Trial Innovation Network
Vision for Hub Liaison Teams

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MARCH 7, 2017

NCATS

NIH National Center for Advancing Translational Sciences

Trial Innovation Network
Site Based Research – Challenges

Clinical Research Sites—The Underappreciated Component of the Clinical Research System

Robert M. Califf, MD

Developing a clinical trial unit to advance research in an academic institution

Ivana T. Croghan [a,*], Steven D. Viker [a], Andrew H. Limper [a], Tamara K. Evans [a], Alissa R. Cornell [b], Jon O. Ebbert [a], Morie A. Gertz [a]

Site-Based Research in Acute Heart Failure

Patricia A. Adams, BSN, RN

- Complex institutional systems
- Regulatory and bureaucratic burdens
- Variability in training and expertise of local study teams
- Lack of harmonization of processes
- Staff turnover

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Hub Liaison Teams – The Vision

- Hub Liaison Teams as clinical trial "catalysts"
  - Work with local study teams to expedite clinical trials
  - Develop harmonized processes
  - Increase quality and efficiency
- Responsibilities
  - Support TIN CIRBs, master contracts, recruitment, study operations
- Standardized training and competencies for Hub Liaison Teams

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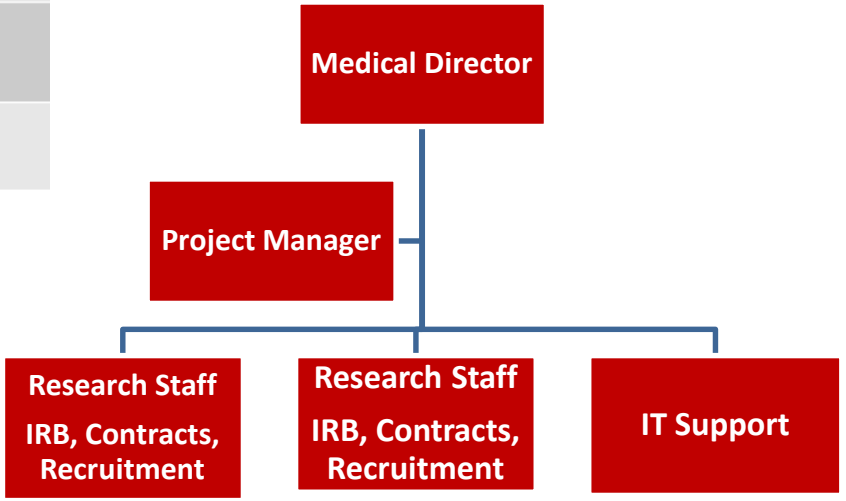
Hub Liaison Teams – A Potential Model

Hub Liaison Team Members	Number of Sites with Identified Team Members
Medical/Faculty Director	61
Hub Admin or Project Manager	59
Primary Research Staff for Central IRB, Standard Agreements, Recruitment Analyses	57
Additional Research Staff for Central IRB, Standard Agreements, Recruitment Analyses	30

- *Most Hubs have identified members of their Hub Liaison Teams*
- *Hub Liaison Teams – a "catalytic" unit*
- *Some Hubs may be able to leverage local infrastructure – clinical trials units*

Potential Hub Liaison Team Organization



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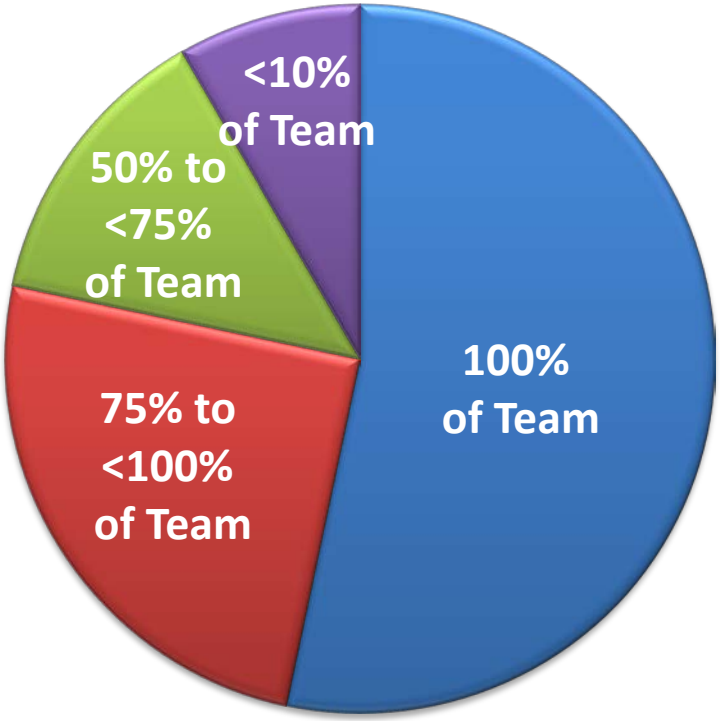
Standardized Training

