ABSTRACT: This study tests a socioecological model of relapse and recovery using latent class growth mixture modeling to identify neighborhood, social network and individual-level predictors of alcohol dependence trajectories among a large, longitudinal sample of problem drinkers recruited from substance use treatment settings. We identified four distinct alcohol dependence trajectories: Stable Recovery/Low (Class 1); Relapsing/Rising (Class 2); Late Recovery/Declining (Class 3); and Chronic/High (Class 4). Neighborhood context (poverty and density of bars), social network characteristics (less involvement with Alcoholics Anonymous [AA], continued affiliation with heavy drinkers), and individual predisposing (psychiatric severity) and need (returning to treatment) characteristics each distinguished individuals in the Relapsing/Rising class from individuals in the Stable Recovery/Low class. Social network characteristics (AA involvement and continued affiliation with heavy drinkers) were the primary distinguishing factors for individuals in the Chronic/High class compared to the Late Recovery/Declining class. Study findings can be used to promote recovery and help prevent relapse by: guiding development of community-level interventions to improve social and physical environments; identifying potentially modifiable factors (social network support for sobriety, participation in self-help) to reduce negative consequences among problem drinkers who remain in high-risk neighborhoods; and contributing to ongoing discussions about new and continued licensing of alcohol outlets and regulation of alcohol sales to prevent alcohol problems in high-risk areas and among high-risk people.

KEYWORDS: alcohol dependence, neighborhood environment, social networks, relapse, recovery, socioecological model, longitudinal trajectory

Alcohol problems, including social, legal, and other negative consequences resulting from heavy alcohol use, often are described as part of a chronic, reoccurring condition that requires ongoing care and support. Over the past decade, attention has increasingly turned to understanding and characterizing successful recovery from alcohol problems, with clinical researchers examining factors that contribute to attainment of long-term stability and remission of problems. Definitions of recovery vary, but they often include abstinence from or non-problematic use of alcohol (and perhaps other drugs), as well as an acknowledgement of other dimensions of mental and physical health, well-being, and quality of life. Using data from an existing longitudinal study of alcohol treatment clients supplemented with new information on the neighborhood environment, we develop and test a socioecological model of relapse and recovery from alcohol problems to describe how neighborhood, social network, and individual factors relate to alcohol problems experienced over a seven-year period.

Theoretical framework

Our study is informed by models of relapse prevention and environmental prevention theory. Environmental and community systems prevention focuses on individuals in their environments, which can inform interventions to alter the social, cultural, economic, and physical context to shift conditions away from those that promote relapse to those that support sustained recovery from alcohol problems. In addition to affecting targeted individuals at a given time, these approaches have the added benefit of potentially helping individuals who later enter the improved environment. Similarly, recovery-oriented systems of care emphasize neighborhood-based delivery of services and development of community-based recovery support systems that provide ongoing, integrated services to prevent relapse. These notions are consistent with relapse prevention paradigms encouraging a shift from a model of treatment that includes repeated episodes of acute care toward a model of chronic care focused on longer term recovery management.

We propose a socioecological model that builds on prior epidemiologic models and integrates these perspectives to contextualize relapse and recovery over time. Studies of alcohol use patterns, negative consequences of use, and dependence suggest that neighborhood contexts impact alcohol outcomes, but few studies have examined the longitudinal effects of neighborhoods on recovery from alcohol problems. In our conceptual model (Figure), individuals reside in neighborhood contexts that contribute to the probability of recovery after treatment. These principles also have been capitalized on by the Oxford House movement in efforts to promote housing after substance use disorder treatment in group settings located in safe and stable neighborhoods in proximity to recovery resources. In our model, relevant neighborhood attributes

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include locations and characteristics of substance use disorder treatment facilities (public and private), self-help resources (Alcoholics Anonymous meetings), alcohol outlets (off-premise liquor stores, as well as on-premise bars and restaurants), and socioeconomic resources. These are accompanied by relevant social network (drinkers/drug users in network, sober network members, participation in self-help, and pressure to stop drinking or get treatment) and individual characteristics (demographics, mental health, drug use, and treatment history) that affect long-term outcomes.

Neighborhood factors related to relapse and recovery

Jacobson detailed neighborhood features related to completion of substance use disorder treatment, which also may impact sustained recovery. Neighborhood drug availability can lead to substance use, alcohol dependence, and relapse into heavy drug use and problems. Similarly, living in neighborhoods with easy access to alcohol could increase individuals’ likelihood of relapse, as alcohol outlet density is positively associated with alcohol use and alcohol problems. However, relatively few studies of treatment clients have used individual-level data on alcohol problems coupled with area-based measures of outlet density, much less with longitudinal data. We examine effects of alcohol outlets on trajectories of alcohol problems, and expect a higher density of alcohol outlets will predict relapse into alcohol problems.

Neighborhood socioeconomic disadvantage also is associated with heavy drinking and alcohol problems, and it predicts alcohol use disorders up to 12 years after exposure. Consequences of heavy substance use, including overdose and incarceration, also may be higher for people living in disadvantaged areas. Few longitudinal studies have examined neighborhood SES effects on alcohol problems experienced after someone completes treatment, so we examine effects of neighborhood disadvantage on recovery from alcohol problems. We hypothesize neighborhood disadvantage will predict relapse into problem drinking.

Community resources may impact treatment utilization, self-help involvement, relapse, and recovery from alcohol problems over time. Several studies suggest proximity to mental health and substance use disorder treatment is an important determinant of treatment utilization and quality of care. Alcoholics Anonymous (AA) attendance and involvement also are important predictors of abstinence over time. One study of male veterans who received inpatient treatment found that a city’s number of weekly self-help meetings was positively related to self-help involvement post-treatment; they did not report effects on relapse or recovery. We examine effects of locations and characteristics of substance use disorder treatment facilities and self-help resources on longitudinal trajectories of alcohol problems. We hypothesize that proximity to recovery resources will be positively associated with recovery.
Social network factors related to relapse and recovery

Because it can be difficult to change neighborhoods, we also seek to identify modifiable social network factors that help prevent relapse. Prior work with these data has found that heavy drinkers and drug users in the social network is one of the strongest predictors of higher alcohol volume, continued problem drinking, continued problem drinking, and binge drinking. Although some models of relapse prevention include high-risk situations, these typically have been limited to a person's social network, without also considering the influence of the neighborhood context on an individual's alcohol use and problem trajectories. We did not find any studies comparing effects of social networks and neighborhood contexts jointly influence alcohol use. Understanding the neighborhood and social network context of alcohol problems among heavy and dependent drinkers can inform service planning to provide continuing care after treatment and support long-term recovery.

