



2023
CDISC TMF
INTERCHANGE

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The TMF Trinity and How Simple is Best

Presented by Jake McDonald and Rob Jones of Phlexglobal

Meet the Speakers

Jake McDonald

Title: Manager of Business Analysis & Solutions

Organization: Phlexglobal

Has spent almost 9 years in the TMF world, starting in processing and moving to the solutions side in 2018



Rob Jones

Title: Director of Professional Services

Organization: Phlexglobal

Has been in the pharmaceutical and drug development industry for almost 15 years, starting off as one of those lab nerds before moving up to the cool kids table and TMF





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PHLEXGLOBAL
A ▶ PHARMALEX Company

- Established in 1997
- Life Sciences Industry Focus
- Software Solutions & Expert Services

▶ **PHARMALEX**

- A PharmaLex Group company with over 2,300 employees, 53 offices in 25 countries and more than 1,000 satisfied clients worldwide



- 10,000 users
- > 500 studies executed
- >7M documents, >30M pages processed
- > 30,000 products managed
- > 4000 submissions published



- Pharmaceuticals
- CRO's
- Biotech's
- Generics and API
- Medical Devices

Three is a magic number

All Good
Things Come
In Three

Three is a magic number



All Good
Things Come
In Three

Three is a magic number



All Good
Things Come
In Three



Three is a magic number



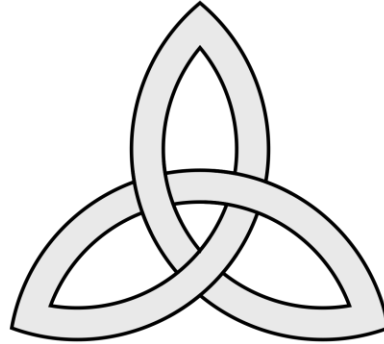
All Good
Things Come
In Three



Three Lions



A measurement of what is expected versus what is there



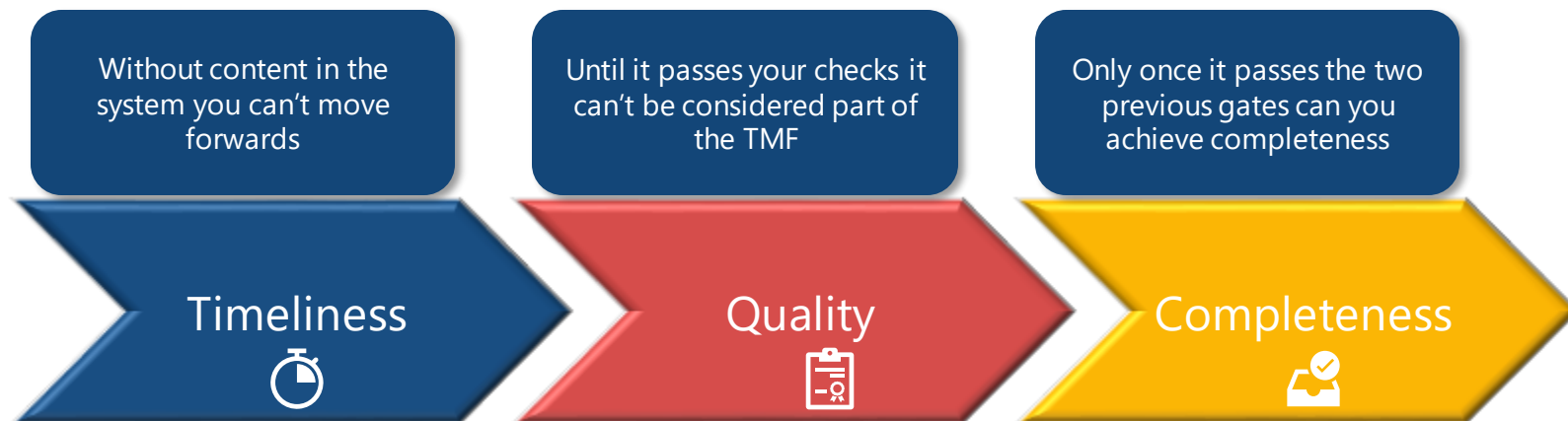
The pass rate of each document entering the system



