

Epidemiology of HCV Infection Among People who Inject Drugs: Where do we Really Need to Focus Our Efforts to prevent HCV (and generate better evidence)?

Matt Hickman, Sarah Larney, Natasha Martin, Peter Vickerman, Hannah Fraser, Zoe Ward, Louisa Degenhardt

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BRISTOL Acknowledgements & Disclosure/Col

- NIHR Health Protection Research Unit in Evaluation of Interventions
- Health Protection Scotland: HCV Action Plan
- European Commission Drug Prevention and Information Programme (DIPP) "Treatment as Prevention in Europe: Model Projections [JUST/2013/DPIP/AG/4812]
- NIHR (HS&DR) (12/3070/13) Assessing the impact and cost-effectiveness of NSP
- MH received honoraria from Abbvie, MSD, Janssen, Gilead.

The views expressed are those of the author(s) and not necessarily those of the NHS, the NIHR or the Department of Health.

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Global prevalence of injecting drug use and sociodemographic characteristics and prevalence of HIV, HBV, and HCV in people who inject drugs: a multistage systematic review

Prof Louisa Degenhardt, PhD, Amy Peacock, PhD, Samantha Colledge, BPsychSc[Hons], Janni Leung, PhD, Jason Grebely, PhD, Prof Peter Vickerman, PhD, Jack Stone, MMathStat, Evan B Cunningham, BSc[Hons], Adam Trickey, MSc, Kostyantyn Dumchev, MD, Prof Michael Lynskey, PhD, Paul Griffiths, MSc, Prof Richard P Mattick, PhD, Prof Matthew Hickman, PhD, Sarah Larney, PhD

> The Lancet Global Health Volume 5, Issue 12, Pages e1192-e1207 (December 2017) DOI: 10.1016/S2214-109X(17)30375-3

Global, regional, and country-level coverage of interventions to prevent and manage HIV and hepatitis C among people who inject drugs: a systematic review

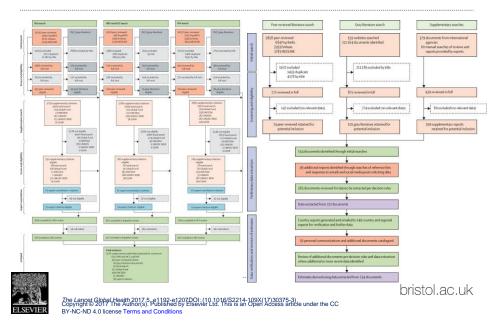
Dr Sarah Larney, PhD, Amy Peacock, PhD, Janni Leung, PhD, Samantha Colledge, BPSc, Prof Matthew Hickman, PhD, Prof Peter Vickerman, PhD, Jason Grebely, PhD, Kostyantyn V Durchev, MD, Paul Griffiths, MSc, Lindsey Hines, PhD, Evan B Cunningham, BSc, Prof Richard P Mattick, PhD, Prof Michael Lynskey, PhD, Prof John Marsden, PhD, Prof Sir John Strang, FMedSci, Prof Louisa Degenhardt, PhD

> The Lancet Global Health Volume 5, Issue 12, Pages e1208-e1220 (December 2017) DOI: 10.1016/S2214-109X(17)30373-X

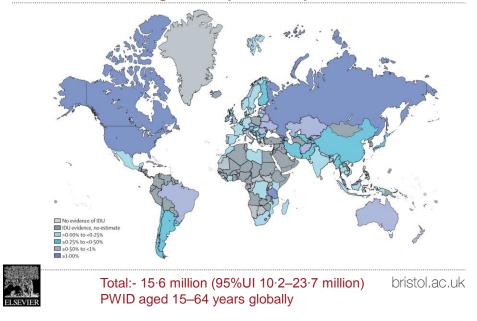


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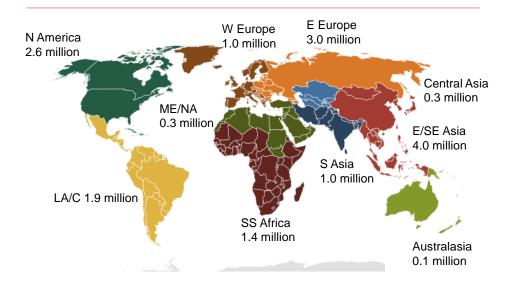
Flowchart presenting no. of sources BRISTOL from identification to inclusion

