Health inequalities in infectious diseases: a systematic overview of reviews


ABSTRACT

Objectives The aim of this systematic overview of reviews was to synthesise available evidence on inequalities in infectious disease based on three dimensions of inequalities; inclusion health groups, protected characteristics and socioeconomic inequalities.

Methods We searched MEDLINE, Embase, Web of Science and OpenGrey databases in November 2021. We included reviews published from the year 2000 which examined inequalities in the incidence, prevalence or consequences of infectious diseases based on the dimensions of interest. Our search focused on tuberculosis, HIV, sexually transmitted infections, hepatitis C, vaccination and antimicrobial resistance. However, we also included eligible reviews of any other infectious diseases. We appraised the quality of reviews using the Assessment of Multiple Systematic Reviews V.2 (AMSTAR2) checklist. We conducted a narrative data synthesis.

Results We included 108 reviews in our synthesis covering all the dimensions of inequalities for most of the infectious disease topics of interest, however the quality and volume of review evidence and consistency of their findings varied. The existing literature reviews provide strong evidence that people in inclusion health groups and lower socioeconomic status are consistently at higher risk of infectious diseases, antimicrobial resistance and incomplete/delayed vaccination. In the protected characteristics dimension, ethnicity, and sexual orientation are important factors contributing to inequalities across the various infectious disease topics included in this overview of reviews.

Conclusion We identified many reviews that provide evidence of various types of health inequalities in different infectious diseases, vaccination, and antimicrobial resistance. We also highlight areas where reviews may be lacking. The commonalities in the associations and their directions suggest it might be worth targeting interventions for some high risk-groups that may have benefits across multiple infectious disease outcomes rather than operating purely in infectious disease siloes.

INTRODUCTION

The WHO regards experiencing the highest possible standard of health as a fundamental human right of every individual regardless of personal or social circumstances. Nevertheless, avoidable inequalities exist in the prevalence of diseases and illnesses, general health status and access to healthcare between various social groups. A complex interaction between structural (e.g. income and wealth distribution) and individual-level (eg, health behaviours and living conditions) determinants of health contributes to the increased vulnerability to poorer health among particular social groups.

Infectious diseases pose a significant health burden with substantial health inequalities globally. In the UK, infectious diseases constitute 7% of deaths alongside 4% of lost life years. The economic burden of infectious diseases in the UK is estimated to be around £30 billion per year. Although infectious diseases impose substantial, negative health and economic consequences within populations, many infectious diseases are vaccine-preventable and avoidable through adequate control measures. However, some groups remain under-vaccinated and other
control measures may be difficult or impossible to implement for some, depending on circumstances.9 Traditionally, policymakers often target infectious diseases individually, but it is known that specific groups are often at higher risks regardless of specific infectious diseases.10–14 In efforts to tackle the observed disparities and to reduce the burden of infectious diseases, a strategic approach that tackles infectious diseases among high-risk groups is required.15 To inform the development of needs-tailored public health policies and initiatives to achieve this goal, a comprehensive synthesis of evidence is required, highlighting the inequalities in infectious diseases according to varying personal and social characteristics.

This project was commissioned by Public Health England (PHE) to gain an overview of the available evidence on health inequalities relating to key infectious disease topics in the UK from a population perspective. PHE had specific interest in three dimensions of inequalities: inclusion health groups (socially excluded and vulnerable populations), protected characteristics and socioeconomic inequalities. The infectious disease topics of interest were tuberculosis (TB), HIV, sexually transmitted infections (STIs), hepatitis C virus (HCV), vaccination and antimicrobial resistance (AMR). Therefore, the aim of this systematic overview of reviews was to describe the existing literature, relevant to the UK, relating to inequalities in the prevalence/incidence of key infectious disease topics as specified by PHE.

METHODS
We conducted a systematic overview of reviews, preregistered in PROSPERO, an international prospective register of systematic reviews (2020 CRD42020220203 https://www.crd.york.ac.uk/prospero/display_record.php?ID=CRD42020220203).

