

RESEARCH DATA ALLIANCE



Adoption of RDA outputs to support Reproducible Health Research



Learning objectives

How RDA is supporting adoption of outputs and recommendations

2. How RDA recommendations and outputs build both <u>social</u> and technical bridges to enable open sharing and re-use of data as exemplified by work across health research

 How to engage with RDA to develop, adopt, or learn more about recommendations or outputs



Contributors

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Health Data IG

https://www.rd-alliance.org/groups/health-data.html

Reproducible Health
Data Services WG

https://www.rd-alliance.org/groups/reproducible-health-data-services-wg

RDA Adoption Support Task Force





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Trust-IT Services Ltd.



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Outreach & Impact Support

Göttingen State and University Library



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Rensselaer Polytechnic Institute



Agenda

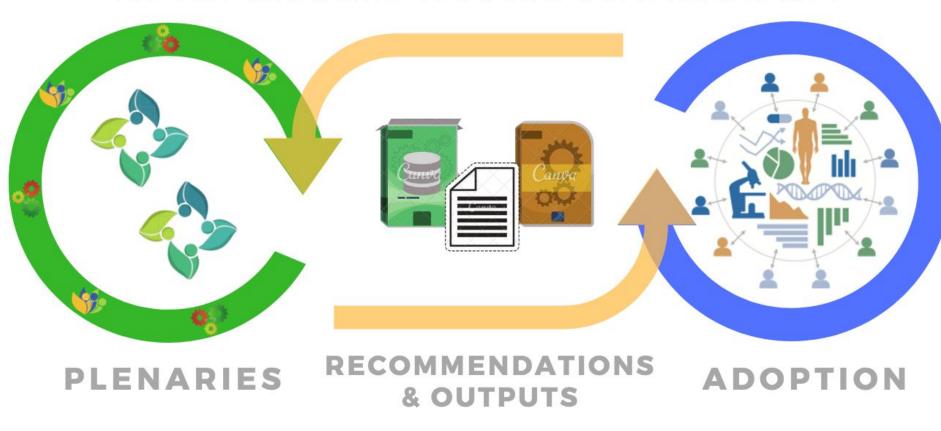
- What is RDA?
- How do we support adoption and impact?
- Introduction of social and technical issues
- Stanford University and the Center for Expanded Data Annotation and Retrieval (CEDAR)
- Washington University's EHR implementation of Dynamic Data Citation
- How to get involved with RDA
- How to learn more about adopting RDA outputs and recommendations

——————— Question & Answer

What is RDA?

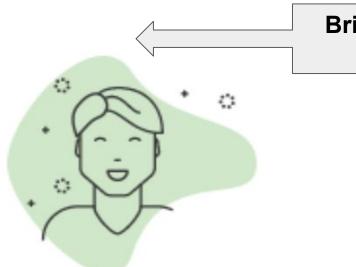


RESEARCH DATA ALLIANCE

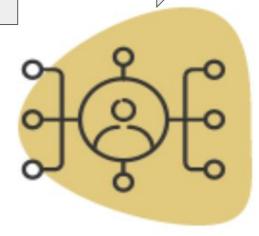


Vision: Researchers and innovators <u>openly share data</u> across technologies, disciplines, and countries to address the grand challenges of society.









- Preservation and stewardship
- Improving usefulness of data
- Data sharing culture
- Education and training
- Data security, legal interoperability, and compliance

- Data discovery
- Provenance, reuse, and interconnection
- > Issues of scale
- Governing and maintaining infrastructure
- Data access and interoperability



What are the issues surrounding data collection and data sharing for reproducible health research?



Assumptions

- 1. Process of sharing biomedical data not comprehensively documented
- 2. Effective (and ethical?) health interventions require verifiable and reproducible evidence
- 3. The responsibility of ensuring reproducible research involves multiple stakeholders across the research workflow



What is reproducibility?

Computational Reproducibility:

If we took your data and code/analysis scripts and re-ran it, we can reproduce the numbers/graphs in your paper

Empirical Reproducibility:

We use your exact methods and analysis, but collect new data, and we get the same results

Replicability (Results Reproducibility):

We have enough information to re-run the experiment the way it was originally conducted



Reproducible Scientific Workflows

"Reproducibility implies repetition and thus a requirement to also move back – to

retrace one's steps, question or change assumptions, and move forward again."

Millman, K. J., & Perez, F. (2014). Developing Open-Source Scientific Practice (V. Stodden, F. Leisch, & R. D. Peng, Eds.). In Implementing Reproducible Research (CRC the R series, pp. 149-183). Boca Raton, FL: Taylor & Francis Group, LLC.



The Grand Why

"The construction of a scientific heritage

where anyone can validate the work of others and build upon it."1

Millman, K. J., & Perez, F. (2014). Developing Open-Source Scientific Practice (V. Stodden, F. Leisch, & R. D. Peng, Eds.). In Implementing Reproducible Research (CRC the R series, pp. 149-183). Boca Raton, FL: Taylor & Francis Group, LLC.

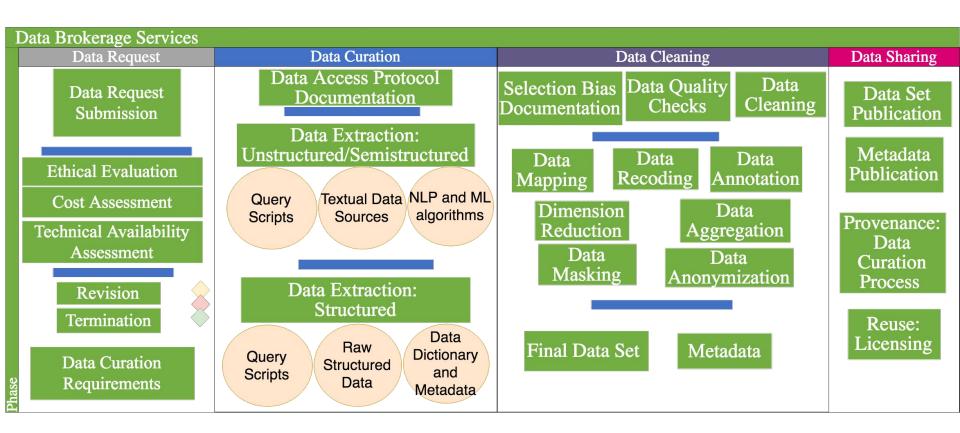


The Concern

How do we ensure all stages of data collection and preparation within the scientific compendium are fully transparent and appropriately accessible to achieve reproducibility?

