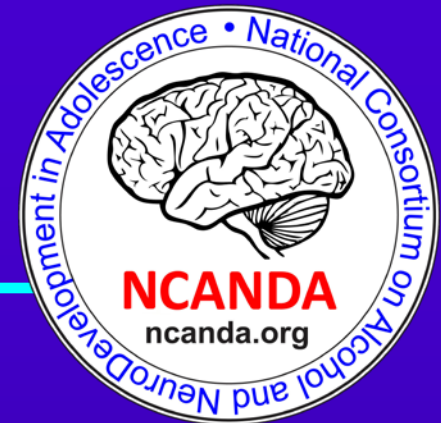


NCANDA: Characterization of the Sample

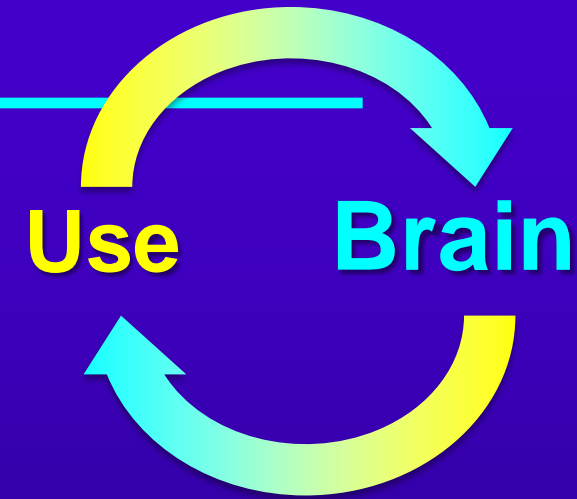
Susan Tapert, Ty Brumbach,
Kristin Tomlinson, Kevin Cummins,
Fiona Baker, Duncan Clark, Ian Colrain,
Michael De Bellis, Bonnie Nagel, Weiwei Chu,
Torsten Rohlfing, Killian Pohl, Edith Sullivan,
Adolf Pfefferbaum, & Sandra Brown

NCANDA Overview



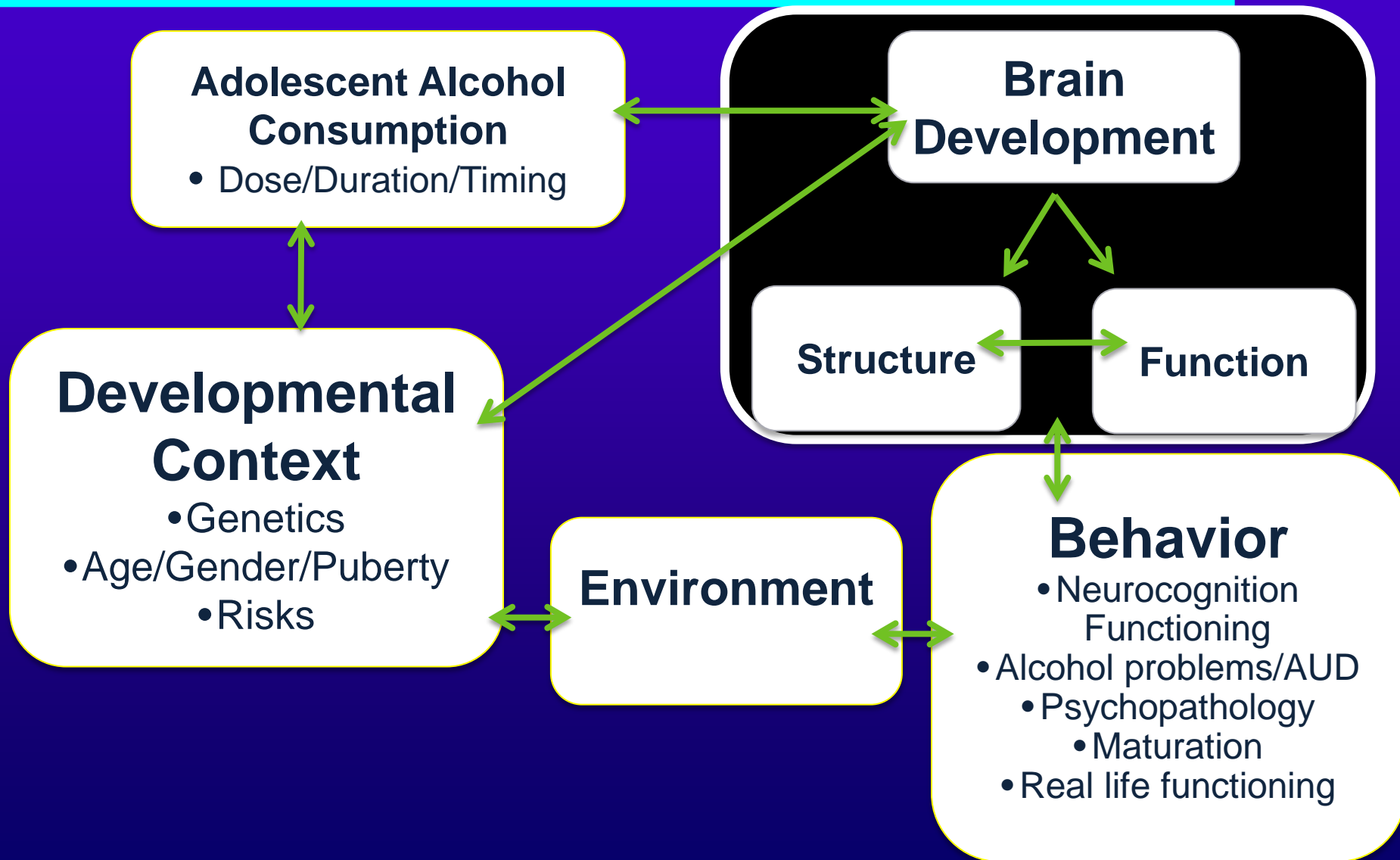
- ◆ Aims
- ◆ Recruitment
- ◆ Protocol
- ◆ Sample characteristics
- ◆ Next directions

