Safety reporting flowchart

Adverse Event Reporting: UK Open Label Trial

PI assesses causality[1]

Related → **Adverse Reaction (AR)**

Not related → **Adverse Event (AE)**

PI assesses seriousness

Not serious[2] → **Adverse Event/Reaction (AE/R)**
PI records and notifies sponsor as per protocol

Serious SAE/R

PI checks protocol to confirm whether SAE/R requires expedited reporting

NO → **'Non-Expeditable' (SAE/R)**
PI records and notifies sponsor as per protocol

YES

PI notifies sponsor of SAE/R within 24 hours

Sponsor's assessment of causality[3]

Unrelated to IMP → **Serious Adverse Event (SAE)**
Sponsor keeps records and follows up until resolution

Related to IMP

Serious Adverse Reaction (SAR)

Sponsor's assessment of expectedness using the RSI[4]

Unexpected → **Suspected Unexpected Serious Adverse Reaction (SUSAR)**

Sponsor to report to MHRA and Ethics Committee:

- Fatal or life threatening SUSARs within 7 days
- All other SUSARs within 15 days

SUSARs reported to PIs as per protocol

Expected

Expected Serious Adverse Reaction (SAR)
Sponsor keeps records and follows up until resolution

Adverse Event (AE):
Any untoward medical occurrence in a clinical trial subject administered a medicinal product and which does not necessarily have a causal relationship with this treatment.

Adverse Reaction (AR):
Any untoward and unintended response to an IMP which is related (a reasonable causal relationship) to any dose administered.

Serious Adverse Event/Reaction (SAE/R):

- Results in death,
- is life-threatening,
- requires hospitalisation or prolongation of existing hospitalisation,
- results in persistent or significant disability or incapacity,
- is a congenital anomaly or birth defect,
- any other safety issues considered medically important.

PI should **actively seek** follow-up information on reported SAE/Rs.

Footnotes

[1] PI or delegate.
[2] Notable or safety critical events must be reported as per protocol.
[3] Sponsor cannot downgrade the PI's causality assessment, but can upgrade it.
[4] Reference Safety Information (RSI) in IB or SmPC.