Advancing Toward Recovery from Post-Acute Sequelae of SARS-CoV-2 Infection (PASC)

NIH RECOVER Initiative

Technical Assistance Workshop for Applicants to the RECOVER Clinical Trials Research Opportunity Announcement

May 6, 2022

RECOVER
Researching COVID to Enhance Recovery

NIH National Institutes of Health
Researching COVID to Enhance Recovery

Zoom Orientation

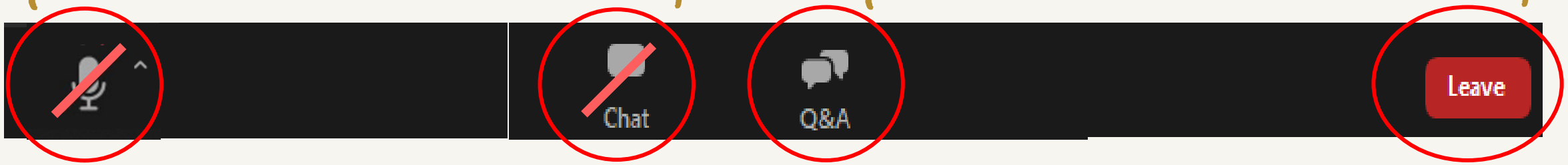
The graphic below highlights the Zoom Bar features that you have as a registrant.

*Your microphone is **automatically muted**. If you have a question, please submit it via the "Q&A" icon.*

*Chat functionality is **unavailable during the webinar.** To ask a question, use the "Q&A" icon.*

*To **ask a question during the webinar**, click on the "Q&A" icon to submit your question.*

*Click here if you would like to **leave the meeting.***



Technical Assistance Webinar (TAW) Overview

Purpose

To enhance potential applicant understanding of the RECOVER initiative, the Clinical Trials Research Opportunity Announcements (ROAs), and to facilitate preparation of responsive applications.

Objectives

- Gain an understanding of the vision and specific objectives of the **RECOVER** initiative.
- Outline the **key scientific & research elements** of the ROAs—including the specific research components.
- Review the **OTA framework**, application process, and requirements.
- Address **prospective applicant questions**.

Agenda

1	**Zoom Introduction**	*Deloitte Zoom Staff*	4:00 - 4:01
2	**Technical Assistance Webinar Introduction and Welcome**	*Clint Wright*	4:01 - 4:03
3	**NIH RECOVER Overview & Mission**	*Clint Wright*	4:04 - 4:10
4	**Clinical Trials Data Coordinating Center (CT – DCC) Overview**	*Tony Punturieri*	4:11 - 4:16
5	**Clinical Trials ROA**	*Gail Weinmann*	4:17 - 4:25
6	**Other Transactions Authority (OTA) Discussion**	*Benjamin Sakovich*	4:26 - 4:33
7	**Q+A**	*Michelle Olive, Moderator*	4:34 - 4:44
8	**Closing Remarks and Next Steps**	*Clint Wright*	4:44 - 4:45

NIH Researching COVID to Enhance Recovery (RECOVER) Initiative on Post-Acute Sequelae of SARS-CoV-2 Infection (PASC)

Overview

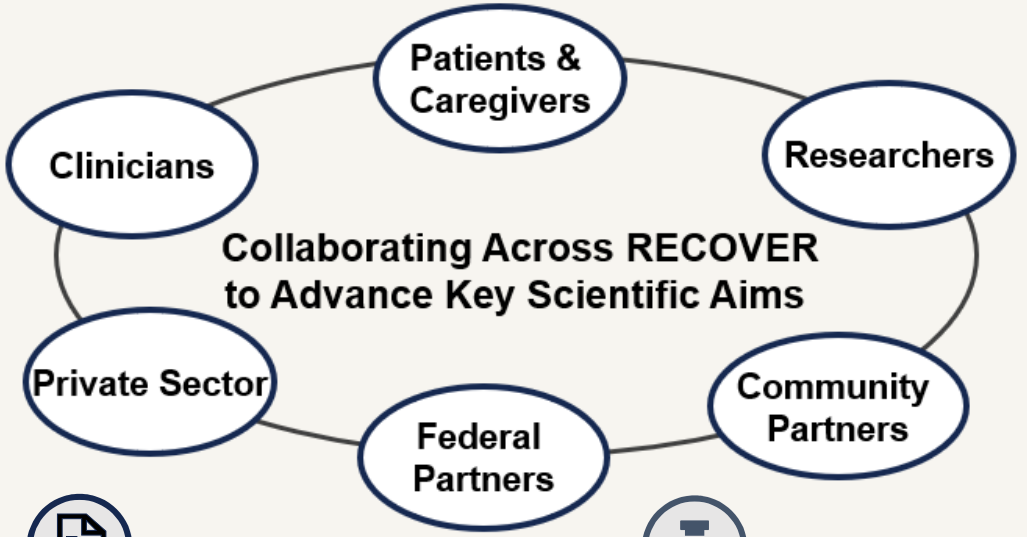
NIH RECOVER Initiative

Goal

Rapidly improve our **understanding** of and ability to **predict, treat, and prevent** PASC.