Individual factors related to relapse and recovery

Key aspects of individuals themselves, such as mental health and co-occurring drug use, also impact relapse and recovery from alcohol problems. As such, our study also is informed by Aday and Andersen’s model of health services utilization. Enabling factors and need characteristics indicating severity of alcohol problems are associated with abstinence outcomes, as well as trajectories of alcohol use in these data, and other research also supports the consideration of these factors in longitudinal studies of relapse and recovery. Individual-level risk factors such as psychological problems may indicate someone has fewer resources to cope with stress, and thus they may relapse or continue to drink heavily, even after completing treatment. No identified studies have examined the role of these individual-level risk factors for relapse in concert with neighborhood and social network characteristics in a longitudinal study of adult treatment clients, however. Our study aims to address these gaps.

Aims of current study

Our primary goals were to characterize neighborhoods that support recovery and identify modifiable social network and individual factors that help prevent relapse of alcohol problems in the seven years after attending alcohol treatment. Capitalizing on unique data available from a large sample of problem and dependent drinkers recruited from treatment centers, we used latent class growth mixture models to describe longitudinal trajectories of alcohol dependence symptoms in a sample of over 700 adults recruited from alcohol treatment programs, and then assessed predictors of class membership to inform a socioecological model of relapse and recovery.

Methods

Data

Participants were from a longitudinal study conducted between 1995 and 2006 that recruited clients meeting criteria for problem drinking (defined below) from 10 public and private detoxification, residential, and outpatient alcohol and drug treatment programs in one Northern California county. Additional information on recruitment and data collection procedures is provided elsewhere. Clients who were at least 18 years of age and gave their written informed consent participated in an in-person structured interview administered by trained interviewers. A total of 926 clients were recruited for the study within three days after entering treatment (80% participation rate). Data on the neighborhood environment were added later (see methods described below).

The initial inclusion criterion required at least two of three problem drinking criteria over the prior 12 months: (1) drinking five or more drinks per day at least once a month for men OR three or more drinks per day at least once a week for women, (2) one or more of eight alcohol-related social consequences in the past year, and/or (3) one or more of nine alcohol dependence symptoms in the past year. These criteria are consistent with prior studies. Follow-up interviews were conducted one, three, five, and seven years after baseline, using computer assisted telephone interviewing; 783 respondents (84.6%) completed at least one follow-up interview. Respondents were tracked every three months using postcard mailings and telephone check-ins.

Participant addresses at each survey timepoint and each interim tracking check-in were geocoded and linked to neighborhod data using ArcGIS 10.2 (ESRI, 2013). For linkage with US Census data on sociodemographic characteristics, neighborhoods were defined by census tract boundaries. For linkage with all other types of neighborhood data (such as alcohol outlets and recovery resources), the geocoded address was used. Values were assigned based on the weighted average across all known addresses during the 12 months prior to each follow-up interview. For the present analysis, due to data availability limitations, eligible geographic areas of residence were restricted to the four Northern California counties which contained the majority of study participants over the follow-up period and for which we had compiled historical data on recovery resources, including 12-step meeting locations and weekly meeting frequency, as well as availability of alcohol/drug treatment. All study procedures were approved by the Institutional Review Boards at the University of California, San Francisco (IRB #11-05204) and the Public Health Institute, Oakland, CA (IRB #199-015 & I11-021e).
The final analytic sample included all participants who had valid neighborhood data at baseline and for at least one follow-up interview (n = 722; 92.2% of those completing one or more follow-up interviews). The analytic sample was 66% male, 54% White and 32% Black/African American (6% Hispanic/Latino and 8% some other race/ethnicity), with an average age at baseline of 38.6 years. Of the respondents in the analytic sample, 76% participated in all four follow-up interviews, 11% completed three interviews, 8% completed two interviews and 5% completed one follow-up after baseline. Men were less likely than women to participate in at least three follow-up interviews (chi-square(df = 1) = 4.55, P = .03) and respondents with more heavy drinkers in their social networks at baseline were marginally less likely to participate in at least three follow-up interviews (t-test = 1.27, P = .10). No significant differences were found on type of treatment program from which the respondent was recruited, baseline 12-step involvement, or baseline neighborhood context (poverty, alcohol availability, 12-step meeting availability or treatment availability) between those who participated in fewer than three follow-ups and those who participated in at least three follow-ups.

Measures

The outcome modeled in the trajectory analyses was alcohol dependence symptoms experienced over the past year as reported at each follow-up interview. This measure was drawn from the eight alcohol-related social consequences and nine alcohol dependence symptoms that were used to establish initial eligibility for the longitudinal study. Social consequences included events such as traffic accidents while drinking and driving, serious arguments with family members or close others about their drinking, and complaints in the workplace about their drinking. Dependence symptoms included desire to cut down/stop drinking, drinking to relieve withdrawal symptoms, drinking interfering with role responsibilities, and blackouts due to heavy drinking. The combination of social problems and physiological symptoms approximates the criteria used in the DSM-5 classification of alcohol use disorder. Scores represented alcohol dependence severity, ranging from 0 to 16 symptoms reported, with an average of 5.59 at baseline, 3.47 at one year, 2.80 at three years, 2.39 at five years, and 2.26 at seven years.

We included predictor variables at the neighborhood level, as well as social network characteristics and individual-level predisposing and need factors (see Tables 1 and 2). Time-varying characteristics were averaged over the follow-up period so they could be entered as predictors of trajectories of alcohol dependence symptoms (described in Analysis section below).

