

Providence St. Joseph Health

Providence St. Joseph Health Digital Commons

Articles, Abstracts, and Reports

4-1-2017

A Randomized Controlled Trial of Ethyl Glucuronide-Based Contingency Management for Outpatients With Co-Occurring Alcohol Use Disorders and Serious Mental Illness.

Michael G McDonell

Emily Leickly

Sterling McPherson

Jordan Skalisky

Debra Srebnik

See next page for additional authors

Follow this and additional works at: <https://digitalcommons.psjhealth.org/publications>



Part of the Behavioral Medicine Commons

Authors

Michael G McDonell, Emily Leickly, Sterling McPherson, Jordan Skalisky, Debra Srebnik, Frank Angelo, Roger Vilaradaga, Jenny R Nepom, John M Roll, and Richard K Ries



Published in final edited form as:

Am J Psychiatry. 2017 April 01; 174(4): 370–377. doi:10.1176/appi.ajp.2016.16050627.

A randomized controlled trial of ethyl glucuronide-based contingency management for outpatients with co-occurring alcohol use disorders and serious mental illness

Michael G. McDonell, Ph.D.^{1,2,3}, Emily Leickly, B.A.^{1,3}, Sterling McPherson, Ph.D.^{2,3,4}, Jordan Skalisky, B.A.^{1,3}, Debra Srebnik, Ph.D.⁵, Frank Angelo, M.A.⁶, Roger Vilardaga, Ph.D.⁵, Jenny R. Nepom, B.S., B.A.^{1,3}, John M. Roll, Ph.D.^{2,3}, and Richard K. Ries, M.D.⁵

¹Initiative for Research and Education to Advance Community Health, Washington State University, Spokane WA

²Elson S. Floyd College of Medicine, Washington State University, Spokane WA

³Program for Excellence in Addiction Research, Washington State University, Spokane WA

⁴Providence Medical Research Center, Providence Health Care, Spokane WA

⁵Department of Psychiatry and Behavioral Sciences, University of Washington School of Medicine, Seattle WA

⁶School of Public Health, University of Washington, Seattle WA

Abstract

Objective—To determine if a contingency management intervention using the ethyl glucuronide (EtG) alcohol biomarker resulted in increased alcohol abstinence in outpatients with co-occurring serious mental illnesses. Secondary objectives were to determine if contingency management was associated with changes in heavy drinking, treatment attendance, drug use, cigarette smoking, psychiatric symptoms, and HIV-risk behavior.

Method—Seventy-nine (37% female, 44% non-white) outpatients with serious mental illness and alcohol dependence receiving treatment as usual completed a 4-week observation period and were randomized to 12-weeks of contingency management for EtG-negative urine samples and addiction treatment attendance, or reinforcement only for study participation. Contingency management included the variable magnitude of reinforcement “prize draw” procedure contingent on EtG-negative samples (<150 ng/mL) three times a week and weekly gift cards for outpatient treatment attendance. Urine EtG, drug test, and self-report outcomes were assessed during the 12-week intervention and 3-month follow-up periods.

Results—Contingency management participants were 3.1 times (95% CI: 2.2, 4.5) more likely to submit an EtG-negative urine test during the 12-week intervention period, attaining nearly 1.5

Correspondence should be addressed to: Michael G. McDonell, Ph.D., Elson S Floyd College of Medicine, PO Box 1495, Washington State University, Spokane, WA 99210-1495, Phone: 509 368-6967.

www.clinicaltrials.gov, NCT01567943

Disclosures: Dr. Ries has been on the speaker bureaus of Janssen and Alkermes. Drs. McPherson and Roll have received research funding from the Bristol Myers Squibb Foundation. All other authors have no disclosures to report.

weeks of additional abstinence relative to controls. Contingency management participants had significantly lower mean EtG levels, reported less drinking and fewer heavy drinking episodes, and were more likely to submit stimulant-negative urine and smoking-negative breath samples, relative to controls. Differences in self-reported alcohol use were maintained at the 3-month follow-up.

Conclusions—This is the first randomized trial utilizing an accurate and validated biomarker (EtG) to demonstrate the efficacy of contingency management for alcohol dependence in outpatients with serious mental illness.

Alcohol misuse is the third leading preventable cause of death in the U.S. and accounts for an estimated \$223.5 billion annually (1, 2), yet 85% of those with an alcohol use disorders never receive treatment (3). Some populations, such as those with serious mental illnesses, including schizophrenia spectrum, bipolar, and major depressive disorders, are at high risk of suffering from alcohol use disorders, with lifetime prevalence as high as 44% (4). Among those with serious mental illness, alcohol use negatively impacts psychiatric symptoms and contributes to high rates of homelessness, psychiatric hospitalization, HIV infection, cigarette smoking, and drug abuse (5, 6). Psychosocial treatments demonstrate reductions in alcohol and drug use in adults with serious mental illness (7, 8); however, few are widely available due to their relatively high costs, need for extensive training and fidelity monitoring, and organizational barriers (9, 10).