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



NCANDA Aims

1. 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
5. Brain features → addiction & psychopathology

NCANDA

Design

Administration:

- Sandy Brown – Coordinator
- Susan Tapert – Scientific Director

Data:

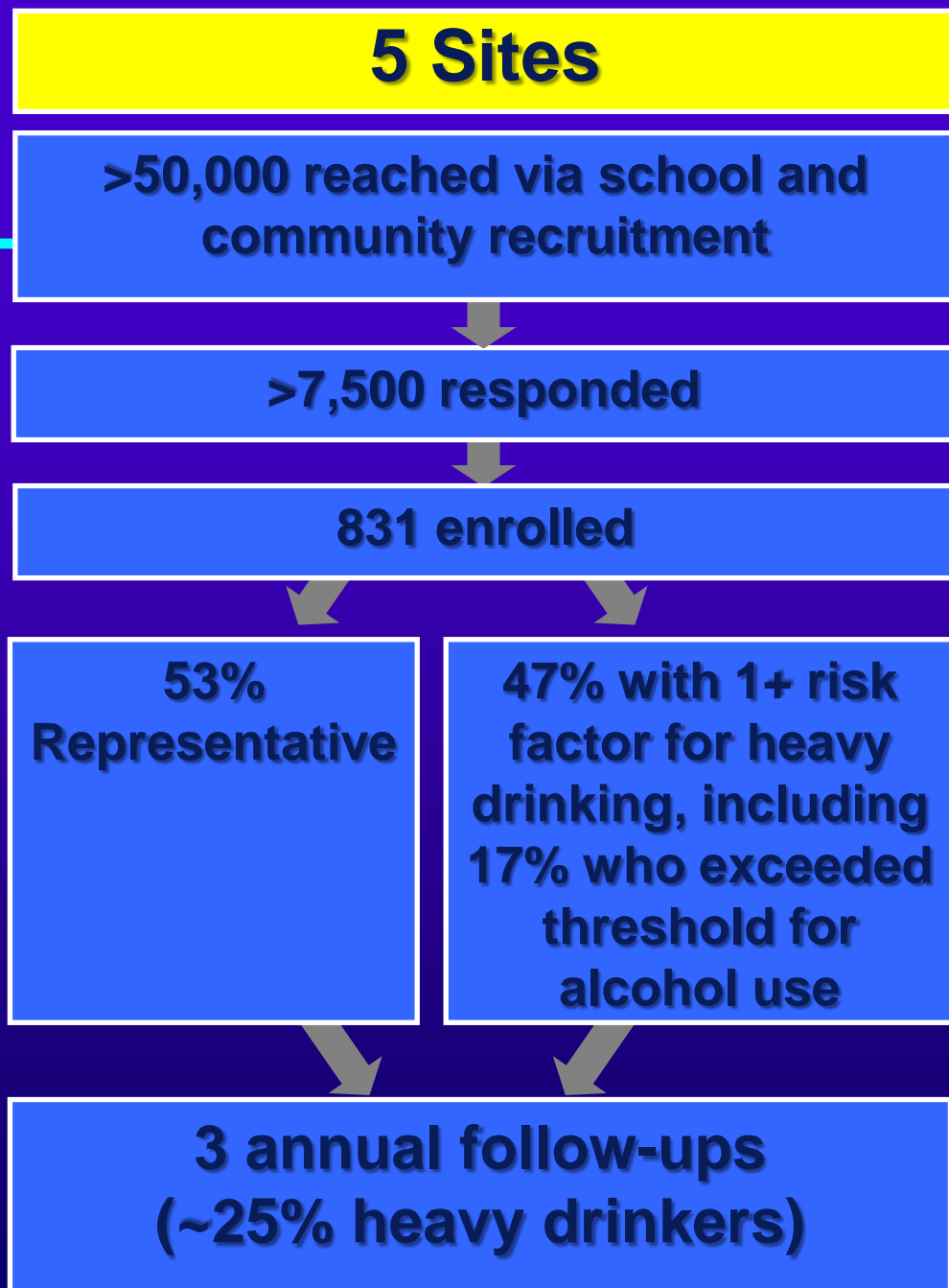
- Dolf Pfefferbaum
- Kilian Pohl
- Edie Sullivan

Sites:

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

Scientific Advisors:

- Ken Sher
- Raquel Gur
- Andrea Hussong
- Arpana Argrawal
- Bob Zucker





Exclusions at Project Entry

- ◆ 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

For classification as Non/Low Drinker:

