STUDY PROTOCOL

Risk factors for completed suicide among people who use drugs: A scoping review protocol [version 2; peer review: 1 approved, 2 approved with reservations]

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Abstract

Background: Research over the past several decades has shown an increased risk for completed suicide among people who use drugs (PWUD). However, no study to date has attempted to summarise the available literature on the variety of risk factors associated with this increased risk. This paper presents a protocol for a scoping review that aims to systematically map and synthesise the extent and nature of published, unpublished and grey literature related to risk factors for suicide among PWUD.

Methods: The following six-stage methodological framework for scoping reviews proposed by Arksey and O'Malley with enhancements by Levac and colleagues will be used: (1) identifying the research question, (2) identifying relevant studies, (3) study selection, (4) charting/mapping the data, (5) collating, summarising and reporting results and (6) expert consultation. The review will be conducted and reported in accordance with the PRISMA Extension for Scoping Reviews (PRISMA-ScR). Key inclusion and exclusion criteria will be developed to guide literature screening and data charting. Three reviewers will conduct the initial screening of published, unpublished and grey literature. Identified risk factors will be collated, summarised and categorised iteratively by two independent reviewers. Stakeholder consultation will occur with experts from a national steering committee, a national advisory group, a national suicide prevention centre and a European drug monitoring centre.

Conclusion: Collating and thematically categorising the various risk factors for suicide among this high-risk group will hold important implications for future research, policy and practice. The research will be disseminated through publication in a peer-reviewed academic journal and a conference presentation, and by sharing the findings with key stakeholders working within research, policy-making and professional practice contexts.
**Keywords**
Suicide, people who use drugs, substance-related disorders, risk factors, scoping review

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Amendments from Version 1

In this updated version of the protocol, we have included a more detailed overview of how the data will be collated and mapped (i.e. according to modifiable and non-modifiable risk factors, proximal and distal risk factors, type of drugs used and quantity of drugs used). We have also acknowledged that some risk factors may vary as a function of others, which will be considered during the data collating process.

The definition of the population is quite broad. However, the inconsistent use of terms in the literature related to people who use drugs makes it quite difficult to develop a strict inclusion/exclusion criterion around population type (e.g. studies referring to people with substance dependence vs. those referring to drug users). We included a note that, where possible, the data would be collated and interpreted according to population type.

Both reviewers expressed concern over the lack of quality review process incorporated into the protocol. However, performing quality appraisal as part of a scoping review is debated in the literature. For our purposes, it is important to obtain a broad sense of the field given that there has been little empirical research to date. In this sense, we anticipate that a substantial proportion of our final sources will come from the grey literature - including policy papers, reports, expert input etc. We believe that quality appraisal would not make a substantial contribution to the overall findings of this particular review. However, we have amended objective three of the review in the Introduction section to acknowledge that how the findings might inform policy and practice should be considered in the absence of a review of evidence quality.

Finally, we clarified the stakeholders that we will consult, including the WHO and a steering committee that includes community representation for families affected by drug-related deaths.

Any further responses from the reviewers can be found at the end of the article.

Introduction

Suicide is a significant public health concern. According to the World Health Organisation (WHO), over 800,000 people die by suicide every year, which amounts to approximately one suicide death every 40 seconds1,2, and 34 million years of life lost annually3. In 2016, suicide accounted for 1.4% of all deaths worldwide, making it the 18th leading cause of death globally4.

In addition to the significant emotional toll experienced by the families and friends of people who die by suicide, there are considerable economic costs to suicide incurred by individuals, families and society more generally. This includes direct monetary costs linked to suicide (e.g. the cost of emergency services, medical care, medicolegal costs and funeral expenses) and indirect costs associated with loss of life (e.g. loss in productive activity and loss of earnings due to premature mortality)5-7. In its Comprehensive Mental Health Action Plan8, the WHO commit to reducing global suicide mortality by 10% between 2012 and 2020. Under objective three of this plan (‘to implement strategies for the promotion and prevention in mental health’), a defined action for WHO Member States is the development and implementation of a comprehensive national suicide prevention strategy, with particular emphasis on vulnerable groups9.

Among those populations known to be at increased risk of suicide are people who use drugs (PWUD)10-17. Evidence from epidemiological and clinical research indicates a 7- to 22-fold increase in suicide mortality among PWUD relative to that expected in the general population14-17. The factors that contribute to suicide are complex, wide-ranging and multi-faceted18,19, acting at multiple levels (i.e. individual, familial, communal and societal) and varying across groups and over time20. Here, we outline a protocol for a scoping review with the primary aim of providing a comprehensive overview of existing literature on risk factors for completed suicide, specifically among PWUD. The particular objectives of the review are:

(a) To map the extent, range and nature of available evidence on risk factors for suicide among PWUD.

(b) To identify knowledge gaps and limitations in this body of literature and make recommendations for addressing them.

(c) To inform suicide prevention policy and best practice guidelines for working with PWUD.

Study rationale

A scoping review approach was deemed suitable for several reasons. Notwithstanding several (unsystematic) literature reviews13,14,21-24 on risk factors for suicide among PWUD, and previous systematic reviews22,25-27 and meta-analyses3,15,28 that aimed to quantify the association of problem drug use with suicide mortality, no study has sought to systematically identify and thematically map the available evidence on risk factors for suicide among PWUD. This is surprising, given the breadth of empirical research on the predictors, patterns, outcomes and implications of problem drug use29-33, academic consensus that problem drug use remains a significant risk factor for suicide34,35 and widespread recognition of PWUD as a high-risk group for suicide in national and international reports and suicide prevention strategies36,37.

