

NCANDA: Characterization of the Sample

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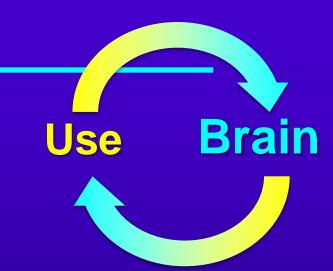
NCANDA Overview

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- ◆ Aims
- Recruitment
- Protocol
- Sample characteristics
- Next directions

Background

- Adolescence:
 - Neuromaturation
 - Escalations in substance use



- Deleterious effects of heavy drinking on adolescent neuromaturation suggested
 - Cross-sectional analyses
 - Smaller longitudinal studies
- Verify in large, representative sample

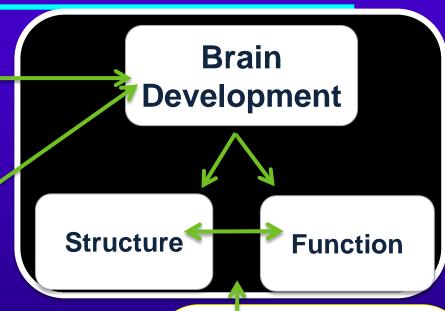
NCANDA Conceptual Model

Adolescent Alcohol Consumption

Dose/Duration/Timing

Developmental Context

- Genetics
- Age/Gender/Puberty
 - Risks



Environment

Behavior

- Neurocognition
 Functioning
- Alcohol problems/AUD
 - Psychopathology
 - Maturation
 - Real life functioning

NCANDA Aims

- Drinking → adolescent neurodevelopment
- 2. Effects of dose, duration, and age of drinking
- 3. Resolution of effects with abstinence
- 4. Modulating factors:
 - Pubertal stage
 - -Sleep
 - -Sex
 - Psychopathology
 - Family history of alcoholism
- Brain features → addiction & psychopathology

NCANDA Design

Administration:

- Sandy Brown Coordinator
- Susan Tapert Scientific Director

Data:

- Dolf Pfefferbaum
- Kilian Pohl
- Edie Sullivan

Sites:

- U Pittsburgh Duncan Clark
- SRI Ian Colrain & Fiona Baker
- Duke Univ Michael DeBellis
- OHSU Bonnie Nagel
- UCSD Susan Tapert

Scientific Advisors:

- Ken SherRaquel Gur
- Andrea HussongArpana Argrawal
- Bob Zucker

5 Sites

>50,000 reached via school and community recruitment

>7,500 responded

831 enrolled

53% Representative 47% with 1+ risk factor for heavy drinking, including 17% who exceeded threshold for alcohol use

3 annual follow-ups (~25% heavy drinkers)

Exclusions at Project Entry

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- ♦ Not age 12.0 21.9 years
- No parental consent
- Factors that preclude valid participation
- Early developmental problems
- Major psychiatric disorder
- Medications
- Serious medical problem
- Excessive substance use

Substance Use Criteria

Lifetime

For classification as Non/Low Drinker:

Age	Days Drinking ^a	Occasion a		Cigarette Use ^b	Marijuana Use ^b	Other Drug Use ^b
		Male	Female			
12-12.9	≤ 5	≤3	≤3	≤10	≤5	≤1
13-13.9	≤ 5	≤3	≤3	≤10	≤10	≤2
14-14.9	≤ 5	≤4	≤3	≤20	≤15	≤3
15-15.9	≤ 5	≤4	≤3	≤30	≤20	≤4
16-16.9	≤11	≤4	≤3	≤40	≤25	≤5
17-17.9	≤23	≤4	≤3	≤50	≤30	≤6
18-18.9	≤51	≤4	≤3	≤60	≤35	≤7
19-19.9	≤51	≤4	≤3	≤70	≤40	≤8
20-20.9	≤51	≤5	≤3	≤80	≤45	≤9
21-21.9	≤51	≤5	≤3	≤90	≤50	≤10

^a NIAAA, 2008

Lifetime

Lifetime

Lifetime

^b SAMHSA, 2013

NCANDA Screening

Step-down screening approach >50,000 reached, >7,500 responded

1110 Ineligible

2548 Completed Screens

> 1438 Eligible

252: MRI / Physical / No parent

310: Substance use

548: Meds / Prenatal / LD

607 not enrolled (target N met with representation & ~50% at-risk)

