### SNARKY: Sharing Needles As a Result of Kowtowing to Yuppies

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#### Abstract:

Aim: People who inject drugs (PWID) in public have a higher risk of homelessness, incarceration, rushed injection, and overdose than people who do not inject drugs in public. There is limited research on the relationship between public injection and syringe sharing, a risk factor for infectious diseases. The aim of this study was to investigate longitudinal associations between frequency of public injection and syringe sharing among PWID in San Francisco and Los Angeles, California. Methods: PWID (N=984) were recruited using targeted sampling and completed surveys during three study visits (baseline, 6-month, and 12-month) in 2016 and 2017. The explanatory variable was frequency of having injected in public (never, occasionally/sometimes, or usually/always), and the outcome variable was number of times having used syringes that had already been used by someone else in the past six months. Explanatory and outcome variables were measured across all three timepoints. We used a longitudinal negative binomial mixed-effects model to assess the relationship of interest.

Results: At baseline, 78% of participants had injected publicly in the past six months, with 38% of the sample reporting that they usually/always injected publicly. Over a quarter (26%) had injected with a previously used syringe in the six months preceding baseline, with an average of 20.9 instances of receptive syringe sharing during this period. The majority (83%) reported current homelessness at baseline. In multivariable analysis across all timepoints, participants who usually/always injected publicly had 7.99 (95% CI: 5.11-10.88, p < .001) times higher counts of receptive syringe sharing over the past six months, compared to those who never publicly injected, controlling for age, gender, homelessness, stimulant injecting, opioid injecting, and binge drinking. Participants who occasionally/sometimes injected publicly had 3.28 (95% CI: 0.50-6.06, p < .001) times higher counts of receptive syringe sharing over the past six months, compared to those who never publicly injected, controlling for covariates.

<u>Conclusion</u>: Injecting publicly is associated with a higher frequency of receptive syringe sharing among  $\overline{PWID}$ , suggesting the need for safer environment interventions, such as supervised consumption sites.

## Research Question

Among people who inject drugs (PWID), is frequency of public injection (e.g. on a sidewalk) associated with increased frequency of injecting with a syringe previously used by someone else?

## Description of Study

This study was a randomized controlled trial to determine efficacy of an intervention to prevent people from helping others to inject drugs for the first time. 984 participants were recruited in San Francisco and Los Angeles using targeted sampling, and completed surveys at three study visits from 2016-2017.

For this analysis we ignored study arm and treated data over the three study visits (baseline, 6-month, and 12-month follow-up) as repeated measures. Surveys included data on demographic variables (age, gender, race), as well as time-varying variables including months homeless, types of substances used and routes of administration, frequency of substance use, frequency of injecting in public, and number of times in the past 6 months using syringes that were already used by someone else.

## Description of Sample

One of the biggest challenges of this analysis was appropriately dealing with informative censoring with large loss-to-follow-up.

		Baseline	6-month F/U	12-month F/U
Total		978 (100%)	595 (60.8%)	532 (54.4%)
Age	Min	18	18	18
	Median	43	45	45
	Max	76	76	71
Gender	Male	739 (75.6%)	437 (73.4%)	387 (72.7%)
	Female	225 (23%)	149 (25%)	139 (72.7%)
	Transgender	14 (1.4%)	9 (1.5%)	6 (1.1%)
Race	Black	196 (20%)	141 (23.7%)	126 (23.7%)
	Latinx	228 (23.3%)	131 (22%)	114 (21.4%)
	White	413 (42.2%)	238 (40%)	219 (41.2%)
	Other	141 (14.4%)	85 (14.3%)	73 (13.7%)
Public Injection	Always/Usually	376 (38.4%)	171 (28.7%)	125 (23.5%)
	Sometimes/Occasionally	390 (39.9%)	171 (35.8%)	193 (36.3%)
	Never	212 (21.7%)	183 (30.8%)	170 (32%)
Times Sharing	0	723 (73.9%)	507 (121.5%)	462 (135.9%)
	1 - 15	207 (21.2%)	68 (34.8%)	50 (38.9%)
	16 - 30	21 (2.1%)	3 (3.5%)	3 (3.9%)
	31+	27 (2.8%)	8 (4.5%)	8 (5.1%)
Months Homeless	0	168 (17.2%)	152 (25.5%)	176 (33.1%)
	1 - 6	138 (14.1%)	33 (5.5%)	346 (65%)
	7 - 12	112 (11.5%)	67 (11.3%)	1 (0.2%)
	13+	560 (57.3%)	343 (57.6%)	9 (1.7%)
Substance Use	Used stimulants	674 (68.9%)	330 (55.5%)	253 (47.6%)
	Used opiates	834 (85.3%)	424 (71.3%)	341 (64.1%)
	Binge drinking	292 (29.9%)	149 (25%)	85 (16%)

### Statistical Model

We converted our research question into a parameter of interest using a series of longitudinal regression models where:

$$log(E[Y|X]) = \beta_0 + \beta_{0i} + (\beta_1 + \beta_{1i} + \beta_{1j})X_{1ij} + (\beta_2 + \beta_{2i} + \beta_{2j})X_{2ij} + \beta_3X_3 + \beta_4X_4 + (\beta_5 + \beta_{5i} + \beta_{5j})X_{5ij} + (\beta_6 + \beta_{6i} + \beta_{6j})X_{6ij} + (\beta_7 + \beta_{7i} + \beta_{7j})X_{7ij}$$

#### where:

- Y = number of times shared syringes (outcome)
- $\triangleright$   $X_1$  = frequency of public injection (exposure)
- $ightharpoonup X_2 = \text{homelessness status}$
- $X_3 = \text{gender}$
- $X_4 = age$
- $\triangleright$   $X_5$  = any stimulant injection
- $ightharpoonup X_6 =$ any opiate injection
- $X_7 =$ any binge drinking

All variables have a recall period of 6 months, where applicable.