Neighborhood variables were all time-varying, and they included the proportion of residents below poverty, number of bars and pubs within one mile of the respondent’s residence (representing approximately a 10-minute walk; this measure of density was more strongly related to the outcome than either the distance in miles or time in minutes to the nearest bar or pub), number of Alcoholics Anonymous (AA) meetings within 10 miles of the respondent’s residence (weekly meeting total tallied across all meeting sites), inpatient treatment availability (measured by the number of treatment programs within 10 miles of the respondent’s residence, the distance to nearest inpatient program in miles, and the time to the nearest inpatient program in minutes), and outpatient treatment availability (measured by the number of treatment programs within 10 miles of the respondent’s residence, the distance to nearest outpatient program in miles, and the time to the nearest outpatient program in minutes). We also tested a composite measure of neighborhood disadvantage (based on the proportion of residents without a high school diploma, proportion residents over age 16 in blue-collar jobs, percent with income below poverty and proportion without access to a car), as well as measures of off-premise alcohol availability (based on the number of liquor and convenience stores in proximity to the participants’ residential address), but these were not as informative as the measures of neighborhood poverty and on-premise alcohol availability (respectively), so they were dropped from further consideration.

Social network characteristics were heavy drinkers in the network (number of people who supported the respondent’s continued drinking at each wave, which was more strongly related to the outcome than the number of non-drinkers in the network, which was dropped from further consideration), AA involvement (this included meeting attendance and engagement with prescribed behaviors including doing service, having a sponsor, being a sponsor, and reading literature, with composite scores calculated for each wave), and suggestions to seek treatment (from medical, mental health, legal or social welfare service providers in the past 12 months, possible range = 0-4 at each wave). Other measures of pressure to get help—the number of friends and number of family members suggesting respondent seek treatment (coded as 0 compared with at least 1)—were less informative than suggestions from formal service providers, so they were dropped from further consideration.

Time-varying individual-level variables were drug and psychiatric co-morbidities (scores on the Addiction Severity Index or ASI67 for past 30-day drug severity and psychiatric severity at each wave) and receipt of treatment (from either a driving under the influence (DUI) program or alcohol or drug specialty treatment program in the past 12 months; at baseline, this referred to the year prior to the enrollment event). Time-invariant individual characteristics were collected at baseline, including the type of treatment program from which the respondent was recruited (detoxification, inpatient, or outpatient, used as referent), gender (female as referent), age (continuous), race/ethnicity (White compared to Black/African American, Hispanic/Latino, or other), educational attainment (less than high school and more than high school education, with high school diploma or equivalent as referent), marital status (married/partnered compared to all others who were separated/divorced, widowed, and never married), household
income (greater than or equal to $25,000 compared to all others), and early onset of regular alcohol use (15% reported using alcohol 3 or more times per month by age 14, and 26% reported using alcohol 3 or more times per month by age 16).

**Analysis**

We used Mplus version 7.4 to generate latent class growth mixture models to depict trajectories of our outcome measure (count of dependence symptoms and social consequences) in the post-treatment period and to simultaneously determine predictors of class membership. In latent class growth analysis (LCGA), the mixture corresponds to different latent trajectory classes wherein no variation across individuals is allowed within classes, that is, the variance and covariance estimates for the growth factors within each class are assumed to be fixed to zero. The goal is to find classes that differ with respect to their average trajectories, with individual variation around each trajectory regarded as random error. We first fit a series of unconditional latent trajectory models (without covariates) to determine the number of classes that best fit our symptom count outcome. Goodness of fit was determined by comparing the AIC, BIC, and sample-size adjusted BIC indices (lower is better) across models, as well as entropy (.80 desirable) and trajectory shapes and sizes. For these unconditional models, we used data from all 783 cases that had data at baseline and at least one follow-up interview, as the neighborhood-level covariates were not required.

We next fit a model wherein the four-class latent trajectory model was re-fit, now including auxiliary predictor variables. Predictors were allowed to influence the intercept and slope, as well as class membership. It was at this stage that we limited our analysis to the 722 cases that had neighborhood data for at least one follow-up interview. Candidate predictors of class membership were added individually or as conceptual blocks (eg, demographic covariates); those with predictive value were retained in a final model where all predictors were entered simultaneously in the regression model. A potential disadvantage of this approach is that the addition of auxiliary variables may affect the latent class formation, and some individuals may be assigned to a different class. If the classification changes dramatically when covariates are included, further analysis is suggested to determine reasons for changes in class membership. However, this was not the case with our analysis; the classification structure was similar for the unadjusted and adjusted models. All analyses were weighted to account for differential recruitment probability across the treatment program universe in the target county at baseline and to adjust for non-response and drop-out over time.

**Results**

**Trajectory class structure**

We determined that a 4-class solution provided the best fit and overall interpretation. Fit statistics for the unconditional models are displayed in Table 3. Entropy was good in all models. While the 5-class solution provided lower fit statistics, the 4-class solution provided a more parsimonious conceptual representation of the data. In particular, the 5-class solution would have divided the declining class into two smaller groups based on whether declines in alcohol dependence symptoms began three or five years after baseline. Figure 2 displays the unconditional class trajectories for the 4-class model.

Table 4 displays information for the conditional 4-class model. Class sizes ranged from 19% to 34%. We labeled these as low (stable recovery; n = 249), rising (relapsing; n = 156), declining (late recovery; n = 137), and high (chronic; n = 172). See Figure 3. Entropy remained high and similar to that in the unconditional model.

We found some movement of individuals between classes when covariates were included in the conditional joint model. Shapes of the four latent class trajectories, however, remained consistent across models. Compared with the unconditional model, intercept values and slopes varied slightly in the conditional model (see Table 5 notes). The number of drinkers in the social network, returning to treatment, early onset of regular alcohol use, and low
Table 2. Time-varying measures tested as predictors of class membership.