Consequently, there is limited clarity on the extent (i.e. size and breadth), range (i.e. variety) and nature (i.e. characteristics and contexts)38 of the evidence regarding risk factors for suicide among PWUD, as well as ambiguity regarding the overall progress and direction of this field of research. Scoping reviews are an increasingly popular form of knowledge synthesis that aim to systematically search and map the breadth of available evidence (including evidence in published and grey literature), categorise key concepts, identify knowledge gaps and research deficits, and propose recommendations to guide future research39,40. In this sense, a scoping review is an ideal approach toward a comprehensive understanding of suicide among PWUD, particularly given the breadth of grey literature (e.g. policy papers, governmental and organisational reports, etc.) on this topic. Moreover, limited clarity on risk factors for suicide among PWUD has implications for policy and practice; understanding risk and the contexts in which risk may be amplified are critical precursors to developing targeted interventions and prevention strategies for any group, including PWUD39. A key characteristic of the scoping review is the incorporation of stakeholder consultation into the methodological framework to both inform and validate the study findings39. This process provides opportunity
for knowledge transfer and exchange with experts working at the intersection of research, policy and practice.

Protocol

The review will be guided by the methodological framework for scoping reviews outlined by Arksey and O’Malley48, with subsequent enhancements by Levac and colleagues49. This framework involves six stages, which are discussed in further detail below: (1) identifying the research question, (2) identifying relevant studies, (3) selecting studies, (4) mapping/charting the data, (5) collating, summarizing, and reporting the results, and (6) expert consultation. The scoping review will also be reported in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) Extension for Scoping Reviews (PRISMA-ScR)48.

Stage 1: Identifying the research question

Scoping review research questions are expected to be broad enough in nature to capture the breadth of research on a given topic48, while also encompassing a clearly articulated scope of enquiry49. As per PRISMA-ScR guidelines48, the research question was guided by the Population, Concept, Context (PCC) mnemonic48, albeit tailored to include the outcome of interest for the purposes of this scoping review. Thus, the following research question was identified based on the overarching aim of the scoping review: What is the extent, range, and nature of evidence regarding risk factors for suicide among PWUD? Addressing this research question will allow us to ascertain current knowledge gaps and research deficits in this field of research and propose recommendations for future research, policy and practice.

Population. The review team note the variety of terms used to describe PWUD in this body of literature. For the purposes of the scoping review, PWUD is considered an umbrella phrase under which various terms indicative of problem drug use are subsumed, including, but not limited to, any of the following:

(a) People who use, misuse, or abuse drugs (including non-medical use of prescribable drugs and illicit drug use).
(b) People with a diagnosis of substance use disorder (SUD) / drug use disorder (DUD).
(c) People with drug dependence.
(d) People who are regular or ‘casual’ users of drugs.
(e) People who report recent drug use.

Concept. For the purposes of the scoping review, a risk factor is defined as any characteristic, fixed or modifiable, that makes it more likely that an individual will die by suicide48. Given the wide range of possible risk factors48, the scope of the review will be kept deliberately broad so that risk factor categories emerge during the process of the systematic search and data charting. However, risk factors for suicide can be broadly grouped as occurring at the individual, relationship, community, society, and systemic levels48. We anticipate that some evidence sources may not explicitly identify a correlate or predictor of suicide as a risk factor per se, and so decisions on the parameters of what constitutes a risk factor will be made iteratively as we become increasingly familiar with the breadth and nature of the field.

Outcome. In relation to suicide as an outcome for PWUD, studies may refer to one or more of the following:

(a) Suicide, defined as “Different manners of non-natural death have different numbers of undetermined cases in terms of intent; for example, in a hanging or a shooting it is usually easy to differentiate between a suicide or a trauma (or a crime), while for drowning, traffic accidents or intoxication it is more difficult. Circumstantial findings, such as suicide notes, expressed intent or other findings such as self-inflicted cutting of the wrist followed by drowning, are suggestive of the intent” (E950-953, X60-84-ICD–International Classification of Disease–9 and 10).
(b) Undetermined suicide, defined as “When crime can be ruled out and it cannot be established whether the manner of death is a suicide or an accident, the manner of death is recorded as death of undetermined intent” (E980, Y10-34, ICD–International Classification of Disease–9 and 10).
(c) Probable suicide. Following previous recommendations42-44, all deaths with a diagnosis of suicide or undetermined suicide can be considered probable suicide deaths.

The scoping review methodology is an iterative process49, and so the research question may be refined, or additional questions identified, as the review team become increasingly familiar with the body of literature.

Stage 2: Identifying relevant research

Information sources and search strategy. A comprehensive search strategy to identify relevant literature will be developed in consultation with a health information specialist, and in accordance with the Joanna Briggs Institute (JBI) Reviewer’s Manual for scoping reviews48. The strategy will involve systematic searching of published, unpublished and grey literature, and the entire search strategy, including the rationale behind any decisions made, will be included in the final manuscript.

As recommended48, a three-step strategy will be utilised to identify published literature. To ensure that all appropriate index terms (i.e. MeSH), keywords and phrases are included in the main search, Step 1 involves an initial limited search of two electronic databases and an analysis of the keywords and phrases contained in the titles and abstracts of retrieved papers, as well as of the index terms used to describe the articles. The following two electronic databases will be searched: Medline (EBSCO) and CINAHL (EBSCO). Search strings combining keywords, phrases and index terms using Boolean operators will be developed in collaboration with a health information specialist. In Step 2, the search strings will be adapted and applied across all included databases, which are: Medline (EBSCO), CINAHL (EBSCO), PsycINFO (Ovid), SOCIndex (EBSCO), the Cochrane Database of Systematic Review and the Campbell Collaboration Database of Systematic Reviews. Several key academic journals will be
identified by the review team and hand searched for relevant published articles that may not be returned in database searching. This step includes initial screen of all titles/abstracts returned from database searching, followed by full-text screening of all evidence sources that meet the eligibility criteria (see below). Following full-text screening, Step 3 involves searching the reference lists of all evidence sources included in the review for additional sources missed in Step 2.

