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Work characteristics, socioeconomic position and health: a systematic review of mediation and moderation effects in prospective studies

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ABSTRACT

Social inequalities in health persist in modern societies. The contribution of adverse work and employment conditions towards their explanation is analysed by two approaches, mediation and moderation. Yet the relative significance of each approach remains unclear in respective research. We set out to study this question by conducting a systematic literature review. We included all original papers based on prospective observational studies of employed cohorts that were published between January 1980 and October 2012 meeting our search criteria, by using major databases and by observing established quality criteria. 26 reports were included after quality assessment. 17 studies examined the mediation hypothesis and nine studies tested the moderation hypothesis. Moderate support was found for the mediation hypothesis where OR or HR of health according to socioeconomic position (SEP) were reduced in a majority of analyses after introducing work characteristics in multivariate models. Evidence in favour of the moderation hypothesis was found in some studies, demonstrating stronger effects of adverse work on health among people with low SEP. Despite some support in favour of the two hypotheses future research should aim at reducing the heterogeneity in defining and measuring core variables and at applying advanced statistical analyses. Policy recommendations would benefit from a higher degree of consistency of respective research evidence.

INTRODUCTION

There is robust evidence of a social gradient of major chronic diseases and other health measures, based on prospective observational studies of employed populations in a majority of modern societies.^{1–3} The lower people's socioeconomic positions (SEP) are, the higher is their risk of morbidity and impaired health. Work and employment conditions play a prominent role in scientific attempts towards explaining this social gradient, given their primary impact on adult everyday life, its obligations, resources and rewards.^{4–6} Although SEP is often measured by occupational characteristics, such as job status, job classification, or employment grade, this information has limited explanatory power as it does not offer a more refined understanding of the ways in which occupational position affects health. Moreover, as these effects are bidirectional research has to assess the relevance of each one of the two pathways, causation and selection. Recent life course research emphasises a triggering role of adverse early life circumstances, including poor health, on occupational

What this paper adds

- To what extent adverse working conditions contribute towards explaining social inequalities in health has been analysed by studies testing the mediation and moderation hypotheses.
- So far, the relative significance of each hypothesis remains unclear, and the majority of available evidence was derived from cross-sectional studies.
- Based exclusively on prospective cohort studies, the results of this systematic review provide moderate support of the mediation hypothesis and modest support of the moderation hypothesis.
- Given a high degree of heterogeneity of core measurements analysed in these studies we propose specific recommendations for more standardised procedures in future research.

careers.⁷ Yet several findings from longitudinal studies indicate that the pathway from work to health ('causation') adds more to the explanation of the social gradient than the reverse pathway from health to work ('selection').^{8, 9}

Traditionally, physical stressors and occupational hazards were considered major causes of work-related health risks. Several of these stressors and hazards continue to follow a social gradient in developed western economies, but evidence indicates that they strongly cluster among low skilled occupations and people with atypical or precarious employment.^{6, 10, 11} To date, given profound changes in the nature of employment and work (expansion of service sector, growing impact of information technology, flexibility of employment arrangements), health-adverse work time arrangements and stressful psychosocial work environments are affecting large parts of the workforce.¹² With the advent of economic globalisation stressful psychosocial work environments became even more important in terms of health-adverse effects.¹³ The experience of work pressure in combination with threats to job stability is no longer confined to low skilled occupational groups, but increasingly affects better trained occupations.¹⁴ It is therefore important to know more exactly how work characteristics in modern economies affect the health of working people and to what extent these associations explain the social gradient of health.

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Occupational health research has tackled this latter challenge by applying two strategies of analysis: mediation and moderation. The mediation hypothesis claims that the strength of association between SEP and health is abolished or substantially weakened if the effect of work characteristics on health is estimated in multivariate regression models. A respective reduction in effect size is interpreted as a partial explanation of the social gradient of health by the work characteristic under study.^{15 16} Traditionally, in a majority of cases, epidemiological studies tested the mediation hypothesis by applying stepwise multivariate regression analysis. However, pathway analysis or structural equation modelling seem more appropriate statistical approaches as they allow for combined estimation of the direct and indirect effects of SEP and work characteristics on health.¹⁷

According to the moderation hypothesis the effect of a predicting variable (work characteristic) on a criterion variable (health) varies according to the level of a third variable (SEP). In this case, stronger effects of adversity at work on health are expected among employed people in less privileged as compared to more privileged socioeconomic conditions. To this aim, stratified analyses are performed, and an interaction term (of work and SEP on health) is assessed. Despite the direct policy implications of this latter approach—higher susceptibility to the exposure among people with low SEP points to priorities in administering interventions—the moderation hypothesis has been tested less frequently than the mediation hypothesis in research on work characteristics and social inequalities in health⁹ (see below).

To the best of our knowledge, no systematic review focusing exclusively on prospective observational cohort studies that tested the mediation and moderation hypothesis of the social gradient of health with reference to work characteristics is available so far. In this contribution, we set out to fill this gap. Our main aim was to document whether and to what extent findings from prospective studies lend support to the two hypotheses (mediation and moderation), even given considerable heterogeneity of work-related exposures and health outcomes in these studies. As both hypotheses address potential entry points for preventive activities at work that aim at reducing health inequalities,¹² it is important to assess the quality of available empirical evidence. This aim to some extent conflicts with another important aim of systematic reviews, and specifically meta-analyses, that is, to provide knowledge based on well comparable measures of specific exposures and specific outcomes. Throughout the paper, this tension will be discussed, and recommendations are proposed on how to strengthen further scientific evidence and its potential policy impact.

