

### ORIGINAL RESEARCH ARTICLE

# Association of occupational exposures with cardiovascular disease among US Hispanics/Latinos

Catherine M Bulka,<sup>1,2</sup> Martha L Daviglus,<sup>2</sup> Victoria W Persky,<sup>1</sup> Ramon A Durazo-Arvizu,<sup>2</sup> James P Lash,<sup>3</sup> Tali Elfassy,<sup>4</sup> David J Lee,<sup>4</sup> Alberto R Ramos,<sup>5</sup> Wassim Tarraf,<sup>6</sup> Maria Argos<sup>1</sup>

#### ABSTRACT

► Additional material is published online only. To view please visit the journal online (http://dx.doi.org/10.1136/ heartjnl-2018-313463).

For numbered affiliations see end of article.

#### Correspondence to

Dr Maria Argos, Division of Epidemiology and Biostatistics, School of Public Health, University of Illinois at Chicago, Chicago IL 60612, USA; argos@ uic.edu

Received 30 April 2018 Revised 6 September 2018 Accepted 9 September 2018 Published Online First 11 December 2018



► http://dx.doi.org/10.1136/ heartjnl-2018-314130

#### Check for updates

© Author(s) (or their employer(s)) 2019. Re-use permitted under CC BY-NC. No commercial re-use. See rights and permissions. Published by BMJ.

To cite: Bulka CM, Daviglus ML, Persky VW, et al. Heart 2019;105:439–448. **Objective** Cardiovascular disease (CVD) is a leading cause of mortality and morbidity in the USA. The role of occupational exposures to chemicals in the development of CVD has rarely been studied even though many agents possess cardiotoxic properties. We therefore evaluated associations of self-reported exposures to organic solvents, metals and pesticides in relation to CVD prevalence among diverse Hispanic/Latino workers.

**Methods** Cross-sectional data from 7404 employed individuals, aged 18–74 years, enrolled in the Hispanic Community Health Study/Study of Latinos (HCHS/ SOL) were analysed. Participants from four US cities provided questionnaire data and underwent clinical examinations, including ECGs. CVD was defined as the presence of at least one of the following: coronary heart disease, atrial fibrillation, heart failure or cerebrovascular disease. Prevalence ratios reflecting the relationship between each occupational exposure and CVD as well as CVD subtypes were calculated using Poisson regression models.

**Results** Hispanic/Latino workers reported exposures to organic solvents (6.5%), metals (8.5%) and pesticides (4.7%) at their current jobs. Overall, 6.1% of participants had some form of CVD, with coronary heart disease as the most common (4.3%) followed by cerebrovascular disease (1.0%), heart failure (0.8%) and atrial fibrillation (0.7%). For individuals who reported working with pesticides, the prevalence ratios for any CVD were 2.18 (95% CI 1.34 to 3.55), coronary heart disease 2.20 (95% CI 1.31 to 3.71), cerebrovascular disease 1.38 (95% CI 0.62 3.03), heart failure 0.91 (95% CI 0.23 to 3.54) and atrial fibrillation 5.92 (95% CI 1.89 to 18.61) after adjustment for sociodemographic, acculturation, lifestyle and occupational characteristics. Metal exposures were associated with an almost fourfold (3.78, 95% CI 1.24 to 11.46) greater prevalence of atrial fibrillation. Null associations were observed for organic solvent exposures.

**Conclusions** Our results suggest that working with metals and pesticides could be risk factors for CVD among Hispanic/Latino workers. Further work is needed to evaluate these relationships prospectively.

#### BACKGROUND

Recent studies have linked working with certain chemicals to markers of adverse cardiovascular health such as increased blood pressure, arterial stiffness and decreased heart rate variability.1-5 However, it remains unknown if exposures to these workplace hazards could manifest as cardiovascular disease (CVD). Cohort studies of aircraft maintenance workers, battery factory workers, miners and pesticide applicators exposed to solvents, metals and pesticides have found inconsistent relationships with rates of CVD death.<sup>6-9</sup> Although studies confined to specific occupations can be useful in identifying the health issues faced by particular types of workers, they are often limited by small sample sizes, ascertainment of clinical parameters, collection of lifestyle factors that could confound exposure-disease relationships and generalisability to wider populations. In addition, such studies are prone to bias from the healthy worker survivor effect since many occupations with high probabilities for exposure require employees meet certain standards of physical health.<sup>10</sup> As a result, measures of association for deleterious exposures can appear underestimated, or even biased so much that harmful risk factors appear protective. A community-based approach that captures a variety of occupations and industries may mitigate some of this bias by including individuals employed in less strenuous positions.

The assessment of occupational exposures in community-based studies is rare, in part because they are logistically difficult to validly measure on a large scale. A common approach is to link job titles to job-exposure matrices to infer probable exposures. However, misclassification can arise given that exposures may vary widely among individuals with the same title.<sup>11</sup> Moreover, the use of generic exposure matrices within a study comprised wholly of Hispanics/Latinos may not fully capture the work experiences of this distinct population. Hispanic/Latino workers may be disproportionately exposed to cardiotoxic agents compared with other racial/ethnic groups and uniquely vulnerable to workplace hazards because of language barriers, low educational attainment and limited resources. A lack of culturally appropriate job safety training coupled with fears surrounding job security and immigration status could make Hispanic/Latinos engaged in hazardous jobs even more susceptible to occupational illnesses. As an initial strategy, we used interviewer-administered questionnaires to examine cross-sectional associations of self-reported organic solvent, metal and pesticide occupational exposures with CVD.

BMJ



## METHODS

#### Study population

The Hispanic Community Health Study/Study of Latinos (HCHS/SOL) is a community-based prospective cohort study designed to evaluate risk factors and chronic diseases among the rapidly growing Hispanic/Latino population across four cities in the USA (Bronx, New York; Chicago, Illinois; Miami, Florida; and San Diego, California). HCHS/SOL included 16415 men and women, aged 18-74 years, who self-identified as Hispanic/ Latino at baseline (2008–2011). The sampling design and cohort selection have been previously described.<sup>12</sup> <sup>13</sup> Briefly, participants were recruited through a multistage probability sampling design. Sampling weights were established to represent the probability of selection at each stage. Baseline examinations included occupational information, sociodemographics, medical history, medication use, tobacco use, alcohol use, physical activity, dietary recall, anthropometry, blood pressure, phlebotomy and electrocardiography.

