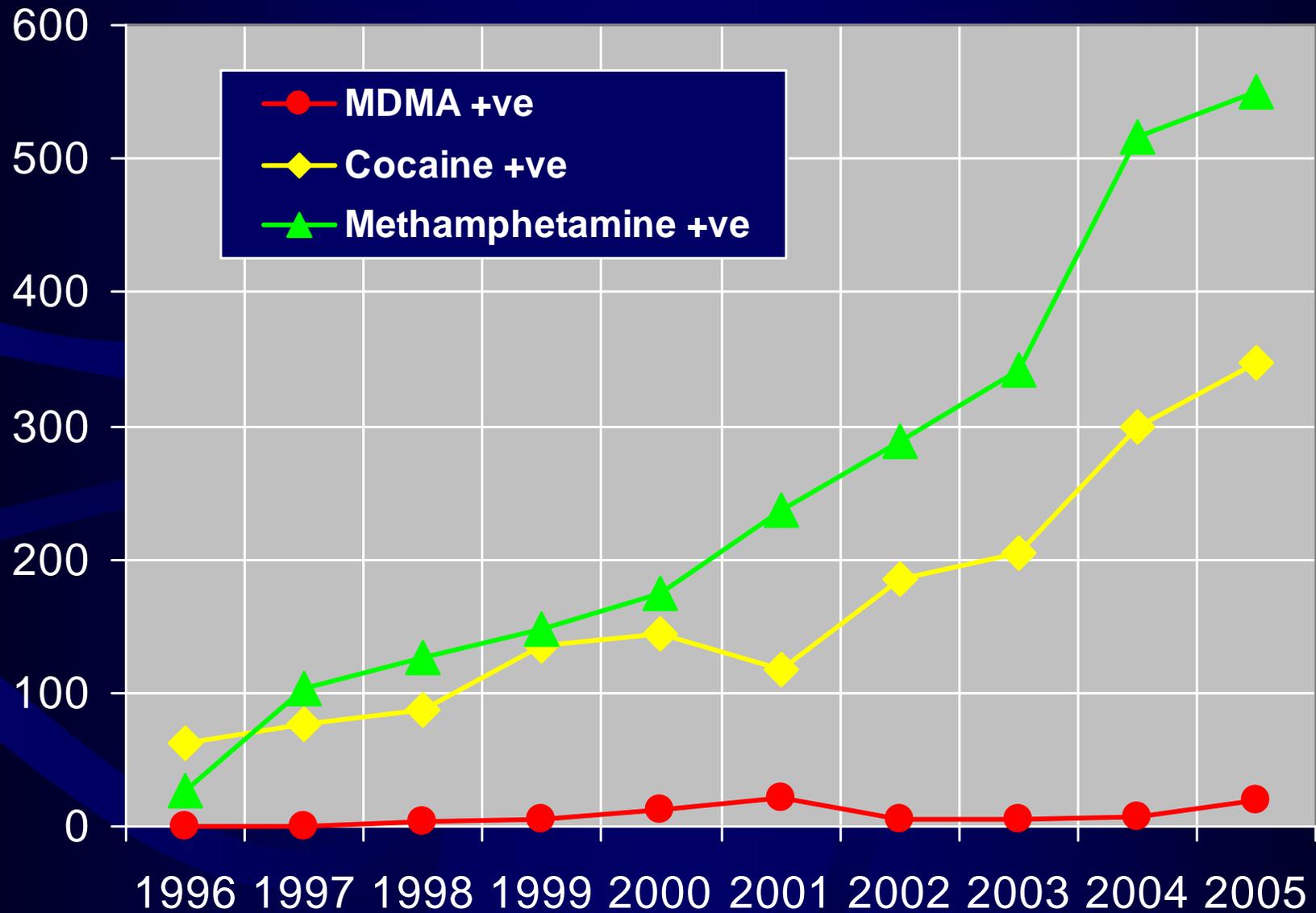


The Downside of Methamphetamine

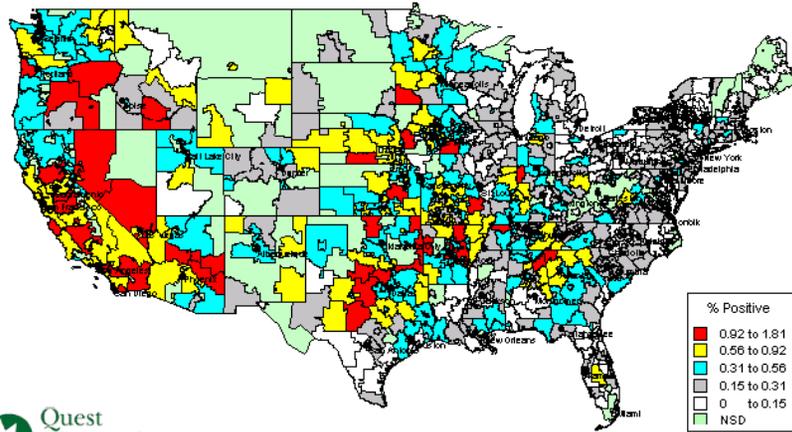
Barry K. Logan Ph.D., DABFT
Washington State Toxicology Laboratory,
Forensic Laboratory Services Bureau,
Washington State Patrol, Seattle, Washington

DUID Stimulants (WA)



Amphetamine Positivity by 3-Digit Zipcode

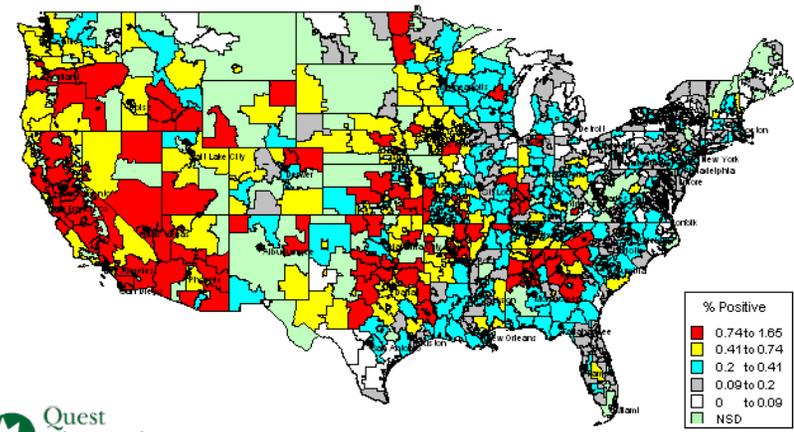
January - December 2002



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Amphetamine Positivity by 3-Digit Zipcode

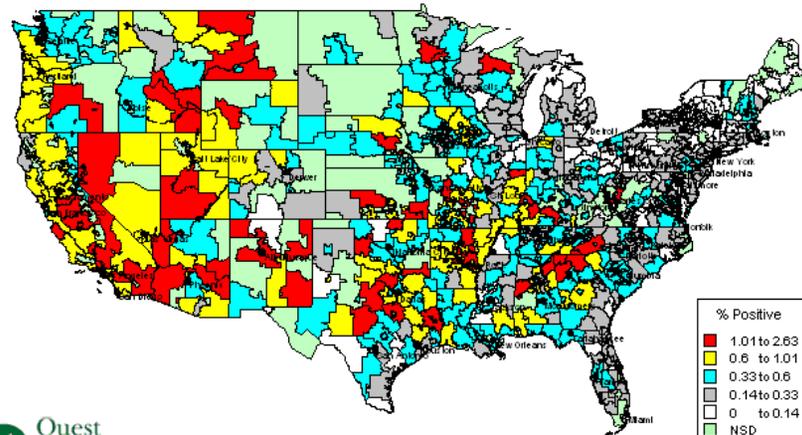
January - December 2003



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Amphetamines Positivity by 3-Digit Zipcode

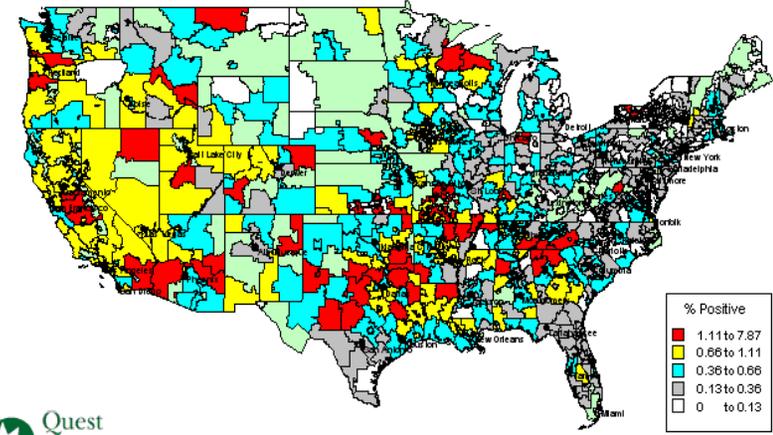
January - December 2004



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Amphetamines Positivity by 3-Digit Zipcode

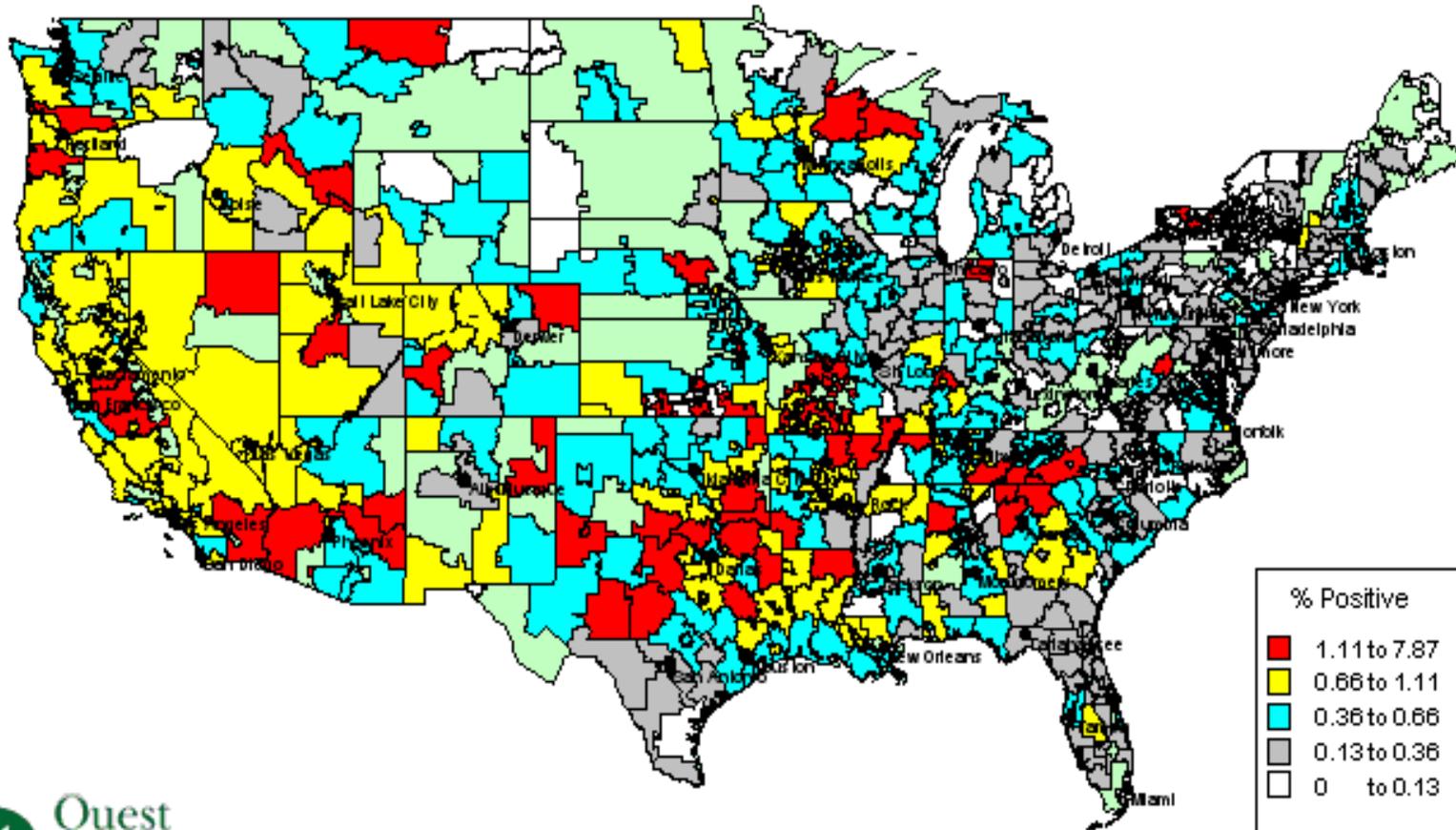
January - June 2005



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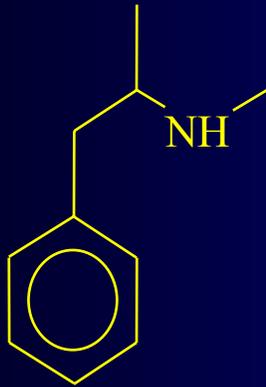
Amphetamines Positivity by 3-Digit Zipcode