Patient and public involvement
This study had no direct patient or public engagement.

Search strategy and study selection
We developed a search strategy using synonyms and Medical Subject Headings (MeSH) terms for inequalities, inclusion health groups, protected characteristics and socioeconomic factors which were combined with synonyms and MeSH terms for infectious diseases and synonyms for reviews (online supplemental file 1). We searched electronic databases from inception to November 2021; MEDLINE, Embase, Web of Science and to identify relevant grey literature, we searched OpenGrey database (http://www.opengrey.eu/) and contacted experts in our network.

We exported citations into EndNote, removed duplicates and then exported them into a web-tool, Rayyan (https://www.rayyan.ai/) to facilitate citation screening. Articles were screened against predefined eligibility criteria (table 1). Titles and abstracts were screened by a single reviewer and 10% were double screened by a second reviewer. Full texts of the potentially relevant articles were obtained and screened independently by two reviewers. Discrepancies were resolved by discussion.

Data extraction
We designed and piloted a data extraction form in Microsoft Excel to extract information including: First author’s last name, publication year, corresponding author’s country, review methodology, inclusion and exclusion criteria, infectious disease(s), population(s) included, dimension(s) of inequality, outcomes, conclusions and strengths and limitations. Data extraction was performed by one reviewer and checked by another.

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<th>Inclusion criteria</th>
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<td>Population: Review including studies from the UK population or other high-income countries. Exposure:</td>
<td>We excluded reviews of qualitative studies and articles that are not systematic reviews as defined above. We excluded review protocols, but we searched the titles to check if the findings had been published. We excluded reviews on COVID-19, as advised by PHE, to avoid overlap with other reviews. We also excluded articles focused on travel-related infections. Reviews which excluded the UK in their eligibility criteria or had not included populations relevant to the UK population (e.g. papers where all results were from low-income countries) were excluded.</td>
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<td>Socioeconomic status: This includes education, income, occupation, social class and deprivation measured at individual or aggregated level.</td>
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<td>Protected characteristics: Age, disability, gender reassignment, marriage and civil partnership, pregnancy and maternity, race, religion or belief, sex, and sexual orientation.</td>
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<td>Inclusion health groups: Vulnerable migrants, people experiencing homelessness and rough sleeping, people who engage in sex work and Gypsy Roma and Traveller communities.</td>
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<td>Outcome: Inequalities relating to incidence, prevalence and consequences of infectious diseases. Despite specific interest in TB, HIV, STIs, HCV, immunisation, and AMR, we included reviews relating to any infectious diseases, except reviews focused on COVID-19. Types of reviews: Any literature review which reports all the following (a) explicit objectives, (b) clear search strategies and (c) eligibility criteria. Publication date: Published from the year 2000 onwards. Language: No language restrictions.</td>
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AMR, antimicrobial resistance; HCV, hepatitis C virus; PHE, Public Health England; STIs, sexually transmitted infections; TB, tuberculosis.
Quality assessment
Two reviewers independently assessed review quality using the Assessment of Multiple Systematic Reviews V.2 (AMSTAR2) checklist. Disagreements were resolved by discussion. Due to the multidimensional nature of this overview of reviews, inclusion of various types of reviews with diverse aims and outcomes, we did not perform an overall rating of confidence for each review. To provide a sense of overall quality of evidence, we calculated the proportion of reviews which fulfilled each AMSTAR2 item.

Data synthesis
We tabulated the dimension of inequalities against the infectious disease topic to create an evidence matrix which was used to highlight areas where reviews already exist and where there may be gaps. Data were synthesised narratively based on the dimensions of inequalities.

RESULTS
Figure 1 shows the study selection. We retrieved 14713 citations from the electronic database searches and 11135 titles and abstracts were screened after the removal of duplicates. One of the experts we contacted sent an article which highlighted UK-based evidence for several inclusion health groups, but did not fulfil other criteria for inclusion. After examining 437 full-text articles against the eligibility criteria, 108 were included in our synthesis.