#### **Exposure assessment**

An occupational questionnaire designed to assess work environments and exposures was administered to all employed participants by a trained interviewer in either English or Spanish, based on the participant's preference. Participants were asked 'At the job you currently work the majority of your work hours per week, how often are you exposed to any type of organic solvents, for example styrene, trichloroethylene, toluene, or xylene?' and 'At the job you currently work the majority of your work hours per week, how often are you exposed to metals such as manganese, lead, or mercury?'. Response options were none of the time, 25% of the time, 50% of the time, 75% of the time, 100% of the time, occasionally and do not know. For this analysis, responses were collapsed into yes (occasional or 25%-100% of the time) or no (0%) because of data sparseness. Participants were also asked 'In your current job(s), are you exposed to pesticides?', with responses recorded as yes or no. Participants unfamiliar with the term 'pesticides' were classified as not exposed.

#### **Outcome assessment**

CVD prevalence was assessed using self-reported information from the medical history questionnaire or from a digital 12-lead ECG performed at baseline. Details regarding HCHS/SOL ECG procedures and subsequent readings performed at a centralised location (EPICARE, Wake Forest University School of Medicine, Winston-Salem, North Carolina) have been described elsewhere.<sup>14</sup> Prevalent CVD was defined as at least one of the following: coronary heart disease, heart failure, atrial fibrillation or cerebrovascular disease. Participants who self-reported that a doctor told them they had angina, a myocardial infarction or reported undergoing coronary bypass surgery, balloon angioplasty or stent placement in coronary arteries, or had ECG evidence of major Q wave abnormalities or minor Q, QS waves with ST and T abnormalities were considered to have coronary heart disease. Atrial fibrillation was defined as either a self-reported physician diagnosis or ECG evidence of atrial fibrillation or flutter. Heart failure was based solely on self-report (a doctor told the participant he or she had heart failure). Cerebrovascular disease was defined as self-reported physician diagnosis of stroke or transient ischaemic attack.

#### Sociodemographic characteristics

The interviewer-administered enrolment questionnaire asked participants for sociodemographic information. Hispanic/Latino

background groups were categorised as Dominican, Central American, Cuban, Mexican, Puerto Rican, South American and more than one heritage/other heritage. Educational attainment was categorised as not having a high school diploma or General Equivalency Diploma (GED), having a diploma or GED or attaining an education beyond a high school equivalent (ie, college or vocational). Health insurance was defined as current coverage through an employer, individual plan, Medicaid/Medicare, military, Indian Health Services or another provider.

#### Lifestyle characteristics

Alcohol intake was classified as none, low/moderate (<7 drinks/ week for females; <14 drinks/week for males) or heavy  $(\geq 7 \text{ drinks/week for females}; \geq 14 \text{ drinks/week for males})$ . Cigarette smoking status was categorised as never, former or current. A modified Global Physical Activity Questionnaire based on the original developed by the WHO was used to collect information on physical activity in a typical week across three domains: work, recreation and transport. Total physical activity across these domains was categorised as low, moderate or high based on established criteria.<sup>15</sup> Dietary quality was assessed using the 2010 Alternative Health Eating Index (AHEI-2010), which was derived from up to two 24-hour dietary recalls. The index combines intakes of vegetables, fruit, whole grains, sugar-sweetened beverages and fruit juice, nuts and legumes, red/processed meats, trans fats, long-chain (n-3) fatty acids, polyunsaturated fatty acids, sodium and alcohol intakes, with higher scores representing a better quality diet.<sup>16</sup>

#### Acculturation characteristics

Residential duration in the mainland USA and the Short Acculturation Scale for Hispanics (SASH) language subscale were used as measures of acculturation. The SASH language subscale is based on five-point Likert-type scale questions; the average score was calculated with higher scores indicating a higher degree of language acculturation.<sup>17</sup>

#### **Occupational characteristics**

Full-time employment was defined as working more than 35 hours per week in one job or more than one job; part-time employment was defined as working 35 or fewer hours per week. Primary occupations were categorised into five groups: non-skilled workers (typically does not require a high school degree and involves manual labour), service workers (provides personal services to individuals), skilled workers (requires specific skills that typically requires technical training or certifications), professional/technical workers (requires a professional degree) and other. Participants were asked whether their current job was the same as their longest held job, and if so, how many years they worked at that job.

#### **CVD** risk factors

Participants were instructed to bring all medications (prescription and non-prescription) taken in the past month with them to the examination where they were scanned and documented. Medications were inventoried and classified using a Master Drug Data Base (Medispan MDDB). Three blood pressure measurements were performed using an automated sphygmomanometer (Omron model HEM-907 XL, Omron Healthcare, Bannockburn, Illinois, USA), with the average of the second and third readings calculated. Hypertension was defined as a systolic or diastolic blood pressure greater than or equal to 140/90 mm Hg, or the use of antihypertensive medications. Body weight

#### Selected characteristics in relation to occupational exposures among participants currently employed in HCHS/SOL (n=7404) Table 1 Current occupational exposure Solvents† Metals† Pesticides† Overall\* % P values % P values % P values Overall 6.5 8.5 4.7 \_ 0.01 < 0.01 Centre < 0.01 Bronx 25.2 4.9 7.6 6.5 18.9 Chicago 8.5 12.5 4.2 Miami 28.1 7.4 6.5 3.1 5.7 San Diego 27.9 8.5 5.0 0.09 < 0.01 0.12 Background Dominican 9.2 4.3 7.4 5.8 **Central American** 8.9 9.1 9.0 4.1 Cuban 17.3 7.6 7.3 2.8 Mexican 42.0 6.5 10.4 4.8 Puerto Rican 12.0 5.9 7.0 6.3 South American 6.0 5.5 4.3 3.8 Other/mixed 4.6 4.4 5.7 6.8 < 0.01 0.11 0.61 Age (years) 18-24 15.4 4.2 8.0 3.4 6.9 25-34 25.8 11.2 5.1 35-44 24.6 6.3 8.7 5.0 45-54 20.9 7.9 7.0 4.6 ≥55 13.3 6.4 5.7 4.8 < 0.01 < 0.01 < 0.01 Gender 45.1 2.9 3.5 2.8 Female Male 54.9 9.5 12.5 6.2 Residential duration in USA 0.58 0.87 0.03 29.6 6.1 <10 years 8.6 3.5 ≥10 years 70.4 6.6 8.4 5.2 < 0.01 < 0.01 0.38 **Employment status** Part time (≤35 hours/week) 34.2 4.5 6.0 4.2 Full time (>35 hours/week) 65.8 7.5 9.7 4.9 < 0.01 < 0.01 0.28 Primary occupation Non-skilled worker 27.8 7.4 10.3 4.9 Service worker 18.9 4.6 3.4 4.0 Skilled worker 22.6 9.7 12.1 5.1 Professional/technical 13.9 2.9 4.8 3.7 Other 16.8 6.4 8.0 6.0 Educational attainment 0.11 < 0.01 0.33 26.3 7.0 9.9 No high school diploma/GED‡ 5.6 High school diploma/GED 28.1 7.5 10.4 4.0 Greater than high school/GED 45.6 5.6 6.4 4.6 0.15 Current health insurance < 0.01 0.02 No 53.2 7.0 9.6 3.8 Yes 46.8 5.9 7.2 5.6 SASH language subscale 0.05 0.04 0.28 1-<3 (low) 21.9 6.9 9.0 4.9 ≥3 (high) 78.1 5.1 6.7 3.9 Physical activity level < 0.01 < 0.01 < 0.01 18.9 3.8 4.6 3.0 Low Medium 46.4 6.2 6.7 4.5 34.7 12.3 20.0 8.4 High Alternative healthy eating index < 0.01 0.03 0.09 ≤43.8 (low) 31.5 4.4 6.6 3.9 >43.8-50.2 (medium) 32.3 7.3 8.8 4.2

