Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eTable 1. Individual Ind	dicators of Cognitive Despair and Their Definitions	
Item	Definition	Relation to DSM disorders
Loneliness	Feeling of being alone and/or friendless.	No disorder
Hopelessness	Bleak, negative, pessimistic view of future and little hope that his/her situation will improve.	Dysthymic disorder
Helplessness	Feels there is little or nothing s/he can do to improve his/her situation	No disorder
Low self-esteem	Unjustified feeling of inferiority to others or severe hostility against him/herself.	Dysthymic disorder
Frequent worries	Painful, unpleasant, or uncomfortable thoughts about future, past behavior, competence, calamities, money, appearance that cannot be stopped voluntarily	Generalized anxiety disorder
Feels sorry for self	A feeling that life or people have been unfairly unpleasant or troubling.	No disorder
Feels unloved	Generalized feeling of being unloved and uncared for.	Major depressive disorder

eTable 2. Definition	eTable 2. Definitions of Different Types of Suicidal Thoughts and Behaviors						
Construct	Definition						
Passive ideation	Thoughts about wanting to be dead or death. Include thoughts about not being able to go on any longer and life not being worth living. To code, thoughts must be intrusive into at least two activities.						
Active ideation	Thoughts specifically about killing oneself, by whatever means, with some intention to carry them out. Do not include suicidal plans						
Suicidal Plans	Suicidal thoughts that contain plans of a suicidal act and some intent to carry them out. If suicidal attempt has been made, determine whether a plan was present prior to the attempt.						
Suicide Attempt	Episodes of deliberately self-harmful behavior involving some intention to die at the time of the attempt. Rate here, no matter how unlikely the attempt was to cause death, so long as the child's intention was to die. If parent unsure about intention to die, code if the parent can describe a clear self-harmful event.						

eTable 3. Associa	ations of Lagged D	espair Scores With Y	oung Adult (age 25, 3	0) Outcomes						
		Despair scores								
	Total	0	1	2	3+					
	n (%)	n (%)	n (%)	n (%)	n (%)	P Value				
Total	2424 (100)	1937 (80.5)	275 (12.0)	94 (4.1)	107 (3.5)					
Suicide										
Yes	160 (5.9)	102 (4.2)	23 (9.7)	16 (15.3)	19 (24.1)					
No	2253 (94.1)	1907 (95.8)	247 (90.3)	57 (84.7)	42 (75.9)	0.001				
Alcohol										
disorder										
Yes	132 (7.6)	109 (7.5)	11 (5.8)	7 (3.9)	5 (3.6)					
No	2292 (92.4)	1910 (92.5)	260 (94.2)	66 (96.1)	56 (96.4)	0.13				
Illicit drug use										
Yes	137(4.7)	102 (3.8)	17 (3.7)	6 (9.6)	12 (27.1)					
No	2247 (95.3)	1885 (96.2)	248 (96.3)	66 (90.4)	48 (72.9)	<0.001				
Illicit drug disorder										
Yes	88 (2.9)	68 (2.6)	10 (1.5)	4 (2.7)	6 (9.9)					
No	2336 (97.1)	1951 (97.4)	261 (98.5)	69 (97.3)	55 (90.1)	0.68				
Opioid Use	, ,	, ,	, ,	, ,	, ,					
Yes	55 (2.2)	41 (1.7)	6 (1.0)	2 (6.8)	6 (16.5)					
No	2328 (97.8)	1945 (98.3)	259 (98.9)	70 (93.2)	54 (83.5)	0.002				

Table is based upon 2424 observations of N=1266 individuals. All percentages are weighted and n is unweighted. Despair score is treated as a count variable. Covariates included sex, race/ethnicity, educational status, and poverty. Bolded values are significant at p < 0.05.

eTable 4. Models Predicting Young Adult Outcomes (ages 25 and 30) Comparing Different Levels of Lagged Despair (0 vs 1, 2, 3)								
		0	vs 1	0 vs. 2		0 vs. 3		
Outcome	Predictors	OR	95% CI	OR	95% CI	OR	95% CI	
Suicidality	Despair	1.7	0.8-3.8	1.6	0.3-7.3	3.4	1.2-9.9	
Illicit drug use	Despair	0.9	0.3-2.3	3.0	0.7-12.2	5.6	1.7-18.5	
Opioid Use	Despair	0.7	0.2-2.1	4.3	0.9-21.2	7.2	1.5-35.6	

Table is based on 2424 observations of 1266 individuals. Covariates included sex, race/ethnicity, educational status, poverty. Bolded values are significant at p < 0.05. OR= odds ratio; CI = 95% confidence intervals.