Characteristics of included reviews are summarised in online supplemental file 2. Included reviews were published between 2005 and 2020 with 95% published after 2010. Fifty-eight (54%) included meta-analysis while the remaining studies used narrative/descriptive synthesis approaches. The reviews covered the three dimensions of inequalities across various infectious disease topics. A summarised version of the evidence matrix, showing how many reviews were identified for each cell is presented in figure 2. The full evidence matrix is presented in online supplemental file 3.

Methodological quality of included reviews
Assessment of the methodological quality of each included review is presented in online supplemental file 4 and the proportion of included reviews that met each AMSTAR2 criteria are presented in figure 3. Many reviews fulfilled
criteria such as including components of PICO (population intervention, comparator group and outcome) in their research questions and inclusion criteria (87%), performing duplicate study selection (55%), discussing heterogeneity (70%) and disclosure of conflicts of interest (83%). Only a few of the reviews (19%) clearly indicated that the review methods were established a priori and 37% performed risk of bias assessment using satisfactory

Figure 2  Matrix showing the number of reviews identified for each dimension of inequality and infectious disease topic. Colour ranges from red which indicates where no review was identified, up to green for a maximum number of reviews (19). AMR, antimicrobial resistance; STIs, sexually transmitted infections.

Figure 3  Assessment of Multiple Systematic Reviews V.2 results for included reviews. PICO, population, intervention, comparator group and outcome.
Evidence relating to inclusion health groups

Of the 108 included reviews, 43 reported on inclusion health groups. The evidence was generally consistent across these groups showing that people who belong to inclusion health groups are often at higher risk of infectious diseases, AMR and under-vaccination. For example, many reviews reported on general migrants, and vulnerable migrants such as asylum seekers, refugees and trafficked sex workers. The reviews often reported that migrants have a higher risk of infectious diseases than the host population, though the magnitude of the association may vary for different geographical regions and different infectious diseases. For example, in the UK, 35% of people with chronic HCV are migrants despite being just 9% of the general population. Other reviews showed that HIV and STIs were more prevalent among migrants. The prevalence of HIV-TB co-infection was higher among immigrants compared with nationals in various countries including England and Wales although the immigrant group reported slightly better survival/lower mortality which authors commented may be due to the possible healthy migrant effect. In another review, migrants were reported to be at higher risk of TB death. Evidence from the UK showed an increasing number of migrants contracted HIV after they arrived in the UK between 2002 and 2011 suggesting that the higher prevalence of infectious diseases among migrants is not limited to pre-migration infection. A meta-analysis showed that refugees were more likely to have chronic hepatitis B virus (HBV) compared with general migrants (OR 1.42, 95% CI 1.01 to 1.99). Some reviews reported no clear evidence that immigrant sex workers had higher risk of HIV and STIs compared with non-migrant sex workers. However, one reported that trafficked sex workers were at a higher risk of HIV and STIs compared with female sex workers in general. The prevalence of Helicobacter pylori among immigrants varied according to continent of origin and the prevalence is higher among migrants compared with their children.

Several reviews reported lower vaccination rates or delayed/incomplete vaccination among migrants and refugees in Europe. However, the association may vary depending on the type of migrant group. For example, authors have reported that the uptake of vaccination among refugees was lower compared with asylum seekers. In a meta-analysis, migrants had increased odds of multidrug-resistant (MDR)-TB incidence compared with non-migrants (OR 3.91, 95% CI 2.98 to 5.14). In two other reviews, AMR carriage and infection were reported to be more prevalent among migrants in Europe.

Three reviews examined Gypsy Roma and Traveller communities. One showed that Roma and Irish Travellers in the UK were often under-vaccinated. Another reported that Roma in Barcelona had a TB incidence 5.3 times greater than Spain’s national TB incidence. Higher prevalence of HIV has also been reported among Iranian, Roma and Peruvian Indigenous populations compared with the general population. We did not identify any reviews that examined the association between being from Gypsy Roma or Traveller communities and AMR.

