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Trends in occupational respiratory conditions with short latency in the UK

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Background: Occupational short-latency respiratory disease (SLRD; predominantly asthma, rhinitis, hypersensitivity pneumonitis, and occupational infections) prevalence is difficult to determine but certain occupations may be associated with increased susceptibility. **Aims:** This study aimed to examine which occupations and industries are currently at high risk for SLRD and determine their respective suspected causal agents.

Methods: SLRD cases reported to the SWORD scheme between 1999 and 2019 were analysed to determine directly standardized rate ratios (SRR) by occupation against the average rate for all other occupations combined.

Results: 'Bakers and flour confectioners' and 'vehicle spray painters' showed significantly raised SRR for SLRD in general, mostly due to occupational rhinitis (234.4; 95% CI 200.5–274.0) and asthma (63.5; 95% CI 51.5–78.3), respectively. Laboratory technicians also showed significantly raised SRR for occupational rhinitis (18.7; 95% CI 15.1–23.1), primarily caused by laboratory animals and insects. Metal machining setters and setter-operators showed increased SRR for occupational hypersensitivity pneumonitis (42.0; 95% CI 29.3–60.3), largely due to cutting/soluble oils. The occupation mostly affected by infectious disease was welding trades (12.9; 95% CI 5.7–29.3), mainly attributable to microbial pathogenicity.

Conclusions: This study identified the occupational groups at increased risk of developing an SLRD based on data recorded over a recent twodecade period in the UK. Occupational asthma and rhinitis were identified as the prevailing conditions and hypersensitivity pneumonitis as a potentially rising respiratory problem in the metalworking industry.

INTRODUCTION

Disorders of the respiratory system with short latency manifest rapidly after risk factor exposure, as opposed to long-latency diseases (e.g. pneumoconioses or cancers) which can have latent periods of up to 50 years [1]. Short-latency respiratory diseases (SLRD) are caused by the inhalation of airborne particles, gases, fumes or vapours, and can affect a range of structures within the respiratory tract from the nose to alveoli. Clinical symptoms may result from allergic, inflammatory, infective or irritant pathophysiological processes.

Exposure circumstances can be varied and include occupational and environmental sources which need to be considered when diagnosing respiratory disease. Workplace hazards may cause or worsen respiratory health, regardless of the duration of the exposure or of the latent period from the cessation of exposure to disease onset. Whilst having less effects on mortality, SLRD are evidently associated with significant morbidity [2, 3], job loss and wider socioeconomic impacts [4].

A short latency period between workplace exposure and onset of symptoms might facilitate the establishment of a causal relationship between a suspected trigger and a disease. However, there is frequently causal misattribution due to complex exposure circumstances and the precise timing of symptoms in relation to exposure. Numerous substances have been recognized as respiratory sensitizers or irritants causing SLRD [5–7]. Notwithstanding, novel substances with the potential to sensitize the respiratory tract are constantly being developed and introduced in workplace environments [8]. It is thus essential to keep track of the causal agents associated with these conditions to prevent exposure and minimize risk. Secondary prevention of SLRD is frequently possible by removing individuals from exposure to these agents.

The UK-based SWORD scheme (Surveillance of Workrelated and Occupational Respiratory Disease) is a national reporting system for physicians that permits the collation of cases of occupational lung disease [9], along with contextual information that includes co-existing diagnoses, likely causative exposure, occupation and work sector.

Given that comparatively little is reported about shorter latency conditions from reporting schemes [10-13], with perhaps

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Key learning points

What is already known about this subject:

 Short-latency respiratory disease is very commonly diagnosed worldwide, yet not much is known about the occupations at high risk of developing shortlatency respiratory disease in the UK.

What this study adds:

 This study found significantly raised standardized rate ratios for occupational rhinitis, asthma, and hypersensitivity pneumonitis, respectively associated with bakers and flour confectioners, vehicle spray painters, and metal machining operatives.

