## Treatment of Hepatitis C in People Who Use Drugs (PWUD)

Andrew Seaman, MD Oregon HCV Update Meeting December, 2018

## Conflicts of interest

2016 received <8% of my salary from an investigator initiated, Merck funded trial (makers of elbasvir/grazoprevir)

I am highly influenced by the opinions, life experience, and knowledge my patients bring to the table (many of whom inject drugs)

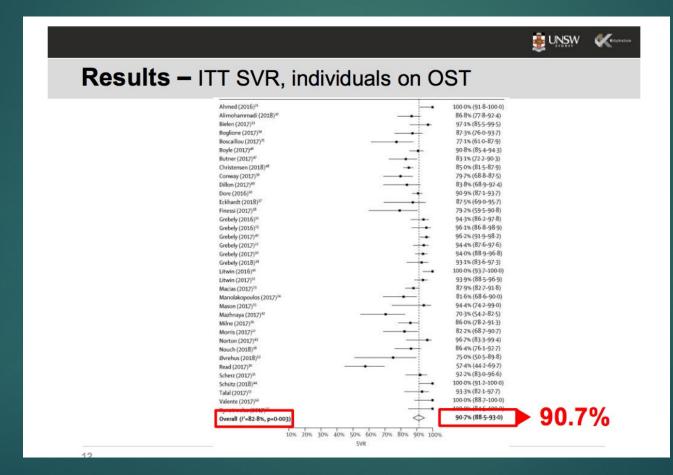
## Objectives

Brief update literature treatment of PWIDs for HCV

Brief review of the Old Town Clinic / Outside In / OHSU pilot trial for HCV treatment in People Who Inject Drugs

Brief discussion of the Old Town Clinic treatment program (if time)

## Meta-analysis 3,634 PWUD treated with DAAs



Hajarizadeh, The Lancet Gastroenterology & Hepatology 2018. Adapted INHSU 2018 with permission.

## Strongly biased by 75% observational studies

		ISW 帐			
Results – Meta-regression, ITT SVR					
	Adjusted model OR (95% CI)	Ρ			
Participants with recent drug use					
Participants receiving OST	1.04 (0.96-1.12)	0.364			
Men	1.07 (0.82-1.39)	0.612			
Median/mean age	1.07 (1.02-1.12)	0.008			
Participants with HIV co-infection	0.96 (0.86-1.07)	0.427			
Study design					
Observational	1.00				
Clinical Trial	2.18 (1.27-3.75)	0.006			

Hajarizadeh, The Lancet Gastroenterology & Hepatology 2018. Adapted INHSU 2018 with permission.

## OTC – OI – OHSU Pilot Study

- Prospective, non-randomized real world clinical trial using elb/graz to treat people who inject drugs with GT 1 or 4 HCV and an APRI <0.7 who:</p>
  - Arm 1: engage with Medication Assisted Therapy (Methadone/Bupe), n=25, Old Town Clinic
  - Arm 2: are actively using and engage with needle exchange program, n=25, Outside In
  - Arm 3: matched cohort in OHSU hepatology clinic, n=50

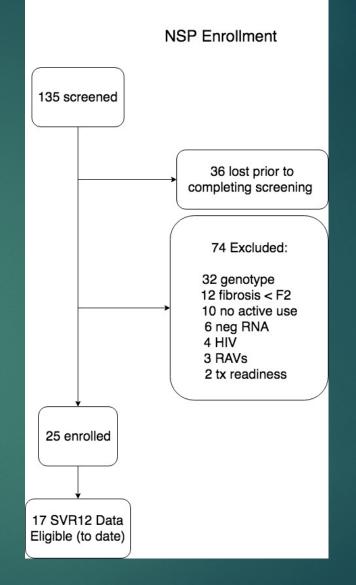
### NSP vs OAT HCV Pilot Study

	OAT	NSP	PWID Standar d	Com. Standar d
SVR 12, ITT % / (N)	<u>96% (24)</u>	<u><b>59% (10)</b></u> (p<0.001)	<u>89% (32)</u>	<u>94% (47)</u>
SVR12, Per Protocol % / (N)	100% (24)	90% (10)	100% (32)	100% (47)
Treatment Failures % / (N)	0%	6%* (1)	0%	0%
<b>Adherence</b> (% ≤ 7 pills missed)	92%	65%	81%	98%



Seaman et al. INHSU. 2018.