Key Scientific Aims

1. Understand clinical spectrum/biology underlying recovery over time.
2. Define risk factors, incidence/prevalence, and distinct PASC sub-phenotypes.
3. Study pathogenesis over time and possible relation to other organ dysfunction/disorders.
4. Identify interventions to treat and prevent PASC.



Guiding Principles

Patient-centered, participants as partners

National scale with inclusive, **diverse** participation & community engagement

Platform protocols, standardized methodologies, and common data elements

Adaptive approaches based on emerging science

RECOVER Study Components

RECOVER Cores

Clinical Trial Data Coord. Center	Clinical Science Core	Data Resource Core	Biorepository Core

Elements





RECOVER Clinical Trials	RECOVER Enrolling Cohorts	EHR/ Health Systems Studies	Pathobiology Studies	Tissue Pathology Studies
Clinical Platform with Multi-therapeutic domains	**~40,000** participants	**60 million+** records; **~4 million+** COVID cases	Mechanistic studies of pathogenesis	**50+** tissue types

Data Resources

Clinical	Imaging	Mobile and Digital Health	EHR / Other Real-World Data	Pathology

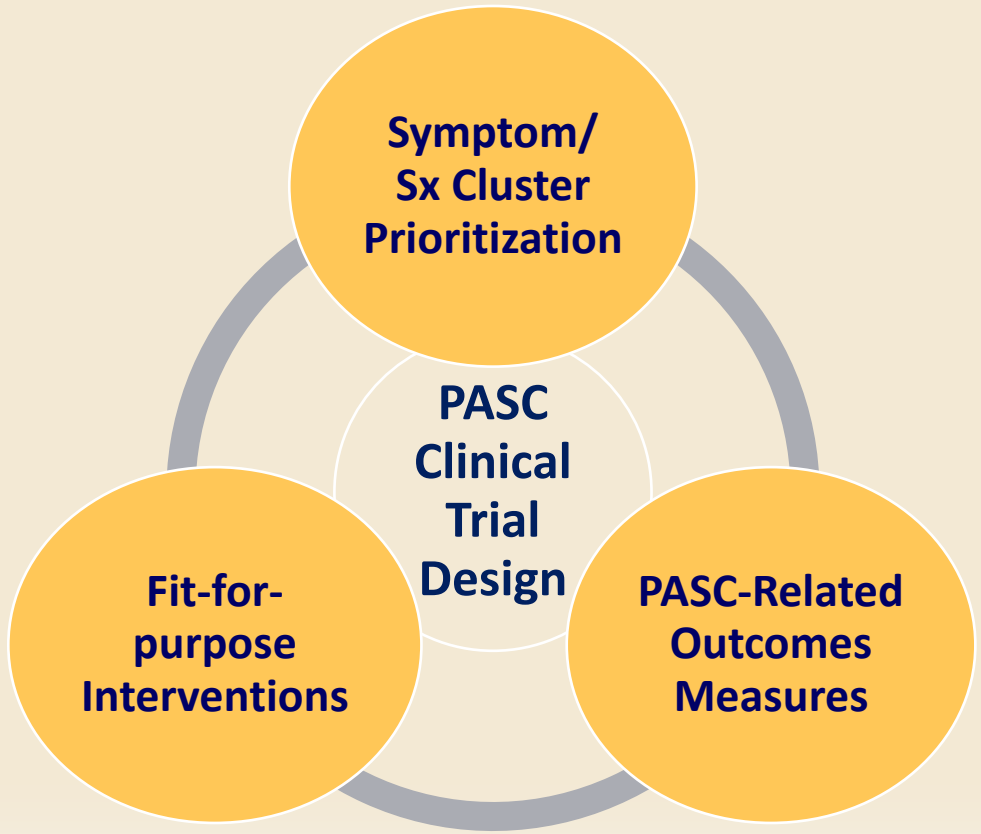
RECOVER Clinical Trial Development and Design: Critical Inputs

• *Patients* • *Clinicians* • *FDA*
• *Clinical Researchers* • *CMS* • *PCORI*

Input from Partners

Phase IIb-III clinical trials that leverage fit-for-purpose design strategies to maximize rigor, efficiency, and flexibility.