What impact this may have on practice or policy:

• Given that the majority of short-latency respiratory diseases have the potential to improve and even resolve when the occupational hazard is mitigated or removed, the identification of occupational groups and agents associated with short-latency respiratory disease is a promising approach to guide future control and preventive measures.

the exception of occupational asthma, and the comparative lack of data related to both occupational rhinitis [14,15] and infections [16], we chose to report here information related to all such cases reported to SWORD over the period 1999–2019.

METHODS

Cases of occupational SLRD reported to the SWORD scheme [17,18] between 1999 and 2019, were used for the purposes of this analysis. Cases were reported anonymously by respiratory physicians in the UK, either as a core or sample reporter. Core reporters return all cases of occupational respiratory disease diagnosed monthly throughout the year, while sample reporters return only cases diagnosed during a single randomly selected month per year.

Information included in the report to SWORD includes age, gender, occupation, industry, first half of postcode, and up to three suspected causal agents attributed to the case. Data are coded according to their respective system classifications: occupation by the Standard Occupational Classification 2010 [19], industry of work by the Standard Industrial Classification [20] and an in-house causal agent classification system is used to collate exposure type.

A total estimated number of cases per year was calculated by adding the actual reported cases from the core reporters to the number of cases reported by the sample reporters factored by 12. The methodologies adopted have previously been described in more detail [17].

Cases extracted for this work included (i) asthma, (ii) rhinitis, (iii) hypersensitivity pneumonitis and (iv) infectious diseases. All other cases were also assessed for completeness, and a priori categorized as (v) 'other'. All short-latency disease cases for which there was a co-diagnosis with a long-latency disease were discarded, as well as those for which the causal agent was specified as asbestos. Inhalation accidents [21], bronchitis/ emphysema (COPD), non-malignant pleural disease (predominantly plaques, predominantly diffuse, and asbestos-related pleural effusion), mesothelioma, lung cancer, pneumoconiosis (asbestosis, silicosis, and coal workers' pneumoconiosis), and unspecified cases, which were unclear about the latency were all excluded from this analysis. Data from 2019 onward were excluded from these calculations due to the atypical reporting pattern during the coronavirus disease 2019 (COVID-19) pandemic.

Directly standardized rate ratios (SRR) per occupation were calculated for all relevant diagnostic categories. For each occupation, the incidence rate of a particular diagnosis (or individual diagnostic category) was obtained by averaging the number of total cases of the individual occupations weighted by their relative sizes according to employment-related statistics in the UK [22]. Considering that the population distribution is not homogeneous regarding occupational groups, and to allow for a meaningful comparison between groups of different professional occupations, rates for each occupation were directly standardized by the average rate of all other occupations combined. Approximate 95% confidence intervals (CIs) were estimated using a Taylor series variance estimator to account for the fluctuation inherent to the weighting of sample reports [23], and a finite population correction factor of 0.3 was used to adjust for the proportion of eligible chest physicians that report to SWORD [24].

The Health and Occupation Research (THOR) network (including EPIDERM and SWORD) has National Health Service ethics approval given by the Northwest (Haydock) Research Ethics Committee (22/NW/0082).

RESULTS

A total of 3476 cases (core plus unweighted sample cases) were reported over the study time period. Table 1 shows the number of actual and estimated cases and diagnoses by each disease category. Overall, cases were predominantly reported in men (71%) with a mean age of 43 ± 12.2 (standard deviation [SD]) years, and predominantly reported in the manufacturing industry (54%), particularly in the manufacture of motor vehicles, trailers and semi-trailers (22%) and the manufacture of food products (19%). The next most frequent industrial sectors were professional, scientific and technical activities (9%), where the majority of cases (86%) were within scientific research and development.

The most reported occupations were skilled trades occupations (34%) and process, plant and machine operatives (25%). Of the former, the commonest occupational groups were food preparation trades (27%) and metal machining, fitting and instrument making trades (21%). Of the latter category, 47% were process operatives and 28% were assemblers and routine operatives.