Reproducible Health Data Services (Social)





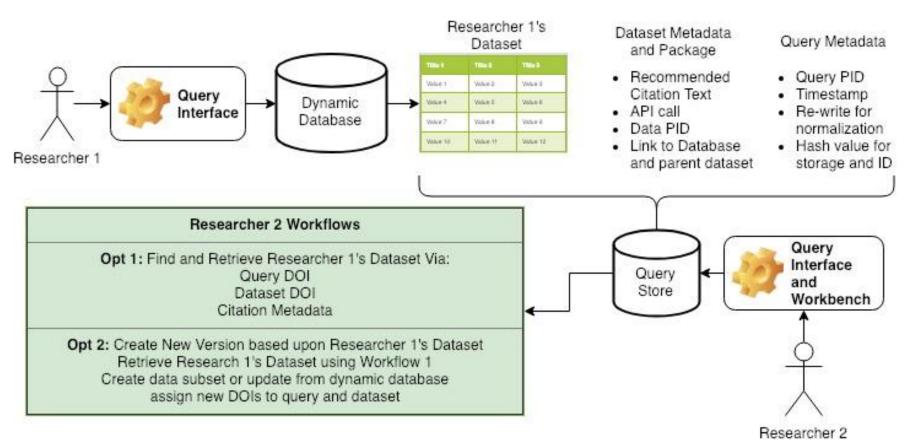


The Concern

Big Data evolves rapidly in volume, variety, and velocity. How do we ensure traceable provenance of a dataset from an evolving data source?

Scalable Dynamic Data Citation (Technical)





RDA Data Citation WG: https://rd-alliance.org/groups/data-citation-wg.html
RDA WG Case Statement: https://www.rd-alliance.org/filedepot?fid=102
Executive Summary of Output:

https://docs.google.com/document/d/1SUer28B30Gg4yNNHfmztRawP7zLJqmx2lpEZSceeBMs/edit?usp=sharing

Health Data IG

&

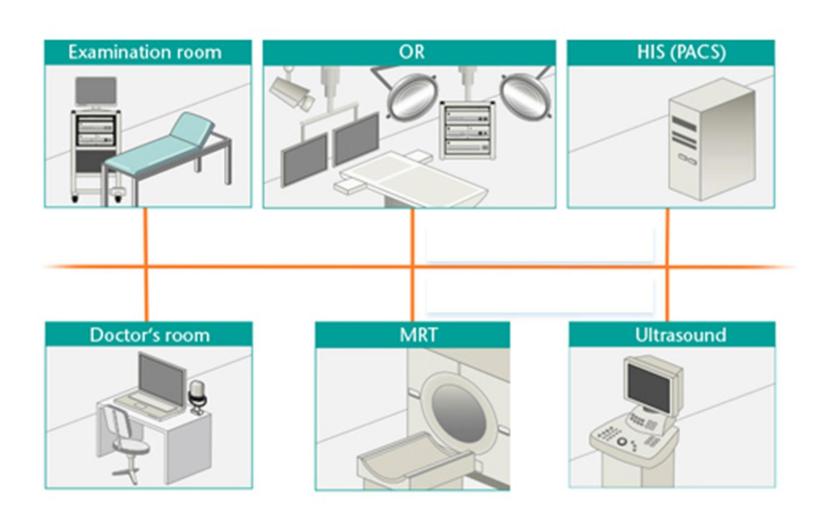
Reproducible Health Data Services WG

Learning from bedside data...

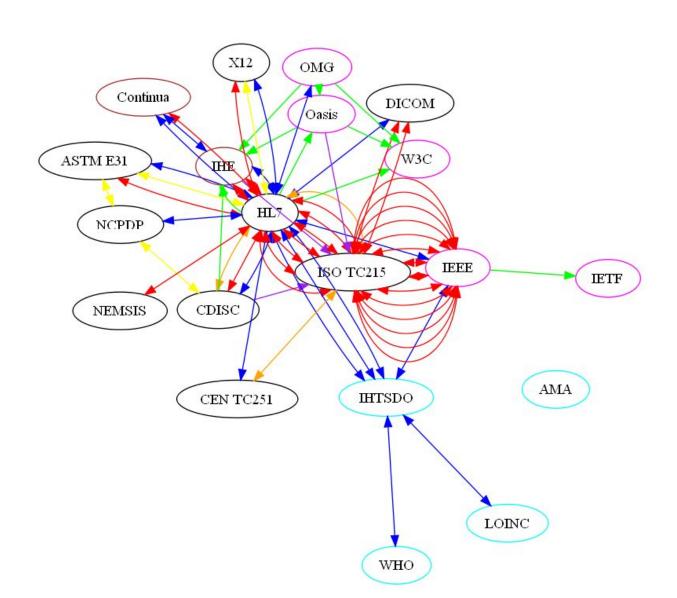
how to create FAIR research data from routine care data?



