

Steve Hamburg



Considerations for Implementing CDISC ARS: Balancing Flexibility and Standards



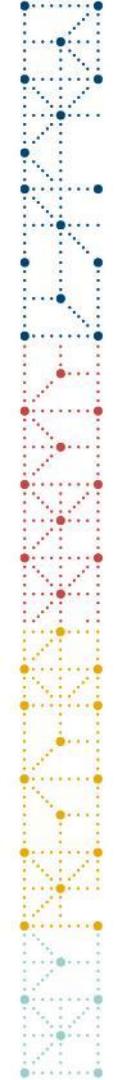
Meet the Speakers

Steve Hamburg

Title: Director, Statistical Programming • Head of Clinical Analytics & Reporting

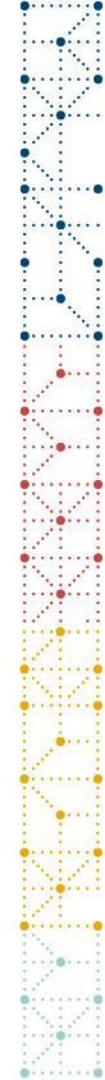
Organization: Jazz Pharmaceuticals

Steve Hamburg is the Head of Clinical Analytics and Reporting at Jazz Pharmaceuticals and has over 20 years of industry experience leading clinical trial reporting and CDISC standards implementation. He spent several years contributing to the CDISC ADaM team as well as various sub-teams, and has recently led the development of multiple AI solutions for automating statistical programming workflows and gleaning insights from analysis results.



Disclaimer

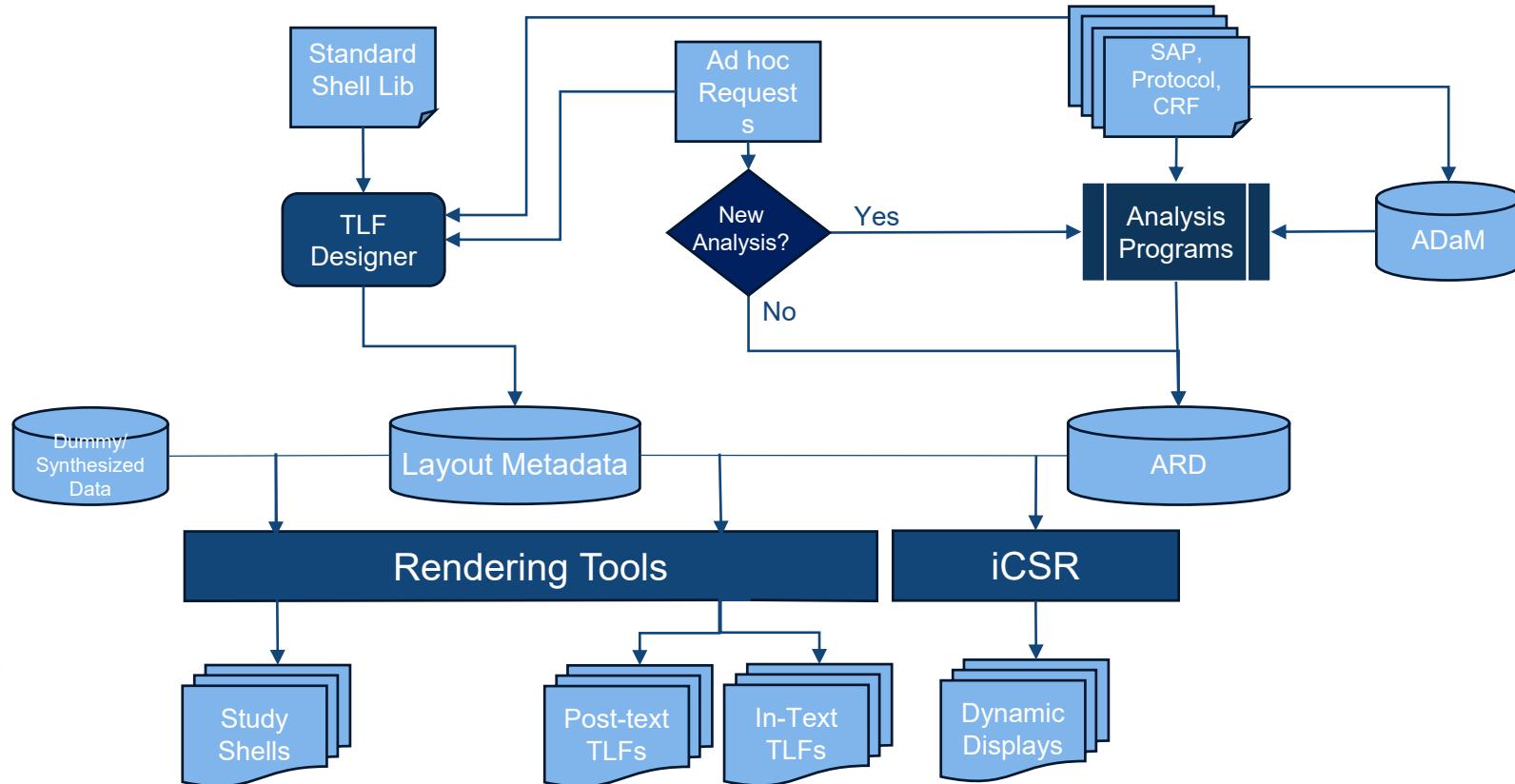
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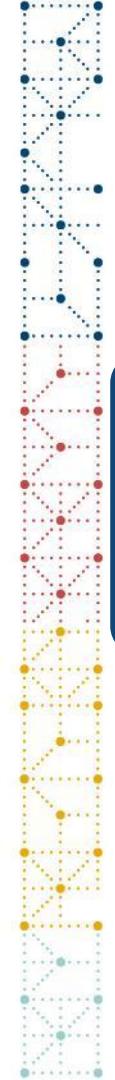