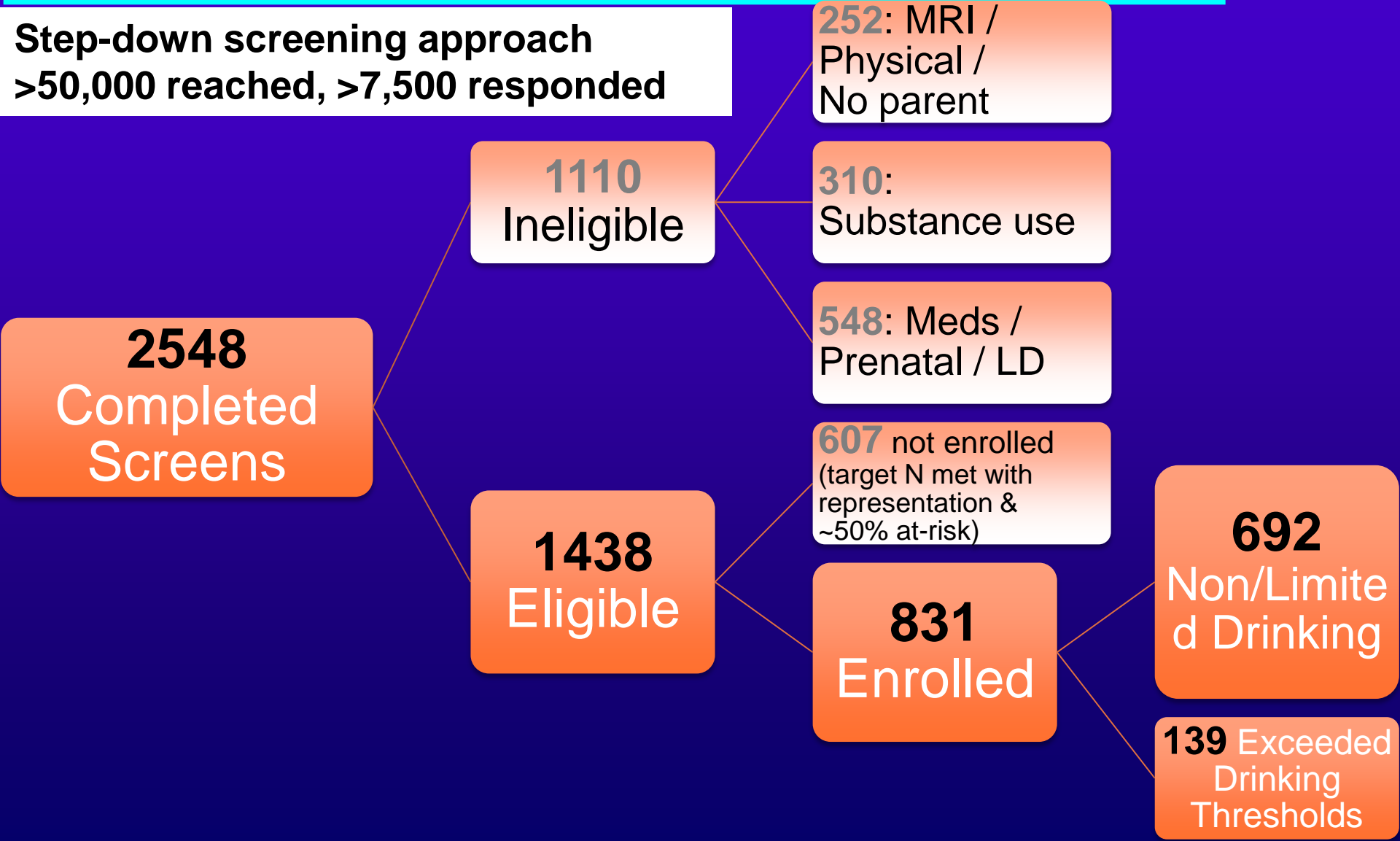
Age	Lifetime Days Drinking ^a	Maximum Drinks on One Occasion ^a		Lifetime Days Cigarette Use ^b	Lifetime Days Marijuana Use ^b	Lifetime Other Drug Use ^b
		Male	Female			
12-12.9	≤5	≤3	≤3	≤10	≤5	≤1
13-13.9	≤5	≤3	≤3	≤10	≤10	≤2
14-14.9	≤5	≤4	≤3	≤20	≤15	≤3
15-15.9	≤5	≤4	≤3	≤30	≤20	≤4
16-16.9	≤11	≤4	≤3	≤40	≤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

^b SAMHSA, 2013

NCANDA Screening

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



1110
Ineligible

252: MRI /
Physical /
No parent

310:
Substance use

548: Meds /
Prenatal / LD

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

2548
Completed
Screens

1438
Eligible

831
Enrolled

692
Non/Limited
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

Other Data Collected:

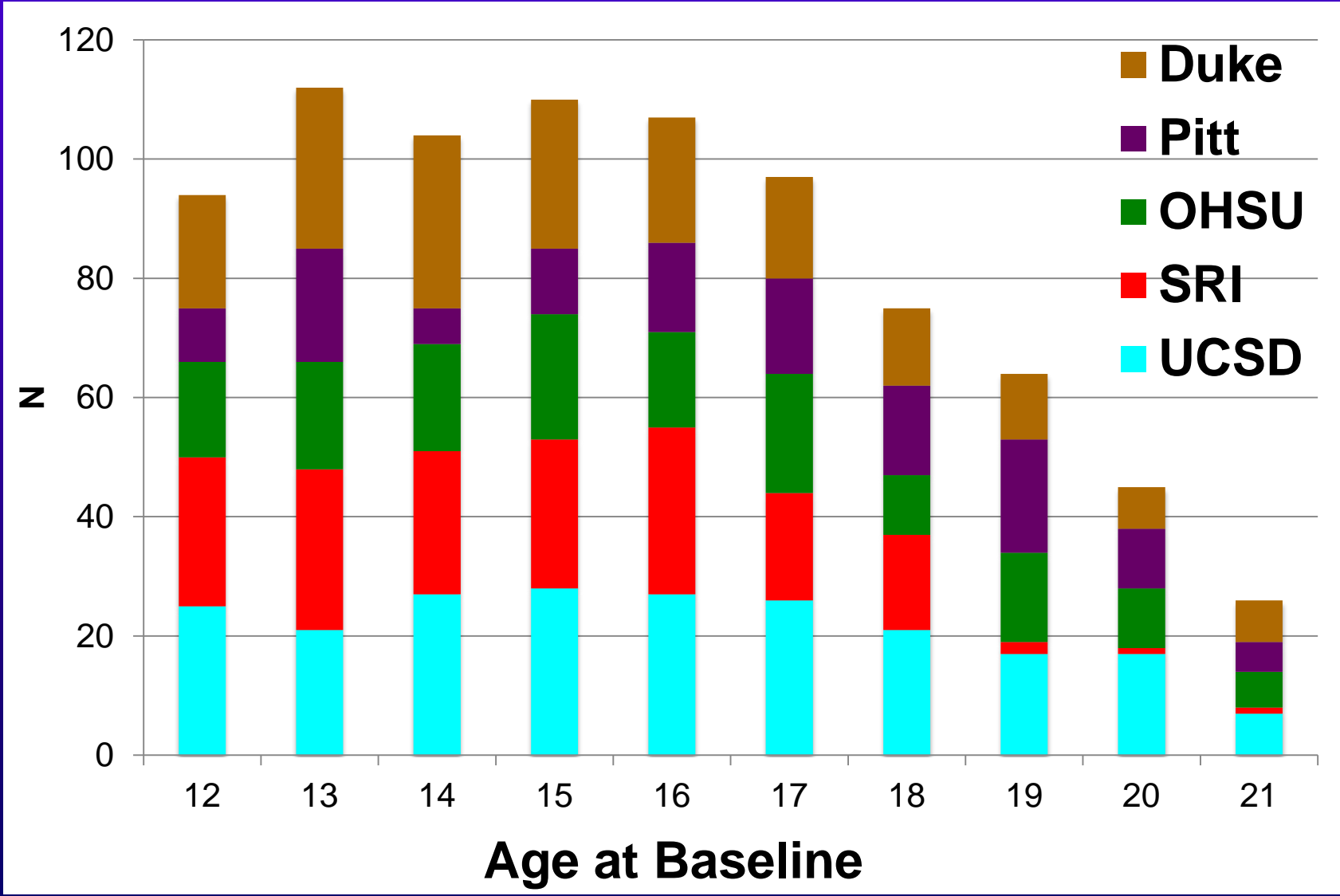
All Sites	MRI: T1, T2, DTI, & resting state fMRI
	Neuropsychological assessment
	Samples for genetic and epigenetic analyses, pubertal hormones, and drug screening
Specialty Projects at 2 Sites	Sleep studies – SRI & Pittsburg
	Stroop fMRI task – SRI & UCSD
	Anti-saccade fMRI task – Duke & Pittsburg
	Recovery protocol – UCSD & Duke



