

## TRIPOD Checklist: Prediction Model Development and Validation

| Section                   | Item |     | Checklist description  | Reported on Page Number/Line Number | Reported on Section/Paragraph      |
|---------------------------|------|-----|--|-------------------------------------|------------------------------------|
| <b>Title and abstract</b> |      |     |  |                                     |                                    |
| Title                     | 1    | D;V | Identify the study as developing and/or validating a multivariable prediction model, the target population, and the outcome to be predicted.   | Page 1/Line 1-2                     | Title Section                      |
| Abstract                  | 2    | D;V | Provide a summary of objectives, study design, setting, participants, sample size, predictors, outcome, statistical analysis, results, and conclusions.  | Page 1-3/Line 20-48                 | Abstract Section                   |
| <b>Introduction</b>       |      |     |  |                                     |                                    |
| Background and objectives | 3a   | D;V | Explain the medical context (including whether diagnostic or prognostic) and rationale for developing or validating the multivariable prediction model, including references to existing models. | Page 3-4/Line 52-75                 | Introduction Section/Paragraph 1-2 |
|                           | 3b   | D;V | Specify the objectives, including whether the study describes the development or validation of the model or both.  | Page 4/Line 76-81                   | Introduction Section/Paragraph 3   |
| <b>Methods</b>            |      |     |  |                                     |                                    |
| Source of data            | 4a   | D;V | Describe the study design or source of data (e.g., randomized trial, cohort, or registry data), separately for the development and validation data sets, if applicable.                          | Page 4-5/Line 84-95                 | Method Section/Paragraph 1-2       |
|                           | 4b   | D;V | Specify the key study dates, including start of accrual; end of accrual; and, if applicable, end of follow-up.   | /                                   | /                                  |
| Participants              | 5a   | D;V | Specify key elements of the study setting (e.g., primary care, secondary care, general population) including number and location of centres.   | /                                   | /                                  |
|                           | 5b   | D;V | Describe eligibility criteria for participants.  | /                                   | /                                  |
|                           | 5c   | D;V | Give details of treatments received, if relevant.  | /                                   | /                                  |
| Outcome                   | 6a   | D;V | Clearly define the outcome that is predicted by the prediction model, including how and when assessed.   | Page 5-6/Line 103-113               | Method Section/Paragraph 4         |

|             |    |     |   |                     |                              |
|-------------|----|-----|---|---------------------|------------------------------|
|             | 6b | D;V | Report any actions to blind assessment of the outcome to be predicted.  | /                   | /                            |
| Predictors  | 7a | D;V | Clearly define all predictors used in developing or validating the multivariable prediction model, including how and when they were measured. | Page 6/Line 115-125 | Method Section/Paragraph 5-6 |
|             | 7b | D;V | Report any actions to blind assessment of predictors for the outcome and other predictors.  | /                   | /                            |
| Sample size | 8  | D;V | Explain how the study size was arrived at.  | /                   | /                            |

|                              |     |     |   |                     |                            |
|------------------------------|-----|-----|---|---------------------|----------------------------|
| Missing data                 | 9   | D;V | Describe how missing data were handled (e.g., complete-case analysis, single imputation, multiple imputation) with details of any imputation method.  | /                   | /                          |
| Statistical analysis methods | 10a | D   | Describe how predictors were handled in the analyses.   | Page 7/Line 135-139 | Method Section/Paragraph 8 |
|                              | 10b | D   | Specify type of model, all model-building procedures (including any predictor selection), and method for internal validation.   | Page5/line 103-113  | Method Section             |
|                              | 10c | V   | For validation, describe how the predictions were calculated.   | Page5-6/line103-125 | Method Section             |
|                              | 10d | D;V | Specify all measures used to assess model performance and, if relevant, to compare multiple models.   | Page5-7             | Method Section             |
|                              | 10e | V   | Describe any model updating (e.g., recalibration) arising from the validation, if done.   | /                   | /                          |
| Risk groups                  | 11  | D;V | Provide details on how risk groups were created, if done.   | Page5/line 103-113  | Method Section             |
| Development vs. validation   | 12  | V   | For validation, identify any differences from the development data in setting, eligibility criteria, outcome, and predictors.   | Page5-7             | Method Section             |
| <b>Results</b>               |     |     |   |                     |                            |
| Participants                 | 13a | D;V | Describe the flow of participants through the study, including the number of participants with and without the outcome and, if applicable, a summary of the follow-up time. A diagram may be helpful. | Page 7/Line 143-147 | Result Section/Paragraph 1 |

|                          |     |     |  |                         |                                  |
|--------------------------|-----|-----|--|-------------------------|----------------------------------|
|                          | 13b | D;V | Describe the characteristics of the participants (basic demographics, clinical features, available predictors), including the number of participants with missing data for predictors and outcome. | /                       | /                                |
|                          | 13c | V   | For validation, show a comparison with the development data of the distribution of important variables (demographics, predictors and outcome).   | Page 8-9/Line 174-186   | Result Section/Paragraph 5       |
| Model development        | 14a | D   | Specify the number of participants and outcome events in each analysis.  | Page 8/Line 162-173     | Result Section/Paragraph 4       |
|                          | 14b | D   | If done, report the unadjusted association between each candidate predictor and outcome.   | /                       | /                                |
| Model specification      | 15a | D   | Present the full prediction model to allow predictions for individuals (i.e., all regression coefficients, and model intercept or baseline survival at a given time point).                        | Page 9-10/Line 189-200  | Result Section/Paragraph 6-8     |
|                          | 15b | D   | Explain how to use the prediction model.   | Page 11-12/line 240-244 | Result Section                   |
| Model performance        | 16  | D;V | Report performance measures (with CIs) for the prediction model.   | Page 10-11/Line 202-241 | Result Section/Paragraph 9-13    |
| Model-updating           | 17  | V   | If done, report the results from any model updating (i.e., model specification, model performance).  | /                       | /                                |
| <b>Discussion</b>        |     |     |  |                         |                                  |
| Limitations              | 18  | D;V | Discuss any limitations of the study (such as nonrepresentative sample, few events per predictor, missing data).   | Page 15/Line 315-319    | Discussion Section/Paragraph 7   |
| Interpretation           | 19a | V   | For validation, discuss the results with reference to performance in the development data, and any other validation data.  | Page 12-13/Line 262-280 | Discussion Section/Paragraph 3   |
|                          | 19b | D;V | Give an overall interpretation of the results, considering objectives, limitations, and results from similar studies, and other relevant evidence.   | Page 13-15/Line 281-313 | Discussion Section/Paragraph 4-6 |
| Implications             | 20  | D;V | Discuss the potential clinical use of the model and implications for future research.  | Page 14-15/Line 305-313 | Discussion Section/Paragraph 6   |
| <b>Other information</b> |     |     |  |                         |                                  |

|                           |    |     |   |                         |                                   |
|---------------------------|----|-----|---|-------------------------|-----------------------------------|
| Supplementary information | 21 | D;V | Provide information about the availability of supplementary resources, such as study protocol, Web calculator, and data sets. | Page 16/Line 335-337    | Declaration Section/Paragraph 3   |
| Funding                   | 22 | D;V | Give the source of funding and the role of the funders for the present study.   | Page 15-16/Line 329-334 | Declaration Section/Paragraph 1-2 |

\* Items relevant only to the development of a prediction model are denoted by D, items relating solely to a validation of a prediction model are denoted by V, and items relating to both are denoted D;V. We recommend using the TRIPOD Checklist in conjunction with the TRIPOD Explanation and Elaboration document.

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\*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.

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