The review team are aware of the breadth of potential sources of grey and unpublished literature. To ensure a systematic search of grey and unpublished literature relating to risk factors for suicide among PWUD, a number of steps will be taken. First, grey literature databases (e.g. Open Grey) will be searched using keywords and phrases identified in published literature. As per previous recommendations, only the first 100 hits (as sorted by relevance) from searches performed using grey literature search engines will be screened, as further screening will unlikely result in additional relevant literature. Second, specific types of evidence sources will be sought and screened, including abstracts submitted to flagship conferences on both substance abuse and suicide, postgraduate theses and dissertations (e.g. Electronic Theses Online Service), preprints (e.g. OSF Preprints), policy documents and governmental and organisational reports. Finally, the review team will attempt to contact academic experts, professional societies and relevant organisations to ascertain the availability of any additional evidence sources not identified in previous searches of published, unpublished or grey literature.

**Eligibility criteria.** All peer-reviewed and non-peer-reviewed articles and reports published in the English language will be eligible for inclusion in the review. Careful consideration was given to the timeframe of the search; as one objective of the scoping review is to make recommendations for policy and practice, more up-to-date research was considered most appropriate. Therefore, searches will be limited to evidence sources published over the past 20 years, between 2000 and 2020, inclusive. Review articles that provide new insights will be eligible for inclusion. This includes evidence syntheses, narrative reviews, rapid reviews, systematic reviews and meta-analyses. No limitations will be placed on study design (i.e. cross-sectional, case-control, cohort, prospective, retrospective studies etc. will all be eligible for inclusion). Where the review team identify evidence sources with obvious overlap in either participant samples (e.g. multiple publications from the same prospective study) or datasets (e.g. studies or reports that draw from the same dataset or information system), sources that provide the most information relevant to the aims of the scoping review will be prioritised. Thus, any articles, reports and evidence syntheses that do not provide new information will be excluded.

As per PRISMA-ScR guidelines, the search strategy will also be underpinned by key eligibility criteria based on the PCC mnemonic, which again includes the primary outcome of interest for purposes of this scoping review. Inclusion and exclusion criteria relating to (a) Population (PWUD), (b) Concept (risk factors), (c) Outcome (suicide), and (d) Context (region, drug treatment settings, etc.) are listed in Table 1. It is noted that these criteria may be refined throughout the process of the scoping review owing to increasing familiarity with the body of literature and subject matter.

**Stage 3: Selecting studies**

All search results will be imported into Endnote X7 (Mendelay reference manager software can be used as a freely available alternative) and any duplicates removed. As per scoping review guidelines, reviewers will meet at the start, during and at the end of each stage of selecting studies and any disagreements on article inclusion will be discussed. Firstly, the entire review team will independently apply the eligibility criteria to a random sample of 25 titles/abstracts and then meet to discuss discrepancies and make modifications to the criteria to ensure complete agreement, if required. Next, one reviewer (LM) will apply the eligibility criteria to all titles/abstracts of all retrieved sources; those that are deemed unsuitable for progression to full-text review will be excluded and the reason for exclusion recorded. Two second reviewers (SL and MO’S) will then independently review all excluded titles/abstracts to ensure accuracy. Results will be compared between all reviewers until consensus is reached. If conflict remains, a fourth reviewer (EL) will be consulted until consensus is reached. Finally, two independent reviewers (LM and SL or MO’S) will independently apply the eligibility criteria to full text publications; those that are deemed unsuitable for progression to Stage 4 (mapping/charting the data) will be excluded and the reason for exclusion recorded. Any disagreements will be discussed and, if required, a third reviewer (EL) will be consulted until full consensus on inclusion and exclusion is achieved.

Throughout this selection process, queries for discussion will be recorded and all queries and associated verdicts will be included in the final manuscript as an appendix. Reasons for exclusion of sources following title/abstract and full text review will be reported in the PRISMA-ScR flow diagram in the final manuscript.

**Stage 4: Mapping/charting the data**

In scoping review methodology, data charting is the process of extracting relevant data from sources deemed eligible for inclusion. A bespoke data charting tool will be developed by the review team a priori, guided by recommendations pertaining to data charting in the JBI Reviewer’s Manual for scoping reviews and by the specific aims and objectives of the review. The following types of information will be collected: study characteristics (e.g. year of publication and country), the overall aim/purpose of the study/report, the study design, study/report setting, population characteristics (e.g. age, gender and ethnicity), the use of diagnostic inclusion criteria for drug use (e.g. DSM or other criteria) or the authors definition of drug use, the presence/absence of a control or comparison group, the definition of suicide (e.g. probable or undetermined), the risk factors examined (e.g. correlates or predictors of suicide) and how they were measured, the main findings and information pertaining to the analyses (e.g. adjustments for covariates), the interpretation of the findings, recommendations for future research, policy or practice, and study limitations.
Table 1. Inclusion and exclusion criteria for study selection.