Heterogeneous system with multiple types of data

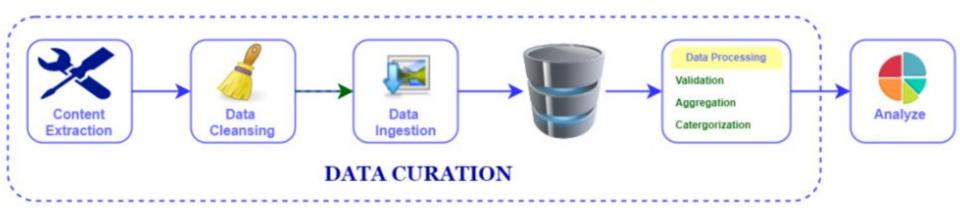


Heterogeneous standards - if any at all



Challenges of Health Data Services

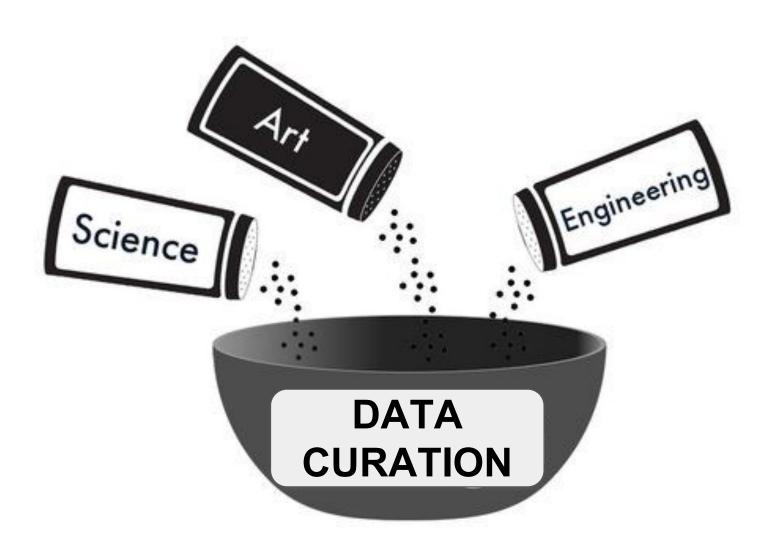
- Data curation processes
 - requires to interact with multiple systems
 - needs manual effort and crafting to map, transform, clean the data
 - typically queries and scripts are written case bases
 - ETL processes varies highly depending on the task and the curator



Challenges of Health Data Services

- Documenting metadata
 - very limited documentation if any
 - there is no recommendation / guideline on what to document
 - too time consuming
- How to capture the metadata
 - no guidance on what is useful (problem of granularity)
 - what are the available standards?
- Sharing health data curation metadata
 - no standard way to find, access and interpret this metadata

Art of curating data



"to improve the reuse of health data by providing recommendations for reproducible data curation and brokerage workflow services"

FAIR + Repeatability
Fitness for Purpose
Trustability ...

Deliverable 1: Recommendation Statement perform a gap analysis

- Relevance, maturity levels, ...
- Identify needs for further standards and methods
- Communicate the need with relevant groups

TASKS

- Identify data curation processes
 - which activities are carried out at the different phases of the data services
- Document challenges related to activities
- Explore the available standard stack
 - RDA, W3C, ISO, research communities

Deliverable 2:

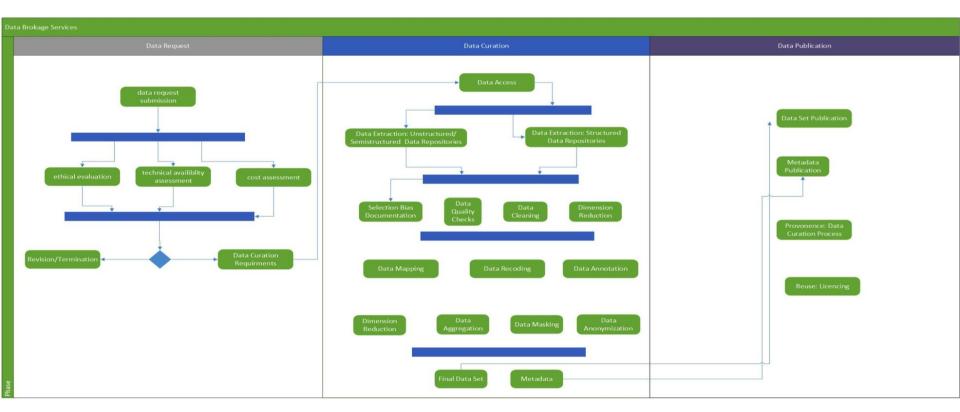
Adoption and Training Guide

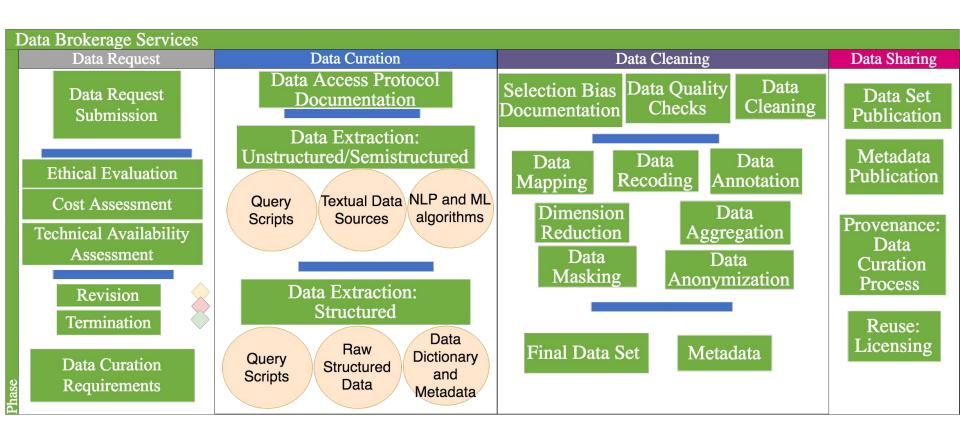
- Documentation of state-of-the-art methods and standards for clinical data curation
- Best practices for capturing and storing data curation metadata
- Identify use cases

