

Safety reporting in clinical trials with drugs

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Agenda

- > Presentation/Promotion SOP
 - Overview on events
 - Overview on SOP: process and procedures of safety reporting
 - Highlights and examples of the safety reporting core procedures: detection, assessment, documentation, reporting
- > Q & A, Discussion

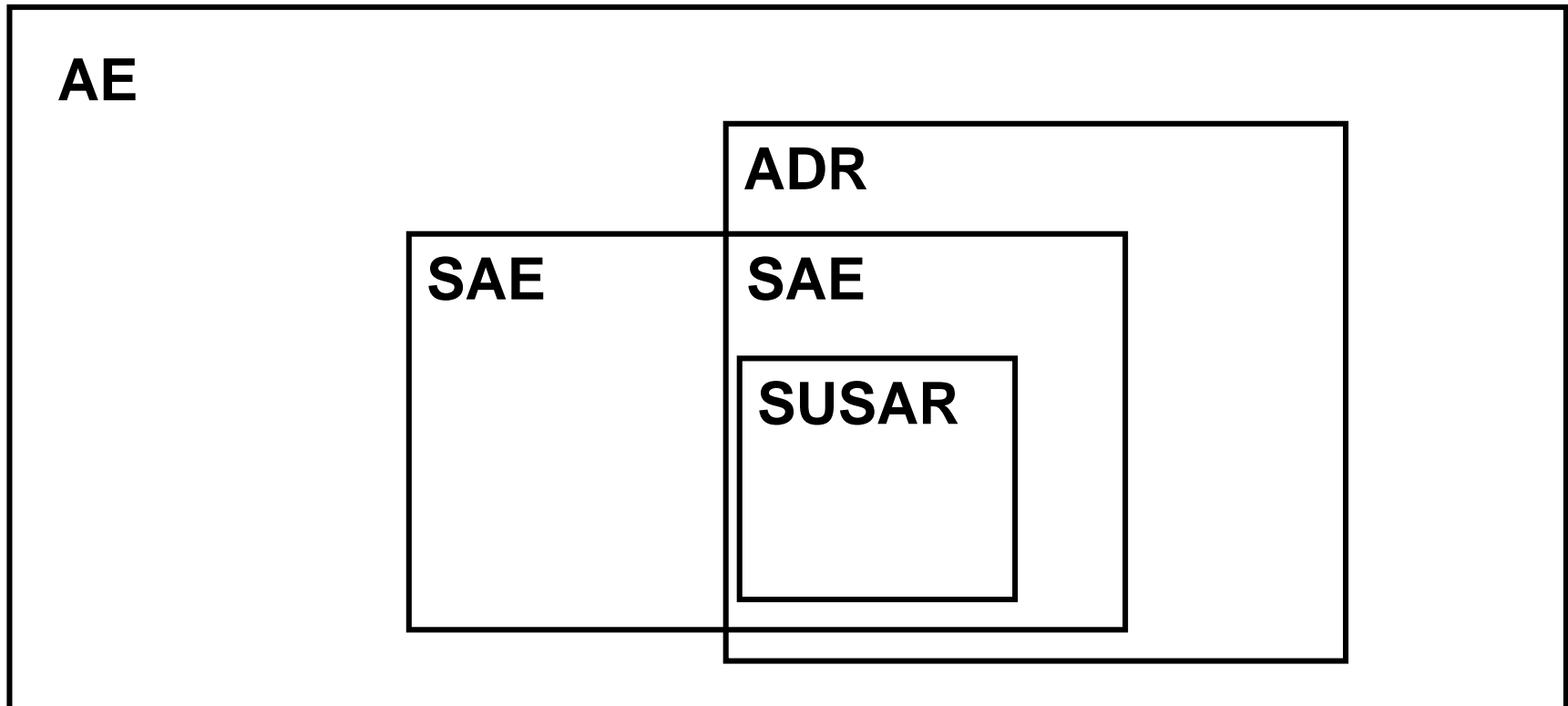
Limitations:

- > No in-depth definitions
 - > Not a step by step instruction
 - > Limited to IMP trials
-

Overview on «events»