| Included                                                                 | Excluded                                                                 |
|--------------------------------------------------------------------------|--------------------------------------------------------------------------|
| **Population: People who use drugs (PWUD)**                              | **Population: People who use drugs (PWUD)**                              |
| • Sources in which it is made explicit that the participant group (or a subgroup) were PWUD | • Sources in which it is not made explicit that the deceased (or a subgroup) were PWUD |
| • Sources that include a participant group (or subgroup) who use, abuse or are dependent on drugs only OR | • Sources that include a participant group (or subgroup) who use, abuse or are dependent on alcohol only OR |
| • Sources that include a participant group (or subgroup) who use, abuse or are dependent on both drugs and alcohol | • Sources in which the participant group (or a subgroup) is only identified as having substance use problems, which could be related to alcohol alone, drug(s) alone or a combination of both |
| • Sources involving adult participants or participants in late adolescence | • Sources involving children or early adolescents (below 15) only |

| **Concept: Risk factors**                                                | **Concept: Risk factors**                                                |
|--------------------------------------------------------------------------|--------------------------------------------------------------------------|
| • Sources that explicitly identify a variable, or several variables, as risk factors for suicide among PWUD OR | • Sources that do not explicitly analyse risk factors for suicide among PWUD OR |
| • Sources in which risk factors can be inferred (e.g. sources that report sex segregated data) | • Sources in which risk factors cannot be inferred |

| **Outcome: Suicide**                                                     | **Outcome: Suicide**                                                     |
|--------------------------------------------------------------------------|--------------------------------------------------------------------------|
| • Sources in which the primary outcome variable (or one of several outcome variables) is completed suicide | • Sources that focus on suicide ideation, non-fatal attempted suicide, non-fatal deliberate self-harm, or accidental overdose only OR |
| | OR | • Sources that focus on the cause of mortality among PWUD only OR |
| | OR | • Sources that focus on the means of suicide death only |
| • Sources that include death via overdose (or poisoning) as an outcome and analyse deliberate overdose (or poisoning) deaths as a distinct subgroup | • Sources in which overdose (or poisoning) is a primary outcome but intentionality is not made explicit (i.e. no differentiation between intentional or accidental overdose deaths) |

| **Context**                                                             | **Context**                                                             |
|-------------------------------------------------------------------------|-------------------------------------------------------------------------|
| • Sources that provide insight into risk factors for suicide among PWUD across all settings, including before, during and after drug treatment, psychiatric treatment and incarceration, and other legal or social care contexts | • Sources in which the illicit use of a drug or drugs was solely to complete suicide (i.e. intentional injecting of insulin, which was not prescribed to the individual, for the purpose of completing suicide) |
| • Sources from any geographic region                                     |                                                                         |

Two independent reviewers (LM and MO’S) will pilot the data chart on a random selection of 10 publications, and then meet to discuss the comprehensiveness of the data chart and determine accuracy and consistency in the data being extracted. It is expected that data charting will be an iterative process, although the data chart will be developed a priori, it may be refined throughout the piloting and charting process. Consultation will take place between the reviewers throughout the data charting process. Any disagreements will be discussed and, if required, resolved in consultation with a third reviewer (SL or EL). Data charting will be conducted using Microsoft Excel. Authors of studies will be contacted if further clarification of the information in any evidence source is required. The data chart headings will be presented in a summary table in the final manuscript, and all charted data will be made publicly available.

Stage 5: Collating, summarizing, and reporting the results

The data will be collated and summarised in accordance with the overall aim and objectives of the scoping review and the PRISMA-ScR checklist will be used for reporting the results. The data will be analysed and aggregated using quantitative (i.e. frequency analysis) and qualitative (i.e. thematic analysis) methods. First, the characteristics of included studies will be summarised and presented. This includes information pertaining to the geographic distribution, publication dates, types of population samples and methodologies of included evidence sources, as well as the types of evidence sources available (e.g. empirical research, policy documents, reports etc.). Variation in relation to methodology and study design, among other characteristics, is expected. Second, a thematic analysis of risk factors for suicide...
among PWUD will be conducted and presented, with a focus on the types of risk factors being examined and the key variables/concepts within each risk factor category. Where possible, trends will be analysed according to specific drugs, as well as the number of drugs reported, albeit this is contingent upon the scope of the individual studies included in the review. We anticipate that certain risk factors (e.g. age range) may vary as a function of other factors (e.g. sex), and this will be a consideration during the data mapping and collating processes. Risk factors will be characterised as modifiable (e.g. current drug use) or non-modifiable (family history of drug use) during data mapping and collation, and distinctions will be drawn between proximal (e.g. polydrug use) and distal (early traumatic experience) risk factors. Moreover, evidence sources will be categorised and analysed according to the description of the population provided. For instance, evidence sources that specifically focus on people with drug dependence or drug use disorder will be analysed separately and compared to those that refer to ‘drug users’ or ‘people who use drugs’, without an indication of the severity of use.

Finally, the results will conclude with a narrative overview of research limitations, knowledge gaps and areas have been under-researched to inform directions for future research and considerations for policy and practice. Recommendations for policy and practice extracted from reports and policy documents will also be charted, summarised and integrated into the review findings. It is anticipated that the process of mapping and analysing the data will be an iterative one, with enrichment and refinement of the review findings and resulting recommendations following expert consultation (see below). The methods used to collate and summarise the data, as well as the rationale behind all decisions pertaining to data handling and analysis, will be described in detail in the final manuscript.

Stage 6: Expert consultation
Expert consultation is an important component of the scoping review methodology as a means by which to engage important stakeholders with expertise in research, policy and practice, and thus enhance the methodological rigour and applicability of the review
Consultations will occur with national experts from the Irish National Drug-Related Deaths Index (NDRDI) Steering Committee, which includes community representation for families affected by drug-related deaths, and the Technical Advisory Group of the National Office of Suicide Prevention, as well as international experts from the European Monitoring Centre for Drugs and Drug Addiction (EMCDDA) and the World Health Organization. Finally, experts from other relevant organisations, professional societies, research centres and institutes will be contacted (e.g. the National Suicide Research Foundation in Ireland, the National Drug Research Institute in Australia, and Harm Reduction International). As recommended, the preliminary findings from Stage 5 will be used as a foundation from which to inform consultations
The purposes of expert consultation in this instance are three-fold. Specifically:

(a) To obtain additional input regarding important evidence sources not identified in published and grey literature searches.