- Adaptive platform design and classic RCT as appropriate.
- Frequentist/Bayesian approaches as needed.



Clinical trials developed through a consultative process with engagement of patient, practitioner, and research communities.

Clinical trial proposals solicited from broad research community, OTA-21-015H.

Goal: Clinical trials launched Q3/Q4 2022

RECOVER Clinical Trials Data Coordinating Center (CT-DCC) Overview and Mission

RECOVER Clinical Trials Data Coordinating Center (CT-DCC)

Roles

Support simultaneous multi-intervention platform trials targeting adult and/or pediatric populations, focused on cognitive/behavioral, rehabilitation, complementary alternative medicine, drug, or device interventions.

Operationalized through provision of:

- Program-wide Infrastructure
- RECOVER Clinical Trials Patient Registry
- Clinical Trials Support

RECOVER CT-DCC Responsibilities

Program Infrastructure

- Scientific & operational leadership.
- Enterprise-wide project oversight & management.
- QA/QC compliance & auditing.
- Subcontractor & vendor selection and oversight.

Registry

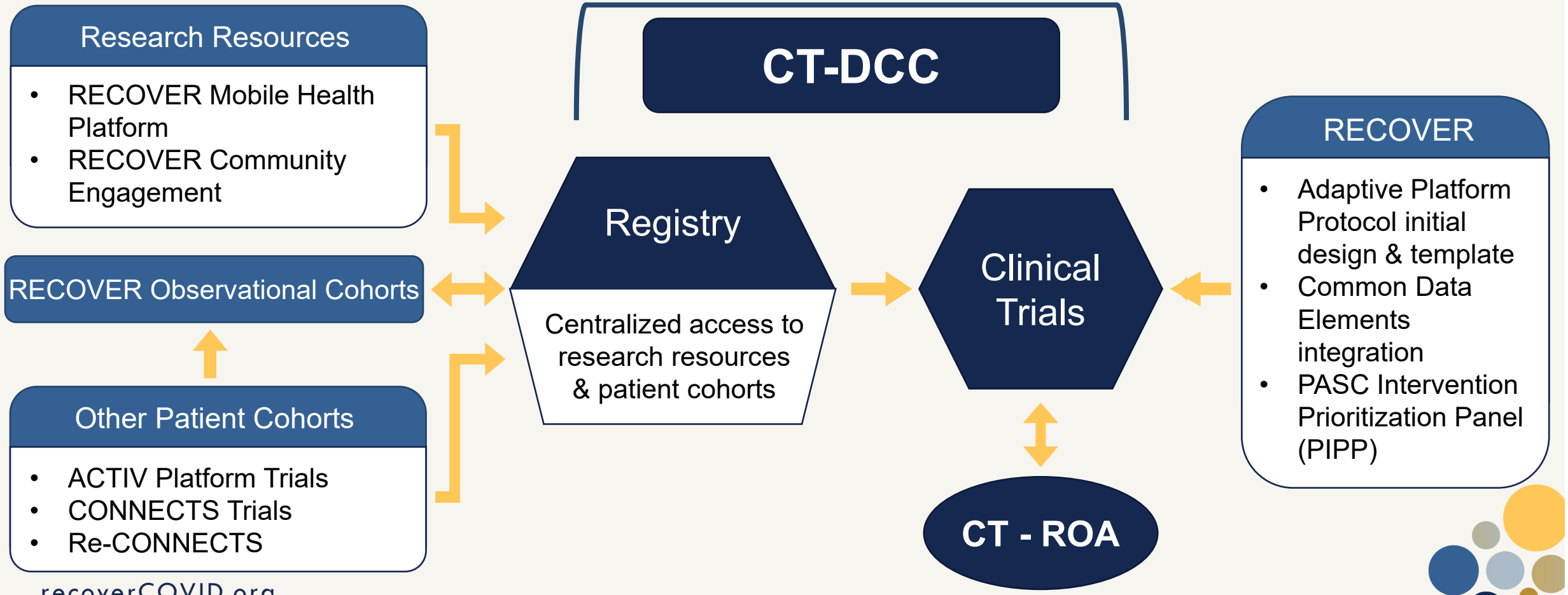
- Support participant recruitment, screening, diverse enrollment, consent & assent.
- Coordinate efforts across existing RECOVER and ACTIV platforms for centralized CT access.
- OEC education materials
- Integrate with RECOVER mobile health platform for device data collection.