There were major differences in the incidence of SLRD between occupations, with the highest incidences seen in bakers and flour confectioners (SRR 58.9; 95% CI 51.8–66.9) and vehicle spray painters (SRR 48.9; 95% CI 39.7–60.2). Although still significantly raised, all other occupations had considerably lower incidence rates than these two occupations, in general. For example, metal working production and maintenance fitters (SRR 2.2; 95% CI 1.8–2.8), nurses (SRR 1.9; 95% CI 1.5–2.3), and cleaners and domestics (SRR 1.5; 95% CI 1.2–1.9) showed the lowest SLRD incidence rates (Figure 1).

Asthma was the most frequently reported condition (2784 actual cases), comprising about 75% of all actual cases, followed by rhinitis (16%). Rhinitis was also the most commonly reported co-diagnosis, usually alongside asthma. Asthma cases were mostly male (72%), with a mean age of 44 ± 11.7 (SD) years, and with a predominance in the manufacture of food products and manufacture of motor vehicles, trailers and semi-trailers industries. Occupations with the highest incidence rates were bakers and flour confectioners (SRR 59.9; 95% CI 51.6–69.5), vehicle spray painters (SRR 63.5; 95% CI 51.5–78.3), and chemical and related process operatives (SRR 21.0; 95% CI 16.9–26.1; Table 2). Isocyanates (24%) and flour (19%) were the most important agents causing occupational asthma, as shown amongst other data relating to all other diagnoses (Figure 2).

Rhinitis cases (588 actual cases) were also mostly male (68%), with a mean age of 36 ± 10.6 (SD) years. Bakers and flour confectioners (SRR 234.4; 95% CI 200.5–274.0) showed the highest incidence of rhinitis, indicating that there is a significantly higher risk of developing rhinitis than of developing asthma. There was also a suggestion that chemical and related process operatives (SRR 29.5; 95% CI 24.3–35.7) and laboratory technicians (SRR 18.7; 95% CI 15.1–23.1) are occupations with an increased risk of occupational rhinitis. The incidence of rhinitis was significantly lower in metal working production and maintenance fitters (SRR 0.7; 95% CI 0.4–1.2) and cleaners and domestics (SRR 0.1; 95% CI 0.0–0.2) than in all other occupations combined (Table 2). Laboratory animals and insects (34%) and flour (30%) were the most important causes of rhinitis (Figure 2).

Table 1. Number and percentage of short-latency respiratory disease (SLRD) cases reported by chest physicians to SWORD (1999-2019)

Disease category	Actual cases ^a , n (%)	Estimated cases ^b , <i>n</i> (%)	Age ^c (mean ± SD) (years)	Sex ^c	
				Female, <i>n</i> (%)	Male, <i>n</i> (%)
Asthma	2784 (75)	5941 (73)	44 ± 11.7	770 (28)	2011 (72)
Rhinitis	588 (16)	731 (9)	36 ± 10.6	189 (32)	399 (68)
Hypersensitivity pneumonitis	209 (6)	770 (10)	52 ± 12.1	46 (22)	163 (78)
Infectious disease	72 (2)	501 (6)	43 ± 12.8	30 (42)	41 (57)
Other SLRD	80 (2)	157 (2)	48 ± 11.5	26 (33)	54 (68)
Total diagnoses ^d	3733	8100	43 ± 12.2	1061 (28)	2668 (71)
Total cases	3476	7818	44 ± 12.1	987 (28)	2485 (71)

Mean age and sex are presented for all actual cases (core plus unweighted sample cases).

^aActual cases refer to cases reported by core reporters plus unweighted cases reported by sample reporters.

^bEstimated cases refer to cases reported by core reporters plus 12 × cases reported by sample reporters.

Refers to cases where patient age/sex was reported.

^dDiagnoses refer to all disease categories reported per case.

