

The Cancer Genomics Cloud powered by Seven Bridges: a secure and scalable cloud-based platform to access, share and analyze multi-omics datasets



May 14, 2021

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Agenda

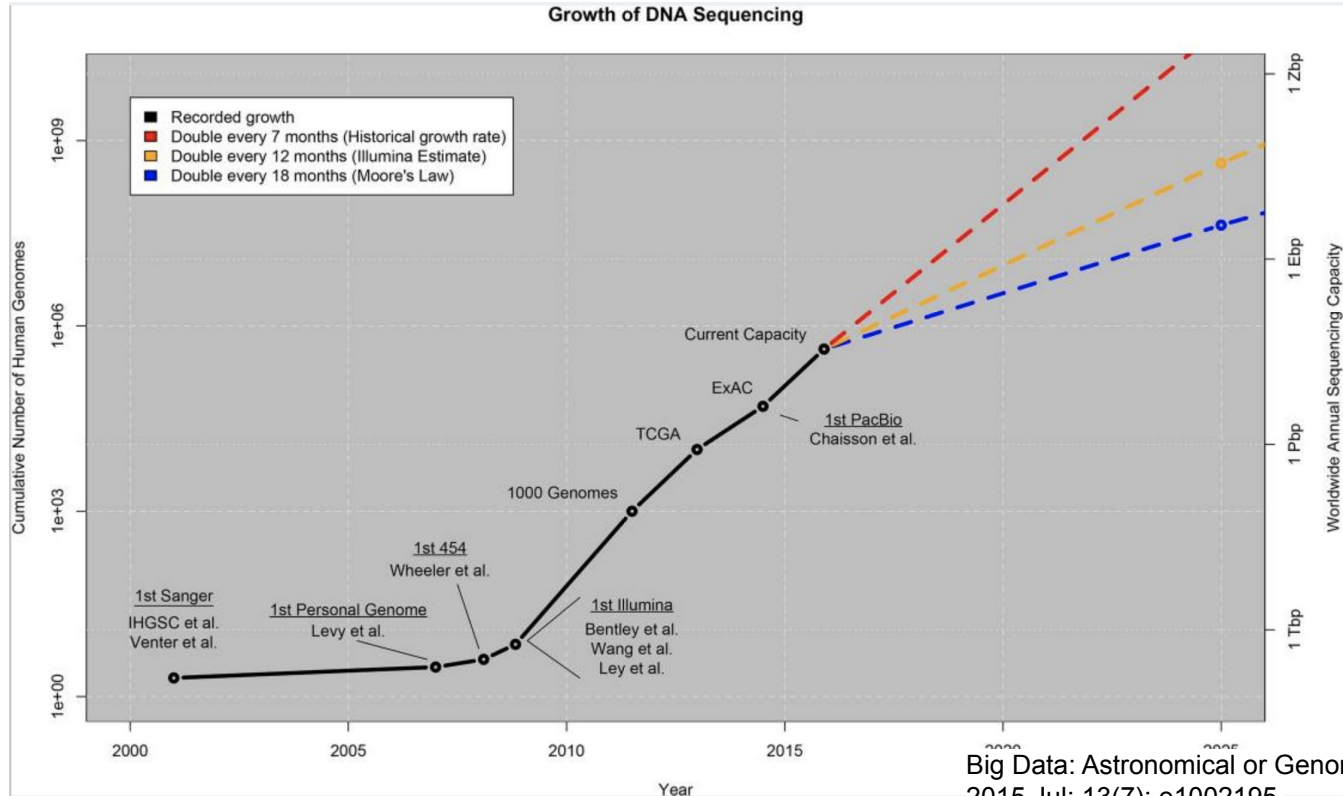
- Background
- Access multi-omics datasets in the CGC
- Use Case: Exploring the human proteome - Analysis of CPTAC datasets
 - Features to control cloud costs
- Ongoing Projects - New workflows for novel science
- Questions/Discussion



Background



Explosion of genomics data with ease of sequencing



Big Data: Astronomical or Genomical? Stephens et al; [PLoS Biol.](https://doi.org/10.1371/journal.pbio.1002195) 2015 Jul; 13(7): e1002195.

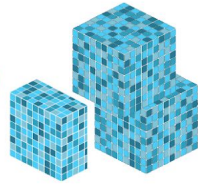
Increasingly large datasets bring challenges to data analysis

NATIONAL CANCER INSTITUTE THE CANCER GENOME ATLAS

TCGA BY THE NUMBERS

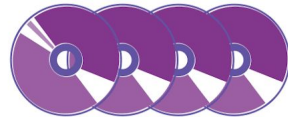
TCGA produced over

2.5
PETABYTES
of data



To put this into perspective, **1 petabyte** of data is equal to

212,000
DVDs



TCGA data describes



33
DIFFERENT
TUMOR TYPES

...including

10
RARE
CANCERS

...based on paired tumor and normal tissue sets collected from



11,000
PATIENTS

...using

7

DIFFERENT
DATA TYPES



www.cancer.gov/ccg

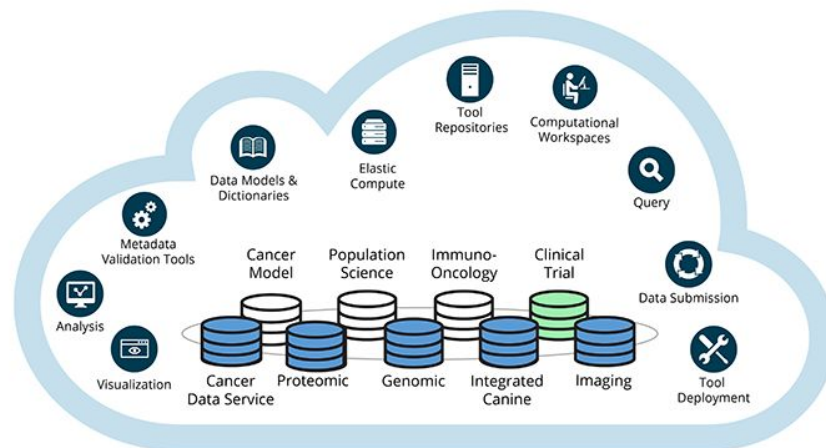
The Seven Bridges Cancer Genomics Cloud (CGC)

NCI Cancer Research Data Commons (CRDC)



A Cloud Resource within the NCI Cancer Research Data Commons for secure storage, sharing & analysis of petabytes of public, multi-omic cancer datasets

