

The effect of depression on missed HIV medical visits among patients in the CFAR Network of Integrated Systems (CNICS) cohort in the United States

Brian Pence*, Angela Bengtson, Katerina Christopoulos, Stephen Cole, Heidi Crane, Bradley Gaynes, Amy Heine, W. Christopher Mathews, Richard Moore, Sonia Napravnik, Conall O’Cleirigh, Steven Safren, and Michael Mugavero for the CNICS Collaboration

* Associate Professor, Dept. of Epidemiology, UNC-Chapel Hill

Depression is common and consequential

- 20-30% prevalence among HIV-infected patients
- Strongly and consistently associated with
 - Reduced ARV adherence
 - Lack of viral suppression
 - Clinical progression
 - Mortality
- Relationship with missed HIV visits less well understood

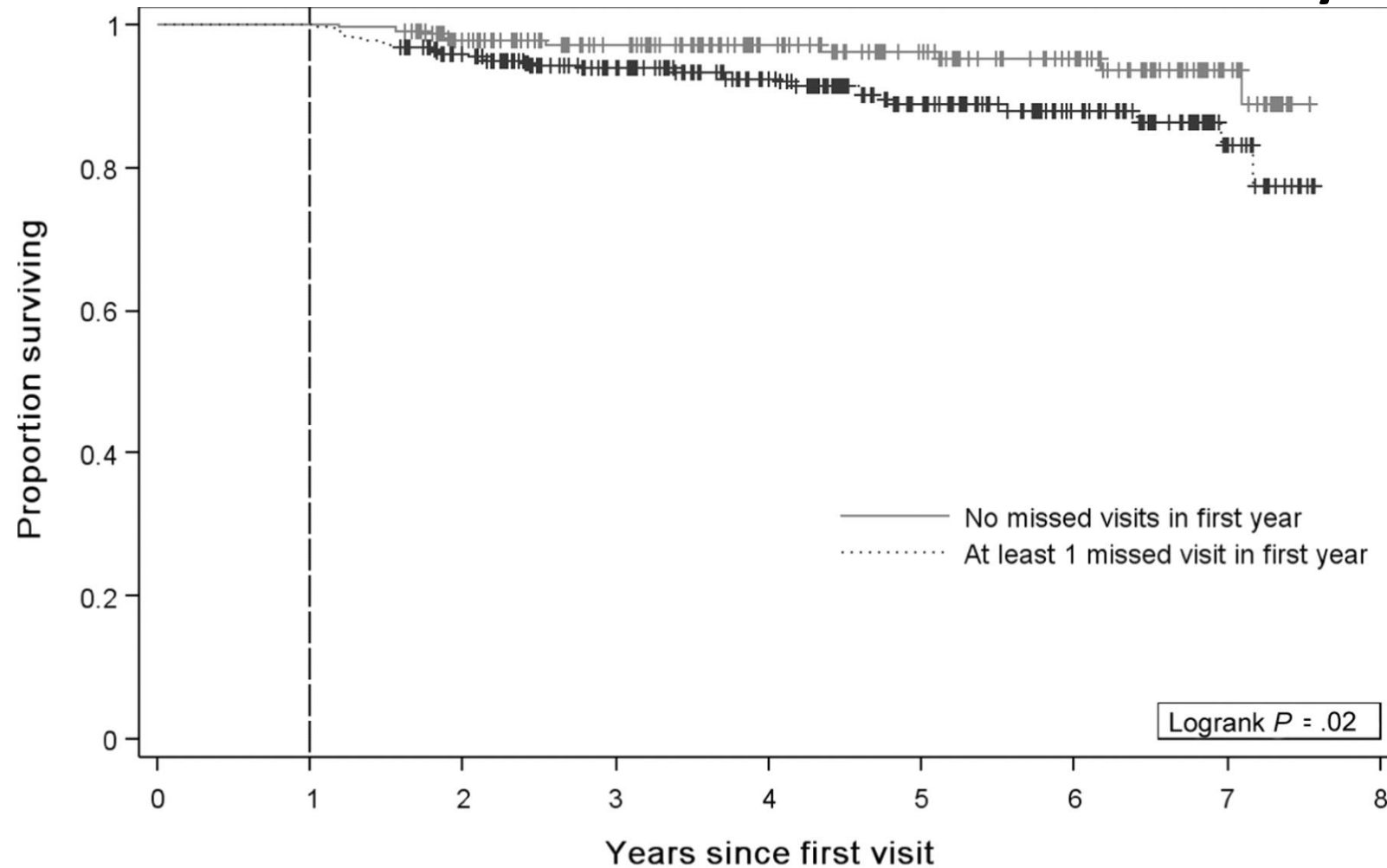
Missed HIV Visits are Common

Indicators of HIV care attendance among 10,053 HIV-infected patients at 6 HIV clinics over 12 months, 2008-2009

Indicator	Percent of patients or appointments
≥1 no-show visit	67%
Missed visit proportion	31%
No 4-month constancy	49%
≥6 month gap between appointments	32%
Not retained by HRSA HAB measure (≥2 visits ≥90 days apart)	23%

Missed Visits Matter...