Contingency management is an intervention in which reinforcers such as vouchers or prizes are provided, typically multiple times per week, when individuals demonstrate drug abstinence (11). Point-of-care urine drug tests allow for implementation of contingency management because they are inexpensive, provide immediate results, and verify abstinence for two days or more. While contingency management is well-established as an intervention for reducing drug use (11, 12), few studies have investigated contingency management as a treatment for alcohol use disorders (13–15). This is primarily due to the lack of an alcohol biomarker that can verify abstinence for two days or more. Previous studies have used alcohol breath tests, including mobile video recorded breathalyzers, which can only detect use for approximately 12 hours after ingestion (13, 15), or transdermal alcohol sensors, which provide continuous monitoring of alcohol use (14). While these studies have observed reductions in drinking, to date, no randomized trial has investigated the efficacy of contingency management for alcohol use disorders using a biomarker that can verify abstinence for two days or more and can be feasibly conducted in a treatment setting.

Ethyl glucuronide (EtG) is an alcohol metabolite produced by the liver that at low cutoffs (100–200 ng/mL) can be detected in urine for up to five days after drinking (16, 17). EtG tests can be conducted in reference laboratories, in clinical settings using a benchtop analyzer, or, more recently, using a point-of-care dipcard. We have demonstrated that an EtG immunoassay can be conducted accurately at low cutoffs (100–200 ng/mL) using a benchtop analyzer in an outpatient clinic (18) and that EtG can be used to implement a contingency management intervention targeting alcohol (19).

Previous research indicates that contingency management is a feasible and effective intervention for adults with substance use disorders and serious mental illness (20–22). In

the only large trial (N=176) of contingency management for outpatients with drug dependence and serious mental illness, those who received contingency management for stimulant abstinence were 2.4 times more likely to submit a stimulant negative urine sample during the intervention, relative to those in the control condition (22). Contingency management participants also reported higher levels of stimulant abstinence during a three month follow-up and had lower levels of alcohol use, cigarette smoking, psychiatric symptoms, and inpatient psychiatric hospitalizations, relative to controls (22, 23). A cost-benefit analysis indicated that these clinical benefits offset the costs of the intervention (24). Despite these positive findings, a weakness of this study was a contingency management attrition rate of 41%.

The objective of this study was to determine if a contingency management intervention that provided reinforcers for alcohol abstinence assessed by EtG, and reinforcers for addiction treatment attendance resulted in increased alcohol abstinence in adults with alcohol use disorders and serious mental illness receiving treatment as usual in a community mental health center. Secondary outcomes were other measures of alcohol use, addiction treatment attendance, drug use, psychiatric symptoms, cigarette smoking, and HIV-risk behavior. A pre-randomization induction period was used, along with other strategies, to reduce attrition in the randomized sample. To our knowledge, this is the first adequately powered randomized trial of contingency management for alcohol use disorders using an appropriate alcohol biomarker and the second of contingency management in serious mental illness among adults.

METHOD

Participants

Participants were recruited from a multisite community mental health and addiction treatment agency in Seattle, WA, and met DSM-IV-TR criteria for alcohol dependence, and schizophrenia, schizoaffective disorder, bipolar I or II, or recurrent major depressive disorder assessed by the *Mini-International Neuropsychiatric Interview*. Other eligibility criteria were alcohol use on >4 of the last 30 days and enrollment in outpatient addiction group treatment at the agency. Exclusions were comorbid drug dependence, except nicotine, or medical or psychiatric severity that would compromise safe participation.

As described in the consort diagram (see supplemental figure), 237 individuals were screened for eligibility. Of the 121 eligible participants, 84 successfully completed the 4-week induction period (described below). Five of these individuals were allocated to the contingency management intervention to set the initial magnitude of reinforcement for the non-contingent condition and were not included in analyses. Seventy-nine participants were randomized to intervention groups and comprise the intent-to-treat sample. Participants provided written informed consent and procedures were approved by the University of Washington's Human Subjects Division.

Design and Procedure

This study employed a 4-week induction observation period (weeks 1–4), designed to increase retention after randomization, followed by a 12-week randomized controlled trial of contingency management (weeks 5–16), including a qualitative interview at week 16, with a 3-month follow-up (weeks 17–28). Participants received treatment-as-usual throughout the study and were randomized to contingency management, where they received reinforcers for EtG-negative urine samples and addiction treatment attendance (n=40) or the non-contingent control group (non-contingent; n=39), in which they received reinforcers regardless of EtG results and treatment attendance. The Urn-randomization procedure was used to allocate participants to intervention groups balanced on mood vs. psychotic disorder, level of alcohol use 30 days before baseline, and drug use 30 days before baseline. Randomized participants completed monthly follow-up interviews for three months.

Measures

At the baseline interview, participants completed the *Mini-International Neuropsychiatric Interview* to assess for alcohol use, drug use, mood and psychotic disorders, and an assessment of demographics. At baseline and weeks 4, 8, 12, 16, 20, 24, and 28, participants completed the *Addiction Severity Index-Lite* to assess days of alcohol use, drinking to intoxication and drug use, the *Clinical Monitoring Form*, *Brief Symptom Inventory*, and *Positive and Negative Syndrome Scale for Schizophrenia* to assess psychiatric symptom severity, the *HIV Risk-Taking Behavior Scale*, and the *Fagerström Test for Nicotine Dependence*. Participants received \$20–\$30 for these interviews.

Participants provided urine and breath samples three times per week during the induction and intervention, and once every four weeks during follow-up. The primary alcohol use outcome was urine EtG level as measured by the Diagnostic Reagents Incorporated EtG enzyme immunoassay conducted onsite using a Thermo Fisher Indiko analyzer (Fremont, CA). Samples were considered alcohol negative if EtG < 150 ng/mL (16, 17). Tests were conducted using EtG 100 ng/mL, 500 ng/mL, 1000 ng/mL, 2000 ng/mL, and Negative calibrators and EtG 100 ng/mL and 375 ng/mL controls. Calibrations occurred weekly. Samples were analyzed on the day of collection. Participants were instructed to avoid non-beverage sources of ethanol, such as mouthwash and cough syrup. Participants were escorted to the bathroom and samples were checked for appropriate temperature after collection.