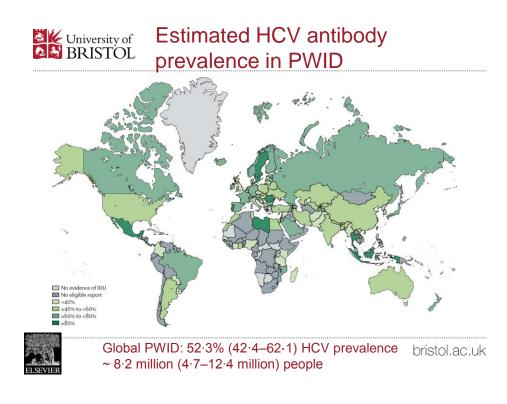


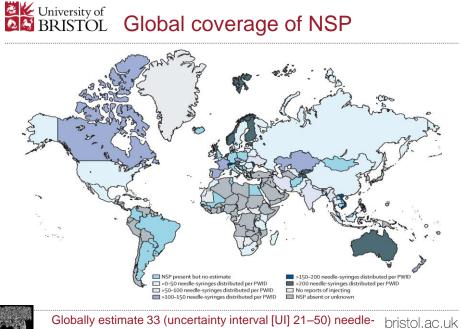
Estimated prevalence of injecting BRISTOL drug use by country

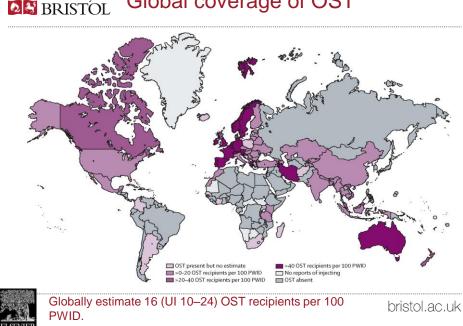


15.6 million people inject drugs globally



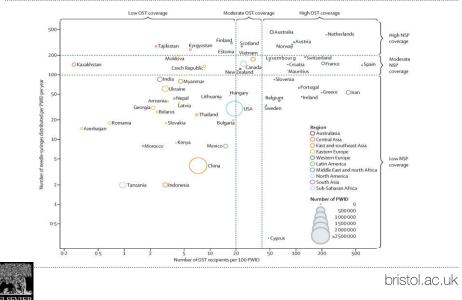






BRISTOL Global coverage of OST



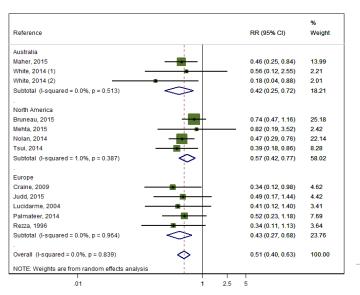




Needle and syringe programmes and opioid substitution therapy for preventing HCV transmission among people who inject drugs: findings from a Cochrane Review and meta-analysis

Lucy Platt¹ ⁽¹⁰⁾, Silvia Minozzi², Jennifer Reed³, Peter Vickerman⁴ ⁽¹⁰⁾, Holly Hagan³, Clare French⁴, Ashly Jordan³, Louisa Degenhardt⁵ ⁽¹⁰⁾, Vivian Hope⁶ ⁽¹⁰⁾, Sharon Hutchinson⁷, Lisa Maher⁸ ⁽¹⁰⁾, Norah Palmateer⁷ ⁽¹⁰⁾, Avril Taylor⁹, Julie Bruneau¹⁰ & Matthew Hickman⁴ ⁽¹⁰⁾

Impact of current OST BRISTOL exposure (adjusted estimates)



- 12 studies:
- 6361 participants
- 1030 HCV cases
- 50% reduction in risk of HCV
- Little
 heterogeneity
- GRADE: Low Evidence.

