**Supplementary Table.** Definition of safety endpoints. AE, adverse event; ADE, adverse device effect; CIP, clinical investigation plan; DD, device deficiencies; SAE, serious adverse event; SADE, adverse device effect; USADE, unanticipated serious adverse device effects.

| Safety endpoint | Definition |
|---|---|
| **AE** | Untoward medical occurrence, unintended disease or injury, or untoward clinical signs (including abnormal laboratory findings) in **subjects**, users or other persons, whether or not related to the **investigational medical device** and whether anticipated or unanticipated.<br>Note 1 to entry: This definition includes events related to the investigational medical device or the **comparator**.<br>Note 2 to entry: This definition includes events related to the procedures involved.<br>Note 3 to entry: For users or other persons, this definition is restricted to events related to the use of investigational medical devices or comparators. |
| **SAE** | **Adverse event** that led to any of the following:<br>a) death,<br>b) serious deterioration in the health of the **subject**, users, or other persons as defined by one or more of the following:<br>1) a life-threatening illness or injury, or<br>2) a permanent impairment of a body structure or a body function including chronic diseases, or<br>3) in-patient or prolonged hospitalization, or<br>4) medical or surgical intervention to prevent life-threatening illness or injury, or permanent impairment to a body structure or a body function,<br>c) foetal distress, foetal death, a congenital abnormality, or birth defect including physical or mental impairment.<br>Note 1 to entry: Planned hospitalization for a pre-existing condition, or a procedure required by the CIP, without serious deterioration in health, is not considered a serious adverse event. |
| **ADE** | Related to the use of an investigational **medical device.**<br>Note 1 to entry: This definition includes adverse events resulting from insufficient or inadequate instructions for use, deployment, implantation, installation, or operation, or any **malfunction** of the investigational medical device.<br>Note 2 to entry: This definition includes any event resulting from **use error** or from intentional misuse of the investigational medical device.<br>Note 3 to entry: This includes '*comparator*' if the comparator is a medical device. |
| **SADE** | Adverse device effect that has resulted in any of the consequences characteristic of a serious adverse event. |

Source: ISO 14155:2020 [43].

**Supplementary Table.** Definition of safety endpoints. AE, adverse event; ADE, adverse device effect; CIP, clinical investigation plan; DD, device deficiencies; SAE, serious adverse event; SADE, adverse device effect; USADE, unanticipated serious adverse device effects.

| Safety endpoint | Definition |
|---|---|
| **USADE** | Serious adverse device effect which by its nature, incidence, severity or outcome has not been identified in the current risk assessment.<br>Note 1 to entry: Anticipated serious adverse device effect (ASADE) is an effect which by its nature, incidence, severity or outcome has been identified in the risk assessment. |
| **DD** | Inadequacy of a **medical device** with respect to its identity, quality, durability, reliability, usability, safety or performance<br>Note 1 to entry: Device deficiencies include **malfunctions**, **use errors**, and inadequacy in the information supplied by the manufacturer including labelling.<br>Note 2 to entry: This definition includes device deficiencies related to the **investigational medical device** or the **comparator**. |

Source: ISO 14155:2020 [46].