We identified eight reviews which examined the association of homelessness with infectious diseases. They all reported a higher risk of various infectious diseases or AMR among people experiencing homelessness compared with those who were not homeless. We did not identify any reviews that examined the association between vaccination and homelessness. Eight reviews explored infectious disease risks among those engaging in sex work compared with the general population. The evidence suggests higher risks of various infectious diseases, such as HBV, hepatitis D virus (HDV), HIV and human papillomavirus (HPV), among sex workers. A review which examined factors associated with HBV vaccination among men who have sex with men (MSM) found mixed evidence relating to sexual risk-taking including involvement in sex trade.

Evidence relating to protected characteristics

Seventy-four reviews reported on protected characteristics, however, our synthesis only found clear evidence for inequalities by ethnicity and sexual orientation. Inequalities in infectious diseases relating to race and ethnicity were explored in 19 reviews. The available evidence suggests a higher rate of various infectious diseases such as TB, HIV and STIs and under-vaccination in people who belong to an ethnic minority group. For example, a meta-analysis found that recent transmission of TB was associated with being of ethnic minority (OR 3.03, 95% CI 2.21 to 4.16). A meta-analysis indicated that on average young black women were less likely to initiate HPV vaccination than young white women (combined OR 0.89, 95% CI 0.82 to 0.97). In a meta-analysis of studies from Europe, children from parents of ethnic minorities (compared with the majority) were less likely to be vaccinated for measles, mumps and rubella (MMR) (OR 0.89, 95% CI 0.86 to 0.93 in a fixed effect model). However, the effect disappeared in the random effects model (OR 1.03, 95% CI 0.79 to 1.34), probably due to heterogeneity between studies. Seasonal influenza vaccine uptake among older people was associated with being white (combined OR 1.30, 95% CI 1.14 to 1.49). Only one review on race and AMR was identified and it reported that people from some black ethnic groups in the USA and
Europe, and Aboriginal ethnic groups living in Canada and Australia are less likely to have AMR-Neisseria gonorrhoea (AMR-NG) than the white majority population.  

Ten included reviews examined the association of sexual orientation with infectious disease topics, mostly focused on MSM.  

In a review, MSM were found to be at risk of acquiring HIV post-migration. However, in a network meta-analysis the highest risk of advanced HIV disease among people living with HIV was found in those with heterosexual contact compared with MSM as well as injection drug use. Some reviews examined disparities of HIV in MSM but did not compare risk between MSM and other populations. Other infectious diseases such as HBV, HCV and HDV are also higher in MSM than in the general population. AMR-NG was reported to be more common among MSM than heterosexual men in England and Wales (OR 5.47, 95% CI 3.99 to 7.48). We did not identify any reviews which assessed the association of sexual orientation with vaccination.

From the reviews identified, inequalities in infectious disease topics based on other protected characteristics, such as age and sex are mixed and for other protected characteristics the synthesised evidence is scant and inconclusive. The association of infectious diseases with age has been reported in various reviews. However, the association varied. Infectious diseases such as HIV, STIs and TB have been reported to be associated with younger age in some reviews, while HCV and hepatitis E virus were associated with older age. Seasonal influenza vaccine uptake was higher in older age groups. A review reported that many studies found an association between HBV vaccination and younger age. Suboptimal vaccination compliance was associated with mother’s younger age. On the other hand, another review reported that HPV vaccine intention and initiation were positively associated with younger parent’s age. Four included reviews examined the association of AMR with age. In one review, MDR-TB was associated with being younger than 65 years (pooled OR 2.53, 95% CI 1.94 to 3.32) while another review reported that AMR-NG was more common in those 25 years or older than in younger adults. Overall, age group classifications often varied between reviews which made it difficult to identify a clear pattern.

Twenty-eight reviews explored inequalities based on sex. The findings varied depending on specific infectious disease. For example, TB transmission was reported to be associated with being male while the prevalence of chlamydia was slightly higher in women than in men. Other reviews reported that seasonal influenza vaccine uptake is often higher in elderly men compared with elderly women, but the differences are not statistically significant in multivariate regression analysis. A meta-analysis of studies from Europe showed that male patients were more likely to have MDR-TB (OR 1.38, 95% CI 1.16 to 1.65).