METHODS

The review was performed by observing the criteria defined in the PRISMA statement.¹⁸ Below we explain eligibility criteria concerning study design, sample and data analysis in more detail.

Study design

This review is restricted to prospective observational cohort studies as we aim at targeting the best available quality of data with regard to potential causal associations in epidemiological studies.¹⁹ Therefore we excluded cross-sectional studies. We also excluded longitudinal studies with a follow-up duration less than 1 year as a short time interval confers a high risk of reverse causation. At study entry, participants had to be free from the disease outcome under study, or baseline health measures had to be adjusted for in subsequent analyses. Furthermore, we

undertook a quality assessment regarding appropriate handling of adjustment procedures for relevant (eg, sociodemographic) confounders and appropriate statistical methods (eg, test for interaction in moderation analyses). To prevent multiple consideration of the same study in different papers, we selected the paper with highest data quality (eg, validity of outcomes and exposure, duration of longitudinal study and general quality assessment), but we included more than one report from the same study (in the case of the British Whitehall II and the French GAZEL study) if different health outcomes or different work characteristics (eg, different work stress models) were tested.

Study sample

We included studies of working age populations who were employed at entry. Studies with a sample size of less than 1000 were excluded for reasons of limited statistical power.

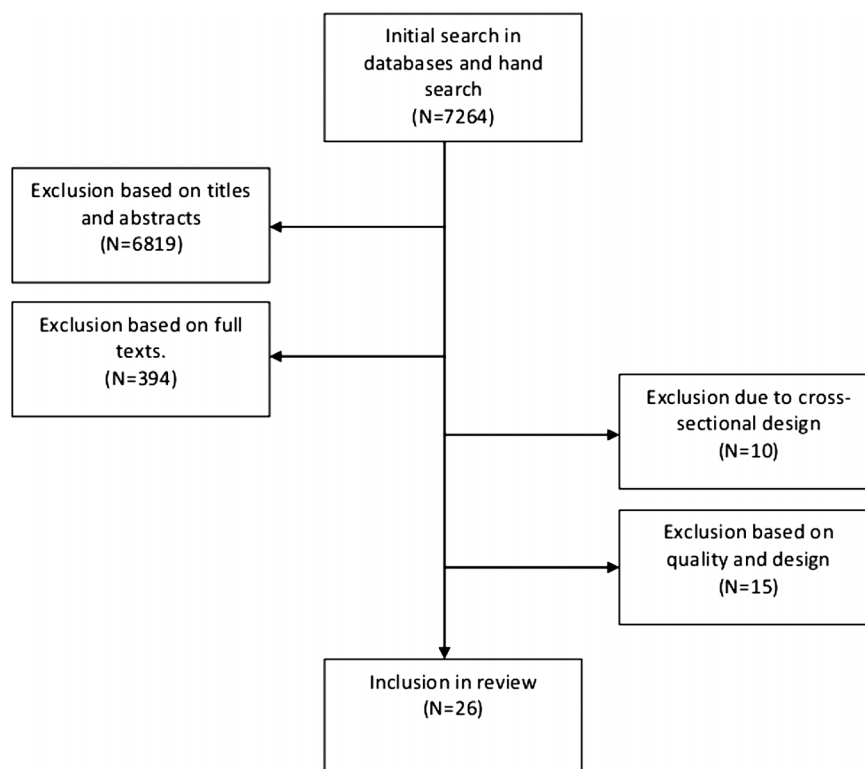
Exposures and health outcomes

Our selection criteria targeted studies with available data on SEP and on distinct health-adverse physical and chemical hazards or psychosocial working conditions. These data had to be assessed at baseline or during follow-up. Furthermore, the studies had to provide data on SEP and on health-adverse working conditions. SEP was measured by occupational position or employment grade in a majority of studies. However, several important investigations used education as a measure of SEP, and some studies applied both indicators. One study only focused on income as an SEP indicator. In view of the importance of each one of these internationally established indicators of SEP¹ we decided to include studies that applied at least one of them. Concerning the measures of stressful psychosocial work characteristics we included all available operational approaches as long as at least short versions of psychometrically validated scales or established indicators (eg, job insecurity) were used. With regard to health outcomes all available respective measures were included, whether based on self-report or on more objective information. Yet as common method variance between self-reported exposures and self-reported outcomes cannot be excluded, we gave priority to studies using more objective health measures by classifying these study results at the top in respective tables.