<https://datacommons.cancer.gov/cancer-research-data-commons>



Authentication & Authorization



Legend

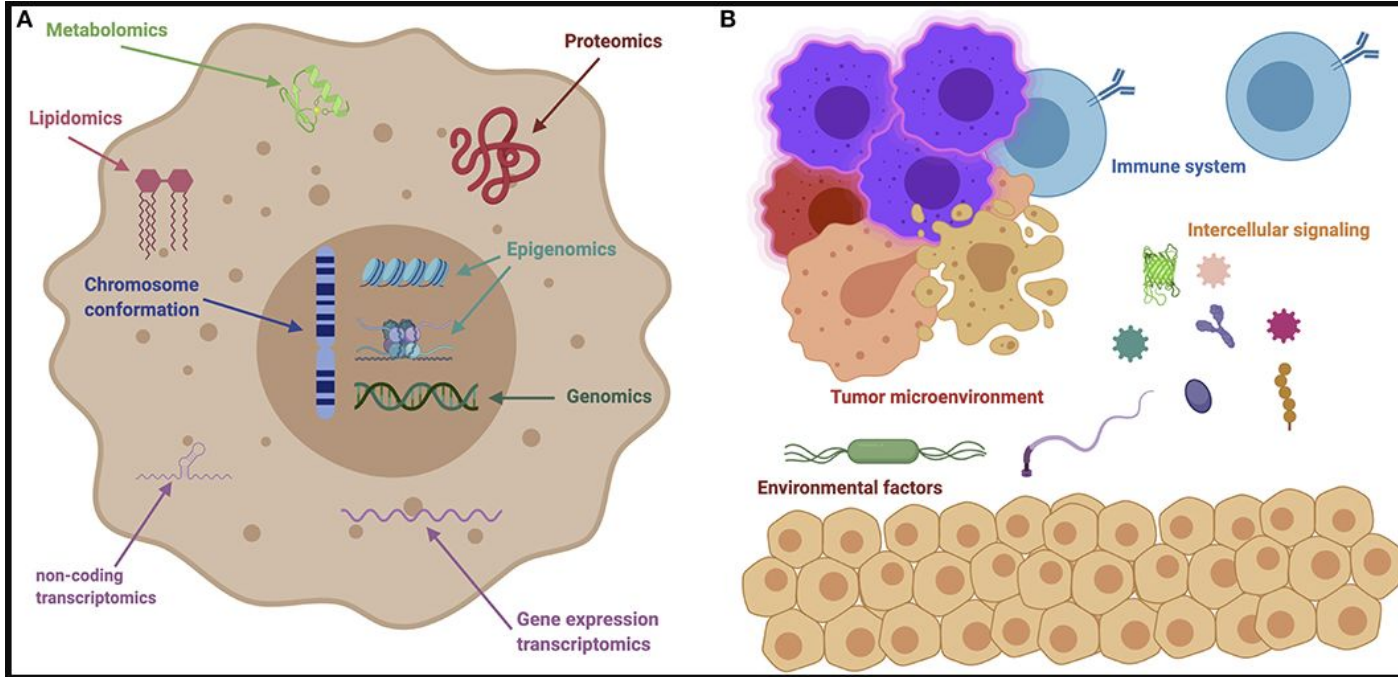
-  Available to researchers
-  Development
-  Future nodes



Data Contributors & Consumers



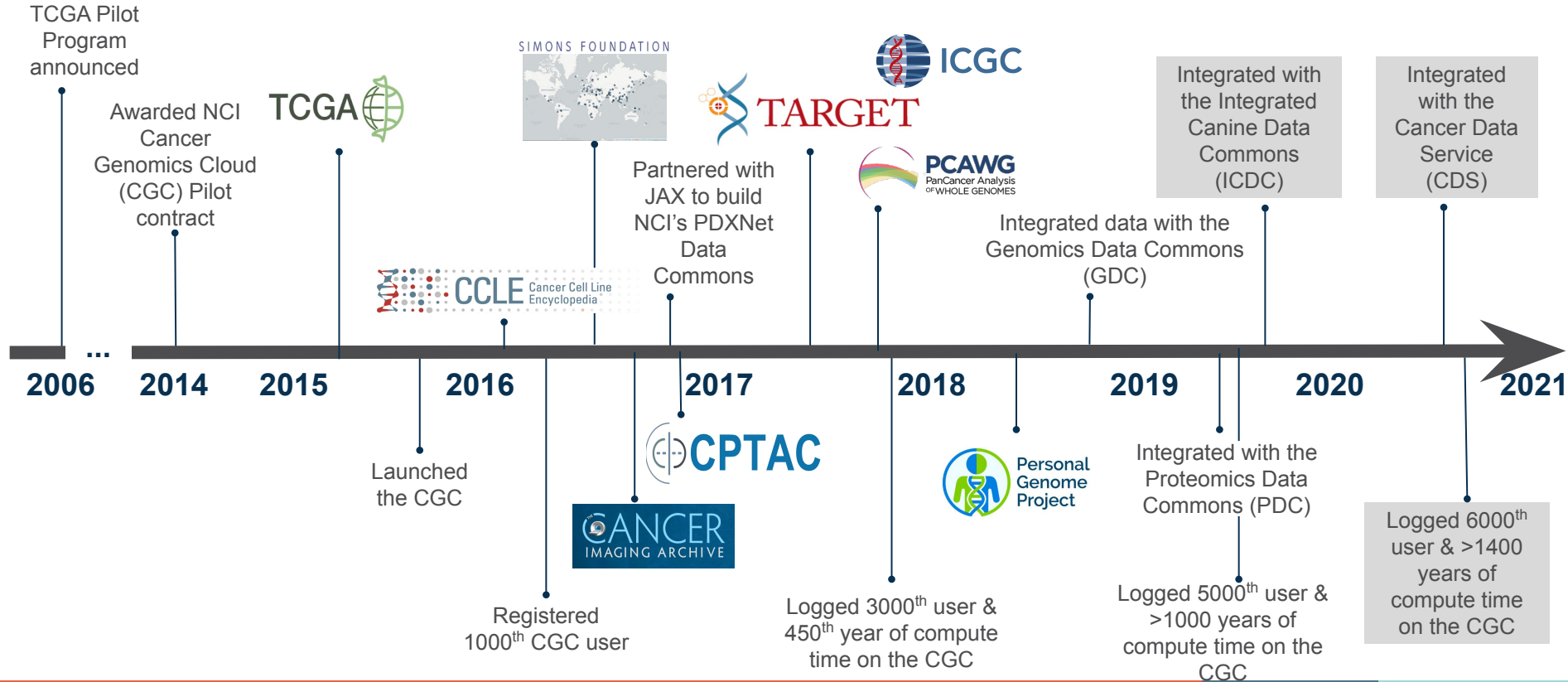
Multi-omic data is critical for cancer research



Cancer is a complex disease!