831 Enrolled **692**Non/Limite
d Drinking

139 Exceeded

Drinking

Thresholds

Modified Accelerated Longitudinal Design

- Samples subjects from range of ages
- Oversampled age 12-15
- Allowed ~15% to exceed drinking thresholds
 - Mostly 18-21
 - Can estimate trajectories representing continuum from non-drinking to heavy drinking
 - Accelerated time scale

NCANDA Protocol: Interview

outh

Youth assent (age 12-17) or consent (18+)

Computerized Semi-Structured Assessment for the Genetics of Alcoholism *NCANDA Version

Demographic Interview

Family History Assessment Module

MacArthur Sociodemographic Questionnaire (18+)

Customary Drinking & Drug Use Record

Traumatic Brain Injury Identification Method-Short Form

Youth Self-Report (12-17) or Adult Self-Report (18+) – dimensional psychopathology

Exit interview

Locator update and payment

Parent consent

Computerized Semi-Structured Assessment for the Genetics of Alcoholism-Parent (12-17)

Demographic Interview-Parent

Traumatic Brain Injury Identification Method-Short Form

Family History Assessment Module

MacArthur Sociodemographic Questionnaire - Parent

Child Behavior Checklist (12-17)

Access to Substances (12-17)

Exit interview

Locator update and Payment



60-90

min

NCANDA Self-Report Measures:

Access to Substances & Neighborhood Strength (12-17)
Alcohol Expectancy Questionnaire-Adolescent version
Behavior Rating Inventory of Executive Function
California Healthy Kids Survey (12-18) 45-60
Child Trauma Questionnaire min
Driving & Riding Behavior & History Questionnaire (15+)
Drug Taking Confidence Questionnaire
Hangover Symptom Survey
Life Events Questionnaire
Motives for Abstaining from Alcohol Questionnaire
Parent Warmth, Monitoring, Knowledge, Control & Relationship (12-18)
Parental Monitoring (12-18)
Peer & Romantic Relationships
Peer Group Deviance
Prosocial Tendencies
Pubertal Development Scale
Responses to Stress Questionnaire
Sexual Behavior Questionnaire
Single-Item Self-Esteem Scale
Sleep: self-reported sleep problems, total sleep, chronicity, and weekday-weekend sleep difference
Social Support Questionnaire
Ten Item Personality Inventory
UPPS-P Impulsive Behavior Scale
Exit interview

Other Data Collected:

All Sites

MRI: T1, T2, DTI, & resting state fMRI

Neuropsychological assessment

Samples for genetic and epigenetic analyses, pubertal hormones, and drug screening