Impact of high NSP by region University of BRISTOL (unadjusted analyses)

Reference	RR (95% CI)	% Weight	• 7 studies
North America Bruneau, 2015 Hagan, 1999 Patrick, 2001 Subtotal (I-squared = 89.5%, p = 0.000)	 0.77 (0.50, 1.19) 1.42 (0.64, 3.14) 3.69 (2.12, 6.43) 1.58 (0.57, 4.42) 	16.10 14.04 15.48 45.62	 High heterogeneity (I²=79%) Weak evidence overall – RR 0.77
Europe Hope, 2011	 0.11 (0.02, 0.54) 0.99 (0.21, 4.63) 0.73 (0.04, 12.63) 0.55 (0.05, 6.15) 0.26 (0.08, 0.86) 0.62 (0.30, 1.29) 0.44 (0.24, 0.80) 0.77 (0.38, 1.54) 	9.27 9.31 4.42 5.59 11.37 14.42 54.38 100.00	 In Europe NSP associated with 66% reduction in HCV Grade: very low evidence
NOTE: Weights are from random effects analy	1 5		bristol.ac.uk

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University of BRISTOL Impact of NSP and OST

			%
Reference		RR (95% Cl)	Weight
Jnadjusted			
3runeau, 2015	-	0.63 (0.37, 1.07)	35.79
lope, 2011 (1)		0.17 (0.02, 1.54)	10.33
Palmateer, 2014 (2)		0.24 (0.10, 0.59)	27.85
/an Den Berg, 2007 (1)		0.15 (0.06, 0.40)	26.03
Subtotal (I-squared = 64.4%, p = 0.038)	\diamond	0.29 (0.13, 0.65)	100.00
NOTE: Weights are from random effects analysis			
.0	1 1	5	
.0	01 1	5	%
.(Reference	1 1	5 RR (95% CI)	% Weight
)1 1 		
Reference			
Reference Unadjusted		RR (95% CI)	Weight
Reference Unadjusted Hope, 2011 (2)		RR (95% Cl)	Weight
Reference Unadjusted Hope, 2011 (2) Palmateer, 2014 (1)		RR (95% CI) - 1.08 (0.31, 3.82) 0.48 (0.24, 0.95)	Weight 16.66 41.23

1 5 .01

High NSP with OST

- 4 studies
- 3356 participants •
- 518 HCV cases ٠ **Reduced HCV by**

71%

moderate ٠ heterogeneity

Low NSP with OST

- · 3 studies
- 3071 participants ٠
- 449 HCV cases, •
- **Reduced HCV by** • 24%
- · GRADE: low evidence



MODELLING HCV TREATMENT AS PREVENTION

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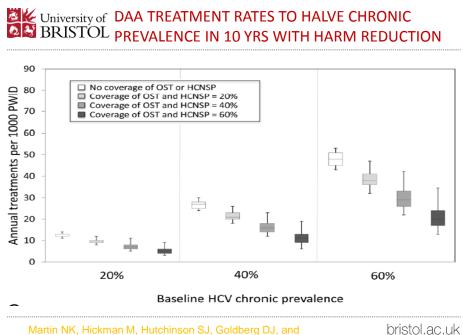




Combination Interventions to Prevent HCV Transmission Among People Who Inject Drugs: Modeling the Impact of Antiviral Treatment, Needle and Syringe Programs, and Opiate Substitution Therapy

Natasha K. Martin,^{1,2} Matthew Hickman,¹ Sharon J. Hutchinson,^{3,4} David J. Goldberg,⁴ and Peter Vickerman²

Modeling HCV Combination Interventions • CID 2013:57 (Suppl 2) • S39



Vickerman P. C. Clinical Infectious Diseases 2013

University of BRISTOL

ARE CURRENT HCV TREATMENT RATES SUFFICIENT?