Reproducible Workflows for Health Data Services

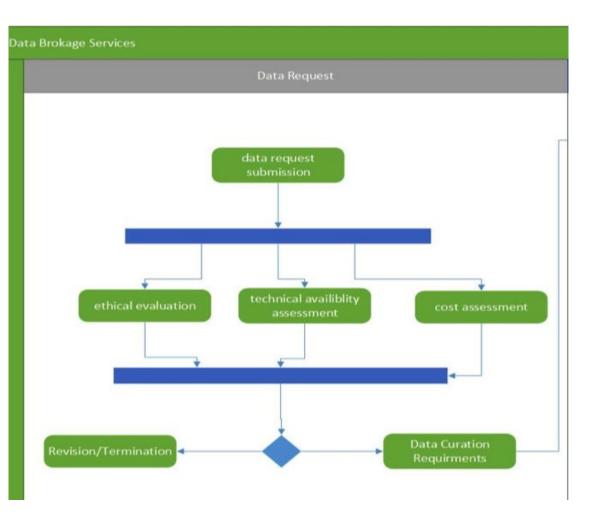
Data Request Data Curation

Data Sharing





Data Request Elements



Ethical Assessment

- IRB protocol approval
- Consent form
- Approval date
- Reviewing institution

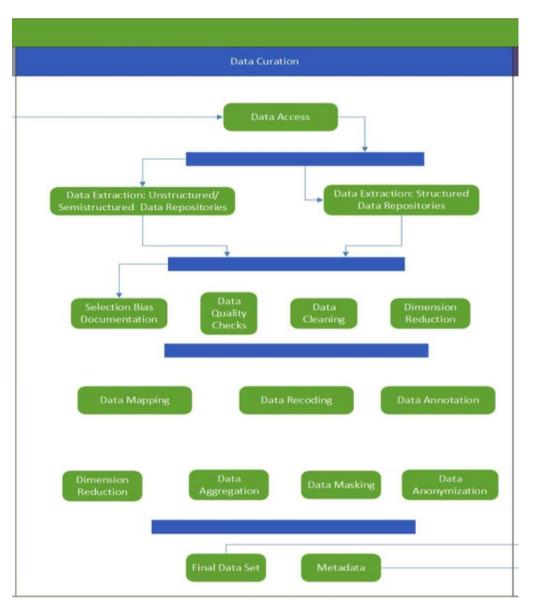
Cost Assessment

- Grant information
- Center expected hours of work

Feasibility Assessment

- Is the requested data available?
- Does it satisfy the clinical question?

Data Curation Processes



What is the source of data?

- DOI and Data citation?
- Multi-Site/Multi-registry Collection

Are query scripts FAIR?

- Interoperable
- Commented

How are participant inclusion and exclusion criteria operationalized?

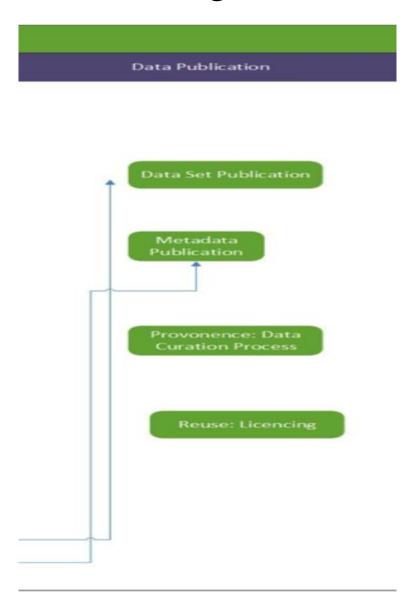
- Ontologies and standardized vocabs
- Limitations

How are data cleaned, mined, and merged?

FAIR software code

Is their a FAIR final dataset or data dictionary?

Data Sharing and Publication



Can we apply FAIR principles to data products of the health data service workflows?

- Rich metadata
- Persistent identifiers
- Licensing
- Common protocols to access

How to share data curation metadata together with the data?

Is it possible to have a minimum information reporting standard for data services?

How to make the data curation workflow metadata FAIR?

Draft Working Document

Current Health Data Service Center Workflow

https://docs.google.com/spreadsheets/d/1-uSocVpju4_fBcMDBgW2LxG3EpvBNRedOQuings2Cok/edit?usp=sharing

- Phases of Data Services
- Curation Activity
- Explanation
- Reproducibility Challenges
- Possible Metadata Formats
- Relevant Community Resources/Standards

Future Work

WG case statement:

https://docs.google.com/document/d/1wrpxYnIdvJHKN21J70esdFhVEabqizjaNXJePAp2SnM/edit?usp=sharing

- Participate in the working group
- Vet the framework of elements
- Interested in adoption testing?

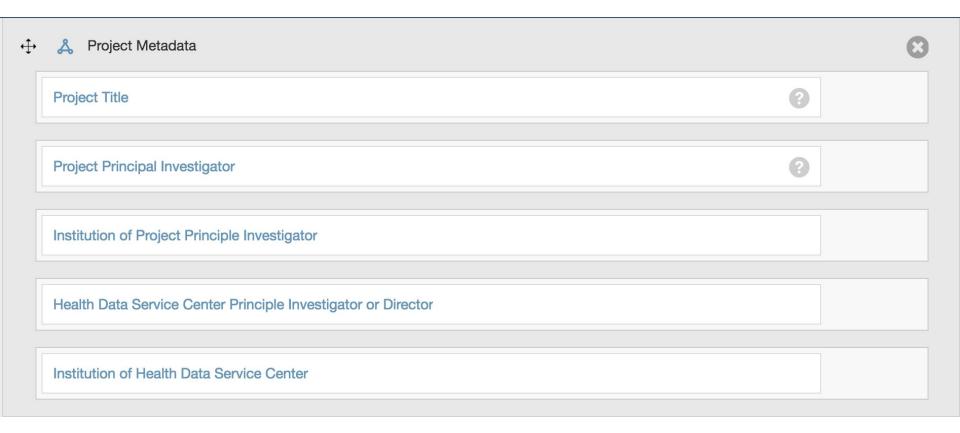
Stanford Adoption Case Study

	Title	Created
#	Cost Assessment	1/9/18 2:15 AM
#	Data Access	1/10/18 7:00 AM
#	Data Extraction - Structured	1/10/18 6:13 PM
#	Data Extraction_Unstructured Data	1/10/18 6:16 PM
#	Ethical Assessment Review	1/9/18 1:50 AM
#	Participant Exclusion Criteria	2/9/18 8:55 PM
#	Participant Inclusion Criteria	1/11/18 2:32 AM
#	Project Metadata	1/11/18 6:59 AM
	RDA_Health Data Service Center_Data Brokerage Workflow	1/11/18 7:21 AM

Types of Data

- Text and numeric descriptors
 - Names, titles, journal of publication...
- Software files (or associated links)
 - Query, cleaning, nlp...
- Data files (or associated links)
 - tabular, clinical notes, data dictionaries
- Identifiers (hopefully persistent)
 - DOIs, URLs, Grant IDs...
- Clinical Ontology Variables
 - Diagnosis, Procedure, Medication, Labs...