Information sources and data extraction

Systematic search was undertaken in PubMed and Scopus. The search was based on specified terms (see below), and original papers in peer-reviewed international journals published in English, French or German language between January 1980 and October 2012 were targeted. Conference papers and government-commissioned reports were not considered. Search terms included work environment factors and related health outcomes. In the first stage of the search strategy, based on titles and abstracts, we did not include terms measuring social inequalities, given a lack of consistent documentation of respective terms in abstracts. However, in a second stage, information on SEP was screened by reading all abstracts and full texts of respective articles. Search terms regarding work environment were: physical exposures, toxic exposures, psychosocial working conditions, job demands, job control, social support at work, job strain, effort–reward imbalance, and organisational justice. Search terms for health outcomes were: all-cause mortality, morbidity, cardiovascular disease, ischaemic heart disease, diabetes, metabolic diseases, myocardial, ischaemic, diabetes, work-related injuries, occupational accidents, suicide, musculoskeletal

Figure 1 Selection process for identification of studies.



disorders, health functioning, disability, disability pension, mental health, depression, and self-rated health. In addition to the systematic review procedure hand search research was conducted by contacting scientific teams involved in current work on social inequalities, work and health. This was done in order to include most updated evidence.

In a first stage, all records were judged by the first author on the basis of titles and abstracts. In difficult cases, full texts were consulted, and ambiguities were discussed between the authors (see flowchart figure 1). In a second stage both authors examined the remaining papers independently and searched for studies analysing the mediation or moderation hypothesis. To this aim, full texts were available, and the two independent ratings were compared. The few discordant cases were resolved by in-depth discussion. Data were extracted in a standardised format, according to categories indicated in supplementary tables S1 and S2 (available online only).

In most studies, mediation effects were evaluated using multivariate regression analysis with stepwise adjustments for confounders and mediators. In this procedure, the contribution of the mediating variable(s) to the explanation of social inequalities in health was estimated by the change in OR or HR after inclusion of the variable(s) in the model. The respective formula was $100 \times (\text{OR}_{\text{extended model}} - \text{OR}_{\text{Model 1}}) / (\text{OR}_{\text{Model 1}} - 1)$.²⁰ In case of reduction of OR or HR in the extended compared to the previous model, the percentage gives an estimate of how much the mediating variable(s) account for the social gradient of the health indicator under study. This percentage is interpreted as partial mediation contributing to the explanation of social inequalities in health. One study only applied path analysis to test the mediation hypothesis²¹ estimating the direct and indirect (work-related) effects of SEP on health. The moderation hypothesis was evaluated by stratified analyses in which the effect size on health was calculated for different SEP groups, assuming higher effect sizes among lower SEP groups.

RESULTS

Our search strategy provided 7264 initial records screened by the first author. Six thousand eight hundred and nineteen titles were excluded based on title and abstract content. The remaining 443 records were checked more precisely in full texts and were additionally evaluated according to whether relevant information (on SEP, on test of mediation or moderator hypothesis) was available. Given the restrictive selection criteria several interesting studies had to be excluded as they were based on a cross-sectional design ($n=10$), or did not meet additional quality or design criteria ($n=15$). Therefore, as demonstrated in figure 1, studies from 26 reports finally fulfilled all selection criteria to a sufficient extent. As one report contained results from two studies, 17 prospective studies tested the mediation hypothesis (see supplementary table S1, available online only), and nine prospective studies tested the moderation hypothesis (see supplementary table S2, available online only). Overall, it is apparent that relatively few papers investigated these mediation or moderation effects in the frame of cohort studies, whereas separate effects of socioeconomic or of work-related factors on health have been explored abundantly.

Mediation

In 13 studies, the most widely used model of a health-adverse psychosocial work environment, demand-control (or its single dimensions), was studied as the mediating construct.²² Two studies tested the effort-reward imbalance model²³ in addition to the demand-control model, and one study analysed the effort-reward imbalance model exclusively. One study tested a job-exposure matrix considering psychosocial working conditions.²⁴ Seven studies analysed physical demands or biomechanical strains in addition to the demand-control model, whereas one study was restricted to physical exposures and chemical substances.²⁵ The latter study was the only one that included a

large number of European countries, while the remaining studies were conducted in single countries of northern, southern or western Europe or Canada.

Health outcomes in those studies can be divided into more objective and more subjective measures. The former include cardiovascular diseases, lung cancer, disability pension and sickness absence, whereas the latter mainly concern self-rated health, depression, low back pain, or functional limitations. In a majority of studies SEP was assessed by occupational category (eg, ISCO-88) or employment grade (12 studies).^{20 21 24 26–34} In order to synthesise the current knowledge we included studies using education (two studies),^{16 25} income (one study),³⁵ or the combination of employment grade and education (one study).³⁶ In addition, one study tested occupation and education as alternative indicators.³⁷ As mentioned, one study only applied pathway analysis, while the majority of reports tested the mediation hypothesis using stepwise regression analysis. It should be noted that four out of the 16 studies were based on data from the British Whitehall II study, and two reports were based on the French GAZEL cohort.

All studies report a social gradient of health. In addition, poor working conditions are more prevalent among employed people with low skill level or low occupational standing. As evident from supplementary table S1 (available online only) OR or HR of low versus high SEP are generally higher in reports that used self-reported health outcomes compared to those using medical diagnoses (except Toivanan and Hemström).³⁵ The crucial information about the mediation hypothesis is provided by comparing the percentage reduction in OR or HR between the first step of a regression model (SEP and health) and the second step of the model, adjusting additionally for work characteristics. In a majority of cases, a percentage reduction between the two OR or HR is observed, as assumed by the mediation hypothesis. However, the amount of percentage reduction, that is, the strength of a mediation effect, varies considerably. This variation is of interest in three respects, first with regard to the applied SEP indicator, second with regard to objective versus subjective health indicators, and third with respect to the mediating construct, that is, work characteristics, although these conditions overlap within the studies.