January - June 2005



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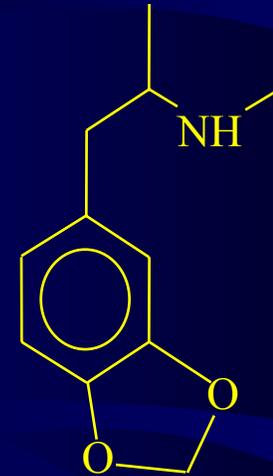
Chemistry



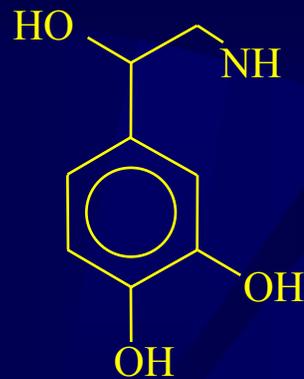
Methamphetamine



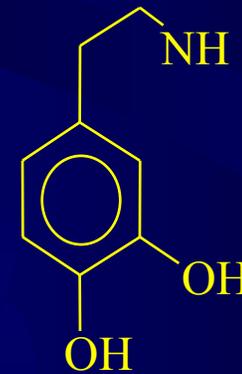
Amphetamine



3,4 MDMA



Norepinephrine



Dopamine

Pharmacology

Peripheral effects

α and β adrenergic agonist effects

pupillary dilation

bronchial muscle dilation

vasoconstriction

coronary dilatation

bladder contraction

increased heart rate/blood pressure

effects on sexual function

Pharmacology - Amphetamines

CNS - Sympathomimetic

promotes synthesis and release of:

Norepinephrine

alerting, anorectic, locomotor effects

Dopamine

locomotor stimulating effects

psychosis, disturbances in perception

5-HT

delusions, psychosis

Methamphetamine

Advances in Pharmacology

Administration of methamphetamine causes long term changes in dopaminergic systems, including decreases in transporter numbers, dopamine concentrations and tyrosine hydroxylase activity.

Dopamine transporter activity is rapidly and reversibly decreased after single use.

Dopamine transporter activity recovers after 24 hours in chronic use, but declines again 8 days post use.

Woolverton et al. Brain Res. 1989;486:73-78

Fleckenstein et al. J Pharm Exp Ther 1997;282(2):834-838

Methamphetamine

Advances in Pharmacology

Methamphetamine concentrations in the brain are eight-fold higher than in the serum during the first hour following IV administration.

In tolerant subjects, serum levels are higher, while brain levels are lower.

Riviere et al. J Pharm Exp Ther 2000;292(3):1042-1047

Gygi et al Neuropharmacology. 1996 Jun;35(6):751-757

Patterns of Use

Clinical use (Desoxyn®):

5 - 60mg (q.i.d.) orally – narcolepsy

2.5 – 40mg (Ext. Rel.) - ADHD

Abuse:

Occasional users – ± 60 mg

Heavy users 250 - >5000 mg/day,

Orally, IV, IN, IM, smoked



Methamphetamine



Crank, Crystal, Speed, Meth

Potent CNS stimulant

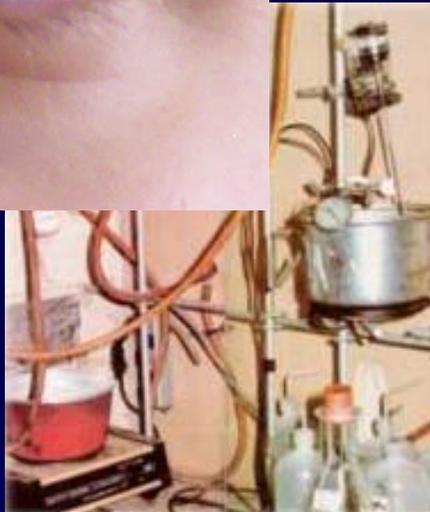
Euphoria

Excitation

Alertness

Agitation

Motor restlessness



Pharmacodynamics

The methamphetamine binge

Characterized by high dose, often IV use

Little or no sleep, no appetite

Use is compulsive and uncontrolled

Repeated administration at 1 - 5 hourly intervals

Binge can persist for days or weeks

Methamphetamine Intoxication

The Rush:

5 minutes intense euphoria

“Orgasmic” pleasure

Rapid flight of ideas

Sexual stimulation

High energy

Obsessive/compulsive activity

Thought blending

Word salad

Dilated pupils

Methamphetamine Intoxication

The Shoulder:

Less intense euphoria

Hyperactivity

Rapid flight of ideas

Obsessive/compulsive activity

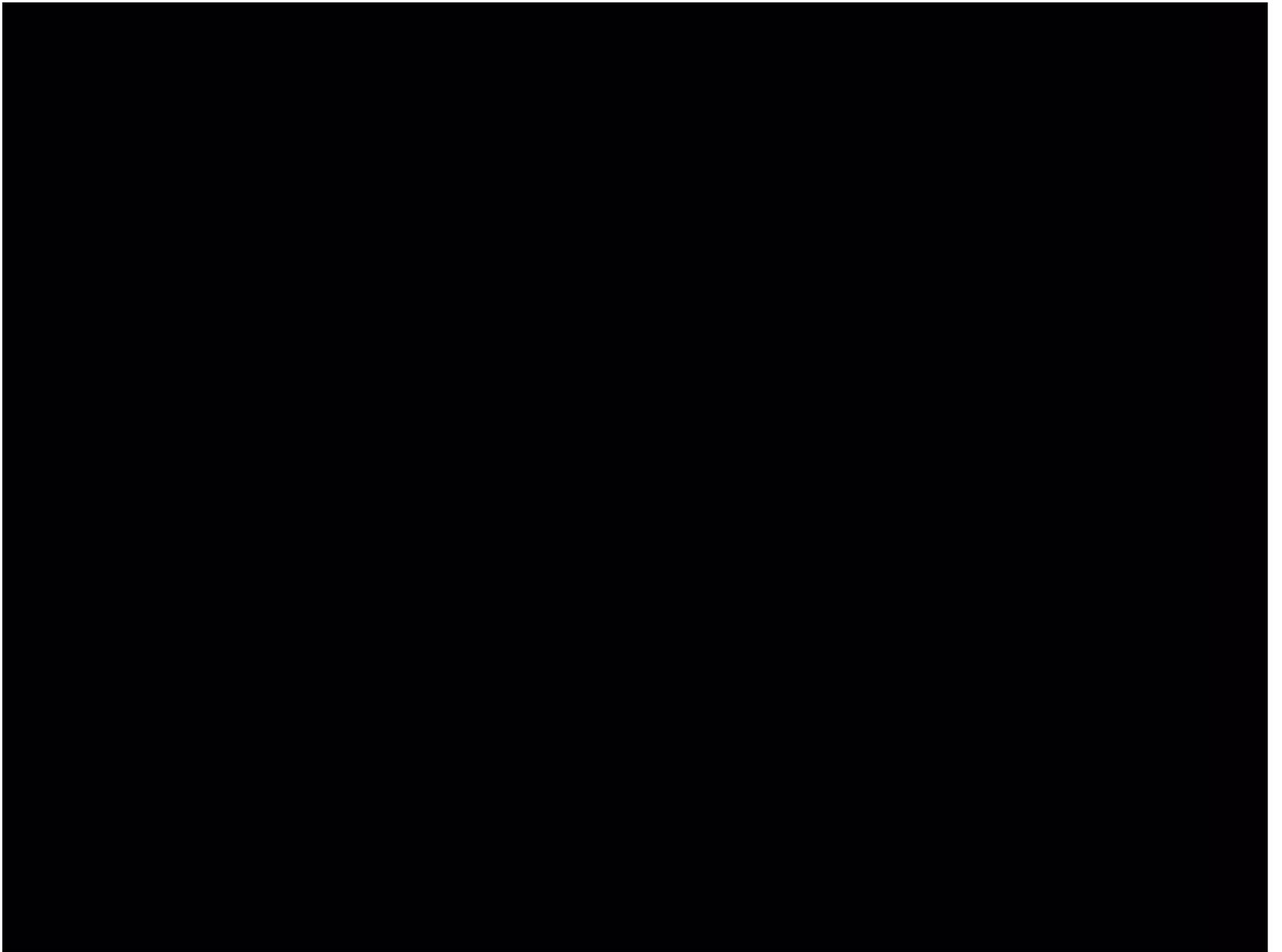
Thought blending

Word salad

Dilated pupils

Shift from seeking of High

to avoidance of Low



Methamphetamine Intoxication

“Tweaking”:

Dysphoria

Scattered, disorganized thought

Intense craving

Paranoia/Anxiety/Irritability

Hypervigilance

Auditory, tactile hallucinations

Delusions

Pupils normal

Methamphetamine Intoxication

Dealing with Tweakers

Keep your distance

Stay within central field of vision

No bright lights

Keep your hands in plain sight

Engage them in reassuring dialogue

Talk slowly with deep pitch

Remind them that it's the drug talking

Care with restraints

Methamphetamine



Crank, Crystal, Speed, Meth

Meth withdrawal

Fatigue

Sleepiness

Irritability

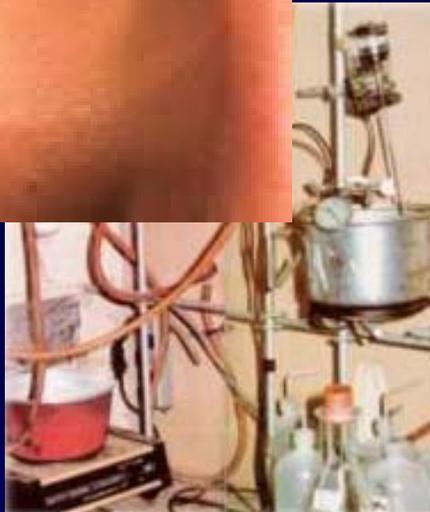
Drug cravings

Anxiety

Depression

Paranoia

Delusions



Methamphetamine Intoxication

The Crash:

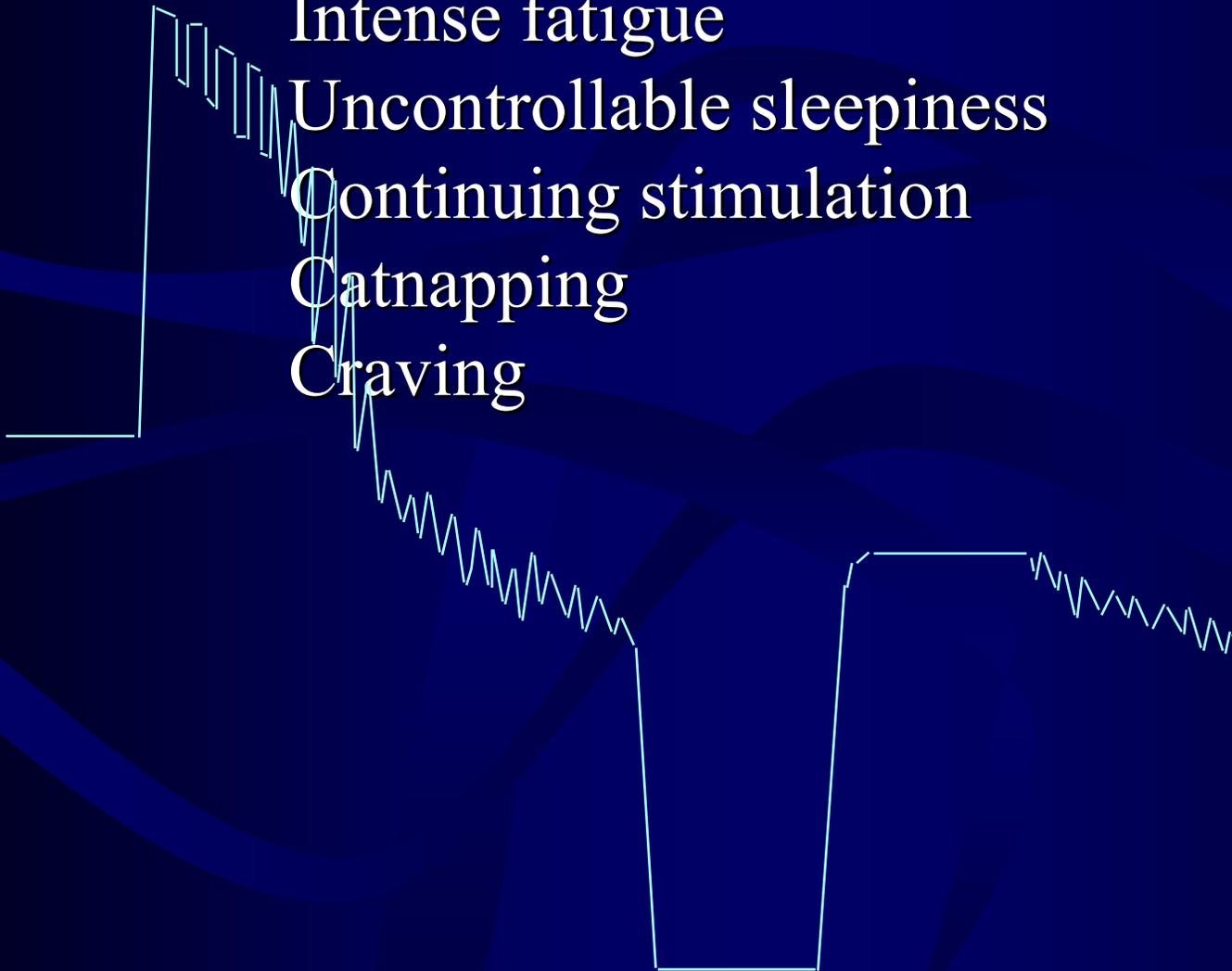
Intense fatigue

Uncontrollable sleepiness

Continuing stimulation

Catnapping

Craving



Methamphetamine Intoxication

Withdrawal:

Anergia

Anhedonia

Waves of intense craving

Environmentally cued

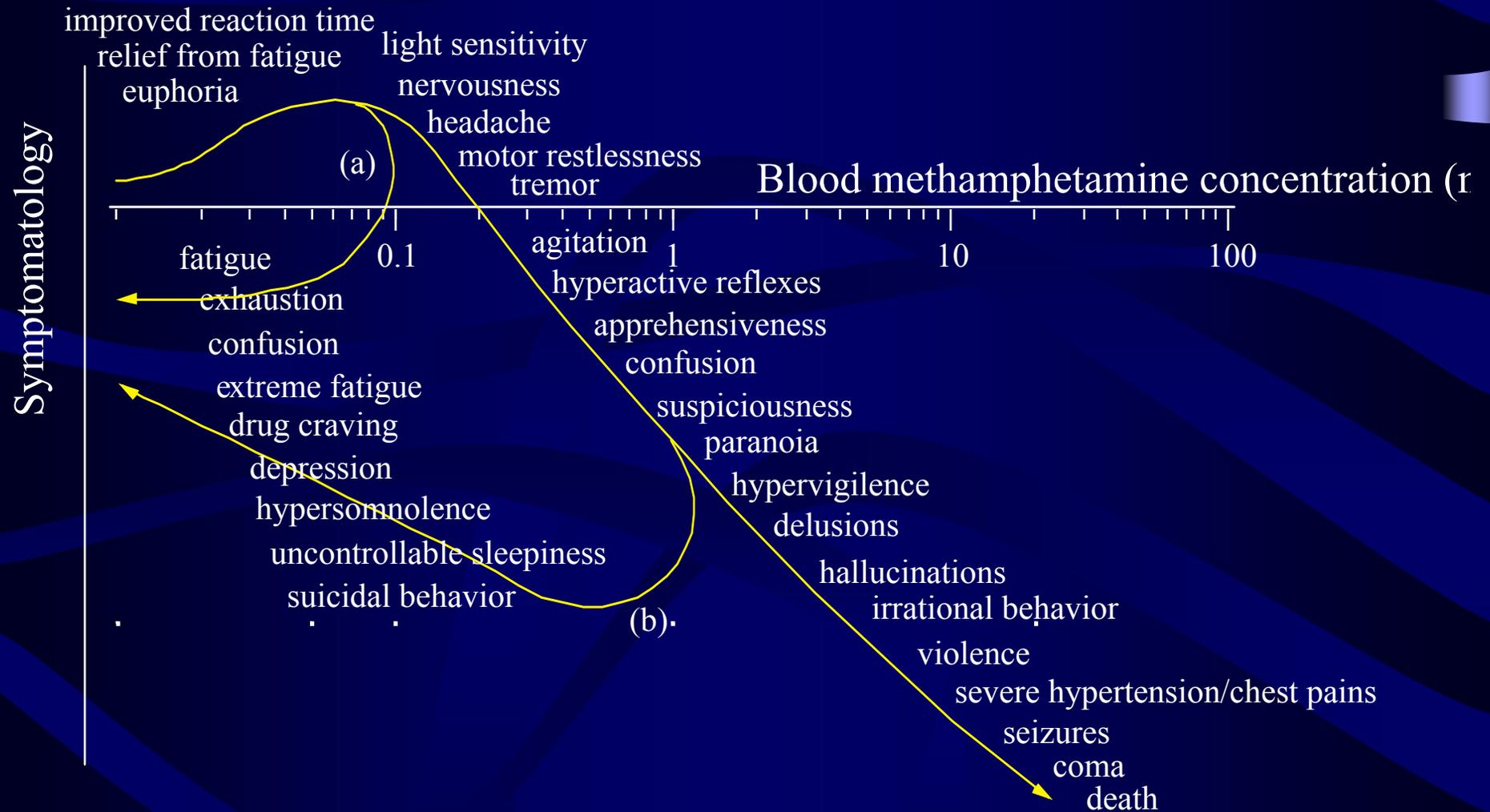
Endogenous

Stress

Inadequately treated withdrawal

Inadequately treated mental illness

Methamphetamine Hysteresis



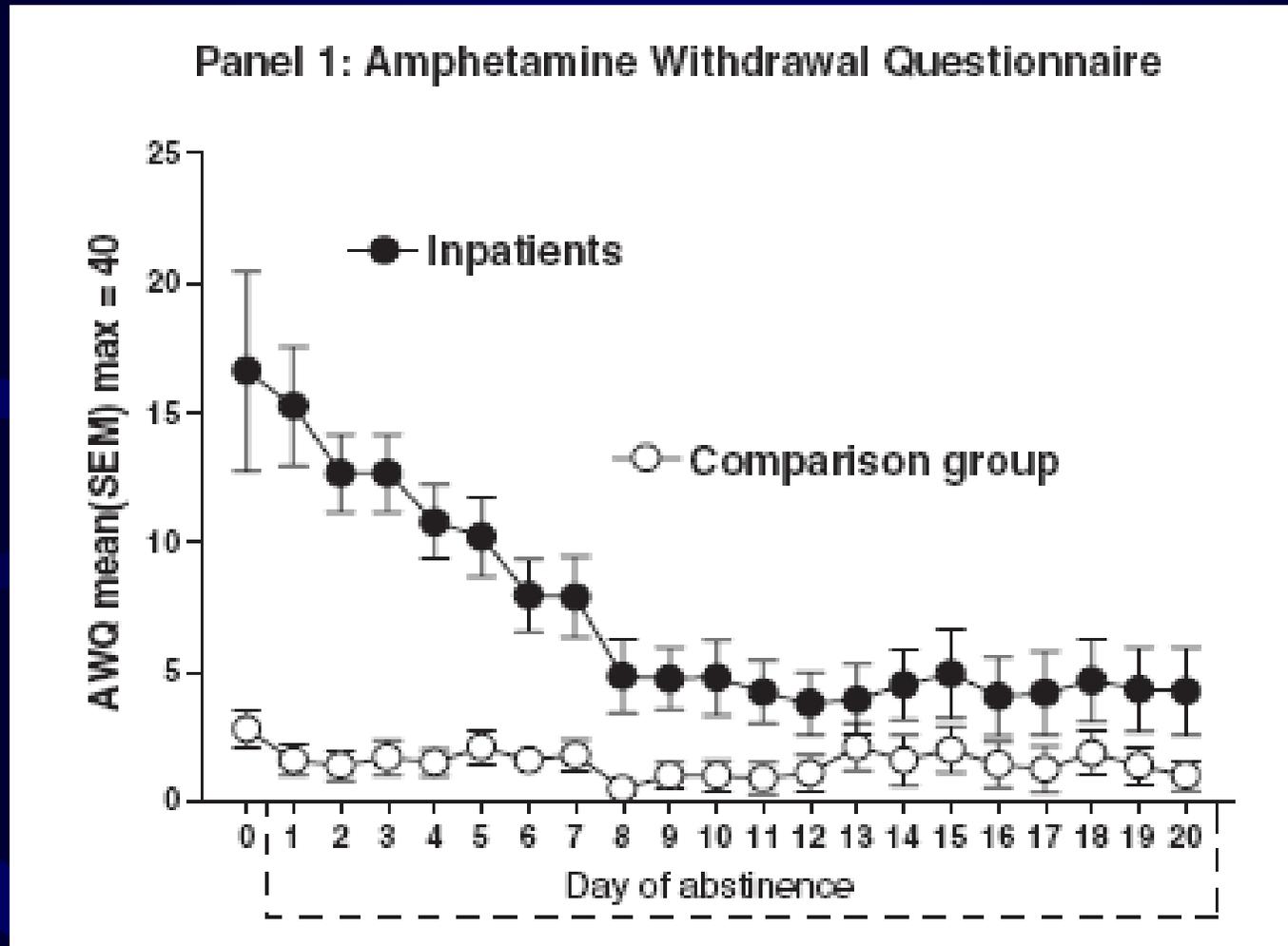
Methamphetamine withdrawal

The nature, time course and severity of methamphetamine withdrawal.

McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

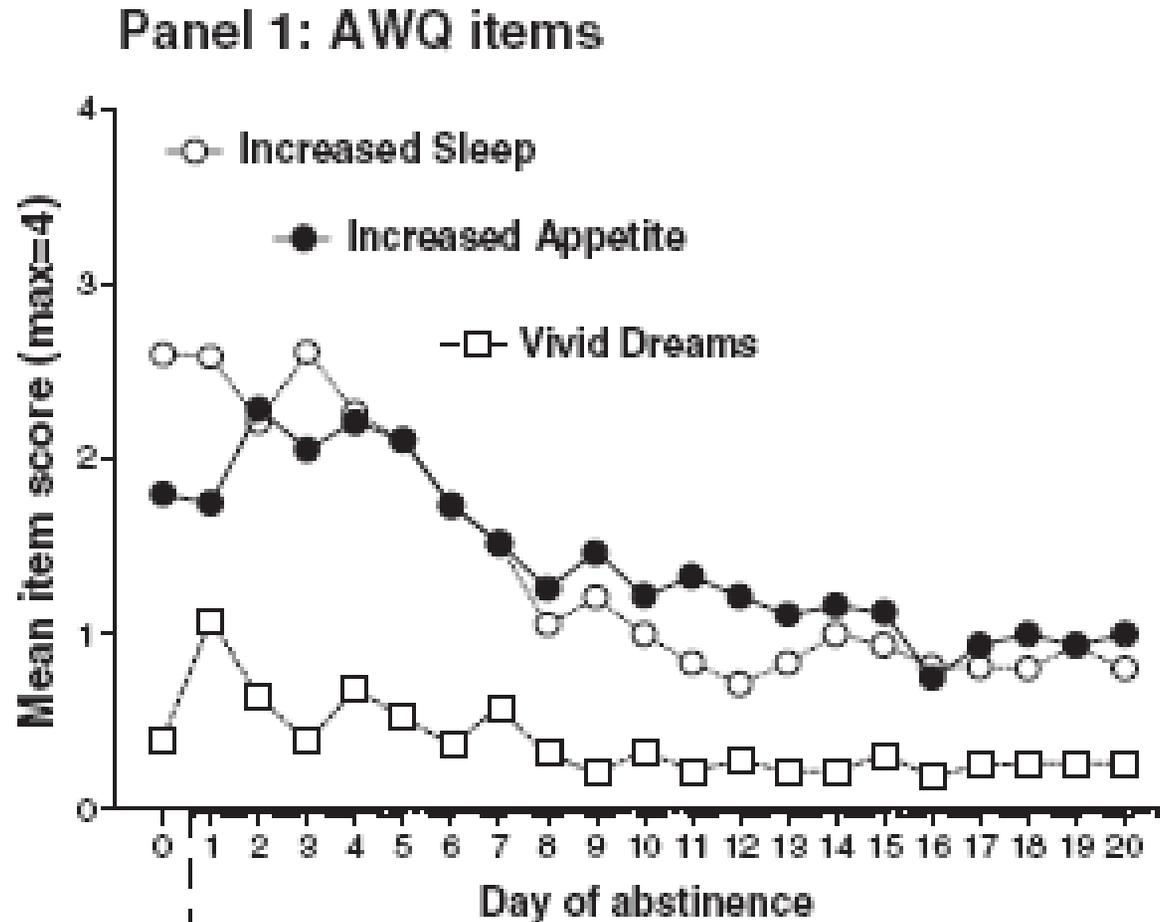
- Studied 21 patients undergoing withdrawal.
- Documented peak withdrawal within 24 hours
- Withdrawal characterized by:
 - Fatigue
 - Hypersomnia
 - Drug Craving
 - Food craving
 - Poor concentration
 - Tension
 - Depression

Methamphetamine withdrawal



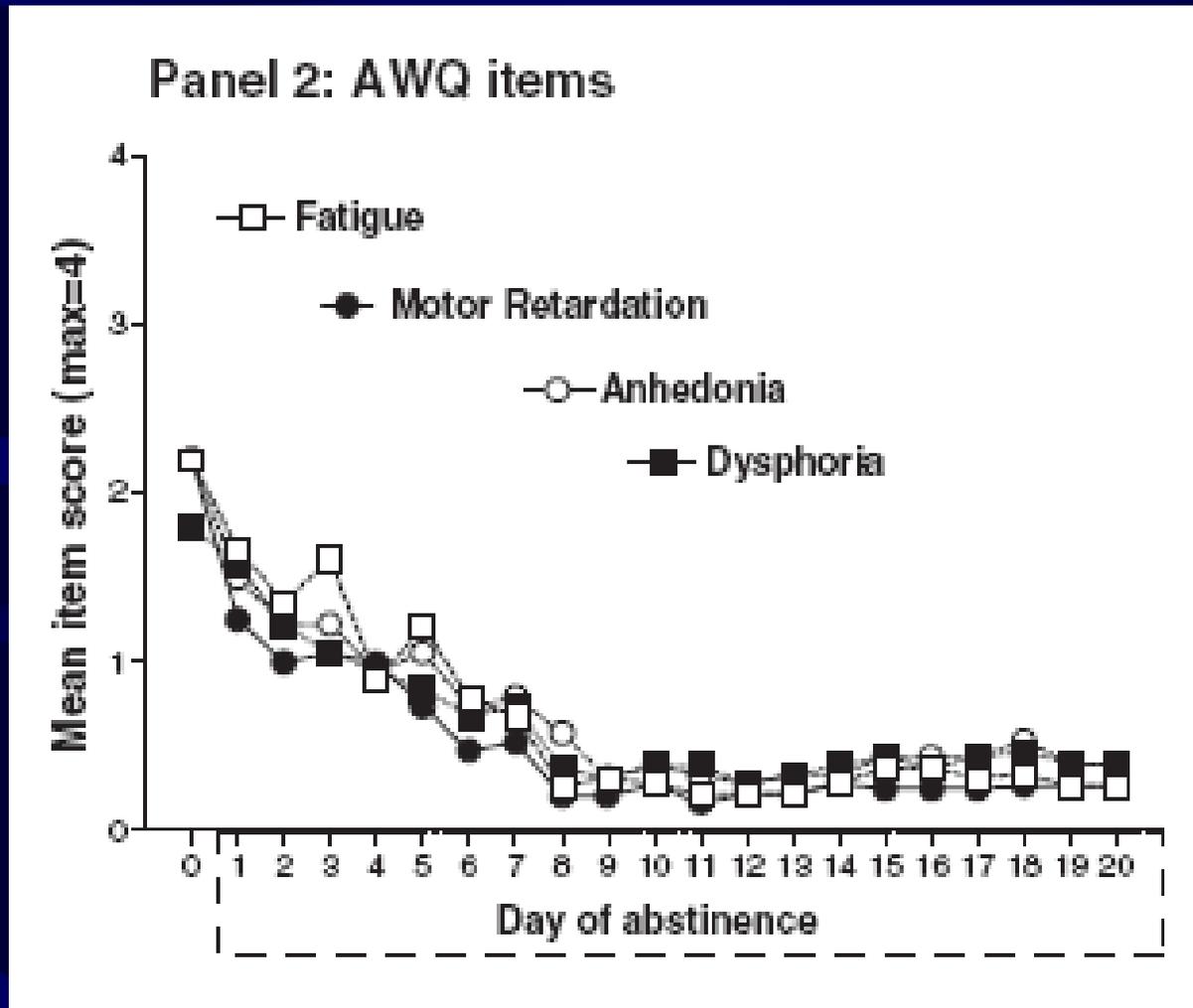
The nature, time course and severity of methamphetamine withdrawal.
McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

Methamphetamine withdrawal



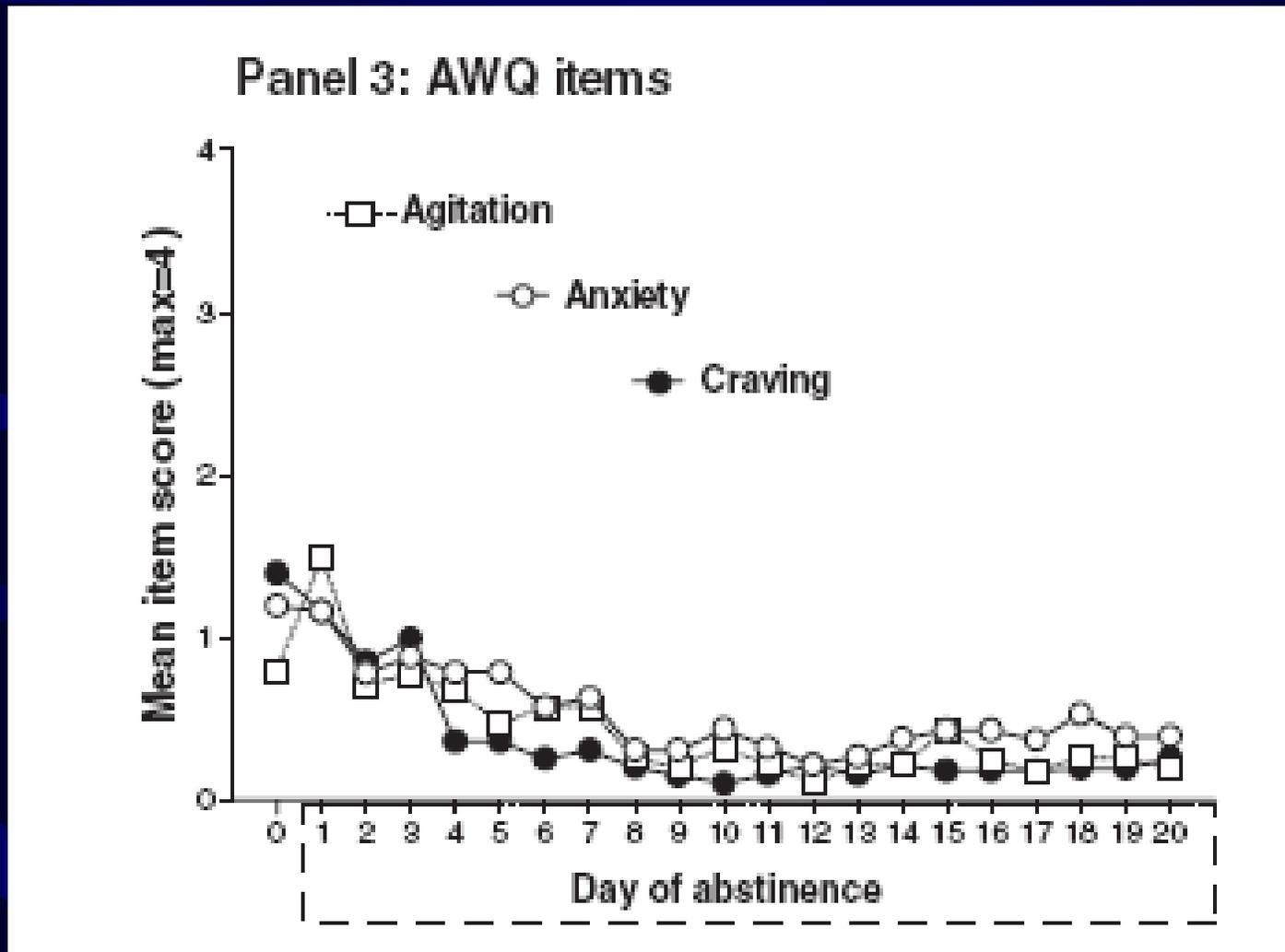
The nature, time course and severity of methamphetamine withdrawal.
McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