Specialty Projects at 2 Sites Sleep studies – SRI & Pittsburg

Stroop fMRI task – SRI & UCSD

Anti-saccade fMRI task – Duke & Pittsburg

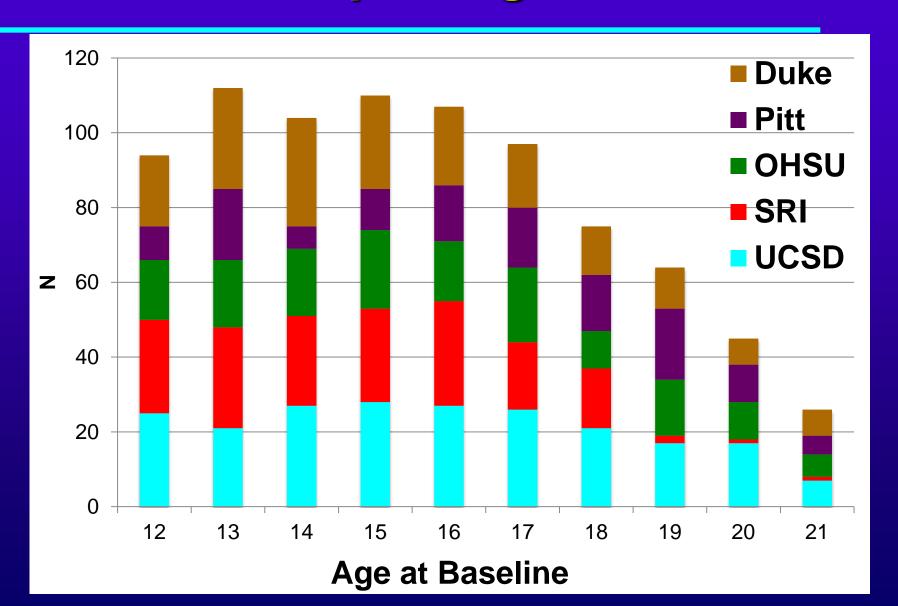
Recovery protocol – UCSD & Duke

Staff Training & QA

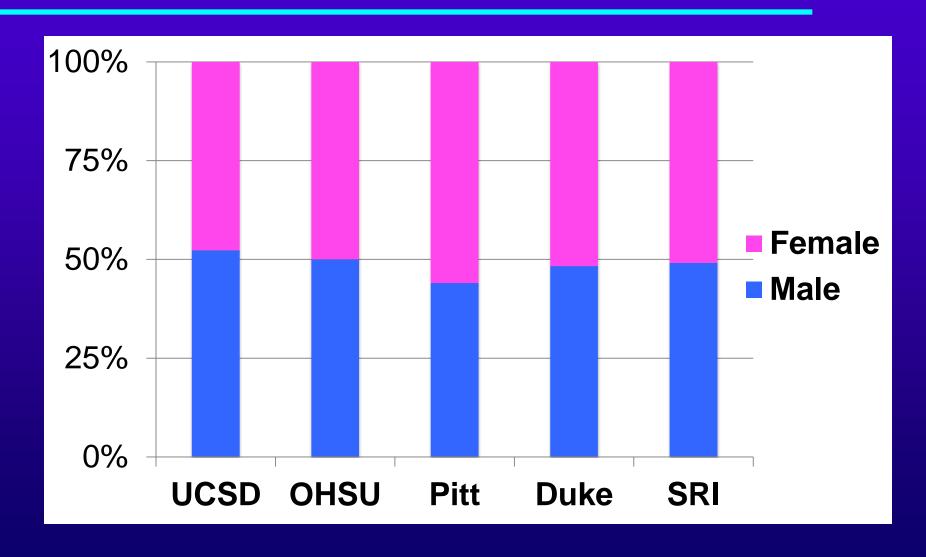
- Senior staff at each site
- Training process:
 - 1. Readings and observation
 - 2. Repeat mock sessions observed by senior staff with feedback
 - 3. Mock session approved by senior-level staff member
 - 4. Observed assessment with real subjects
- QA:
 - Annual calibration at each site
 - Check for interviewer drift and confirm training of new staff
- Additional in-person and Skype training and reliability checks
 - Dr. Schuckit provided video-recorded training on SSAGA
- Manuals: clinical, neuropsych, MRI, and data management