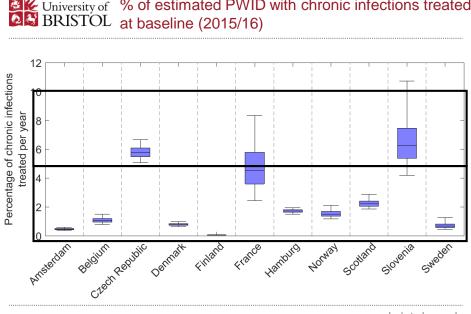
Research Article Viral Hepatitis

JOURNAL **OF HEPATOLOGY**

Model projections on the impact of HCV treatment in the prevention of HCV transmission among people who inject drugs in Europe

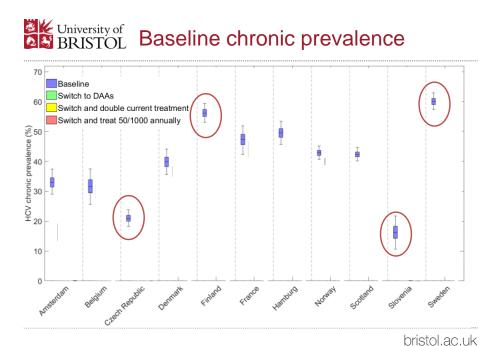
Hannah Fraser^{1,*}, Natasha K. Martin^{2,1}, Henrikki Brummer-Korvenkontio³, Patrizia Carrieri^{4,5}, Olav Dalgard^{6,7}, John Dillon⁸, David Goldberg⁹, Sharon Hutchinson^{10,9}, Marie Jauffret-Roustide^{11,12}, Martin Kåberg¹³, Amy A. Matser^{14,15}, Mojca Matičič^{16,17}, Havard Midgard⁶, Viktor Mravcik^{18,19,20}, Anne Øvrehus²¹, Maria Prins^{14,22}, Jens Reimer^{23,24}, Geert Robaeys^{25,26,27}, Bernd Schulte²⁴, Daniela K. van Santen¹⁴, Ruth Zimmermann²⁸, Peter Vickerman^{1,1}, Matthew Hickman^{1,1}

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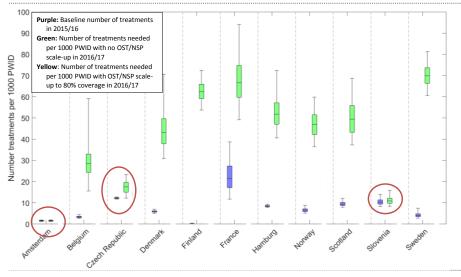
With chronic infections treated at baseline (2015/16)

Fraser et al, (2017) Journal of Hepatology



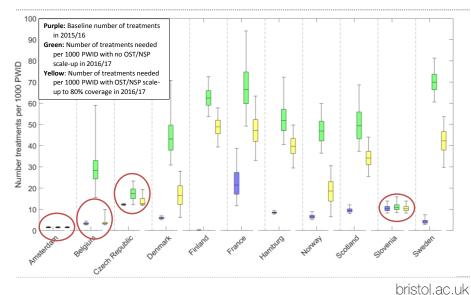
Fraser et al, (2017) Journal of Hepatology





Fraser et al, (2017) Journal of Hepatology, In press

Treatment needed per 1000 PWID to reduce BRISTOL HCV to 2% by 2026 IF scale-up OST/NSP



Fraser et al, (2017) Journal of Hepatology



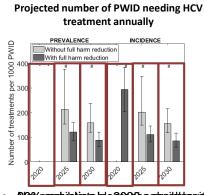


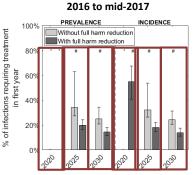
Scaling-up HCV prevention and treatment interventions in rural United States—model projections for tackling an increasing epidemic

Hannah Fraser¹ ⁽¹⁾, Jon Zibbell², Thomas Hoerger², Susan Hariri³, Claudia Vellozzi³, Natasha K. Martin^{4,1}, Alex H. Kral² ⁽¹⁾, Matthew Hickman¹ ⁽¹⁾, John W. Ward³ & Peter Vickerman¹ ⁽¹⁾