Comprehensively understanding the full picture of a research question requires examining multiple modalities

Growth of the Cancer Genomics Cloud Ecosystem



The CGC democratizes complex analyses in a FAIR data ecosystem

Cloud-based Environment for Collaborative Research and Bioinformatics Data Analysis

- A stable, secure, and highly **customizable** cloud storage and computing platform
- Promotes a **F**indable, **A**ccessible, **I**nteroperable and **R**eusable (FAIR) data ecosystem
- A user-friendly portal for **collaborative** analysis of petabytes of public data alongside private data
- An optimized venue for **reproducible data analysis** using validated tools and pipelines



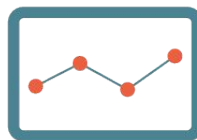
Easy data
management



Secure
collaboration &
managed billing



Flexible & fully
reproducible
methods



Optimized
bioinformatics
algorithms



Scalable
computation



Extensible &
developer
friendly tools

Accelerating cancer research

- Detect aberrant splice junctions and splicing profiles across patient populations
- Identify neoantigens arising from novel gene fusion events
- Profile miRNA expression across patient populations
- Conduct HLA typing to identify neoantigens
- Compare viral infection patterns across patient populations
- Detect novel gene fusions from RNA-Seq data
- Identify cis-regulatory region variants across patient populations
- ...and much more



High impact publications on the CGC

nature communications

Explore our content ▾ Journal information ▾

nature > nature communications > articles > article

Article | [Open Access](#) | Published: 02 June 2020

AGO-bound mature miRNAs are oligouridylated by TUTs and subsequently degraded by DIS3L2

Acong Yang, Tie-Juan Shao, Xavier Boffill-De Ros, Chuanjiang Lian, Patricia Villanueva, Lisheng Dai & Shuo Gu 

Nature Communications **11**, Article number: 2765 (2020) | [Cite this article](#)

2767 Accesses | **1** Citations | **11** Altmetric | [Metrics](#)

Genome Medicine

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Research | [Open Access](#) | Published: 17 February 2020

The pan-cancer landscape of prognostic germline variants in 10,582 patients

[Ajay Chatrath](#), [Roza Przanowska](#), [Shashi Kiran](#), [Zhangli Su](#), [Shekhar Saha](#), [Briana Wilson](#), [Takaaki Tsunematsu](#), [Ji-Hye Ahn](#), [Kyung Yong Lee](#), [Teressa Paulsen](#), [Ewelina Sobierajska](#), [Manjari Kiran](#), [Xiwei Tang](#), [Tianxi Li](#), [Pankaj Kumar](#), [Aakrosh Ratan](#) & [Anindya Dutta](#) 





Genome Medicine **12**, Article number: 15 (2020) | [Cite this article](#)

2844 Accesses | **1** Citations | **78** Altmetric | [Metrics](#)

Oncogene
<https://doi.org/10.1038/s41388-020-01507-5>

ARTICLE

Genetic alterations of *SUGP1* mimic mutant-*SF3B1* splice pattern in lung adenocarcinoma and other cancers

[Samar Alsafadi](#) ^{1,2} · [Stephane Dayot](#)² · [Malcy Tarin](#)¹ · [Alexandre Houy](#) ² · [Dorine Bellanger](#)² · [Michele Cornella](#)² · [Michel Wassef](#)^{3,4} · [Joshua J. Waterfall](#) ^{1,2} · [Erik Lehnert](#)⁵ · [Sergio Roman-Roman](#)¹ · [Marc-Henri Stern](#) ² · [Tatiana Popova](#)²

nature genetics

Explore our content ▾ Journal information ▾

nature > nature genetics > articles > article

Article | Published: 07 January 2021

Conservation of copy number profiles during engraftment and passaging of patient-derived cancer xenografts

Xing Yi Woo, Jessica Giordano, Anuj Srivastava, Zi-Ming Zhao, Michael W. Lloyd, Roebi de Bruijn, Yun-Suh Suh, Rajesh Patidar, Li Chen, Sandra Scherer, Matthew H. Bailey, Chieh-Hsiang Yang, Emilio Cortes-Sanchez, Yuanxin Xi, Jing Wang, Jayamanna Wickramasinghe, Andrew V. Kossenkov, Vito W. Rebecca, Hua Sun, R. Jay Mashl, Sherri R. Davies, Ryan Jeon, Christian Frech, Jelena Randjelovic, Jacqueline Rosains, Francesco Galimi, Andrea Bertotti, Adam Lafferty, Alice C. O'Farrell, Elodie Modave, Diether Lambrechts, Petra ter Brugge, Violeta Serra, Elisabetta Marangoni, [Rania El Botty](#), Hyunsoo Kim, Jong-Il Kim, Han-Kwang Yang, Charles Lee, Dennis A. Dean II, Brandi Davis-Dusenbery, Yvonne A. Ervard, James H. Doroshov, Alana L. Welm, Bryan E. Welm, Michael T. Lewis, Bingliang Fang, Jack A. Roth, Funda Meric-Bernstam, Meenhard Herlyn, Michael A. Davies, Li Ding, Shunqiang Li, Ramaswamy Govindan, Claudio Isella, Jeffrey A. Moscow, Livio Trusolino, Annette T. Byrne, Jos Jonkers, Carol J. Bult, Enzo Medico , Jeffrey H. Chuang , PDXNET Consortium & EuroPDX Consortium - Show fewer authors

Nature Genetics **53**, 86–99(2021) | [Cite this article](#)

618 Accesses | **42** Altmetric | [Metrics](#)

Abstract

Patient-derived xenografts (PDXs) are resected human tumors engrafted into mice for preclinical studies and therapeutic testing. It has been proposed that the mouse host affects

CANCER RESEARCH

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Tumor Biology and Immunology

Systematic Establishment of Robustness and Standards in Patient-Derived Xenograft Experiments and Analysis

Yvonne A. Ervard, Anuj Srivastava, Jelena Randjelovic, The NCI PDXNet Consortium, James H. Doroshov, Dennis A. Dean II, Jeffrey S. Morris, and Jeffrey H. Chuang

DOI: 10.1158/0008-5472.CAN-19-3101 Published June 2020 

Find a growing list of publications at: <https://www.cancergenomicscloud.org/publications>



Participating in open standards groups helps make us more FAIR



Global Alliance
for Genomics & Health
Collaborate. Innovate. Accelerate.



BioCompute
Objects



COMMON
WORKFLOW
LANGUAGE



docker



How do I get an account on the CGC?