(b) To gain perspectives and insights beyond those acquired during data charting and analysis, as well as suggestions for knowledge translation.

(c) To present the preliminary findings of the scoping review to research and policy stakeholders in the field.

The insights acquired throughout this phase will be analysed, interpreted and integrated into the review findings and recommendations.

Study status
At the time of publication of this protocol, informal preliminary searches of the literature had been undertaken primarily to help to identity keywords, phrases and index terms in order to develop the full search strategy.

Conclusion and dissemination
The overarching aim of the scoping review is to gain a comprehensive understanding of the literature on risk factors for suicide among PWUD over the past two decades. A scoping review will achieve several important outcomes that will contribute to the overall progress and direction of this field, including a narrative overview of the types of evidence available, a thematic summary of what is currently known about risk factors for suicide among PWUD, identification of knowledge gaps and research limitations, and recommendations for future research avenues, policy development and professional practice. To our knowledge, this will be the first scoping review of risk factors for suicide among this particularly high-risk group. Key strengths of the review will be the use of the most up-to-date methodological guidelines and recommendations and the process of stakeholder consultation. A limitation of the scoping review methodology is that it does not typically include an assessment of publication bias or research quality appraisal; however, relative to other types of reviews such as a systematic review, a scoping review has the capacity to capture the breadth of available evidence on a given topic from a large variety of sources including grey and unpublished literature, ascertain the types of evidence available across the body of literature and make appropriate recommendations for future research, policy and practice.

The results of the scoping review will be disseminated widely. The review will be submitted for publication in a peer reviewed academic journal, presented at an interdisciplinary conference and disseminated to key experts and stakeholders, including those identified during the review process as well as those in the network of the review team.

Data availability
No data are associated with this article.
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Alexandra Pitman
1 Division of Psychiatry, University College London, London, UK
2 Camden and Islington NHS Foundation Trust, London, UK

This study protocol describes the plan to conduct a scoping review of the literature on risk factors for suicide among people who use drugs. It has already had a first round of peer reviewer comments, to which the authors have responded appropriately. The key compromise, as with all scoping reviews, lies in covering the breadth of research but at the expense of time available to conduct critical appraisal of included articles. This has been commented on by other reviewers and responded to.

The searches will start from a date set at 2000, yet as other reviewers point out this seems quite late, and would miss out potentially useful studies. The rationale for this date is that in a previous similar review most of the studies identified were published after 2000, and that the author of a EMCDDA report had observed differences in the patterns of drug use since 1995. However this does not seem a robust justification, unless people had stopped using certain substances. I would suggest an earlier search limit of 1980 given that the outcome is restricted to completed suicide.

A couple of minor comments:

Page 6 mentions thematic analysis as a way of summarising the literature, but this is a qualitative method and would be more appropriate as a narrative synthesis or summary.

Dissemination plans would be improved if a blog was planned with a major non-profit (e.g. Addaction) to gain wide reach beyond academic audiences.

Is the rationale for, and objectives of, the study clearly described?
Yes

Is the study design appropriate for the research question?
Yes
Are sufficient details of the methods provided to allow replication by others?
Yes

Are the datasets clearly presented in a useable and accessible format?
Not applicable

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Suicide and self-harm epidemiology.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Reviewer Report 08 February 2021

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Anne Reneflot
Mental and Physical Health, Norwegian Institute of Public Health, Skøyen, Oslo, Norway

I have no further comments.

Is the rationale for, and objectives of, the study clearly described?
Not applicable

Is the study design appropriate for the research question?
Not applicable

Are sufficient details of the methods provided to allow replication by others?
Not applicable

Are the datasets clearly presented in a useable and accessible format?
Not applicable

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Suicide, epidemiology, demography.

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.
Thank you for the opportunity to review the paper Risk factor for completed suicide among people who use drugs: A scoping review protocol

This paper presents a protocol for a scoping review of the risk factors for completed suicide among people who use drugs. The increased suicide risk associated with problematic drug use is well documented, and the planned scoping review has the potential to make an important contribution to the existing literature. However, we believe the protocol would benefit from some important revisions:

1. **Objectives of the review**: The third objective of the planned scoping review is to inform suicide prevention policy and best practice guidelines for working with PWUD. However, this objective is considerably limited by the lack of quality assessment of eligible studies included in the planned scoping review. We find it particularly problematic that the authors plan to include unpublished and grey literature. This challenge should be acknowledged, and the authors must either adjust the third objective of the planned scoping review, ensure a kind of quality appraisal of the included studies or conduct a systematic review instead.

2. **Definition of the population**: We find the protocol unclear as to which population this scoping review will cover. Will the planned scoping review focus on people who use drugs or on people with problematic drug use? Will the former definition of the population also include people with recreational drug use? We believe the planned scoping review would benefit from limiting the population to people with problematic drug use or a drug abuse disorder. With a very broadly defined population, it will be challenging to identify risk factors particularly pertaining to groups with a high risk of suicide. Further, we question why the research group have decided to exclude people with problematic alcohol use and alcohol use disorder? Is it reasons to believe that the risk factors for suicide in this group is very different from those with a problematic drug use and drug use disorders? At least, we believe this should be discussed in more detail in the protocol.

3. **Eligibility criteria**: The authors have decided to limit the search from 2000 and onwards. Is it reason to believe that risk factors for completed suicide among people who use drugs have changed after the turn of the century? By excluding studies published before year 2000 the authors risk to miss seminal studies. Further, evidence sources with obvious
overlap in either participant samples or datasets, only sources that provide the most
information relevant to the aims of the scoping review will be prioritized. This is common in
systematic reviews, but in a scoping review the aim is among others to map the existing
literature and we believe this restriction is unnecessary.