Association of missed visits with mortality

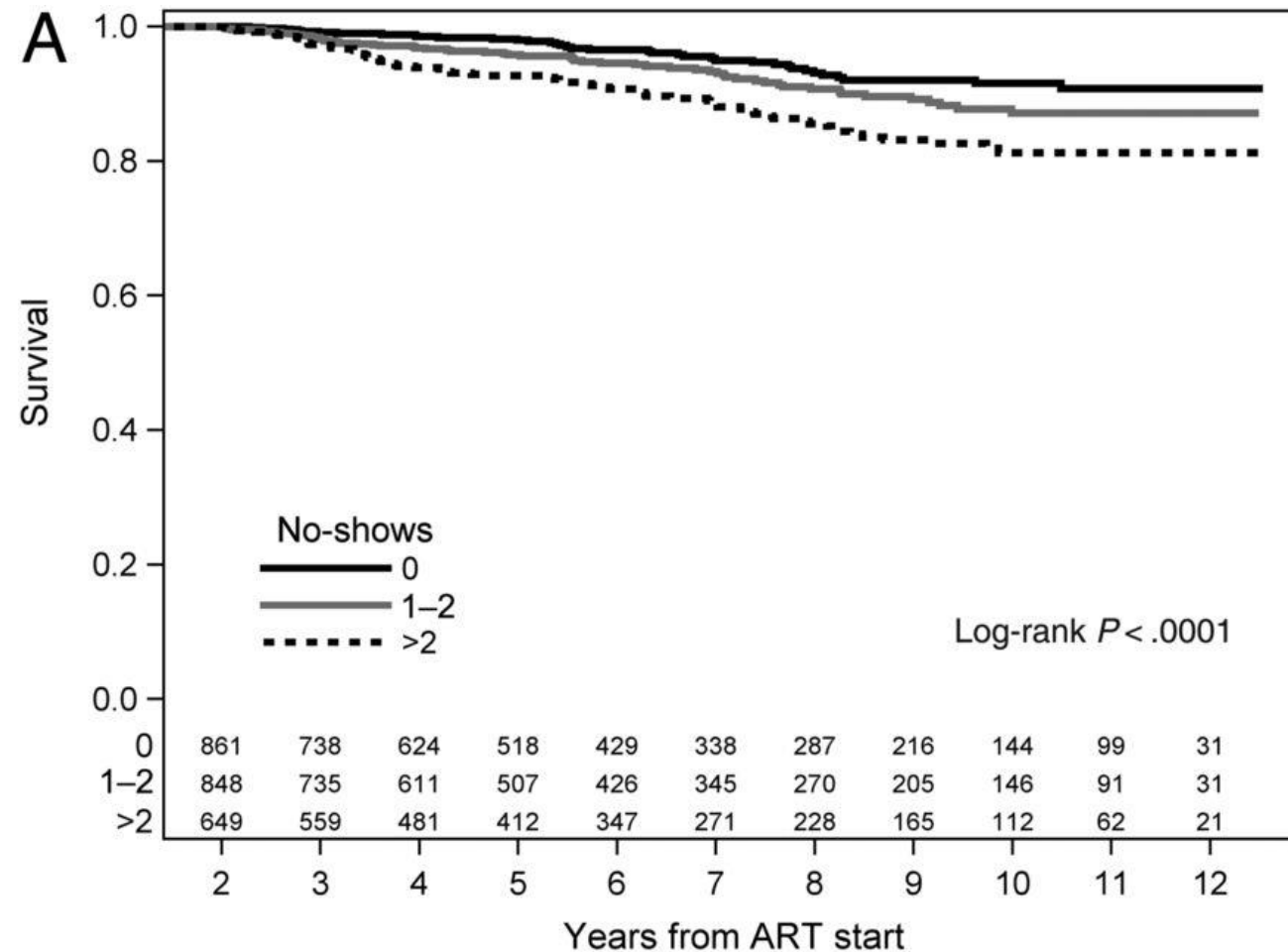


Mugavero
CID 2009

	Patients	Died	Censored
No missed visits in first year	218	5% (10)	95% (208)
At least 1 missed visit in first year	325	10% (32)	90% (293)