Hub Liaison Teams
Current Status of GCP Training



Hub Liaison Teams
Training Goals

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Site Based Research – Goals

- Sites/Hubs are the frontline of clinical research
- Organize, optimize, and recognize site-based research efforts
- Create a national harmonized site-based research system
- Catalyze clinical trials

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Central IRB Process and Development

March 7, 2017

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CIRB Discussion Objectives

Present the process of the Central IRB for the Trial Innovation Network, including:

1. Initial Protocol Submission Process
2. Development of tools and guidance
3. Suggested roles and responsibilities related to the Central IRB for the CTSA Hub Liaison Team

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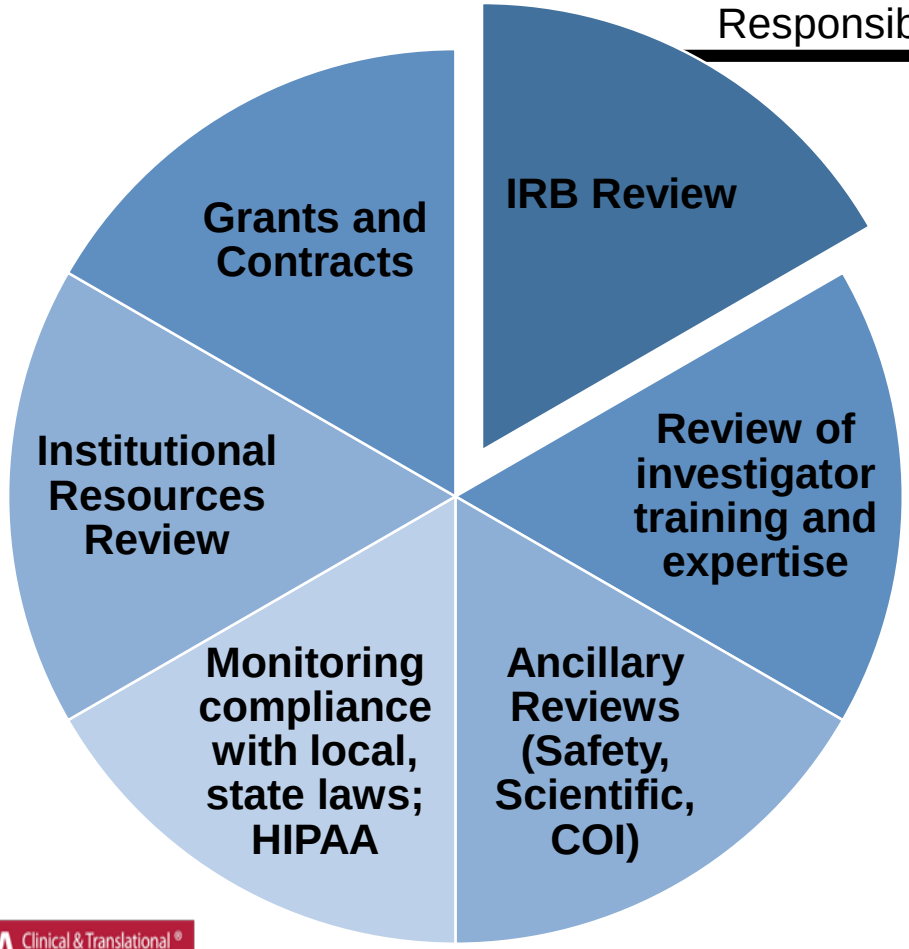
Human Research Protection Program (HRPP)



Responsibilities given to a CIRB

- Initial/new study
- Continuing review
- Amendments to the study
- Local changes (changes to funding, key study personnel, site-specific requests)
- Potential unanticipated problems

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CIRB Roles & Stakeholders

1 Central IRB

Johns Hopkins University | Vanderbilt University Medical Center | University of Utah

2 Relying Site HRPP

Local IRB Professionals | Research Leadership | CTSA Hub Liaisons Ancillary Review Professionals | General Counsel

3 Lead Study Team

4 Participating Study Team

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Central IRB Process for the Trial Innovation Network

