

## Needle syringe programmes and opioid substitution therapy for prevention HCV transmission among people who inject drugs: Cochrane Systematic Review

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#### Acknowledgements & Disclosure/Col

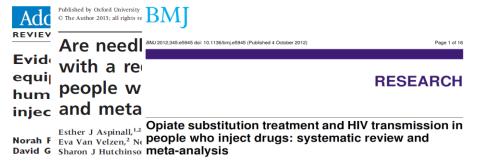
- NIHR (HS&DR) (12/3070/13) Assessing the impact and cost-effectiveness of NSP
- NIHR Health Protection Research Unit in Evaluation of Interventions
- European Commission Drug Prevention and Information Programme (DIPP)
  "Treatment as Prevention in Europe: Model Projections
  [JUST/2013/DPIP/AG/4812]
- 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.



#### Background

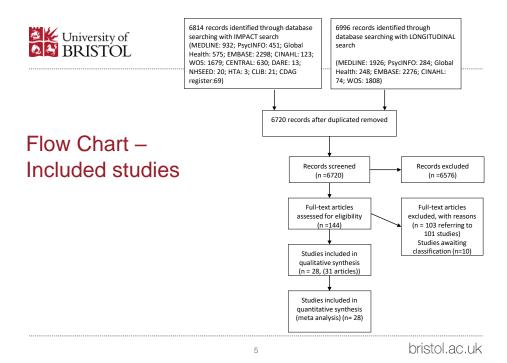
- NSP review level evidence:-
  - sufficient that reduce injecting related risk behaviour
  - insufficient that prevent HCV transmission;
  - tentative that prevent HIV transmission;
- OST associated with reduce risk of HIV transmission





#### Methods

- Search
  - Medline, PsycInfo, Embase, Cochrane and Web of Science up to July 2014; Reference lists; Grey literature
- Inclusion
  - observational and intervention studies measuring exposure to NSP and/or OST (compared to no intervention) among PWID and HCV incidence
- Risk of bias: <u>www.riskofbias.info/</u>
  - Studies classified as Low, Moderate, Serious, Critical risk according to 7 domains
  - · Critical confounders: duration of injection or age; injecting frequency
- Analyses
  - Meta-analysis using random effects models & heterogeneity explored
  - Separate analyses for adjusted and unadjusted, exclusion of studies classified as 'critical' and subgroup analyses, geographical region, recruitment site, type of drug injected, frequency of injecting





### Description of studies - I

- 28 studies
  - · 21 published, 7 unpublished
  - 13 North America, 5 UK, 4 other Europe, 5 Australia and 1 China
- 1817 HCV infections 8806.95py follow-up.
  - HCV incidence 0.09 to 42 cases per 100 py
- · Risk of Bias
  - Only 2 at moderate overall risk of bias, 17 at serious risk, 7 at critical risk; 2 unpublished studies no RoB
- No RCT intervention effects so typically evidence quality categorised as low.



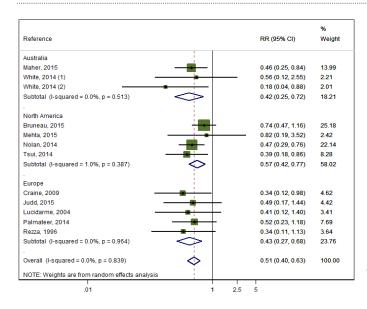
#### Description of studies - II

- Interventions
  - current OST (within last 3-6 months)
  - High NSP coverage (regular NSP attendance/ all injections covered by a new needle/syringe)
- Variation across studies:
  - Sample size (range 46-2788)
  - Method of recruitment: street outreach; RDS; snowball sampling; service users
  - Study design: Prospective cohort (20); Cross-sectional (3); case-control studies (2); 2 retrospective cohort, 1 serial crosssectional
  - Published 1995-2014
  - Follow-up time 1 and 22 years

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# Impact of current OST exposure (adjusted estimates)



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



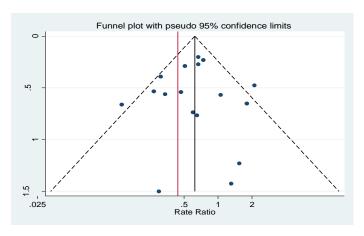
### Impact of current OST

- Effect maintained if exclude unpublished estimates, studies at critical or no information on risk of bias, cross-sectional studies
  - Effect strengthened if exclude unpublished estimates and weakened slightly if pool unadjusted studies (10,647 participants, RR =0.57, I<sup>2</sup>=32.4)
- No evidence that effectiveness varied by geographic region or site of recruitment.
  - Intervention effect reduced with 10% increase in sample of PWID who were women

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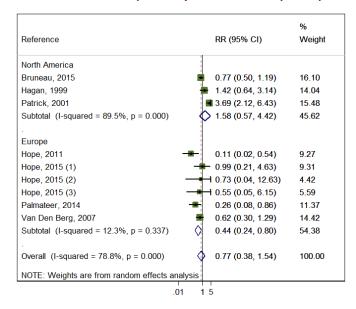
## No evidence of publication bias



Funnel plot – OST meta-analysis - Egger's bias coefficient (-0.87 p=0.106)



## Impact of high NSP by region (unadjusted analyses)

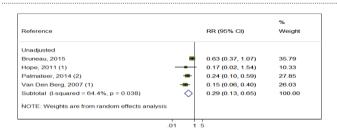


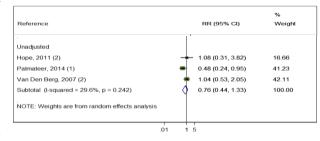
- 7 studies
- High heterogeneity (I<sup>2</sup>=79%)
- Weak evidence overall – RR 0.77
- In Europe NSP associated with 66% reduction in HCV
- Grade: very low evidence

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### Impact of NSP and OST





#### **High NSP with OST**

- · 4 studies
- 3356 participants
- 518 HCV cases Reduced HCV by 71%
- moderate heterogeneity

#### **Low NSP with OST**

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



#### Impact of current NSP

- Meta-regression
  - differential impact by region N. America vs Europe (Ratio of rate ratios= 3.73, p=0.06).
- No evidence
  - publication bias
  - differential impact by proportion of female participants in the sample, homelessness or experience of prison.

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## Impact of current NSP: Key limitations & potential explanations

- · heterogeneity in measurement of NSP exposure
  - In North America studies had varied definitions of NSP use relating to frequency of attendance at NSPs.
  - In Europe studies tend to measure 100% NSP coverage "person reports that they receive sufficient or a greater number of sterile syringes per injecting frequency"
- confounding/ selection bias
  - lower overall coverage in US may mask intervention effect
  - higher proportion of stimulant injectors in US studies that without OST may contribute to lower impact
  - regular NSP attenders at greater risk of HCV acquisition
  - US studies conducted when incidence was higher



#### **Implications**

- strong consistent evidence that OST reduces HCV transmission
- weaker evidence for high coverage NSP
  - more heterogeneity
  - NSP highly cost effective/ cost saving
    - See Zoe Ward talk
- corroborates importance of combining interventions (NSP and OST)
  - Model evidence that OST/NSP enhance HCV TasP & minimize re-infection

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#### **Implications**

- Lack of published data from outside Europe, (Australia) and North America
- Studies with high level of bias
  - No RCT possible
- Global coverage estimates confirm that NSP/OST scale-up needed in most countries
- Improve reporting of observational studies
  - Better natural experiments
  - Strengthen evidence base as improve coverage



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#### **Review information**

Review type: Intervention

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