4. **Outcome**: in the planned scoping review, only studies that examine risk factors associated
with completed suicide will be included. A problem with this strategy is that the authors risk
ending up with very few included studies. We acknowledge that the risk factors for
completed suicide and non-fatal suicidal behavior may differ, but suicide attempt is an
important risk factor and precursor for suicide and it may therefore be informative to
include studies of risk factors associated with suicide attempts as well.

**Is the rationale for, and objectives of, the study clearly described?**
Partly

**Is the study design appropriate for the research question?**
Yes

**Are sufficient details of the methods provided to allow replication by others?**
Yes

**Are the datasets clearly presented in a useable and accessible format?**
Not applicable

*Competing Interests:* No competing interests were disclosed.

*Reviewer Expertise:* Suicide, epidemiology, demography.

We confirm that we have read this submission and believe that we have an appropriate level
of expertise to confirm that it is of an acceptable scientific standard, however we have
significant reservations, as outlined above.

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**Author Response 20 Nov 2020**

Lisa Murphy, Health Research Board, Dublin 2, Ireland

We would like to thank both reviewers for their careful read and review of our manuscript.
We know that this takes significant time and effort and appreciate the recommendations
they have made, which we believe have improved the overall quality of the protocol.

**Reviewer 1: Holly Wilcox**

1. In the Background section of the Abstract suicide add "AMONG" or "in" PWUD.

Many thanks for catching this error, we have now included ‘among’.
2. The intro reports old data - most recent data is from 2018 not 2016.

Assuming the reviewer is referring to WHO data, we have checked the most recent reports and information from the WHO regarding suicide and believe that the 2016 report is the most recent to specifically highlight suicide as accounting for 1.4% of deaths worldwide and being the 18th leading cause of death globally. Subsequent reports that state the same cite this 2016 report. The most recent information provided in a 2019 WHO factsheet indicates that 800,000 deaths can be attributed to suicide and is cited at the very beginning of the introduction section.

3. Great that you are including grey literature and including stakeholders. Who are the stakeholders? Do they include people with lived experience of drug use and suicide attempts or family member who lost a loved one to suicide and drug overdose?

The stakeholders are identified in the ‘Stage 6’ subsection:

“Consultations will occur with national experts from the Irish National Drug-Related Deaths Index (NDRDI) Steering Committee…the Technical Advisory Group of the National Office of Suicide Prevention and international experts from the European Monitoring Centre for Drugs and Drug Addiction (EMCDDA) and the World Health Organization.”

While we very much appreciate the importance of exploring the lived experience of suicide attempts among people who use drugs, it is not the focus of this review. We hope that this review can serve as a steppingstone to continued work on this topic, including exploration of risk factors for attempted suicide among people who use drugs.

Regarding consultation with family members, we have now made clearer in the protocol where this form of consultation will occur (in the ‘Stage 6’ subsection):

“Consultations will occur with national experts from the Irish National Drug-Related Deaths Index (NDRDI) Steering Committee, which includes community representation for families affected by drug-related deaths (including suicides)...”

4. Age range of the population could differ by risk factors.

We agree with the reviewer and have now noted this in the protocol as a factor to consider when mapping and analysing the data (paragraph 2 in the ‘Stage 5’ subsection).

5. In terms of the population will you include those with alcohol use/misuse/abuse? Will you include those who try to intentionally overdose to kill themselves?

The purpose of the review is to examine risk factors for complete suicide among PWUD, which includes people who use both drugs and alcohol, but not people who use/misuse/abuse alcohol only. This is noted in Table 1 which outlines inclusion exclusion
Regarding suicide methods, including sources that refer to intentional overdose, this is also detailed in Table 1 (under ‘Outcome’).

6. It would be ideal to separately list modifiable and non-modifiable risk factors as modifiable risk factors are actionable; it would be ideal to also draw a distinction between proximal and distal risk factors.

Many thanks for this suggestion. We have included a note on (paragraph 2 in the ‘Stage 5’ subsection) to indicate that we intend to characterise risk factors according to these categories during the data charting and analysis process.

7. Will this be done for each drug or drug users lumped together? Impact of the review would be better if results were provided by drug and multidrug users.

Thank you for this suggestion. We agree and have now included a note that when possible, trends will be analysed according to specific drugs as well as the number of drugs reported (paragraph 2 in the ‘Stage 5’ subsection). However, we note that this depends on the level of detail included in the studies included in the review.

8. These groups are ok but if you want this to have international impact (not just Ireland or Europe), WHO seems better: Consultations will occur with national experts from the Irish National Drug-Related Deaths Index (NDRDI) Steering Committee, the Technical Advisory Group of the National Office of Suicide Prevention and international experts from the European Monitoring Centre for Drugs and Drug Addiction (EMCDDA). Those with lived experience should provide input.

Many thanks to the reviewer for drawing our attention to the fact that this stage of the review process is not described clearly in the protocol. We are contacting experts and organisations/groups worldwide, including people at the WHO. This is now revised on in the ‘Stage 6’ subsection.

Regarding including those with lived experience, we refer to our response to the reviewers’ point no. 3 above.

9. It would be ideal to include an assessment of publication bias or research quality appraisal.

Thank you for raising this point. One of the principals of a scoping review is not to include a quality appraisal of the items included, although we acknowledge that this is debated in the literature. We also acknowledge that some researchers conducting scoping reviews are incorporating quality appraisal into their protocol. However, we believe this should be
determined on a case by case basis. For our purposes, it is important to obtain a broad sense of the field, given that there has been little empirical research to date. In this sense, we anticipate that a substantial proportion of our final sources will come from the grey literature - including policy papers, reports, expert input etc. We believe that quality appraisal would not make a substantial contribution to the overall findings of this particular review. However, we will acknowledge this in the discussion section of the review manuscript.