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General Stages of CIRB Reliance & Review

1

Protocol & Consent Development

- Consent/authorization document and process
- Recruitment plan
- Data and safety monitoring plan

2 **CIRB Reliance Decision**

3 **Initial Review**

4 **Ongoing Review & Oversight**

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CIRB Reliance Decision

Responsibility of the CIRB and the Relying HRPP

Reliance Agreements delineate what the obligations of the CIRB and the Relying HRPP

Standardized/Harmonized Agreements & Multi-lateral Agreements

SMART IRB Agreement is used for the Trial Innovation Network

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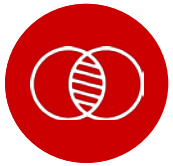
Submission Models



Lead site and other sites can be approved during initial submission

Additional sites can be added via amendment

Only lead site approved during initial submission; all additional sites added via amendment

Used by the Trial Innovation Network

CTSA Clinical & Translational Science Awards

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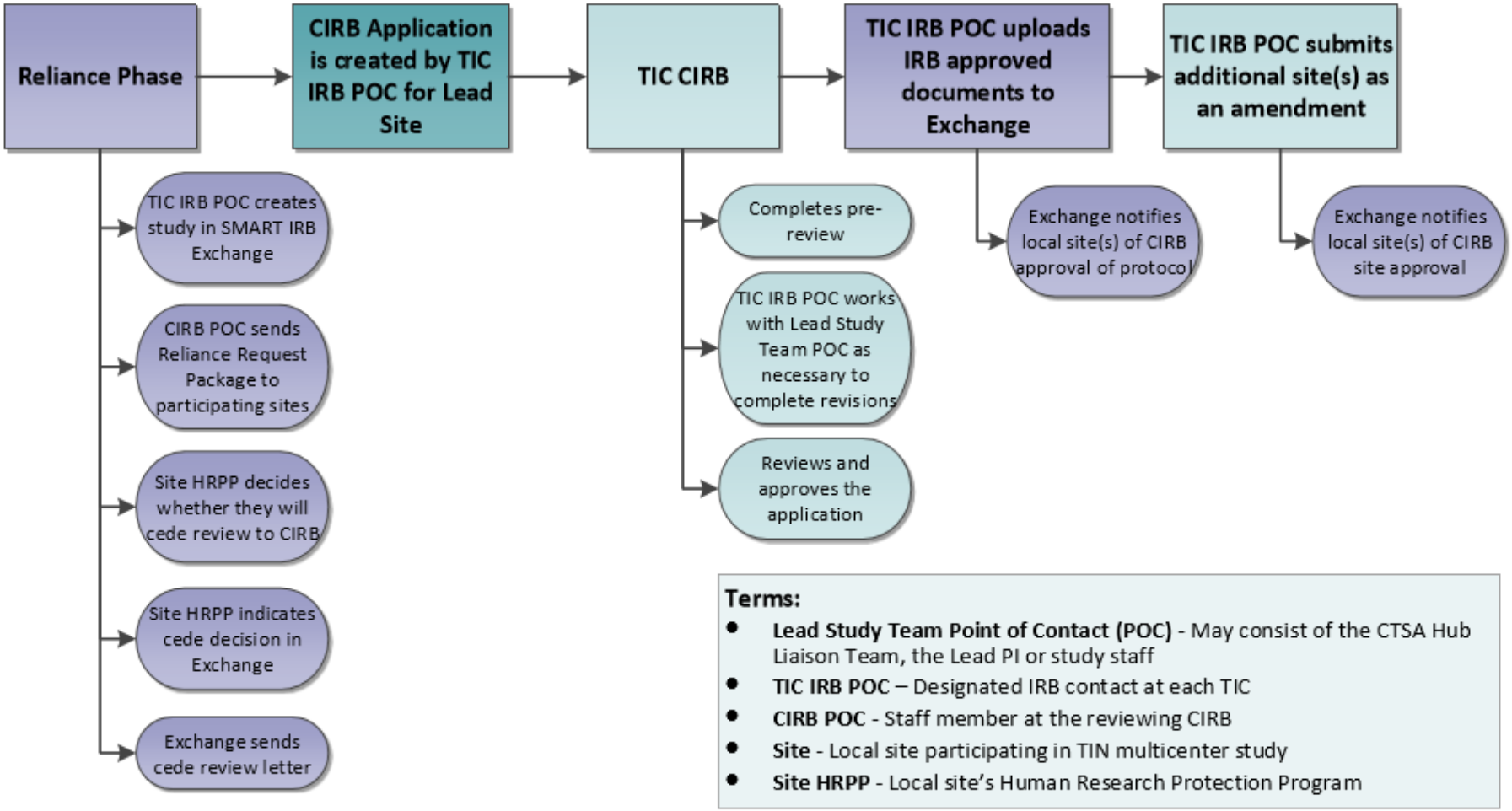
Reliance Phase
CIRB Application is created by TIC IRB POC for Lead Site
TIC CIRB
TIC IRB POC uploads IRB approved documents to Exchange
TIC IRB POC submits additional site(s) as an amendment
TIC IRB POC creates study in SMART IRB Exchange
CIRB POC sends Reliance Request Package to participating sites
Site HRPP decides whether they will cede review to CIRB
Site HRPP indicates cede decision in Exchange
Exchange sends cede review letter
Completes pre-review
TIC IRB POC works with Lead Study Team POC as necessary to complete revisions
Reviews and approves the application
Exchange notifies local site(s) of CIRB approval of protocol
Exchange notifies local site(s) of CIRB site approval
Terms:
Lead Study Team Point of Contact (POC) - May consist of the CTSA Hub Liaison Team, the Lead PI or study staff
TIC IRB POC – Designated IRB contact at each TIC
CIRB POC - Staff member at the reviewing CIRB
Site - Local site participating in TIN multicenter study
Site HRPP - Local site's Human Research Protection Program

