**S2 Appendix. Causality assessment of AEs.**

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| Causality term  | Assessment criteria |
| Certain | * Event or laboratory test abnormality, with plausible time relationship to drug intake
* Cannot be explained by disease or other drugs
* Response to withdrawal plausible (pharmacologically, pathologically)
* Event definitive pharmacologicallyor phenomenologically (i.e. an objective and specific medical disorderor a recognised pharmacological phenomenon)
* Rechallenge satisfactory, if necessary
 |
| Probable / Likely | * Event or laboratory test abnormality, with reasonable time relationship to drug intake
* Unlikely to be attributed to disease or other drugs
* Response to withdrawal clinically reasonable
* Rechallenge not required
 |
| Possible | * Event or laboratory test abnormality, with reasonable time relationship to drug intake
* Could also be explained by disease or other drugs
* Information on drug withdrawal may be lacking or unclear
 |
| Unlikely | * Event or laboratory test abnormality, with a time to drug intake that makes a relationship improbable (but not impossible)
* Disease or other drugs provide plausible explanations
 |
| Conditional / Unclassified | * Event or laboratory test abnormality
* More data for proper assessment needed, or
* Additional data under examination
 |
| Unassessable / Unclassifiable | * Report suggesting an adverse reaction
* Cannot be judged because information is insufficient or contradictory
* Data cannot be supplemented or verified
 |