Many barriers to HCV treatment in harm reduction environments.



# Small difference in SVR, partially explained by trial type, LTFU

Meta-analysis of 38 studies of PWUDs, n=3,634 included

	<b>Treatment Completion</b>	Svr12
All PWUDs	98%	8.8%
PWUDs on OAT	97%	91%
Recent IDU	97%	87.5%

Meta-Regression Analysis: Clinical trials associated with OR 2.2 (1.27-3.75) of achieving SVR12

#### ► ITT analysis

Hajarizadeh B. Lance Gastro & Hepatology. V3 (11), Nov 2018.

## Old Town Clinic Treatment Program

#### Multidisciplinary

- Medical director + two providers
- ► HCV coordinator
- Clinical pharmacist
- CADC as indicated
- Weekly committee meetings
  - Decision made on need for treatment candidacy, Substance Used Disorder support, adherence support
  - Drug, labs ordered and PA process started by coordinator
  - First, last, and SVR visit by provider, remainder by pharmacist

### We treat... everyone

#### Treatment candidacy

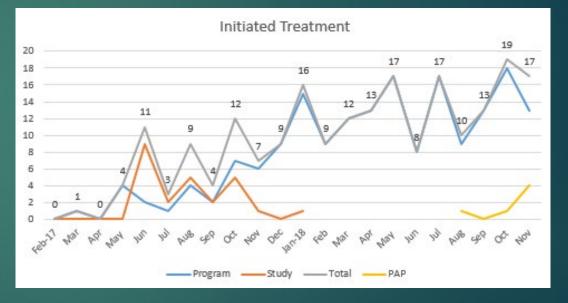
- Made 2/3 last appointments <u>or</u> subjective adherence measure (whichever lower barrier)
- Desires treatment
- Any engagement in treatment for substance use disorder, usually in primary care setting

## HCV Tx at OTC: ~2 years in

#### Patients Referred

#### Patients Initiating Tx



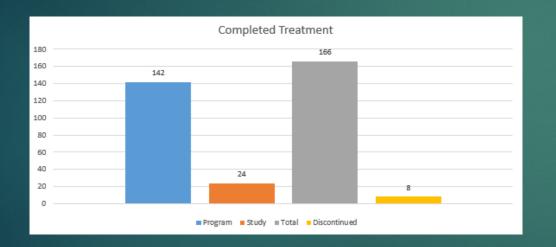


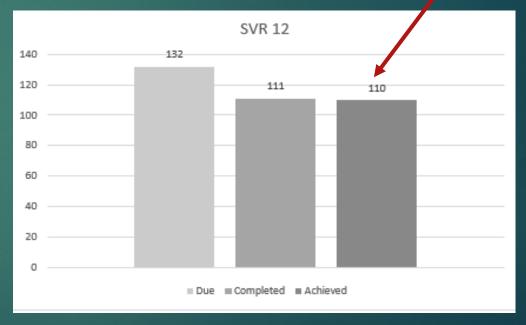
## HCV Tx at OTC: SVR12

#### Patients Completing Tx

#### Patients Achieving SVR12

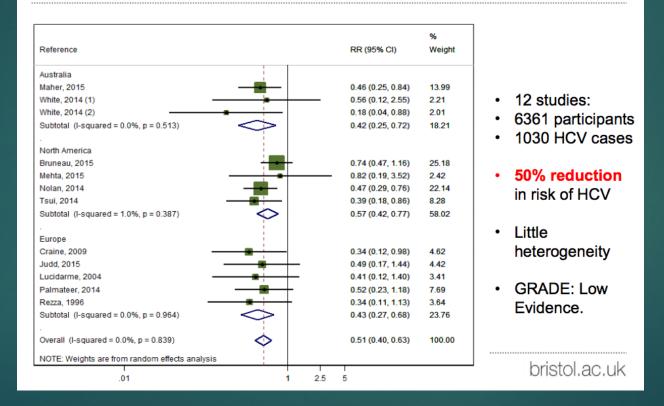
#### 1 tx failure





## OAT associated with 50% reduction HCV transmission

Impact of current OST BRISTOL exposure (adjusted estimates)