Continued

#### Table 1 Continued

		Current occupational exposure						
		Solvents†		Metals†		Pesticide	st	
	Overall*	%	P values	%	P values	%	P values	
>50.2 (high)	36.2	7.5		9.7		5.8		
Current alcohol intake			0.17		<0.01		<0.01	
None	42.0	5.6		6.0		3.3		
Low/moderate	51.3	7.1		9.9		5.6		
Heavy	6.6	7.4		13.2		6.5		
Smoking status			0.05		<0.01		0.05	
Never	64.2	5.8		7.5		4.2		
Former	17.6	7.1		8.6		4.2		
Current	18.2	8.3		11.7		6.9		
Hypertension			0.01		0.72		0.60	
No	83.1	6.0		8.5		4.6		
Yes	16.9	9.0		8.2		5.1		
Hypercholesterolaemia			<0.01		0.29		0.31	
No	60.8	5.6		8.1		4.3		
Yes	39.2	7.9		9.0		5.2		
Obese			0.25		0.31		0.42	
No	63.0	6.8		8.8		4.4		
Yes	37.0	5.9		7.9		5.1		
Diabetes mellitus			0.27		0.92		0.57	
No	89.9	6.4		8.5		4.6		
Yes	10.1	7.6		8.3		5.2		

\*The overall column shows column percentages for all 7404 currently employed HCHS/SOL participants.

The current occupational exposure columns show row percentages corresponding to currently employed HCHS/SOL participants who self-reported exposures to solvents, metals or pesticides, respectively.

GED, General Equivalency Diploma; HCHS/SOL, Hispanic Community Health Study/Study of Latinos; SASH, Short Acculturation Scale for Hispanics.

and height were measured by trained research technicians. Body mass index (BMI) was calculated as weight in kilograms divided by height in metres squared. Obesity was defined as a BMI of  $30.0 \text{ kg/m}^2$  or greater.

Total cholesterol, high-density lipoprotein (HDL) cholesterol, low-density lipoprotein (LDL) cholesterol, glucose and glycated haemoglobin were measured from fasting blood samples at the HCHS/SOL Central Laboratory. Laboratory assay methodologies are further described in the HCHS/SOL Manual.<sup>18</sup> Hypercholesterolaemia was defined as a total cholesterol  $\geq$ 240 mg/ dL, LDL cholesterol  $\geq$ 160 mg/dL, HDL cholesterol <40 mg/dL or antihyperlipidaemic medication use. Diabetes mellitus was defined as a fasting plasma glucose  $\geq$ 126 mg/dL, 2-hour postload plasma glucose  $\geq$ 200 mg/dL, glycated haemoglobin  $\geq$ 6.5% or antidiabetic medication use.

#### **Statistical analyses**

The prevalence of occupational exposures, CVD and CVD subtypes were calculated and compared across categorical variables using the Pearson  $\chi^2$  test. Poisson regression models were used to investigate the relationships between occupational exposures and CVD prevalence. Confounders were identified using a directed acyclic graph based on subject matter knowledge and the relevant epidemiological literature and were sequentially entered into models according to their hypothesised strengths of association with exposure status and CVD (online supplementary figure 1).<sup>19</sup> Adjusted model 1 included basic sociodemographic characteristics considered to be most strongly related to both exposure and CVD: age, gender, field centre and Hispanic/Latino background. Adjusted model 2 further adjusted for lifestyle and behavioural

factors that were additionally considered to confound exposure-CVD associations, although not as strongly sociodemographic characteristics. These included employment status, current health insurance status, cigarette use, alcohol use level, years of residential duration in the USA, physical activity level and the AHEI-2010 and the SASH language subscale. Adjusted model 3 included all variables in adjusted model 2 and additionally included hypertension, hypercholesterolaemia, BMI and diabetes mellitus as these were hypothesised to be potential intermediates on the causal pathways that could potentially explain exposure-CVD relationships. In all models, restricted cubic splines with four equally spaced knots at the 5th, 35th, 65th and 95th percentiles were used for continuous variables (age, years of residential duration in the USA, AHEI-2010, SASH language subscale and BMI) to allow for non-linearity. P values less than 0.05 were considered statistically significant. All analyses were conducted in Stata V.14.2. The Stata svy prefix was used to incorporate the HCHS/SOL sampling weights and account for the complex sampling design.

#### Sensitivity analyses

Since solvent, metal and pesticide exposures in the participants' current job may not accurately reflect chronic exposures, we performed a sensitivity analysis restricting to participants who indicated their current held job was their longest held. The models were identical to those used in our main analyses, with the addition of years worked as a covariate.

#### RESULTS

Participants who did not know their solvent or metal exposure status were excluded from analyses (275 for solvents; 133 for

