CENTRALISED MONITORING OF DATA

Anna Lindsjö, Specialist, Clinical Data and Standards Management

15th June 2016, Phuse Copenhagen

DISCLAIMER: The views and opinions expressed in this presentation are those of the author and are not necessarily those of H. Lundbeck A/S

AGENDA

Fast Track to Approval: Speed and Efficiency

- Why start to implement a risk-based approach
- Risk-based approach and centralised monitoring
- What have we done so far in Lundbeck
- Examples of metrics and centralised monitoring
- Next steps
- Summary

What is a risk-based approach?

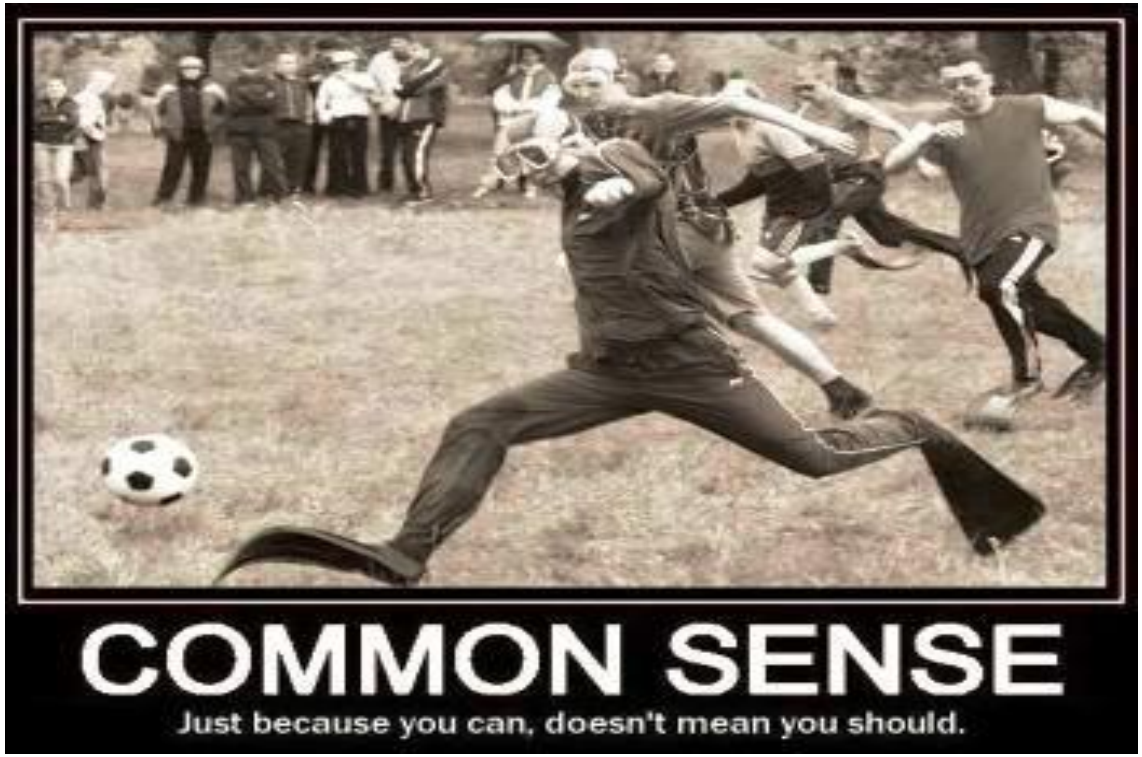
Targeted?

Planned?

Adapted?

Intelligent?

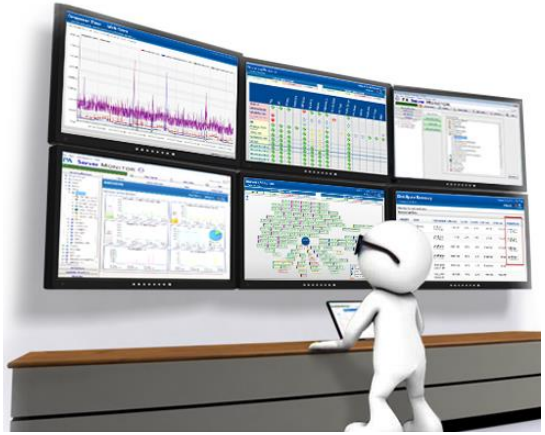
Focused?



Why start to implement a risk-based approach?

