NCI's Clinical Trials Reporting Program (CTRP)

NIH NATIONAL CANCER INSTITUTE

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What is NCI's Clinical Trials Reporting Program (CTRP)?

- Comprehensive database containing regularly updated information, including accrual, on interventional trials and observational studies either directly or indirectly funded by NCI
- Supports NCI clinical trials portfolio management
  - Identify gaps in portfolio
  - Prioritize clinical research opportunities
- Supports registration and results reporting to ClinicalTrials.gov for NCI sponsored trials consistent with NIH policies and FDAAA (and 42 CFR Part 11)

CTRP: Key Attributes

- Consistent terminology and standardized data elements to optimize search and retrieval of cancer clinical trials information
- Inclusion of structured biomarker and disease information
- Standard representation of persons and organizations
- Identification of associated NCI awards and contracts
- Regular updates to reflect protocol amendments, as well as participating site and status changes
  - Automated NCI system data feeds update participating site information on trials directly funded and managed by NCI nightly
- Quarterly reporting of accrual, including participant-level demography

Scope of CTRP – Trial Registration

- Trials taking place in at least one NCI-Designated Cancer Center, including industrial trials
  - Interventional* clinical trials open to accrual on or after 1/1/2009
  - Observational studies open to accrual on or after 1/1/2020
- Trials sponsored by NCI, as well as trials sponsored by other entities
- More than 85% of interventional cancer clinical trials open to patient accrual in the United States found in ClinicalTrials.gov are also in CTRP (as of September 2022)

*Studies in human beings in which individuals are assigned by an investigator, based on a protocol, to receive specific interventions. Subjects may receive diagnostic, therapeutic, behavioral or other types of interventions. The assignment of the intervention may or may not be random. The individuals are followed and biomedical and/or health outcomes are assessed. (Source: http://prsinfo.clinicaltrials.gov/definitions.html and http://cancercenters.cancer.gov/documents/CCSGDataGuide508C.pdf)

CTRP – Source and Management

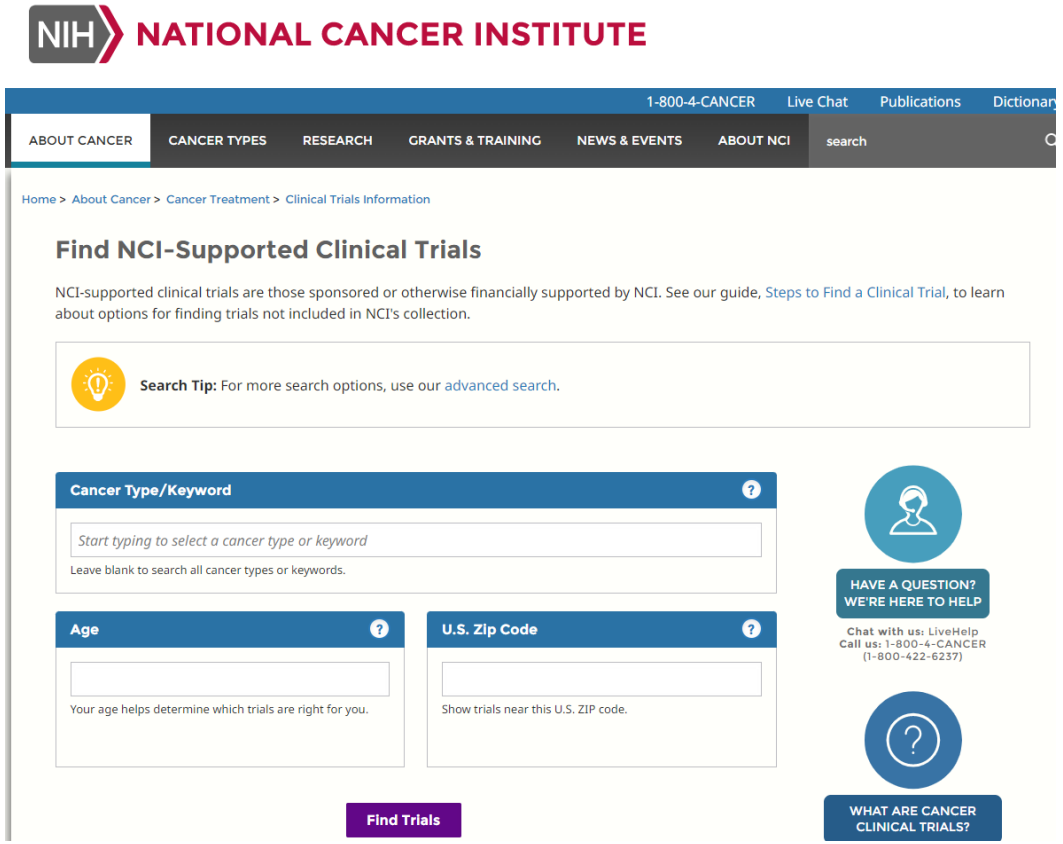
- Source
  - Internal NCI programs, including CCR[1], CTEP[2], DCP[3], e.g., NCORP[4] , NCTN[5] trials
  - NCI Awardees, e.g., trials ongoing in NCI-Designated Cancer Centers
- Management
  - Protocol document required for registration (except for industrial trials)
    - Trial abstract prepared by professional scientific abstractors
    - Coding with cancer specific terminology adds structure to disease, biomarkers, and interventions
  - Regularly updated to reflect protocol amendments, as well as participating site and status changes
  - Data abstraction harmonized with ClinicalTrials.gov
    - Supports registration with CTRP content in ClinicalTrials.gov by trial sponsors, including NCI

