STROBE Statement—checklist of items that should be included in reports of observational studies

	Item No.	Recommendation		Page No.	Relevant text from manuscript
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	2		Lines 25-31; see also Figure 1
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found	2		Lines 25-44
Introduction					
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	5-6		
Objectives	3	State specific objectives, including any prespecified hypotheses	6		Lines 111-117
Methods					
Study design	4	Present key elements of study design early in the paper	8		Lines 147-158; see also Figure
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	7-8		Lines 119-142; lines 156-179
Participants	6	<ul> <li>(a) Cohort study—Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up</li> <li>Case-control study—Give the eligibility criteria, and the sources and methods of case ascertainment and control selection. Give the rationale for the choice of cases and controls</li> <li>Cross-sectional study—Give the eligibility criteria, and the sources and methods of selection of participants</li> <li>(b) Cohort study—For matched studies, give matching criteria and number of exposed and unexposed</li> <li>Case-control study—For matched studies, give matching criteria and the number of controls per case</li> </ul>	8		Lines 156-179
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable	13		Lines 284-295
Data sources/	8*	For each variable of interest, give sources of data and details of methods of assessment	9-12		
measurement		(measurement). Describe comparability of assessment methods if there is more than one group			
Bias	9	Describe any efforts to address potential sources of bias	13-14		
Study size	10	Explain how the study size was arrived at			Exploratory study; power not formally predefined

Quantitative	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which	13-14	
variables		groupings were chosen and why		
Statistical	12	( <i>a</i> ) Describe all statistical methods, including those used to control for confounding	13-14	
methods		(b) Describe any methods used to examine subgroups and interactions	13-14	
		(c) Explain how missing data were addressed	14	
		(d) Cohort study—If applicable, explain how loss to follow-up was addressed	8	
		Case-control study-If applicable, explain how matching of cases and controls was addressed		
		Cross-sectional study—If applicable, describe analytical methods taking account of sampling		
		strategy		
		( <u>e</u> ) Describe any sensitivity analyses		None.
Results				
Participants	13*	(a) Report numbers of individuals at each stage of study-eg numbers potentially eligible, examined	8; 14	
		for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed		
		(b) Give reasons for non-participation at each stage	8	
		(c) Consider use of a flow diagram		Not used in this study.
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and information on	14	
		exposures and potential confounders		
		(b) Indicate number of participants with missing data for each variable of interest	13-14	Not indicated, but total of cases
				analysed clearly stated (see also
				Tables 1-3).
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)		
Outcome data	15*	Cohort study-Report numbers of outcome events or summary measures over time		
		Case-control study-Report numbers in each exposure category, or summary measures of exposure	13-14	
		Cross-sectional study-Report numbers of outcome events or summary measures		
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and their precision	Tables 1-3	
		(eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were		
		included		
		(b) Report category boundaries when continuous variables were categorized		
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time		
		period		

Other analyses	17	Report other analyses done-eg analyses of subgroups and interactions, and sensitivity analyses	None.
Discussion			
Key results	18	Summarise key results with reference to study objectives	22-27
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss	22-27
		both direction and magnitude of any potential bias	
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of	22-27
		analyses, results from similar studies, and other relevant evidence	
Generalisability	21	Discuss the generalisability (external validity) of the study results	22-27
Other informati	ion		
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the	Information provided
		original study on which the present article is based	

\*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

**Note:** An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.