In general, mediation effects are more consistently observed in studies based on employment grade as a measure of social inequality, compared to those using alternative SEP indicators. This conclusion is supported by the study that provides an alternative test of both SEP indicators and observes stronger percentage reduction in the case of employment grade.³⁶

Concerning a comparison of mediation effects between studies using more objective versus subjective health outcomes an interesting finding becomes obvious. With two exceptions, mediation effects tend to be somewhat stronger if objective health outcomes are analysed. One of these exceptions relates to depression in which a combination of both psychosocial work stress models contributes towards explaining the social gradient quite remarkably, especially among men.³² The other exception is observed in a study focusing on biomechanical stressors with regard to low back pain.³³ This latter finding is best understood in terms of the third source of variation, work characteristics. It appears that studies that combine psychosocial and physical work stressors achieve relatively stronger mediation effects although two studies only analysed these two effects separately.^{28 33} As the demand–control model was applied in a majority of studies no comparison of mediation effects with alternative psychosocial exposures was possible. Some study results confirm that this model, or its single components, makes

a distinct contribution towards explaining the social gradient of health.

Moderation

Five of the nine studies included in this part of the review used the demand–control model,^{9 38–41} two studies were based on the effort–reward imbalance model,^{42 43} one study analysed several aspects of work-related social support,⁴⁴ and one study applied a job-exposure matrix of different occupational solvents.⁴⁵ Given the relatively small number of studies, a further differentiation according to SEP indicators or health outcomes is not feasible. It should also be kept in mind that any generalisation of findings related to the moderation hypothesis is limited by the fact that three out of nine reports are based on the same cohort, the British Whitehall II study. Although they address different health outcomes and different time frames within the longitudinal design there is a clear risk of over-reporting.

According to the moderation hypothesis stronger effects of work characteristics on health outcomes are expected in low SEP as compared to high SEP groups. This hypothesis finds empirical support in four studies. Two of them tested the effort–reward imbalance model,^{42 43} one the demand–control model,⁴⁰ and one the chemical exposure of different solvents.⁴⁵ A fifth investigation reports that a mitigating effect of a favourable psychosocial work environment (social support and job security) on the amount of experienced distress is confined to the subgroup with low socioeconomic standing, while it is absent among higher SEP groups.⁹ Three of the four remaining studies observed higher OR or HR among more privileged as compared to less privileged occupational groups, thus contradicting the general hypothesis.^{38 39 44} However, in one of those studies this only holds true for men,³⁸ and in another study this effect is restricted to one out of three work characteristics entering statistical analysis⁴⁴ (see supplementary table S2, available online only).

In view of the small number of studies and the heterogeneity of relevant measures it is difficult to find a consistent pattern of results with regard to the moderation hypothesis. Yet slightly more results are in favour of the hypothesis, whereas negative findings from at least three studies challenge this assumption.

DISCUSSION

To our knowledge, this is the first systematic review of available cohort studies testing the mediation or the moderation hypothesis of adverse working conditions in the context of social inequalities in health. Based on an analysis of 27 prospective investigations we found some empirical support in favour of either assumption. However, given the heterogeneity of measures applied in these studies the degree of consistency of findings was clearly restricted. The decision on including reports using different indicators (specifically of SEP and of health outcomes) was based on the premise that these two hypotheses offer important entry points of preventive activities at work with potential benefit for reducing health inequalities and, thus, that systematic knowledge on a broad spectrum of current evidence would instruct these activities.

Several findings deserve attention. Concerning the mediation hypothesis, first, a majority of studies support the notion that adverse working conditions to some extent mediate the association of SEP with health. Respective percentage reductions in OR were sometimes modest, but in a few instances rather substantial, in particular if physical stressors were included or combined with psychosocial stressors. Second, studies that

used occupational categories or employment grades as indicators of SEP provided more robust findings than those using education or income as indicators. Third, with two noticeable exceptions, studies based on objective health measures demonstrated somewhat stronger mediation effects than studies based on self-reported health measures. Concerning the moderation hypothesis, results are less consistent as five out of nine studies only support the hypothesis of stronger effects among lower versus higher socioeconomic groups. Three of those studies used the effort–reward imbalance model or the demand–control model, and one applied a job exposure matrix of chemical solvents. On balance, in view of moderate support of the mediation hypothesis and modest support of the moderation hypothesis, it seems appropriate to draw preliminary rather than firm conclusions from the available evidence, given the relative paucity of studies, the heterogeneity of definition criteria and measures of core variables, and the limitations of applied statistical analyses.

There is a remarkable contrast of the consistency and strength of reported effects between the prospective investigations analysed in this contribution and the cross-sectional studies that tested the two hypotheses so far. In this latter case, a rather high degree of consistency of findings was observed, supporting either hypothesis (for review).^{5 6 9 37 46 47} It is unclear to what extent this discrepancy in the consistency of results according to study design is due to methodological problems (eg, higher risk of reporting bias or common method variance in cross-sectional studies) or to different quality of studies (less heterogeneity of concepts and measures in cross-sectional studies, in combination with more systematic hypothesis testing). In view of the relevant contribution of employment and working conditions to social inequalities in health⁴⁸ there is an urgent need for further clarifying this divergent trend.