|                              | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|------------------------------|----------|--------------------------|----------------------|--------------------|----------------------|
|                              | MEAN     | LCL 95% CI               | UCL 95% CI           | MEAN   | LCL 95% CI               | UCL 95% CI           | MEAN   | LCL 95% CI               | UCL 95% CI           | MEAN   | LCL 95% CI               | UCL 95% CI           |
| ASI drug severity score      | 0.13     | 0.12 0.14                | 0.05 0.04            | 0.04   | 0.03 0.05                | 0.05 0.05            | 0.05   | 0.04 0.05                | 0.05 0.05            | 0.05   | 0.04 0.05                | 0.05 0.05            |
| ASI psychiatric score        | 0.39     | 0.37 0.41                | 0.21 0.19            | 0.20   | 0.19 0.22                | 0.22 0.24            | 0.21   | 0.19 0.23                | 0.21 0.23            | 0.21   | 0.19 0.23                | 0.21 0.23            |
| Went to treatment in past 12 months, proportiona | 0.70     | 0.67 0.73                | 0.40 0.36            | 0.29   | 0.25 0.32                | 0.22 0.25            | 0.22   | 0.18 0.25                | 0.19 0.23            | 0.19   | 0.15 0.23                | 0.23 0.23            |
| Alcoholics Anonymous Involvement | 1.29     | 1.20 1.37                | 1.68 1.56            | 1.25   | 1.13 1.38                | 1.07 1.20            | 1.13   | 1.00 1.26                | 1.31 1.26            | 1.31   | 0.63 0.63                | 1.26 1.31            |
| Suggestions to get help      | 0.41     | 0.36 0.46                | 0.66 0.59            | 0.36   | 0.31 0.41                | 0.31 0.37            | 0.28   | 0.23 0.33                | 0.26 0.33            | 0.26   | 0.23 0.33                | 0.23 0.33            |
| Drinkers in social network   | 0.31     | 0.11 0.50                | 0.45 0.32            | 0.28   | 0.16 0.40                | 0.14 0.18            | 0.31   | −0.02 0.63               | 0.10 0.18            | 0.31   | −0.02 0.63               | 0.18 0.63            |
| Neighborhood residents below poverty (%) | 12.59     | 11.81 13.38              | 12.47 11.66          | 13.05  | 12.17 13.92              | 12.92 12.05          | 12.86  | 11.90 13.81              | 12.86 13.81          | 12.86  | 11.90 13.81              | 12.86 13.81          |
| Bars and pubs within 1 mile  | 4.64     | 3.54 5.74                | 3.92 2.88            | 5.52   | 3.99 7.04                | 5.19 6.66            | 5.18   | 3.29 7.06                | 5.18 7.06            | 5.18   | 3.29 7.06                | 5.18 7.06            |
| AA meetings each week within 10 miles | 151.14  | 137.75 164.54            | 141.75 127.13        | 195.15  | 174.91 215.39            | 192.10 171.70        | 206.59  | 183.77 229.41            | 206.59 229.41        | 206.59  | 183.77 229.41            | 206.59 229.41        |
| Inpatient treatment programs within 10 miles | 6.59     | 6.12 7.06                | 7.49 6.94            | 6.39   | 5.77 7.00                | 6.65 7.30            | 7.44   | 6.72 8.16                | 7.44 8.16            | 7.44   | 6.72 8.16                | 7.44 8.16            |
| Outpatient treatment programs within 10 miles | 11.48    | 10.91 12.05              | 11.41 10.77          | 10.06  | 9.39 10.73               | 10.33 11.25          | 12.25  | 11.16 13.34              | 12.25 13.34          | 12.25  | 11.16 13.34              | 12.25 13.34          |
| Time to nearest outpatient program (minutes) | 4.16     | 3.84 4.49                | 4.27 3.89            | 4.75   | 4.37 5.12                | 5.03 5.48            | 4.31   | 3.95 4.67                | 4.31 4.67            | 4.31   | 3.95 4.67                | 4.31 4.67            |
| Distance to nearest outpatient program (miles) | 2.12     | 1.94 2.30                | 2.16 1.95            | 2.44   | 2.23 2.65                | 2.70 2.98            | 2.21   | 2.01 2.41                | 2.21 2.41            | 2.21   | 2.01 2.41                | 2.21 2.41            |
| Time to nearest inpatient program (minutes) | 6.58     | 6.05 7.10                | 5.63 5.20            | 9.11   | 8.40 9.82                | 8.23 8.94            | 8.05   | 7.32 8.78                | 8.05 8.78            | 8.05   | 7.32 8.78                | 8.05 8.78            |
| Distance to nearest inpatient program (miles) | 3.67     | 3.32 4.02                | 3.02 2.74            | 5.33   | 4.84 5.81                | 4.90 5.41            | 4.72   | 4.22 5.23                | 4.72 5.23            | 4.72   | 4.22 5.23                | 4.72 5.23            |

aAt baseline, refers to 12 months prior to study enrollment.  
LCL, lower confidence limit; UCL, upper confidence limit; CI, confidence interval.
levels of education were associated with trajectory intercepts, while neighborhood poverty, suggestions to get help and early onset were associated with trajectory slopes over time.

Table 3. Comparing class structure from unconditional LCGA models (N = 783 cases).

| NUMBER OF CLASSES | 1       | 2       | 3       | 4       | 5       |
|-------------------|---------|---------|---------|---------|---------|
| AIC               | 17,960  | 12,866  | 11,964  | 11,310  | 10,974  |
| BIC               | 17,974  | 12,898  | 12,016  | 11,379  | 11,062  |
| ABIC              | 17,965  | 12,876  | 11,980  | 11,332  | 11,002  |
| Log likelihood    | −8977   | −6426   | −5971   | −5640   | −5468   |
| Entropy           | NA      | .929    | .870    | .853    | .836    |
| LRT               | NA      | P < .05 | P < .05 | P > .05 | P > .05 |

AIC, Akaike information criterion; BIC, Bayesian information criterion; ABIC, adjusted Bayesian information criterion; LRT, Lo-Mendell-Rubin likelihood ratio test.

Table 4. Conditional 4-class model (N=722 cases).

| FINAL CLASSIFICATION | NUMBER OF CASES (%) | MEAN OF THE CLASS-MEMBERSHIP PROBABILITIES IN CLASS (%) | MEAN EST. (SE) | INTERCEPT (SE) | SLOPE (SE) |
|----------------------|---------------------|----------------------------------------------------------|----------------|----------------|------------|
|                      |                     | CLASS 1 | CLASS 2 | CLASS 3 | CLASS 4 |                     |               |               |               |
| Class 1 (low)        | 249 (34.4)          | .937    | .039    | .023    | .000    | .046 (.055)         | −1.96 (.824) | −.36 (.496)  |
| Class 2 (rising)     | 165 (22.8)          | .033    | .908    | .031    | .029    | −.087 (.036)        | −1.69 (.580) | .95 (.263)   |
| Class 3 (declining)  | 137 (18.9)          | .022    | .029    | .925    | .024    | −.022 (.033)        | 1.61 (.338)  | −.33 (.217)  |
| Class 4 (high)       | 172 (23.8)          | .000    | .031    | .038    | .930    | .017 (.012)         | 1.39 (.324)  | −.12 (.100)  |

AIC = 9938; BIC = 10396; ABIC = 10078; entropy = .872

Figure 2. Latent classes from 4-class unconditional LCGA models (sample mean scores for alcohol dependence symptoms at each follow-up interview). Proportions in each class: low = 38.5%; declining = 20.7%; rising = 20.5%; high = 20.3%.