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HRPPs and CTSA Hub Liaison Teams

Working hand in hand

Each HRPP and CTSA Hub Liaison Team should work together to decide where the CTSA can be the most helpful.

All components of the HRPP should be considered.

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Suggested Roles for the CTSA Hub Liaison Teams

Protocol Development

Help investigators identify CTSA Hub sites that may be a good fit for a study.

Help identify local policies or practices that could impede the implementation of a protocol at the site.

Initial Review

Help the HRPP provide information to the TIC CIRB about about local laws institutional policies.

Help local study teams respond to the TIC IRB POC in a timely manner.

Facilitate HRPP initial review components.

Ongoing Review & Oversight

Help local study teams remember when certain information is due to the TIC CIRB (e.g., continuing reviews, reportable events).

Facilitate any local audit/QA activity that may be required by the TIC CIRB.

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Suggested Roles for the CTSA Hub Liaison Teams

EFIC: Exception From Informed Consent

1

Design of public disclosure and community consultation plans

2

Advise TIC CIRB on local context

3

Engage with the community and gather feedback

4

Present feedback:

- **TIC CIRB**
- **Establish precedent**

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Development of Tools & Guidance

Phase I Deliverables

- Local Context Information from Relying HRPPs
  - Worksheet for HRPPs and accompanying instruction document
- Initial Central IRB Review Procedures
  - Step-by-step workflow diagram
  - Guidance document
- TIN Indemnification Agreement
  - Still in progress

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Development of Tools & Guidance

Phase II Deliverables

- Developing guidance and instructions for CIRB review of
  - HIPAA authorization and other determinations
  - Informed consent
  - Investigators' financial conflicts of interest
- Methodology for collecting metrics on the CIRB process
  - CTSA Common Metrics

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Development of Tools & Guidance

Phase III Deliverables

- Enhancement of electronic systems for CIRB processes
  - SMART IRB Exchange (ongoing)
  - IRB systems used at individual TICs
- Developing guidance and instructions for reporting of problems and events
- QI and monitoring plan
  - Study level
  - CIRB level

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CIRB Webpage

- Phase I deliverables nearly complete and to be posted ASAP
- Phase II deliverables anticipated completion in the next several months
- Phase III deliverables completion TBD

All documents and processes developed will be tested by the TICs during 2017 and adjustments and improvement will be made.

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Q&A

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Standard Agreements & Streamlined Budgeting Working Group

Hub Liaison Team Meeting

March 7, 2017

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Standard Agreements & Streamlined Budgeting Working Group - Organization

Standard Agreements & Streamlined Budgeting Working Group

Leads: Gordon Bernard, Libby Salberg & Todd Wilson

Budgeting Tools Subgroup

Leads: Bree Burks & Pavel Kruchek

Standard Agreements Development (SA Dev) Subgroup

Leads: Colleen Lawrence

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Summary of Network Priority Deliverables

Deliverable	Responsible Group	Status
• **3 TICs Agree to Use FDP-CTSA Agreement** • **3 TICs Agree to Trial Innovation Network Addendum to FDP-CTSA Agreement**	Standard Agreement Development Team TIC PIs	Complete
• **Sign On CTSA Hubs to FDP-CTSA Agreement with Trial Innovation Network Addendum**	Standard Agreement Development Team	19/64 Hubs Signed On
• **Development of Standard Agreement Workflow**		In progress

