### 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:

- Was a consecutive or random sample of patients enrolled?  
  - Yes/No/Unclear
- Was a case-control design avoided?  
  - Yes/No/Unclear
- Did the study avoid inappropriate exclusions?  
  - Yes/No/Unclear

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:

- Were the index test results interpreted without knowledge of the results of the reference standard?  
  - Yes/No/Unclear
- If a threshold was used, was it pre-specified?  
  - Yes/No/Unclear

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