Six reviews described the influence of being married or in civil partnerships on vaccination. Generally, those who were married had higher vaccination uptake although some studies found no association or higher uptake among those who were never married. No included review examined inequalities in infectious disease prevalence or AMR based on marital status. Only four included reviews reported the prevalence of infectious diseases (HBV, HCV, latent or acute toxoplasma infection) in pregnant women compared with the general population and the findings were mixed. We found no reviews that examined the association of pregnancy with vaccination uptake or risk of AMR.

Six reviews reported on inequalities relating to religion and meningococcal disease, as well as vaccination. A recent meta-analysis of two studies showed that religious events attendance was significantly associated with a decreased risk of invasive meningococcal disease (OR 0.47 (95% CI 0.28 to 0.79, p=0.0004)). Meta-analyses showed no strong evidence between various vaccination and religion including frequency of attendance at a place of worship. Jewish Orthodox people in the UK and Belgium and Orthodox Protestants in The Netherlands were described as being under-vaccinated. We did not identify any studies on the association of religion with the risk of any of the key infectious diseases or AMR.

Two reviews examined the association between gender reassignment and the risk of infectious diseases (HIV and HBV). The prevalence of HIV was significantly higher among transgender female sex workers compared with biologically female sex workers (relative risk=4.02, 95% CI 1.60 to 10.11). However, in another review, transgender persons had lower prevalence of HBV compared with other groups such as sex workers, injection drug users, MSM and pregnant women. We did not identify any reviews examining the association of gender reassignment with other infectious diseases of interest, vaccination or AMR.

We did not identify any reviews that reported the association of disability with our key infectious diseases topics. However, we identified one review which reported that disability was associated with a higher incidence of listeriosis.

Evidence relating to socioeconomic inequalities

Fifty reviews explored socioeconomic status. The evidence consistently shows that those with lower level of income, lower educational attainment, unemployment, higher area level deprivation, lower socioeconomic status or poor living situations are at higher risk of infectious diseases, AMR, and lower vaccine uptake. For example, many reviews highlighted that low income, poverty and unemployment were associated with various infectious diseases including, HIV, STIs, TB, HBV, and HCV among others.

Level of education, income or occupation are often associated with vaccination uptake. Reviews have also reported an association of MDR-TB
and AMR with lower level of income or education. Many reviews examined the association of infectious disease topics of interest with real/socioeconomic status, deprivation, migration status, and areal level deprivation. Although the evidence is scant, the findings suggest that those living in deprived areas or poor living conditions may be at higher risk of AMR.

**DISCUSSION**

This overview of reviews provides a broad synopsis of three dimensions of inequalities (inclusion health groups, protected characteristics and socioeconomic inequalities) across several infectious disease topics. We synthesised the existing evidence based on the dimension of inequalities. Of the three dimensions of inequalities assessed, the evidence relating to people in the inclusion health groups is the most consistent although the volume of evidence identified for each group varied. Most of the reviews identified under this dimension were on migration status, with a higher prevalence of infectious diseases, AMR and lower vaccine uptake among migrants compared with non-migrants. Vulnerable migrants (such as refugees, asylum seekers and trafficked persons) are at higher risk when compared with general migrants and the size of inequalities varied depending on the country of origin. Although few reviews were identified for the remaining inclusion health groups, the evidence suggests that homelessness is associated with risk of infectious diseases and AMR; Gypsy Roma/Traveller communities are often under-vaccinated and are also at greater risk of infectious diseases; and people who engage in sex work are at greater risk of some infectious diseases.