Not drug related

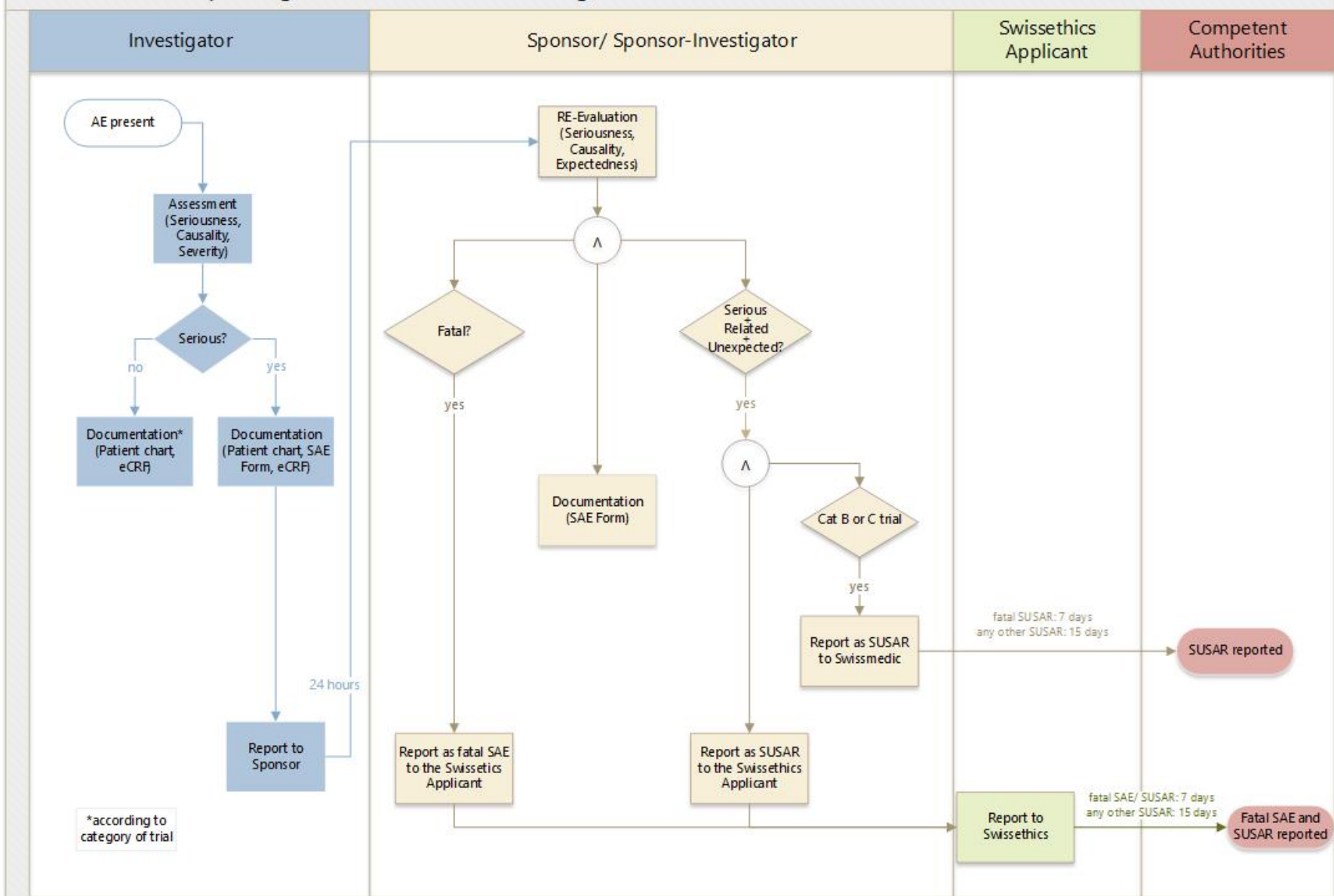
Drug related



*A=Adverse; E=Event; D= Drug R=Reaction;
S=Serious/Suspected; U=Unexpected*

Overview on the process

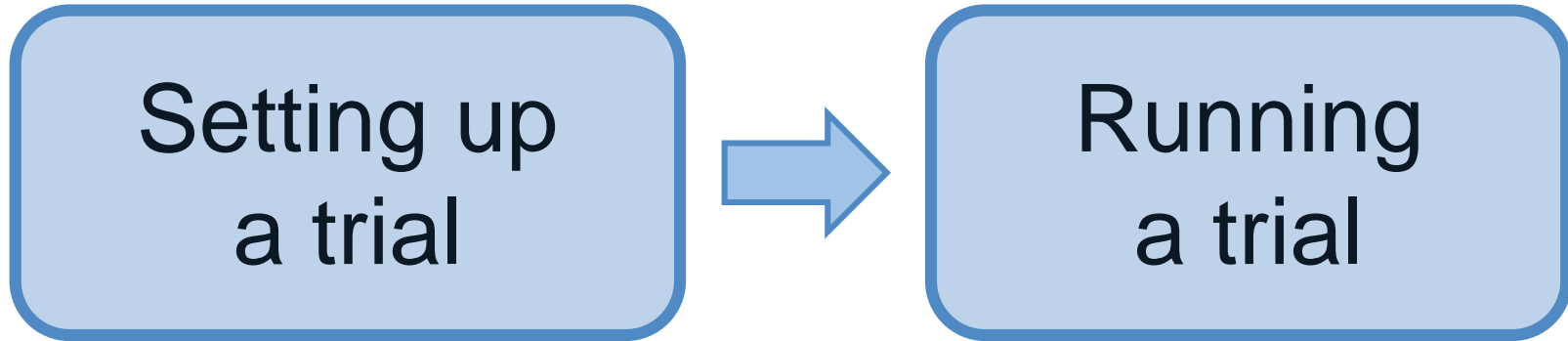
Adverse event reporting in clinical trials with drugs



Overview of procedures

Who		What	How
Investigator	Sponsor	Detection	Where the documentation of AEs/ADRs applies (as per category of the study or as required in the protocol), detect them based on spontaneous reports (e.g. by the subject), (...)
		Assessment/ Re-Evaluation	Assess all AEs/ADRs occurring during a clinical trial, regardless of whether they occur under the investigational medicinal product, comparative treatment, placebo, or no treatment (e.g. follow-up, wash-out phase). (...)
		Documentation	Document AEs/ADRs in source documents (e.g. Patient charts, specific AE form), and record them in the Case Report Forms (CRF). (...)
	Swissthethics Applicant	Reporting	Report AEs/ADRs to the Sponsor via (e)CRF as required by the protocol.

