RISK-BASED MONITORING: SEEING THE FOREST FOR THE TREES WHEN DESIGNING AND IMPLEMENTING AN ENTERPRISE PROCESS

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Topics Covered

Why embark on an RBM journey?

Starting with the end in mind
  • Process
  • Technology

Implementation examples & learnings

Value – what is the ‘return on RBM’?
Today’s Biggest RBM Risk

Key success factor: differentiating between enablers, elements and outputs – and solutions and objectives
WHY EMBARK ON AN RBM JOURNEY?
RBM: Coming to a Trial Near You – and Soon…

Current use of RBM solutions

Expected use of RBM solution in the next 5 years*

Expected use of RBM solution in the next 10 years*

- Increased quality expectations
- Development cost increases
- RBM process maturation
- Regulatory mandate

RBM will become the new normal

Source: Industry interviews commissioned by Covance. * Note: Not a quantitative forecast
Covance Definition of RBM

- Holistically identifies risk of failure
- Designs monitoring approaches to prevent or mitigate risk
- Differentiates critical vs. non-critical
- Delivers fit for purpose quality – flexible to study and regulatory need
- End to end trial conduct transformation
- An extension to smart study design

RBM is a logical next step to enhance efficiency and quality
Covance RBM: Transformation – and Continuity
The Journey to Develop RMB Core Expertise and Systems

- Predictive, proactive and preventative operational platform established
- EMA Draft Reflections Paper and FDA Draft Guidance on RBM
- FDA Final Guidance on RBM
- TransCelerate Updated Paper on RBM

Various academic research model studies e.g. more than 8,000 patients targeted monitoring

More than 8,000 patients cardiovascular trial
- Reduced source data verification analytics: data action reports targeted & reduced monitoring

New site quality risk reports
Expanded informatics capability
New clinical quality control process

2005
2009
2010
2011
2012
2013
2014

Launch of Covance RBM dashboard and process

Strategic partner portfolio
‘Fit-for-purpose’ monitoring tailored monitoring & SDV in multiple therapy areas and studies

More than 12,000 patients
Full RRM implementation incl. tSDV web-based analytics solution
Governance: Defining the Team to Get Us There

Core Development Team

Functional Teams

- Process & Documents
- IT & Technical Requirements
- Organization Design
- Commercial

Business-Led Transformation Team
PROCESS:
TYING IT ALL TOGETHER
OR -
STARTING WITH THE END
IN MIND
Start with the End in Mind

Design & Delivery: Fit for purpose throughout the continuum
RBM – If You Start with Monitoring it’s Too Late

Starts with designing out risk: the protocol is the blueprint for quality
Start with the end in mind: reliable data withstanding Agency scrutiny
# Translating Risk into Action

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<th>Examples</th>
<th>Management Approaches</th>
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<tr>
<td><strong>Intrinsic Risks</strong></td>
<td>• Average on-site visit interval</td>
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<td>• Maturity of safety profile</td>
<td>• Source Data Verification level</td>
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<td>• Phase of development</td>
<td>• Proportion remote/on site visits</td>
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<td><strong>Design Risks</strong></td>
<td>• Quality by design improvements</td>
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<td>• Clarity of protocol</td>
<td>• Central data surveillance</td>
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<td>• Primary endpoint approach</td>
<td>• Supplementary training (Site &amp; CRA)</td>
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<td>• Fit vs. standard of care</td>
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<td><strong>In-Flight Risks</strong></td>
<td>• Adjusted site-specific intervention levels</td>
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<td>• Site experience</td>
<td>• Turnover management &amp; training plans</td>
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<tr>
<td>• Protocol deviation rate</td>
<td>• Issue management plans</td>
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<td>• Investigator oversight</td>
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Appropriate risk management: Not ‘one size fits all’
Full Portfolio of Actionable Tools

- Risk Assessment Tools
- RBM Templates
- Control Charts
- Risk Libraries
- Site Risk Dashboards
- Process Integrations
- Training Materials
- Process Manuals
- Functional Process Tools
- Web-Based Visualizations
- Clinical Data Repository
- RBM Teams

Implementation-Ready, Tried and Tested
TECHNOLOGY: A KEY ENABLER
Effective RBM is Multifaceted – Technology is Not the Be All And End All
IMPLEMENTATION: EXAMPLES & LEARNINGS
Overcoming Functional Barriers

Functional View

• “RBM is all about clinical cost reduction”
• “Quality will suffer on variables not monitored” – risk prevention not considered
• “There is nothing wrong with how we’ve done things before”
• ‘Burning platform’ challenge is only for Clinical Operations

Holistic View

• Quality of critical data & processes enhanced through preferential focus
• Cost saving is a consequence of monitoring what matters
• Focus on Quality by Design and Risk, not SDV reduction
• Areas reduced are not critical from an outcome perspective

Focused change management
Experiences from Implementation

**Example #1**
- Cardiovascular outcomes study
- Full RBM Design
- tSDV implemented
- Ongoing, enrolling

**Example #2**
- Phase III Pivotal Study
- Reduced SDV
- Targeted RMV strategy
- NDA/MAA approved

**Example #3**
- Partner portfolio
- ‘Fit-for-purpose’ monitoring
- Tailored RMV/SDV
- Multiple studies and TAs
Implementation Learnings

**Sponsor**
- Variable needs and priorities
- Tailored solution needed, consistent but flexible

**Site**
- Managing monitor support expectations
- Expectation-setting and alternative methodologies

**Monitors**
- Resource management
- Change management
- Access to information
VALUE:
WHAT IS THE ‘RETURN ON RBM’?
RBM Value, It’s All In The Design

- Protocol simplification
- Quality-by-design
- Risk-based site selection
- Systematic mitigation
- Centralized data surveillance
- Risk-targeted on-site review

Quality and Efficiency go Hand-in-Hand
RBM can transform risks into returns

- Effective, adaptive resources allocated targeting maximum impact and value
- Proactive planning and focus on primary quality and state of control
- Efficiency gains a consequence of fitness-for-purpose
For more information please visit
www.covance.com/RBM