Bibliographic and Project Metadata



Ethical Assessment and Review



Data Access and Collection

- **Data Access** Database Name Database DOI Database URL Institution Hosting Database Institutional Department or Center Overseeing Database
- Data Extraction Structured Query Script_Structured Data_File Name Link to Shared Query Script_Structured Data Query Script_Structured Data File Format Query Script_Structured Data_Software Language Query Script_Structured Data_File Version # Query Script_Structured Data_Author(s) Query_Structured Data Execution Date Data Extraction_Unstructured Data

Learning Effective Treatment Pathways for Type-2 Diabetes from a clinical data warehouse

Rohit Vashisht, PhD, 1 Ken Jung, PhD, 1 and Nigam Shah, MBBS, PHD1

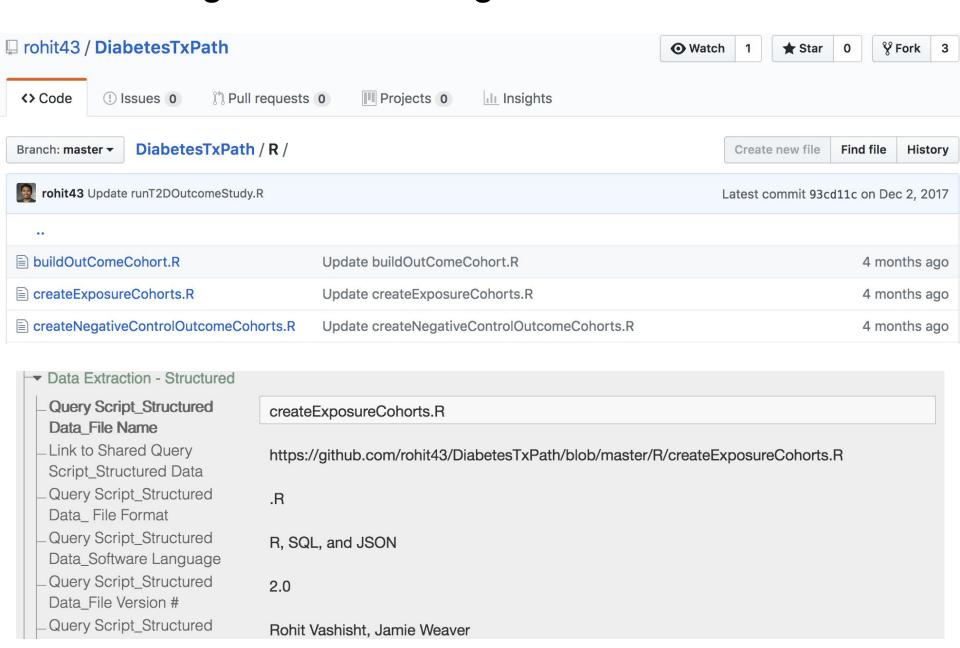
rohit43 modified temp table names



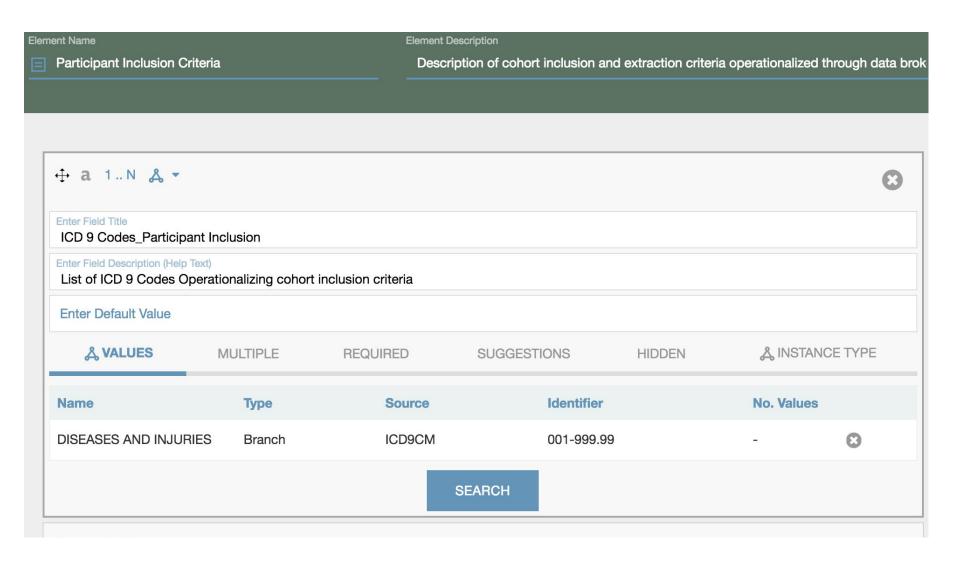
Latest commit d34242a on Dec 12 2017

Tome-o mounted temp table names		Latest commit u34242a on Dec 12, 2017
■ R	Update runT2DOutcomeStudy.R	4 months ago
extras	second commit	11 months ago
inst inst	modified temp table names	3 months ago
■ man	updated R functions	11 months ago
■ DESCRIPTION	Update DESCRIPTION	11 months ago
■ DiabetesTxPath.Rproj	first commit	a year ago
NAMESPACE	first commit	a year ago
README.Rmd	Update README.Rmd	4 months ago
README.html	updated R functions	11 months ago

Linking and describing cohort collection files



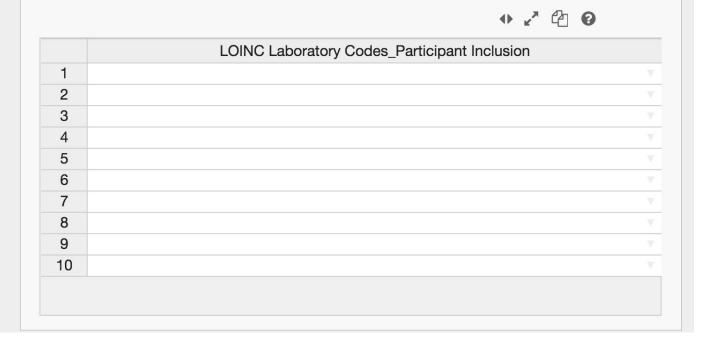
Mapping Ontologies, Vocabularies, and Standards



- → Data Extraction_Unstructured Data
- Inclusion Criteria
 - _ICD 9 Codes_Participant Inclusion
 - _ICD10 Codes_Participant Inclusion
 - _HCFA/HCPCS Procedure Codes_Participant Inclusion
 - CPT Codes_Participant
 Inclusion
 - RXNORM Medication
 Codes_Participant Inclusion
 - _SNOMED Medication
 Codes_Participant Inclusion
 - LOINC Laboratory

