Title and abotes 4	No	Recommendation	
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract	
		[Within the Title Page, page I and Method section of the Abstract] (b) Provide in the abstract an informative and balanced summary of what was done	
		and what was found [See Results Section of the Abstract]	
Introduction			
Background/rational	le 2	Explain the scientific background and rationale for the investigation being reported [See Introduction Section of the Manuscript file, page 2]	
Objectives	3	State specific objectives, including any prespecified hypotheses	
M-41-1-		[See Introduction Section of the Manuscript file, page 2]	
Methods Study design	4	Present key elements of study design early in the paper	
		[See Methods Section of the Manuscript file, page 2-6]	
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection	
Participants	6	[N/A] (a) 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 [See Methods Section of the Manuscript file, Pre-treatment Behavioral Appraisal page 4, Post-treatment Behavioral Follow-Up page 5] (b) For matched studies, give matching criteria and the number of controls per case	
		[N/A]	
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable [See Methods Section of the Manuscript file, Pre-treatment Behavioral	
		Appraisal page 4, Post-treatment Behavioral Follow-Up page 5]	
Data sources/	8*	For each variable of interest, give sources of data and details of methods of	
measurement		assessment (measurement). Describe comparability of assessment methods if there i more than one group	
Bias	9	[N/A] Describe any efforts to address potential sources of bias	
		[See Discussion Section of the Manuscript file, page 8]	
Study size	10	Explain how the study size was arrived at [Need for reduction to a minimum as possible (20 rats) the number of animal	
		employed for ethical purposes, 3R rules: Reduction, Refinement, Replacement	
Quantitative variable	es 11	Explain how quantitative variables were handled in the analyses. If applicable,	
		describe which groupings were chosen and why [N/A]	
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding	
		[See Method Section of the Manuscript file, Statistical Analysis, page 7]	
		(b) Describe any methods used to examine subgroups and interactions [N/A]	
		(c) Explain how missing data were addressed	
		[N/A]	
		(d) If applicable, explain how matching of cases and controls was addressed [N/A]	
		(e) Describe any sensitivity analyses	
		[N/A]	
Results	13*	(A) Provident Control of Control	
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially eligible, examined for eligibility, confirmed eligible, included in the study, completing follow-up, and analysed [See Methods Section of the Manuscript file, Pre-treatment Behavioral	
		Appraisal page 4] (b) Give reasons for non-participation at each stage	
		(b) Orve reasons for non-participation at each stage [N/A] (c) Consider use of a flow diagram	
		[See Fig. 1]	
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and	
		information on exposures and potential confounders [See Methods Section of the Manuscript file, Pre-treatment Behavioral	
		Appraisal page 4] (b) Indicate number of participants with missing data for each variable of interest	
Outcome data	15*	[N/A]	
oncome data	15*	Report numbers in each exposure category, or summary measures of exposure	
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and	
		their precision (eg, 95% confidence interval). Make clear which confounders were adjusted for and why they were included	
		[See Method Section of the Manuscript file, Statistical Analysis, page 7]	
		(b) Report category boundaries when continuous variables were categorized	
		[N/A] (c) If relevant, consider translating estimates of relative risk into absolute risk for a	
		meaningful time period	
Other analyses 1	7 Report of	[N/A] her analyses done—eg analyses of subgroups and interactions, and sensitivity analyse	
Discussion	[
Key results 1		se key results with reference to study objectives	
Limitations 1		ussion Section of the Manuscript file, page 8-9] mitations of the study, taking into account sources of potential bias or imprecision.	
Zaramuono I	Discuss b	Discuss immutations of the study, taking into account sources of potential ones or imprecision. Discuss both direction and magnitude of any potential bias [See Discussion Section of the Manuscript file, page 8-9]	
Interpretation	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity		
	of analyses, results from similar studies, and other relevant evidence		
Generalisability 2		ussion Section of the Manuscript file, page 8-9] ne generalisability (external validity) of the study results	
		ussion Section of the Manuscript file, page 8-9]	
Other information			
Funding 2		ource of funding and the role of the funders for the present study and, if applicable, ginal study on which the present article is based	
*Give information s	eparately for	cases and controls.	
		ation article discusses each checklist item and gives methodological background and t reporting. The STROBE checklist is best used in conjunction with this article (freely	

Supplemental Digital Content 1: STROBE Statement—Checklist of items that should be included in reports of case-control studies