<u>Materials Design Analysis Reporting (MDAR)</u> Checklist for Authors

The MDAR framework establishes a minimum set of requirements in transparent reporting applicable to studies in the life sciences (see Statement of Task: doi:10.31222/osf.io/9sm4x.). The MDAR checklist is a tool for authors, editors and others seeking to adopt the MDAR framework for transparent reporting in manuscripts and other outputs. Please refer to the MDAR Elaboration Document for additional context for the MDAR framework.

accession number if available, and source

Materials

| Antibodies | Yes (indicate where provided: section/paragraph) | |
|--|--|-----|
| For commercial reagents, provide supplier name, catalogue number and RRID, if available. | Not involved in this article. | |
| Cell materials | Yes (indicate where provided: section/paragraph) | n/a |
| Cell lines: Provide species information, strain. Provide accession number in repository OR supplier name, catalog number, clone number, OR RRID | Not involved in this article. | |
| Primary cultures: Provide species, strain, sex of origin, genetic modification status. | Not involved in this article. | N/A |
| Experimental animals | Yes (indicate where provided: section/paragraph) | n/a |
| Laboratory animals: Provide species, strain, sex, age, genetic modification status. Provide accession number in repository OR supplier name, catalog number, clone number, OR RRID | Not involved in this article. | N/A |
| Animal observed in or captured from the field: Provide species, sex and age where possible | Not involved in this article. | N/A |
| Model organisms: Provide Accession number in repository (where relevant) OR RRID | Not involved in this article. | N/A |
| Plants and microbes | Yes (indicate where provided: section/paragraph) | n/a |
| Plants: provide species and strain, unique accession number if available, and source (including location for collected wild specimens) | Not involved in this article. | N/A |
| Microbes: provide species and strain, unique | Not involved in this article. | N/A |

| Human research participants | Yes (indicate where provided: section/paragraph) | n/a |
|---|---|-----|
| Identify authority granting ethics approval (IRB or equivalent committee(s), provide reference number for approval. | Yes. Page 7, line 91-92. Methods/ Study subjects/ paragraph 1 | |
| Provide statement confirming informed consent obtained from study participants. | Yes. Page 7, line 100-101. Methods/ Study subjects/ paragraph 1 | |
| Report on age and sex for all study participants. | Yes. Page 7, line94, line 105-106. Methods/ Study subjects/ paragraph 1 | |

Design

| Study protocol | Yes (indicate where provided: section/paragraph) | n/a |
|---|--|-----|
| For clinical trials, provide the trial registration number OR cite DOI in manuscript. | Not involved in this article. | |
| Laboratory protocol | Yes (indicate where provided: section/paragraph) | n/a |
| Provide DOI or other citation details if detailed step- by-step protocols are available. | Yes. Page 7-9, line 110-136. Methods/ SNP selection and genotyping/ paragraph 1-3. | |
| Experimental study design (statistics details) | Yes (indicate where provided: section/paragraph) | n/a |
| State whether and how the following have been done, or if they were not carried out. | | |
| Sample size determination | Yes. Page 8, line 127-136. Methods/ SNP selection and genotyping/ paragraph 3. | |
| Randomisation | Not indicated in this article. | N/A |
| Blinding | Yes. Page 8, line 129-130. Methods/ SNP selection and genotyping/ paragraph 3. | |
| Inclusion/exclusion criteria | Yes. Page 8, line 133-134. Methods/ SNP selection and genotyping/ paragraph 3. | |
| Sample definition and in-laboratory replication | Yes (indicate where provided: section/paragraph) | n/a |
| State number of times the experiment was | Not indicated, data was generated from the | N/A |
| replicated in laboratory | HumanExome Beadchip array. | |
| Define whether data describe technical or biological replicates | Not indicated. | N/A |
| Ethics | Yes (indicate where provided: section/paragraph) | n/a |
| Studies involving human participants: State details of authority granting ethics approval (IRB or equivalent committee(s), provide reference number for approval. | Yes. Page 7, line91-92. Methods/ Study subjects/ paragraph 1 | |
| Studies involving experimental animals: State details of authority granting ethics approval (IRB or equivalent committee(s), provide reference number for approval. | Not involved in this article. | N/A |
| Studies involving specimen and field samples: State if relevant permits obtained, provide details of authority approving study; if none were required, explain why. | Not involved in this article. | N/A |
| Dual Use Research of Concern (DURC) | Yes (indicate where provided: section/paragraph) | n/a |
| If study is subject to dual use research of concern, state the authority granting approval and reference number for the regulatory approval | Not involved in this article. | N/A |