 Codes_Participant Inclusion

{"ConceptSets":[{"id":0,"name":"HbA1c_v2","expression ":{"items":[{"concept":{"CONCEPT_ID":3004410, "CONCEPT_NAME":"Hemoglobin A1c (Glycated)", "STANDARD_CONCEPT":"S","INVALID_REASON":"V", "CONCEPT_CODE":"4548-4","DOMAIN_ID":"Measure ment", "VOCABULARY_ID":"LOINC","CONCEPT_CLASS_ID":"Lab Test","INVALID_REASON_CAPTION":"Valid","STAND ARD CONCEPT CAPTION":"Standard"}}



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"Data Extraction_Unstructured Data": {
  "@context": {
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    "Text Extract Script_Unstructured Data_Software Language": "https://schema.metadatacenter.org/properties/719ec416
    "Query Script Unstructured Data File Name": "https://schema.metadatacenter.org/properties/51645b80-69c3-4e51-9b94
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    "Query script_Unstructured Data_Software Language": "https://schema.metadatacenter.org/properties/3e9e8745-4b96-4
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    "Text Mining Script Unstructured Data Date Executed": "https://schema.metadatacenter.org/properties/4166f856-17ba-
    "Link to Shared Query Script_Unstructured Data": "https://schema.metadatacenter.org/properties/6a4d573b-a83d-4e02-
    "Link to Share Text Mining Script Unstructured Data": "https://schema.metadatacenter.org/properties/9865a4cd-b9e0-
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    "LOINC Laboratory Codes_Participant Inclusion": "https://schema.metadatacenter.org/properties/08e85ded-623c-4fec-
    "HCFA/HCPCS Procedure Codes_Participant Inclusion": "https://schema.metadatacenter.org/properties/5afc336c-5dc1-4
    "ICD10 Codes_Participant Inclusion": "https://schema.metadatacenter.org/properties/867ab168-46d9-48c5-8f59-941a11
  },
  "ICD 9 Codes_Participant Inclusion": [
   {}
  1,
  "CPT Codes Participant Inclusion": [
```

{}

Potential Future Use Cases



Australian Government

Australian Digital Health Agency







Treatment Pathways in Chronic Disease

Objective: The objective of this study is to characterize the prevalence of different treatment pathways for three chronic diseases: Hypertension, Type II Diabetes, and Depression. We will systematically summarize the treatment pathways observed among patients who have at least 3 years of continuous observation and persistent treatment following initiation. We will stratify the results by year to evaluate temporal trends, and will further stratify by data source to determine if treatment pathways vary by population, geography, and data capture process.

Rationale: While numerous treatment guidelines exist for chronic conditions, there is a paucity of data on the real-world treatment pathways that patients experience in practice. Understanding these pathways is essential for establishing context around questions of drug utilization, effectiveness, and adherence.

Project Leads: Patrick Ryan, Jon Duke, George Hripcsak, Martijn Schuemie, Nigam Shah

Coordinating Institution(s): Janssen R&D, Columbia University, Regenstrief Institute, Stanford University

Additional Participants:

Full Protocol: W Hypertension Treatment Pathways 12-4-2014

Initial Proposal Date: 12/3/2014

Launch Date: 12/5/2014

Study Closure Date: 12/31/2014

Results Submission: Email or SFTP

Requirements

CDM: V4 or V5

Database Dialect: SQL Server, Postgres, Oracle

Software: SQL as above, R (optional)

Proposed Next Steps

- Assess impact and scalability with more use cases
- Iterative reviews with other members of the RDA to vet granularity and interoperability
- Continued collaboration with CEDAR to test and develop functionality
- Link brokerage metadata to additional scientific outputs across repositories

Washington University Center for Biomedical Informatics Adoption Case Study

Incorporating Data Citation in a Biomedical Repository