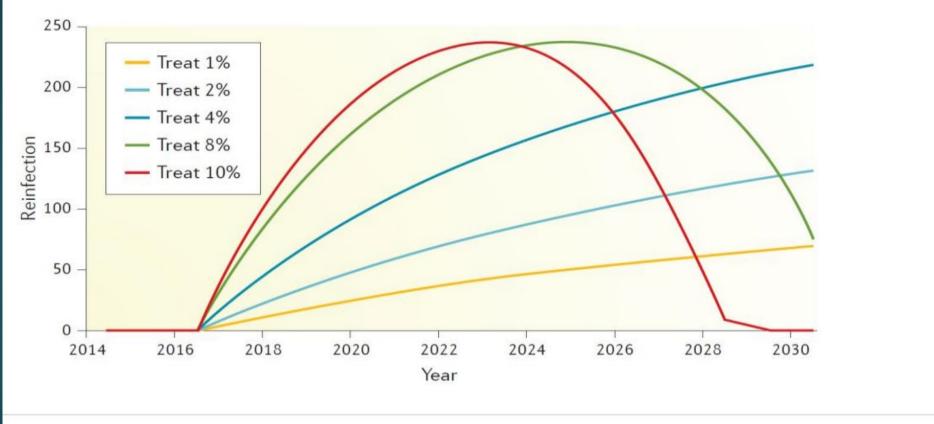
Hickman et al. Lancet Global Health. Vol 5 (2): e1192-e1207.

## NSP associated with 20% reduction HCV transmission

Reference	RR (95% CI)	% Weight	7 studies
North America			
Bruneau, 2015	0.77 (0.50, 1.19)	16.10	High heterogeneit
Hagan, 1999	1.42 (0.64, 3.14)	14.04	(l <sup>2</sup> =79%)
Patrick, 2001	3.69 (2.12, 6.43)	15.48	
Subtotal (I-squared = 89.5%, p = 0.000)	1.58 (0.57, 4.42)	45.62	Weak evidence     overall – RR 0.77
Europe			
Hope, 2011	0.11 (0.02, 0.54)	9.27	<ul> <li>In Europe NSP</li> </ul>
Hope, 2015 (1)		9.31	associated with
Hope, 2015 (2)		4.42	66% reduction in
Hope, 2015 (3)	- <b>1</b> 0.55 (0.05, 6.15)	5.59	HCV
Palmateer, 2014	0.26 (0.08, 0.86)	11.37	1101
Van Den Berg, 2007	<b>0.62 (0.30, 1.29)</b>	14.42	Condex and the
Subtotal (I-squared = 12.3%, p = 0.337)	0.44 (0.24, 0.80)	54.38	<ul> <li>Grade: very low evidence</li> </ul>

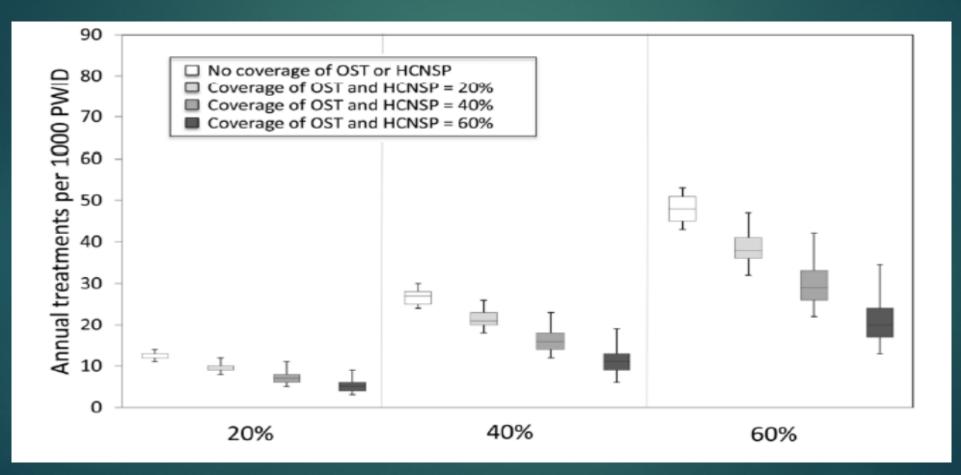
Hickman et al. Lancet Global Health. Vol 5 (2): e1192-e1207.

## Treat more high risk pts, end HCV sooner



Grebely J, Hajarizadeh B, Dore GJ. Nature Reviews Gastro Hepatol 2017

## DAA tx rates to half chronic prevalence in 10 years with HR.



Martin, et al Clinical Inf Diseases 2013. Adapted from Hickman, INHSU 2018.

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### **References** Continued

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