QUADAS-2

Phase 1: State the review question:

<table>
<thead>
<tr>
<th>Patients (setting, intended use of index test, presentation, prior testing):</th>
</tr>
</thead>
<tbody>
<tr>
<td>Index test(s):</td>
</tr>
<tr>
<td>Reference standard and target condition:</td>
</tr>
</tbody>
</table>

Phase 2: Draw a flow diagram for the primary study
Phase 3: Risk of bias and applicability judgments

QUADAS-2 is structured so that 4 key domains are each rated in terms of the risk of bias and the concern regarding applicability to the research question (as defined above). Each key domain has a set of signalling questions to help reach the judgments regarding bias and applicability.

### DOMAIN 1: PATIENT SELECTION

#### A. Risk of Bias

Describe methods of patient selection:

<table>
<thead>
<tr>
<th>Question</th>
<th>Yes/No/Unclear</th>
</tr>
</thead>
<tbody>
<tr>
<td>Was a consecutive or random sample of patients enrolled?</td>
<td>Yes/No/Unclear</td>
</tr>
<tr>
<td>Was a case-control design avoided?</td>
<td>Yes/No/Unclear</td>
</tr>
<tr>
<td>Did the study avoid inappropriate exclusions?</td>
<td>Yes/No/Unclear</td>
</tr>
</tbody>
</table>

Could the selection of patients have introduced bias? **RISK: LOW/HIGH/UNCLEAR**

#### B. Concerns regarding applicability

Describe included patients (prior testing, presentation, intended use of index test and setting):

Is there concern that the included patients do not match the review question? **CONCERN: LOW/HIGH/UNCLEAR**

### DOMAIN 2: INDEX TEST(S)

If more than one index test was used, please complete for each test.

#### A. Risk of Bias

Describe the index test and how it was conducted and interpreted:

<table>
<thead>
<tr>
<th>Question</th>
<th>Yes/No/Unclear</th>
</tr>
</thead>
<tbody>
<tr>
<td>Were the index test results interpreted without knowledge of the results of the reference standard?</td>
<td>Yes/No/Unclear</td>
</tr>
<tr>
<td>If a threshold was used, was it pre-specified?</td>
<td>Yes/No/Unclear</td>
</tr>
</tbody>
</table>

Could the conduct or interpretation of the index test have introduced bias? **RISK: LOW/HIGH/UNCLEAR**

#### B. Concerns regarding applicability

Is there concern that the index test, its conduct, or interpretation differ from the review question? **CONCERN: LOW/HIGH/UNCLEAR**
DOMAIN 3: REFERENCE STANDARD

A. Risk of Bias

Describe the reference standard and how it was conducted and interpreted:

- Is the reference standard likely to correctly classify the target condition? Yes/No/Unclear
- Were the reference standard results interpreted without knowledge of the results of the index test? Yes/No/Unclear

Could the reference standard, its conduct, or its interpretation have introduced bias? RISK: LOW /HIGH/UNCLEAR

B. Concerns regarding applicability

Is there concern that the target condition as defined by the reference standard does not match the review question? CONCERN: LOW /HIGH/UNCLEAR

DOMAIN 4: FLOW AND TIMING

A. Risk of Bias

Describe any patients who did not receive the index test(s) and/or reference standard or who were excluded from the 2x2 table (refer to flow diagram):

Describe the time interval and any interventions between index test(s) and reference standard:

- Was there an appropriate interval between index test(s) and reference standard? Yes/No/Unclear
- Did all patients receive a reference standard? Yes/No/Unclear
- Did patients receive the same reference standard? Yes/No/Unclear
- Were all patients included in the analysis? Yes/No/Unclear

Could the patient flow have introduced bias? RISK: LOW /HIGH/UNCLEAR