

ICMJE Form for Disclosure of Potential Conflicts of Interest

Instructions

The purpose of this form is to provide readers of your manuscript with information about your other interests that could influence how they receive and understand your work. The form is designed to be completed electronically and stored electronically. It contains programming that allows appropriate data display. Each author should submit a separate form and is responsible for the accuracy and completeness of the submitted information. The form is in six parts.

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5. Intellectual Property.

This section asks about patents and copyrights, whether pending, issued, licensed and/or receiving royalties.

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Royalties: Funds are coming in to you or your institution due to your patent

ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Lai-Rong 2. Surname (Last Name) Song 3. Date 27-February-2020

4. Are you the corresponding author? ☐ Yes ☒ No Corresponding Author's Name Jun-Ting Zhang

5. Manuscript Title Prognostic and predictive value of an immune infiltration signature in diffuse lower-grade gliomas

6. Manuscript Identifying Number (if you know it) 133811-INS-CMED-RV-2

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Are there any relevant conflicts of interest? ☐ Yes ☒ No

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Dr. Song has nothing to disclose.

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Section 1. Identifying Information

1. Given Name (First Name) Jian-Cong 2. Surname (Last Name) Weng 3. Date 27-February-2020

4. Are you the corresponding author? ☐ Yes ☒ No Corresponding Author's Name Jun-Ting Zhang

5. Manuscript Title
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6. Manuscript Identifying Number (if you know it)
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Section 1. Identifying Information

1. Given Name (First Name)
Cheng-Bei

2. Surname (Last Name)
Li

3. Date
27-February-2020

4. Are you the corresponding author? ☐ Yes ☒ No
Corresponding Author's Name
Jun-Ting Zhang

5. Manuscript Title
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6. Manuscript Identifying Number (if you know it)
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Section 1. Identifying Information

1. Given Name (First Name) Xu-Lei	2. Surname (Last Name) Huo	3. Date 27-February-2020
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Jun-Ting Zhang
5. Manuscript Title Prognostic and predictive value of an immune infiltration signature in diffuse lower-grade gliomas		
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Li

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27-February-2020

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ICMJE Form for Disclosure of Potential Conflicts of Interest

Section 1. Identifying Information

1. Given Name (First Name) Shu-Yu	2. Surname (Last Name) Hao	3. Date 27-February-2020
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Jun-Ting Zhang
5. Manuscript Title Prognostic and predictive value of an immune infiltration signature in diffuse lower-grade gliomas		
6. Manuscript Identifying Number (if you know it) 133811-INS-CMED-RV-2		

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Dr. Hao has nothing to disclose.

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1. Given Name (First Name) Zhen	2. Surname (Last Name) Wu	3. Date 27-February-2020
4. Are you the corresponding author? <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No		Corresponding Author's Name Jun-Ting Zhang
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1. Given Name (First Name)
Liang

2. Surname (Last Name)
Wang

3. Date
27-February-2020

4. Are you the corresponding author? ☐ Yes ☒ No Corresponding Author's Name
Jun-Ting Zhang

5. Manuscript Title
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Section & Topic	No	Item	Reported on page #
TITLE OR ABSTRACT			
	1	Identification as a study of diagnostic accuracy using at least one measure of accuracy (such as sensitivity, specificity, predictive values, or AUC)	AUC and c-index were used.
ABSTRACT			
	2	Structured summary of study design, methods, results, and conclusions (for specific guidance, see STARD for Abstracts)	Yes
INTRODUCTION			
	3	Scientific and clinical background, including the intended use and clinical role of the index test	Included
	4	Study objectives and hypotheses	Included
METHODS			
<i>Study design</i>	5	Whether data collection was planned before the index test and reference standard were performed (prospective study) or after (retrospective study)	Retrospective study
<i>Participants</i>	6	Eligibility criteria	Included
	7	On what basis potentially eligible participants were identified (such as symptoms, results from previous tests, inclusion in registry)	LGGs with molecular and clinical information
	8	Where and when potentially eligible participants were identified (setting, location and dates)	NA
	9	Whether participants formed a consecutive, random or convenience series	Convenience series
<i>Test methods</i>	10a	Index test, in sufficient detail to allow replication	Yes
	10b	Reference standard, in sufficient detail to allow replication	Yes
	11	Rationale for choosing the reference standard (if alternatives exist)	NA
	12a	Definition of and rationale for test positivity cut-offs or result categories of the index test, distinguishing pre-specified from exploratory	Yes
	12b	Definition of and rationale for test positivity cut-offs or result categories of the reference standard, distinguishing pre-specified from exploratory	NA
	13a	Whether clinical information and reference standard results were available to the performers/readers of the index test	Yes
	13b	Whether clinical information and index test results were available to the assessors of the reference standard	Yes
<i>Analysis</i>	14	Methods for estimating or comparing measures of diagnostic accuracy	Time-dependent ROC curves, Harrell's C statistic
	15	How indeterminate index test or reference standard results were handled	NA
	16	How missing data on the index test and reference standard were handled	Deletion
	17	Any analyses of variability in diagnostic accuracy, distinguishing pre-specified from exploratory	NA
	18	Intended sample size and how it was determined	NA
RESULTS			
<i>Participants</i>	19	Flow of participants, using a diagram	Figure 1
	20	Baseline demographic and clinical characteristics of participants	Table S1
	21a	Distribution of severity of disease in those with the target condition	Figure 4
	21b	Distribution of alternative diagnoses in those without the target condition	NA
	22	Time interval and any clinical interventions between index test and reference standard	Figure 2
<i>Test results</i>	23	Cross tabulation of the index test results (or their distribution) by the results of the reference standard	Table S1
	24	Estimates of diagnostic accuracy and their precision (such as 95% confidence intervals)	Figure 4, Figure S5, Figure 8
	25	Any adverse events from performing the index test or the reference standard	NA
DISCUSSION			
	26	Study limitations, including sources of potential bias, statistical uncertainty, and generalisability	Included
	27	Implications for practice, including the intended use and clinical role of the index test	Included