Pre-ARS vision

- All Analysis Results live in a single repository
- Robust metadata contain labels and by-vars
- Analysis programs reuse results where available; update/create as needed
- Key Principle: Create analysis results data once, then flexibly map to multiple output formats:
 - Post-text tables
 - In-text summaries
 - Interactive R Shiny dashboards
 - Publication tables
 - Posters
 - Regulatory submissions

Pre-ARS Vision





The Crossroads

Custom ARD vs. CDISC ARS

Buy vs Build

Need to balance
immediate operational
needs with long-term
industry alignment

Option 1 - Custom ARD Solution

- Advantages:

- Full control over design and implementation
- Simple, fit-for-purpose structure tailored to organizational needs
- Flexible many-to-many mapping of results to layout metadata

- Challenges:

- Requires dedicated time and specialized skill sets to build
- Resource constraints and competing priorities
- Proprietary solution without industry standardization – Lack of interoperability
- Unknown regulatory acceptance

Custom ARD Solution (PoC)



Jazz Pharmaceuticals®
Innovation that performs

Horizon - Integrated E2E Analytics Platform

[Home](#) [Digital Shell](#)

Metadata Model Data Layer Processing Shell Renderer

Shell Metadata

T_DEMOG_001

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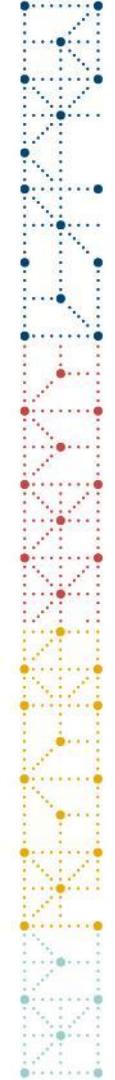
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Shell Preview Shell ARD Preview

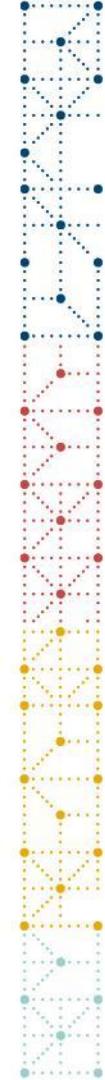
Table 9.1.2.1
Summary of Demographic Data
(Safety Analysis Set)

Characteristic	Placebo (N=25)	JZP999 - 4mg (N=25)	JZP999 - 10mg (N=25)	JZP999 - 20mg (N=25)	Overall (N=100)
Age (years)					
n	25	25	25	25	100
Mean (SD)	57.5 (18.38)	59.0 (21.02)	51.8 (23.89)	56.1 (22.06)	56.1 (21.28)
Median	57.0	62.0	50.0	63.0	58.5
Q1, Q3	45.0, 72.0	44.0, 76.0	33.0, 79.0	34.0, 72.0	38.0, 76.5
Min, Max	18, 84	19, 89	19, 88	25, 89	18, 89
Sex at birth, n (%)					
Female	13 (52.0)	11 (44.0)	15 (60.0)	11 (44.0)	50 (50.0)
Male	12 (48.0)	14 (56.0)	10 (40.0)	14 (56.0)	50 (50.0)
Race, n (%)					
American Indian or Alaska Native	0	0	0	0	0
Asian	13 (52.0)	12 (48.0)	13 (52.0)	9 (36.0)	47 (47.0)
Black or African American	8 (32.0)	8 (32.0)	4 (16.0)	8 (32.0)	28 (28.0)
Native Hawaiian or Other Pacific Islander	1 (4.0)	0	0	0	1 (1.0)
White	3 (12.0)	5 (20.0)	8 (32.0)	8 (32.0)	24 (24.0)
Other	0	0	0	0	0
Multiple	0	0	0	0	0
Ethnicity, n (%)					
Non Hispanic or Latino	23 (92.0)	24 (96.0)	22 (88.0)	24 (96.0)	93 (93.0)
Hispanic or Latino	2 (8.0)	1 (4.0)	3 (12.0)	1 (4.0)	7 (7.0)
Not Reported	0	0	0	0	0
Unknown	0	0	0	0	0
Height (cm) at Baseline					