[1]CCR = Center for Cancer Research; [2]CTEP = Cancer Therapy Evaluation Program; [3]DCP = Division of Cancer Prevention; [4]NCORP = NCI Community Oncology Research Program; [5]NCTN = National Clinical Trials Network

CTRP – Data Distribution

- **Clinical Trials Search API (CTS API)**
  - NCI Cancer.gov website for clinical trial searching (trials.cancer.gov)
  - More than 150 academic, advocacy, and industry groups
- **Reporting**
  - CTRP-Generated Cancer Center Support Grant Data Table 4*
    - Competing applications – interventional trials
    - Non-Competing applications – interventional trials and observational studies
- **ClinicalTrials.gov**
  - Source of Registration Records for NCI-Sponsored Trials
  - PRS Upload from CTRP – facilitates use of CTRP record for use by other sponsors for registration in ClinicalTrials.gov.

*Data Table 4 serves as a report of the cancer-related hypothesis-driven clinical research studies open at the Cancer Center during a center-defined 12-month reporting period.
https://cancercenters.cancer.gov/GrantsFunding/DataGuide



CTRP and ClinicalTrials.gov

Trial Registration

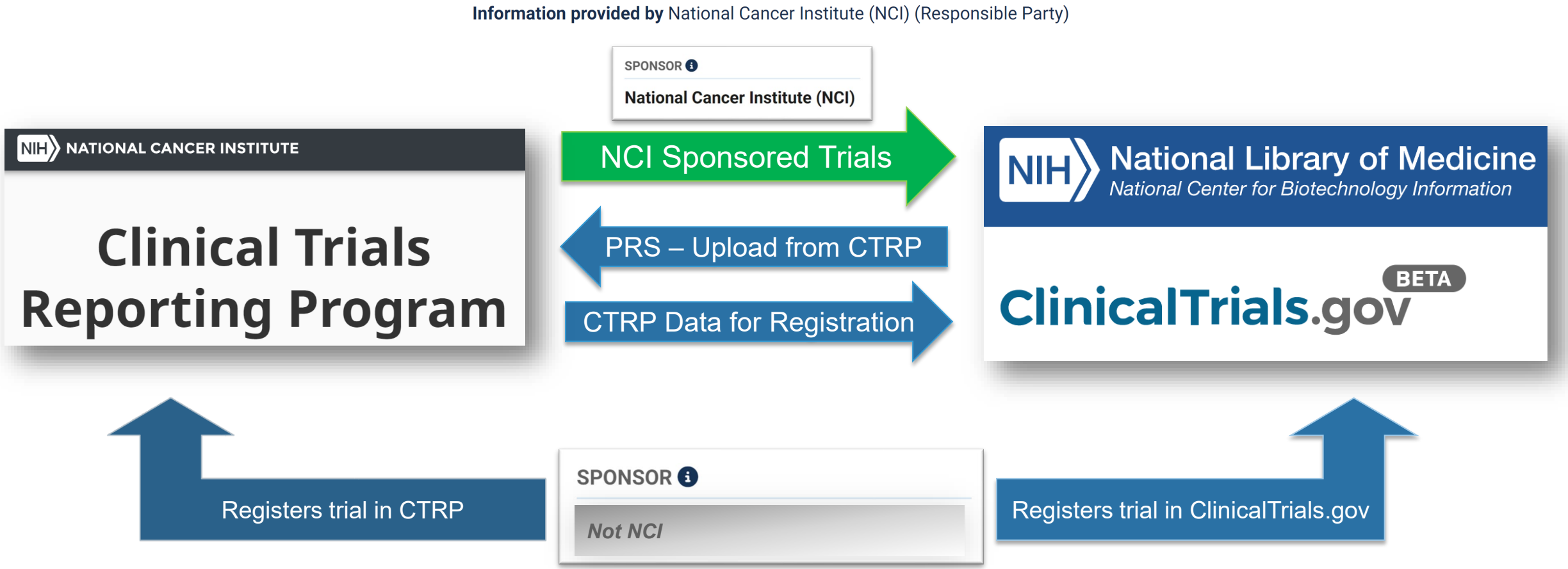
CTRP Supports ClinicalTrials.gov Registration

- When NCI is the sponsor (per FDAAA[1]):
  - NCI registers and updates the trial in ClinicalTrials.gov
- When another entity, e.g., grantee is the sponsor (per FDAAA)
  - The grantee can use the CTRP protocol abstract for ClinicalTrials.gov registration and updates using PRS Upload from CTRP[2]

[1] Food and Drug Administration Amendments Act of 2007 (FDAAA) https://www.fda.gov/regulatory-information/selected-amendments-fdc-act/food-and-drug-administration-amendments-act-fdaaa-2007

[2] Upload from CTRP https://prsinfo.clinicaltrials.gov/prs-users-guide.html#section9