The time taken from the content ready to file to available within the TMF



ABC, easy as 123





KISS

Keep It Simple Stupid

Time Warp

- Allows you to measure the efficiency of teams and individuals with submission to the TMF
- Timeliness is often measured from the Document Date to the Date of Final within your system
 - There are some documents that are removed from this such as CVs, Licenses etc. as their document date is often further in the past
- Standard Measurement
 - Number of documents available within the agreed time period vs those that were not



Quality Street

- Gives you insight into the effective classification and metadata assignment to content
- It is NOT just a quality check of content, this should be performed by the Document Owner prior to upload e.g. is the Protocol suitable for this study?
 - You will check for missing signatures and blank pages, standard GDP
- Standard measurement:
 - Number of uploads that passed first time versus those that did not



QUALITY



DURABLE

Complete Mess

- Allows you to confirm that you have everything you need within your TMF
- This can only be done by having an accurate idea of what is expected within the TMF, placeholders available
- The most accurate Completeness is measured against the current milestone of the study to remove content not yet required
- Standard Measurement:
 - Calculation of what is available in a Final state in the TMF versus the expected content



Count On Me

Timeliness

If $\geq 85\%$	For Timeliness to be Green for the previous month, at least 85% of documents submitted must be Live in the TMF within 35 days of the Document Date.
If 75-84%	If 75-84% of documents are filed within 35 days of the Document Date, Timeliness drops to Amber and needs improvement.
If $< 75\%$	If less than 75% of documents submitted were within 35 days, Timeliness is Red and an Action Plan must be agreed for future submissions.

Quality

If $\geq 90\%$	For a study to be considered in the green, 90% of documents must pass QC first time, without a Query being raised.
If 82-89%	If Quality is between 82-89%, it is considered Amber and needs improvement.
If $< 82\%$	If fewer than 82% of documents pass QC first time, Quality is in the Red and further training may be required.

Completeness

If $\geq 90\%$	For a study to be considered in the green, 90% of documents expected to-date should be Final.
If 83-89%	If completeness is between 83-89%, it is considered Amber and needs improvement.
If $< 83\%$	When completeness falls below 83%, the study is not considered compliant and immediate action should be taken.



Dig a Little Deeper

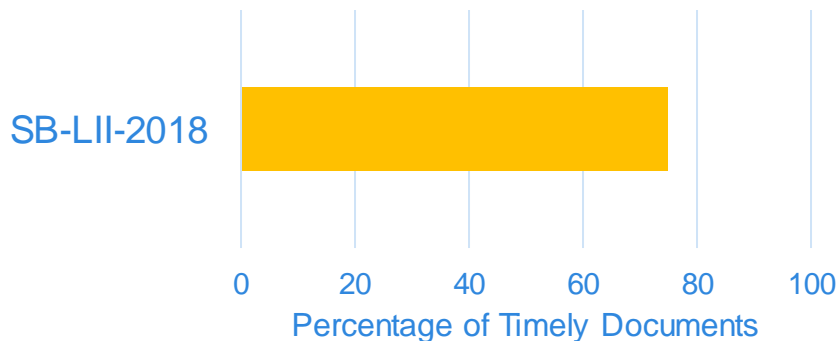
Too Little Too Late

- Timeliness going into the Amber or Red can alert you to a problem
- It can be a trigger to delve further
 - Certain Functions or Sites that are struggling
 - Artifacts that are dragging down the metrics