OTHER INFORMATION			
	28	Registration number and name of registry	NA
	29	Where the full study protocol can be accessed	Yes
	30	Sources of funding and other support; role of funders	Included

STARD 2015

AIM

STARD stands for “Standards for Reporting Diagnostic accuracy studies”. This list of items was developed to contribute to the completeness and transparency of reporting of diagnostic accuracy studies. Authors can use the list to write informative study reports. Editors and peer-reviewers can use it to evaluate whether the information has been included in manuscripts submitted for publication.

EXPLANATION

A **diagnostic accuracy study** evaluates the ability of one or more medical tests to correctly classify study participants as having a **target condition**. This can be a disease, a disease stage, response or benefit from therapy, or an event or condition in the future. A medical test can be an imaging procedure, a laboratory test, elements from history and physical examination, a combination of these, or any other method for collecting information about the current health status of a patient.

The test whose accuracy is evaluated is called **index test**. A study can evaluate the accuracy of one or more index tests. Evaluating the ability of a medical test to correctly classify patients is typically done by comparing the distribution of the index test results with those of the **reference standard**. The reference standard is the best available method for establishing the presence or absence of the target condition. An accuracy study can rely on one or more reference standards.

If test results are categorized as either positive or negative, the cross tabulation of the index test results against those of the reference standard can be used to estimate the **sensitivity** of the index test (the proportion of participants *with* the target condition who have a positive index test), and its **specificity** (the proportion *without* the target condition who have a negative index test). From this cross tabulation (sometimes referred to as the contingency or “2x2” table), several other accuracy statistics can be estimated, such as the positive and negative **predictive values** of the test. Confidence intervals around estimates of accuracy can then be calculated to quantify the statistical **precision** of the measurements.

If the index test results can take more than two values, categorization of test results as positive or negative requires a **test positivity cut-off**. When multiple such cut-offs can be defined, authors can report a receiver operating characteristic (ROC) curve which graphically represents the combination of sensitivity and specificity for each possible test positivity cut-off. The **area under the ROC curve** informs in a single numerical value about the overall diagnostic accuracy of the index test.

The **intended use** of a medical test can be diagnosis, screening, staging, monitoring, surveillance, prediction or prognosis. The **clinical role** of a test explains its position relative to existing tests in the clinical pathway. A replacement test, for example, replaces an existing test. A triage test is used before an existing test; an add-on test is used after an existing test.

Besides diagnostic accuracy, several other outcomes and statistics may be relevant in the evaluation of medical tests. Medical tests can also be used to classify patients for purposes other than diagnosis, such as staging or prognosis. The STARD list was not explicitly developed for these other outcomes, statistics, and study types, although most STARD items would still apply.

DEVELOPMENT

This STARD list was released in 2015. The 30 items were identified by an international expert group of methodologists, researchers, and editors. The guiding principle in the development of STARD was to select items that, when reported, would help readers to judge the potential for bias in the study, to appraise the applicability of the study findings and the validity of conclusions and recommendations. The list represents an update of the first version, which was published in 2003.

More information can be found on <http://www.equator-network.org/reporting-guidelines/stard>.