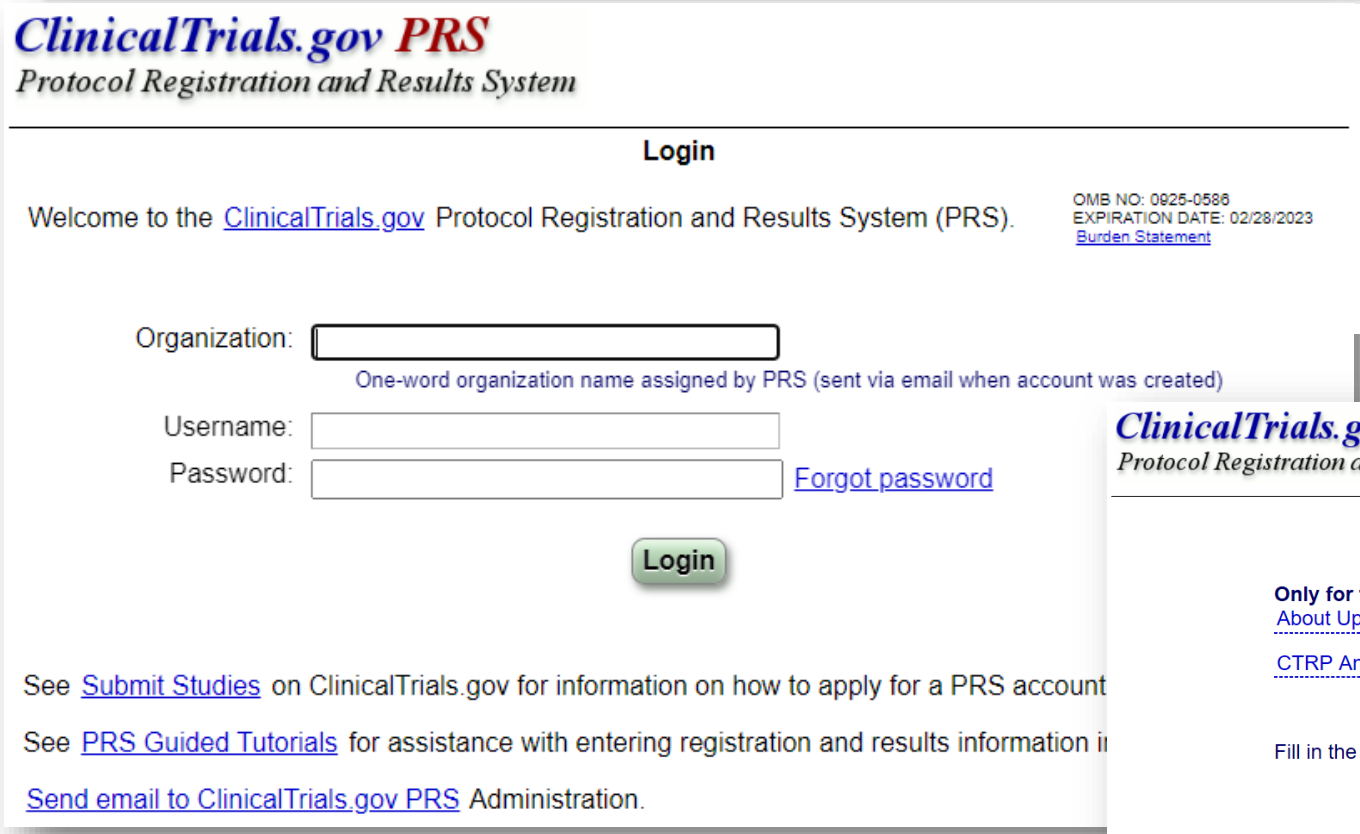
CTRP and ClinicalTrials.gov: Registration