eTable 5. Lagged Longitudinal Models Testing Interactions Between Despair and Each of, Sex, Race/Ethnicity, Poverty, and Educational Attainment in the Prediction of Each Outcome

	Sex	Race/ethnicity	Poverty	Educational
				Attainment
	P Value	P Value	P Value	P Value
Outcome				
Suicide	0.03	0.53	0.01	0.44
Alcohol disorder	0.75	0.11	0.84	0.08
Illicit drug use	0.08	0.03	0.71	0.07
Illicit drug disorder	0.99	0.36	0.90	0.008
Opioid Use	0.09	0.55	0.62	0.13

Bolded values are significant at p < 0.05.

eTable 6. Lagged Models Predicting Young Adult Outcomes Separately by Poverty Status								
		Impoverished	No	ot impoverish	ed			
Outcome	OR	95% CI	р					
Suicide	2.2	1.3-3.5	0.78	0.8	0.5-1.3	0.41		
Alcohol disorder	0.7	0.4-1.3	0.28	0.8	0.5-1.4	0.51		
Illicit drug use	1.7	1.0-3.0	0.06	1.7	0.9-3.2	0.14		
Illicit drug disorder	0.8	0.4-1.6	0.52	1.3	0.8-2.0	0.31		

Poverty status was defined based upon Federal definitions incorporating household size and income.

eTable 7. Lagged Models Predicting Young Adult Outcomes Separately by Educational Attainment							
		Low attainment	Н	igher attainme	ent		
Outcome	OR	95% CI	р				
Suicide	1.5	0.6-3.9	0.38	1.6	1.0-2.4	0.03	
Alcohol disorder	0.5	0.3-1.1	0.07	1.1	0.7-1.7	0.85	
Illicit drug use	1.2	0.6-2.3	0.65	2.6	1.7-4.2	<0.001	
Illicit drug disorder	0.6	0.2-1.6	0.30	1.9	1.3-2.8	<0.001	

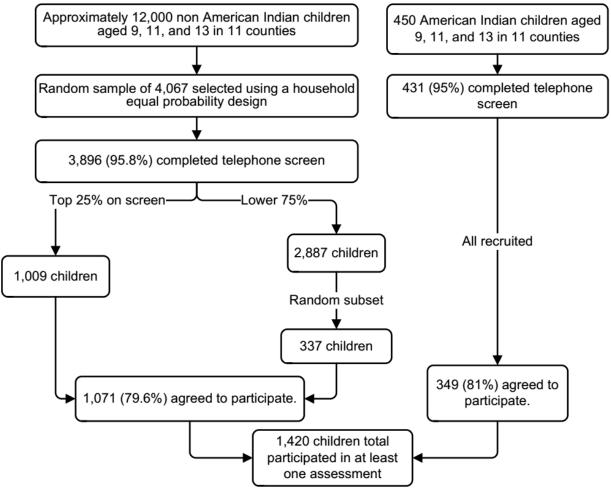
High educational attainment was defined as finishing at least some college; low attainment was defined failing to complete high school or obtaining a high school degree only.

eTable 8. Lagged Models Predicting Young Adult Outcomes Separately by Race/Ethnicity								
		White			lative America	n		
Outcome	OR	95% CI	р					
Suicide	1.3	0.9-1.9	0.12					
Alcohol disorder	0.8	0.5-1.2	0.25	1.6	0.9-2.9	0.10		
Illicit drug use	1.6	1.1-2.4	0.02	0.7	0.4-1.5	0.38		
Illicit drug disorder	1.1	0.6-1.9	0.87	0.8	0.5-1.3	0.81		

Only the comparison between white participants and Native American participants was tested. This table excluded African Americans (n=79) as that group is too small for stand-along comparisons.

eTable 9. Lagged Models Predicting Young Adult Outcomes Separately by Sex								
	Male				Female			
Outcome	OR	95% CI	р					
Suicide	2.3	1.4-3.8	<0.001	1.1	0.7-1.5	0.79		
Alcohol disorder	0.7	0.5-1.2	0.22	0.7	0.3-1.5	0.30		
Illicit drug use	2.2	1.4-3.3	<0.001	1.2	0.7-2.1	0.53		
Illicit drug disorder	1.1	0.5-2.7	0.81	1.0	0.5-1.9	0.95		

eFigure. Ascertainment of the Original Great Smoky Mountains Study Sample



Telephone screen assessed for psychopathology risk.