	Cardiovascular (			Coronary heart		Cerebrovascular					
	disease*		diseaset		disease‡		Atrial fibrillation§		Heart failure¶		
Characteristic	%	P value	%	P value	%	P value	%	P value	%	P valu	
Overall	6.1		4.3		1.0		0.7		0.8		
Centre		<0.01		<0.01		<0.01		0.02		0.09	
Bronx	8.9		6.3		1.6		1.5		0.9		
Chicago	5.0		3.7		1.1		0.2		0.3		
Miami	6.1		4.2		1.0		0.6		0.7		
San Diego	4.4		3.0		0.3		0.3		1.3		
Background		<0.01		0.13		<0.01		<0.01		0.44	
Dominican	6.8		4.4		0.8		1.4		0.7		
Central American	5.5		3.6		1.6		0.3		0.5		
Cuban	6.6		5.1		0.9		0.7		0.8		
Mexican	5.0		3.7		0.6		0.2		0.9		
Puerto Rican	10.6		6.4		2.3		2.6		1.4		
South American	4.9		3.3		1.0		0.1		1.1		
Other/mixed	3.4		3.1		0.4		0.2		0.0		
Age (years)		<0.01		<0.01		<0.01		0.63		< 0.01	
18–24	4.0		2.8		0.4		0.6		0.1		
25–34	3.1		2.3		0.4		0.4		0.2		
35–44	5.7		4.1		0.8		0.7		1.1		
45–54	7.3		4.5		1.7		0.6		1.3		
≥55	13.2		10.1		2.0		1.4		1.8		
Gender		0.05		< 0.01		0.91		0.51		0.28	
Female	5.2		3.1		1.0		0.5		1.0		
Male	6.9		5.2		1.0		0.8		0.7		
Residential duration in USA (years)		0.12		0.07		0.58		0.93		0.79	
<10	5.1		3.4		0.8		0.6		0.8		
≥10	6.5		4.6		1.0		0.7		0.9		
Employment status		0.58		0.49		0.37		0.70		0.58	
Part time (≤35 hours/week)	5.8		3.9		1.2		0.6		0.9		
Full time (>35 hours/week)	6.3		4.5		0.9		0.7		0.8		
Primary occupation		0.85		0.27		0.85		0.14		0.07	
Non-skilled worker	5.9		3.9		1.0		1.2		0.6		
Service worker	5.9		4.5		0.9		0.1		1.1		
Skilled worker	6.0		3.9		1.1		0.6		1.1		
Professional/technical	5.8		3.4		0.9		0.9		1.4		
Other	7.1		6.0		0.7		0.4		0.2		
Educational attainment		0.18		0.3		0.32		0.20		0.85	
No high school diploma/GED	7.0		5.1		1.3		0.8		0.7		
High school diploma/GED	5.0		3.6		0.9		0.2		0.9		
Greater than high school/GED	6.3		4.3		0.8		0.9		0.9		
Current health insurance		<0.01		<0.01		0.45		<0.01		0.14	
No	4.6		3.2		0.9		0.2		0.6		
Yes	7.8		5.6		1.1		1.2		1.1		
SASH language subscale		0.84		0.27		0.24	_	0.03		0.40	
1-<3 (low)	6.2		4.5		0.9		0.5		0.9	55	
≥3 (high)	5.9		3.5		1.3		1.5		0.6		
Physical activity level	0.0	0.03	0.0	0.23		0.25	,	0.27		0.07	
Low	5.0		3.6	0.20	0.7	0.20	0.4	,	0.9	5.07	
Medium	7.2		4.8		1.2		0.9		1.0		
High	5.5		4.0		0.9		0.5		0.3		
Alternative healthy eating index	5.5	0.06	7.2	0.09	0.5	0.64	0.5	0.04	0.5	0.66	
≤43.8 (low)	6.5	0.00	4.3	0.03	1.0	0.04	1.3	0.04	0.9	0.00	
<ul><li>&lt;43.8 (IOW)</li><li>&gt;43.8–50.2 (medium)</li></ul>	6.5 4.7		4.5 3.3		0.8		0.5		0.9		
>50.2 (high)	4.7		5.2		1.1		0.5		0.7		

#### Table 2 Continued

Characteristic		Cardiovascular disease*		Coronary heart disease†		Cerebrovascular disease‡		Atrial fibrillation§		Heart failure¶	
	%	P value	%	P value	%	P value	%	P value	%	P value	
Current alcohol intake		0.02		0.09		<0.01		0.01		0.28	
None	6.6		4.3		1.4		1.0		1.1		
Low/moderate	5.2		3.9		0.7		0.3		0.7		
Heavy	10.2		7.4		0.3		2.1		0.4		
Smoking status		0.25		0.06		0.13		0.50		0.90	
Never	5.7		3.6		0.9		0.8		0.8		
Former	6.2		5.0		0.7		0.4		1.0		
Current	7.5		5.9		1.4		0.6		0.8		
Hypertension		<0.01		<0.01		<0.01		<0.01		<0.01	
No	4.6		3.3		0.7		0.5		0.6		
Yes	13.4		9.2		2.6		1.7		2.1		
Hypercholesterolaemia		<0.01		<0.01		0.05		0.95		0.09	
No	5.1		3.2		0.8		0.7		0.7		
Yes	7.7		6.0		1.3		0.7		1.1		
Obese		<0.01		<0.01		0.08		0.79		0.48	
No	5.2		3.3		0.8		0.7		0.8		
Yes	7.7		5.9		1.3		0.6		1.0		
Diabetes mellitus		<0.01		<0.01		<0.01		0.09		<0.01	
No	5.3		3.7		0.8		0.6		0.6		
Yes	13.7		9.7		2.3		1.6		3.1		

\*Cardiovascular disease was defined as at least one of the following: coronary heart disease, cerebrovascular disease, heart failure or atrial fibrillation; data are expressed as row percentages.

+Coronary heart disease was defined as self-reported angina, myocardial infarction, coronary bypass surgery, balloon angioplasty or stent placement in coronary arteries, or ECG evidence of major Q wave abnormalities or minor Q, QS waves with ST and T abnormalities; data are expressed as row percentages.

\*Cerebrovascular disease was defined as self-reported stroke or transient ischaemic attack; data are expressed as row percentages.

§Heart failure was defined as self-reported heart failure; data are expressed as row percentages.

¶Atrial fibrillation was defined as self-reported atrial fibrillation or ECG evidence of atrial fibrillation or flutter; data are expressed as row percentages.

GED, General Equivalency Diploma; HCHS/SOL, Hispanic Community Health Study/Study of Latinos; SASH, Short Acculturation Scale for Hispanics.

metals); however, they did not differ substantially from those who provided a definitive response in terms of Hispanic/Latino background, age, gender, educational attainment, language acculturation, nativity or CVD status (online supplementary table 1). After excluding these participants and those with incomplete data, our analytic sample included 7404 (91.1%) participants of the 8156 employed at baseline. Occupational exposures and participant characteristics are displayed in table 1. The most commonly reported exposure was to metals (8.5%), followed by solvents (6.5%) and pesticides (4.7%). Male gender, higher physical activity levels and current smoking were consistently associated with a higher probability of reporting working with chemical hazards. Overall, 6.1% of participants had CVD, of which the majority of cases were coronary heart disease (table 2). CVD was more common among Bronx field centre, Puerto Rican background, older age, insured, heavy alcohol consumption, hypertensive, hypercholesterolaemic, obese and diabetic participants.