Methamphetamine withdrawal



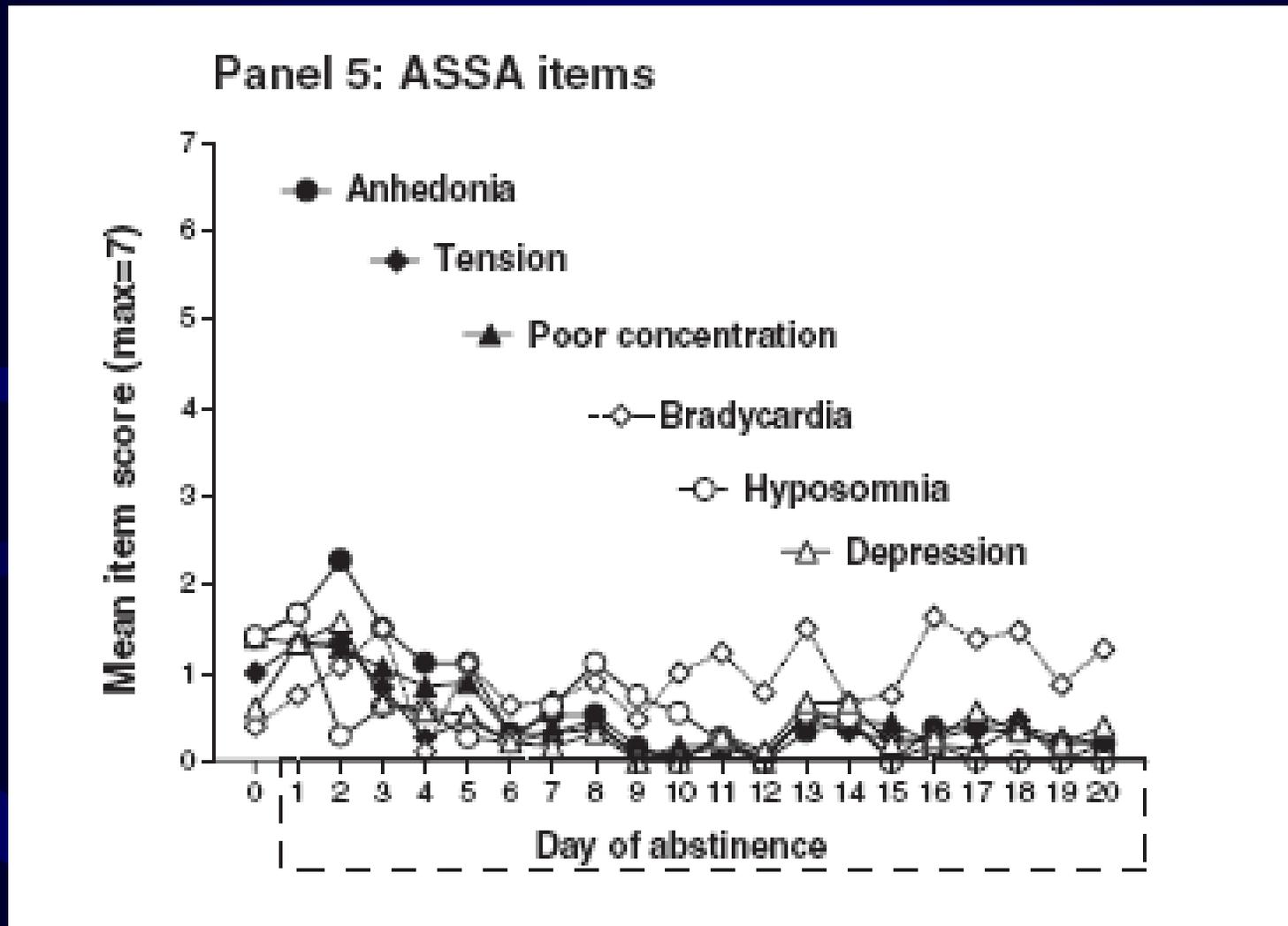
The nature, time course and severity of methamphetamine withdrawal.
McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

Methamphetamine withdrawal



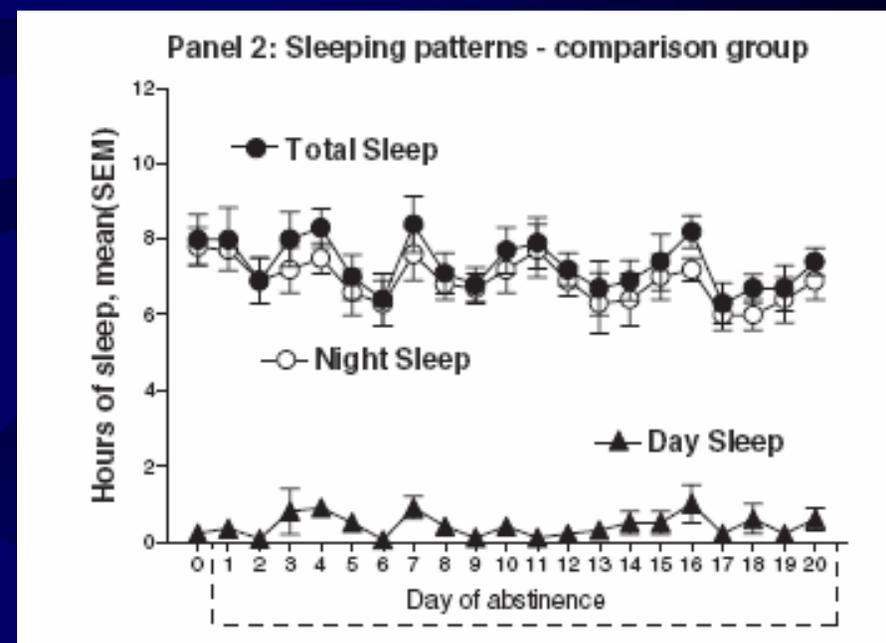
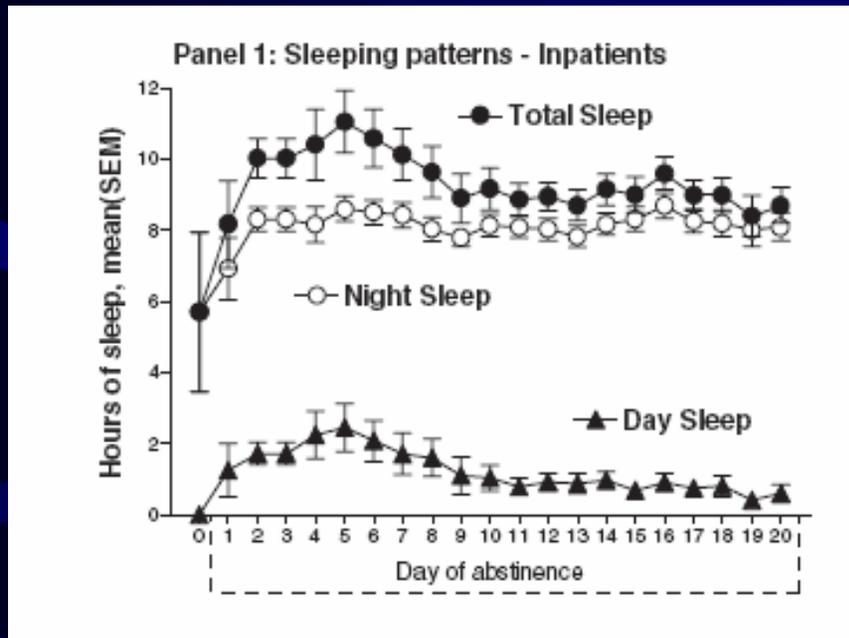
The nature, time course and severity of methamphetamine withdrawal.
McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

Methamphetamine withdrawal



The nature, time course and severity of methamphetamine withdrawal.
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Methamphetamine withdrawal



The nature, time course and severity of methamphetamine withdrawal.
McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

Methamphetamine withdrawal

The nature, time course and severity of methamphetamine withdrawal.

McGregor et al. *Addiction*. 2005 Sep;100(9):1320-9.

- Sleep patterns were disrupted.
 - More daytime sleeping.
 - Less clearheaded on awakening.
 - Poorer quality of sleep.
 - More frequent awakenings at night.
- Acute withdrawal period lasted 9 days.

Methamphetamine: the DRE Assessment

	Acute	Downside
HGN	No	No
VGN	No*	No
Lack of convergence	No	No
Pupil size	Dilated	Normal/constricted
Rxn to light	Slow	Slow
Pulse	Up	Normal to slow
BP	Up	Normal
Temp.	Up	Normal

Methamphetamine: the DRE Assessment

	Acute	Downside	Narcotic
HGN	No	No	No
VGN	No*	No	No
Lack of convergence	No	No	No
Pupil size	Dilated	Normal/constricted	Constricted
Rxn to light	Slow	Slow	Little/none
Pulse	Up	Normal to slow	Down
BP	Up	Normal	Down
Temp.	Up	Normal	Down

Methamphetamine: the DRE Assessment

Other Indicators:

- Speech may be slurred/slow/
- May be “on the Nod”.
- May cycle from alert/agitated to asleep.
- May be lethargic
- May be suicidal/depressed

Methamphetamine and Driving

Review of 101 DRE cases

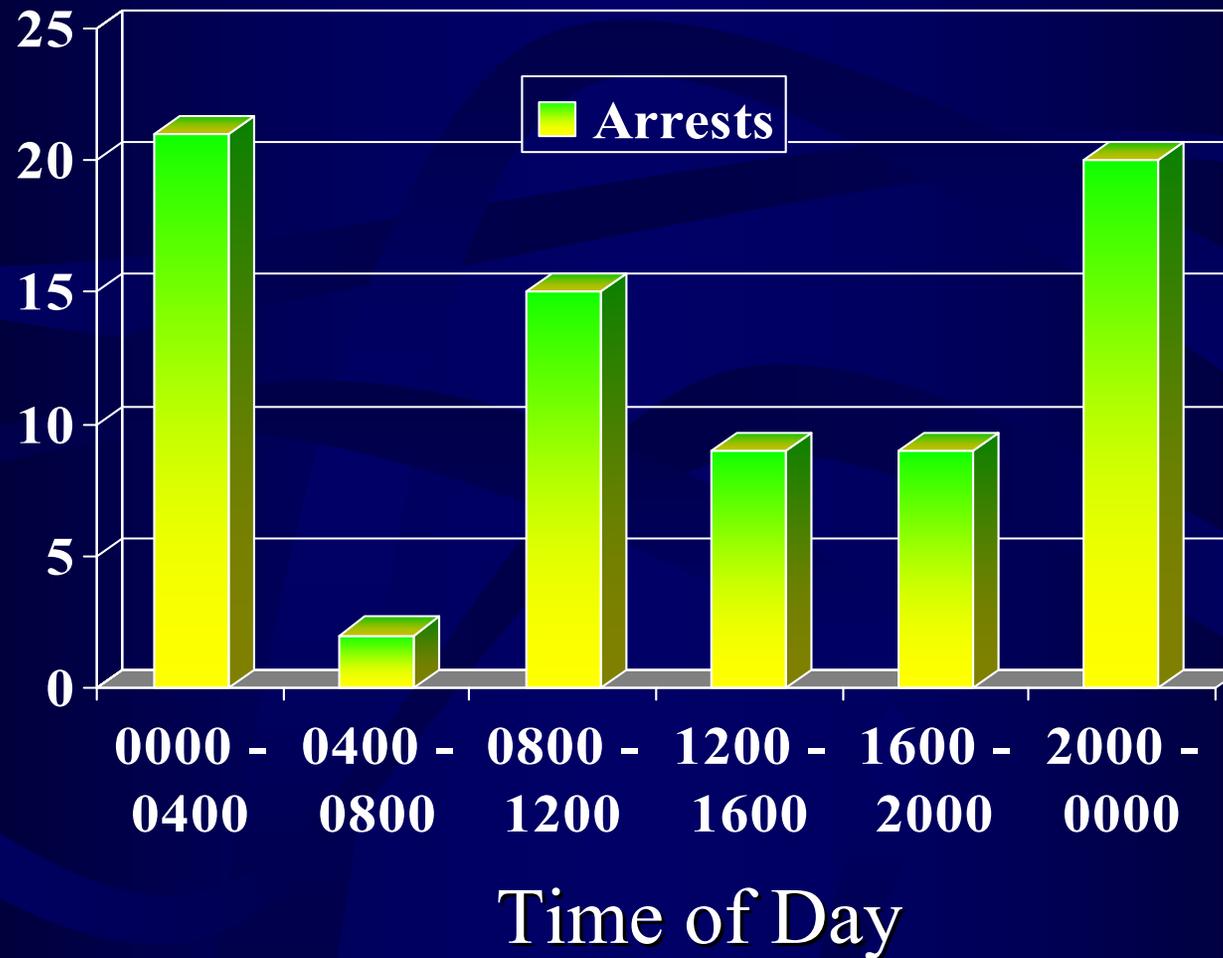
101 Cases Methamphetamine ONLY

Alcohol <0.02g/100mL, Blood cannabinoids <10ng/mL

	Count	Mean age	Mean (mg/L)	Median (mg/L)	Range (mg/L)
♂	74	30.6	0.36	0.27	<0.05 – 2.34
♀	27	31.9	0.34	0.19	<0.05 – 2.36