<u>Analysis</u>

| Attrition | Yes (indicate where provided: section/paragraph) | n/a |
|--|---|-----|
| State if sample or data point from the analysis is | Yes. Page 8-9, line 133-134. Methods/ SNP selection and | |
| excluded, and whether the criteria for exclusion were determined and specified in advance. | genotyping/ paragraph 3 | |

| Statistics | Yes (indicate where provided: section/paragraph) | n/a |
|--|--|-----|
| Describe statistical tests used and justify choice of tests. | Yes. Page 9, line 138-149. Methods/Statistical analyses/ | |
| | paragraph 1 | |

| Data Availability | Yes (indicate where provided: section/paragraph) | |
|--|---|-----|
| State whether newly created datasets are available, including protocols for access or restriction on access. | Not involved in this article. | N/A |
| If data are publicly available, provide accession number in repository or DOI or URL. | Yes. Page 9-10, line 154-158, line 160-166. Methods /Bioinformatics analysis&Online Kaplan-Meier plotter /paragraph 1,2,3 | |
| If publicly available data are reused, provide accession number in repository or DOI or URL, where possible. | No publicly available data are reused. | N/A |

| Code Availability | Yes (indicate where provided: section/paragraph) | n/a |
|---|--|-----|
| For all newly generated code and software essential for | | |
| replicating the main findings of the study: | | |
| State whether the code or software is available. | Not involved in this article. | N/A |
| If code is publicly available, provide accession number in repository, or DOI or URL. | Not involved in this article. | N/A |

Reporting

| Adherence to community standards | Yes (indicate where provided: section/paragraph) | |
|--|---|--|
| MDAR framework recommends adoption of discipline- | | |
| specific guidelines, established and endorsed through | | |
| community initiatives. Journals have their own policy | | |
| about requiring specific guidelines and recommendations | | |
| to complement MDAR. | | |
| State if relevant guidelines (eg., ICMJE, MIBBI, ARRIVE) | Yes, the paper follows the ICMJE recommendations. | |
| have been followed, and whether a checklist (eg., | STROBE Checklist is provided with the manuscript. | |
| CONSORT, PRISMA, ARRIVE) is provided with the | | |
| manuscript. | | |

Article information: http://dx.doi.org/10.21037/atm-20-6108

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

| Section/item | Item No | Recommendation | Reported on Page Number/Line Number | Reported on Section/Paragraph |
|------------------------------|------------|--|---|----------------------------------|
| Title and abstract | 1 | (a) Indicate the study's design with a commonly used term in the title or the abstract | Title page | Title |
| | | (b) Provide in the abstract an informative and balanced summary of what was done and what was found | Page 3~4/line 7~27 | Abstract/Paragraph 2~3 |
| Introduction | | | | |
| Background/ rationale | 2 | Explain the scientific background and rationale for the investigation being reported | Page 4~6/line 34~72 | Introduction/Paragraph 1~2 |
| Objectives | 3 | State specific objectives, including any prespecified hypotheses | Page 6/line 74~86 | Introduction/Paragraph 3 |
| Methods | | | | |
| Study design | 4 | Present key elements of study design early in the paper | Page 7/line 91~108 | Methods/ Paragraph 1~2 |
| Setting | 5 | Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection | Page 7/line 96-106 | Methods/ Paragraph 1 |
| Participants | 6 | (a) Cohort study —Give the eligibility criteria, and the sources and methods of selection of participants. Describe methods of follow-up 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 Cross-sectional study —Give the eligibility criteria, and the sources and methods of selection of participants | Page 7/line 91~106 | Methods/ Paragraph 1 |
| | | (b) Cohort study —For matched studies, give matching criteria and number of exposed and unexposed Case-control study —For matched studies, give matching criteria and the number of controls per case | N/A | N/A |
| Variables | 7 | Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable | Page 7/line 101~105 | Methods/ Paragraph 1 |
| Data sources/ measurement | 8* | For each variable of interest, give sources of data and details of methods of assessment (measurement). Describe comparability of assessment methods if there is more than one group | Page 7/ line 101~105 | Methods/ Paragraph 1 |
| Bias | 9 | Describe any efforts to address potential sources of bias | Page 7/line 105~106 | Methods/ Paragraph 1 |
| Study size | 10 | Explain how the study size was arrived at | Page 8~9/line 127~136 | Methods/ Paragraph 5 |
| Quantitative variables | 11 | Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why | Page 9/line 139~149 | Methods/ Paragraph 6 |