Whereas prospective observational cohort studies represent the gold standard in this area of research, cross-sectional investigations may be in a better position to contribute to conceptual and methodological innovations, given a shorter time frame of overall research investment. For instance, one recent important cross-sectional study extended the frame of analysis by testing the contribution of material and psychosocial factors in occupational life and in additional life domains, thus quantifying their respective contribution towards explaining social inequalities in health.⁴⁶ Moreover, that study included data from 28 European countries, thus providing opportunities of testing differential associations with regard to distinct national social and labour policies (see below).

This systematic review has several limitations. We restricted the search strategy in terms of type of publication (original research articles), language (English, French, German), number of databases consulted (PubMed, Scopus), and choice of search terms. Therefore, we cannot exclude the possibility that some studies bypassed our review. Second, it is entirely possible that due to publication bias studies are overrepresented that reported positive findings. Third, given more sophisticated statistical approaches towards testing the mediation hypothesis,¹⁷ the majority of studies on which this review is based applied the conventional approach of stepwise regression modelling, which may not provide definite evidence on true mediation effects. Finally, as mentioned, given the heterogeneity of measures across studies, a meta-analysis comparing effect strengths of different study findings was not feasible. This fact restricts the quality of this review, which nevertheless classifies and quantifies available data.⁴⁹

In view of a high amount of heterogeneity concerning core measures applied in the cohort studies and in view of

potential advances of more recently developed statistical approaches, it seems timely to call for a higher degree of standardisation and methodological sophistication in future research in this field. In particular, the following criteria might be observed in designing and conducting respective empirical studies.

First, studies should test an explicit theoretically justified hypothesis, rather than exploring what variables may produce statistically significant results. This requires an a priori definition of the core variables entering the analysis of mediation or moderation hypotheses. Second, given the paucity of studies measuring physical and chemical hazards at work, and given the fact that the effects of these exposures were generally strong, there is an urgent need to analyse the combined and separate effects of physical/chemical and psychosocial work characteristics in future interdisciplinary research. Moreover, concerning a health-adverse psychosocial work environment, appropriate tests of respective theoretical models based on psychometrically validated scales should be performed. Along these lines one can argue that respective analyses may even be broadened by including distinct extra-work conditions (as exemplified in one recent study).⁴⁶ Third, with regard to health outcomes, more emphasis could be put on objective outcomes of functioning, morbidity and mortality, thus improving criterion validity. As mentioned, in this review the effect sizes of studies analysing objective health indicators tended to be stronger than those based on self-reported health outcomes. Fourth, while the generation of cumulative knowledge based on established theoretical models, such as the demand–control or the effort–reward imbalance model, is an important scientific goal, these models may need some extension or modification in order to capture more recent trends of health-adverse work and employment appropriately (eg, atypical work, contract work, self-employment; see Clougherty *et al*).⁶ There is a danger that results from cohort studies (which are predominantly conducted in large companies) are biased by overemphasising those stable work and employment conditions that are more prevalent in large organisations, thereby disregarding the health burden of unstable, precarious work. Fifth, study designs and statistical analyses should take account of the complexity and dynamics of work life in current societies. Among others, structural equation modelling, path analysis and multilevel analysis offer opportunities for respective extensions. Event history methods and multiple longitudinal exposure assessments are desirable, when feasible. The same holds true for the inclusion of contextual measures of workplaces, departments, firms and organisational environments. Finally, in times of economic globalisation, comparative research on the role of work and employment conditions in explaining health inequalities between countries is considered a high priority. With a few exceptions study findings so far are restricted to economically developed countries, mostly in northern or western Europe. Therefore, it will be important to analyse the two hypotheses in non-western countries and in rapidly developing societies. Preliminary results from internationally comparative studies testing the demand–control and/or the effort–reward imbalance model of adverse work with regard to health indicate that its main results can be replicated in eastern European countries,^{50 51} in rapidly developing countries (for China see Li *et al*⁵² and Xu *et al*),⁵³ and in countries with markedly different cultures (for Japan see Tsutsumi *et al*⁴⁰ and Siegrist *et al*).⁵⁴ Moreover, cross-country analyses provide an opportunity of testing the impact of distinct national labour and social policies on the quality of work and its effects on workers' health.^{55 56}

In conclusion, the current state of research, as reflected by findings from prospective observational cohort studies, provides some support in favour of the mediation and moderation hypotheses in analysing associations of work characteristics and SEP with health. Given a high degree of heterogeneity of concepts, measurements and methods of data analysis, more standardised research procedures are needed. To this end, several propositions for the generation of improved knowledge were made. It is hoped that, as a result, improved scientific evidence will be available to instruct work and employment-related policies that aim at reducing social inequalities in health.

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Table 1 Mediation effects in prospective studies.