In crude regression models, the prevalence of any CVD was more than twice (prevalence ratio (PR): 2.51, 95% CI 1.35 to 4.69) as high among participants who reported exposures to pesticides in their current job(s) compared with participants who did not (table 3). This association persisted after adjustment for confounders (adjusted model 2; PR: 2.18, 95% CI 1.34 to 3.55). The corresponding adjusted PRs for pesticide exposure were 2.20 (95% CI 1.31 to 3.71) for coronary heart disease, 1.38 (95% CI 0.62 to 3.03) for cerebrovascular disease, 5.92 (95% CI 1.89 to 18.61) for atrial fibrillation and 0.91 (95% CI 0.23 to 3.54) for heart failure. The adjusted PR for self-reported metal exposures in the primary job and any CVD from model 2 was 1.24 (95% CI 0.80 to 1.94). Metal exposures were significantly associated with atrial fibrillation (adjusted model 2; PR: 3.78, 95% CI 1.24 to 11.46), but not coronary heart disease, cerebrovascular disease or heart failure. Associations for self-reported solvent exposure in the primary job were largely null. Including hypertension, hypercholesterolaemia, BMI and diabetes mellitus (ie, adjusted model 3) in our models did not appreciably change the results (table 3; figures 1–3).

A total of 5080 participants were currently working at their longest held job, with the median duration of employment in that position being 10 years (IQR: 5–20 years). In sensitivity analyses of these individuals, the magnitude of the observed associations for metal and pesticide exposures was increased. For example, in our main analyses, self-reported metal exposure was associated with an approximately fourfold greater prevalence of atrial fibrillation (table 3), whereas for those whose current job was their longest held, the relative association was almost fivefold (table 4). Again, we observed no appreciable attenuation after adjusting for hypertension, hypercholesterolaemia, BMI or diabetes mellitus.

#### DISCUSSION

Between 4.7% and 8.5% of Hispanic/Latino workers in urban areas of the US report being exposed to organic solvents, metals or pesticides at their jobs. Occupational exposures to pesticides,

	Crude model	Adjusted model 1	Adjusted model 2	Adjusted model 3	
	PR (95% CI)*	PR (95% CI)†	PR (95% CI)‡	PR (95% CI)§	
Cardiovascular disease					
Solvents	1.03 (0.64 to 1.66)	0.93 (0.58 to 1.51)	0.94 (0.58 to 1.52)	0.92 (0.58 to 1.47)	
Metals	1.21 (0.80 to 1.83)	1.26 (0.81 to 1.95)	1.24 (0.80 to 1.93)	1.22 (0.79 to 1.89)	
Pesticides	2.51 (1.35 to 4.69)	2.25 (1.26 to 4.01)	2.18 (1.34 to 3.55)	2.18 (1.34 to 3.56)	
Coronary heart disease					
Solvents	1.12 (0.67 to 1.86)	0.97 (0.59 to 1.60)	0.96 (0.59 to 1.57)	0.96 (0.59 to 1.57)	
Metals	1.05 (0.65 to 1.69)	1.01 (0.61 to 1.67)	0.97 (0.59 to 1.59)	0.98 (0.60 to 1.59)	
Pesticides	2.85 (1.30 to 6.26)	2.47 (1.24 to 4.90)	2.20 (1.31 to 3.71)	2.24 (1.29 to 3.90)	
Cerebrovascular disease					
Solvents	0.70 (0.25 to 1.97)	0.59 (0.20 to 1.70)	0.59 (0.20 to 1.76)	0.60 (0.22 to 1.64)	
Metals	0.59 (0.24 to 1.42)	0.65 (0.25 to 1.67)	0.67 (0.26 to 1.76)	0.64 (0.25 to 1.65)	
Pesticides	1.48 (0.67 to 3.24)	1.33 (0.58 to 3.04)	1.38 (0.62 to 3.03)	1.42 (0.64 to 3.14)	
Atrial fibrillation					
Solvents	1.34 (0.20 to 8.97)	1.46 (0.21 to 10.23)	1.62 (0.32 to 8.31)	1.25 (0.28 to 5.47)	
Metals	2.45 (0.73 to 8.27)	2.88 (0.81 to 10.29)	3.78 (1.24 to 11.46)	3.93 (1.41 to 11.00)	
Pesticides	4.00 (1.10 to 14.51)	3.52 (0.80 to 15.46)	5.92 (1.89 to 18.61)	5.61 (1.97 to 16.03)	
Heart failure					
Solvents	0.36 (0.10 to 1.30)	0.38 (0.10 to 1.41)	0.40 (0.11 to 1.48)	0.41 (0.11 to 1.51)	
Metals	0.91 (0.18 to 4.49)	1.27 (0.22 to 7.15)	1.53 (0.27 to 8.69)	1.56 (0.28 to 8.64)	
Pesticides	0.85 (0.22 to 3.30)	0.86 (0.23 to 3.27)	0.91 (0.23 to 3.54)	0.92 (0.23 to 3.70)	

\*Prevalence ratio (PR) with 95% CI comparing self-reported exposure to the respective agent to no exposure.

+PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for age (restricted cubic splines with four knots), gender (male/ female), field centre (Bronx, Chicago, Miami or San Diego) and Hispanic/Latino background (Dominican, Central American, Cuban, Mexican, Puerto Rican, South American or other/mixed).

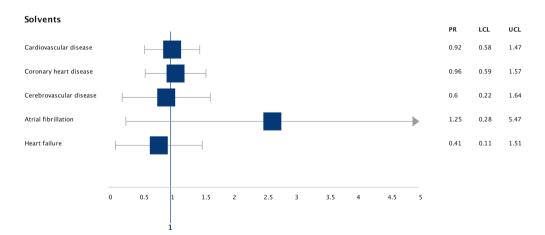
\*PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for the variables listed in † with additional adjustments for current health insurance status (yes/no), cigarette use (current, former or never), alcohol use level (none, low/moderate or high) years of residential duration in the USA (restricted cubic splines with four knots), employment status (full time or part time), physical activity level (low, medium or high), alternative healthy eating index (restricted cubic splines with four knots) and the Short Acculturation Scale for Hispanics – language subscale (restricted cubic splines with four knots).

§PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for the variables listed in † with additional adjustments for hypertension (yes/no), hypercholesterolaemia (yes/no), body mass index (restricted cubic splines with four knots) and diabetes mellitus (yes/no).

and to a lesser degree metals, were positively associated with CVD, while no associations were observed for organic solvents. Given the stronger results found when restricting analyses to participants whose current job was their longest held, it appears that cumulative exposures are more harmful. Prior work within the HCHS/SOL cohort suggests workplace metals and pesticide exposures are unrelated to cardiometabolic conditions.<sup>5</sup> The lack

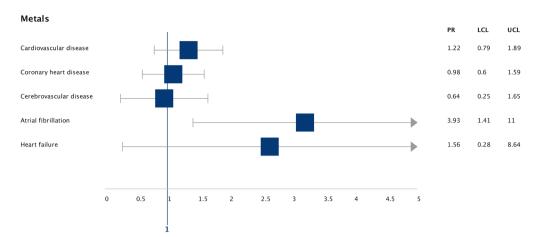
of attenuation by hypertension, hypercholesterolaemia, diabetes and BMI in the present analysis further supports this hypothesis and suggests that alternative biological mechanisms may play a role in the development of metal-association and pesticide-associated CVD.