Smoke from a Distant Fire

Study Timeliness



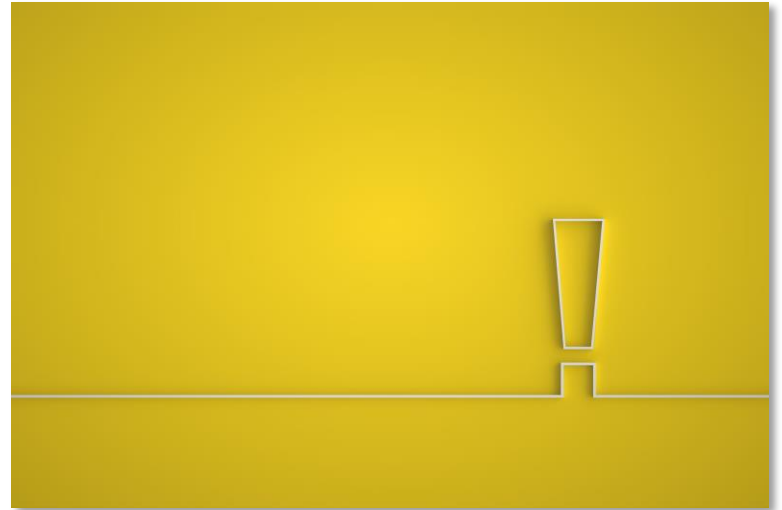
High level timeliness metrics only help you smell the smoke

Oversight and analysis needs to be done at multiple levels to see where the fire is

Protocol	Country	Site	Percentage of Timely Docs
SB-LII-2018	United States	100 - Smith	91
SB-LII-2018	United States	101 - Jacobs	95
SB-LII-2018	United States	102 - Matthews	80
SB-LII-2018	United States	103 - Hewlett	82
SB-LII-2018	United States	104 - Brady	68
SB-LII-2018	United States	105 - Belichick	52

Issues

- Perhaps your Quality is dropping every month
- This can help guide you
 - Is there a set of artifacts causing the issue?
 - Is retraining needed with specific groups?
 - What can be done to remediate before it becomes an issue



Step Back in Time

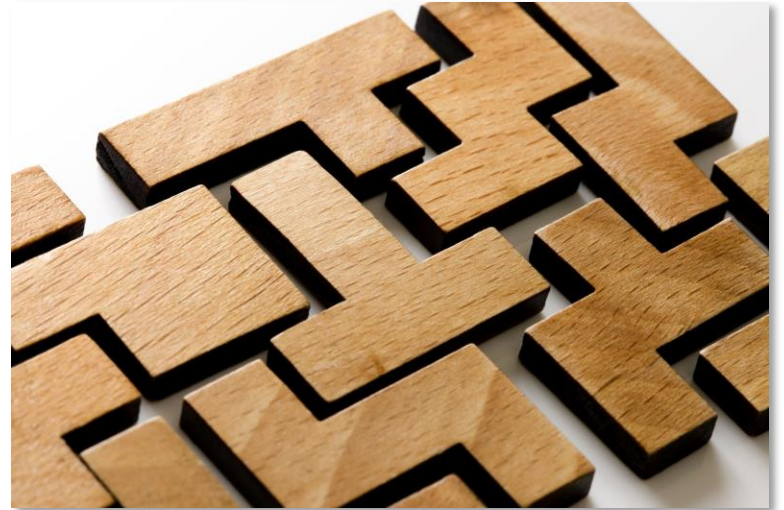
<u>Quality Percentage Over Time</u>			Jan-2023	Feb-2023	Mar-2023
Protocol	Country	Site			
SB-LII-2018	United States	100 - Smith	95	91	91
SB-LII-2018	United States	101 - Jacobs	95	95	96
SB-LII-2018	United States	102 - Matthews	80	84	68
SB-LII-2018	United States	103 - Hewlett	82	93	89
SB-LII-2018	United States	104 - Brady	68	82	87
SB-LII-2018	United States	105 - Belichick	71	73	73

- Reviewing quality trends mitigates the need to look at every site every month
- Look at quality over time to identify the problem areas