Treatment scale-up to reduce HCV to low levels



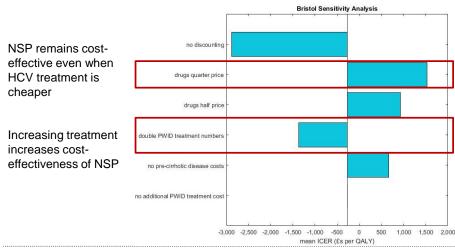


Percentage of infections treated in mid-

- 900% pressbilde to to by 2028 popyeid be available of the state of the
- 90% decrease inincidence may be possible (65%) of simulations) if MATU and SSP and even up.

Only a pupplinul for sets achieved the target. * <5% of parameter sets achieved the target.

BRISTOL NSP Cost-effectiveness



Ward et al, (2018) Addiction in press.

BRISTOL Implications – mixture of evidence

- Global evidence that OST/NSP coverage is low
 - BUT evidence incomplete especially in LMIC
- Empirical evidence OST/NSP reduces HCV
 - NSP & OST highly cost-effective
 - BUT evidence considered moderate/weak and needs to be strengthened
- Dynamic and Economic Models show that:
 - HCV treatment scale-up critical for HCV prevention
 - OST/NSP avert HCV infections & increase TasP impact
 - Early treatment of PWID cost-effective

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BRISTOL Implications – mixture of evidence

- Model projections suggest treatment scale-up needed to observe change in HCV transmission
 - Substantial increases needed in most countries to achieve WHO elimination targets
 - Strengthen PH surveillance so can detect change
- Promote HCV TasP and OST/NSP scale-up to prevent HCV transmission - EVALUATE
 - Natural experiments
 - Revive and strengthen evidence for Harm Reduction
 - Re-infection density case control

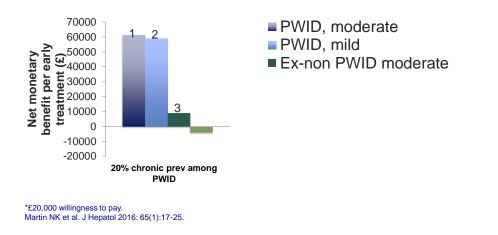


Image: Distribution of HCV treatment in the direct-acting antiviral era: Lace to Material 20

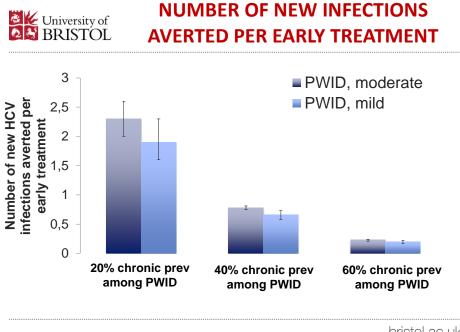
Natasha K. Martin^{1,2,*}, Peter Vickerman², Gregory J. Dore³, Jason Grebely³, Alec Miners⁴, John Cairns⁴, Graham R. Foster⁵, Sharon J. Hutchinson^{6,7}, David J. Goldberg^{6,7}, Thomas C.S. Martin⁸, Mary Ramsay⁹, the STOP-HCV Consortium, Matthew Hickman²

¹Division of Global Public Health, University of California San Diego, San Diego, USA; ²School of Social and Community Medicine, University of Bristol, UK; ³Kirby Institute, UNSW Australia, Sydney, Australia; ⁴Faculty of Public Health and Policy, London School of Hygiene and Tropical Medicine, UK; ⁵Queen Mary's University of London, UK; ⁶Glasgow Caledonian University, UK; ⁷Health Protection Scotland, UK; ⁸Guy's and St Thomas's NHS Foundation Trust, London, UK; ⁹Public Health England, UK

Image: Construction of the second s



Economic modeling supports treatment for and prioritization of PWID – essential for achieving elimination targets



Martin NK et al. J Hepatology 2016