Author, year	Journal	Study design and setting	Socioeconomic position	Work characteristics	Health outcomes	Adjustments	Main findings %-Reduction by mediation
Marmot et al. 1997 ²⁶	The Lancet	London-based Civil servants N=7372 Mean follow-up: 5.3 yrs	Employment grade: high, medium, low	job control, effort-reward imbalance	CHD (Angina Pectoris, Severe chest pain, diagnosed ischaemia)	Age, separated by sex	Reduction of OR after adjustment for job control in men of low employment grade (ref.: high): 64%* (OR: 1.50 to 1.18). In women: 51%* (OR: 1.47 to 1.23).
Andersen et al. 2004 ²⁴	Occupational and Environmental Medicine	Copenhagen, Denmark N=16216 Follow-up: 4-22 yrs	Occupational position in 5 categories	JEM: psychosocial working conditions: skill discretion and decision authority	Myocardial infarction	Cohort of investigation, age, cohabitation, sex, smoking, BMI, alcohol intake, physical activity, Systolic blood pressure, cholesterol	Reduction of HR after adjustment for skill discretion in unskilled workers (ref.: executive managers) by 79%* (HR: 1.33 to 1.07 (n.s.)). Reverse trend after adjusting for decision authority.
Chandola et al. 2005 ²¹	Occupational and Environmental Medicine	London-based Civil servants N= 3697 Follow-up: up to 16 yrs	Employment grade in 3 categories	Effort-reward imbalance	Angina	Age, obesity, blood pressure. Separated by sex.	Men: Small but significant direct and indirect effects (through imbalance at phase 5) of employment grade on angina. The effect of grade on angina is in part mediated by ERI. Women: direct effects of grade on angina, but no indirect effects through ERI.
Melchior et al. 2005 ²⁷	American Journal of Public Health	France Employees of a gas and electricity company N=11733 Follow-up: 6 yrs	Employment grade in 6 categories	Physical work factors (postural complaints Occupational hazards Night work Outdoor work activities Customer contact) Job strain	All-cause Sickness absence	Age, demographics, health behaviours, stressful life events Separated by sex	Reduction of occupational class gradient of all-cause sickness absence after adjustment for work characteristics in male manual workers (ref.: managers) by 23% (RR: 3.06 to 2.95). In male clerks: 24% (RR: 2.88 to 2.42), female clerks (ref.: managers): 26% (RR: 2.76 to 2.30).
Hagen et al. 2006 ¹⁶	Social Science & Medicine	Nord-Trøndelag, Norway N=26823 Follow-up: 7 yrs	Education	Physical demands, concentration and attention, stress and tension, authority to plan own work, job satisfaction	Disability pension caused by Back Pain	Age, separated by sex	Reduction of HR (men: 0.77; women: 0.76) after adjustment for work characteristics: 24% (HR: 0.82) in men and 13% (HR: 0.79) in women.

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Toivanen et al. 2006 ³⁵	International Journal of Behavioral Medicine	Sweden - ULF survey Age: 40-64 N=6405 follow-up: 6 yrs - Swedish Census Age 40-64 N= 1,858,373 Follow-up: 5 yrs	Income	Job control Physical work demands	Cardiovascular disease (ULF) Mortality from Cardiovascular disease (Census)	-Year, age, sex, smoking -Age, sex	ULF: Reduction in OR after adjustment for work characteristics: 8-10% in income quartiles (OR of lowest income quartile (ref.: highest): 3.63-3.43). Swedish Census: Reduction in RR after adjustment for dichotomized job control in all income quartiles 10% each (OR of lowest income quartile (ref.: highest): 2.10-1.99), job control in quartiles: reduction amounts to 14-18% (OR of lowest income quartile (ref.: highest): 2.10-1.95).
Huisman et al. 2008 ³⁷	Social Science & Medicine	southeastern Netherlands N=5757 Follow-up: up to 12 yrs	Education Occupation in 3 categories	Job strain	Myocardial infarction	Sex, age, marital status	Reduction of HR of education after adjustment for job strain: 23%* HR: (1.97 to 1.75). Reduction in HR of occupation after adjustment for job strain in manual workers (ref.: non-manual): 44%.* (HR: 1.62 to 1.35(n.s.)).
Laaksonen et al. 2009 ²⁸	Journal of Epidemiology and Community Health	Helsinki, Finland N=6934 Mean follow-up: 3,9 yrs	Occupational class in 4 categories	Work arrangements (working hours, shift work, permanent and temporary work contract), physical (physical work load, exposure to hazardous substances, sedentary work and computer use) and psychosocial (job demands, job control, social support at work, job satisfaction, workplace climate) working conditions	Sickness absence	age	Men: HR of work arrangements among manual workers (ref.: managers and professionals) is 2.78, physical working conditions: 2.08, psychosocial working conditions: 2.87. Adjustment for work arrangements slightly narrowed the class differences. Physical working conditions attenuated the social gradient by about 40 %. Women: HR of work arrangements among manual workers (ref.: managers and professionals) is 2.98, physical working conditions: 2.00, psychosocial working conditions: 3.41. Adjustment for work arrangements and physical working conditions narrowed the class differences, and psychosocial working conditions widened them.
Menvielle et al. 2010 ²⁵	International Journal of Cancer	Europe 23 centers in 10 countries Men N=88,265 Mean Follow-up: 8.4 yrs	Education	Exposure to asbestos, heavy metals, and PAH	First primary lung cancer	Smoking, dietary habits	Reduction of HR after adjustment for job exposures in lowest education: 12%* (HR: 1.6 to 1.53).