- Sign up with your email
 - <https://www.cancergenomicscloud.org/>
- Option to connect with eRA Commons to access controlled data
- **\$300 of pilot funding** to get your project started
- Comprehensive online documentation and training resources
- Technical support from a team of scientists, bioinformaticians, and engineers

Log in



Log in with eRA Commons










[Log in with username and password](#)

New to the CGC? [Create an account](#)



Access multi-omics datasets in the CGC

Access and search large public datasets on the CGC

Dataset	Description	Experimental setup	File types
 TCGA	Rich dataset of tumor and normal tissues from 11,000 patients, covering 33 cancer types	WES, RNAseq, miRNAseq, methylation, genotyping, ATACseq, imaging, WGS, ..	BAM, VCF, MAF, TXT, TSV, SVS, XML
 TARGET	Dataset of genomic changes in childhood cancers	RNASeq, WGS, WES, miRNAseq	BAM, MAF, TSV, VCF, XLSX, TXT
 CANCER IMAGING ARCHIVE	Imaging data from many 21 tumor types	Imaging	DCM
 CPTAC	Proteomics of 10 tumor types and associated genomic data	Proteomics, WGS, WES, RNAseq	BAM, TSV, VCF, mzML.gz, mzid.gz, raw, tar.gz
 International Cancer Genome Consortium	Consortium of many datasets, 20 studies on CGC	WGS, RNASeq	BAM, VCF
 CCLE Cancer Cell Line Encyclopedia	Dataset of 1457 cancer cell lines	WGS, WES, RNAseq	BAM
 SIMONS FOUNDATION	Genome sequencing of 130 populations	WGS	BAM, VCF
 Personal Genome Project	Crowdsourced genomics, datasets from 10 individuals	WGS, WGBS, RNAseq, methylation	BAM, FASTQ, IDAT, TBI, VCF
 HUMAN CELL ATLAS	Single-cell genomics of healthy tissues	scRNASeq	FASTQ

CGC connects with several CRDC data repositories

Coming soon!

REPOSITORIES



Cancer Data Service (CDS)

Store and share NCI-funded data that are not hosted elsewhere to further advance scientific discovery across a broad range of research areas.



Clinical Trial Data Commons (CTDC)

Store and share data from NCI Clinical Trials. The resource is expected to launch in 2020.



Genomic Data Commons (GDC)

Share, analyze, and visualize harmonized genomic data, including TCGA, TARGET, and CPTAC.



Imaging Data Commons (IDC)

Share, analyze, and visualize multi-modal imaging data from both clinical and basic cancer research studies.



Integrated Canine Data Commons (ICDC)

Share data from canine clinical trials, including the PRE-medical Cancer Immunotherapy Network Canine Trials (PRECINCT) and the Comparative Oncology Program.



Proteomic Data Commons (PDC)

Share, analyze, and visualize proteomic data, such as CPTAC and The International Cancer Proteogenome Consortium (ICPC).

Harmonized Cancer Datasets
Genomic Data Commons Data Portal

Get Started by Exploring:

- Projects
- Exploration
- Analysis
- Repository

Search: e.g. BRAF, Breast, TCGA-BLCA, TCGA-A5-ADG2

Data Portal Summary Data Release 27.0 - October 29, 2020

PROJETS	PRIMARY SITES	CASES
67	68	84,392

FILES	GENES	MUTATIONS
596,758	23,399	3,287,299

Cases by Major Primary Site

- Adrenal Gland
- Bladder
- Bone
- Bone Marrow
- Brain
- Breast
- Cervix
- Colorectal



NIH NATIONAL CANCER INSTITUTE
Proteomic Data Commons

HOME BROWSE ANALYZE

62 Studies

24 TB Data volume

81,275 Data files

> 357 M Spectra

NIH NATIONAL CANCER INSTITUTE
Cancer Research Data Commons

REPOSITORY
Cancer Data Service (CDS)

Enabling secure and flexible storage and sharing of data

Learn More

Overview

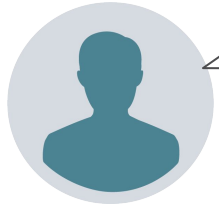
The Cancer Data Service (CDS) is a data repository under the Cancer Research Data Commons (CRDC) infrastructure for storing cancer research data generated by NCI funded programs. The CDS provides secure and authorized storage and data sharing capabilities in the cloud for studies that can fall under either of the categories below:



Use Case: Exploring the human proteome - Analysis of CPTAC datasets

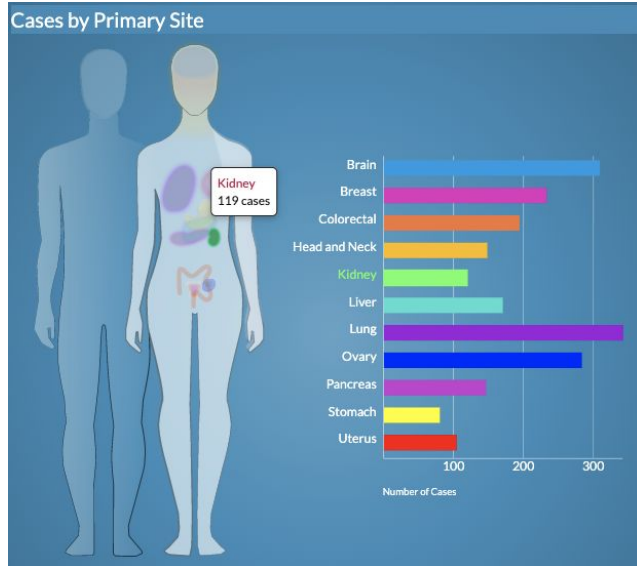


Proteogenome characterization of ccRCC



How do I analyze the spectra data of the proteome in clear cell renal cell carcinoma cases?

Kidney cancer is among the 10 most common cancers in both men and women
~73,000 new cases with >14,000 deaths in 2020
<https://seer.cancer.gov/statfacts/html/kidrp.html>



- Proteome and phosphoproteome data from the ccRCC tumors is available in PDC along with peptide spectrum analyses (PSMs) and protein summary reports from the CPTAC common data analysis pipeline (CDAP).
- Using the CGC, process high-throughput data-dependent acquisition (DDA) tandem mass spectrometry data acquired from peptides labeled with TMT tags.
- The multiplexed labeling, as applied in the CPTAC program, allows for differential quantitation across multiple tumor/normal samples.

Typical User Flow

Create a Project

Organizational unit
within the CGC

Find datasets of interest

Many ways to find and
bring in data:

- Data Browser
- Desktop
uploader
- Command line
uploader
- Volumes

Bring/Build tools or workflows

Tools, workflows, and
software packages

- Public Apps
Gallery
- Tools or
workflows
wrapped in
CWL
- R packages
- Python libraries