Little is known about the effects of chronic, low-level pesticide and metal exposures on the development of CVD. Of



**Figure 1** PRs with 95% CI for current occupational exposure to solvents and cardiovascular disease after adjustment for age, gender, field centre, Hispanic/Latino background, current health insurance status, cigarette use, alcohol use level, years of residential duration in the USA, employment status, physical activity level, alternative healthy eating index, Short Acculturation Scale for Hispanics – language subscale, hypertension, hypercholesterolaemia, body mass index and diabetes mellitus. LCL, lower confidence limit; PR, prevalence ratio; UCL, upper confidence limit.

#### Cardiac risk factors and prevention

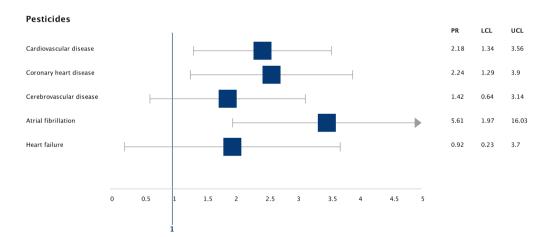


**Figure 2** PRs with 95% CI for current occupational exposure to metals and cardiovascular disease after adjustment for age, gender, field centre, Hispanic/Latino background, current health insurance status, cigarette use, alcohol use level, years of residential duration in the USA, employment status, physical activity level, alternative healthy eating index, Short Acculturation Scale for Hispanics – language subscale, hypertension, hypercholesterolaemia, body mass index and diabetes mellitus. LCL, lower confidence limit; PR, prevalence ratio; UCL, upper confidence limit.

the CVD subtypes, the strongest cross-sectional relationships with pesticide use were observed for coronary heart disease and atrial fibrillation. The underlying mechanisms of pesticide cardiotoxicity relevant for such diseases include degeneration of myocardial fibres, modification of sodium channels and reduced intracellular uptake of calcium, all of which have been observed in animal studies.<sup>20-22</sup> For metals, epidemiological studies have produced suggestive evidence of a causal relationship with CVD, but the number of studies is small.<sup>23 24</sup> In our analyses, the only CVD subtype significantly associated with occupational metal exposures was atrial fibrillation. This is a novel finding and is biologically plausible since metals can induce both systemic (eg, oxidative stress and inflammation) and direct effects on the heart and vasculature (eg, impaired left ventricular function and degeneration of arterial compliance).<sup>23</sup> While there is some evidence that exposure to certain organic solvents adversely affects cardiac health and function, we did not observe any significant associations in our analyses.<sup>156</sup>

Though our results are intriguing, this work is not without limitations. We relied on self-reported information about

exposures from questionnaires, which may have been misreported. The questionnaires used recognised strategies for accurate responses by prompting participants with recognisable terms (eg, 'metals') that did not allow for open-ended responses.<sup>25</sup> We dichotomised exposure as yes/no because the data regarding frequency of exposure were sparse. Thus, we were unable to conduct dose-response analyses. One potential benefit of collapsing exposure status is that we may have limited exposure misclassification as the reliability of self-reported frequency of exposures tends to be lower than the reliability of self-reported exposure status itself.<sup>26</sup> Ultimately, we have no reason to believe that any exposure misclassification would be associated with CVD status. Some of the CVD endpoints used in this study were also ascertained via self-report; however, many have been shown to have high validity and were combined with electrocardiographic evidence.<sup>27-29</sup> Nevertheless, there remains the potential for recall bias and/or reverse causality if those with existing disease were more likely to report their occupational exposures or if diseased individuals were more



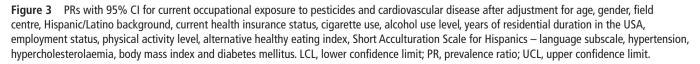


 Table 4
 Associations between occupational exposures and cardiovascular disease and subtypes among participants whose current job was their longest held (n=5080)

	Crude model	Adjusted model 1	Adjusted model 2	Adjusted model 3	
	PR (95% CI)*	PR (95% CI)†	PR (95% CI)‡	PR (95% CI)§	
Cardiovascular disease					
Solvents	1.18 (0.67 to 2.07)	1.14 (0.66 to 1.97)	1.20 (0.70 to 2.08)	1.16 (0.69 to 1.98)	
Metals	1.38 (0.85 to 2.23)	1.44 (0.85 to 2.43)	1.43 (0.85 to 2.38)	1.44 (0.88 to 2.35)	
Pesticides	3.36 (1.71 to 6.60)	2.80 (1.55 to 5.05)	2.61 (1.64 to 4.14)	2.59 (1.67 to 4.03)	
Coronary heart disease					
Solvents	1.32 (0.72 to 2.44)	1.19 (0.67 to 2.11)	1.20 (0.69 to 2.08)	1.17 (0.68 to 2.02)	
Metals	1.21 (0.69 to 2.13)	1.12 (0.61 to 2.06)	1.09 (0.63 to 1.88)	1.13 (0.66 to 1.94)	
Pesticides	4.25 (1.82 to 9.90)	3.20 (1.73 to 5.91)	2.44 (1.61 to 3.70)	2.47 (1.64 to 3.73)	
Cerebrovascular disease					
Solvents	0.90 (0.29 to 2.80)	0.84 (0.26 to 2.77)	1.03 (0.35 to 3.06)	1.07 (0.37 to 3.09)	
Metals	0.62 (0.22 to 1.76)	0.72 (0.23 to 2.24)	0.78 (0.26 to 2.34)	0.82 (0.30 to 2.25)	
Pesticides	1.32 (0.48 to 3.66)	1.25 (0.44 to 3.56)	1.38 (0.51 to 3.75)	1.46 (0.52 to 4.06)	
Atrial fibrillation					
Solvents	1.37 (0.20 to 9.28)	1.77 (0.26 to 12.02)	2.98 (0.87 to 10.26)	2.41 (0.98 to 5.95)	
Metals	2.42 (0.69 to 8.51)	3.38 (1.04 to 11.02)	4.89 (2.01 to 11.95)	5.15 (2.15 to 12.38)	
Pesticides	4.36 (1.17 to 16.21)	3.73 (0.80 to 17.48)	8.21 (3.21 to 21.02)	7.34 (3.17 to 16.96)	
Heart failure					
Solvents	0.30 (0.06 to 1.45)	0.33 (0.07 to 1.64)	0.37 (0.08 to 1.79)	0.36 (0.07 to 1.82)	
Metals	0.83 (0.13 to 5.41)	1.11 (0.15 to 8.08)	1.29 (0.18 to 9.11)	1.29 (0.25 to 6.65)	
Pesticides	0.71 (0.13 to 3.91)	0.80 (0.15 to 4.30)	0.84 (0.15 to 4.76)	0.82 (0.13 to 5.04)	

\*Prevalence ratio (PR) with 95% CI comparing self-reported exposure to the respective agent with no exposure.

†PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for age (restricted cubic splines with four knots), gender (male/ female), field centre (Bronx, Chicago, Miami or San Diego) and Hispanic/Latino background (Dominican, Central American, Cuban, Mexican, Puerto Rican, South American or other/mixed).

\*PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for the variables listed in † with additional adjustments for current health insurance status (yes/no), cigarette use (current, former or never), alcohol use level (none, low/moderate or high) years of residential duration in the USA (restricted cubic splines with four knots), employment status (full time or part time), physical activity level (low, medium or high), alternative healthy eating index (restricted cubic splines with four knots) and the Short Acculturation Scale for Hispanics – language subscale (restricted cubic splines with four knots).

§PR with 95% CI comparing self-reported exposure to the respective agent to no exposure with adjustment for the variables listed in † with additional adjustments for hypertension (yes/no), hypercholesterolaemia (yes/no), body mass index (restricted cubic splines with four knots) and diabetes mellitus (yes/no).

willing to work with hazardous substances. Because we did not collect agent-specific information, we are unable to attribute the observed associations to individual metals or pesticides. Lastly, our questionnaires only queried participants with regards to their chemical exposures at work, so we were unable to assess the contribution of any exposures from leisure-time activities.

As a community-based study, this work was the first to evaluate the role of workplace chemical exposures in CVD among urban Hispanic/Latino workers. Though relatively rare in this population, metal and pesticide exposures in the workplace were cross-sectionally associated with marked elevations in the prevalence of CVD. These results should be considered preliminary and interpreted with caution given the limitations of our study design; namely, our reliance on self-reported exposure status and cross-sectional data. However, our results do suggest occupational exposures could be important contributors to CVD for Hispanics/ Latinos. The findings from the present analysis are consistent with prior studies within the HCHS/SOL cohort and highlight the substantial burden of chronic disease risk factors and CVD in the US Hispanic/Latino population.<sup>30</sup> A major discovery is that US Hispanics/Latinos are not a homogenous group; there exists substantial heterogeneity by Hispanic/Latino background and gender.<sup>30</sup> Notably, Puerto Rican individuals and men were significantly more likely to suffer from CVD than their peers. In our study, the proportion of Puerto Ricans reporting workplace exposures was

on par with most of the other background groups; however, men reported exposures 2–3 fold more than women. Since workplace chemical hazards are known risk factors for other

#### Key messages

#### What is already known on this subject?

Some industrially used chemicals are known cardiotoxins. Both acute and long-term exposures can result in damages to the cardiovascular system.

#### What might this study add?

- Here, we estimate the proportion of exposed US Hispanic/ Latino workers, a large and growing minority population that may especially vulnerable and quantify associations with prevalent cardiovascular disease. Our findings suggest between 5% and 9% of employed Hispanics/Latinos are exposed to solvents, metals or pesticides in the workplace.
- Occupational exposures to metals or pesticides, in particular, were associated with an elevated prevalence of coronary heart disease and atrial fibrillation.

#### How might this impact on clinical practice?

Clinicians should consider the value of taking an occupational history during medical examinations, as doing so could assist in identifying workplace hazards that need to be controlled.

#### Cardiac risk factors and prevention

serious illnesses, including cancers, respiratory and neurological conditions, efforts should be undertaken to minimise or eliminate exposures. Such workplace interventions may want to consider first targeting male workers, especially Puerto Rican males given their double burden of occupational exposures and existing CVD.

#### Author affiliations

<sup>1</sup>Division of Epidemiology and Biostatistics, School of Public Health, University of Illinois at Chicago, Chicago, Illinois, USA

<sup>2</sup>Division of Academic Internal Medicine, Department of Medicine, Institute for Minority Health Research, University of Illinois at Chicago, Chicago, Illinois, USA <sup>3</sup>Division of Nephrology, Department of Medicine, College of Medicine, University of Illinois at Chicago, Chicago, Illinois, USA

<sup>4</sup>Department of Public Health Sciences, Miller School of Medicine, University of Miami, Miami, Florida, USA

<sup>5</sup>Department of Neurology, Miller School of Medicine, University of Miami, Miami, Florida, USA

<sup>6</sup>Department of Healthcare Sciences, Wayne State University, Detroit, Michigan, USA

**Acknowledgements** The authors would like to thank the staff and participants of Hispanic Community Health Study/Study of Latinos (HCHS/SOL) for their important contributions. Investigators website: http://www.cscc.unc.edu/hchs/.

**Contributors** CMB performed all statistical analyses and wrote the manuscript. MLD, VWP, RAD-A and MA were integrally involved in the study design and manuscript preparation. JPL, TE, DJL, ARR and WT each critically revised the manuscript with regards to data interpretation.

**Funding** CMB was supported by the National Heart, Lung, and Blood Institute (NHLBI) T32-HL125294. The Hispanic Community Health Study/Study of Latinos was carried out as a collaborative study supported by contracts from the NHLBI to the University of North Carolina (N01-HC65233), University of Miami (N01-HC65234), Albert Einstein College of Medicine (N01-HC65235), University of Illinois at Chicago (HHSN268201300003I), Northwestern University (N01-HC65236) and San Diego State University (N01-HC65237). The following institutes/centres/offices contribute to the HCHS/SOL through a transfer of funds to the NHLBI: National Institute on Minority Health and Health Disparities, National Institute on Deafness and Other Communication Disorders, National Institute of Deatal and Craniofacial Research, National Institute of Diabetes and Stroke and NIH Institution-Office of Dietary Supplements.

**Disclaimer** The views expressed in this manuscript are those of the authors and do not necessarily represent the views of the National Heart, Lung, and Blood Institute; the National Institutes of Health; or the U.S. Department of Health and Human Services.

Competing interests None declared.

Patient consent Not required.

**Ethics approval** University of Illinois at Chicago IRB; University of Miami IRB; San Diego State University IRB; Albert Einstein College of Medicine IRB; University of North Carolina at Chapel Hill IRB.

Provenance and peer review Not commissioned; externally peer reviewed.