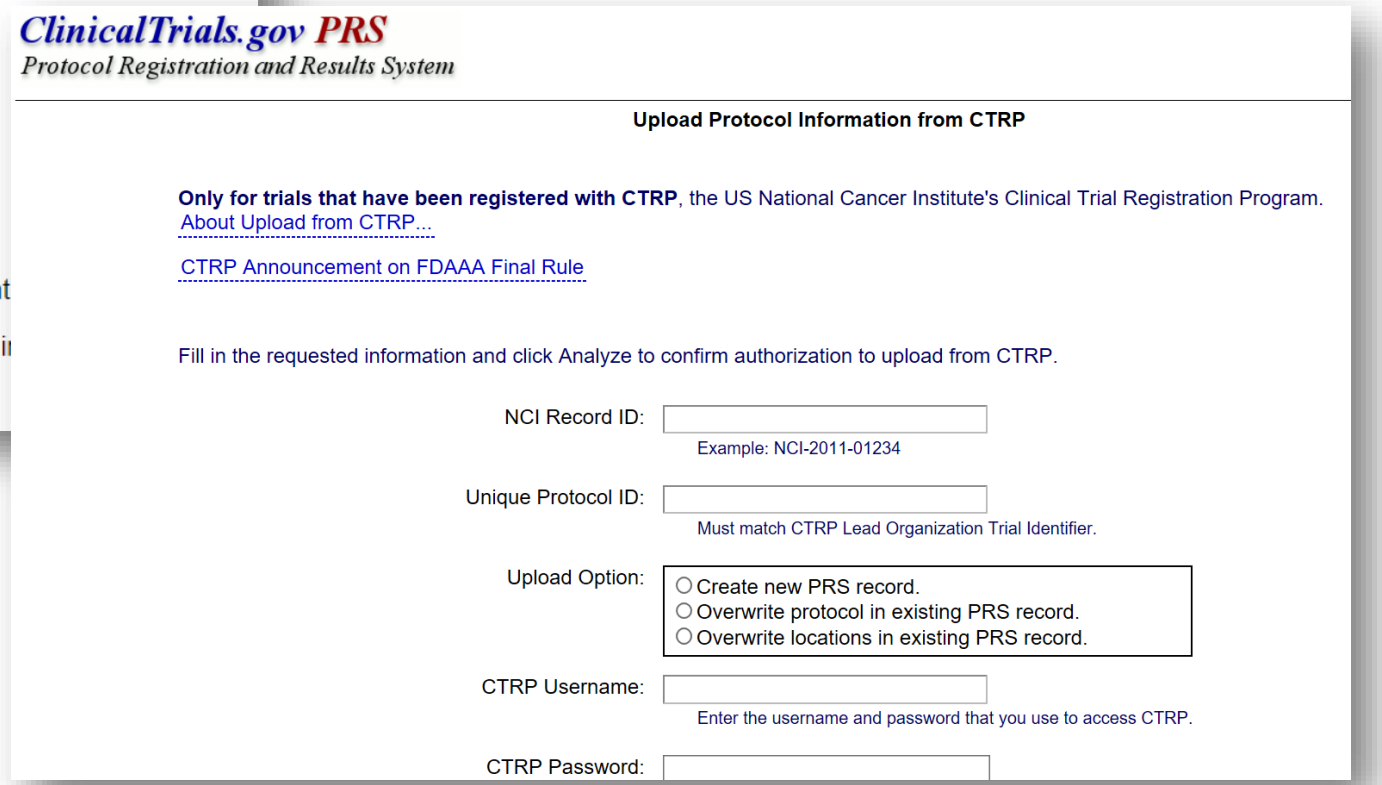
Registration of NCI Sponsored Trials in ClinicalTrials.gov