An Implementation Use Case

Snehil Gupta
Connie Z. Regan
Brian Romine

Daniel Vianello Cynthia Hudson Vitale Leslie McIntosh



Funding Support

MacArthur Foundation 2016 Adoption Seeds program Foundation through a sub-contract with Research Data Alliance

Washington University Institute of Clinical and Translational Sciences
NIH CTSA Grant Number UL1TR000448 and UL1TR000448-09S1

Siteman Cancer Center at Washington University NIH/NCI Grant P30 CA091842-14

WashU CBMI Research Reproducibility

Resources Repository

https://github.com/CBMIWU/Research_Reproducibility

Slides

http://bit.ly/2nxjNK8

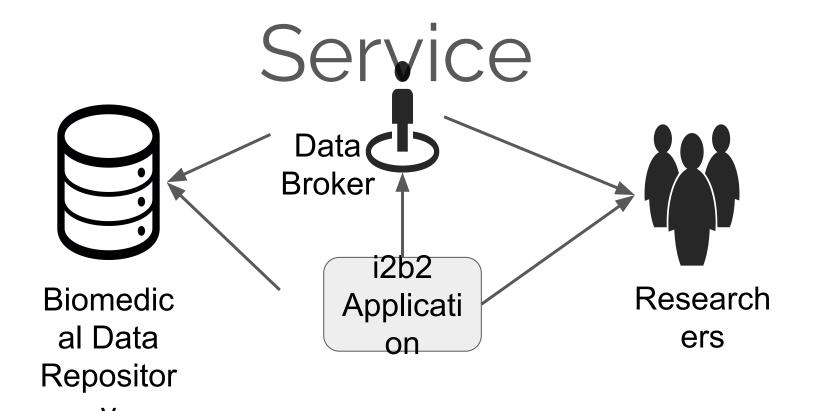
Bibliography

https://www.zotero.org/groups/biomedical_informatics_resrepro

Background

BDaaS

Biomedical Data as a



Move some of the responsibility of reproducibility

Biomedical Biomedical Researcher Pipeline

RDA Data Citation WG Recommendations

- R1: Data Versioning
- R2: Data Timestamping
- R3, R9: Query Store
- R7: Query Timestamping
- R8: Query PID
- R10: Query Citation

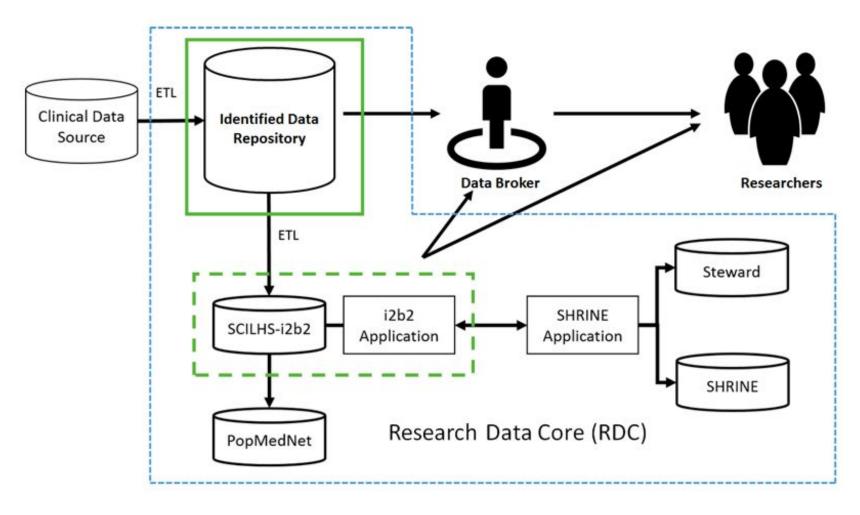
Biomedical Adoption Project Goals

- Implement RDA Data Citation WG recommendation to local Washington U i2b2
- Engage other i2b2 community adoptees
- Contribute source code back to i2b2 community

Approach

- 1. Assess the Center's data infrastructure
- 2. Conduct gap analysis of the infrastructure against RDA-DC recommendations
- 3. Define our local requirements
- 4. Define and evaluate our approach

Infrastructure



Gap Analysis

Table 2: Gap Analysis Summary

Database		Data Versioning (R1)	Data Timestamp (R2)	Query Store (R3/R9)	Query Timestamp (R7)	Query PID (R8)	Citation Text (R10)
Identified	Data	Yes	Yes	No	No	No	No
Repository		(default)	(default)				
i2b2 (Local)		No	No	Yes (i2b2 default)	Yes (i2b2 default)	Yes (i2b2 default)	No
i2b2 (SCILHS	S)	No	No	Yes (i2b2 default)	Yes (i2b2 default)	Yes (i2b2 default)	No
PopMedNet		No	No	No	No	No	No

Internal Implementation Requirements

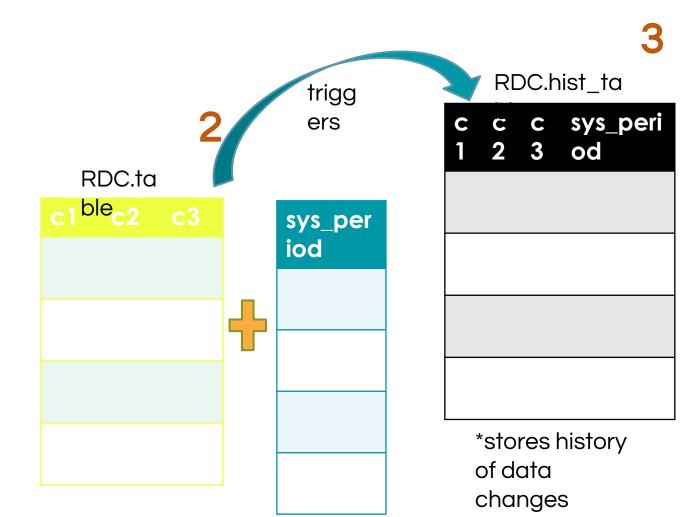
- Scalable
- Available for PostgreSQL
- Actively supported
- Easy to maintain
- Easy for data brokers to use

Implementation

R1 and R2 Implementation



PostgreSQL Extension "temporal_table s"



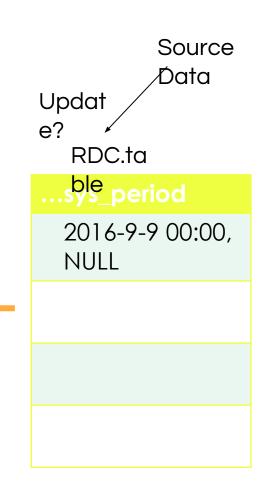
ETL Incrementals

Old

Data

RDC.hist_tabl

sys_period	
2016-9-8	
00:00,	l
2016-9-9	l
00:00	
	1
	1
	1

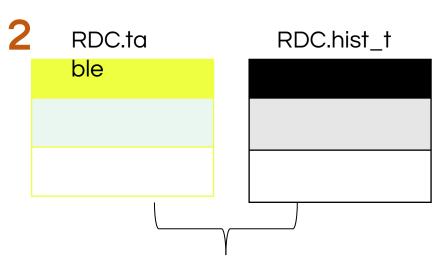




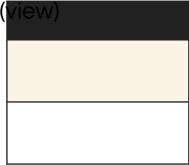
R3, R7, R8, R9, and R10 Implementation



PostgreSQL Extension "temporal_table s"



RDC.table_with_history

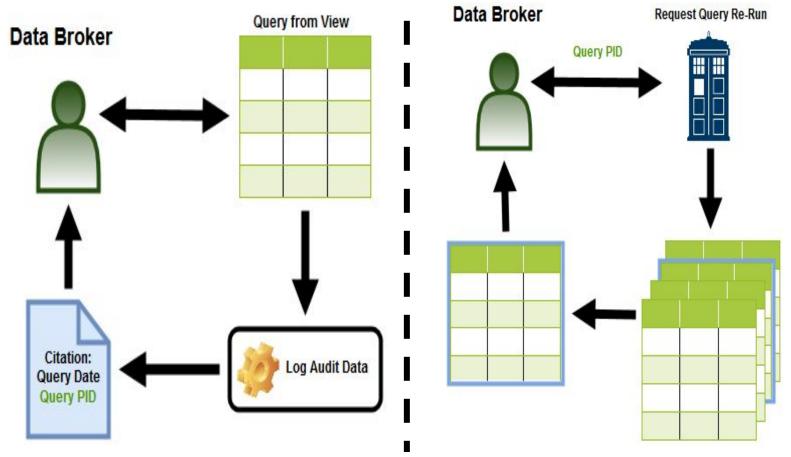


3

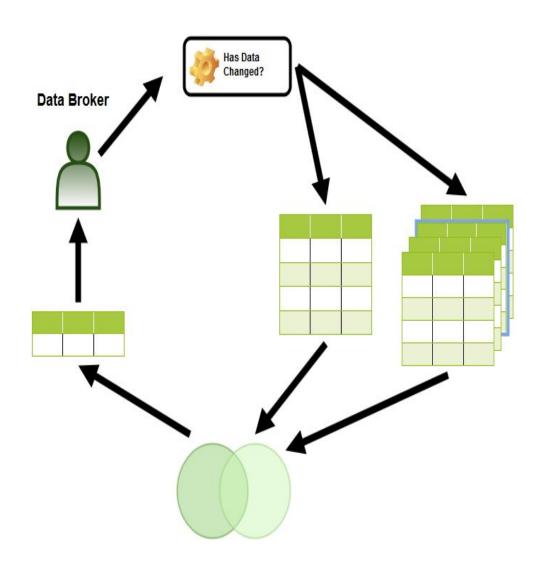
- functions
- triggers
- query audit tables

Data Reproducibility Workflow

TODAY SOME TIME LATER



Bonus Feature: Determine if Change Occurred



Implementation in Practice

Temporal tables

For RDA compliance, we've instituted Temporal tables

This means that each of these tables has a historical analogue.

The naming convention is hist_domain

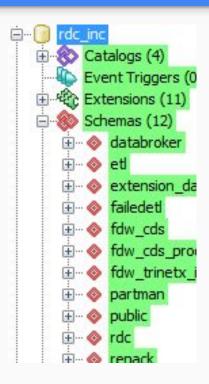
These aren't very helpful for data brokers

You'll use the view domain_with_history

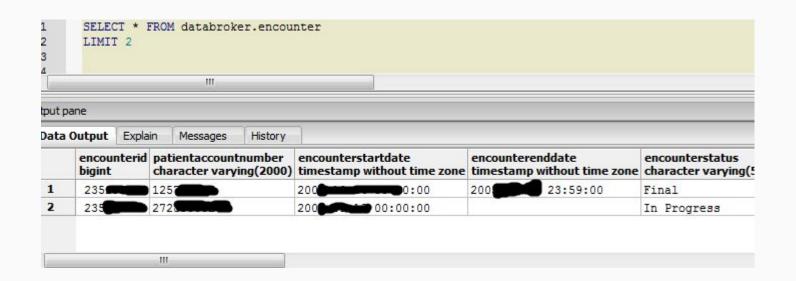
Temporal query workflow

Currently all RDC data is in the RDC schema.

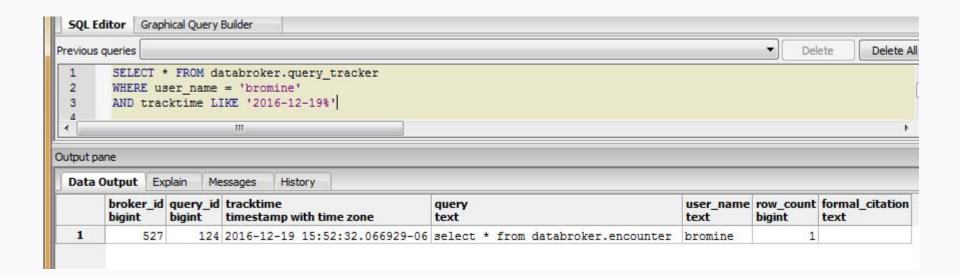
The plan is for databrokers to have access to the databroker schema that will automatically log queries run on it.



Using the databroker schema



Using the databroker schema



Using the databroker schema

revious	s queries [
1 2 3	SELECT databroker.has_my_data_changed('previous', 124)				
4 <u> </u>	III				
utput p	oane				
Data Output		Explain	Messages	History	
	has_m boolea	ıy_data_d ın	changed		
1	f				

Rerunning a historical query

We have a function rerun_histroical_query()

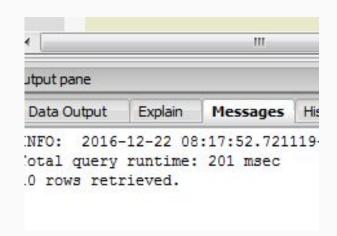
For the earlier example we'll just use these commands