**Data sharing statement** Data from HCHS/SOL can be accessed by submitting proposals for manuscripts through the HCHS/SOL website http://www2.cscc.unc. edu/hchs/.

**Open access** This is an open access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited, appropriate credit is given, any changes made indicated, and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/.

#### REFERENCES

- Attarchi M, Golabadi M, Labbafinejad Y, et al. Combined effects of exposure to occupational noise and mixed organic solvents on blood pressure in car manufacturing company workers. Am J Ind Med 2013;56:243–51.
- 2 Li H, Hedmer M, Kåredal M, et al. A cross-sectional study of the cardiovascular effects of welding fumes. PLoS One 2015;10:e0131648.
- 3 Wong JY, Fang SC, Grashow R, *et al*. The relationship between occupational metal exposure and arterial compliance. *J Occup Environ Med* 2015;57:355–60.

- 4 Fang SC, Eisen EA, Cavallari JM, et al. Acute changes in vascular function among welders exposed to metal-rich particulate matter. *Epidemiology* 2008;19:217–25.
- 5 Bulka CM, Daviglus ML, Persky VW, et al. Occupational Exposures and Metabolic Syndrome Among Hispanics/Latinos: Cross-Sectional Results From the Hispanic Community Health Study/Study of Latinos (HCHS/SOL). J Occup Environ Med 2017;59:1047–55.
- 6 Blair A, Hartge P, Stewart PA, et al. Mortality and cancer incidence of aircraft maintenance workers exposed to trichloroethylene and other organic solvents and chemicals: extended follow up. Occup Environ Med 1998;55:161–71.
- 7 Boffetta P, Sällsten G, Garcia-Gómez M, *et al*. Mortality from cardiovascular diseases and exposure to inorganic mercury. *Occup Environ Med* 2001;58:461–6.
- 8 Kazantzis G, Lam TH, Sullivan KR. Mortality of cadmium-exposed workers. A five-year update. Scand J Work Environ Health 1988;14:220–3.
- 9 Cantor KP, Silberman W. Mortality among aerial pesticide applicators and flight instructors: follow-up from 1965-1988. Am J Ind Med 1999;36:239–47.
- Arrighi HM, Hertz-Picciotto I. The evolving concept of the healthy worker survivor effect. *Epidemiology* 1994;5:189–96.
- 11 Tielemans E, Heederik D, Burdorf A, et al. Assessment of occupational exposures in a general population: comparison of different methods. Occup Environ Med 1999;56:145–51.
- 12 Lavange LM, Kalsbeek WD, Sorlie PD, et al. Sample design and cohort selection in the Hispanic Community Health Study/Study of Latinos. Ann Epidemiol 2010;20:642–9.
- 13 Sorlie PD, Avilés-Santa LM, Wassertheil-Smoller S, et al. Design and implementation of the Hispanic Community Health Study/Study of Latinos. Ann Epidemiol 2010;20:629–41.
- 14 Denes P, Garside DB, Lloyd-Jones D, *et al*. Major and minor electrocardiographic abnormalities and their association with underlying cardiovascular disease and risk factors in Hispanics/Latinos (from the Hispanic Community Health Study/Study of Latinos). *Am J Cardiol* 2013;112:1667–75.
- 15 World Health Organization, 2012. Global physical activity questionnaire (GPAQ) analysis guide http://www.who.int/chp/steps/resources/GPAQ\_Analysis\_Guide.pdf? ua=1 (cited 2016 Sep 27).
- 16 Chiuve SE, Fung TT, Rimm EB, et al. Alternative dietary indices both strongly predict risk of chronic disease. J Nutr 2012;142:1009–18.
- 17 Marin G, Sabogal F, Marin BV, et al. Development of a short acculturation scale for Hispanics. Hisp J Behav Sci 1987;9:183–205.
- 18 Study of Latinos/Hispanic Community Health Study, 2011. Addendum Central Laboratory Procedures https://www2.cscc.unc.edu/hchs/system/files/protocolsmanuals/UNLICOMMManual07AddendumCentralLaboratoryProceduresv1006 222011.pdf (cited 8 Aug 2016).
- 19 Greenland S, Pearl J, Robins JM. Causal diagrams for epidemiologic research. *Epidemiology* 1999;10:37–48.
- 20 Curic S, Gojmerac T, Zuric M. Morphological changes in the organs of gilts induced with low-dose atrazine. *Veterinarski Arhiv* 1999;69:135–48.
- 21 Spencer CI, Yuill KH, Borg JJ, et al. Actions of pyrethroid insecticides on sodium currents, action potentials, and contractile rhythm in isolated mammalian ventricular myocytes and perfused hearts. J Pharmacol Exp Ther 2001;298:1067–82.
- 22 Gress S, Lemoine S, Puddu PE, *et al.* Cardiotoxic Electrophysiological Effects of the Herbicide Roundup(®) in Rat and Rabbit Ventricular Myocardium In Vitro. *Cardiovasc Toxicol* 2015;15:324–35.
- 23 Navas-Acien A, Guallar E, Silbergeld EK, et al. Lead exposure and cardiovascular disease–a systematic review. Environ Health Perspect 2007;115:472–82.
- 24 Navas-Acien A, Sharrett AR, Silbergeld EK, et al. Arsenic exposure and cardiovascular disease: a systematic review of the epidemiologic evidence. Am J Epidemiol 2005;162:1037–49.
- 25 Teschke K, Olshan AF, Daniels JL, et al. Occupational exposure assessment in casecontrol studies: opportunities for improvement. Occup Environ Med 2002;59:575–94. discussion 594.
- 26 Blair A, Tarone R, Sandler D, *et al*. Reliability of reporting on life-style and agricultural factors by a sample of participants in the Agricultural Health Study from Iowa. *Epidemiology* 2002;13:94–9.
- 27 Okura Y, Urban LH, Mahoney DW, et al. Agreement between self-report questionnaires and medical record data was substantial for diabetes, hypertension, myocardial infarction and stroke but not for heart failure. J Clin Epidemiol 2004;57:1096–103.
- 28 Soliman EZ, Howard G, Meschia JF, et al. Self-reported atrial fibrillation and risk of stroke in the Reasons for Geographic and Racial Differences in Stroke (REGARDS) study. Stroke 2011;42:2950–3.
- 29 Lampe FC, Walker M, Lennon LT, et al. Validity of a self-reported history of doctordiagnosed angina. J Clin Epidemiol 1999;52:73–81.
- 30 Daviglus ML, Talavera GA, Avilés-Santa ML, et al. Prevalence of major cardiovascular risk factors and cardiovascular diseases among Hispanic/Latino individuals of diverse backgrounds in the United States. JAMA 2012;308:1775–84.