



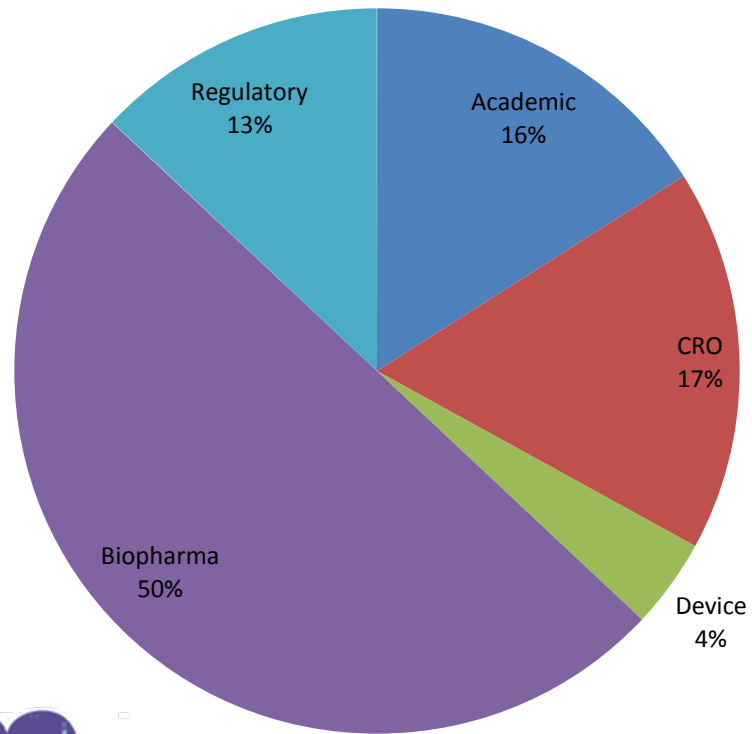
Setting the Stage

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Cardiac Safety Research Consortium

Welcome



Cardiac Safety Research Consortium

Disclosure



- Financial: I am an officer and stockholder of ACI Clinical* which specializes in EACs and DMCs and have served as a member of EACs on a fee for service basis.
- Intellectual: I believe the variability surrounding the establishment and operation of EACs distorts their usefulness, creates unnecessary expense and diverts sponsor and investigative site efforts.

* aka Applied Clinical Intelligence

TD Endpoint Adjudication



		Instant Replay Indicated			
		YES	PROBABLY	POSSIBLY	NO
Can Instant Replay be Performed Properly	5-13 years ago	X	X	X	
	?		X	X	
		X	X	X	



Current Situation		Instant Replay Indicated	
		YES	NO
Can Instant Replay be Performed Properly	YES	Scoring Play Final 2 minutes Legal Coach Request "Reviewable Play"	
	NO	Trained Booth Officials Right Number of Cameras	

CV Endpoint Adjudication



CURRENT SITUATION	Endpoint Adjudication Indicated			
	YES	PROBABLY	POSSIBLY	NO
Can Endpoint Adjudication Performed Properly	X	X	X	
?		X	X	
	X	X	X	

Need for an EAC

- Improve
 - Type of trial/ endpoints
 - Practicality
 - Retrospective vs Prospective

Role of Committee Member

- Independence
- Sponsor Interaction
- Qualifications
- Types of Adjudication
- Interaction

Information Rule

- Minimal quantity of information
- Minimal quality of information
- Impact of Endpoint CRFs
- New Information and Re-adjudication

Event Ascertainment Strategy

- Investigator Reported
- SMQ Trigger Events
- Safety Driven

Case Distribution Rule

- Number of voters
- Multiple therapeutic areas
- Multiple committees
- Majority vs Unanimity
- Investigator Vote

Coverage Rule

- Distribution
- Method
- Prospective 'adaptive' methodologies

Endpoint Definition

- EAC vs Endpoint Documents
- Primary vs Secondary
- Levels of Agreement
- Changes over time

Reliability and Quality Measures

- Voting Process Controls
- Reliability
- Integrity

Unclear Value Statement



- No Appropriate Usage Guidance
- No Taxonomy Of Adjudication
- No Framework For Risk-based Adjudication
- No Framework For Cost-efficiencies
- Burden On Sponsors
- Burden On Sites

Topics for ThinkTank



- Why Do We Need Adjudication
- Event Identification And Ascertainment Strategies
- Clinical Development And CV Endpoints— Potential Impact On Adjudication Strategies
- Practical Implications And Future Directions

Goal of the Think Tank



CURRENT SITUATION	Endpoint Adjudication Indicated			
	YES	PROBABLY	POSSIBLY	NO
Can Endpoint Adjudication Be Performed Properly	X	X	X	
?		X	X	
	X	X	X	



GOAL	Is Endpoint Adjudication Indicated		
	YES	MAYBE	NO
Can Endpoint Adjudication Be Performed Properly	YES	X XX	X
	MAYBE	X XX XX	
	NO	X X	XX

"If you can't describe what you are doing as a process, you don't know what you are doing."

W. Edwards Deming