Urine samples were tested for amphetamine, methamphetamine, cocaine, marijuana, and opiate use (Integrated E-Z Split Key Cup II, Innovacon, Inc, San Diego, CA). Breath samples were also tested for carbon monoxide (CO) analysis (Bedfont-Smokerlyzer Micro-IV) (smoking positive CO 3ppm) (25).

At every study visit the Alcohol Timeline Followback method (26) was used to assess hours since the last drinking episode and the number of standard drinks consumed at the last drinking episode in the prior five days. These data were used to assess the presence of heavy drinking (women > 3 standard drinks, men > 4 standard drinks) during the prior five days.

Participants completed a brief qualitative interview assessing the feasibility and effectiveness of the study at week 16.

Study Interventions

Treatment as Usual—Participants were required to be enrolled in state-certified outpatient addiction treatment where they received the same curriculum during group counseling sessions two to four times weekly for twelve weeks or more. This included psychoeducation about the causes of addiction, relapse prevention, and other relevant topics, such as wellness. Participants could receive mental health care which involved meeting with a case manager once per week and psychiatric medication management, group treatment, and housing and vocational services as appropriate. Self-reported pre-randomization utilization of addiction treatment as usual care did not vary by group.

Induction period—After the baseline interview, participants entered a 4-week induction where they received reinforcers contingent on providing urine samples three times per week regardless of EtG results. Participants received three “prize draws” (described below) for each urine sample provided. The induction period has been used in previous studies to decrease attrition in the randomized sample by selecting for randomization only those who can attend brief study appointments and are currently using alcohol (27, 28). Participants who attended at least one visit during the fourth week and provided at least one EtG-positive urine sample at any time during induction were randomized. Individuals who did not initially meet criteria for randomization were allowed to re-enroll in the study. Re-enrolled individuals completed a second baseline and induction period. Ten participants were re-enrolled and six were randomized.

Contingency Management—Participants in contingency management received the variable magnitude of reinforcement procedure each time they tested negative for EtG. This well-researched procedure involves making “prize draws” from a container of tokens representing different magnitudes of reinforcement (11). Fifty percent read “good job” (no prize). The other 50% were associated with a tangible prize (41.8% read “small” \$1.00 value, 8.0% read “large” \$20.00 value, and 0.2% read “jumbo” \$80.00 value).

Participants provided urine samples three times per week for 12 weeks. They earned at least three prize draws for each EtG-negative urine sample submitted (EtG < 150 ng/mL). One additional prize draw was earned for each week of continuous alcohol abstinence (three consecutive EtG-negative samples). Participants who missed an appointment or had an EtG-positive sample did not earn prize draws and had a “reset” to three prize draws when their next negative sample was submitted. Following a reset, participants could return to the number of prize draws at which the reset occurred by providing three consecutive EtG-negative samples. Participants received gift cards for attending all (\$10) or at least one (\$5) of their addiction groups each week.

Non-Contingent Reinforcement—Non-contingent participants submitted urine samples three times a week for 12 weeks, but received prize draws for each urine sample submitted, regardless of EtG results. Consistent with previous studies (22, 29), the number of prize

draws they received was equal to the average number of prize draws earned by the contingency management group in the previous week. This procedure assured that groups received an equal number of prize draws. Non-contingent participants did not need to attend addiction treatment to earn gift cards; instead they received gift cards equal to those earned by the contingency management group during the previous week.

Reinforcers and Earnings—The total average values of prizes earned by the non-contingent (Mean=\$147.54, SD=\$143.19) and contingency management (Mean=\$175.03, SD=\$183.17) groups were not significantly different.

Attrition—Individuals who missed nine consecutive study visits (3 weeks) during the intervention phase were considered dropouts. These individuals could complete follow-up interviews and were included in the intent-to-treat analyses.

Data Analysis

We used chi-square tests for categorical variables and *t* tests for continuous variables to examine baseline differences across groups. Analyses were conducted on the intent-to-treat sample. Generalized estimation equations were used to analyze outcomes that were collected over time. The maximum likelihood estimation procedure was used (30). To protect against Type-I error, analyses utilized bi-directional tests despite our uni-directional hypotheses. Odds ratios with 95% confidence intervals are presented for binary outcomes, and unstandardized regression coefficients with 95% confidence intervals are presented for continuous outcomes. These methods of analysis have been used in many contingency management trials and are effective and efficient methods of analyzing outcomes across time nested within participants (22, 29). Analyses controlled for baseline levels of the outcome being analyzed.

Multiple imputation procedures were used to handle missing data. This approach has advantages over single imputation, listwise deletion, or other techniques (31, 32) in conjunction with generalized estimation equation analyses. We present listwise deletion analyses first and include the multiple imputation analyses in Table 3 to provide confidence bounds for the location of the true treatment effect. Preliminary analyses identified several variables that predicted missingness due to intervention dropout. We used these variables during the imputation phase to ensure our ‘missing at random’ assumption was tenable. While there is no test for whether missing data are truly ‘missing at random’ versus ‘missing not at random,’ our inclusive strategy for auxiliary variables during imputation made for a tenable assumption that data were ‘missing at random.’ Multiple imputation procedures used a regression-based approach to fill in the missing values to produce multiple datasets. Fifty datasets were analyzed for each analysis to maximize statistical efficiency. Parameters and standard errors were combined using Rubin’s rules (33). Analyses were performed using Stata 14.1 (StataCorp, College Station, TX).