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Ferrario et al. 2011 ²⁹	Occupational and Environmental Medicine	Brianza, Italy N= 1441 Median Follow-up: 12 yrs (8-16 yrs)	Occupational class in 4 categories	Job strain	First CHD event	Age, systolic blood pressure, total cholesterol, HDL-cholesterol, current smoking, diabetes mellitus, alcohol intake, sport physical activity index	Reduction of HR of manual workers (ref.: non-manual) from 1.18 (n.s.) to 1.12 (n.s.). Differences between occupational classes are not significant.
Haukenes et al. 2011 ³⁶	BMC Public Health	Hordaland County, Norway Age: 40-45 N=7031 Follow-up: 7 yrs	Employment grade (ISCO-88 and education) in 4 categories	Job demands Job control Physical demands Paid working hours years in current occupation	Disability pension	Sex, health at baseline	Reduction of HR after adjustment for work characteristics: 24% (HR: 2.78 to 2.12) in unskilled manual workers (ref.: administrators and professionals).
Borg et al. 2000 ²⁰	Social Science & Medicine	Denmark N=4751 Follow-up: 5 yrs	Employment grade in 5 categories, based on job title and education	Job demands, decision authority, social support, skill discretion, conflicts at work, job insecurity, repetitive work ergonomic, chemical, climatic, and physical exposures	Self-rated health	Age, sex, disease, injury or long lasting illness	Reduction of OR (4.23) after adjustment for ergonomic exposures: 34% (OR: 3.13); repetitive work: 24% (OR: 3.44); job insecurity (OR: 3.9), climatic (OR: 3.88), and physical exposures (OR: 3.95) in each case ca. 10% (OR: 3.88 to 3.95). The combined adjustment for this exposures explained 59% (OR: 2.34) of the gradient.
Ferrie et al. 2003 ³⁰	Social Science & Medicine	London-based Civil servants N= depends on model: 1089 to 3239 Follow-up: up to 14 yrs	Employment grade in 2 categories	Job insecurity Financial insecurity	Self-rated health Depression	Age Separated by sex.	Reduction of OR of self-rated health (men: 2.24; women: 4.35) after adjustment for job insecurity by 6% in men and 4% in women. Reduction of OR of self-rated health (men: 2.23; women: 3.92) after adjustment for financial insecurity by 33% in men and 13% in women. Reduction of Diff of depression (men: 0.51; women: 0.43) after adjustment for job insecurity by 9% in men and 16% in women. Reduction of Diff of depression (men: 0.49; women: 0.52) after adjustment for financial insecurity by 49% in men and 14% in women.
Mustard et al. 2003 ³¹	Social Science & Medicine	Canada N=5691 Follow-up: 4 yrs	Employment grade in 4 categories	Work demands/ work control Job strain	Self-perceived general health status	Age, sex, health conditions (chronic conditions, back pain, BMI, psychological distress and depression) at baseline	Reduction of OR of the lowest occupational position (ref.: highest) by 17% (OR: 1.72 to 1.60) in men after adjustment for job demands/control and by 11% (OR: 1.72 to 1.64) after adjustment for job strain. No reduction in women and in the full sample.

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Stansfeld et al. 2003 ³²	Journal of Epidemiology and Community Health	London-based Civil servants N=5912 Follow-up: 11 yrs.	Employment grade as linear term	Decision latitude Job demands Work social supports ERI	Depressive symptoms	Age, separated by sex	Decline of grade gradient in depression after adjustment: 66% (coefficient: 0.105-0.036) in men and 43% (0.096-0.055) in women.
Plouvier et al. 2009 ³³	Scandinavian Journal of Work, Environment & Health	France Male Employees of a gas and electricity company N=1487 Follow-up: 12 yrs	Employment grade in 4 categories	Biomechanical strains (driving, pushing, pulling or carrying heavy loads, bending or twisting) Psychosocial strains (psychological demands, decision latitude, social support)	Low back pain	Age	Reduction of PR after adjustment for biomechanical strains (ref. managers): in blue collar workers: 73% (PR: 1.92 to 1.25). Adjustment for psychosocial strains: in blue collar workers: 11% (PR: 1.92 to 1.81). Reduction after combined adjustment: 77% (PR: 1.92 to 1.21).
Pietiläinen et al. 2012 ³⁴	Journal of Epidemiology and Community Health	Helsinki, Finland N= 7332 Mean Follow-up: 6 yrs	Occupational class in 4 categories	Physical and psychosocial (demand-control-model) working conditions	Physical functioning	Age, separated by sex	Women: Physical working conditions had marginal effects on the difference between occupational classes, adding psychosocial working characteristics narrowed the differences. Men: Adjusting for physical working conditions slightly widened the not significant differences between the occupational classes; psychosocial working conditions slightly narrowed the differences.

*own calculations based on ORs/HRs reported in the paper

OR= Odds Ratio, HR= Hazard Ratio, RR= Rate Ratio, PR= Prevalence Ratio, IRR= Incidence Rate Ratio, Diff= difference, n.s.= not significant, ERI= Effort-reward imbalance, CHD= Coronary heart disease, BMI= Body Mass Index, JEM= Job Exposure Matrix

Table 2 Moderation effects in prospective studies.