You Complete Me

- Is Completeness slipping away?
Now you can delve into why
- Completeness is often a complex topic
 - Maybe you need more expected documents
 - Maybe you actually need less
 - Or perhaps it's just simply your dates are wrong



Piece by Piece

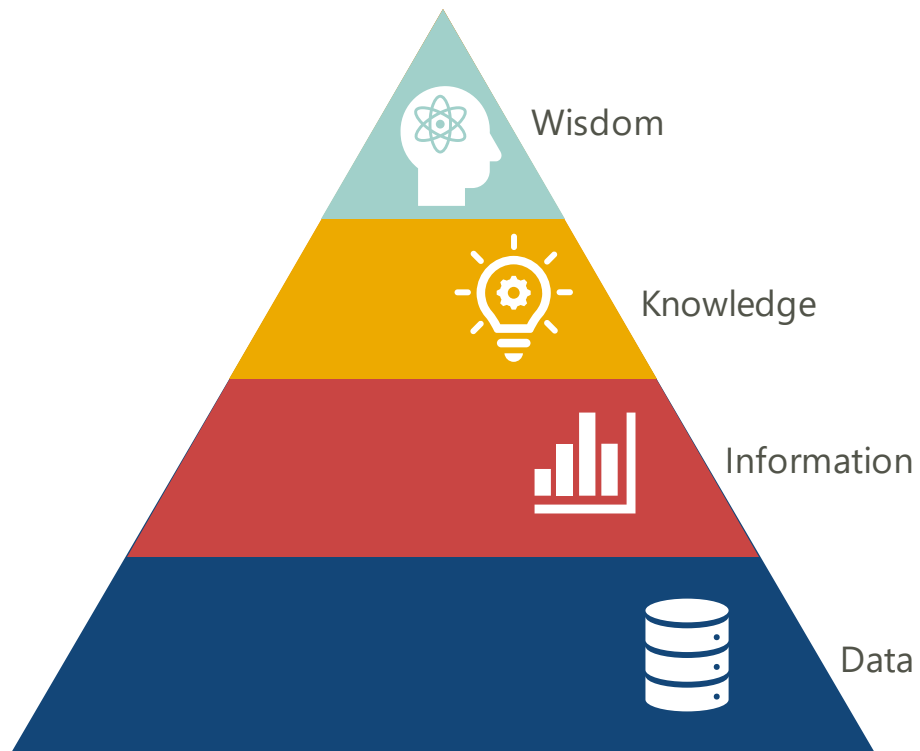
Document TMF Index

Artifact	# of Expected Documents	SB-LII-2018 100 - Smith	SB-LII-2018 101 - Jacobs	SB-LII-2018 102 - Matthews	SB-LII-2018 103 - Hewlett	SB-LII-2018 104 - Brady	SB-LII-2018 105 - Belichick
02.02.03 Informed Consent Form	3	3	2	3	3	3	2
04.01.01 IRB or IEC Submission	1	1	1	1	1	1	1
04.01.02 IRB or IEC Decision	2	1	1	1	1	1	1
05.02.03 Protocol Amendment Signature Page	3	3	3	3	3	3	3
05.03.01 Site Initiation Visit Monitoring Report	3	0	0	0	0	0	0
05.03.03 Site Evidence of Training	6	10	10	10	11	9	10

- Requirements don't just show you what documents are missing
- Sometimes they show you that your requirements may be wrong!