```
select
databroker.rerun_historical_query('cursorna
me', 124); fetch all from cursorname;
```

These need to be run together, the first creates a cursor and the second displays it. They need to be highlighted and run at once.

Citations for queries

After running your query, just click on the messages tab in the output pane on PG admin. There is an info: note raised that will contain the information necessary to cite the dataset you've just generated. Include this when you deliver, so the researcher can cite it. It also contains the information needed to easily re-run the query.



Future Developments

- Develop a process for sharing
 Query PID with researchers in an automated way
- Resolve Query PIDs to a landing page with Query metadata
- Implement research reproducibility requirements in other systems as possible

Outcomes and Support

Return on Investment (ROI) - Estimated

- 20 hours to complete 1 study
- ► \$150/hr (unsubsidized)
- ► \$3000 per study
- ▶ 115 research studies per year
- ▶ 14 replication studies

Center for Biomedical Informatics @WUSTL

NIH-NLM Supplement ucit Research Data

Leslie McIntosh

Cynthia

Hudson-Vitale

Anthony Juehne

Rosalia Alcoser

Xiaoyan 'Sean' Liu

Brad Evanoff

RDA Collaborators

Andreas Rauber

Stefan Pröll

Alliance

Leslie McIntosh

Cynthia

Hudson-Vitale

Anthony Juehne

Snehil Gupta

Connie Zabarovskaya

Brian Romine

Washington University in St. Louis

WashU CBMI Research Reproducibility

Resources Repository

https://github.com/CBMIWU/Research_Reproducibility

Slides

http://bit.ly/2nxjNK8

Bibliography

https://www.zotero.org/groups/biomedical_informatics_resrepro

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https://www.rd-alliance.org/get-involved/individual-membership.html

RDA Recommendations and Outputs

https://rd-alliance.org/recommendations-and-outputs/all-recommendations-and-outputs

Adoption Stories Series

https://www.rd-alliance.org/recommendations-outputs/adoption-stories

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https://rd-alliance.org/add/adoption-stories



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Questions & Discussion