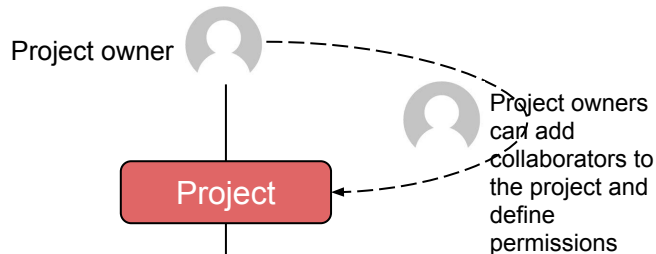
Analyze

Specify how an
analysis will be run

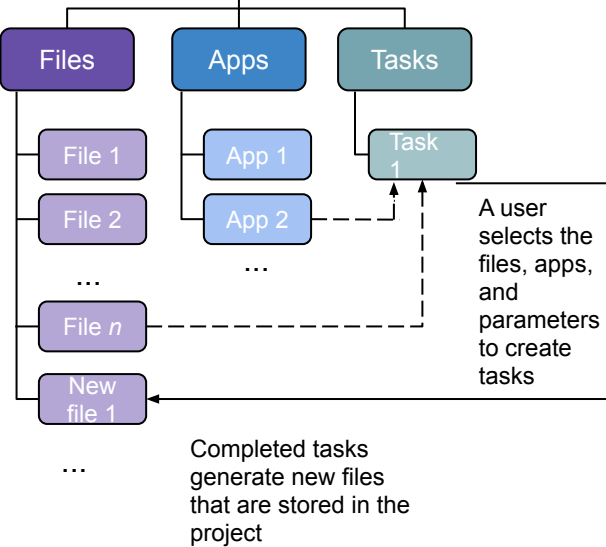
- Task page
- Notebooks in
RStudio or
JupyterLab



Projects organize files, methods, and results



Also known as *workspaces* or *sandboxes*
Easily manage collaborators and permissions



Create a project ✕

Name

Project URL:
<https://cgc.sbgenomics.com/u/sailakss/containers-and-workflows-webinar-demo>

Billing Group
Pilot Funds (sailakss) ▾

Location ⓘ
AWS (us-east-1) ▾

Execution settings:
Spot Instances ⓘ On
Memoization (WorkReuse) ⓘ Off
Network Access ⓘ

Block network access
Executions within the project won't have network access

Allow network access
Executions will have unrestricted network access

This project will contain **CONTROLLED** Data. ⓘ

Cancel Create

- Projects are configurable, e.g.
- Customizable billing group - where costs should be attributed
 - Cloud resources (AWS or GCP)
 - **Spot** (or **preemptible**) instances
 - Memoization - Intermediate file retention
 - Using S3 or Glacier storage

Collaborate and share results quickly and easily

The screenshot displays the CGC Platform interface for a project named "Purdue-Bioinformatics-Class". The top navigation bar includes "Projects", "Data", "Public Apps", "Public projects", "Developer", and "Staff". The user "sailakss" is logged in. The project page is divided into three main sections:

- Description:** Contains a welcome message, an overview of projects, a list of actions users can take within the project, and a note about recording project details. It also includes a citation section.
- Members:** Shows the current member "sailakss" with roles "Write, Copy, Execute, Admin". It features a motivational message: "Don't work alone. The best research happens in teams." and a prominent "Invite new members" button.
- Analyses:** Includes a search bar and a "Tasks" tab. The "Data Cruncher" section is currently empty, with a message: "Your executions will appear here. Before you start, learn more about them."

Description Tags

Welcome to your new project!

Projects are the core building blocks of the CGC Platform. Each project corresponds to a distinct scientific investigation, serving as a container for its data, analysis pipelines, and results. Projects are shared only by designated project members.

Within your project, you can:

- Start exploring public datasets straight away
- Install your tools on the CGC and create workflows
- Upload your own private data and analyze it along with public datasets
- Collaborate securely with other researchers

Please record the details of your project here, such as its aims, experimental context, and any other ideas that you'd like to share with your project members. Remember that details of each pipeline execution you run on the CGC are logged on the task page. This notepad is just for your own notes.

You can also use [markdown](#) here to add formatting to your notes.

Good luck with your research! If you get stuck, take a look at the [Knowledge Center](#)

The Seven Bridges CGC Team

Citation

Project participants agree to acknowledge the funding for the CGC in all publications and external presentations, as follows:

"The Seven Bridges Cancer Research Data Commons Cloud Resource has been funded in whole or in part with Federal funds from the National Cancer Institute, National Institutes of Health, Contract No. HHSN261201400008C and IDIQ Agreement No. 17X146 under Contract No. HHSN261201500003I and 75N011019D00024."

Members

Email notifications

sailakss OWNER
Write, Copy, Execute, Admin

Don't work alone.
The best research happens in teams.

[Invite new members](#)

Share your tools, data, and ideas with collaborators

Analyses

 🔍

Tasks Data Cruncher

Your executions will appear here.
Before you start, learn more about them.

Billing groups

Clear advantages for collaboration and interoperability. Aligned to temporal dynamics of research funding.

Allow users to distribute costs appropriately per function, topic, lab, etc

Use different funding sources (e.g. R24, Pilot Funds, credit card)

SB can reimburse for task failure due to external factors

The screenshot shows the 'Billing Group settings: Pilot Funds (sailakss)' page. The navigation bar includes 'Projects', 'Data', 'Public Apps', 'Public projects', 'Developer', and 'Staff'. The left sidebar has 'Current Usage' and 'Spending Details'. The main content area is divided into 'Info', 'Remaining credits', 'Pilot funds', 'Total charges', 'Analysis usage', and 'Storage usage'.

Info	
Organization	Seven Bridges Genomics
Creator	sailakss
Primary contact	Seven Bridges Genomics
Address	One Broadway, 14th Fl. , Massachusetts, United States

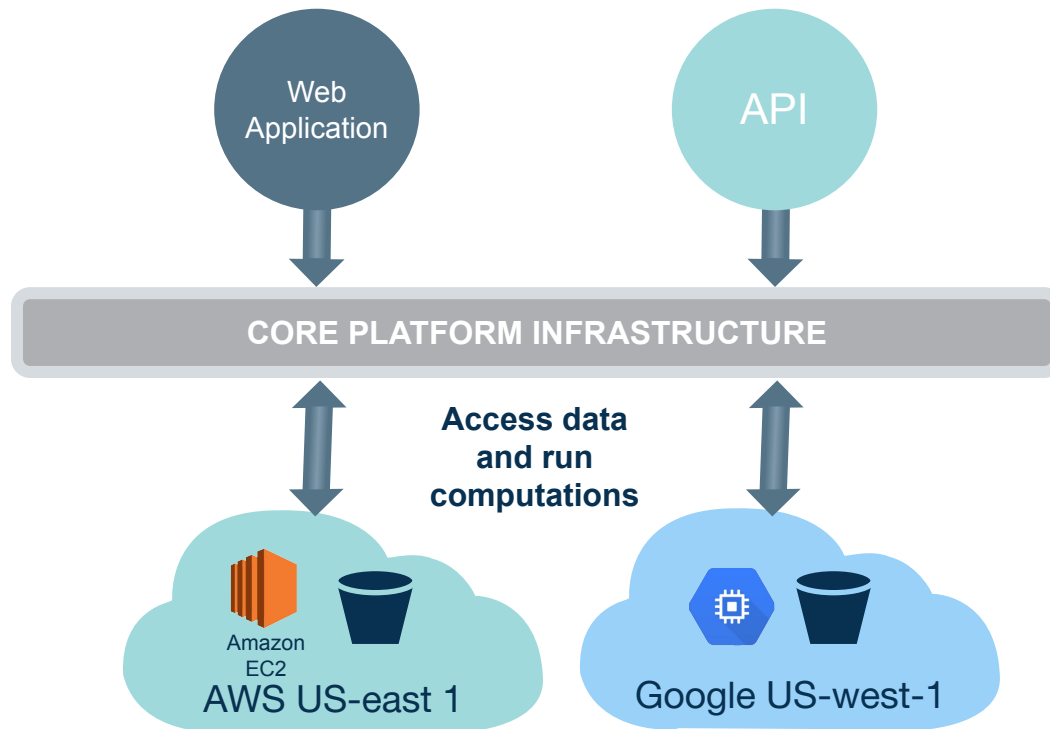
Remaining credits	\$ 296.45
Pilot funds	\$ 300.00
Total charges	\$ 3.55