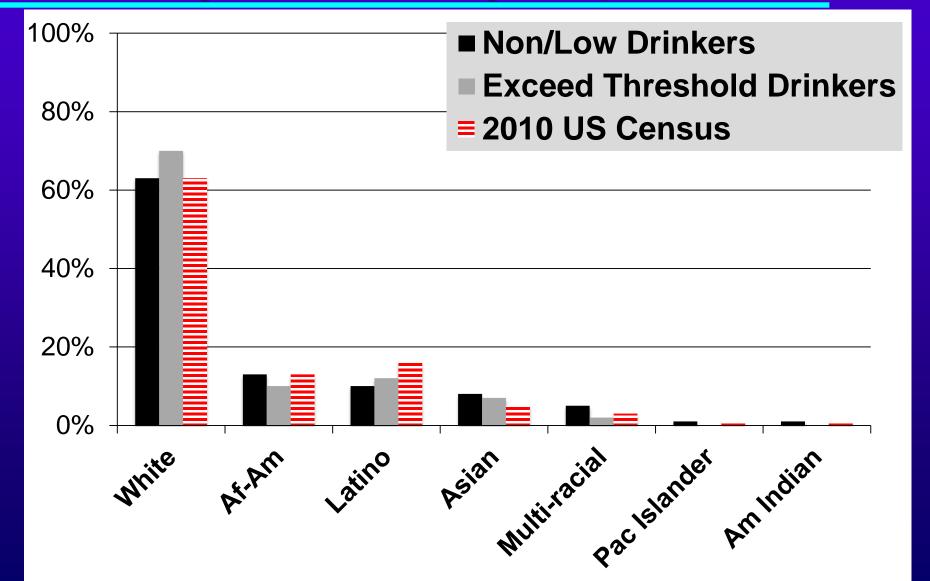
NCANDA Sample: Age Distribution



Balanced Female: Male Ratio



Ethnicity: Nationally Representative



Sample Characteristics N = 831, age = 12-21 years



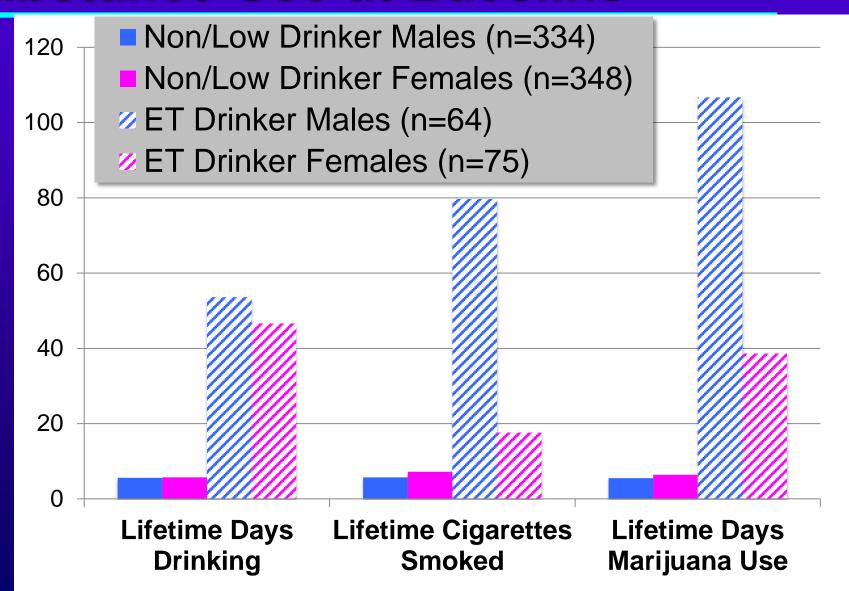
	Non/Low Drinker (N=692)		Exceeded Threshold (N=139)	
	Male	Female	Male	Female
	(n=334)	(n=348)	(n=64)	(n=75)
Age (mean years)	15.6	15.8	18.4	18.5
Pubertal Development Scale (median)	3.0	3.6	3.6	4.0
Socioeconomic status (mean)	17.0	16.6	17.1	17.3
% Right-handed	76%	83%	83%	81%

Sample Description

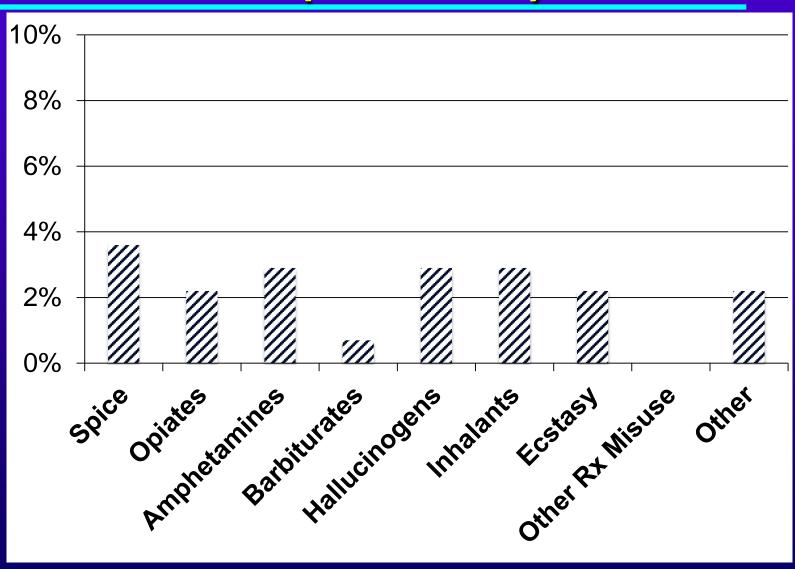
- 88% reside with one or both parents
- 27% parents divorced/separated/single
- ◆ 20% below college degree
- 18% household income <\$50k/yr*</p>
- **♦ 85% normal BMI (13% above, 2% under)**

*higher in ET Drinkers (p<.05)

Substance Use at Baseline



Lifetime Other Drug Use: ET at Baseline (% of 139)