Use of CTRP Record for Non-NCI Sponsors for ClinicalTrials.gov Registration

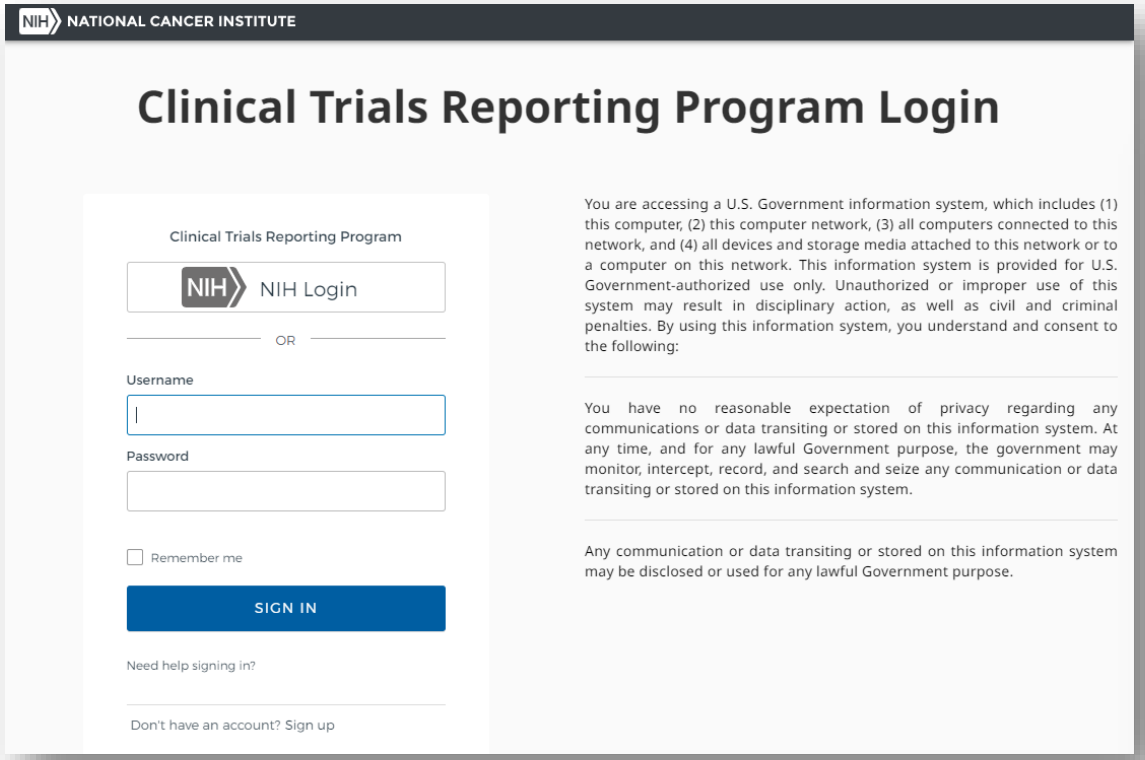
ClinicalTrials.gov: Upload from CTRP

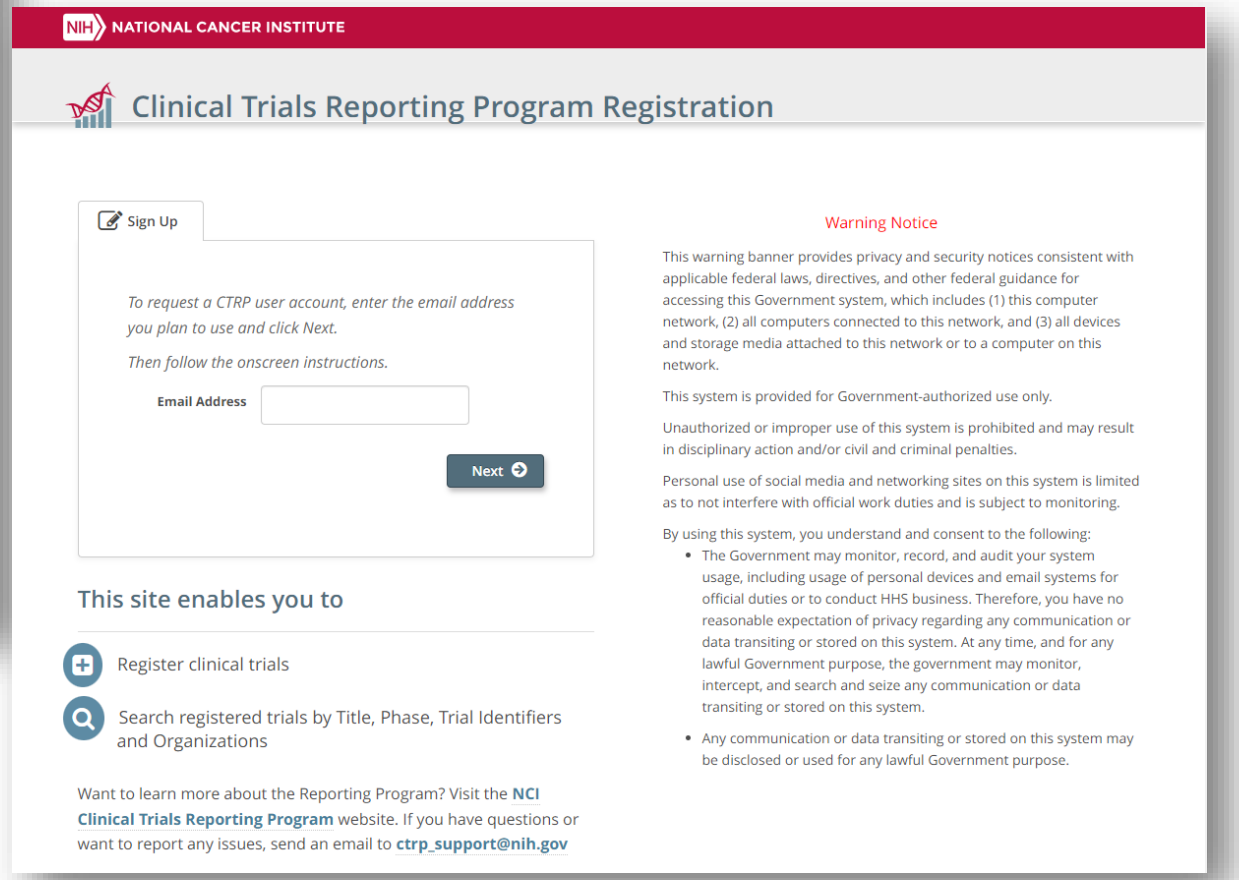


A Sponsor's PRS Admin/User logs into their Sponsor PRS account and with appropriate CTRP access can use the PRS Upload from CTRP feature



CTRP User Account





A CTRP user account is required to use the PRS Upload from CTRP feature – available to NCI-designated Cancer Centers and NCI grantees

Note – Each NCI-Designated Cancer Center has a CTRP Administrator who approves new CTRP accounts for their Center