RESULTS

Twenty-six (65%) contingency management and 29 (74%) non-contingent participants completed the intervention phase, a non-significant difference. Thirty (75%) of the

contingency management participants and 30 (77%) of the non-contingent participants attended at least one of three follow-up assessments and 42 (53%) participants attended all follow-up assessments. Demographic and clinical characteristics of the sample did not differ by group (see Table 1).

EtG and Self-Reported Alcohol Use

As Figure 2 illustrates, individuals randomized to contingency management had a mean longest duration of alcohol abstinence (number of consecutive EtG-negative urine samples) more than twice as long as controls (contingency management: Mean=8.56, SD=12.59; non-contingent: Mean=4.11, SD=1.22; $F(3)=5.55$, $df=1, 76$, $p<0.05$), with approximately 1.5 weeks of additional abstinence. Contingency management participants were 3.13 times (95% CI: 2.18, 4.50, $p<0.05$) more likely to submit an EtG-negative urine sample during the intervention period relative to controls (see Figure 1). Contingency management participants had significantly lower EtG levels during the intervention period relative to controls ($B=325.93$; 95% CI: 213.35, 438.51, $p<0.05$). While group differences on both the binary and continuous versions of EtG persisted into follow-up (see Table 2), these differences were non-significant. Multiple imputation analyses verified results of EtG analyses during intervention and follow-up (Table 3).

Contingency management participants had significantly fewer days of any drinking ($B=8.29$; 95% CI: 3.97, 12.60, $p<0.05$) and drinking to intoxication ($B=6.43$; 95% CI: 2.40, 10.47, $p<0.05$) throughout the intervention period, relative to controls (see Table 2). They were 3.48 times (Odds ratio=3.48; 95% CI: 2.32, 5.23, $p<0.05$) less likely to report recent heavy drinking during the intervention period, relative to controls. These findings persisted into the follow-up period (days of drinking, $B=7.40$; 95% CI: 3.52, 11.29, $p<0.05$, days of intoxication, $B=5.69$; 95% CI: 2.57, 8.80, $p<0.05$, heavy drinking in last 5 days, Odds Ratio=4.90; 95% CI: 1.78, 13.47, $p<0.05$). These differences were significant using the multiple imputations during treatment and follow-up (Table 3).

Secondary Outcomes

Contingency management participants were more likely to submit stimulant-negative urine tests during the intervention (Odds ratio=3.19, 95% CI: 1.99, 5.10, $p<0.05$) and follow-up periods (Odds ratio=4.59, 95% CI: 1.24, 17.08, $p<0.05$), and self-reported fewer days of stimulant drug use during the intervention period ($B=2.22$, 95% CI: 0.34, 4.10, $p<0.05$). Multiple imputation analyses found similar results for stimulant drug tests (Table 3). The multiple imputation analyses replicated the effect size of self-reported days of stimulant drug use, but the treatment effect for the intervention period was not significant (Table 3).

Among the 54 smokers identified by smoking-positive self-report or carbon monoxide samples at baseline, contingency management participants were 5.4 times more likely to submit a smoking-negative breath sample, relative to controls during the intervention period (Odds ratio=5.39, 95% CI: 1.93, 15.0, $p<0.05$). The multiple imputation model represented an attenuation of these findings (Table 3). This difference was not significant during the follow-up period. Group differences on other secondary outcomes including addiction

treatment attendance, opioid and marijuana use, psychiatric symptoms, HIV risk behavior, and self-reported cigarette smoking were not statistically significant.

Patient Perspective

The participant was a 52 year-old African-American man diagnosed with major depressive disorder. He wanted to stop using alcohol but struggled to stop drinking due to his unpredictable housing situation. He was abstinent for 11 weeks during the contingency management intervention and earned over \$700 in gift cards and prizes. He expressed appreciation for the small pragmatic prizes, such as laundry detergent, toothpaste and deodorant, because they lessened his day-to-day worries and expenses. By the end of the intervention, he obtained sober housing and was regularly attending outpatient treatment.

DISCUSSION

This is the first adequately powered randomized trial of contingency management for alcohol use disorders utilizing an accurate alcohol biomarker. Those who received contingency management were three times more likely to submit an EtG-negative urine sample during the intervention period, and they attained approximately 1.5 weeks of additional abstinence, relative to controls. Those who were assigned to contingency management had an average EtG level that was approximately 300 ng/mL lower than controls during the intervention period. These differences are large and clinically significant. Self-report data corroborated EtG results, with contingency management participants reporting fewer days of alcohol use, drinking to intoxication, and episodes of heavy drinking during the intervention and follow-up periods. Results are consistent with reductions in drinking during contingency management (13–15), as well as during follow-up (14) reported by others. This study provides further evidence of post-intervention benefits of contingency management for alcohol use (14).

This is the second trial to investigate the efficacy of a contingency management intervention focused on promoting abstinence from alcohol or drug use in adults with serious mental illness (22). Both trials demonstrated that contingency management is a powerful procedure for reducing substance use when added to treatment as usual at a community mental health center. This study also replicates our previous findings of group differences in non-targeted drug use and cigarette smoking (22, 23). We did not observe group differences in other outcomes, such as psychiatric functioning or HIV-risk behavior. However, this could be due to a lack of statistical power, given the relatively small sample size.