Our Initial Exploration - Custom ARD POC

- Early Proof of Concept Development
 - Demonstrated feasibility of custom approach
 - Aligned with our reusability vision
 - Ultimately set aside to prioritize ARS evaluation



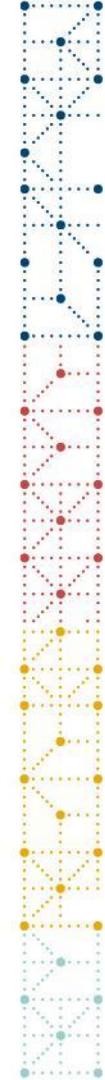
Option 2 - CDISC ARS Compliant Solution

- Advantages:
 - CDISC compliant with common industry structure
 - Future-ready for potential regulatory requirements
 - Early adopters already submitting ARS to FDA proactively
 - Industry momentum and tool ecosystem developing
- Challenges:
 - Extremely complicated design and specification
 - 1-to-1 linking of layout metadata with ARD (limits reusability)
 - Typically represented as separate ARD files per output vs. central repository
 - Diverges from "derive once, render many" vision

ARS Implementation - Buy vs. Build

- Tool Ecosystem for ARS Adoption
- Commercial Solution:
 - Clymb Clinical TFL Designer - Shell design and layout metadata generation
- Open-source R Tools for ARD and TFL generation:
 - siera – ingest ARS metadata to generate R-code for ARD generation
 - CARDS - Create ARD from ADaM
 - TFMRT - Table, Figure, and Listing creation from ARS
- Strategy: Hybrid approach leveraging both commercial and open-source tools

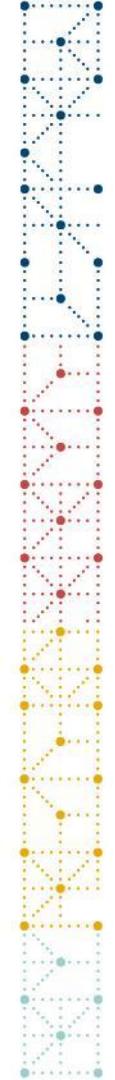




ARS Implementation Insights

- Template and metadata-driven tools [commercial and open-source] are key to generating CDISC ARS-compliant metadata efficiently for a successful ARD generation.
- Prospective metadata is essential to support standardized, automation-ready deliverables from study startup. Metadata setup takes upfront time.
- Iterative refinement - Metadata accuracy improves as teams align to ARS conventions and practices.
- Integrating ARS required change, prompting updates to established workflows and team processes.
- Learning curve - Many nuances not apparent upfront

Key Observation: Successful ARS implementation depends on prospective, metadata-driven design supported by adaptable tools (commercial and open-source) and evolving team practices.



Lessons Learned

- Key Takeaways:
 - Gap between initial ARD vision and ARS reality requires pragmatic workarounds
 - Tool maturity and process integration are critical success factors
 - ARS adoption is a journey, not a destination

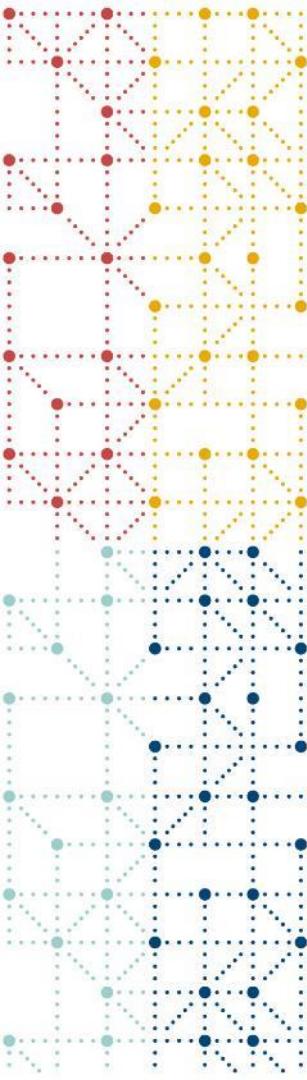
Questions & Discussion



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Thank You!