Predictors of class membership

Predictors of class membership are shown in Table 5. Compared to individuals in Class 1 (stable recovery, with low levels of alcohol dependence symptoms post-treatment), individuals in Class 2 (relapsing, with rising alcohol dependence symptoms...
Table 5. Associations of predictors with class membership.

| Predictor                          | CLASS #1 (LOW) AS REFERENT | P-VALUE | CLASS #2 (RISING) AS REFERENT | P-VALUE | CLASS #3 (DECLINING) AS REFERENT | P-VALUE |
|-----------------------------------|-----------------------------|---------|-----------------------------|---------|-------------------------------|---------|
| C#2 (rising) ON                   |                             |         |                             |         |                               |         |
| Neighborhood poverty              | 0.043(0.020)                | .031    |                             |         |                               |         |
| Bar/pub density                   | 0.040(0.023)                | .084    |                             |         |                               |         |
| Suggestions to get help           | 1.406(0.690)                | .042    |                             |         |                               |         |
| AA involvement                    | −0.376(0.149)               | .012    |                             |         |                               |         |
| Drinkers in social network        | 0.633(0.294)                | .032    |                             |         |                               |         |
| Returned to alcohol/drug treatment| 1.680(0.757)                | .026    |                             |         |                               |         |
| ASI psychiatric severity          | 2.535(1.101)                | .021    |                             |         |                               |         |
| ASI drug severity                 | −2.823(4.097)               |         |                             |         |                               |         |
| Recruited from detoxification program | 0.771(0.513)            |         |                             |         |                               |         |
| Recruited from inpatient program  | 0.363(0.362)                |         |                             |         |                               |         |
| Early onset of regular drinking   | 0.200(0.368)                |         |                             |         |                               |         |
| Male gender                       | −0.216(0.325)               |         |                             |         |                               |         |
| Income above $25K                 | 0.238(0.333)                |         |                             |         |                               |         |
| Married/partnered                      | 0.052(0.308)              |         |                             |         |                               |         |
| Less than high school              | 0.550(0.412)                |         |                             |         |                               |         |
| More than high school              | 0.037(0.355)                |         |                             |         |                               |         |
| Whitea                             | 0.341(0.346)                |         |                             |         |                               |         |
| C#3 (declining) ON                |                             |         |                             |         |                               |         |
| Neighborhood poverty              | −0.004(0.025)               | −0.047 (0.029) |                             |         |                               |         |
| Bar/pub density                   | 0.055(0.024)                | .020    | 0.014 (0.013)               |         |                               |         |
| Suggestions to get help           | 2.563(0.537)                | .000    | 1.157 (0.723)               |         |                               |         |
| AA involvement                    | −0.401(0.142)               | .005    | −0.025 (0.172)              |         |                               |         |
| Drinkers in social network        | −0.273(0.555)               | −0.906 (0.481) | .060 |                             |         |
| Returned to alcohol/drug treatment| 1.127(0.694)                | −0.554 (0.803) |                             |         |                               |         |
| ASI psychiatric severity          | 0.153(1.516)                | −2.382 (1.658) |                             |         |                               |         |
| ASI drug severity                 | 2.293(4.627)                | 5.117 (3.640) |                             |         |                               |         |
| Recruited from detoxification program | 0.830(0.632)             | 0.059 (0.734) |                             |         |                               |         |
| Recruited from inpatient program  | −0.535(0.404)               | −0.897 (0.453) | .048 |                             |         |
| Early onset of regular drinking   | 0.576(0.396)                |         |                             |         |                               |         |
| Male gender                       | 0.489(0.370)                | 0.705 (0.473) |                             |         |                               |         |
| Income above $25K                 | −0.014(0.368)               | −0.252 (0.385) |                             |         |                               |         |
| Married/partnered                      | 0.221(0.313)              | 0.169 (0.386) |                             |         |                               |         |
| Less than high school              | 0.009(0.426)                | −0.541 (0.543) |                             |         |                               |         |
| More than high school              | −0.192(0.379)               | −0.229 (0.496) |                             |         |                               |         |
| Whitea                             | 0.486(0.408)                | 0.145 (0.496) |                             |         |                               |         |

(Continued)
Table 5. (Continued)

|                                   | CLASS #1 (LOW) AS REFERENT | CLASS #2 (RISING) AS REFERENT | CLASS #3 (DECLINING) AS REFERENT |
|-----------------------------------|-----------------------------|-----------------------------|----------------------------------|
|                                   | ESTIMATE (SE) | P-VALUE       | ESTIMATE (SE) | P-VALUE       | ESTIMATE (SE) | P-VALUE       |
| C#4 (high) ON                     |               |               |               |               |               |               |
| Neighborhood poverty              | 0.038 (0.026) | -0.005 (0.025) | 0.042 (0.032) |               |               |               |
| Bar/pub density                   | 0.044 (0.024) | .064          | 0.004 (0.013) | -0.011 (0.013) |               |               |
| Suggestions to get help           | 3.303 (0.914) | .000          | 1.898 (1.223) | 0.741 (0.807) |               |               |
| AA involvement                    | -0.751 (0.216) | .001          | -0.375 (0.236) | -0.350 (0.210) | .095          |               |
| Drinkers in social network        | 0.706 (0.294) | .016          | 0.073 (0.105) | 0.978 (0.483) | .043          |               |
| Returned to alcohol/drug treatment| 2.061 (1.025) | .044          | 0.381 (1.240) | 0.934 (0.891) |               |               |
| ASI psychiatric severity          | 4.904 (1.582) | .002          | 2.369 (1.658) | 4.751 (1.493) |               |               |
| ASI drug severity                 | 1.352 (4.011) |               | 4.176 (2.784) | -0.941 (3.405) |               |               |
| Recruited from detoxification program | -0.157 (0.756) |               | -0.928 (0.875) | -0.987 (0.733) |               |               |
| Recruited from inpatient program  | -0.305 (0.477) |               | -0.667 (0.453) | 0.230 (0.491) |               |               |
| Early onset of regular drinking   | 0.410 (0.443) |               | 0.209 (0.409) | -0.167 (0.388) |               |               |
| Male gender                       | 0.855 (0.394) | .030          | 1.071 (0.419) | .011          | 0.366 (0.480) |               |
| Income above $25K                 | 0.146 (0.399) |               | -0.092 (0.389) | 0.160 (0.414) |               |               |
| Married/partnered*                | -0.132 (0.380) |               | -0.184 (0.410) | -0.353 (0.388) |               |               |
| Less than high school             | -0.347 (0.638) |               | -0.897 (0.783) | -0.356 (0.622) |               |               |
| More than high school             | 0.079 (0.470) |               | 0.042 (0.557) | 0.271 (0.426) |               |               |
| White*                            | -0.330 (0.456) |               | -0.671 (0.500) | -0.353 (0.388) |               |               |