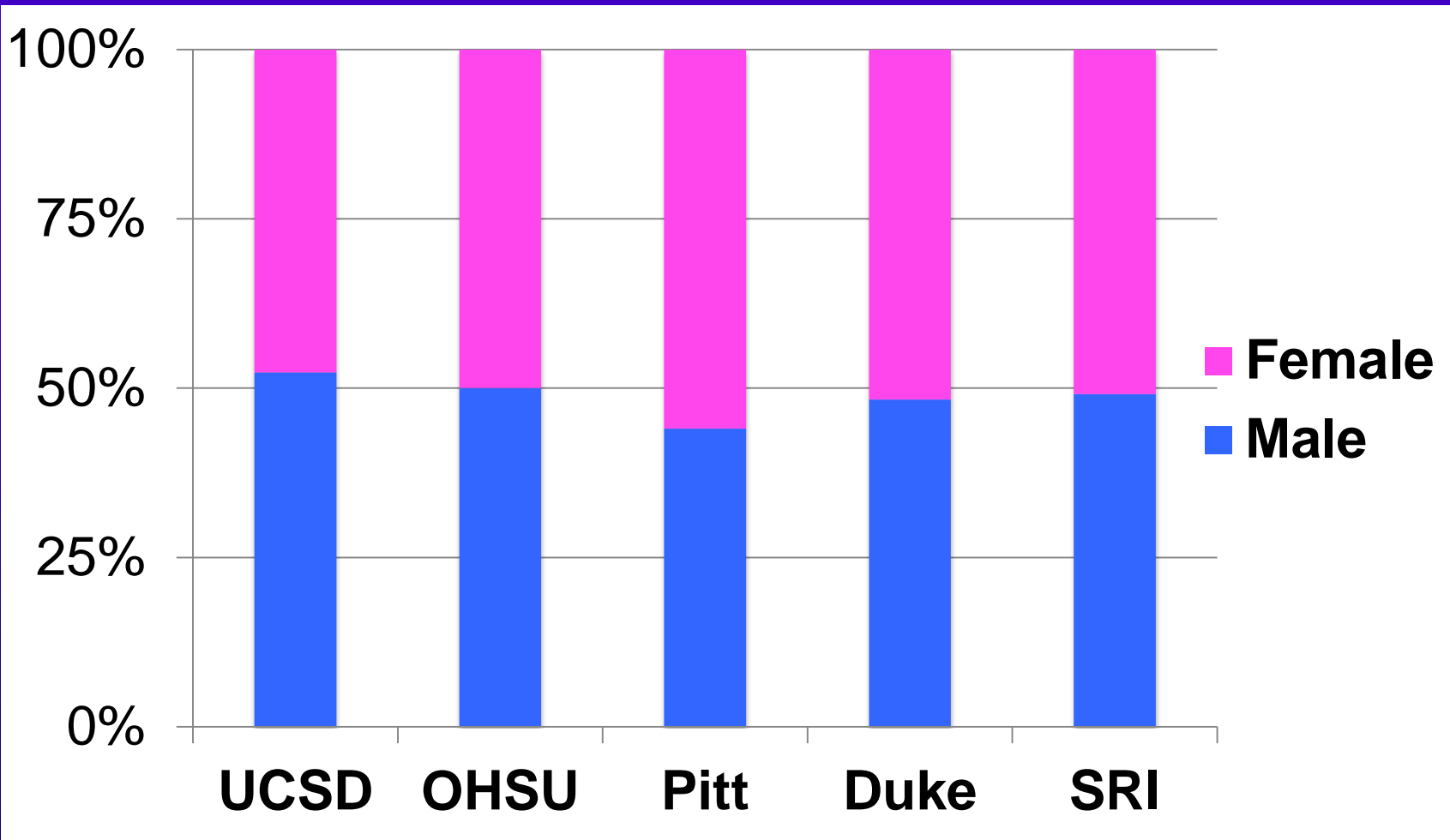
Staff Training & QA

- ◆ 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

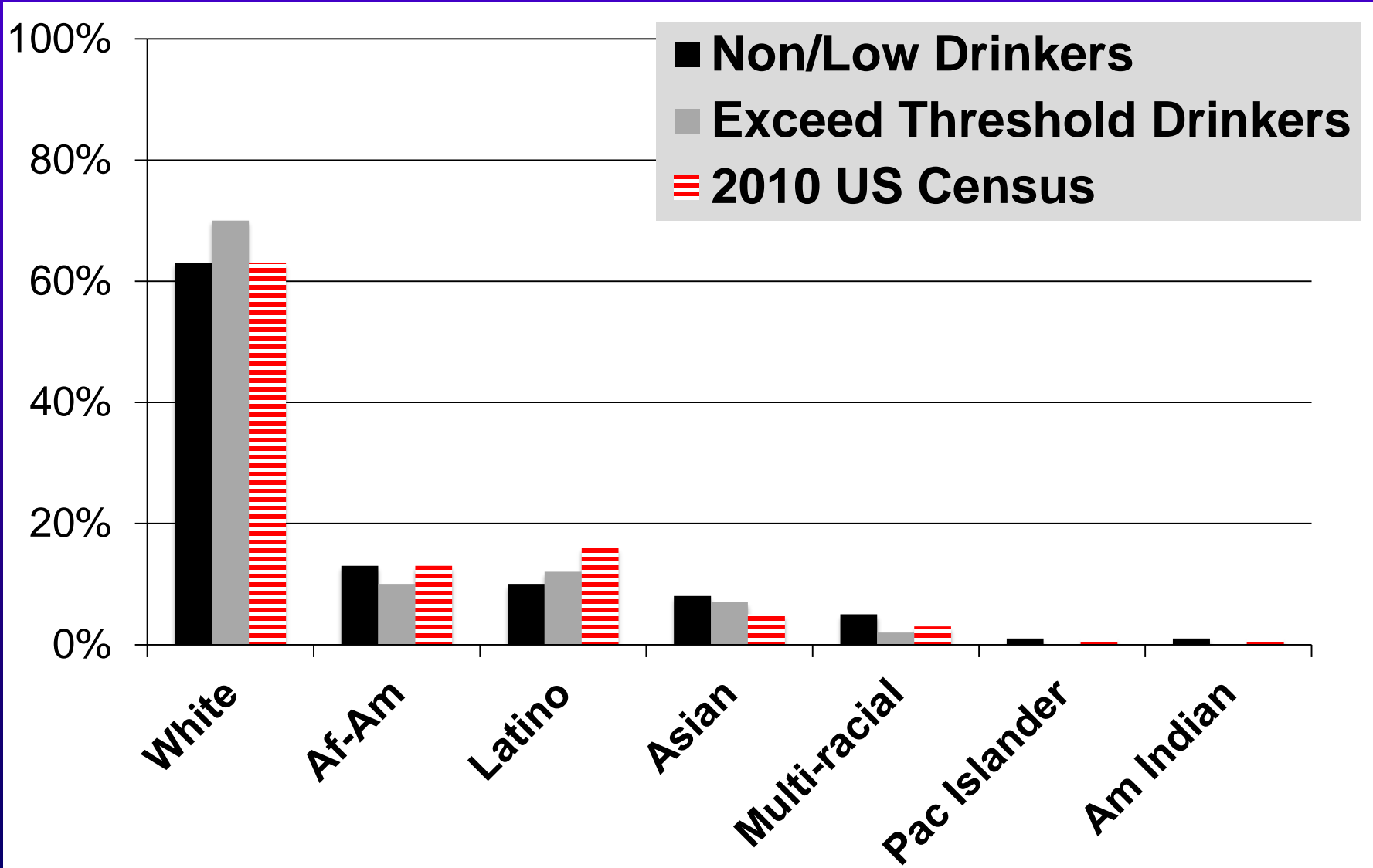
NCANDA Sample: Age Distribution



Balanced Female : Male Ratio



Ethnicity: Nationally Representative





Sample Characteristics

N = 831, age = 12-21 years