Occupation	SRR (95% CI)	4			
Bakers, flour confectioners	58.9 (51.8 - 66.9)	·			
Vehicle spray painters	48.9 (39.7 - 60.2)	·			
Chemical and related process operatives	18.1 (14.9 - 22.0)				
Assemblers electrical products	12.7 (10.0 - 16.0)	_			
Welding trades	11.7 (9.8 - 14.0)				
Metal machining setters and setter-operators	11.2 (9.3 - 13.4)	1			
Food, drink and tobacco process operatives	9.1 (7.7 - 10.8)	- - -			
Metal working machine operatives	8.9 (7.0 - 11.2)	·			
Assemblers vehicles metal goods	6.9 (5.9 - 8.2)				
Laboratory technicians	6.1 (4.8 - 7.6)	·			
Packers, bottlers, fillers	2.9 (2.2 - 3.7)	·			
Metal working production and maintenance fitters	2.2 (1.8 - 2.8)	·			
Nurses	1.9 (1.5 - 2.3)	·			
Cleaners, domestics	1.5 (1.2 - 1.9)	·			
		1 3 10 30			
		SRR with 95% Confidence Interval			

Figure 1. Standardized rate ratios (SRR) and 95% CI of medically reported occupational SLRD incidence reported to SWORD compared with all other employment sectors combined (1999–2019). Please note the *x*-axis log scale.

Occupation	SRR (95% CI)					
	Asthma	Rhinitis	Hypersensitivity pneumonitis	Infectious disease		
Bakers, flour confectioners	59.9 (51.6–69.5)	234.4 (200.5–274.0)	13.1 (4.4–38.6)	_		
Vehicle spray painters	63.5 (51.5–78.3)	-	1.3 (0.4–3.8)	-		
Chemical and related process operatives	21.0 (16.9–26.1)	29.5 (24.3–35.7)	0.6 (0.2–1.8)	-		
Welding trades	13.0 (10.8–15.7)	1.0 (0.5–2.2)	6.8 (2.4–18.8)	12.9 (5.7–29.3)		
Assemblers electrical products	16.4 (12.9–20.8)	1.9 (0.8–4.4)	_	-		
Metal machining setters and setter-operators	9.5 (7.6–11.7)	3.7 (2.4–5.6)	42.0 (29.3–60.3)	-		
Food, drink and tobacco pro- cess operatives	10.0 (8.3–12.0)	14.1 (9.3–21.4)	1.1 (0.6–1.9)	8.6 (3.9–18.7)		
Metal working machine op- eratives	9.0 (6.9–11.6)	2.7 (1.7–4.5)	17.8 (10.1–31.3)	-		
Assemblers vehicles metal goods	8.0 (6.7–9.6)	5.6 (3.8–8.2)	9.6 (6.9–13.3)	-		
Laboratory technicians	5.6 (4.2–7.4)	18.7 (15.1–23.1)	_	8.7 (2.9–25.8)		
Packers, bottlers, fillers	2.1 (1.6–2.8)	2.6 (1.8-3.7)	4.2 (1.9–9.2)	10.2 (4.7–22.2)		
Metal working production and maintenance fitters	2.3 (1.8–2.8)	0.7 (0.4–1.2)	3.9 (1.8–8.3)	0.2 (0.1–0.5)		
Nurses	1.3 (1.1–1.6)	1.6 (0.8–3.1)	0.9 (0.3–2.5)	10.0 (6.4–15.7)		
Cleaners, domestics	1.8 (1.4–2.2)	0.1 (0.0–0.2)	0.4 (0.3–0.7)	1.3 (0.4–3.7)		

Table 2. Standardized rate ratios (SRR) and 95% CI of medically reported occupational SLRD incidence reported to SWORD compared with all other employment sectors combined per diagnostic category (1999–2019).

Hypersensitivity pneumonitis cases (209 actual cases) were again predominantly male (78%), with a mean age of 52 ± 12.1 years, as shown in Table 1. Metalworking industry activities such as metal machining setters and setter-operators (SRR 42.0; 95% CI 29.3-60.3) and metal working machine operatives (SRR 17.8; 95% CI 10.1–31.3) were the most affected occupations. These were followed by grain-based food industries, represented by bakers and flour confectioners (SRR 13.1; 95% CI 4.4–38.6). Occupations such as chemical and related process operatives (SRR 0.6; 95% CI 0.2–1.8), nurses (SRR 0.9; 95% CI 0.3–2.5), and cleaners and domestics (SRR 0.4; 95% CI 0.3-0.7) showed a significantly lower incidence of hypersensitivity pneumonitis than all other occupations, again as shown in Table 2. Metal working fluids, such as cutting and soluble oils, were the main attributed causal agent in hypersensitivity pneumonitis (34% of all cases), followed by exposure to fungi, moulds and yeast (18%) (Figure 2).