Pyramid



The Reason

The quality by design approach to clinical research (Section 3.1) involves focusing on **critical to quality factors** to ensure the protection of the rights, safety, and wellbeing of study participants, the generation of reliable and meaningful results, and the management of risks to those factors using a risk-proportionate approach (Section 3.2). The approach is supported by the establishment of an appropriate framework for the **identification and review of critical to quality factors** (Section 3.3) at the time of design and planning of the study, and throughout its conduct, analysis, and reporting

ICH E8 (R1) §3 DESIGNING QUALITY INTO CLINICAL STUDIES



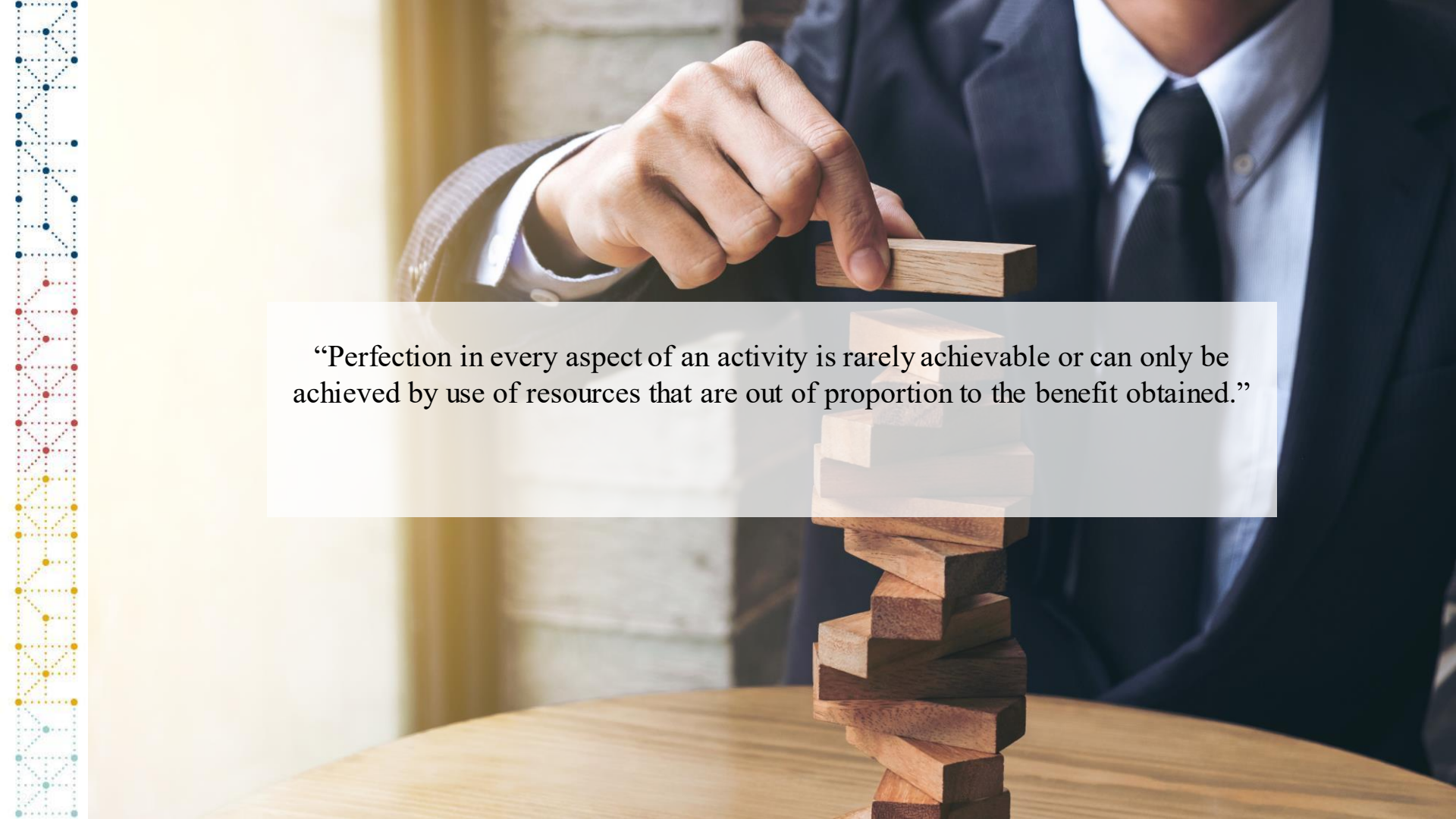
The Reason

The quality by design approach to clinical research (Section 3.1) involves focusing on **critical to quality factors** to ensure the protection of the rights, safety, and wellbeing of study participants, the generation of reliable and meaningful results, and the management of risks to the study. The quality by design approach is an appropriate and review of (Section 3.3) at the time of design and planning of the study, and throughout its conduct, analysis, and reporting.