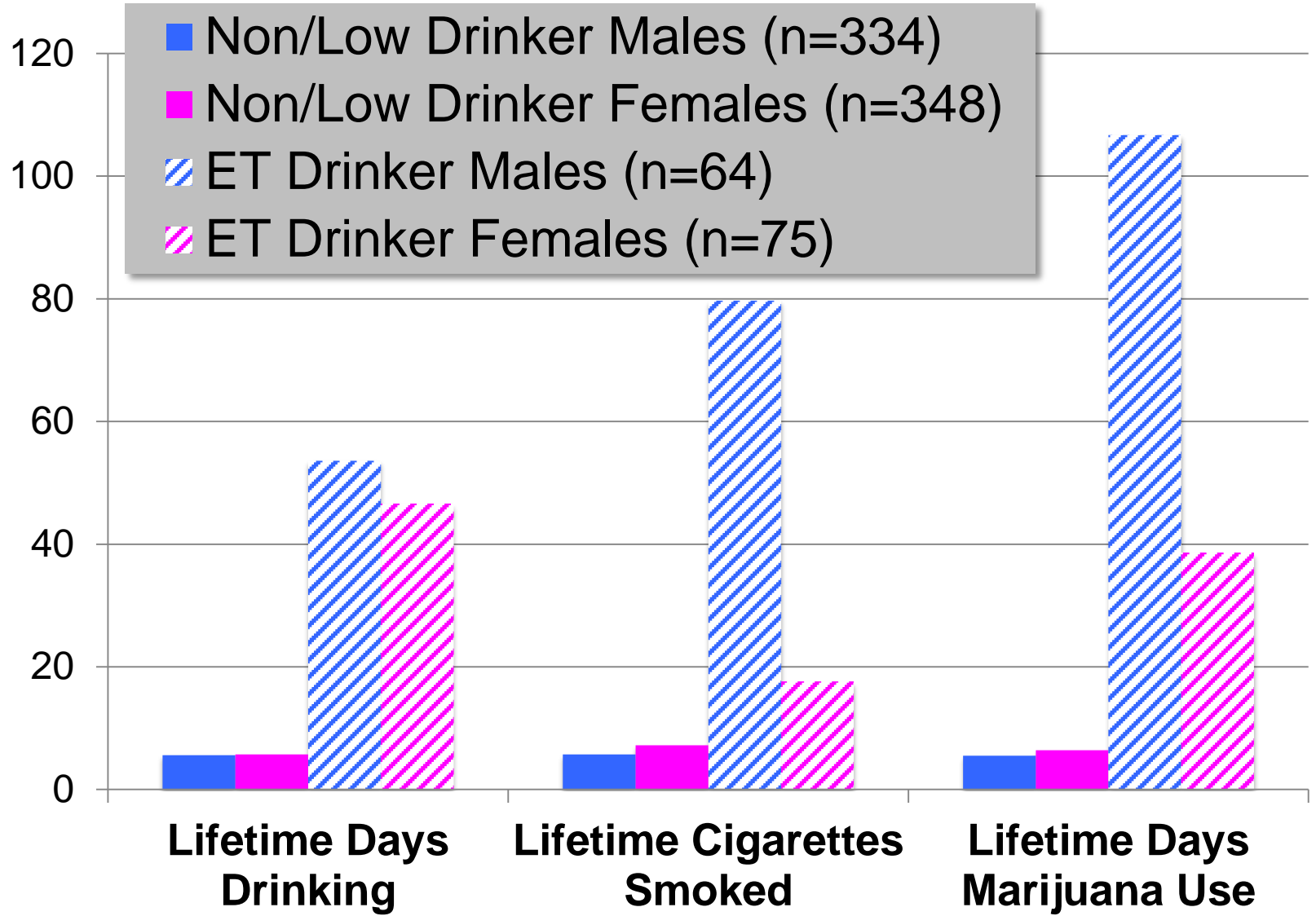
	Non/Low Drinker (N=692)		Exceeded Threshold (N=139)	
	Male (n=334)	Female (n=348)	Male (n=64)	Female (n=75)
Age (mean years)	15.6	15.8	18.4	18.5
Pubertal Development Scale (median)	3.0	3.6	3.6	4.0
Socioeconomic status (mean)	17.0	16.6	17.1	17.3
% Right-handed	76%	83%	83%	81%