Methamphetamine and Driving

Time distribution of arrests

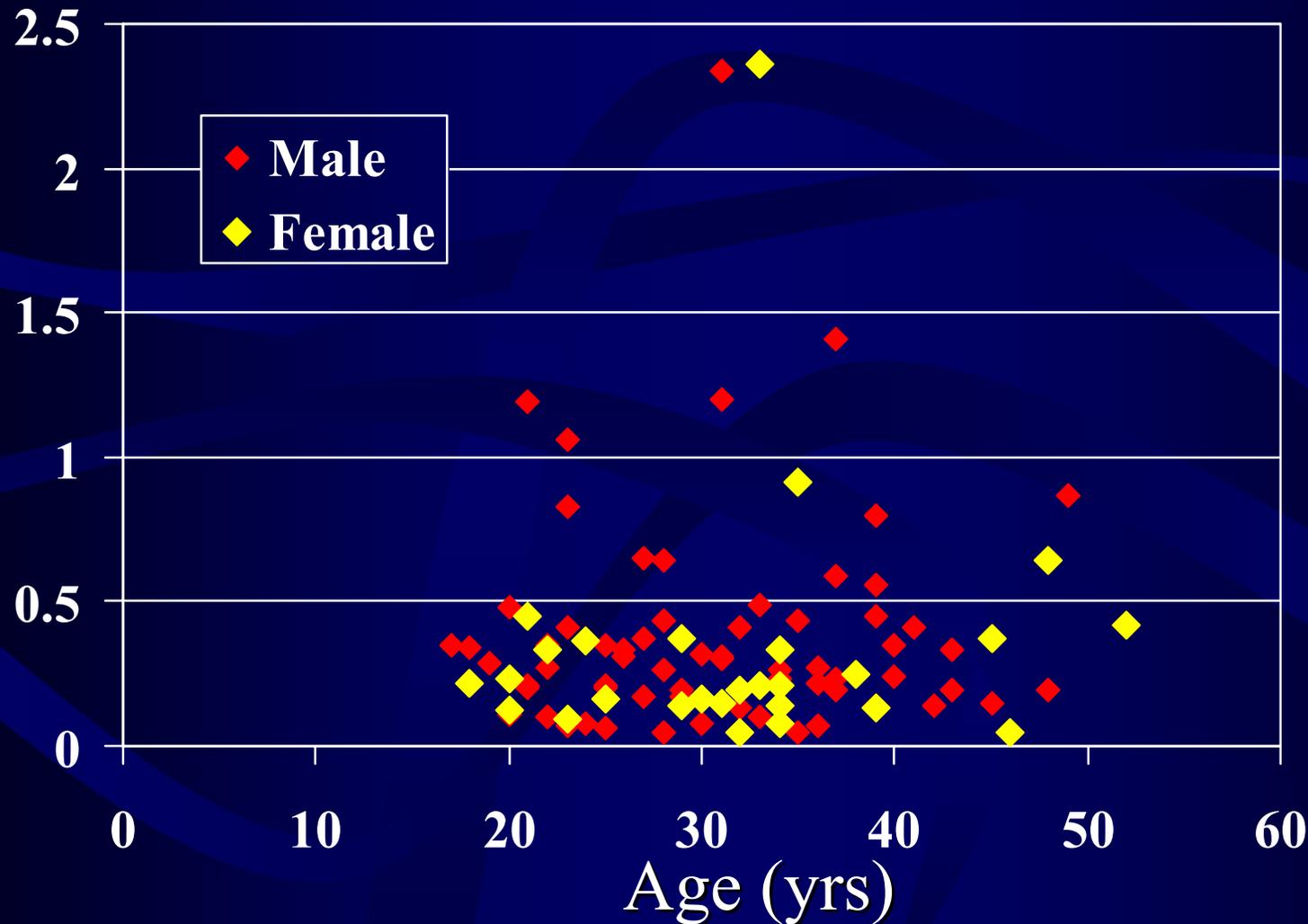


Methamphetamine and Driving

Reason for stop	#	%
Lane Travel (OOL)	23	33
Erratic Driving	9	13
Equipment	9	13
Accident	8	12
Speeding	7	10
CVE	3	4
Hit and Run	2	3
Lane Travel (WL)	2	3
Stolen Vehicle	2	3
Eluding	1	1
Fail to Signal	1	1
Reckless Driving	1	1
Wrong way	1	1

Methamphetamine and Driving

Age/Gender/Concentration Distribution

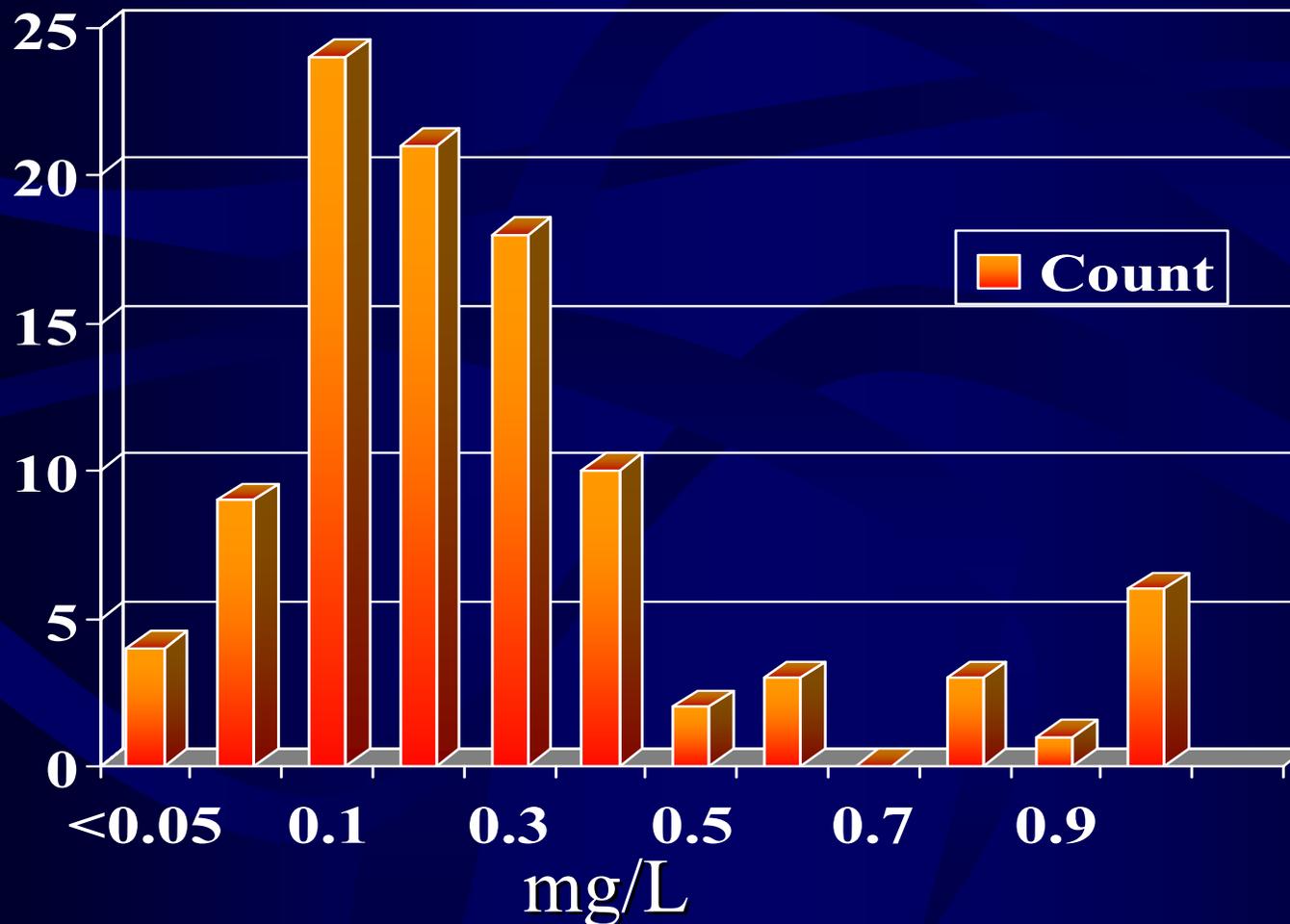


♂ = 74
♀ = 27

Methamphetamine and Driving

Methamphetamine distribution (mg/L)

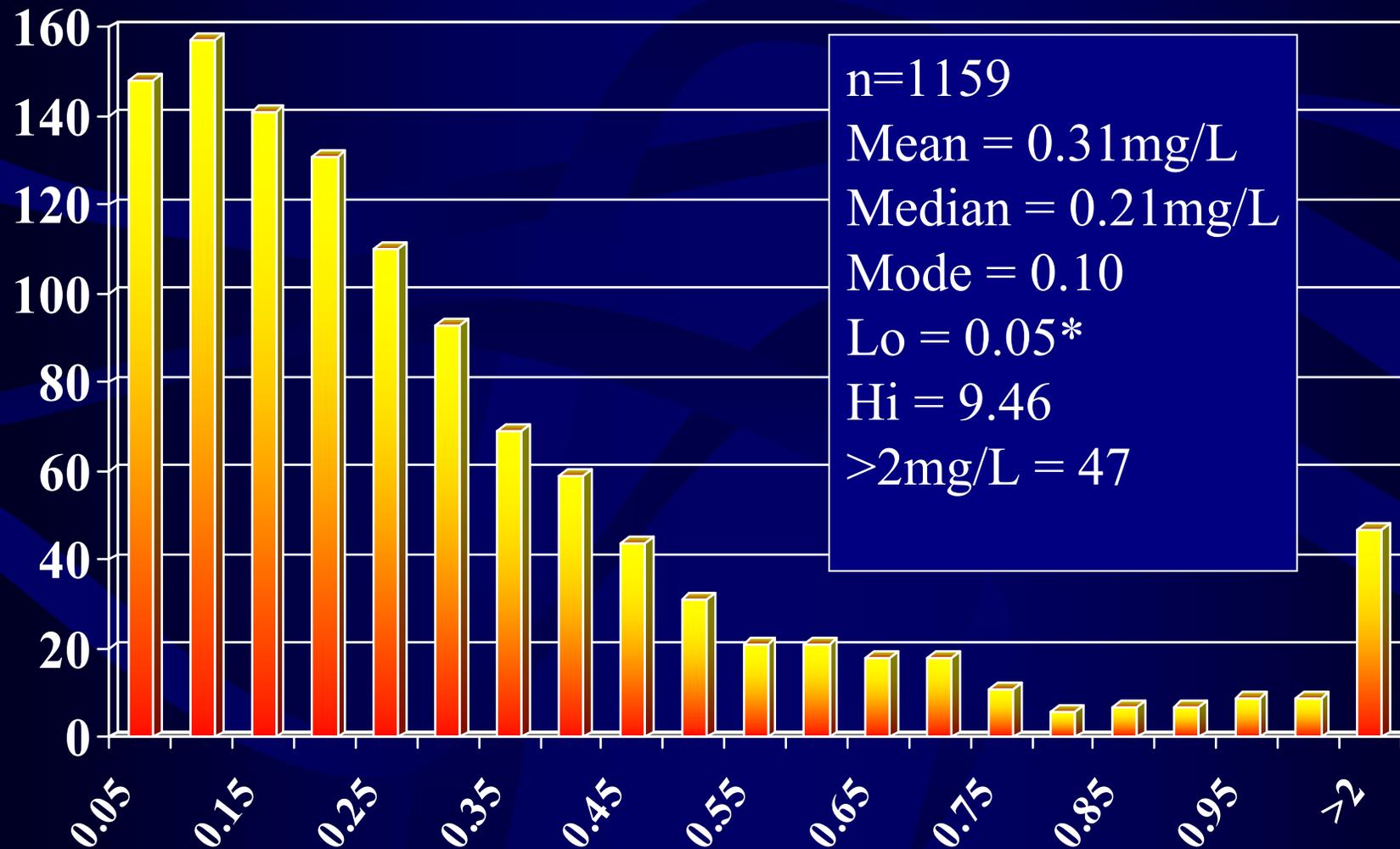
Mean = 0.35mg/L, Median = 0.23mg/L



Methamphetamine Drivers

Methamphetamine Concentrations in Impaired Drivers

Logan (unpublished, 2006)