Author, year	Journal	Study design and setting	Socioeconomic position	Work characteristics	Health outcomes	Adjustments	Main findings
Hemingway et al. 1997 ³⁸	Scandinavian Journal of Work, Environment & Health	London-based Civil servants N= 5620 Mean follow-up: 4 yrs	Employment grade (3 categories in men, 2 categories in women)	Job control	Sickness absence due to back pain	Age, education, housing tenure, access to use of car, BMI, exercise, smoking habits, number of questionnaire reports of back pain at baseline, separated by sex	Men: RR of low job control (ref.: high control) is 3.42 in the top grade, and 0.78 in the lowest grade. Women: RR of low job control is 0.80 in the higher grade and 1.35 in the lower grade.
Kuper et al. 2002 ⁴²	Occupational and Environmental Medicine	London-based Civil servants N= 9870 Mean follow-up: 11 yrs	Employment grade in 3 categories	ERI	CHD	Age, sex	OR of ERI is 1.56 in clerical, 1.26 (n.s.) in professional and 1.19 (n.s.) in administrative workers.
Kuper et al. 2003 ³⁹	Journal of Epidemiology and Community Health	London-based Civil servants N=9746 Median follow-up: 11.2 yrs	Employment grade in 3 categories	Job strain	CHD (non-fatal myocardial infarction, angina events)	Age, sex	HR of high demand/low control (ref. low demand/high control) is 1.31 (n.s.) in clerical, 1.51 in professional and 1.81 in administrative workers.
Tsutsumi et al. 2011 ⁴⁰	Social Science & Medicine	Japan N= 6553 Follow-up: 11 yrs	Occupational class (white-/blue-collar) Occupational position (manager/non-manager)	Job strain	Stroke	Age, educational attainment, smoking status, alcohol consumption, physical activity, separated by gender	Men: HR of high job strain (ref. low strain) is 3.1 among blue collar and 1.4 (n.s.) among white collar workers. HR is 8.9 among non-managers and 2.0 (n.s.) among managers. Women: HR of high job strain is 5.6 among blue collar and 1.0 (n.s.) among white collar workers. HR is 5.3 among non-managers and 0.7 (n.s.) among managers.
von Bonsdorff et al. 2012 ⁴¹	British Medical Journal	Finland N= 5731 Follow-up: 28 yrs	Occupational class (blue- and white collar)	Job strain	Mortality	Age, smoking, alcohol intake, physical activity, prevalent CVD, metabolic disorders and cancer. Separated by sex	Men: HR of high strain (ref.: low strain) is 1.38 (n.s.) in white collar, and 1.09 (n.s.) in blue collar workers. Women: HR of high strain (ref.: low strain) is 0.81 (n.s.) in white collar, and 0.97 (n.s.) in blue collar workers.

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Väänänen et al. 2004 ⁴⁴	Social Science & Medicine	Finland Industrial company N= 2225 Follow-up: 4 yrs	Occupational class (white- and blue-collar)	Experienced change in job position Organizational support Supervisor's support Coworker's support	Functional incapacity	Age, sex, sickness absence, pre-merger functional incapacity	OR of weak Supervisor's support (ref.: strong support) is 2.13 among white collar workers and 1.19 among blue collar workers OR of weak organizational support is 1.71 in both white- and blue-collar workers. OR of weak coworker's support is 1.37 (n.s.) among white collar workers and 1.24 (n.s.) among blue collar workers.
Ibrahim et al. 2009 ⁹	Social Science & Medicine	Canada, N=2556 Follow-up: 2-6 yrs	Employment grade in 2 categories	Job strain Work social support Job insecurity	Depression Distress Self-rated health	Age, sex, marital status and part-time/full-time work hours	Higher social support at work and low job insecurity were each associated with lower distress among the lower occupational group, but not in the higher group. The other pathways showed only limited differences among occupational groups.
Rugulies et al. 2012 ⁴³	European Journal of Public Health	Denmark Follow-up: 5 yrs N=2701	Employment grade in 3 categories	ERI	Severe depressive symptoms	Sex, age, family status, survey method, smoking, heavy alcohol consumption, leisure time physical activity, self-rated health, sleep disturbances, non-severe depressive symptom score at baseline	OR of high ERI is 4.08 among manual workers, 2.11 (n.s.) among low grade non-manual workers and 1.30 (n.s.) among executives, academics and middle managers.
Sabbath et al. 2012 ⁴⁵	Neurology	GAZEL Male French national gas and electricity employees N=4134 Follow-up:	Education (dichotomized)	JEM: Lifetime exposure to 4 solvent types (chlorinated, petroleum, benzene, and nonbenzene aromatic solvents)	Cognition	age, smoking status, alcohol consumption, Occupational grade at age 35	Higher RRs of exposure to solvents among less-educated individuals than in more-educated. Significant associations of exposure to petroleum, benzene, and aromatic solvents among the less-educated, no significant associations among the more-educated.

OR= Odds Ratio, HR= Hazard Ratio, RR= Rate Ratio, n.s.= not significant, ERI= Effort-reward imbalance, CHD= Coronary heart disease, CVD= cardiovascular disease, BMI= Body Mass Index, JEM= Job Exposure Matrix