There is a plethora of evidence from reviews showing higher prevalence of infectious diseases and under-vaccination among minority ethnic groups. We also identified several reviews suggesting higher prevalence of infectious diseases and AMR among MSM. These suggest that ethnicity and sexual orientation are important protected characteristics and targeting or tailoring interventions for such groups may be beneficial to reduce inequalities in infectious diseases. It is important to note that there is inequality in access to vaccinations as shown in reviews included in this overview of reviews and beyond. Since many infectious diseases are vaccine preventable, identified inequalities in infectious diseases that we have noted in this overview of reviews, may also be related to inequalities in access to vaccination.

Many reviews examined the association with age and sex, however, the identified associations varied depending on the specific infectious disease or type of vaccination. In addition, for most of the reviews, the comparator age groups were often unclear. Therefore, we are not able to identify specific age groups with higher risk across various infectious disease topics. Other factors besides equity issues may contribute significantly to associations with age. For example, people in the most sexually active age groups are more likely to contract STIs whereas people of older ages, where immunity is weaker, are more likely to get infectious diseases associated with low immunity. Also, vaccinations are often offered at specific ages so it is expected that uptake would be higher among those groups that are targeted. However, it is important to highlight that we found evidence suggesting that childhood vaccination compliance is lower for those with younger mothers/parents. Based on this review, age and sex may be important for some infectious diseases but the group at higher risk may vary across diseases.

Reviews exploring marital status focused on vaccination, particularly seasonal influenza vaccine in older adults, tetanus vaccination among pregnant women and MMR vaccination in children. Reviews generally reported higher vaccination uptake among adults who are married and children whose parents are married. It is not possible to draw a conclusion regarding the association of religion, disability, transgenderism, and pregnancy with infectious diseases based on the findings of this review as the synthesised evidence is scant and often inconsistent. More evidence is therefore needed to be able to establish the presence and direction of any associations of these factors with infectious diseases.

Several reviews provide compelling evidence of higher risk of infectious diseases, AMR, and lower vaccination uptake among those with lower level of income, lower educational attainment, unemployment, higher area level deprivation, lower socioeconomic status and poor living situations. Although most of the evidence in this dimension is on vaccination, those of lower socioeconomic groups are often at higher risk from infectious diseases and should be targeted for intervention.

**STRENGTHS AND LIMITATIONS**

The protocol used to guide the conduct of this review was designed a priori. We conducted a comprehensive literature search of four electronic databases with no language limits and searched for grey literature. Data extraction was checked by a second reviewer to improve accuracy and quality assessments were performed by two reviewers independently. Due to the time frame required for the work, we could not complete all the initial titles and abstract screening in duplicate, however, full texts of potentially relevant reviews were independently screened.
by two reviewers. Despite our best efforts, we acknowledge that some relevant reviews may have been missed in the study selection process. The lack of synthesised evidence observed in some areas does not necessarily mean a lack of evidence. This is because there may be primary studies in those areas which have not been synthesised in reviews and meta-analyses. Also, this project focused on specific infectious disease topics and we included specific search terms for those topics in the search strategy and included broad terms for infectious diseases. This allowed us to include other infectious diseases that were identified in our search to capture evidence of health inequalities from various infectious diseases (excluding COVID-19). Therefore, this is not intended to provide a comprehensive overview of reviews in those topics which are not the focus of this project. Furthermore, some underserved populations (such as people who inject drugs and prisoners) are not covered in this overview as these are beyond the scope of the work. Notwithstanding, we believe this provides a useful summary of available evidence relating to inequalities in infectious diseases relevant to high-income countries and highlights areas where evidence may be lacking or minimal.

The quality of the included reviews varied significantly as we included various types of reviews, and some were not necessarily systematic reviews for which AMSTAR2 is designed. Heterogeneity between studies was a limitation reported in many of the included reviews where meta-analyses were performed, therefore, pooled estimates should be interpreted with caution.

CONCLUSION

Overall, we provide evidence from many papers with accordant findings, of groups consistently at higher risk of infectious diseases, AMR and under-vaccination. Developing targeted interventions for high-risk groups rather than focusing on individual infections could contribute significantly to reducing infectious disease burden. This review also highlights important evidence gaps that should be considered when commissioning future evidence syntheses or primary studies.

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