- Regulatory guidance
- Change focus from quantity to quality
- Increased number and more complex clinical trials
- Increased costs

Definition of Centralised Monitoring 1(2)



ICH E6 R2 draft version,

Centralised monitoring

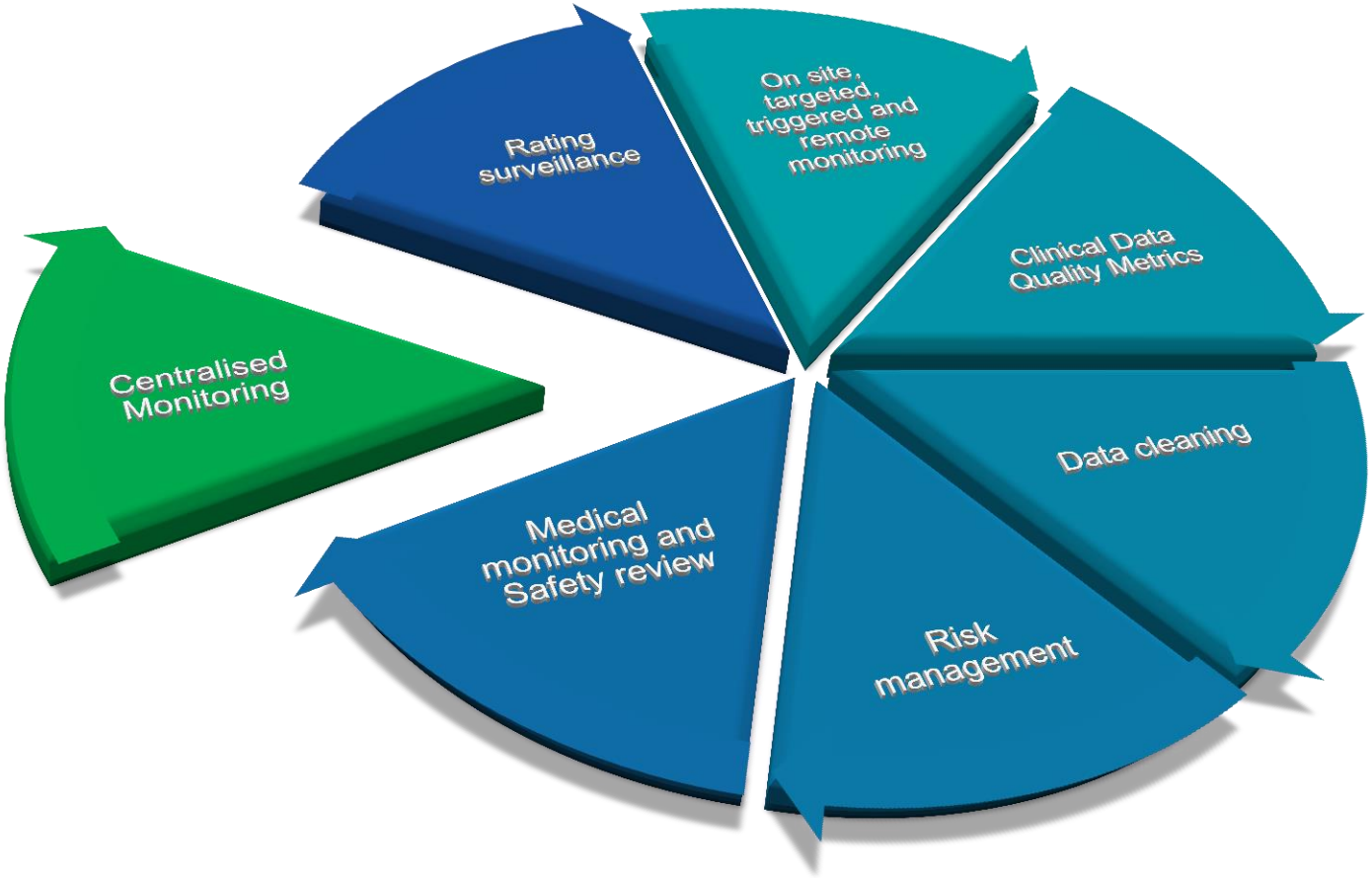
"is a remote evaluation of ongoing and/or cumulative data collected from trial sites, in a timely manner"

Definition of Centralised Monitoring 2(2)

- Support the routine monitoring (targeted approach)
- Routine review of data
- Review of:
  - missing and inconsistent data
  - outliers or unexpected lack of variability
  - protocol deviations
  - etc.
- Site performance, within or across sites

How will this be better?

Combination of automated, analytical and visualization tools and processes



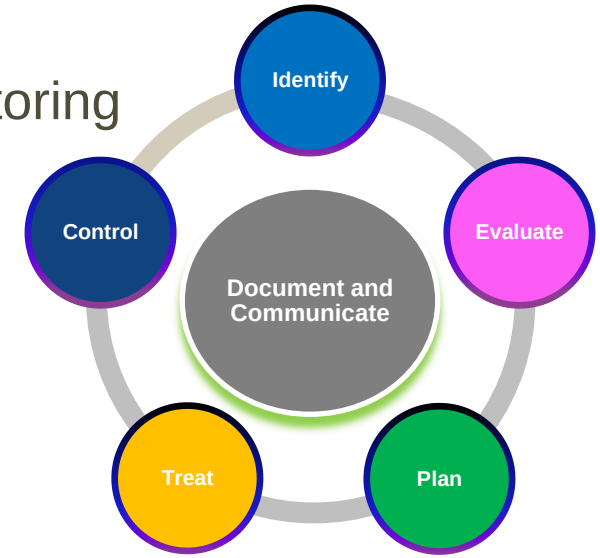
What have we done so far at Lundbeck? 1(2)

