**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 |