Stimulants and Driving

SFST Impairment indicators:

One Leg Stand

*Sways, Uses arms, Hops, Foot touches,
Inches of sway (in. bf/ss)*

Walk and Turn

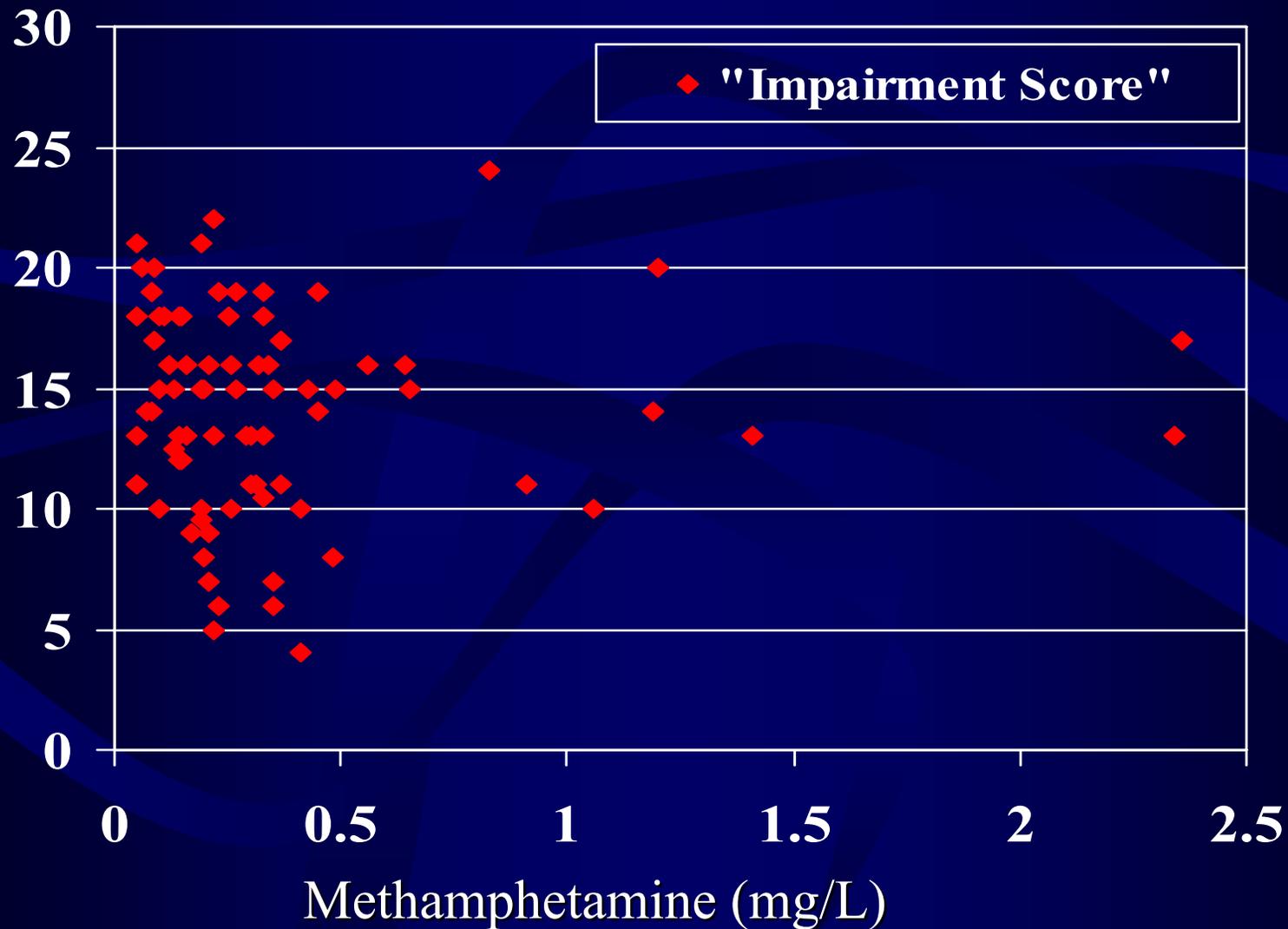
*Can't balance, Starts too soon, misses heel
to toe, walks off line, stops, puts arms up,
incorrect # steps*

Finger to nose

Accuracy on 6 attempts

Methamphetamine and Driving

Impairment vs. methamphetamine conc.



Methamphetamine and Driving

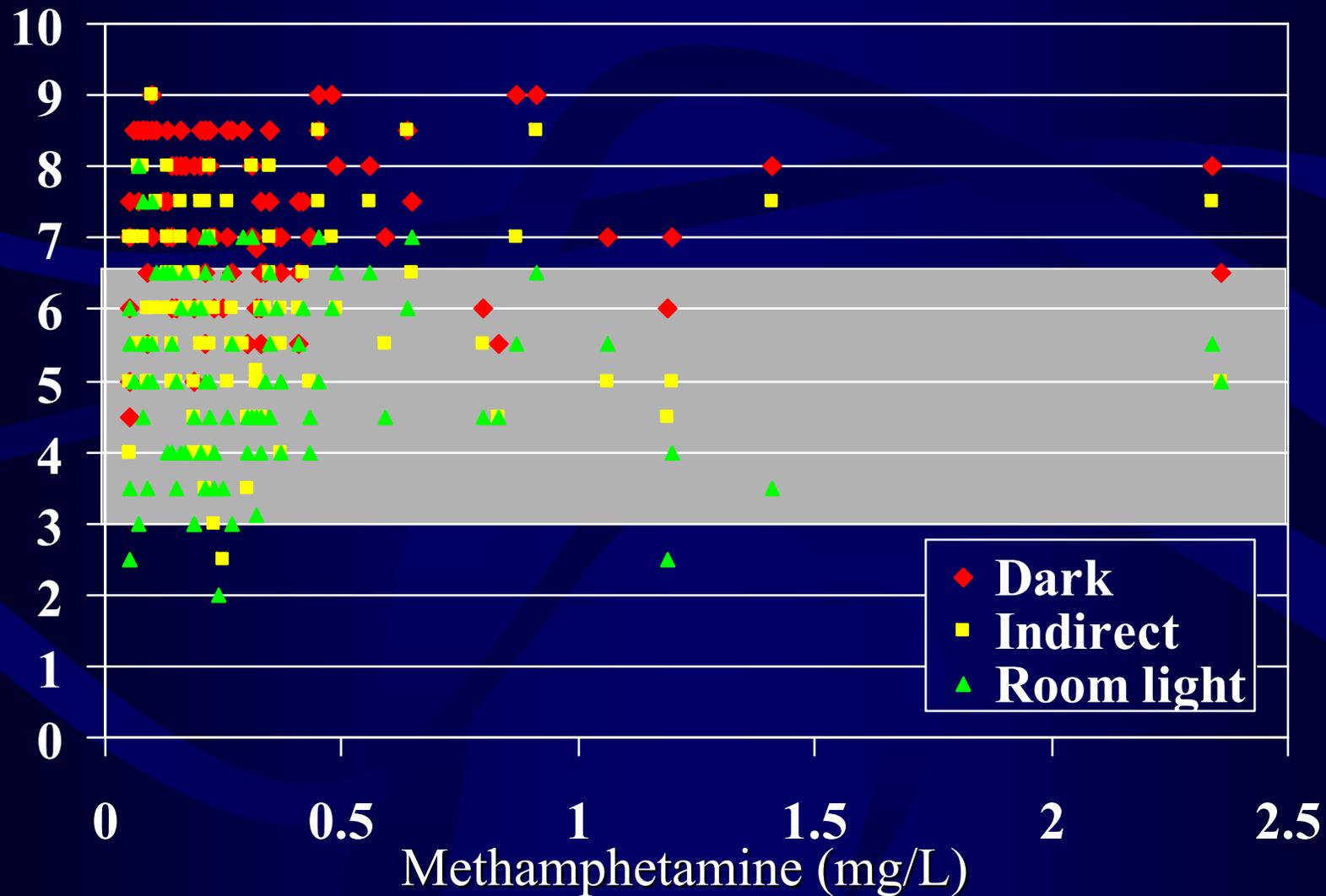
Methamphetamine Behaviors:

Eyes:

- Tracking and pupils equal in all.
- All could follow stimulus.
- Bloodshot, watery (80%), red, glassy.
- Eyelids droopy in about half
- HGN: Only 6 had more than 3 clues and they all had 6.
- VGN: 2 (also had 6 clues in HGN).
- 25 had lack of convergence.