- Identification of risks, triggers and creating of action plans
- Identification of critical data variables and processes

- Communication between team, CRO, site

- On-site, targeted, triggered and remote monitoring

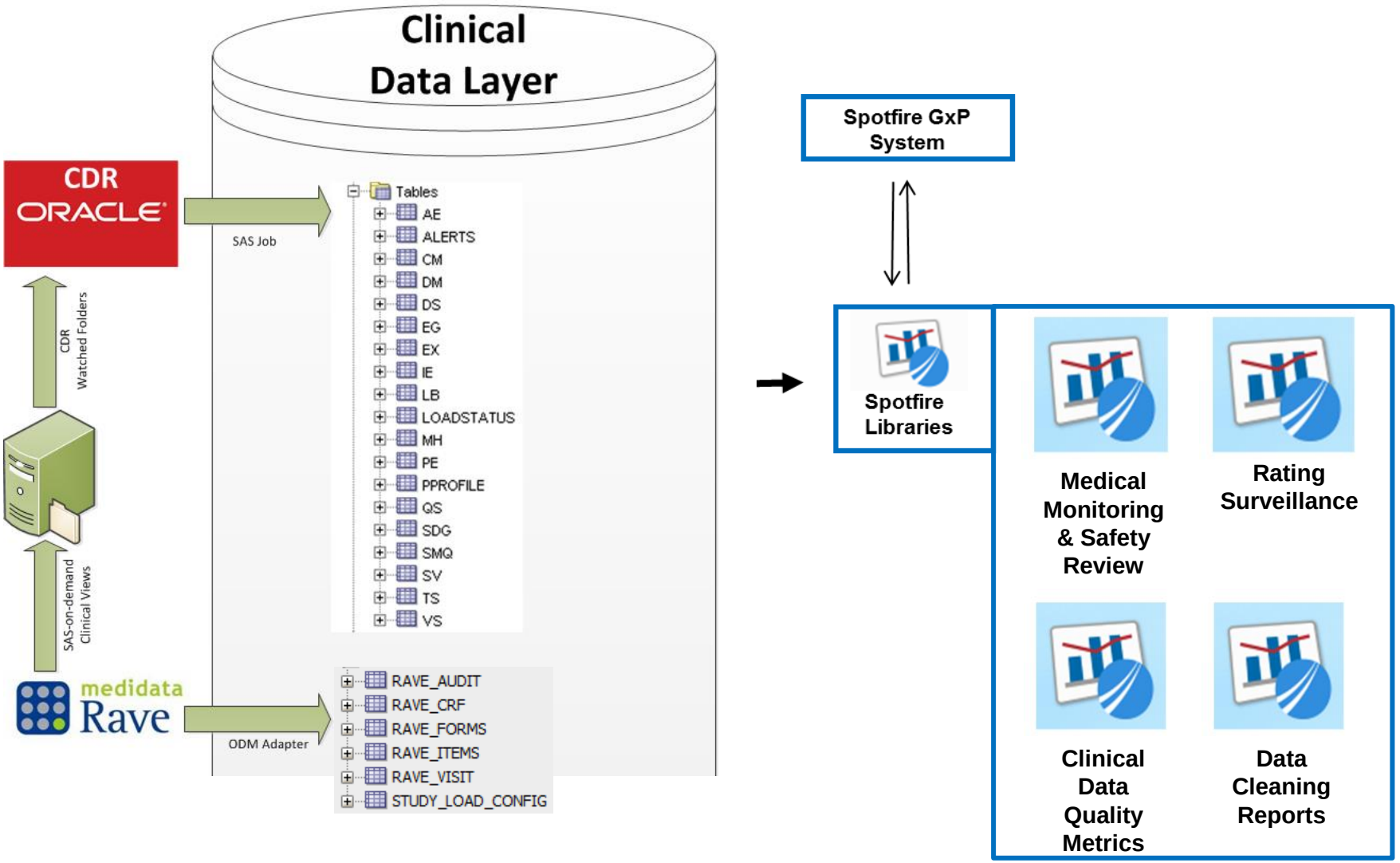
- Regular meetings –
  assess safety and quality trends



Risk management process

What have we done so far at Lundbeck? 2(2)

- Review of clinical data quality metrics
- Medical monitoring and safety review
- Data cleaning activities
- **Centralised monitoring**


