Figure E2: Risk of Bias Assessment for Question 1

| Study                                      | Was the sample frame appropriate to address the target population? | Were study participants sampled in an appropriate way? | Was the sample size adequate? | Were the study subjects and the setting described in detail? | Was the data analysis conducted with sufficient coverage of the identified sample? | Were valid methods used for the identification of the condition? | Was the condition measured in a standard, reliable way for all participants? | Was there appropriate statistical analysis? | Was the response rate adequate, and if not, was the low response rate managed appropriately? | Overall Risk of Bias |
|--------------------------------------------|---------------------------------------------------------------|-----------------------------------------------------|-------------------------------|------------------------------------------------------------|------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| UK MHRA                                    |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| UK MHRA                                    |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| UK MHRA                                    |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| UK MHRA                                    |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Institute of Public Health of Chile²      |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Institute of Public Health of Chile²      |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Institute of Public Health of Chile²      |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Institute of Public Health of Chile²      |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Australia Health Department⁴               |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Australia Health Department⁴               |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Mexico Health Ministry⁴                    |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Singapore Health Ministry⁵                 |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| USA CDC VAERS⁶                             |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| MGB (Boston)⁶                               |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| India Health Ministry⁴                     |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Public Health Agency of Canada⁹⁰           |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Poland Health Ministry¹⁰                   |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| USA NCT04405076⁶                           |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Australia and the United States NCT04360988⁴⁵ |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| USA NCT04537208⁴³                           |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| South Africa NCT04444674⁴⁴                 |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| South Africa NCT04533399⁴⁵                 |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| USA NCT04436276⁴⁶                           |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| USA NCT04470427³⁷                           |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | LOW                                                                    |
| Australia NCT04495933⁴¹                     |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| China NCT04412538⁴⁸                         |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| India NCT04471519²⁰                         |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| India NCT04471519²¹                         |                                                |                                                     |                               |                                                             |                                                                                                           |                                                                                                                                  |                                                                                                                                  |                                                                                                                                  |                                                                                                                                                                                                 | HIGH                                                                   |
| Registration | Region |
|--------------|--------|
| NCT04505722  | Multiple |
| NCT04368988  | Australia |
| NCT04530396  | Russia |
| NCT04368728  | Multiple |
| NCT04421528  | China |
| NCT04405908  | Australia |
| NCT04368728  | USA |
| NCT044450004 | Canada |
| NCT04383574  | China |
| CTR2000032459 | China |
| CTR2000031809 | China |
| NCT04466085  | China |
| NCT0445194   | China |
| NCT04352608  | China |
| NCT04341389  | China |
| NCT043841389 | Israel |
| NCT0431890   | Japan |
| NCT043043189 | Norway |
| NCT0431890   | Romania |
| NCT0431890   | Romania |
| NCT0431890   | Netherlands |
| NCT0431890   | Netherlands |
| NCT0431890   | Switzerland |
| NCT0431890   | Denmark |
| NCT0431890   | Denmark |
| NCT0431890   | Denmark |

**Notes:**
- **LOW** indicates a low level of regulation.
- **HIGH** indicates a high level of regulation.

*Numbers indicate additional registration details.*