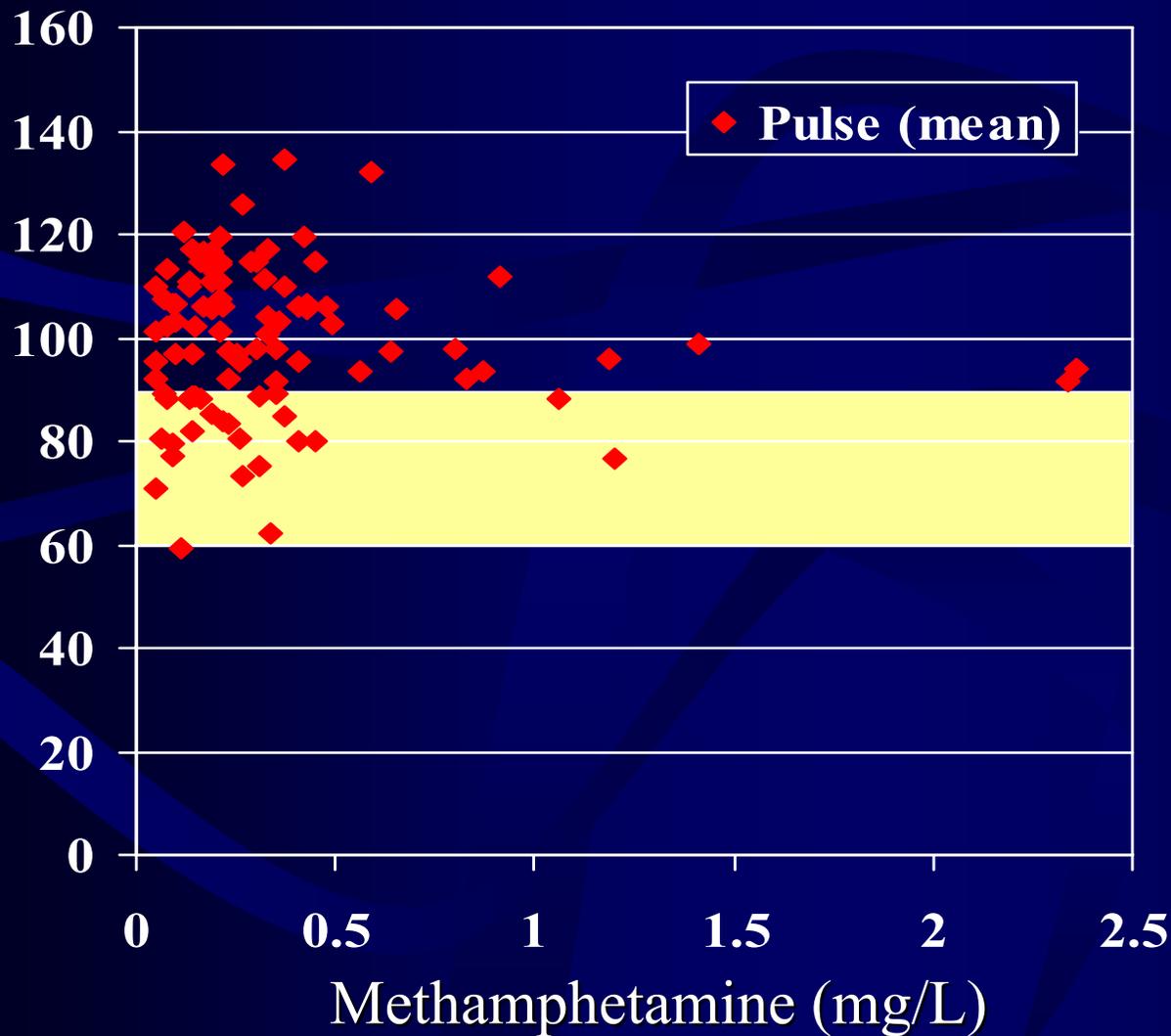
Methamphetamine and Driving

Pupil diameter and methamphetamine conc.



Methamphetamine and Driving

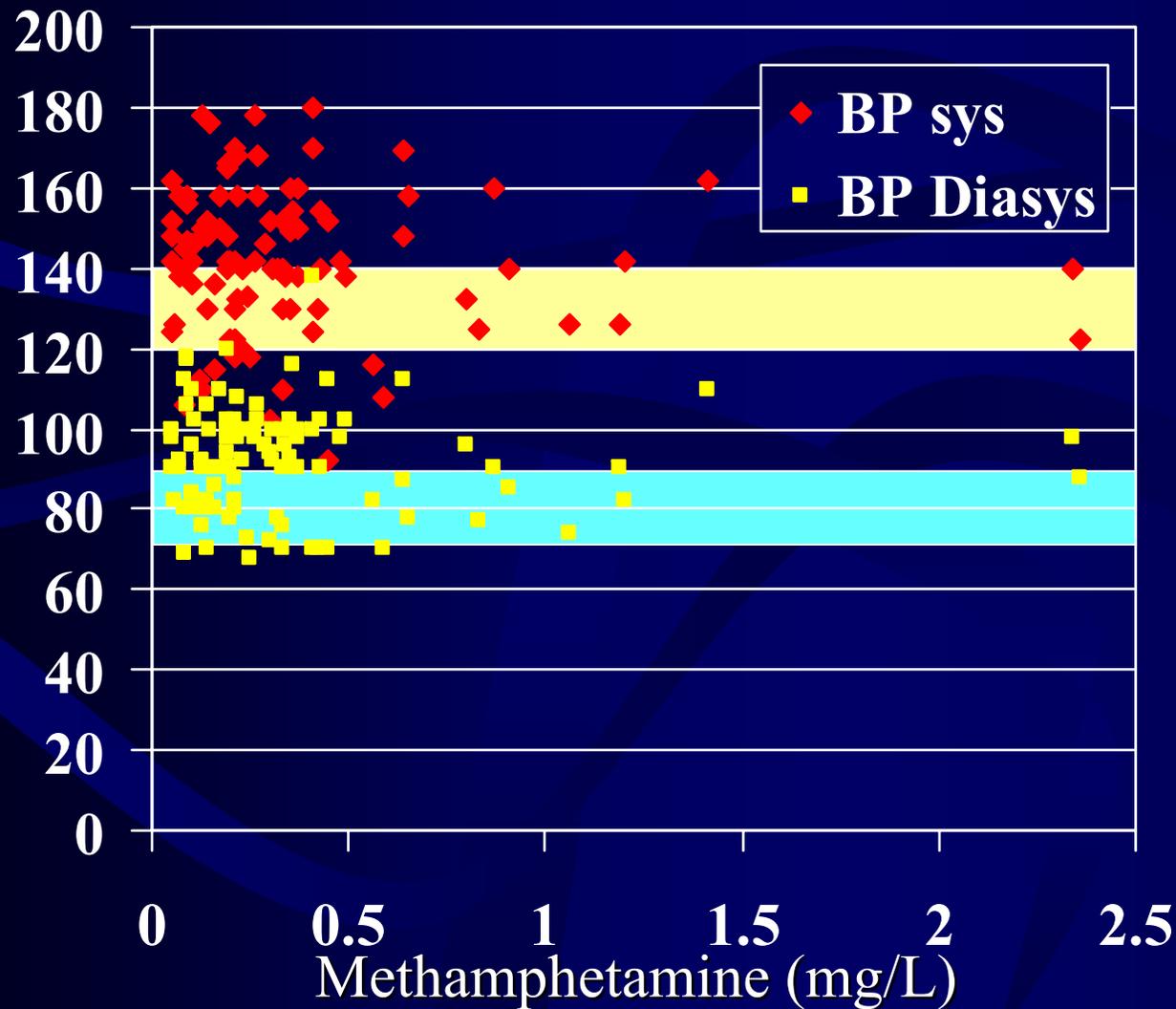
Pulse vs. methamphetamine conc.



74% exceed
normal range
(60-90 BPM)

Methamphetamine and Driving

Blood pressure and methamphetamine conc.

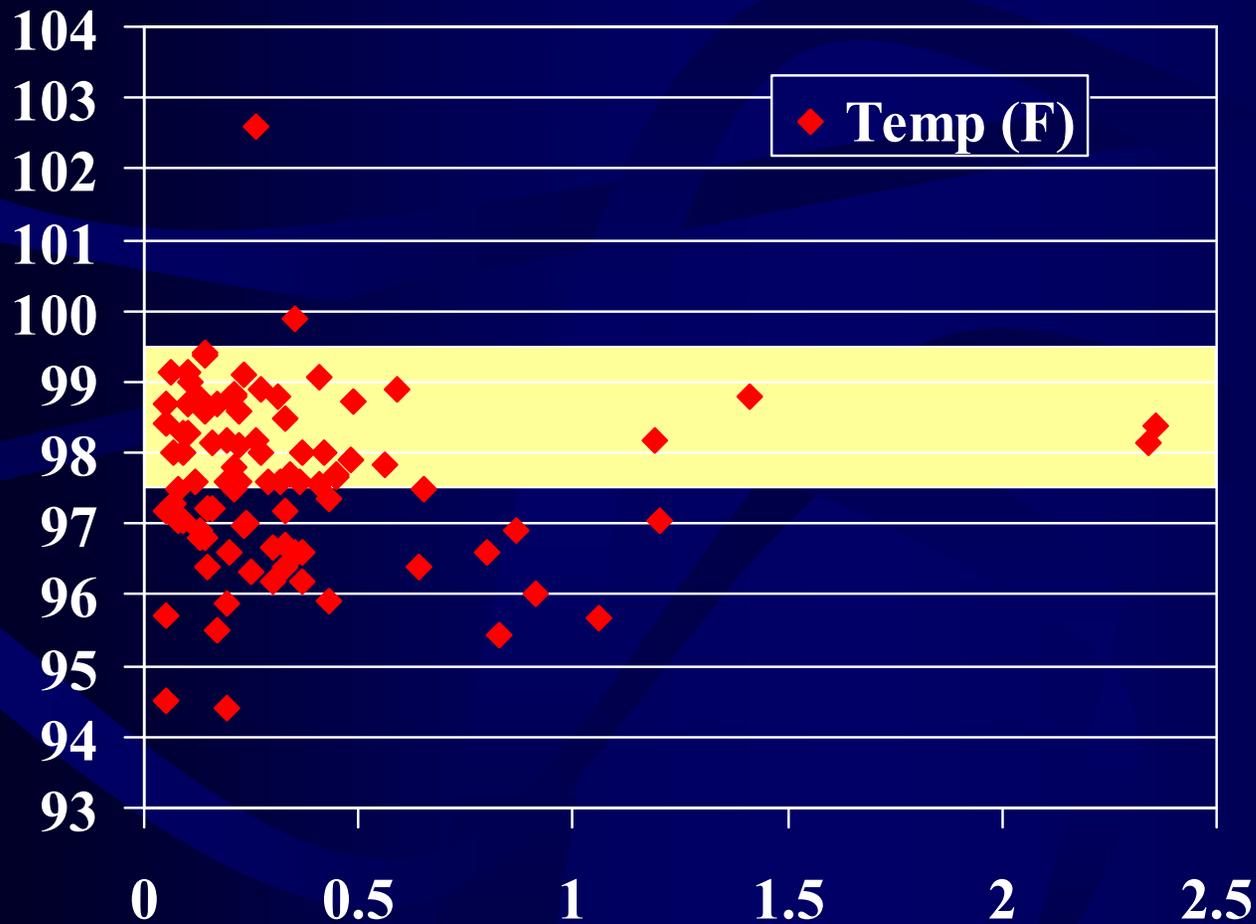


Systolic –
66% exceed
normal

Diasystolic –
50% exceed
normal

Methamphetamine and Driving

Body temperature and methamphetamine



53% fell below
normal
temperature
range of
97.6 – 99.6°F

3% exceeded

Methamphetamine and Driving

Methamphetamine Behaviors:

Coordination:

Poor, jerky, fast movements, jittery, fidgeting, staggering, awkward, unsteady, clumsy, deliberate, slow.

Speech:

Fast, rapid, non-stop, unintelligible, mumbling, low, raspy, hoarse, slow, thick tongued.

Injection marks

29% track marks, 26% recent injections

Same for ♂ and ♀.

Methamphetamine and Driving

DRE Call	All (n= 101)	Adm. (n=76)	No Adm.(n=14)
Stimulant	64	53	7
Stimulant & Cannabis	11	9	1
Stimulant & Narcotic	7	4	1
Stimulant & Depressant	5	3	2
Stimulant, Depressant & Cannabis	3	2	0
Stimulant, PCP, Cannabis	1	1	0
<i>Any Stimulant Call</i>	<i>90%</i>	<i>95%</i>	<i>79%</i>
No Drugs Present	2	1	1
Cannabis	3	2	1
Depressant	1	0	1
Cannabis & Narcotic	1	1	0
Depressant, Narcotic & Cannabis	1	0	0
Depressant & Narcotic	1	0	0

Methamphetamine and Driving

Conclusions

- Methamphetamine impairs during both the acute intoxication phase and the withdrawal or downside phase.
- The upside impairment is more agitated, euphoric, impulsive, risk taking, hypervigilant, paranoid.
- The downside impairment is more like CNS depression, with fatigue, sleepiness, motor retardation, poor concentration.
- The physiological signs may not exactly fit the matrix.