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FDP-CTSA Standard Agreement

- 3 TICs have agreed to use FDP-CTSA Agreement with Trial Innovation Network Addendum for all Trial Innovation Network Trials

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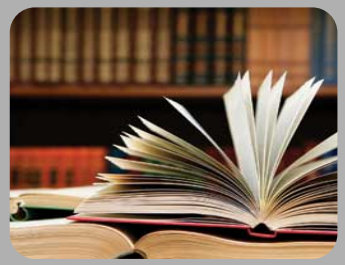
CTSA Hub Call to Action

NOTE: SA-POC (Standard Agreement Point of Contact) is responsible for ensuring Hub Liaison Team is informed regarding SA and Study execution



SA-POC review FDP-CTSA w/ TIN addendum with your Contracts Office

SA-POC Register your Hub for the FDP-CTSA w/ TIN Addendum (REDCap survey)

TIC sends SA package (includes budget, SOW, etc) to CTSA Hub (Study PI) and copies SA-POC, Medical Director, & Hub PI

Study PI submits SA package to Hub Contracts Office and copies SA-POC

Contracts office returns signed SA to TIC for signature and execution and copies responsible parties, including SA-POC

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CTSA Hub Standard Agreement Point of Contact

- CTSA Hub Standard Agreement Point of Contact (SA-POC) ideally should be on the Hub Liaison team
- SA-POC is responsible for working with the TIC, the Study PI, the Hub Contracts Office, and the Hub Liaison Team to move the Standard Agreement to execution
- If there are concerns about using the FDP-CTSA Agreement with Trial Innovation Network Addendum (e.g., need to include State specific language), those concerns can be addressed by the SA Triage team (1 member/TIC) and the Hub SA-POC

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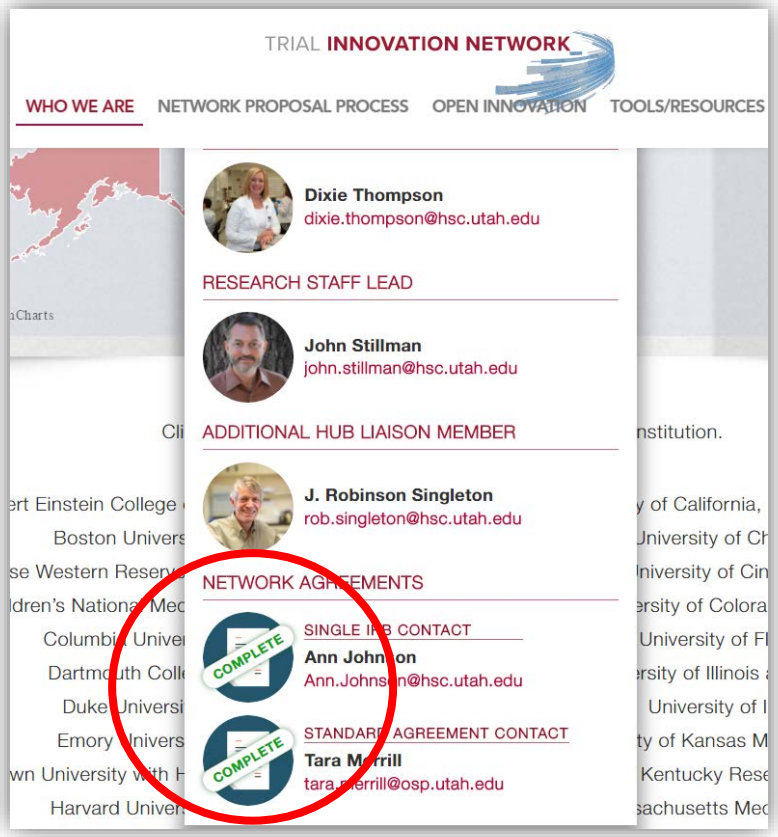
CTSA CALL TO ACTION – Standard Agreement

Registration:

- CTSA Hubs can view and register for the FDP-CTSA Agreement with Network Addendum via a REDCap survey
- Registered sites will be listed on the Network website
- Registration does not commit a Hub to use the agreement, but rather tells other sites that the Hub is willing to use the agreement under the appropriate circumstances

Website view:



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Q&A

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Next Meeting: April 4, 2017, 3-4:30PM ET