Reviewer 2 & 3: Anne Reneflot & Kim Stene-Larsen

1. Objectives of the review: The third objective of the planned scoping review is to inform suicide prevention policy and best practice guidelines for working with PWUD. However, this objective is considerably limited by the lack of quality assessment of eligible studies included in the planned scoping review. We find it particularly problematic that the authors plan to include unpublished and grey literature. This challenge should be acknowledged, and the authors must either adjust the third objective of the planned scoping review, ensure a kind of quality appraisal of the included studies or conduct a systematic review instead.

Many thanks for raising this point. One of the principals of a scoping review is not to include a quality appraisal of the items included, although we acknowledge that this is debated in the literature. We also acknowledge that some researchers conducting scoping reviews are incorporating quality appraisal into their protocol, but we believe this should be determined on a case by case basis. We have, however, revised the phrasing of the third objective in the Introduction.

For our purposes, it is important to obtain a broad sense of the field given that there has been little empirical / published research to date (hence, limited knowledge can be garnered from the published literature alone). In this sense, we anticipate that a substantial proportion of our final sources will come from the grey literature - including policy papers, government reports, expert input etc, which are used to inform policy in this area, including prevention policy. This is acknowledged in the ‘Conclusion and Dissemination’ section at the end of the protocol and will also be acknowledged in the discussion section of the review manuscript.

2. Definition of the population. We find the protocol unclear as to which population this scoping review will cover. Will the planned scoping review focus on people who use drugs or on people with problematic drug use? Will the former definition of the population also include people with recreational drug use? We believe the planned scoping review would benefit from limiting the population to people with problematic drug use or a drug abuse disorder. With a very broadly defined population, it will be challenging to identify risk factors particularly pertaining to groups with a high risk of suicide. Further, we question why the research group have decided to exclude people with problematic alcohol use and alcohol use disorder? Is it reasons to believe that the
risk factors for suicide in this group is very different from those with a problematic drug use and drug use disorders? At least, we believe this should be discussed in more detail in the protocol.

Many thanks to the reviewers for raising this point. While we agree that there is a lack of clarity regarding the population, the inconsistent use of terms in the literature related to people who used drugs makes it quite difficult (particularly in the first stages of the review process) to develop a strict inclusion/exclusion criterion around population type. For this reason, we have kept the parameters of the population deliberately broad. Once we have identified suitable sources, we will be in a better position to categorise and analyse findings according to population type, e.g. studies that specifically focus on people with drug dependence or drug/substance use disorder compared to those that refer to the population sample (or a subgroup of the sample) as ‘drug users’ or ‘people who use drugs’, without reference to drug use severity. We have now included a note on this in paragraph 2 in the ‘Stage 5’ subsection.

3. Eligibility criteria: The authors have decided to limit the search from 2000 and onwards. Is it reason to believe that risk factors for completed suicide among people who use drugs have changed after the turn of the century? By excluding studies published before year 2000 the authors risk to miss seminal studies. Further, evidence sources with obvious overlap in either participant samples or datasets, only sources that provide the most information relevant to the aims of the scoping review will be prioritized. This is common in systematic reviews, but in a scoping review the aim is among others to map the existing literature and we believe this restriction is unnecessary.

While we appreciate the reviewer’s point regarding the timeframe, our rationale for restricting the timeframe is based on notable changes in drug use patterns and trends over time. Of particular note is a systematic review of literature on the topic of non-medical use of prescription drugs; although no date limitations were applied, the review found that most of the literature was post 2000 (Clark et al., 2015*). Moreover, a recent article from the EMCDDA* showing a change in drug use patterns over time states: ‘The world is a very different place in 2020 than it was in 1995’ ‘Strikingly, across the 25 years of reporting, cannabis has gone from being a drug relatively rarely reported in data from drug treatment registries to being the drug that is now most commonly associated with new treatment entries’.

We believe that such changes in drug use patterns, as well as changes across almost all other aspects of society over time, including policy, practice and health-related services, growing socio-economic inequality, etc. are relevant to the study of suicide as they hold implications for how we characterise and assess risk among different populations, including people who use drugs.

*Clark, M., Hamdi-Ghaz, E., Jauffret-Roustide, M., Le Moigne, P., Melpomeni Malliori, M., Simeoni, E., Jasaitis, E., Jovanovic, M., James, K. & Palczak, K. 2015a. The gender dimension of non-medical use of prescription drugs in Europe and the Mediterranean region. Pompidou
“Looking back on 25 years of annual reporting on the drugs problem in Europe By Paul Griffiths, EMCDDA Scientific Director.

In regards excluding evidence sources with obvious overlap, we emphasise on page 7 (under ‘eligibility criteria’) that any evidence sources that do not provide new information will be excluded. We will retain any sources that provide new insights, even if they use the same data. In instances where two sources draw from the same data and one does not provide any additional insight to the other, we will exclude it. We do not anticipate this to be a major obstacle, however, given the limited amount of empirical work that appears to have been done to date.

4. Outcome: in the planned scoping review, only studies that examine risk factors associated with completed suicide will be included. A problem with this strategy is that the authors risk ending up with very few included studies. We acknowledge that the risk factors for completed suicide and non-fatal suicidal behaviour may differ, but suicide attempt is an important risk factor and precursor for suicide and it may therefore be informative to include studies of risk factors associated with suicide attempts as well.

Many thanks to the reviewers for this suggestion. We agree that ultimately, we may end up with few empirical studies / evidence sources. However, we believe that limiting the review to studies of completed suicide provides a clear purpose and scope. Mapping the available literature regarding risk factors for attempted suicide among PWUD is a very worthwhile but ultimately distinct research endeavour, particularly, as the reviewer notes, since the nature and trends of risk factors will vary depending on the outcome of interest. We hope that this review can serve as a steppingstone to continued work on this topic, including exploration of risk factors for attempted suicide among people who use drugs.