Risk Criteria: Target 50%

1. 1st full drink <age 15

(Grant & Dawson, 1997)

- 2. Family history of substance use disorder (Edenberg et al., 1998; Schuckit & Smith, 1996)
- 3. 1+ externalizing symptoms

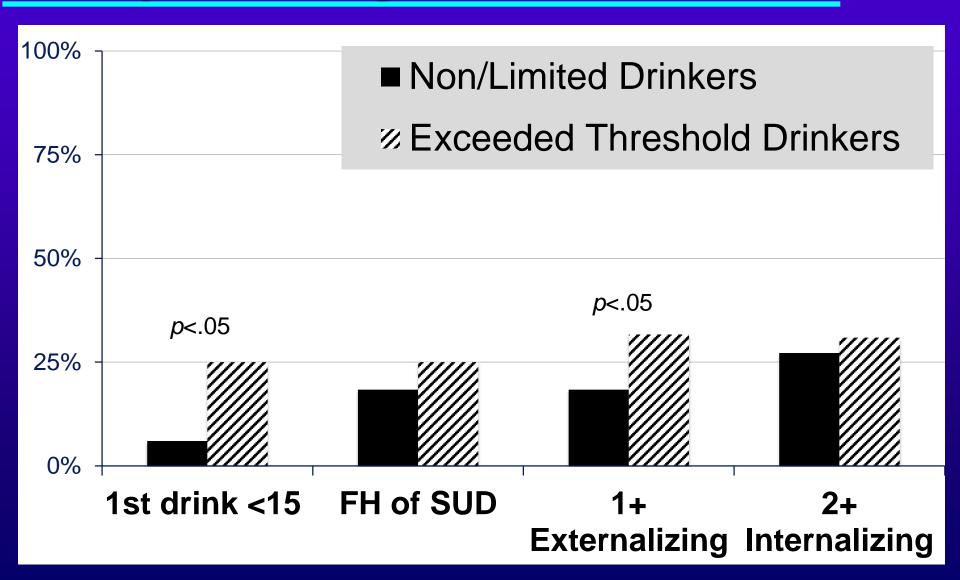
(Brown et al., 1996; Myers et al., 1995; Slutske et al., 1998)

4. 2+ internalizing symptom

(Chassin et al. 2002; Husseng et al. 2014)

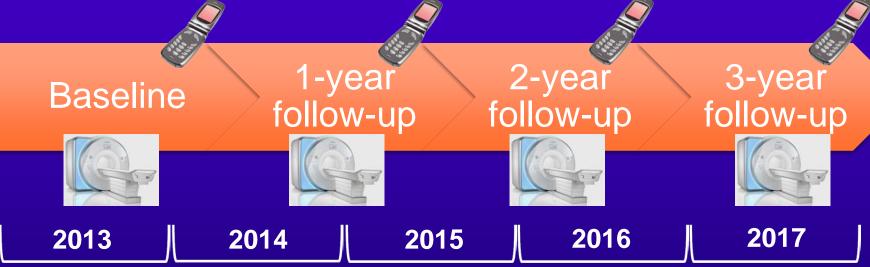
(Chassin, et al., 2002; Hussong et al., 2011)

47% with 1+ Risk Factor for Heavy Drinking



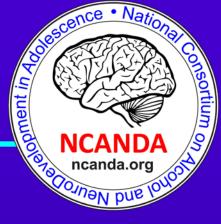
NCANDA Follow-up Design





- 4 assessments with imaging
 + Half-yearly telephone interviews
- 6-month: 98% of 831 done.
- 1-year: in progress, >90% complete.

Next Directions



- Begin to test Aims 1 and 5:
 - Effects of adolescent drinking on trajectory of adolescent brain development
 - Neural, cognitive, and affective markers predicting addiction & psychopathology

NCANDA Summary



- Exceeded recruitment goals on time
- Nationally representative sample
- Procedures and protocol set a standard
- High follow-up rates
- 47% with risk factors
 - Suggests aims can be tested

Acknowledgements

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- ◆U01 AA021690 (DBC)
- ◆U01 AA021691 (BN)
- ◆U01 AA021696 (IMC+FCB)
- ◆T32 AA013525 (TB)



- ◆NIAAA Program Staff
- NCANDA Scientific Advisory Board
- NCANDA Co-Investigators
- NCANDA Research Associates