*Compared to all others.

Notes: Auxiliary variables exerting influence on the latent trajectory intercept (Drinkers in social network, P = .061; Returned to treatment, P = .092; Early onset, P = .070; Less than high school education, P = .066) and slope (Neighborhood poverty, P = .008; Suggestions to get help, P = .086; Early onset, P = .007).

Figure 3. Latent classes from 4-class conditional LCGA models (sample mean scores for alcohol dependence symptoms at each follow-up interview). Proportions in each class: low = 33.5%; declining = 19.9%; rising = 23.3%; high = 23.3%.
Table 6. Time-varying measures retained in latent class growth mixture model as predictive of class membership.

| # Dependence symptoms (LCGA outcome) | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|-------------------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
| Class 1 low                          | 4.0      | 3.4                      | 4.5                  | 0.2                 | 0.1                  | 0.2                 |
| Class 2 rising                        | 5.3      | 4.6                      | 6.0                  | 0.8                 | 0.6                  | 1.0                 |
| Class 3 declining                     | 6.5      | 5.7                      | 7.2                  | 6.4                 | 5.7                  | 7.1                 |
| Class 4 high                          | 8.0      | 7.3                      | 8.7                  | 8.0                 | 7.4                  | 8.6                 |

| ASI drug severity score               | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|--------------------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
| Class 1 low                          | 0.115    | 0.099                    | 0.132                | 0.029               | 0.021                | 0.038               |
| Class 2 rising                        | 0.139    | 0.115                    | 0.164                | 0.033               | 0.021                | 0.045               |
| Class 3 declining                     | 0.110    | 0.088                    | 0.132                | 0.046               | 0.032                | 0.060               |
| Class 4 high                          | 0.135    | 0.115                    | 0.156                | 0.080               | 0.062                | 0.097               |

| ASI psychiatric score                 | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|---------------------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
| Class 1 low                           | 0.326    | 0.295                    | 0.361                | 0.116               | 0.095                | 0.136               |
| Class 2 rising                         | 0.421    | 0.383                    | 0.460                | 0.188               | 0.151                | 0.226               |
| Class 3 declining                      | 0.403    | 0.364                    | 0.442                | 0.174               | 0.130                | 0.219               |
| Class 4 high                           | 0.455    | 0.416                    | 0.494                | 0.348               | 0.309                | 0.386               |

| Suggestions to get help                | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|---------------------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
| Class 1 low                           | 0.27     | 0.20                     | 0.35                 | 0.36                | 0.28                 | 0.44                |
| Class 2 rising                         | 0.55     | 0.41                     | 0.70                 | 0.58                | 0.45                 | 0.71                |
| Class 3 declining                      | 0.43     | 0.30                     | 0.57                 | 0.93                | 0.77                 | 1.09                |
| Class 4 high                           | 0.61     | 0.49                     | 0.72                 | 0.93                | 0.80                 | 1.07                |

| Drinkers in social network            | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|---------------------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
| Class 1 low                           | 0.13     | 0.05                     | 0.22                 | 0.25                | 0.10                 | 0.40                |
| Class 2 rising                         | 0.77     | −0.29                    | 1.83                 | 0.32                | 0.10                 | 0.53                |
| Class 3 declining                      | 0.16     | 0.00                     | 0.33                 | 0.13                | 0.03                 | 0.24                |
| Class 4 high                           | 0.50     | 0.15                     | 0.85                 | 1.12                | 0.68                 | 1.57                |

(Continued)
### Table 6. (Continued)