The contingency management completion rate (65%) was superior to our previous study in adults with serious mental illness and stimulant dependence (41%) (22) and is comparable, if not superior, to contingency management studies in populations without serious mental illness (29). We implemented a number of strategies to reduce attrition including requiring participants be in treatment as usual, increasing the minimum number of prize draws for each abstinent urine sample from one to three, and utilizing a four-week induction period prior to randomization. While differences in completion rates could be due to differences in study populations, and the first two strategies may have reduced attrition, we hypothesize that the induction period played an important part in the improved contingency management

completion observed in this study. Aside from the methodological advantages, the induction period may have clinical relevance. It allows clinicians to identify individuals who are most likely to benefit from contingency management (i.e., current alcohol users who can attend frequent brief intervention visits) and allows individuals to receive reinforcers before abstinence is required. Those who cannot complete the induction are unlikely to benefit from contingency management, but might benefit from other interventions.

Limitations of this study include the relatively small sample size, which may have limited our ability to observe statistically significant differences during follow-up, as well as differences in psychiatric symptoms. Also, while we observed reductions in smoking, we did not observe group differences in smoking quit rates. Recruitment from one agency and the use of the induction period to restrict randomization to individuals best suited to contingency management might limit the generalizability of study results. In terms of age (Mean=45.38, SD=10.20) and gender (37% female), the sample is consistent with other studies of those with co-occurring disorders (21). Another limitation was the use of a \$30,000 onsite urine analyzer to conduct EtG tests, and an additional approximate cost of \$2.50 per test due to chemicals required for testing. The recent development of inexpensive (\$4) point-of-care EtG immunoassay dipcard tests (Confirm Biosciences, San Diego, CA) further improves the feasibility of an EtG-based contingency management intervention. Funding for reinforcers and urine tests can be a barrier to implementation. However, this barrier is likely surmountable as cost-effectiveness studies of contingency management in those with and without serious mental illness support the intervention's cost-savings (24, 34, 35).

Results of this study strongly support the efficacy of an EtG-based contingency management intervention for alcohol use disorders. Group differences in alcohol use observed in this study, and positive outcomes of our previous trial of contingency management targeting stimulant use, suggest that contingency management may be a particularly effective intervention for those with co-occurring serious mental illness and substance use disorders, a high-cost and difficult to engage population.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgments

Author Note: Funding for this study was provided by the National Institute on Alcohol Abuse and Alcoholism, R01 AA AA020248 (Principal Investigator: M.G. McDonnell)

The authors would like to thank the leadership and staff at Community Psychiatric Clinic, including Shirley Havenga, Kelli Nomura, Susan Peacy, Liz Quakenbush, Kurt Davis, and all those who volunteered to participate in the study.

Literature cited

1. Mokdad AH, Marks JS, Stroup DF, Gerberding JL. Actual causes of death in the United States, 2000. *JAMA*. 2004; 291:1238–1245. [PubMed: 15010446]
2. Bouchery EE, Harwood HJ, Sacks JJ, Simon CJ, Brewer RD. Economic costs of excessive alcohol consumption in the U.S., 2006. *Am J Prev Med*. 2011; 41:516–524. [PubMed: 22011424]