Analysis usage		Storage usage	
Analysis charges	\$ 0.84	Storage charges	\$ 2.71
Tasks	\$ 0.13	Active	\$ 0.28
Data Cruncher analyses	\$ 0.71	Downloaded	\$ 0.00
		Storage deduction	\$ 0.00

Instance limits
Total number of instances that can be run in parallel


Current usage: **0** of **80**


Multi-cloud implementation on the CGC




Memoization allows use of previously computed results

COMPLETED Whole Genome Sequencing - BWA + GATK 4.0 (with Metrics) 

 Get support

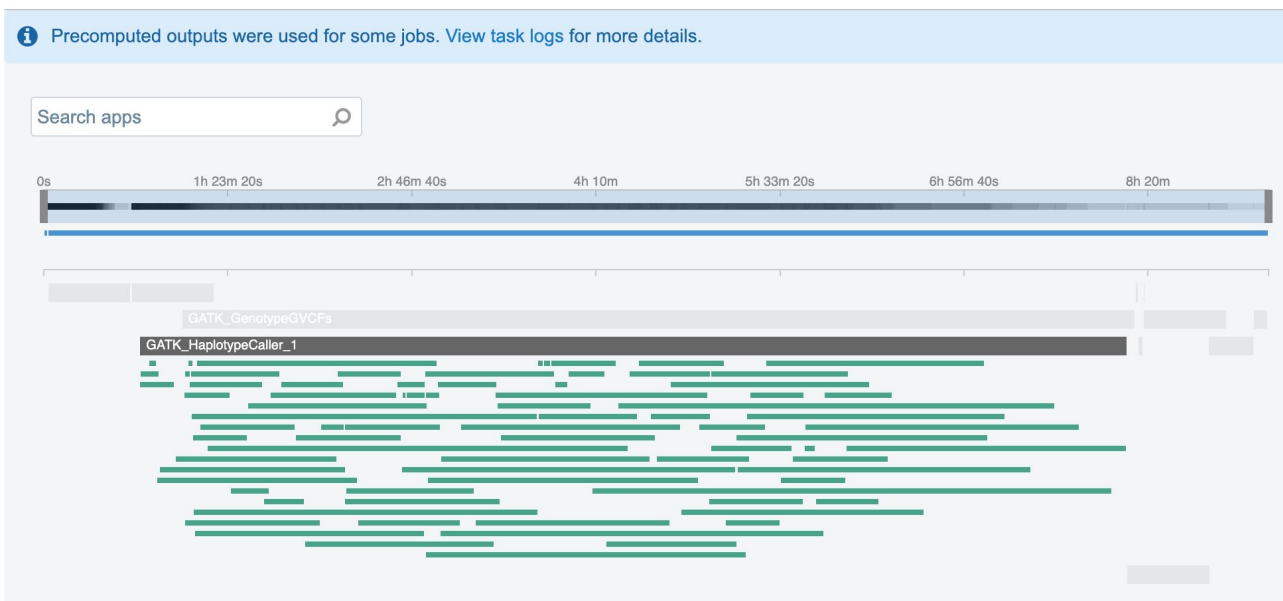
 View stats & logs

 Edit and rerun

Executed on Aug. 23, 2020 22:12 by [sinan.yavuz_demo](#)

Preemptible Instances: **On**  | Memoization (WorkReuse): **On**  | Price: **\$3.45**  | Duration: **9 hours, 14 minutes** 

▼ App: [Whole Genome Sequencing - BWA + GATK 4.0 \(with Metrics\)](#) - Revision: 0



User Flow

Create a Project

Organizational unit
within the CGC

Find datasets of interest

Many ways to find and
bring in data:

- Data Browser
- Desktop
uploader
- Command line
uploader
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Bring/Build tools or workflows

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software packages

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Gallery
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workflows
wrapped in
CWL
- R packages
- Python libraries

Analyze

Specify how an
analysis will be run

- Task page
- Notebooks in
RStudio or
JupyterLab



Different options to bring data

Add files to "Purdue-Bioinformatics-Class" ✕

Case Explorer and Data Browser **Public Files** Projects Your Computer FTP / HTTP Data Tools Volumes Import from a manifest file

Files

Category: All ▾ Type: All ▾ Sample ID: All ▾ Tags: All ▾ + Copy to Project

<input type="checkbox"/>	Name	Size	Type
<input type="checkbox"/>	1000G_omni2.5.b37.vcf <small>(GATK_RESOURCE_BUNDLE) SUGGESTED</small>	192.1 MIB	VCF
<input type="checkbox"/>	1000G_omni2.5.hg19.sites.vcf <small>(GATK_RESOURCE_BUNDLE)</small>	199.1 MIB	VCF
<input type="checkbox"/>	1000G_omni2.5.hg38.vcf <small>(GATK_RESOURCE_BUNDLE)</small>	198.8 MIB	VCF
<input type="checkbox"/>	1000G_phase1.indels.b37.vcf <small>(GATK_RESOURCE_BUNDLE) SUGGESTED</small>	226.7 MIB	VCF
<input type="checkbox"/>	1000G_phase1.indels.hg19.vcf <small>(GATK_RESOURCE_BUNDLE)</small>	230.8 MIB	VCF
<input type="checkbox"/>	1000G_phase1.snps.high_confidence.b37.vcf <small>(GATK_RESOURCE_BUNDLE) SUGGESTED</small>	6.8 GIB	VCF
<input type="checkbox"/>	1000G_phase1.snps.high_confidence.hg19.sites.vcf <small>(GATK_RESOURCE_BUNDLE)</small>	6.9 GIB	VCF
<input type="checkbox"/>	1000G_phase1.snps.high_confidence.hg38.vcf <small>(GATK_RESOURCE_BUNDLE)</small>	6.9 GIB	VCF
<input type="checkbox"/>	20.intervals <small>(TEST)</small>	0.0 KIB	INTERVALS
<input type="checkbox"/>	C835.HCC1143.2.converted.pe_1.fastq <small>(WES) TUMOR SAMPLE</small>	7.1 GIB	FASTQ
<input type="checkbox"/>	C835.HCC1143.2.converted.pe_2.fastq <small>(WES) TUMOR SAMPLE</small>	7.1 GIB	FASTQ
<input type="checkbox"/>	C835.HCC1143.2.converted.realigned.base_recalibrated.bam <small>(WES) TUMOR SAMPLE GRCH37</small>	5.3 GiB	BAM
<input type="checkbox"/>	C835.HCC1143.2.converted.realigned.base_recalibrated.bam.bai <small>(WES) TUMOR SAMPLE GRCH37</small>	2.9 MIB	BAI
<input type="checkbox"/>	C835.HCC1143.BL_4.converted.pe_1.fastq <small>(WES) NORMAL SAMPLE</small>	6.2 GIB	FASTQ