|                          | BASELINE | ONE YEAR AFTER TREATMENT | THREE YEAR FOLLOW-UP | FIVE YEAR FOLLOW-UP | SEVEN YEAR FOLLOW-UP |
|--------------------------|----------|--------------------------|----------------------|---------------------|----------------------|
|                          | MEAN     | LCL 95% CI               | MEAN     | LCL 95% CI               | MEAN     | LCL 95% CI               | MEAN     | LCL 95% CI               | MEAN     | LCL 95% CI               |
| **Went to treatment in past 12 months, proportion** |          |                          |                      |                     |                      |                      |                      |                      |                      |                      |
| Class 1 low              | 0.61     | 0.55 0.67                | 0.28      | 0.21 0.34                | 0.12     | 0.08 0.16                | 0.10     | 0.05 0.14                | 0.09     | 0.05 0.13                |
| Class 2 rising           | 0.66     | 0.58 0.74                | 0.38      | 0.29 0.47                | 0.31     | 0.22 0.39                | 0.23     | 0.14 0.31                | 0.24     | 0.16 0.33                |
| Class 3 declining        | 0.74     | 0.66 0.82                | 0.50      | 0.40 0.60                | 0.34     | 0.24 0.45                | 0.21     | 0.12 0.29                | 0.16     | 0.08 0.25                |
| Class 4 high             | 0.75     | 0.68 0.82                | 0.55      | 0.46 0.63                | 0.45     | 0.36 0.53                | 0.39     | 0.30 0.48                | 0.35     | 0.25 0.44                |
| **Alcoholic anonymous involvement score** |          |                          |                      |                     |                      |                      |                      |                      |                      |                      |
| Class 1 low              | 1.20     | 1.03 1.37                | 1.86      | 1.64 2.08                | 1.39     | 1.15 1.63                | 1.19     | 0.95 1.43                | 1.06     | 0.83 1.29                |
| Class 2 rising           | 1.47     | 1.26 1.68                | 1.76      | 1.48 2.05                | 1.14     | 0.87 1.41                | 0.99     | 0.71 1.27                | 1.24     | 0.95 1.53                |
| Class 3 declining        | 1.34     | 1.11 1.56                | 1.76      | 1.49 2.03                | 1.48     | 1.16 1.80                | 1.09     | 0.77 1.40                | 1.17     | 0.87 1.48                |
| Class 4 high             | 1.36     | 1.16 1.56                | 1.45      | 1.23 1.68                | 1.19     | 0.94 1.44                | 1.11     | 0.94 1.37                | 1.27     | 1.02 1.53                |
| **Neighborhood % below poverty** |          |                          |                      |                     |                      |                      |                      |                      |                      |                      |
| Class 1 low              | 10.10    | 8.98 11.22               | 10.28     | 9.08 11.47               | 11.10    | 9.77 12.44               | 10.73    | 9.40 12.06               | 10.65    | 9.32 11.98               |
| Class 2 rising           | 13.85    | 12.04 15.67               | 14.32     | 12.39 16.25               | 14.26    | 12.18 16.33               | 15.31    | 13.24 17.38               | 14.44    | 12.16 16.72               |
| Class 3 declining        | 12.08    | 10.21 13.96               | 11.69     | 9.85 13.53               | 12.00    | 9.72 14.28               | 11.43    | 9.31 13.56               | 12.77    | 10.24 15.30               |
| Class 4 high             | 14.10    | 12.35 15.84               | 14.28     | 12.34 16.22               | 15.43    | 13.65 17.20               | 15.32    | 13.49 17.16               | 15.32    | 13.15 17.49               |
| **# Bars and pubs within 1mi** |          |                          |                      |                     |                      |                      |                      |                      |                      |                      |
| Class 1 low              | 2.35     | 1.98 2.71                | 3.35      | 1.41 5.30                | 2.57     | 2.15 3.00                | 2.50     | 1.95 3.06                | 2.47     | 1.87 3.07                |
| Class 2 rising           | 3.33     | 2.73 3.94                | 3.13      | 2.53 3.74                | 4.62     | 2.73 6.51                | 6.89     | 2.48 11.30               | 4.73     | 1.40 8.06                |
| Class 3 declining        | 6.01     | 2.27 9.76                | 5.83      | 1.97 9.68                | 6.68     | 1.63 11.74               | 4.86     | 1.35 8.37                | 9.98     | 1.43 18.53               |
| Class 4 high             | 5.06     | 2.89 7.23                | 4.25      | 2.50 6.00                | 9.25     | 4.49 14.01               | 8.21     | 4.23 12.20               | 6.84     | 2.76 10.93               |

LCL, lower confidence limit; UCL, upper confidence limit; CI, confidence interval.
post-treatment) lived in neighborhoods with higher levels of poverty, had more bars/pubs within one mile of their home (marginally significant), received more suggestions to go to treatment, were less involved in AA, had more heavy drinkers in their social network, returned to treatment, and had higher psychiatric severity. Compared to individuals in Class 1 (stable recovery/low symptoms), individuals in Class 3 (late recovery, with declining alcohol dependence symptoms post-treatment) had more bars/pubs near their homes, received more suggestions to go to treatment, and were less involved in AA, and compared to individuals in Class 2 (relapsing/rising symptoms), those in Class 3 (late recovery/declining) had marginally fewer drinkers in their social network and they were significantly less likely to be recruited from an inpatient program (compared to outpatient programs). Finally, compared to individuals in Class 1 (stable recovery/low symptoms), individuals in Class 4 (chronic, with high alcohol dependence symptoms post-treatment) had more bars/pubs near their homes (marginally significant), received more suggestions to go to treatment, were less involved in AA, had more drinkers in their social network, were more likely to return to treatment, had higher psychiatric severity, and were more likely to be male. Compared to individuals in Class 2 (relapsing/rising symptoms), those in Class 4 (chronic/high) were significantly more likely to be male, and compared to individuals in Class 3 (late recovery/declining) had marginally lower AA involvement and significantly more drinkers in their social network.

Table 6 presents values for the time-varying predictors for each trajectory class. At baseline, individuals in Class 1 (stable recovery/low) had the lowest number of dependence symptoms and those in Class 4 (chronic/high) had the highest number of dependence symptoms; those in Classes 1 (stable recovery/low) and 3 (late recovery/declining) had the lowest drug severity and the lowest number of drinkers in their social networks; and those in Class 1 (stable recovery/low) had the lowest psychiatric severity, received the fewest suggestions to get help, lived in neighborhoods with fewer residents below poverty and had fewer bars/pubs close to their home. As shown in Supplemental Table S1, the patterns of past 30-day abstinence and ASI alcohol severity scores followed similar patterns as the dependence symptoms measure across the four trajectory classes, suggesting validity of the classification.

Discussion

We tested a socioecological model of relapse and recovery using latent class growth mixture modeling to identify predictors of longitudinal alcohol dependence trajectories among a large sample of problem drinkers recruited from substance use treatment programs. Based on the seven-year trajectories, we identified four classes. The groups characterized as Stable Recovery/Low (Class 1) and Relapsing/Rising (Class 2) both started with very few dependence symptoms (less than one) in the year after treatment, but the Relapsing/Rising class showed recurrence of problems by five years after treatment. The groups characterized as Late Recovery/Declining (Class 3) and Chronic/High (Class 4) both started with high levels of dependence symptoms (more than 6) in the year after treatment, but the Late Recovery/Declining class showed evidence of improvement by five years after treatment. Prior analyses of these data showed that respondents’ drinking patterns changed over time, and people either became non-problem drinkers or remained problem drinkers, rather than fluctuating between problem and non-problem drinking. Our findings add to our understanding of long-term outcomes of treatment clients, and these results may inform treatment planning in the future, particularly if they are substantiated by further studies in other diverse samples.