Clinical
Data Layer
CDR
ORACLE
CDR
Watched Folders
SAS-on-demand
Clinical Views
medidata
Rave
SAS Job
ODM Adapter
Tables
AE
ALERTS
CM
DM
DS
EG
EX
IE
LB
LOADSTATUS
MH
PE
PPROFILE
QS
SDG
SMQ
SV
TS
VS
RAVE_AUDIT
RAVE_CRF
RAVE_FORMS
RAVE_ITEMS
RAVE_VISIT
STUDY_LOAD_CONFIG
Spotfire GxP
System
Spotfire
Libraries
Medical
Monitoring
& Safety
Review
Rating
Surveillance
Clinical
Data
Quality
Metrics
Data
Cleaning
Reports

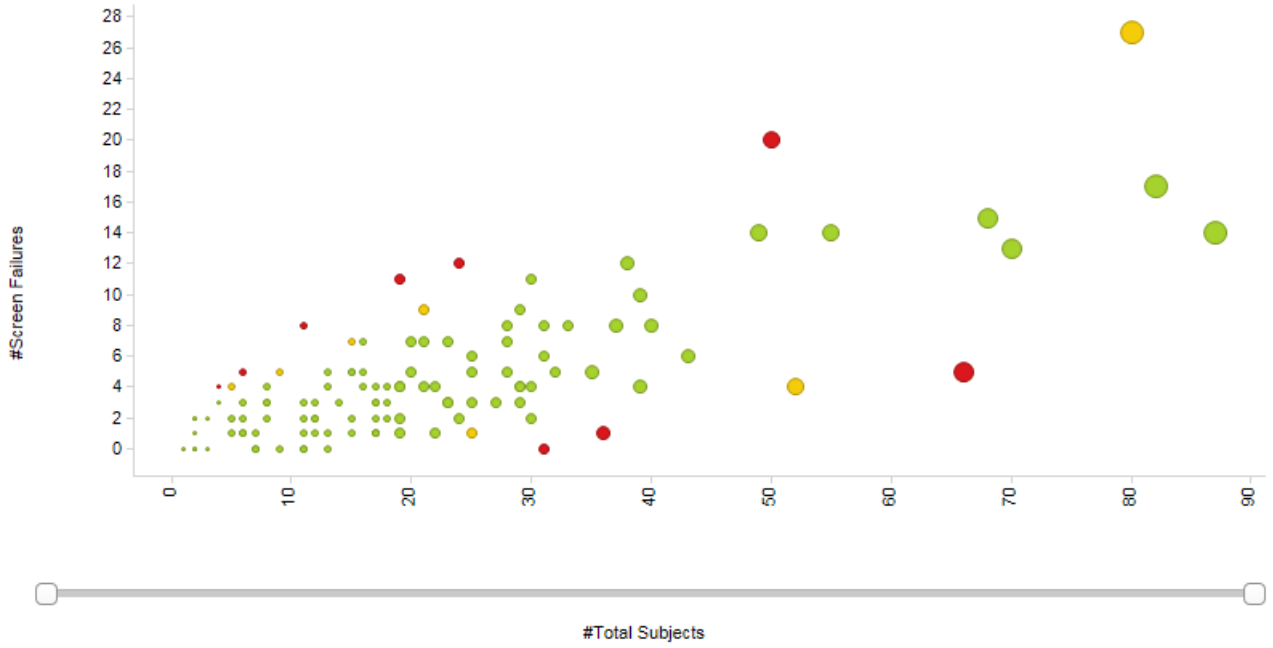
Risk Indicators – Screening Failure Rates