Competing Interests: No competing interests were disclosed.
USA

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Is the rationale for, and objectives of, the study clearly described?
Partly

Is the study design appropriate for the research question?
Partly

Are sufficient details of the methods provided to allow replication by others?
Partly

Are the datasets clearly presented in a useable and accessible format?
Not applicable

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Suicide prevention.

I confirm that I have read this submission and believe that I have an appropriate level of
expertise to confirm that it is of an acceptable scientific standard, however I have significant reservations, as outlined above.

Author Response 20 Nov 2020

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garnered from the published literature alone). In this sense, we anticipate that a substantial proportion of our final sources will come from the grey literature - including policy papers, government reports, expert input etc, which are used to inform policy in this area, including prevention policy. This is acknowledged in the ‘Conclusion and Dissemination’ section at the end of the protocol and will also be acknowledged in the discussion section of the review manuscript.

2. Definition of the population. We find the protocol unclear as to which population this scoping review will cover. Will the planned scoping review focus on people who use drugs or on people with problematic drug use? Will the former definition of the population also include people with recreational drug use? We believe the planned scoping review would benefit from limiting the population to people with problematic drug use or a drug abuse disorder. With a very broadly defined population, it will be challenging to identify risk factors particularly pertaining to groups with a high risk of suicide. Further, we question why the research group have decided to exclude people with problematic alcohol use and alcohol use disorder? Is it reasons to believe that the risk factors for suicide in this group is very different from those with a problematic drug use and drug use disorders? At least, we believe this should be discussed in more detail in the protocol.

Many thanks to the reviewers for raising this point. While we agree that there is a lack of clarity regarding the population, the inconsistent use of terms in the literature related to people who used drugs makes it quite difficult (particularly in the first stages of the review process) to develop a strict inclusion/exclusion criterion around population type. For this reason, we have kept the parameters of the population deliberately broad. Once we have identified suitable sources, we will be in a better position to categorise and analyse findings according to population type, e.g. studies that specifically focus on people with drug dependence or drug/substance use disorder compared to those that refer to the population sample (or a subgroup of the sample) as ‘drug users’ or ‘people who use drugs’, without reference to drug use severity. We have now included a note on this in paragraph 2 in the ‘Stage 5’ subsection.

3. Eligibility criteria: The authors have decided to limit the search from 2000 and onwards. Is it reason to believe that risk factors for completed suicide among people who use drugs have changed after the turn of the century? By excluding studies published before year 2000 the authors risk to miss seminal studies. Further, evidence sources with obvious overlap in either participant samples or datasets, only sources that provide the most information relevant to the aims of the scoping review will be prioritized. This is common in systematic reviews, but in a scoping review the aim is among others to map the existing literature and we believe this restriction is unnecessary.

While we appreciate the reviewer’s point regarding the timeframe, our rationale for restricting the timeframe is based on notable changes in drug use patterns and trends over time. Of particular note is a systematic review of literature on the topic of non-medical use
of prescription drugs; although no date limitations were applied, the review found that most of the literature was post 2000 (Clark et al., 2015*). Moreover, a recent article from the EMCDDA* showing a change in drug use patterns over time states: ‘The world is a very different place in 2020 than it was in 1995’ ‘Strikingly, across the 25 years of reporting, cannabis has gone from being a drug relatively rarely reported in data from drug treatment registries to being the drug that is now most commonly associated with new treatment entries’.

We believe that such changes in drug use patterns, as well as changes across almost all other aspects of society over time, including policy, practice and health-related services, growing socio-economic inequality, etc. are relevant to the study of suicide as they hold implications for how we characterise and assess risk among different populations, including people who use drugs.

*Clark, M., Hamdi-Ghoz, E., Jauffret-Roustide, M., Le Moigne, P., Melpomeni Malliori, M., Simeoni, E., Jasaitis, E., Jovanovic, M., James, K. & Palczak, K. 2015a. The gender dimension of non-medical use of prescription drugs in Europe and the Mediterranean region. Pompidou Group and the Council of Europe. Strasbourg: Council of Europe, Co-operation Group to Combat Drug Abuse and Illicit Trafficking in Drugs (Pompidou Group).

*Looking back on 25 years of annual reporting on the drugs problem in Europe By Paul Griffiths, EMCDDA Scientific Director.

In regards excluding evidence sources with obvious overlap, we emphasise on page 7 (under ‘eligibility criteria’) that any evidence sources that do not provide new information will be excluded. We will retain any sources that provide new insights, even if they use the same data. In instances where two sources draw from the same data and one does not provide any additional insight to the other, we will exclude it. We do not anticipate this to be a major obstacle, however, given the limited amount of empirical work that appears to have been done to date.

4. Outcome: in the planned scoping review, only studies that examine risk factors associated with completed suicide will be included. A problem with this strategy is that the authors risk ending up with very few included studies. We acknowledge that the risk factors for completed suicide and non-fatal suicidal behaviour may differ, but suicide attempt is an important risk factor and precursor for suicide and it may therefore be informative to include studies of risk factors associated with suicide attempts as well.

Many thanks to the reviewers for this suggestion. We agree that ultimately, we may end up with few empirical studies / evidence sources. However, we believe that limiting the review to studies of completed suicide provides a clear purpose and scope. Mapping the available literature regarding risk factors for attempted suicide among PWUD is a very worthwhile but ultimately distinct research endeavour, particularly, as the reviewer notes, since the nature and trends of risk factors will vary depending on the outcome of interest. We hope that this review can serve as a steppingstone to continued work on this topic, including
exploration of risk factors for attempted suicide among people who use drugs.

**Competing Interests:** No competing interests were disclosed.