- * Public files
- * Case Explorer & Data Browser
- * Projects (that you are a member of)
- * FTP/HTTP (signed URLs)
- * Data tools
 - Command Line Uploader
 - Desktop Uploader
 - SBFS: Seven Bridges File System
 - API upload
- * Volumes
- * Import from manifest: ICDC/PDC

Find open access TCGA data with Data Browser

TCGA GRCh38 New query Edited

Create new query

Queries ▾

Search by ID

Copy files to project



Case 183 ▾ Investigation 1 ▾ File 4 ▾

Export ▾

Details

Analytics

- File**
- TCGA.PAAD.somaticsnpier.0ca0083b-4c11-47f2-b672-d74911f50b89.DR-10.0.somatic.m...
 - TCGA.PAAD.mutect.fea333b5-78e0-43c8-bf76-4c78dd3fac92.DR-10.0.somatic.maf.gz
 - TCGA.PAAD.muse.93c525cc-655c-4c1c-b590-18d851473f68.DR-10.0.somatic.maf.gz
 - TCGA.PAAD.varscan.d5737b1c-afc7-4fe7-8a30-e1bc9b44fa26.DR-10.0.somatic.maf.gz

Details for
TCGA.PAAD.somaticsnpier.0ca0083b-4c11-47f2-b672-d74911f50b89.DR-10.0.somatic.maf...

TCGA GRCh38

Access level ⓘ	Open
Created datetime ⓘ	2017-12-01T23:52:47.832Z
Data category ⓘ	Simple Nucleotide Variation
Data format ⓘ	MAF
Data type ⓘ	Masked Somatic Mutation
Experimental strategy ⓘ	WXS

Connections

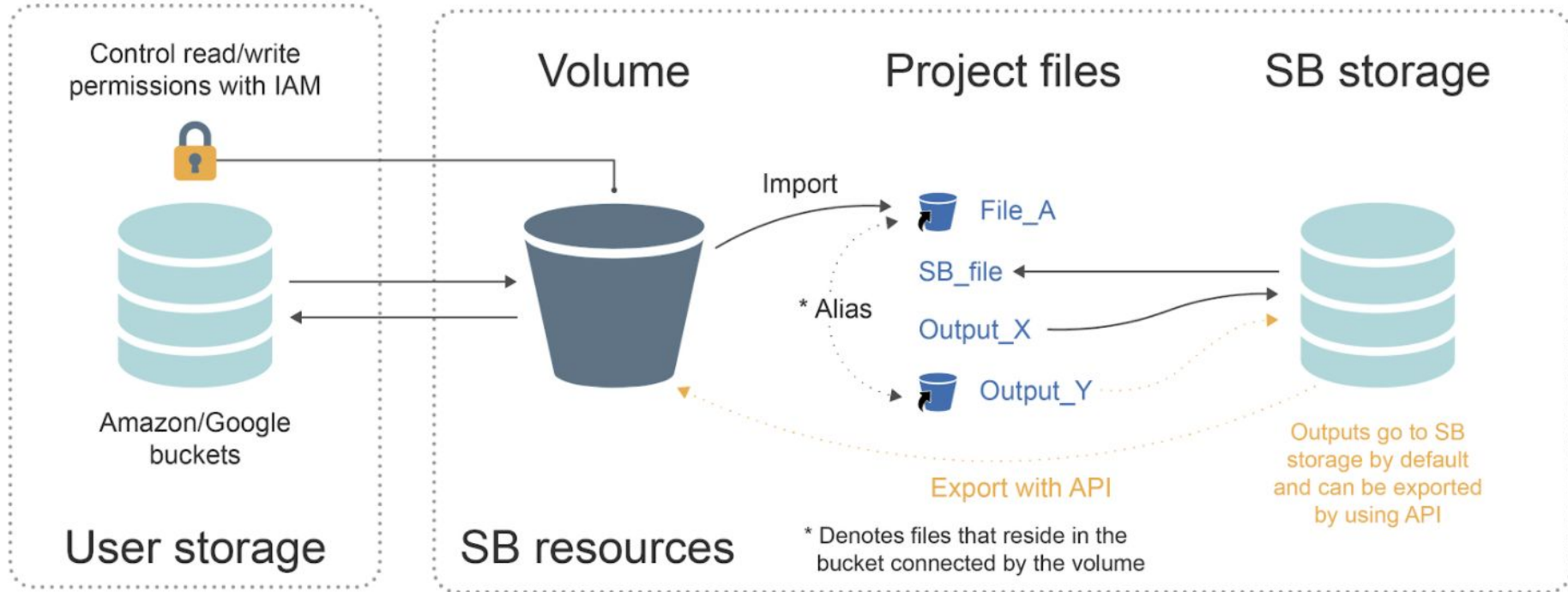
Inbound: Case 183

- 01775b06-5836-469c-8537-120cb8cc94e9
- 02dbd5fa-e31f-4486-8df8-5b851f2e92bd

Outbound:

No outbound connections

Easily connect cloud volumes



Enabling multi-omic research on the CGC through integrating with the PDC, ICDC, CDS

NATIONAL CANCER INSTITUTE Proteomic Data Commons



NATIONAL CANCER INSTITUTE
Integrated Canine Data Commons

Cancer Data
Service
(CDS)

1. User starts on PDC/ICDC/SRA (for CDS) portal to identify cohort of files
2. User downloads **files manifest** of selected cohort