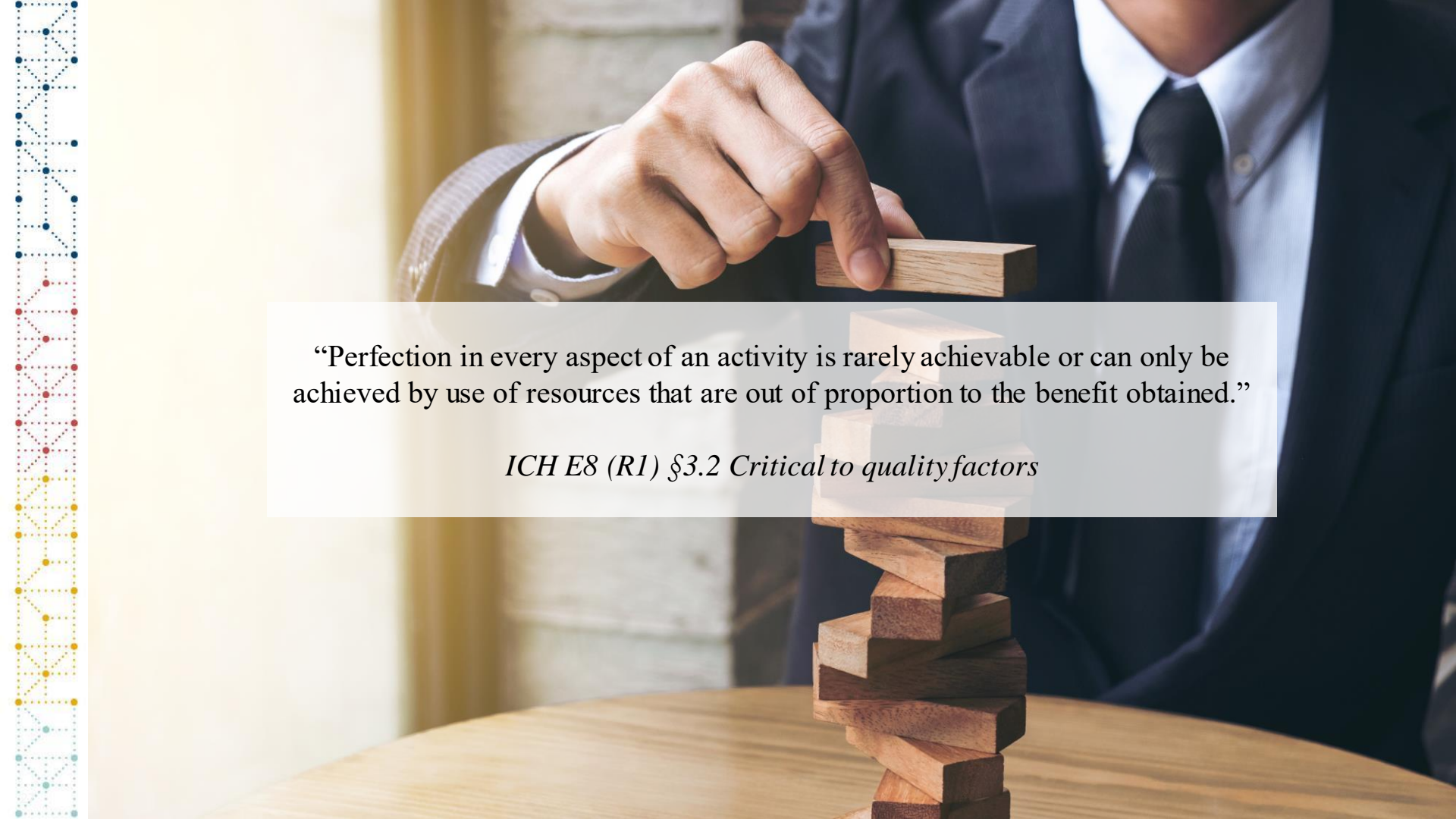
ICH E8 (R1) §3 DESIGNING QUALITY INTO CLINICAL STUDIES