Risk Indicators

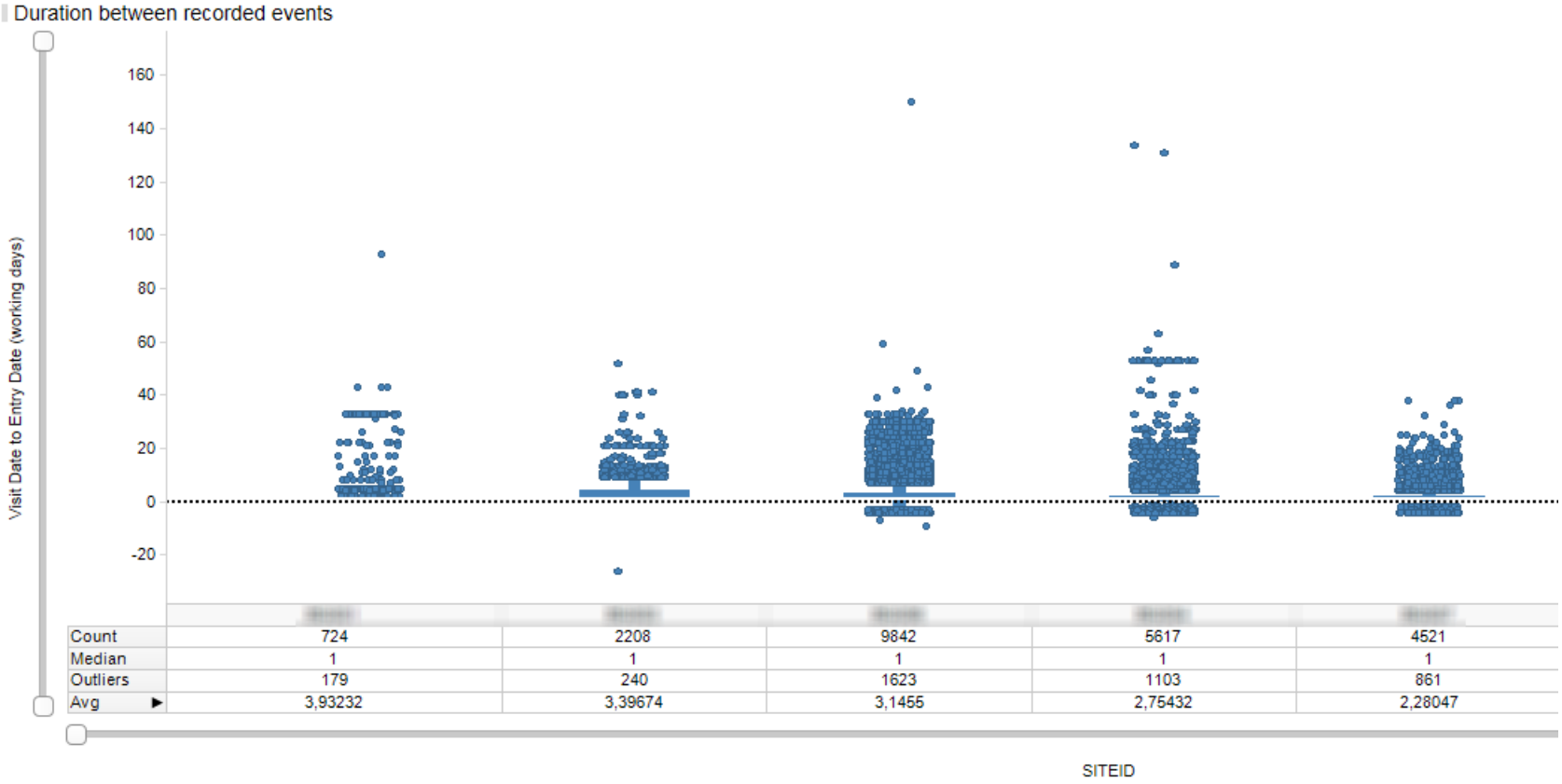
SITEID	Overall Risk Indicator Level	Site AE Rate (per PatientYears)	Site SAE Rate (per PatientYears)	Site CM Rate (per PatientYears)	Site MH Rate (per PatientYears)	Site Withdrawal Rate (per PatientYears)	Site Screening Failure Rate (per Total Subjects)
[redacted]	HIGH	3,77	0,00	1,80	2,60	0,27	0,00
[redacted]	HIGH	2,76	0,00	0,50	2,32	0,06	0,11
[redacted]	HIGH	1,06	0,00	0,23	2,87	0,35	0,22
[redacted]	HIGH	2,52	0,00	0,48	2,62	0,13	0,08
[redacted]	HIGH	3,44	0,00	0,19	3,25	0,32	0,04
[redacted]	HIGH	2,66					
[redacted]	HIGH	1,46					
[redacted]	HIGH	1,37					
[redacted]	HIGH	8,34					
[redacted]	HIGH	1,51					
[redacted]	HIGH	30,42					
[redacted]	HIGH	1,87					
[redacted]	HIGH	3,61					
[redacted]	HIGH	12,21					
[redacted]	HIGH	5,82					
[redacted]	HIGH	2,59					
[redacted]	HIGH	15,46					

Screening Failures

Size is based on the number of subjects at site, and color is based on the risk level.