Clinical Trials

- Protocol & study design for approved interventions leveraging an adaptive protocol template.
- Operations: regulatory, study materials & training, site selection & management, pharmacovigilance support.
- Data management leveraging RECOVER CDEs & statistics
- Study reporting.

CT-DCC Relationships with RECOVER

Play a critical role in fostering and coordinating the collaboration across numerous institutions participating in the RECOVER Program, as well as leveraging and integrating associated resources in support of RECOVER Clinical Trials.



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RECOVER Clinical Trials ROA

RECOVER Clinical Trials Solicitation: Overview

- **Goal: Well-designed clinical trials** to identify safe and effective treatments and preventive strategies for PASC.
  - Protocols **to be finalized and executed rapidly in collaboration with the RECOVER CT-DCC.**
  - **Interventions across multiple domains** (registration/non-registration pharmacologic/non-pharmacologic/devices/behavioral health and lifestyle/intervention strategies with an evidence base for addressing other relevant conditions).
  - **Diverse trial types** are acceptable.
- Awards are anticipated **to be issued as sub-agreements** under the RECOVER CT-DCC.
- **A future ROA is anticipated for trials in children**.
- **NIH puts high priority on obtaining rapid results and expedited data sharing.**

Other Important Provisions

- **The final protocol(s) will be developed in collaboration with the RECOVER CT-DCC to align with the guiding principles below.**
  - The proposed trial may not be conducted as submitted.
  - Multiple awardees may be asked to work to develop a new protocol that incorporates multiple intervention strategies.
  - Additional interventions may be prioritized and selected by NIH for testing in awarded clinical trials.
- **The RECOVER Clinical Trial Data Coordinating Center will provide administrative, data management, monitoring and pharmacovigilance and statistical support across all RECOVER clinical trials, including selection of appropriate sites.**
- **Evidence of active engagement and contribution of people suffering with PASC,** as well as their caregivers, in the development of the research program.

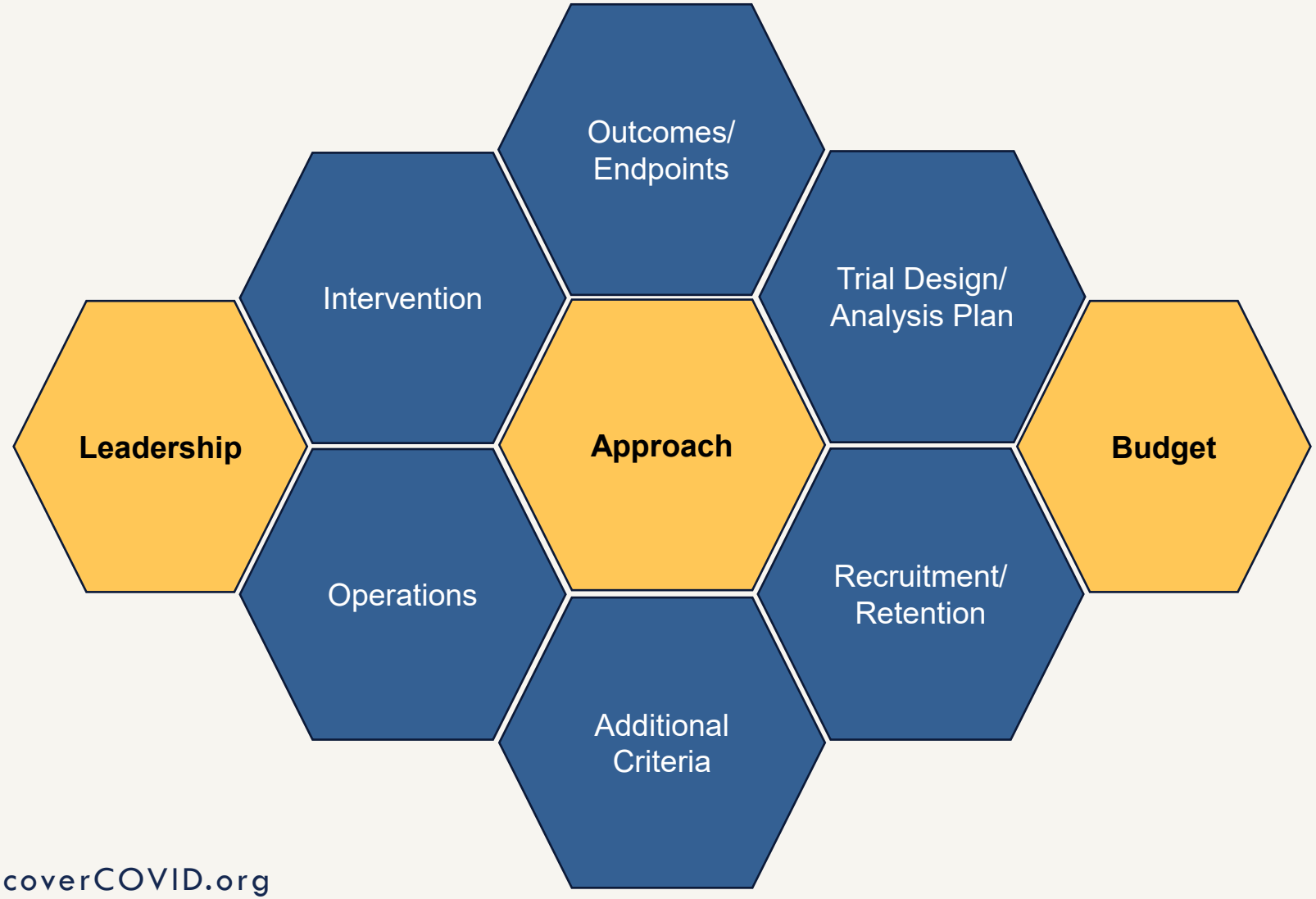
Patient-centered, participants as partners