Infectious disease was the diagnostic category with the lowest number of reported cases (72 actual cases) and the highest female proportion (42%). Welding trades (SRR 12.9; 95% CI 5.7–29.3), packers, bottlers, fillers (SRR 10.2; 95% CI 4.7–22.2) and nurses (SRR 10.0; 95% CI 6.4–15.7) ranked as the top three occupations at the highest risk of being reported as a case and only metal working production and maintenance fitters (SRR 0.2; 95% CI 0.1–0.5) showed a significantly lower incidence of infectious disease than all other occupations (Table 2). As expected, pathogens and micro-organisms were the predominant suspected cause of infectious diseases (80%) (Figure 2). In particular, *Mycobacterium tuberculosis* (53%), other species of bacteria (43%) and fungi (2%) were attributed as causes of individual cases. More information is included

in Figure 1 (available as Supplementary data at *Occupational Medicine* online).

Figure 3 (similar plots for individual diseases are found in Supplementary Material, available as Supplementary data at *Occupational Medicine* online) shows the distribution of the suspected causal agents reported for all SLRD and respective categories communicated to SWORD between 1999 and 2019. The most frequently reported agents across the reporting period were biological substances (42%), followed by chemically ill-defined substances (13%) and isocyanates (10%).

Finally, other less commonly reported cases included bronchiolitis (nine cases), metal fume fever (nine cases), humidifier fever (eight cases), hyperventilation (eight cases) and cough/ laryngitis (four cases each). More information can be found in Table 1 (available as Supplementary data at *Occupational Medicine* online).

DISCUSSION

This study reports the incidence of SLRD in the UK as communicated by chest physicians to the SWORD surveillance scheme over a recent two-decade period (before the COVID-19 pandemic). Overall case numbers indicated that respiratory diseases with short latency account for approximately 20% of all occupational respiratory diseases affecting the UK working population [13] and are comparable to those reported in other European countries [25–28]. In previous years in the UK, the annual incidence rate of SLRD has remained stable with moderate decreases in a few disease groups including asthma [13, 29], until the recent reversal of this trend observed after 2014 [30]. Reports since the first analyses of SLRD in the UK have kept

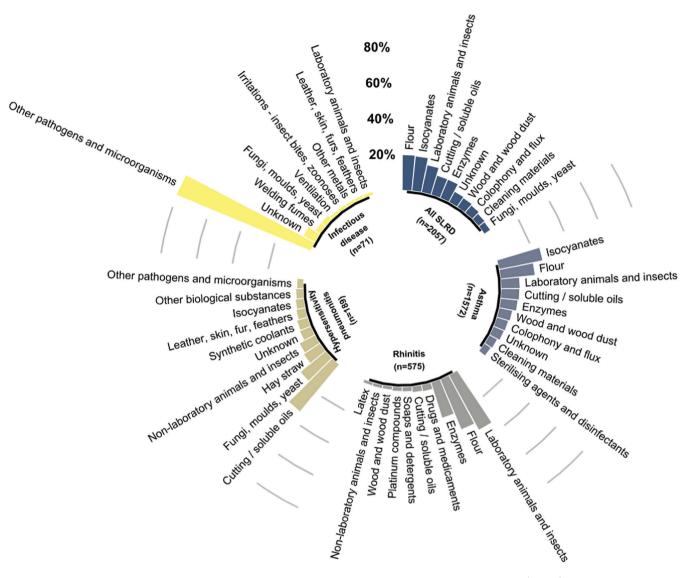


Figure 2. Suspected causal agents associated with the different occupational short-latency respiratory disease (SLRD) categories reported to SWORD (1999–2019). Vertical bars represent the percentage of each agent relative to the total frequency of the top ten agents reported (n), as indicated by the grey concentric lines. The suspected agents' subcategories are not necessarily grouped or ranked according to the hierarchy of the coding system.

unchanged regarding the main disease categories [17]; however, hypersensitivity pneumonitis cases reported in this study do not seem to be as rare as previously described, and the same is true for cases of rhinitis.