3. Cohen E, Feinn R, Arias A, Kranzler HR. Alcohol treatment utilization: findings from the National Epidemiologic Survey on Alcohol and Related Conditions. *Drug Alcohol Depend.* 2007; 86:214–221. [PubMed: 16919401]
4. Buckley PF. Prevalence and consequences of the dual diagnosis of substance abuse and severe mental illness. *J Clin Psychiatry.* 2006; 67(Suppl 7):5–9.
5. Lohr JB, Flynn K. Smoking and schizophrenia. *Schizophr Res.* 1992; 8:93–102. [PubMed: 1360812]
6. DHHS. Substance Abuse Treatment for Persons with Co-occurring Disorders. A Treatment Improvement Protocol TIP 42. Rockville MD: US DHHS; 2005.
7. Bradizza CM, Stasiewicz PR, Dermen KH. Behavioral Interventions for Individuals Dually-Diagnosed with a Severe Mental Illness and a Substance Use Disorder. *Curr Addict Rep.* 2014; 1:243–250. [PubMed: 25530935]
8. Hunt GE, Siegfried N, Morley K, Sitharthan T, Cleary M. Psychosocial interventions for people with both severe mental illness and substance misuse. *Cochrane Database Syst Rev.* 2013; 10:CD001088.
9. Wakefield PA, Randall GE, Richards DA. Identifying barriers to mental health system improvements: An examination of community participation in Assertive Community Treatment programs. *International journal of mental health systems.* 2011; 5:27. [PubMed: 22059856]
10. Brunette MF, Asher D, Whitley R, Lutz WJ, Wieder BL, Jones AM, McHugo GJ. Implementation of integrated dual disorders treatment: a qualitative analysis of facilitators and barriers. *Psychiatr Serv.* 2008; 59:989–995. [PubMed: 18757591]
11. Benishek LA, Dugosh KL, Kirby KC, Matejkowski J, Clements NT, Seymour BL, Festinger DS. Prize-based contingency management for the treatment of substance abusers: a meta-analysis. *Addiction.* 2014; 109:1426–1436. [PubMed: 24750232]
12. Prendergast ML, Hall EA, Roll J, Warda U. Use of vouchers to reinforce abstinence and positive behaviors among clients in a drug court treatment program. *J Subst Abuse Treat.* 2008; 35:125–136. [PubMed: 17997267]
13. Petry NM, Martin B, Cooney JL, Kranzler HR. Give them prizes, and they will come: contingency management for treatment of alcohol dependence. *J Consult Clin Psychol.* 2000; 68:250–257. [PubMed: 10780125]
14. Dougherty DM, Lake SL, Hill-Kapturczak N, Liang Y, Karns TE, Mullen J, Roache JD. Using contingency management procedures to reduce at-risk drinking in heavy drinkers. *Alcohol Clin Exp Res.* 2015; 39:743–751. [PubMed: 25833033]
15. Alessi SM, Petry NM. A randomized study of cellphone technology to reinforce alcohol abstinence in the natural environment. *Addiction.* 2013; 108:900–909. [PubMed: 23279560]
16. Jatlow PI, Agro A, Wu R, Nadim H, Toll BA, Ralevski E, Nogueira C, Shi J, Dziura JD, Petrakis IL, O'Malley SS. Ethyl Glucuronide and Ethyl Sulfate Assays in Clinical Trials, Interpretation, and Limitations: Results of a Dose Ranging Alcohol Challenge Study and 2 Clinical Trials. *Alcohol Clin Exp Res.* 2014
17. McDonnell MG, Skalisky J, Leickly E, McPherson S, Battalio S, Nepom JR, Srebnik D, Roll J, Ries RK. Using ethyl glucuronide in urine to detect light and heavy drinking in alcohol dependent outpatients. *Drug Alcohol Depend.* 2015; 157:184–187. [PubMed: 26475403]
18. Leickly E, McDonnell MG, Vilardaga R, Angelo FA, Lowe JM, McPherson S, Srebnik D, Roll JM, Ries RK. High levels of agreement between clinic-based ethyl glucuronide (EtG) immunoassays and laboratory-based mass spectrometry. *Am J Drug Alcohol Ab.* 2015; 41:246–250.
19. McDonnell MG, Howell DN, McPherson S, Cameron JM, Srebnik D, Roll JM, Ries RK. Voucher-based reinforcement for alcohol abstinence using the ethyl-glucuronide alcohol biomarker. *J Appl Behav Anal.* 2012; 45:161–165. [PubMed: 22403460]
20. Roll JM, Chermack ST, Chudzynski JE. Investigating the use of contingency management in the treatment of cocaine abuse among individuals with schizophrenia: a feasibility study. *Psychiatry Res.* 2004; 125:61–64. [PubMed: 14967553]
21. Bellack AS, Bennett ME, Gearon JS, Brown CH, Yang Y. A randomized clinical trial of a new behavioral treatment for drug abuse in people with severe and persistent mental illness. *Arch Gen Psychiatry.* 2006; 63:426–432. [PubMed: 16585472]

22. McDonnell MG, Srebnik D, Angelo F, McPherson S, Lowe JM, Sugar A, Short RA, Roll JM, Ries RK. Randomized controlled trial of contingency management for stimulant use in community mental health patients with serious mental illness. *Am J Psychiatry*. 2013; 170:94–101. [PubMed: 23138961]
23. McDonnell M, McPherson S, Vilardaga R, Srebnik D, Angelo FN, Leickly E, Saxon AJ, Roll J, Ries R. Preliminary findings: Contingency management targeting psycho-stimulant use results in secondary decreases in smoking for severely mentally ill adults. *Am J Addict*. 2014; 23:407–410. [PubMed: 24961363]
24. Murphy SM, McDonnell MG, McPherson S, Srebnik D, Angelo F, Roll JM, Ries RK. An economic evaluation of a contingency-management intervention for stimulant use among community mental health patients with serious mental illness. *Drug Alcohol Depend*. 2015; 153:293–299. [PubMed: 26026494]
25. Javors MA, Hatch JP, Lamb RJ. Cut-off levels for breath carbon monoxide as a marker for cigarette smoking. *Addiction*. 2005; 100:159–167. [PubMed: 15679745]
26. Sobell, LC., Sobell, MB. *Handbook of Psychiatric Measures*. Washington DC: American Psychiatric Association; 2000. Alcohol timeline followback (TLFB); p. 477-479.
27. Donlin WD, Knealing TW, Needham M, Wong CJ, Silverman K. Attendance rates in a workplace predict subsequent outcome of employment-based reinforcement of cocaine abstinence in methadone patients. *J Appl Behav Anal*. 2008; 41:499–516. [PubMed: 19192855]
28. Holtyn AF, Koffarnus MN, DeFulio A, Sigurdsson SO, Strain EC, Schwartz RP, Leoutsakos JM, Silverman K. The therapeutic workplace to promote treatment engagement and drug abstinence in out-of-treatment injection drug users: a randomized controlled trial. *Prev Med*. 2014; 68:62–70. [PubMed: 24607365]
29. Petry NM, Peirce JM, Stitzer ML, Blaine J, Roll JM, Cohen A, Obert J, Killeen T, Saladin ME, Cowell M, Kirby KC, Sterling R, Royer-Malvestuto C, Hamilton J, Booth RE, Macdonald M, Liebert M, Rader L, Burns R, DiMaria J, Copersino M, Stabile PQ, Kolodner K, Li R. Effect of prize-based incentives on outcomes in stimulant abusers in outpatient psychosocial treatment programs: a national drug abuse treatment clinical trials network study. *Arch Gen Psychiatry*. 2005; 62:1148–1156. [PubMed: 16203960]
30. Zeger SL, Liang KY, Albert PS. Models for longitudinal data: a generalized estimating equation approach. *Biometrics*. 1988; 44:1049–1060. [PubMed: 3233245]
31. Enders CK. Missing not at random models for latent growth curve analyses. *Psychol Methods*. 2011; 16:1–16. [PubMed: 21381816]
32. Muthen B, Asparouhov T, Hunter AM, Leuchter AF. Growth modeling with nonignorable dropout: alternative analyses of the STAR*D antidepressant trial. *Psychol Methods*. 2011; 16:17–33. [PubMed: 21381817]
33. Schafer JL, Graham JW. Missing data: our view of the state of the art. *Psychol Methods*. 2002; 7:147–177. [PubMed: 12090408]
34. Olmstead TA, Petry NM. The cost-effectiveness of prize-based and voucher-based contingency management in a population of cocaine- or opioid-dependent outpatients. *Drug Alcohol Depend*. 2009; 102:108–115. [PubMed: 19324501]
35. Sindelar J, Elbel B, Petry NM. What do we get for our money? Cost-effectiveness of adding contingency management. *Addiction*. 2007; 102:309–316. [PubMed: 17222286]