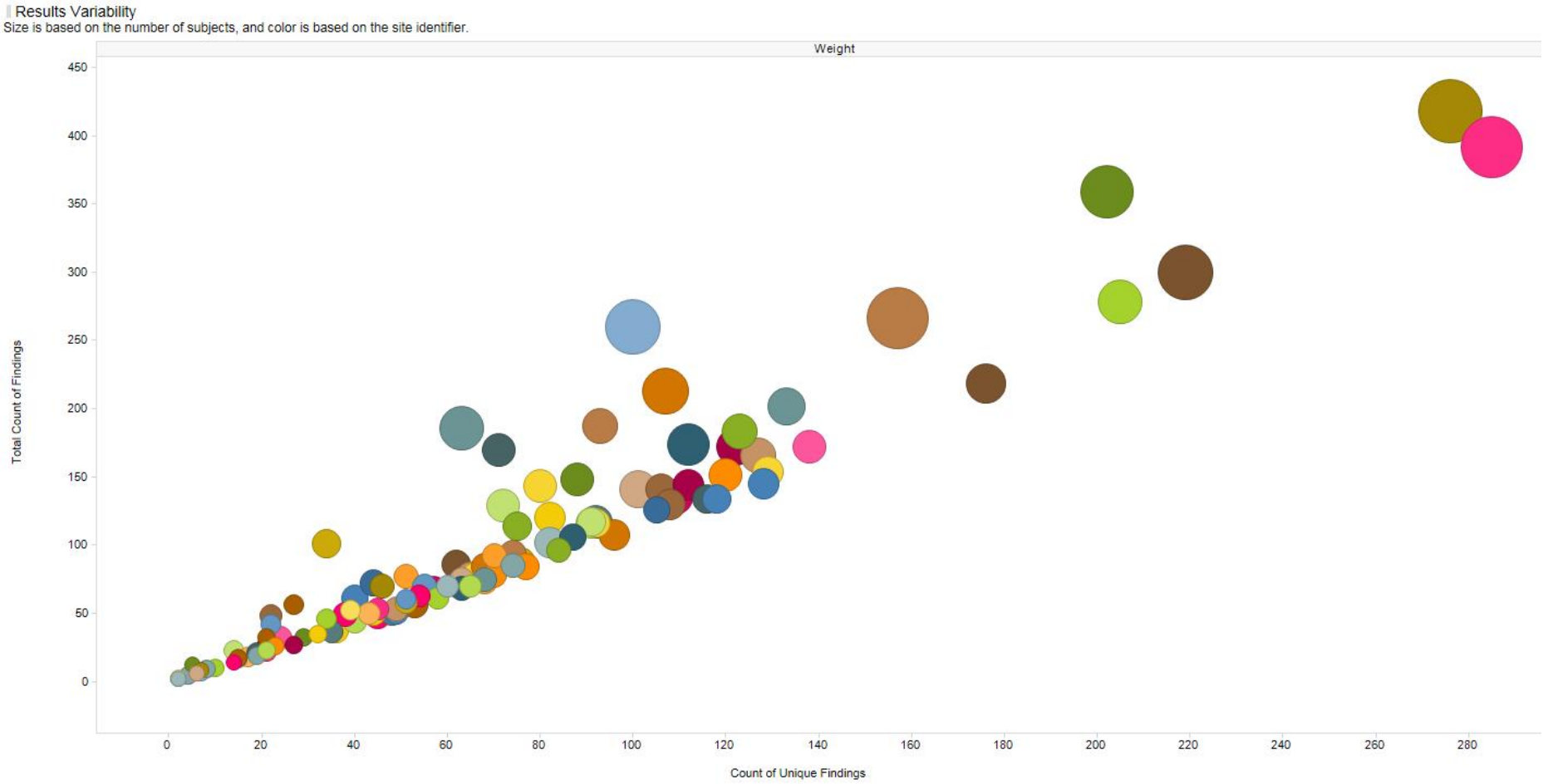


Time from visit to data entry

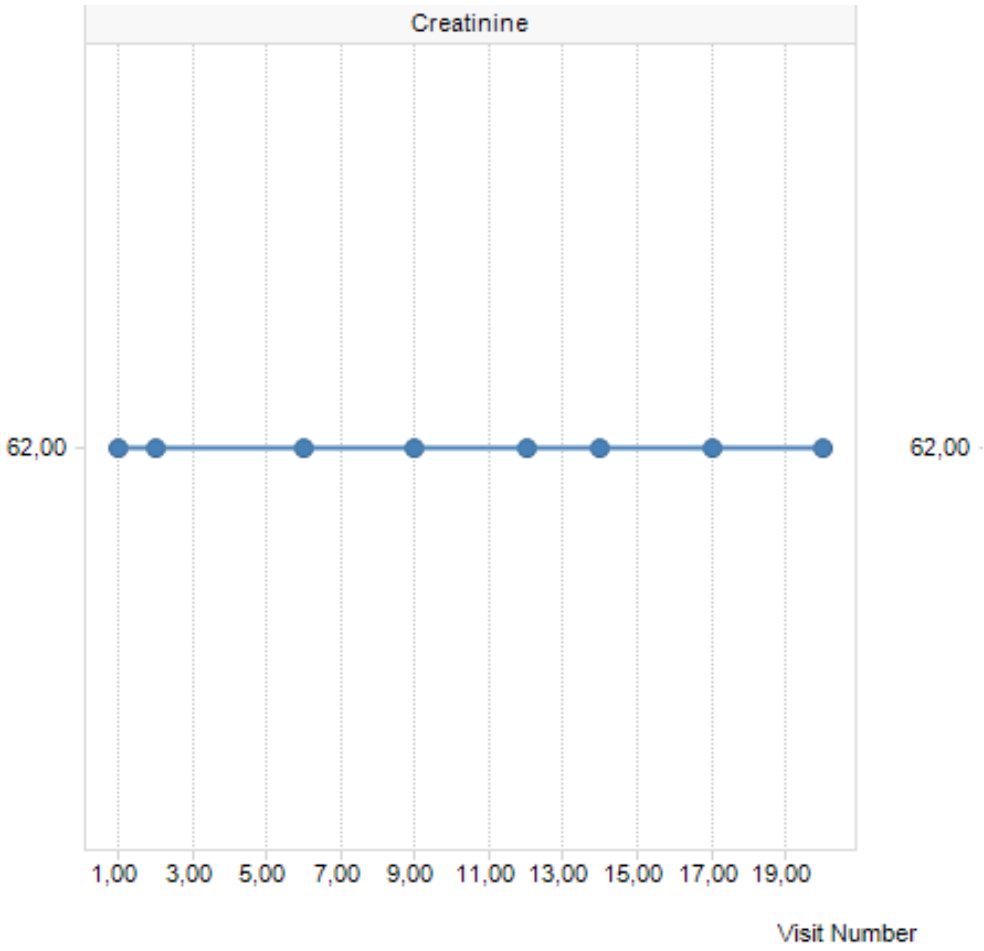


Count	724	2208	9842	5617	4521
Median	1	1	1	1	1
Outliers	179	240	1623	1103	861
Avg ▶	3,93232	3,39674	3,1455	2,75432	2,28047