The findings of this study also indicated that the occupations at the highest risk of developing SLRD are bakers and flour confectioners and vehicle spray painters (Figure 1). Most SWORD reports for these groups include occupational asthma and rhinitis attributed to flour and diisocyanates (Figures 2 and 3, available as Supplementary data at *Occupational Medicine* online). In the case of asthma, although both substances are responsible for causing sensitization, they have been shown to differ in a number of ways in terms of properties and disease mechanisms [31]. For example, flour is a high-molecular-weight substance that usually causes IgE-dependent immunologic reactions [32], while diisocyanates are low-molecular-weight chemicals that are thought to cause asthma by other mechanisms since specific IgE antibodies to diisocyanates are not observed in the serum of the majority of patients [33]. Flour dust has long been recognized as a hazardous substance that can include and combine diverse components used to improve baking, for example, enzymes, amino acids and chemical additives such as bleaching agents and emulsifiers [34]. These substances are potential sensitizing agents and are therefore susceptible to cause allergic and non-allergic respiratory reactions. Moreover, new additives are constantly being developed [35] and as such, the reporting of SLRD becomes an important means of detecting new hazardous agents [36]. Our data are consistent with the observation that occupational rhinitis in bakers is more common (the highest SRR observed in this study) and usually precedes asthma [3] since approximately 8% of asthma cases reported to SWORD have a co-diagnosis of rhinitis [37,38].

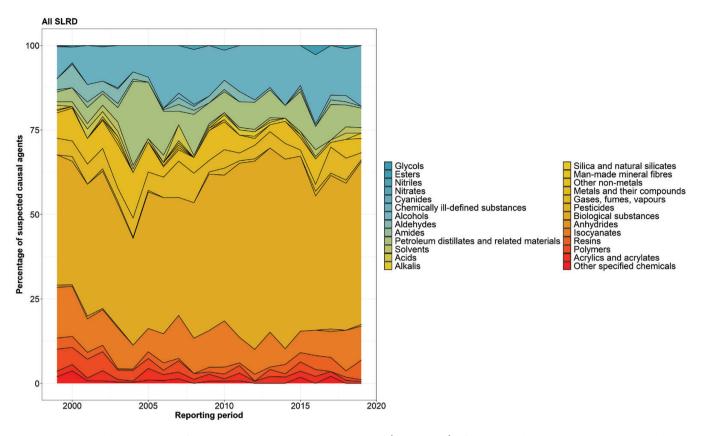


Figure 3. Suspected causal agents of occupational SLRD reported to SWORD (1999–2019). The suspected agents are organized according to the SWORD coding system categories and provide a higher-level classification in terms of their chemical, molecular and structural properties (e.g. flour and enzymes are categorized as 'Biological substances').

Exposure to metalworking fluids such as cutting and soluble oils has also been linked to occupational respiratory diseases, due to the inhalation of fluid mists during metal machining processes [39,40]. When considering hypersensitivity pneumonitis, high incidence rates were observed for metal machining setters and setter-operators and metal working machine operatives, even though this lung condition is typically associated with the inhalation of organic dust particles [41]. Although mineral oil lubricants are thought to be carcinogenic [42] and cause pulmonary fibrosis [43], their relationship with SLRD is perhaps less studied. In fact, reports of hypersensitivity pneumonitis due to metal working fluid aerosols reported before 2000 were relatively uncommon as documented by SWORD and OPRA reports [44] (Figure 4, available as Supplementary data at Occupational Medicine online). Although much focus has been on microbial contamination of metal working fluids as a potential antigenic source that might lead to hypersensitivity pneumonitis this has not been fully established as the cause [45]. Metal working fluids are complex mixtures of chemicals that serve a range of functions, such as emulsification and corrosion inhibition. Some organic compounds within these fluids might have the structural features of lipophilicity and multiple reactive groups that have been associated with hypersensitivity pneumonitis causation [46]. The chemical composition of metal working fluids is constantly evolving and varies between manufacturers, so there is at least a theoretical possibility that some of the outbreaks of hypersensitivity pneumonitis could be caused by specific chemical ingredients.