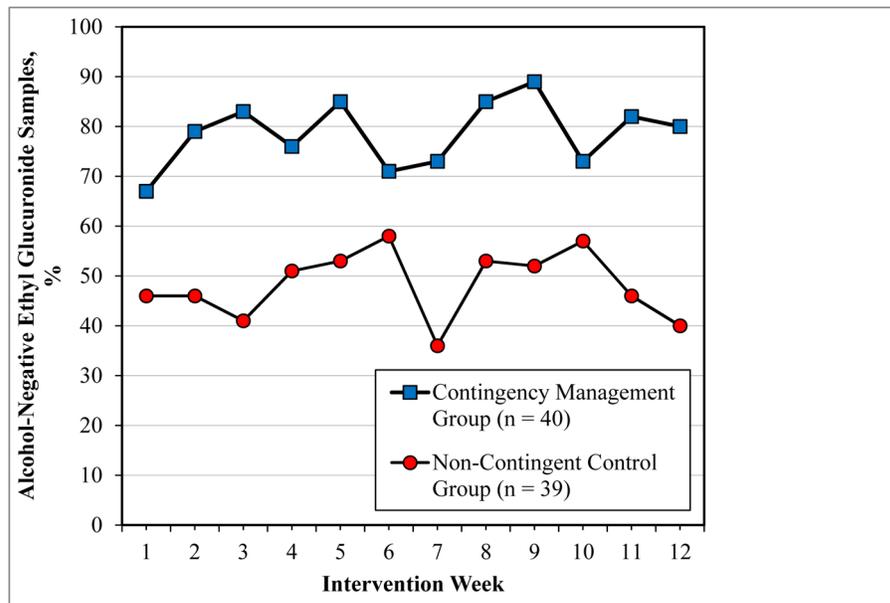


FIGURE 1.

Group Differences in Alcohol-Negative Ethyl Glucuronide Urine Samples across the 12-week Intervention Period.

Note: Participants assigned to the contingency management group were 3.13 (95% CI: 2.18, 4.50, $p < 0.05$) times more likely to submit an alcohol-negative urine sample relative to controls during the 12-week treatment period. While the use of multiple imputation techniques allowed us to include the entire sample ($N=79$) in our inferential statistical tests for treatment outcomes, in this figure we display the original raw data without imputation.

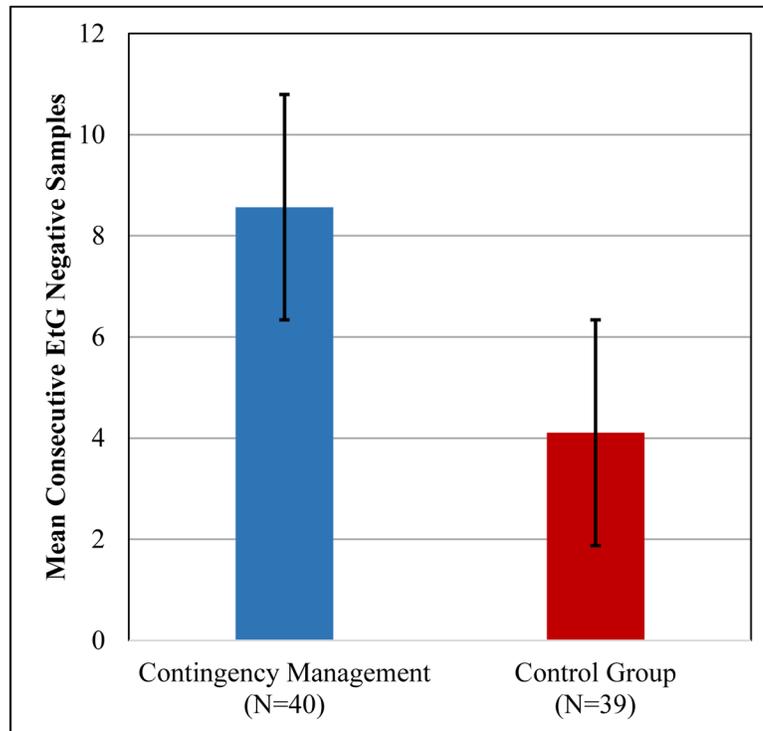


FIGURE 2.