Who, What, How... and When?



... and Why?

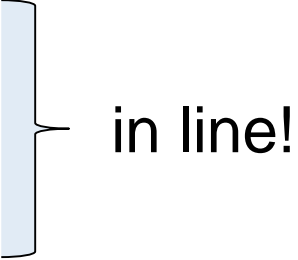
Patient safety and well-being

Who		What	
Investigator	Sponsor	Detection	<p>AE/SAE detection – How?</p> <p>Once you have the starting point the process is quite straightforward</p> <p>AE detection is how to get there...</p>
		Assessment/ Re-Evaluation	
		Documentation	
	SE-Applicant	Reporting	<p>Objective: Supervise known risks Detect unknown risks</p> <p>Danger: Information overload</p>

Who		What	
Investigator	Sponsor	Detection	(ClinO, art.39, art.40) Category A: Only SAEs Category B: SAEs Requested by EC/RA According to protocol
		Assessment/ Re-Evaluation	Category C: All AEs
	Documentation	<div data-bbox="718 668 1711 1076" data-label="Diagram"> <p>The diagram consists of a large black-outlined rectangle labeled 'AEs'. Inside this rectangle, there is a smaller black-outlined rectangle labeled 'SAEs'. This 'SAEs' rectangle is partially enclosed by a light blue oval that overlaps its left and bottom edges.</p> </div>	
SE-Applicant	Reporting	(ClinO, art.40): Events which are not to be reported according to the protocol are exempted.	

Who		What	
Investigator		Detection	<p>Example 1: Heart surgery Solution: Exempt expected SAEs</p>
	Sponsor	Assessment/ Re-Evaluation	<p>Example 2: Botox trial Solution: Define AE detection time window</p>
		Documentation	<p>Example 3: Ocular injection Solution: Only ophthalmic events</p>
	SE-Applicant	Reporting	<p>Warning! Swissethics protocol template: “During the entire duration of the study, all adverse events (AE) and all serious adverse events (SAEs) are collected, fully investigated and documented in source documents and case report forms (CRF).”</p>

Who		What	
Investigator	Sponsor	Detection	<p>Event Investigator</p> <p>Severity Investigator</p>
		Assessment/ Re-Evaluation	<p>Seriousness Investigator + Sponsor</p> <p>Causality Investigator + Sponsor</p> <p>Yes / No related, probably, possibly, unlikely, unrelated Other</p> <p>Related as a precaution ⇔ Missing a related event</p>
	SE-Applicant	Documentation	<p>Expectedness Sponsor</p> <p>Technical assessment IB / product information</p>
		Reporting	

Who		What	
Investigator	Sponsor	Detection	<p>Site documentation, 3 Levels:</p> <ul style="list-style-type: none"> • Source documents • Forms • CRF <p> in line!</p> <p><u>Sponsor oversight</u> Responsible for ongoing safety evaluation and overview of the trial</p> <ul style="list-style-type: none"> • Listing of all AEs & SAEs • Review the situation <ul style="list-style-type: none"> Per AE category, per site, etc. Assess changes to risk benefit ratio Document! (Statement, sign and file) • Interval: Risk profile <ul style="list-style-type: none"> Frequency of reported events
		Assessment/ Re-Evaluation	
	Documentation		
	SE-Applicant	Reporting	

Who		What	
Investigator		Detection	<p>Reporting...</p> <p>Patient safety and well-being</p>
	Sponsor	Assessment/ Re-Evaluation	<p>Ethics committee responsibility (ICH GCP): Ensure the protection of the rights, safety and well-being of human subjects involved in a trial</p>
		Documentation	<p>Swissmedic (home page): We consistently and rigorously apply the requirement to ensure the safety of therapeutic products</p>
	SE-Applicant	Reporting	<p>... to Ethics committees and Swissmedic</p>

Who		What	
Investigator	Sponsor	Detection	<p>Reporting...</p> <p>... to Ethics committees and Swissmedic</p> <p>Swissmedic:</p> <p>SUSARS safety and protective measures</p> <p>Swissethics:</p> <p>lethal SAEs SUSARS safety and protective measures</p> <p>And that's all!</p>
		Assessment/ Re-Evaluation	
		Documentation	
	SE-Applicant	Reporting	<p>+ once a year – Annual Safety Report</p> <p>YOU</p>

Questions & Answers

Discussion