... Even for patients meeting retention benchmarks

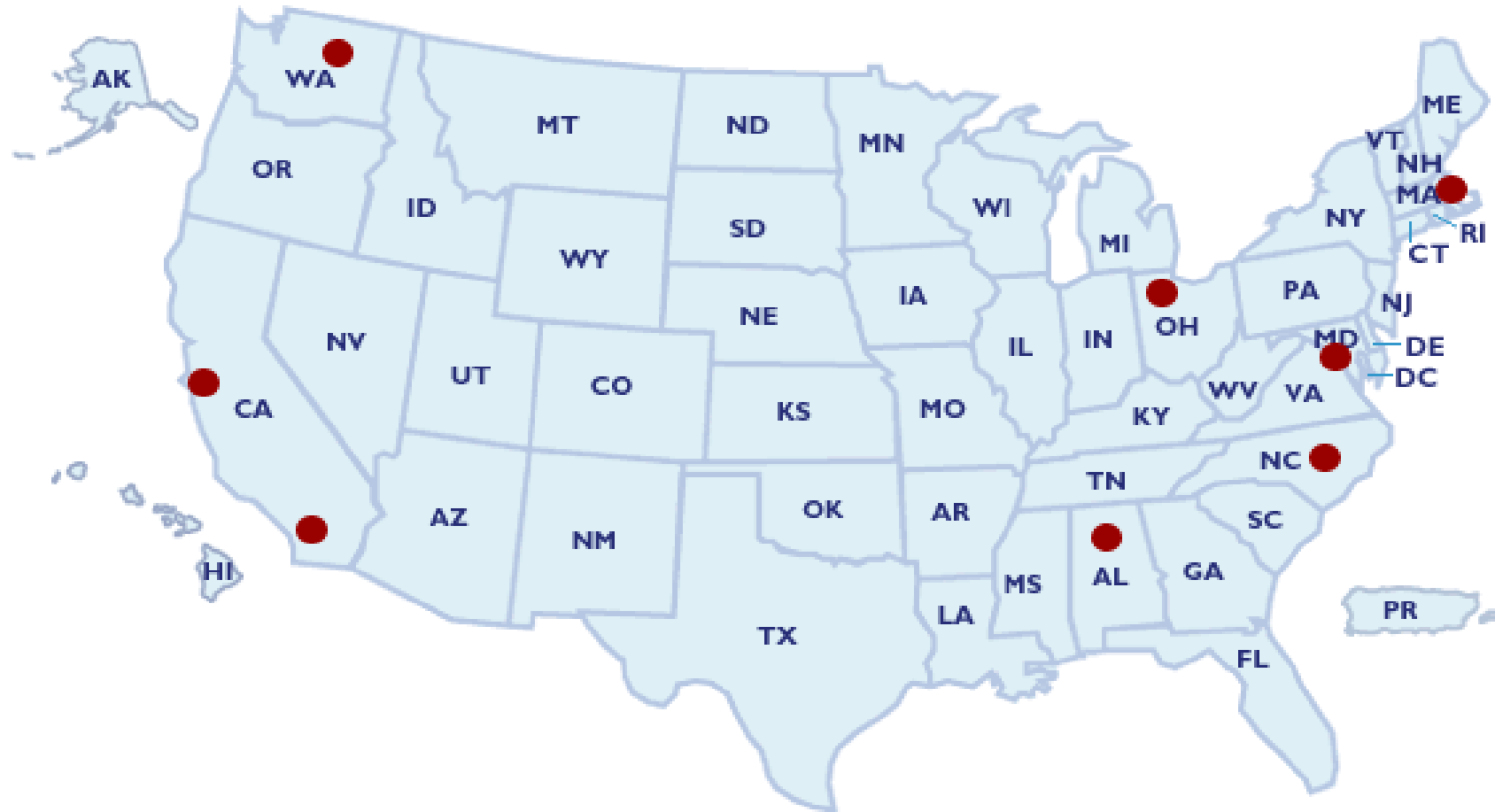
Association of missed visits with mortality among patients meeting HRSA HAB retention criterion



Research question

What is the effect of depression on missed visits and retention in HIV care?

Data source: CFAR Network of Integrated Clinical Systems (CNICS)



CNICS Data Elements

- Electronic Health Records data
 - Demographics
 - Appointment attendance
 - Labs
 - Medications
 - Diagnoses
- Patient-Reported Outcome (PRO) data (~ every 6 months)
 - Depression, anxiety, substance use, alcohol use
 - ARV adherence

Sample

- All patients with ≥ 1 attended HIV medical appointment with ≥ 1 depression assessment (PHQ-9) between 2005-2013
- Patients followed from first PHQ-9 to earliest of:
 - Death
 - Administrative censoring (2014)
 - Loss to care (>12 months with no attended HIV appointment)