Group differences in longest duration of continuous alcohol abstinence.

Note: The Figure represents the mean number of continuous alcohol-negative ethyl glucuronide alcohol tests attained by the contingency management and non-contingent control groups during the intervention period, $F(3)=5.55$, $df=1, 76$, $p<0.05$.

TABLE 1

Baseline demographic and clinical characteristics of the evaluative sample (N=79)

	Contingency Management n=40		Non-Contingent Control n=39	
	n	%	n	%
Demographic Characteristics				
Age			Mean	SD
			44.55	9.92
Male	26	65%	46.23	10.55
Race				
Caucasian	23	57%	21	53%
African American	13	33%	10	26%
Other	4	10%	8	21%
Homeless or unstably housed	16	40%	15	39%
Psychiatric Medications				
Antidepressants	12	30%	11	28%
Mood stabilizers	4	10%	6	15%
Antipsychotics	9	23%	9	23%
Naltrexone	0	0%	1	3%
Diagnosis				
Major depressive disorder	14	35%	12	31%
Bipolar disorder	14	35%	15	38%
Schizophrenia-spectrum disorder	12	30%	12	31%
Substance Use Characteristics				
Self-reported days of alcohol use in the 30 days prior to study entry			14.50	7.75
Self-reported days of alcohol intoxication in the 30 days prior to study entry			12.90	7.83
Positive urine analysis for illicit drugs	15	38%	20	51%
Moderate-high nicotine dependence (assessed by Fagerstrom)	19	24%	14	18%
% Addiction treatment attendance			57%	34%
			53%	37%

Note: All differences are non-significant.

TABLE 2

Descriptive statistics for primary and secondary outcomes during the intervention and post-intervention follow-up time periods (N=79)*

Outcome Variable	Intervention				Follow-up			
	Contingency Management (n=40)	Non-Contingent Control (n=39)	Contingency Management (n=40)*	Non-Contingent Control (n=39)*	Mean	SE	Mean	SE
% Alcohol Negative urine sample**	50%	24%	3%	3%	51%	8%	29%	7%
EtG Value (ng/mL)	408.86	734.79	39.57	40.66	417.45	109.29	777.11	112.30
Self-reported days Alcohol Use Last 30 days	3.72	12.01	1.57	1.52	3.32	1.40	10.73	1.39
Self-reported days Alcohol Intoxication Last 30 Days	2.92	9.35	1.47	1.43	2.22	1.13	7.91	1.12
Self-reported % Heavy drinking last 5 days	13%	34%	2%	3%	12%	5%	41%	7%
% Stimulant Negative Urine Tests	90%	75%	3%	2%	90%	5%	66%	8%
Self-reported Days Stimulant Use in Last 30 Days	0.57	2.79	0.68	0.68	0.34	0.70	1.95	0.70
% Smoking Negative Carbon Monoxide Samples (N=54)	10%	2%	2%	1%	10%	5%	2%	2%

Note: Bold text indicates statistically significant group differences, p<0.05.

* During the follow-up period 25% (n=10) and 23% (n=9) participants in the contingency management and non-contingent control condition, respectively, did not provide data that could be descriptively analyzed.

** Alcohol negative urine sample equals ethyl glucuronide < 150 ng/mL.

*** EtG: ethyl glucuronide immunoassay value ranging from 0 to 2,000 ng/mL

Stimulant drugs include amphetamines, methamphetamine, and cocaine.

Note: While the use of multiple imputation techniques allowed for us to include the entire sample (N=79) in our inferential statistical tests of intervention outcomes, in this Table the original raw data (without imputation) were used to generate descriptive statistics. Estimated marginal means are reported.

TABLE 3

Multiple imputation results for primary and secondary outcomes during the intervention and post-intervention follow-up time periods (N=79).

Outcome Variable	Intervention		Follow-up	
	Estimate	CI (95%)	Estimate	CI (95%)
% Alcohol Negative urine sample *	Odds ratio=2.86	2.12:3.87	Odds ratio=2.60	0.97:6.93
EtG Value (ng/mL) **	B=325.70	223.86:427.53	B=283.27	-58.84:625.18
Self-reported days Alcohol Use Last 30 days	B=7.22	3.30:11.14	B=7.21	3.30:11.14
Self-reported days Alcohol Intoxication Last 30 Days	B=4.58	0.18:8.99	B=5.27	2.30:8.24
Self-reported % Heavy drinking last 5 days	Odds ratio=2.96	2.09:4.18	Odds ratio=4.11	1.54:10.93
% Stimulant Negative Urine Tests	Odds ratio=2.79	2:03:3.84	Odds ratio=2.99	0.98:8.49
Self-reported Days Stimulant Use in Last 30 Days	B=1.86	0.66:5.09	B=2.89	0.98:8.49
% Smoking Negative Carbon Monoxide Samples (N=54)	Odds ratio=2.62	1.16:5.91	Odds ratio=5.58	0.46:67.92

Note: Bold text indicates statistically significant group differences, $p < 0.05$.

* Alcohol negative urine sample equals ethyl glucuronide < 150 ng/mL.

** EtG: ethyl glucuronide immunoassay value ranging from 0 to 2,000 ng/mL