| Statistical methods | 12 | (a) Describe all statistical methods, including those used to control for confounding | Page 9/line 138~149 | Methods/Paragraph 6 | |
|------------------------|--------------|---|--|------------------------|----------------------|
| | | (b) Describe any methods used to examine subgroups and interactions | Page 9/line 145~147 | Methods /Paragraph 6 | |
| | | (c) Explain how missing data were addressed | N/A | N/A | |
| | | (d) Cohort study —If applicable, explain how loss to follow-up was addressed 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 | Page 7/line 105~106 | Methods/ Paragraph 1 | |
| | | (e) Describe any sensitivity analyses | N/A | N/A | |
| Results | | | | | |
| 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 | Page 10/line 169~170 | Results/Paragraph 1 | |
| | | (b) Give reasons for non-participation at each stage | Page 8~9/line133~134 | Methods/ Paragraph 6 | |
| | | (c) Consider use of a flow diagram | N/A | N/A | |
| Descriptive data | 14* | (a) Give characteristics of study participants (eg demographic, clinical, social) and information on exposures and potential confounders | Page 10/line 169~174 | Results/Paragraph 1 | |
| | | (b) Indicate number of participants with missing data for each variable of interest | N/A | N/A | |
| | | (c) Cohort study —Summarise follow-up time (eg, average and total amount) | N/A | N/A | |
| Outcome data | 15* | Cohort study — Report numbers of outcome events or summary measures over time | N/A | N/A | |
| | | Case-control study—Report numbers in each exposure category, or summary measures of exposure | Page 11~12/line 195~204 | Results/Paragraph 4 | |
| | | Cross-sectional study—Report numbers of outcome events or summary measures | N/A | N/A | |
| Main results | 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 | Page 11/line 185~186 | Results/ Paragraph 2 |
| | | (b) Report category boundaries when continuous variables were categorized | Page 11/line 195~200 | Results/Paragraph 4 | |
| | | (c) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period | N/A | N/A | |
| Other analyses | 17 | Report other analyses done - eg analyses of subgroups and interactions, and sensitivity analyses | Page 11/line 200~204 | Results/paragraph 4 | |
| Discussion | | | | | |
| Key results | 18 | Summarise key results with reference to study objectives | Page 13/line 226-233 | Discussion/Paragraph 1 | |
| Limitations | 19 | Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias | Page 16/line 302-308 | Discussion/Paragraph 6 | |

| Interpretation | 20 | Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence | Page 16/line 298~309 | Discussion/Paragraph 6 |
|-------------------|----|--|-------------------------|----------------------------------|
| Generalisability | 21 | Discuss the generalisability (external validity) of the study results | Page 13~16/line 234~297 | Discussion/Paragraph 2~5 |
| Other information | | | | |
| Funding | 22 | Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based | Page 17/line 313-317 | Acknowledgements/ Paragraph 2 |

^{*}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.

Article information: http://dx.doi.org/10.21037/atm-20-6108

^{*}As the checklist was provided upon initial submission, the page number/line number reported may be changed due to copyediting and may not be referable in the published version. In this case, the section/paragraph may be used as an alternative reference.