Measures

- Unit of analysis: Each attended appointment
- Outcomes: At each attended appointment, identified
 - Whether next scheduled appointment was attended or missed (excluding bounced, canceled, and rescheduled visits) (no-show)
 - Whether patient had ≥ 2 visits ≥ 90 days apart over next 12 months (HRSA HAB measure)
 - Missed visit proportion over next 12 months (MVP)
- Only visits with ≥ 12 months of subsequent follow-up before censoring were included for HAB and MVP measures
- Exposure: Probable depression (PHQ-9 total score ≥ 10)

Analysis

- To address confounding and identify causal effect: Marginal structural model (MSM) fit using inverse probability of treatment weights (IPTW)
 - Goal of MSM is to use weights to achieve balance in covariates between exposed and unexposed groups
 - Unadjusted (weighted) analysis then yields causal contrast between exposure groups, akin to RCT
- IPTW created by fitting a model with depression status ($\text{PHQ-9} \geq 10$) on the left and potential confounders on the right
- Weighted analysis mimics intent-to-treat RCT analysis of “randomized to depression at baseline” vs. “randomized to no depression”

Analysis

- To address nonrandom loss to follow-up: Inverse probability of censoring weights (IPCW)
- Both sets of weights stabilized by the appropriate marginal probability (of treatment or censoring)
- Two sets of weights multiplied to create single IPTC weight
- Pooled generalized linear models to estimate risk differences, risk ratios, and mean differences, accounting for multiple observations per person

Inputs into weight models

Treatment weights

- Site
- Age*, gender, race/ethnicity
- CD4, suppressed VL*
- ART status, antidepressant status*
- Anxiety, alcohol use, drug use PROs*
- Chart diagnoses: Mental health, CVD, diabetes
- Time since entry into analysis sample*

Censoring weights

Same inputs as well as

- Depression PRO*

* Time-updated

Sample

	N
Patients	9,752
Person-years	26,155
Age (mean, SD)	43 (11)
Male gender	85%
Black non-Hispanic	29%
Hispanic	16%
Ever depressed	37%

Unweighted characteristics

	Ever depressed person-time	Never depressed person-time
Male gender	84%	86%
Suppressed VL	74%	78%
Recent drug use (PRO)	21%	13%
Anxiety diagnosis	29%	19%
On antidepressants	41%	21%

Effect of weighting

	Unweighted person-time		Weighted person-time	
	Ever depressed	Never depressed	Ever depressed	Never depressed
Male gender	84%	86%	85%	85%
Suppressed VL	74%	78%	77%	79%
Recent drug use (PRO)	21%	13%	18%	19%
Anxiety diagnosis	29%	19%	26%	25%
On antidepressants	41%	21%	35%	30%

Effect of depression on visit attendance (weighted analysis)

Outcome	Exposure	Risk	Difference	Ratio
Next visit missed (no show)	Depressed	16%	3% (1-4%)	1.18 (1.08-1.29)
	Not depressed	13%	0 (ref)	1 (ref)

Effect of depression on visit attendance (weighted analysis)

Outcome	Exposure	Risk / Mean	Difference	Ratio
Next visit missed (no show)	Depressed	16%	3% (1-4%)	1.18 (1.08-1.29)
	Not depressed	13%	0 (ref)	1 (ref)
Missed visit proportion, next 12 mo.	Depressed	16%	3% (1-4%)	n/a
	Not depressed	13%	0 (ref)	n/a

Effect of depression on visit attendance (weighted analysis)

Outcome	Exposure	Risk / Mean	Difference	Ratio
Next visit missed (no show)	Depressed	16%	3% (1,4%)	1.18 (1.08,1.29)
	Not depressed	13%	0 (ref)	1 (ref)
Missed visit proportion, next 12 mo.	Depressed	16%	3% (1,4%)	n/a
	Not depressed	13%	0 (ref)	n/a
Out of care, next 12 mo. (HRSA HAB)	Depressed	18%	-1% (-3,1%)	0.92 (0.84,1.03)
	Not depressed	19%	0 (ref)	1 (ref)

Assumptions for calling this an “effect”

- Exchangeability (no unmeasured confounding)
- Consistency (“depressed” and “not depressed” are well defined and consistent conditions)
- Positivity (no one was structurally unable to be depressed or to be not depressed)
- Good measurement (PHQ-9 ≥ 10 is a good measure of depression)

Interpretation and Conclusions

- Depression had an effect, albeit small in magnitude, on missed visits
- No effect (or possibly a protective effect) on minimum retention in care (HRSA HAB measure)
- Supports other research suggesting that missed visits and minimum retention are separate phenomena
- Suggests that depression care should be a component of a multifaceted strategy to pre-empt no-shows

Many thanks to...

- NIH: R01 MH 100970; CNICS platform grant R24 AI 067039
- Co-authors and other CNICS collaborators
- IAPAC conference organizers
- CNICS participants

Contact: bpence@unc.edu