Sample Description

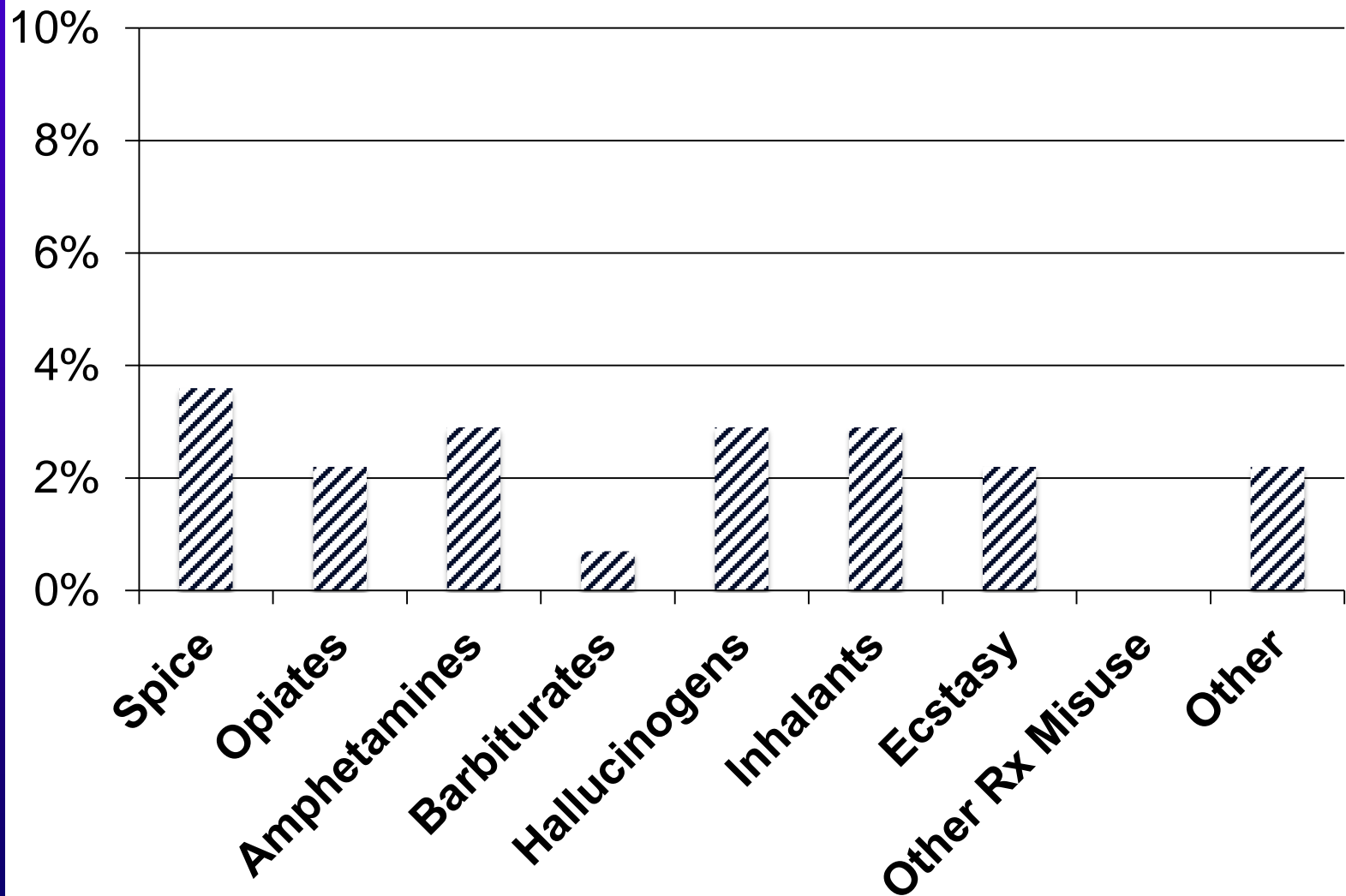
- ◆ 88% reside with one or both parents
- ◆ 27% parents divorced/separated/single
- ◆ 20% below college degree
- ◆ 18% household income <\$50k/yr*
- ◆ 85% normal BMI (13% above, 2% under)

*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)