Challenges of Trial Registration: Master Protocol Research Program (MPRP)[1]

- May involve testing multiple interventions against multiple targets
- Single MPRP may increase efficiency, e.g., shared infrastructure for screening patients for eligibility
- Clinical trial repositories, e.g., CTRP and ClinicalTrials.gov were designed for traditional clinical trials and pose challenges for the registration and reporting of MPRPs

[1] Williams RJ, Dobbins JD, Tse T, et al. Approach for Reporting Master Protocol Study Designs on ClinicalTrials.gov: Qualitative Analysis. BMJ 2022; 377: e067745

NCI-MATCH Trial (Molecular Analysis for Therapy Choice) EAY171 NCT02465060

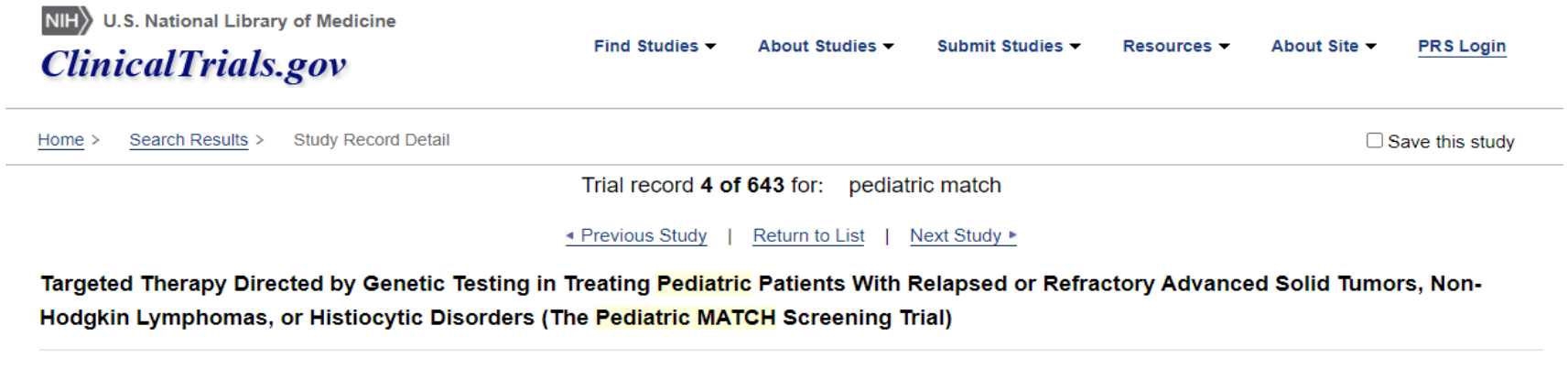
- Patients assigned to treatment with a single drug based on the genetic changes found during screening of their tumors
- Several treatment arms are open at any given time, each enrolling participants whose tumors have a specific genetic change
- Initially registered in CTRP and ClinicalTrials.gov as one trial
- Challenges with one record:
  - Does not display status changes of individual arms opening and closing to accrual
  - Does not support results reporting until all arms with all interventions are closed
  - Includes all eligible disease sites, biomarkers, and interventions studied in all arms