The neighborhood context (poverty and density of bars), social network characteristics (less AA involvement, continued affiliation with heavy drinkers), and individual predisposing (psychiatric severity) and need (returning to treatment) characteristics each distinguished individuals in the Relapsing/Rising class from individuals in the Stable Recovery/Low class. Social network characteristics (AA involvement and continued affiliation with heavy drinkers) were the primary distinguishing factors for individuals in the Chronic/High class compared to the Late Recovery/Declining class. Overall, the individuals in the Stable Recovery/Low class lived in neighborhoods that could better support recovery (fewer residents below poverty and fewer bars/pubs), were in social networks that could help prevent relapse (lowest number of drinkers), and had fewer indicators of complex treatment needs (lowest drug severity, lowest psychiatric severity). Similarly, the individuals in the Late Recovery/Declining class also were in social networks with fewer drinkers, and they had lower drug severity. These results suggest that, particularly for treatment clients living in high-risk neighborhoods, supportive social networks are important to long-term recovery, reiterating earlier findings from an outpatient treatment subsample. Although most treatment programs work to actively engage clients with recovery support resources such as AA after treatment, it is less common that programs address neighborhood risk factors for relapse. Neighborhood-based delivery of services and community-based recovery support systems are key elements in recovery-oriented systems of care to prevent relapse. Without support and engagement in these comprehensive care systems, it is unlikely that clients will be able to make substantial changes in their neighborhood environments on their own.

The individuals in the Chronic/High class were the most complex in terms of co-morbidities (use of other drugs, psychiatric problems), but they also continued to be involved with heavy-drinking social networks and they lived in high-risk neighborhoods. Although they continued to receive suggestions that they should seek help and they repeatedly
re-engaged with the treatment system, their levels of dependence symptoms remained high over time. Without ongoing support to address the neighborhood and social network factors that contribute to alcohol problems, these high-need clients are likely to continue to experience significant symptoms of alcohol dependence for many years, as other studies have shown a strong relationship between serious psychiatric problems and drug use for people living in poverty.84 Despite evidence of effectiveness,85-87 wraparound services, including housing support and other strategies to build recovery capital, are not commonly integrated into drug and alcohol treatment.89 These services may help the most complex and chronic patients recover from their alcohol and other drug problems.

**Study strengths and limitations**

Our study is unique in the longitudinal coverage of alcohol dependence symptoms for seven years after treatment intake. Capitalizing on the large sample, we examined predictors of alcohol problems at the individual, social network, and neighborhood levels, which is another strength of our analysis. However, the results should be interpreted taking some limitations into account. First, we focused on one outcome (alcohol problems in the years after treatment), so further research is needed to examine how neighborhoods and social networks might influence other aspects of successful recovery from alcohol (and drug) problems,89 including mental and physical well-being, developing a sense of purpose and enhancing quality of life, which were not an explicit focus here. There also are some limitations to using the ASI67 to measure drug and psychiatric co-morbidities. For example, although the psychiatric composite score has shown generally high internal consistency across studies, indicators of reliability are lower for the drug composite score, and in some samples, the specificity of the psychiatric score is lower than acceptable.90 Future work could use more detailed measures of mental health co-morbidities to build on our findings. Finally, latent class growth mixture modeling has some noted limitations as well,91 such as an inability to account for within-class variability and a tendency to over-extract mean trajectories with non-normal outcomes.92,93 Thus, our results suggesting that people experience different recovery and relapse trajectories after entering alcohol treatment should be interpreted within these analytic limitations, and future studies should use complementary methods to validate our findings. Qualitative research on elements of recovery among people from these different problem drinking groups over time would be particularly informative.

**Implications**

Despite the aforementioned limitations, our study has practical implications for prevention. Based on the identified risk factors for continued alcohol problems after treatment (exemplified by those in the Chronic/High class) as well as risk factors for relapse after a period of remission (exemplified by those in the Rising/Relapsing class), priorities for intervention should be to increase AA involvement and decrease the number of drinkers in people’s social networks and to provide services to address psychiatric comorbidities. Additionally, to reduce the risk of relapse, it also may be important to include supportive housing services or poverty reduction strategies to accompany treatment programming. Finally, reductions in local bar density or specialized interventions to help clients manage environmental triggers in their home neighborhoods after treatment also may increase the likelihood of long-term recovery.

**Study results also can guide development of effective community-level interventions designed to improve the social and physical environment, which may have synergistic effects that enhance the success of formal substance use disorder treatment.**94 Findings also suggest different intervention approaches may be helpful in high-risk neighborhoods. In terms of treatment, understanding how neighborhoods contribute to alcohol problems after treatment could help health providers and clients become more effective in setting realistic treatment goals and expectations. This also could inform individual treatment planning by identifying and developing strategies to cope with potential neighborhood triggers for relapse. A housing mobility study95 found women who moved from high-poverty neighborhoods to public housing in middle-class neighborhoods often developed new social networks, which is an important step in the recovery process from alcohol problems.48-51

Our evidence about neighborhood triggers for relapse and buffering factors that support recovery can be used by treatment professionals to assist patients as they return home after treatment. Providers could integrate support services for people who want to move to a neighborhood that will support their recovery process, and incentives or ancillary programs could be provided to facilitate housing changes by clients with limited financial resources. We also identified potentially modifiable factors (social network support for sobriety, participation in self-help) that may reduce negative consequences among problem and dependent drinkers who remain in high-risk neighborhoods. Effective interventions such as Making Alcoholics Anonymous Easier (MAAEZ)96 exist to facilitate participation in self-help groups; these could be implemented with treatment programs serving clients from high-risk neighborhoods. Finally, findings will contribute to ongoing discussions about new and continued licensing of alcohol outlets and regulation of alcohol sales to prevent alcohol problems in high-risk areas and among high-risk people.27

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KKJ-J contributed to funding acquisition, data curation, conceptualization, data interpretation, and writing the original draft. JW contributed to conceptualization, formal analysis, data interpretation, and writing the original draft. AAM contributed to data interpretation and writing (review and editing). DLP contributed to data interpretation and writing (review and editing). LAK contributed to funding acquisition, conceptualization, data interpretation and writing (review and editing). All authors materially participated in the work and have approved the final manuscript.

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Supplemental material

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