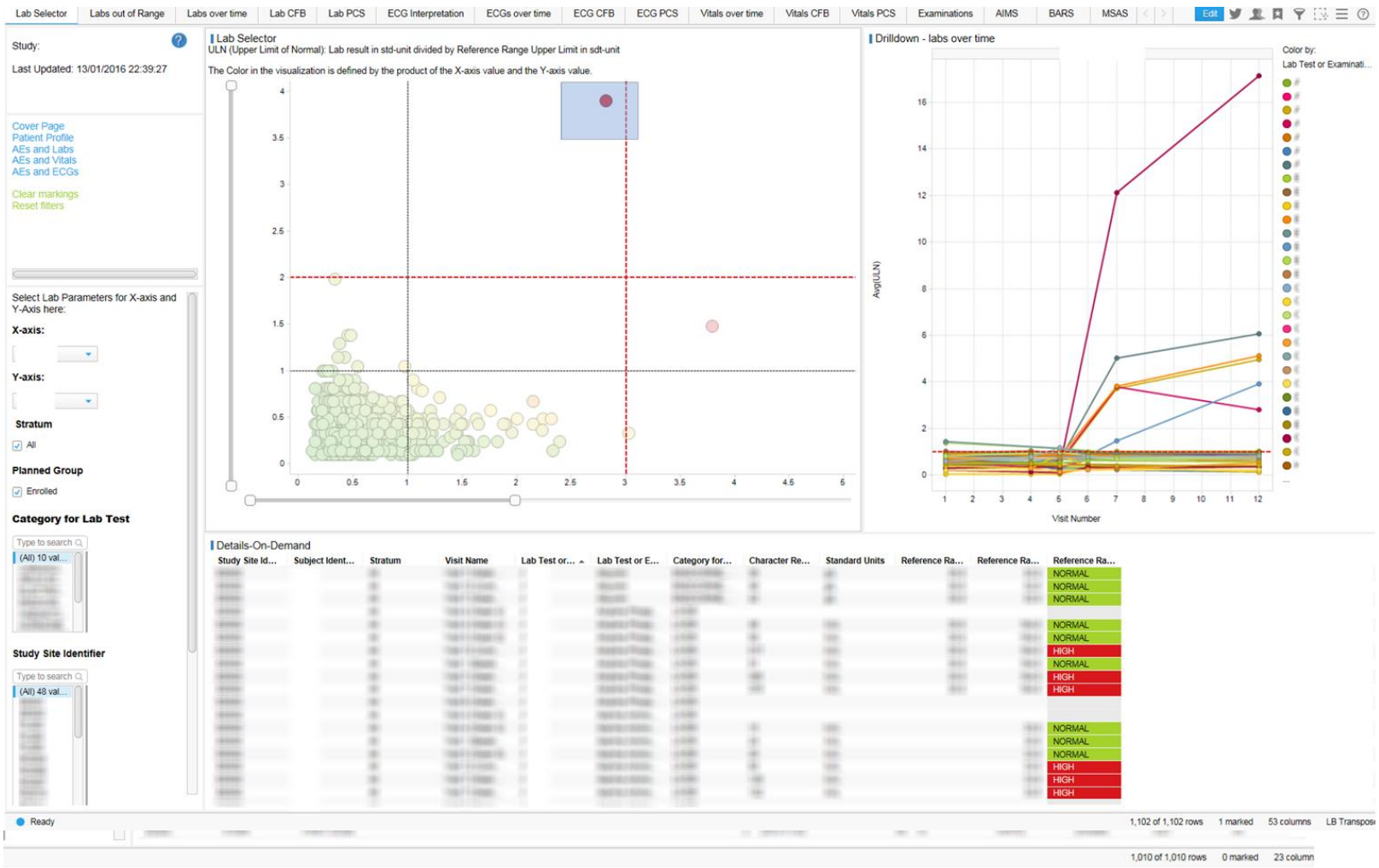
Results Variability - Weight



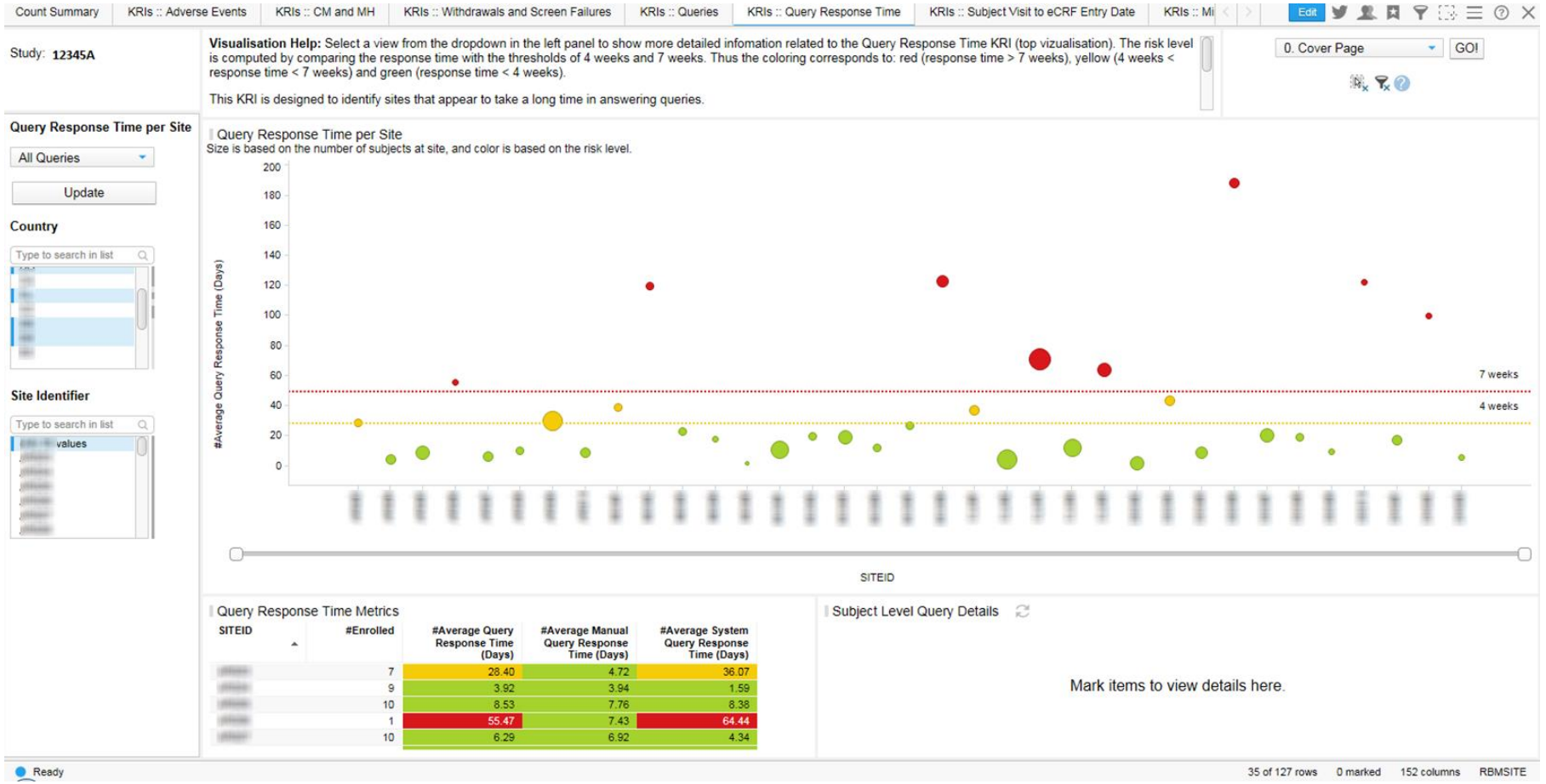
Site Quality – Value propagation Example



Medical Monitoring and Safety Review (MMSR)



Clinical Data Quality Metrics (CDQM)



Next steps

Continue to improve the risk-based process including centralised monitoring and adapt to ICH E6 (R2):

- Develop standard tool/tracker for documenting & tracking discussion/actions taken throughout process
- Further develop tools to assist in monitoring data quality
- Use Spotfire to visualize data based on the Rave clinical views and audit trail as a source

Summary

Risk-based approach and centralised monitoring

Fast Track to Approval: Speed and Efficiency

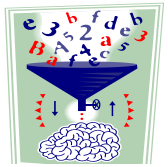
Authorities support

Holistic view across whole process

Quality –

”fit for purpose”

Proactive, targeted risk based approach



Centralised monitoring - Improve data quality, integrity

Greater and focused teamwork

Spent on “right” tasks

Questions

- alen@lundbeck.com