National scale with Inclusive, diverse participation & community engagement

Platform protocols, standardized methodologies, and common data elements

Adaptive approaches based on emerging science

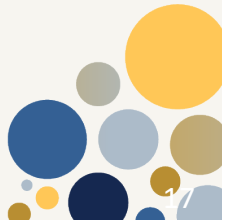
Review Criteria



Additional

- Strong strategies proposed to address potential implications of participant co-enrollment in the RECOVER observational cohort study and other clinical trials.
- Results of the study will have potential applicability to clinical practice.

Other Transactions Authority (OTA) Discussion



Other Transaction Authority (OTA) Framework

An Other Transaction Authority provides the NIH ***greater flexibility to identify and engage nontraditional research partners****, to* ***engage traditional partners in new ways****, and negotiate terms and conditions that will* ***concentrate their efforts****,* ***spur innovation****, and* ***facilitate collaborative problem solving****.*

Defined in the negative:

- Not a grant.
- Not a contract.
- Not a cooperative agreement.

Defined in the positive:

- Is an agreement between the government and a legal entity.
- Is used primarily for R&D.
- Is funded from the NIH *(usually).*

OTA Framework Considerations

Proposal Formatting:

- Proposals **must be submitted as one single .PDF file** and should address the ROA requirements.

- Unless otherwise specified, you have flexibility to make formatting decisions if the content requirements are addressed.

- Additional content in the proposal such as bio-sketches, appendixes, or letters of support **will not count** towards page limits.

OTA Framework Considerations

Budget and Negotiation:

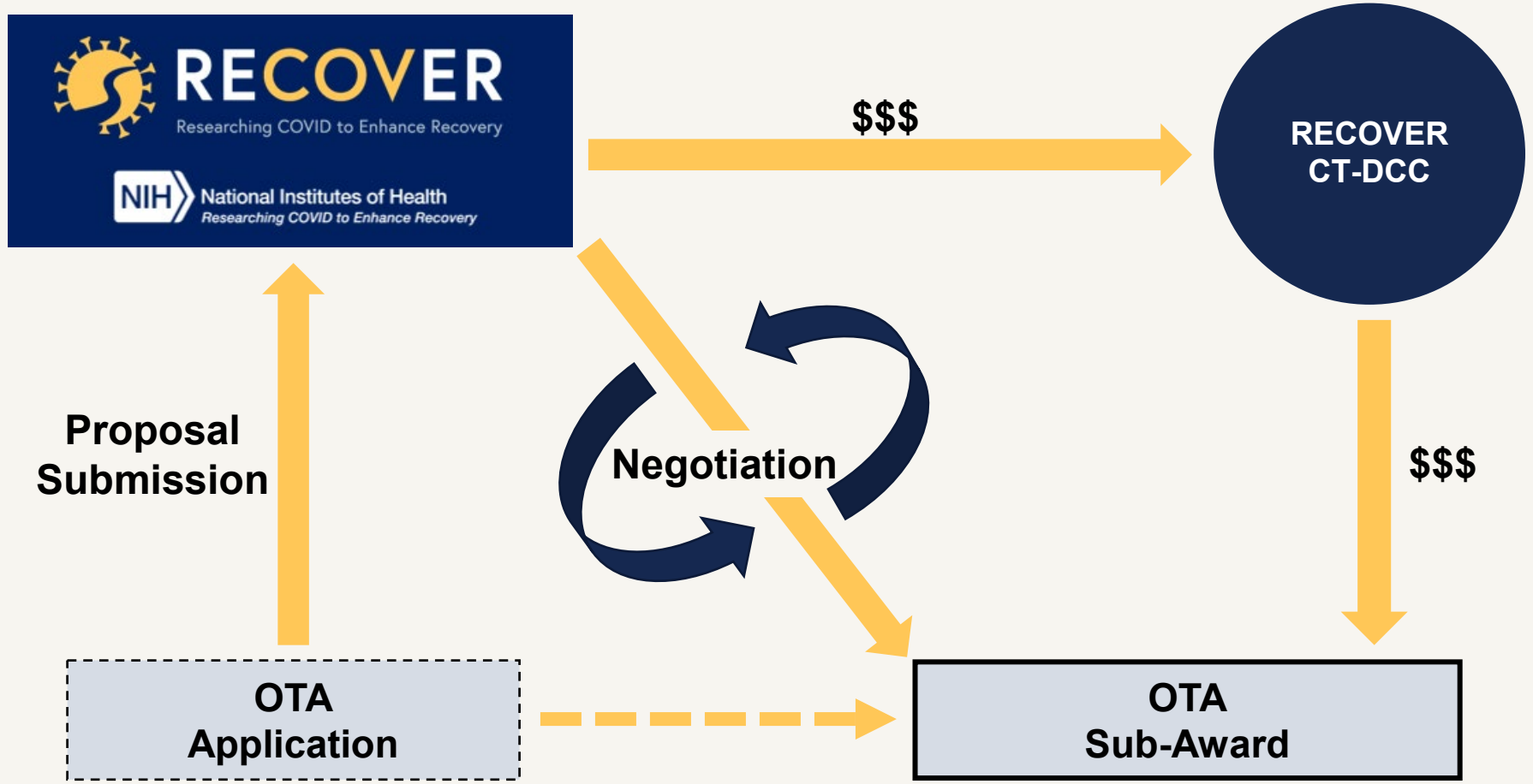
- OTA is reviewed and awarded through negotiation.
- If warranted by review outcome, we will be in touch to discuss.
- Award level is based on the requirement. No predetermined budget has been established for awards.
- Refer to ROA for specific budget instructions, R&R form recommended but not required.
- Awards expected to be issued as sub-awards of the CT-DCC.

OTA Framework Considerations

What do we mean by negotiation?

- Unlike with a grant, you can expect that we will work with you to revise elements of what we fund.
- You should aim to propose what you believe you can readily offer at the time of submission.
- The CT-DCC is able to provide multiple services (e.g., sites, statistical services, IRB, DSMB support, etc).
- The application submitted is not your final opportunity to propose elements to be funded.

OTA Framework Considerations



Submission Information

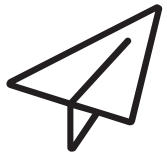
- For best consideration applications are due by **May 19th, 2022.**
- Submit to NHLBI_OTA@mail.nih.gov by an authorized business official of your institution.
- Address financial, administrative, and technical programmatic questions to NHLBI_OTA@mail.nih.gov.
- Reference **OTA-21-015H** in the title of all inquiries.

Applications will be accepted after May 19th and may be given future review consideration but, at this time, that is not guaranteed.

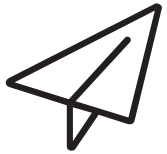
Q&A

Please Post Questions in the Q&A Box

Submitting Additional Questions



For any questions related to the **Clinical Trials ROA**, you can reach out to **NHLBI_OTA@mail.nih.gov**.



As a reminder, questions that we did not cover today will be shared in an FAQ document.

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Closing Remarks & Next Steps

Staying connected with the RECOVER Initiative

Visit the RECOVER Initiative website https://recovercovid.org/.

View the RECOVER Clinical Trials ROA https://recovercovid.org/docs/RECOVER_CT_ROA.pdf

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