Exposure to microbial organisms in the workplace has been associated with diverse infectious respiratory diseases [16]. SWORD reports include infections predominantly caused by bacteria (80%). These are mostly associated with pathogenic mycobacteria and include multidrug-resistant and zoonotic bovine tuberculosis (Figure 1, available as Supplementary data at Occupational Medicine online), with most cases observed in healthcare occupations. However, a shift from these pathogens and micro-organisms to causal agents such as welding fumes, fungi, moulds and yeast, and ventilation has been observed in recent years (Figure 5, available as Supplementary data at Occupational Medicine online). Our results show that nurses have one of the lowest SLRD incidence rates; however, the incidence of infection for these healthcare professionals is significantly raised when compared to all other occupations (Table 2). Nonetheless, according to the results of a cohort study based on healthcare workers in the UK, which found that, although this occupational group has a higher incidence of tuberculosis than non-healthcare workers, this disease is generally not acquired through occupational exposure [47]. Welding trades occupations, on the other hand, are known to be at increased risk of developing pneumococcal pneumonia, in particular lobar pneumonia, which affects one or more lobes of the lung where inflammation and oedema acquire a consolidated pattern. Yet the mechanisms through which metal fume promotes pneumonia remain unclear. Many common welding processes involve intense heat generation, with subsequent interaction of UV radiation with metal surfaces and

surrounding gases, which can lead to the formation of reactive oxygen species. Several studies suggest that oxidative stress mechanisms may play a role in the development of pneumococcal pneumonia [48, 49], predisposing welders to this type of infection.

Taking this into consideration, in 2014, the Health and Safety Executive jointly with the manufacturers' organization Make UK (formerly known as EEF) and the Cast Metals Federation have provided additional guidance on previous advice from the Department of Health issued in 2012, whereby pneumonia vaccination was strongly recommended to employees exposed to welding or metal fumes [50]. In this study, we still find welding trades as the occupation with the highest incidence rate of occupational infectious diseases, and it would be interesting to follow up on the trend in the incidence of pneumonia in welding trades workers after the introduction of this preventive measure.

Clinical diagnosis of occupational SLRD can become challenging when dealing with diseases presenting similar clinical features. SWORD reports of diseases categorized as 'Other SLRD' include cases that may have common causes and overlapping symptoms with the otherwise classified subgroups. This is the case for organic dust toxic syndrome and humidifier fever, which include non-immunologic reactions involving non-specific symptoms.

The SRR used in this study allow for cross-occupational comparisons of incidence rates with all other occupations combined, revealing significant high-risk exposures for SLRD in activities such as bakers and flour confectioners and vehicle spray painters. Although incidence estimates may be improved by adjustments for cases missed due to non-participation of eligible chest physicians and respective response rates [51], we did not take these factors into consideration in this study. However, estimates of variance of total estimated cases due to sample reporting in the numerator were taken into account when calculating confidence intervals. Moreover, potential bias in the LFS-based estimates of the workforce by occupation in the UK is also mitigated by the use of rate ratios.

Unlike long-latency respiratory diseases, the majority of SLRD (with the exception of asthma and chronic hypersensitivity pneumonitis) are preventable. Additionally, they have the potential for significant improvement or resolution upon cessation of work exposure. This offers significant opportunities not just for enhancing workplace hazard prevention through improved risk assessment, but also for implementing more effective interventions in at-risk populations.

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COMPETING INTERESTS

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