NCI-MATCH Trial (Molecular Analysis for Therapy Choice) EAY171 NCT02465060

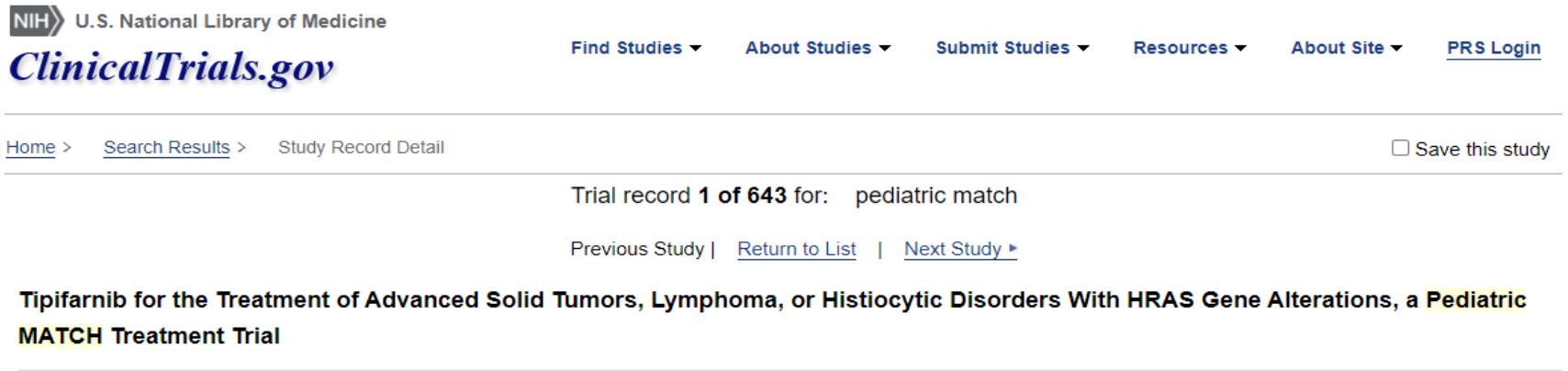
- When a treatment arm closes, the primary outcome of that arm has been reached
- MATCH has 39 treatment arms; waiting until the primary completion date of the last arm would have delayed reporting for many years
- To support results reporting, NCI, the sponsor:
  - Modified the original record to become the “screening” record
  - Registered each treatment arm in a separate record
  - Has been reporting primary outcomes as the primary completion dates for each of the treatment arms/records is reached

Pediatric MATCH

- Screening and treatment are registered in separate records in CTRP and ClinicalTrials.gov at initiation
- Screening record and the treatment record reference each other in the title



Screening Record



One of Several Treatment Records

CTRP and ClinicalTrials.gov

- CTRP includes all trials directly or indirectly funded by NCI
- Upon receipt of a protocol document, CTRP abstracts a file with data suitable for registration in ClinicalTrials.gov
- Use of Upload from CTRP in PRS provides the sponsor, when not NCI, access to the record for ClinicalTrials.gov registration
- Master Protocol Research Program (MPRP) trials pose challenges for registration and results reporting
  - Separate registration records for screening and treatment may help simplify registration and results reporting
    - Referencing the other parts of the clinical trial, e.g., in the title or body of the registration record, can be helpful.
    - Coordination needed when different sponsors manage separate components of the MPRP

Resources & Contact Information

- Search CTRP content on NCI Website: trials.cancer.gov
- NCI's Clinical Trials Search API: clinicaltrialsapi.cancer.gov
- How to obtain a CTRP User Account (required for Upload from CTRP): trials.nci.nih.gov/registry/registerUser.action
- Contact CTRP: ctrp_support@mail.nih.gov

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