### Methods used to fit model and derive inference

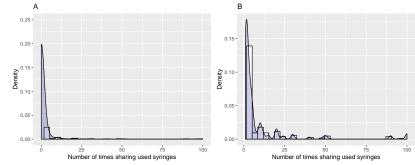
We fit 3 models to answer this question: GEE, and mixed effects models using both poisson and negative binomial distributions.

The **GEE model** estimated the marginal effects of public injection frequency on receptive syringe sharing across the three timepoints, controlling for homelessness, gender, age, stimulant or opiate injection, and binge drinking. GEEs treat the correlation between subjects as a "nuisance" parameter. We did this using the gee package in R, assuming an exchangeable correlation structure and a poisson distribution.

The **mixed effects models** estimate the marginal longitudinal effect of public injection frequency on receptive syringe sharing, controlling for the same confounders, but allow for both a random intercept term and a random effect term for frequency of public injection (our exposure), homelessness, and the substance use variables. These models use maximum likelihood to estimate the coefficients in the model, but rely on correct specification of both the mean model and the correlation structure of the residual errors. We did this using the glmer function in the MASS R package, and using the glmer.nb function in the lme4 R package for the negative binomial model.

# Compare models (1)

Since we have count data, we first created a histogram of the outcome (number of times sharing needles after someone else had already used them, over the prior six months). The left panel (A) includes all participants, and the right panel (B) includes only participants who reported sharing needles at least once during the recall period.



This shows us that the vast majority of participants (more than half) reported zero for the outcome, which may cause problems for a poisson model. It is also extremely right-skewed, also implying that a negative binomial model may be a better fit.

# Compare models (2)

We also want to compare models with a few different tests, to see which appears to have the best fit.

1. Since the mixed models are nested, we can use a likelihood ratio test:

Model	Log Likelihood	P-value
Poisson Neg Binom.	-2231.702 -1223.066	0

This tells us that the negative binomial model is significantly preferred over the poisson model.

2. We can also compare the AIC of the mixed models, though this involves comparing likelihoods that ignore the repeated measures in our study:

Model	AIC			
Poisson	4485.403			
Neg Binom.	2470.132			

This also suggests that the negative binomial model is a better fit.

# Compare models (3)

We are concerned about strong violations of the assumption of no informative censoring, for our GEE model. Yet we are also concerned about overdispersion, as well as the large number of zeros in our outcome, both of which may cause our poisson mixed methods model to be a poor fit.

For these reasons, it is not surprising that the negative binomial model may be the best fit.

We are not able to run comparison tests of fit for GEE compared to our mixed models, so one of the best things to do is to look at our results, including standard error.

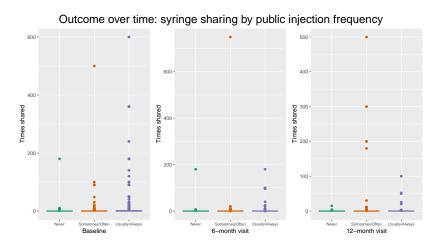
### Results

	GEE (Poisson)		Mixed Model (Poisson)		Mixed Model (Neg Binom.)	
Predictor	Est.	p-value*	Est.	p-value	Est.	p-value
Never publicly inject	ref		ref		ref	
Occasionally/sometimes publicly inject	4.12	0.09	2.44	2e-22	3.28	0.00066
Usually/always publicly inject	9.77	0.01	3.67	5.5e-40	8	7.5e-08
Housed at study visit	ref		ref		ref	
Homeless at study visit	0.27	0.08	1.24	0.05	0.89	0.73
Male	ref		ref		ref	
Female	0.89	0.83	1.01	0.98	1.31	0.47
Transgender	0.28	0.16	0.91	0.94	1.1	0.93
Age (year)	0.98	0.32	0.96	0	0.98	0.08
No stimulant use	ref		ref		ref	
Stimulant use	1.33	0.57	0.66	3.5e-05	1.87	0.03
No opiate use	ref		ref		ref	
Opiate use	0.82	0.73	0.72	0.0016	1.74	0.09
No binge drinking	ref		ref		ref	
Binge drinking	0.72	0.45	0.89	0.053	1.59	0.05

<sup>\*</sup>p-values based on robust SEs

We can see from these results that the inference is similar across models, though the magnitude of the effect of public injection is much stronger in the negative binomial mixed model. The standard error is also much larger for the GEE model, as evidenced by larger p-values.

### Plot of Results



From these plots we can see there are numerous outliers at each visit, which again support the use of the negative binomial model for our findings.

#### Discussion

In multivariable analysis across all timepoints, participants who usually/always injected publicly had 8 (95% CI: 5.11-10.88, p <.001) times higher counts of receptive syringe sharing over the past six months, compared to those who never publicly injected, controlling for covariates. Participants who occasionally/sometimes injected publicly had 3.28 (95% CI: 0.50-6.06, p <.001) times higher counts of receptive syringe sharing over the past six months, compared to those who never publicly injected, controlling for covariates.

<u>Conclusion</u>: Injecting publicly is associated with receptive syringe sharing among PWID, with higher frequency of public injection corresponding to higher frequency of sharing syringes. This suggests that PWID may be rushing their injections in public settings by using shared syringes. Future research should explore the role that police presence plays in this relationship, e.g. as a mediator on the pathway. Findings underscore the need for safer environment interventions, such as supervised consumption sites, where PWID can take their time injecting and use sterile equipment.

<u>Limitations</u>: This study had significant loss-to-follow-up. The authors used complete case analysis for the sake of simplicity, however results may change if other missing data approaches were applied, such as inverse probability of censoring weighting or multiple imputation.