Don't just measure, measure what matters



A person in a dark suit and tie is shown from the chest down, leaning over a round wooden table. They are carefully balancing a single wooden block on top of a tall, wobbly stack of approximately 15 wooden blocks. The stack is precariously balanced, with each block slightly offset from the one below it. The background is softly blurred, showing what appears to be a window with light coming through. On the far left edge of the image, there is a vertical decorative border consisting of a series of colored dots (blue, red, yellow, green) connected by thin lines, forming a grid-like pattern.

“Perfection in every aspect of an activity is rarely achievable or can only be achieved by use of resources that are out of proportion to the benefit obtained.”

A person in a dark suit and tie is shown from the chest down, balancing a single wooden block on top of a tall, precarious stack of other wooden blocks. The stack is on a light-colored wooden table. The background is softly blurred, showing what appears to be a window with light coming through. A semi-transparent white box is overlaid on the center of the image, containing text.

“Perfection in every aspect of an activity is rarely achievable or can only be achieved by use of resources that are out of proportion to the benefit obtained.”

ICH E8 (R1) §3.2 Critical to quality factors



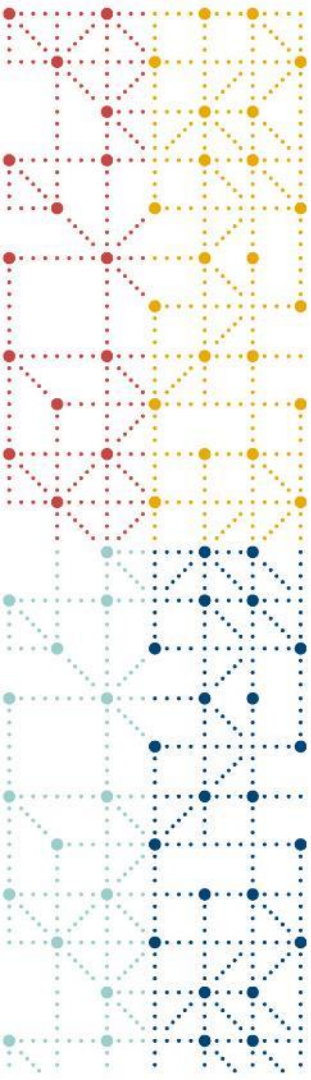
The TMF Trinity

1. Three is the Magic Number – Bob Dorough
2. Three Lions - Baddiel, Skinner & Lightning Seeds
3. ABC, easy as 123 – The Jackson 5
4. Kiss - Prince
5. Time Warp – Rocky Horror Picture Show
6. Quality Street – Van Morrison
7. Complete Mess – 5 Seconds of Summer
8. Count On Me – Bruno Mars
9. Dig a Little Deeper – The Princess and the Frog
10. Too Little Too Late – JoJo
11. Smoke From a Distant Fire – Sanford-Townsend
12. Issues – The Saturdays
13. Step Back in Time – Kylie Minogue
14. You Complete Me – Keyshia Cole
15. Piece by Piece – Kelly Clarkson
16. Pyramid - ALYSS
17. The Reason - Hoobastank

The TMF Trinity



Now On Spotify



Thank You!

cdisc