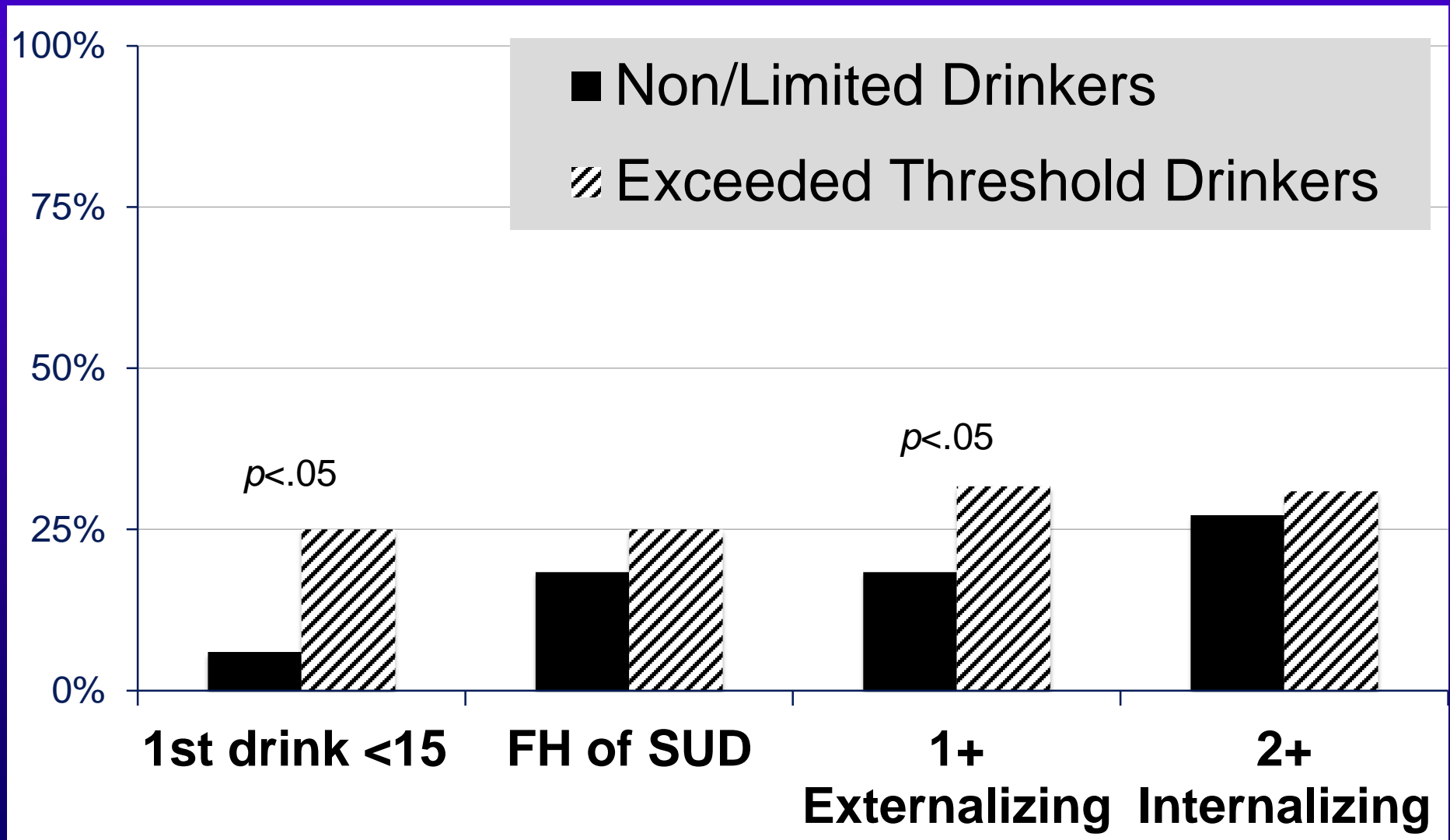
3. 1+ externalizing symptoms

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

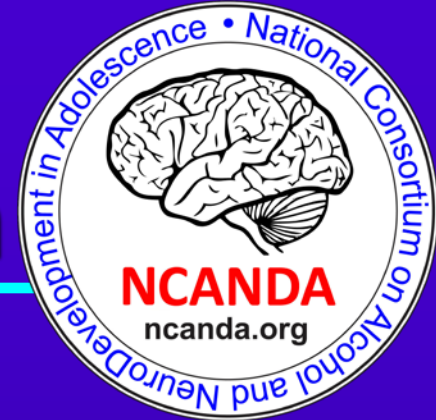
4. 2+ internalizing symptom

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

47% with 1+ Risk Factor for Heavy Drinking



NCANDA Follow-up Design



Baseline



1-year
follow-up



2-year
follow-up



3-year
follow-up



2013

2014

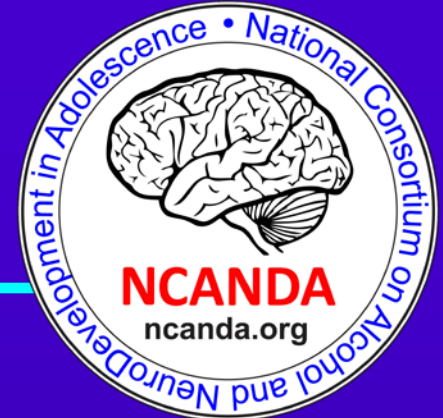
2015

2016

2017

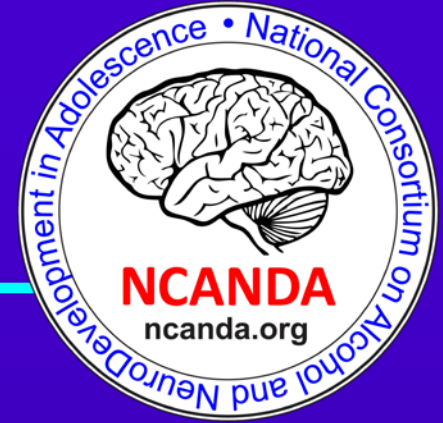
- 4 assessments with imaging + Half-yearly telephone interviews
- 6-month: 98% of 831 done.
- 1-year: in progress, >90% complete.

Next Directions



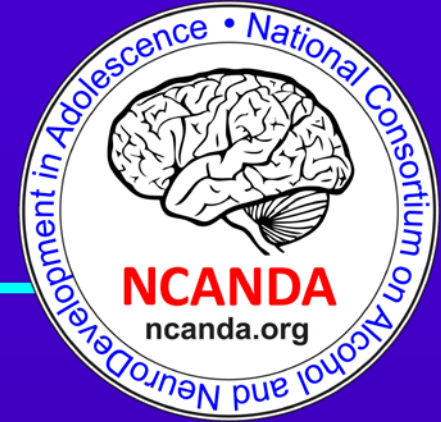
- ◆ **Begin to test Aims 1 and 5:**
 - **Effects of adolescent drinking on trajectory of adolescent brain development**
 - **Neural, cognitive, and affective markers predicting addiction & psychopathology**

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



- ◆ U01 AA021695 (SAB+SFT)
- ◆ U01 AA021697 (AP+KMP)
- ◆ U01 AA021692 (SFT)
- ◆ U01 AA021681 (MDDDB)
- ◆ U01 AA021690 (DBC)
- ◆ U01 AA021691 (BN)
- ◆ U01 AA021696 (IMC+FCB)
- ◆ T32 AA013525 (TB)



National Institute
on Alcohol Abuse
and Alcoholism

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