CANCER GENOMICS
CLOUD

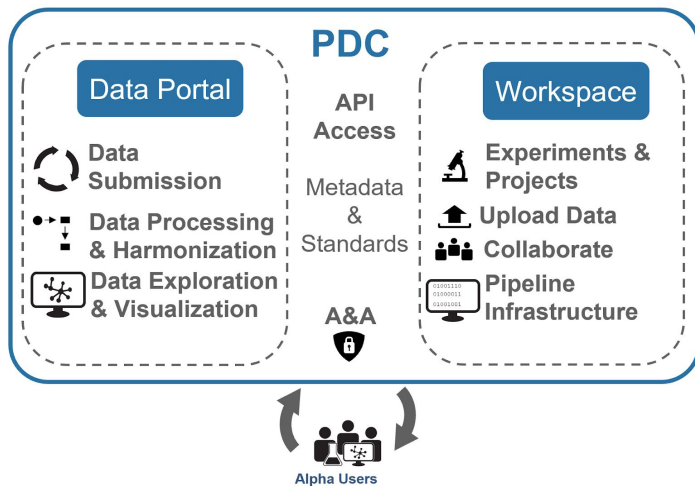
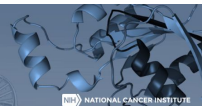
1. User moves to CGC, creates a project
 - a. Files → Add files → Import from a manifest
2. User prompted to upload the manifest from the PDC/ICDC/CDS
3. Data files from PDC/ICDC/CDS copied into user's project
4. Additional metadata accessed via Data Cruncher notebook

Links to doc pages to import data from: [PDC](#), [ICDC](#), [CDS](#)



Integration with the Proteomic Data Commons (PDC)

Proteome Data Commons – democratize access to cancer-related proteomic datasets



Courtesy of Izumi Hlnkson

CGC integration with PDC enables both tool developers and researchers to take full advantage of rich data resources

Proteomics tools can run quickly and efficiently in the cloud, lowering barriers for usage

NIH NATIONAL CANCER INSTITUTE
Proteomic Data Commons

Login

Q e.g. BRCC3, 05BR003, kinase, PDC0001 X

NCI is pleased to release these data to the public. Some data are under an EMBARGO for publication and/or citation.

HOME BROWSE ANALYSIS SUBMIT DATA ABOUT

62
Studies

24 TB
Data volume

81,275
Data files

> 357 M
Spectra

> 1 M
Peptides

15,007
Proteins



CANCER GENOMICS CLOUD
SEVEN BRIDGES

How to import data from PDC

NATIONAL CANCER INSTITUTE Proteomic Data Commons

Search: BRCC3, 058R003, kinase

HOME BROWSE ANALYSIS SUBMIT DATA ABOUT HELP

FILTERS

Start searching by selecting a facet

ANALYTICAL FRACTIONS

- Chloroquine
- Phosphoproteome
- Proteome

DISEASE TYPES

- Metastatic Colorectal Cancer
- Breast Invasive Carcinoma
- Clear Cell Renal Cell Carcinoma
- Core Anaplastic Carcinoma
- Ovarian Serous Cystadenocarcinoma
- Rectal Adenocarcinoma

EXPERIMENT TYPES

- 178424
- Label Free
- TMT10

Files (37170)

Total files: 37170

Study #	File Name #	Run Metadata	Project #	Data Category	File Type	Access	File Size (MB)	Downloadable
✓ PCT_284781_Kshay	GUO2_1138410_001C_SW	SMO2170263	1	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,648,476	Yes
✓ PCT_284781_Kshay	GUO2_1138410_002_SW	SMO2170263	2	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,713,976	Yes
✓ PCT_284781_Kshay	GUO2_1138410_003_SW	SMO2170263	3	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,631,773	Yes
✓ PCT_284781_Kshay	GUO2_1138410_004_SW	SMO2170263	4	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,818,437	Yes
✓ PCT_284781_Kshay	GUO2_1138410_005_SW	SMO2170263	5	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,642,704	Yes
✓ PCT_284781_Kshay	GUO2_1138410_006_SW	SMO2170263	6	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,697,989	Yes
✓ PCT_284781_Kshay	GUO2_1138410_007_SW	SMO2170263	7	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,601,017	Yes
✓ PCT_284781_Kshay	GUO2_1138410_008_SW	SMO2170263	8	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,769,847	Yes
✓ PCT_284781_Kshay	GUO2_1138410_009_SW	SMO2170263	9	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,548,046	Yes
✓ PCT_284781_Kshay	GUO2_1138410_010_SW	SMO2170263	10	Quantitative digital mass of blood samples	Raw Mass Spectra	Proprietary	1,604,427	Yes

Download File Manifest

CANCER GENOMICS CLOUD

FTP / HTTP Data Tools Volumes **Import from a manifest file**

Import files from **Proteomic Data Commons (PDC)**

- Proteomic Data Commons (PDC)
- Integrated Canine Data Commons (ICDC)
- Cancer Data Service (CDS)

Upload a .csv manifest file
See information about .csv manifest files

Select file No file chosen

Some NCI data are under an **EMBARGO** for publication and/or citation. For more details, visit the [NCI Proteomic Data Commons](#) for the study of interest.

Researchers can also bring private data to co-analyze with public resources.

Import

User Flow

Create a Project

Organizational unit
within the CGC

Find datasets of
interest

Many ways to find and
bring in data:

- Data Browser
- Desktop uploader
- Command line uploader
- Volumes

**Bring/Build
tools or
workflows**

Tools, workflows, and
software packages

- Public Apps Gallery
- Tools or workflows wrapped in CWL
- R packages
- Python libraries

Analyze

Specify how an
analysis will be run

- Task page
- Notebooks in RStudio or JupyterLab



FragPipe: A complete proteomics pipeline

Cell

Volume 179, Issue 4, 31 October 2019, Pages 964-983.e31



Resource

Integrated Proteogenomic Characterization of Clear Cell Renal Cell Carcinoma

David J. Clark ^{1, 32}, Saravana M. Dhanasekaran ^{2, 32}, Francesca Petralia ^{3, 32}, Jianbo Pan ^{1, 32}, Xiaoyu Song ^{4, 5, 32}, Yingwei Hu ^{1, 32}, Felipe da Veiga Leprevost ^{2, 32}, Boris Reva ^{3, 32}, Tung-Shing M. Lih ^{1, 32}, Hui-Yin Chang ², Weiping Ma ³, Chen Huang ⁶, Christopher J. Ricketts ⁷, Lijun Chen ¹, Azra Krek ³, Yize Li ⁸, Dmitry Rykunov ³, Qing Kay Li ¹ ... Zhidong Tu

[Show more](#) ✓

Cell

Volume 182, Issue 1, 9 July 2020, Pages 200-225.e35



Resource

Proteogenomic Characterization Reveals Therapeutic Vulnerabilities in Lung Adenocarcinoma

Michael A. Gillette ^{1, 2, 24, 27} ✉, Shankha Satpathy ^{1, 24} ✉, Song Cao ^{3, 25}, Saravana M. Dhanasekaran ^{4, 25}, Suhas V. Vasaikar ^{5, 25}, Karsten Krug ^{1, 25}, Francesca Petralia ^{6, 25}, Yize Li ³, Wen-Wei Liang ³, Boris Reva ⁶, Azra Krek ⁶, Jiayi Ji ⁷, Xiaoyu Song ⁷, Wenke Liu ⁸, Runyu Hong ⁸, Lijun Yao ³, Lili Blumenberg ⁹, Sara R. Savage ¹⁰ ... Zhiao Shi

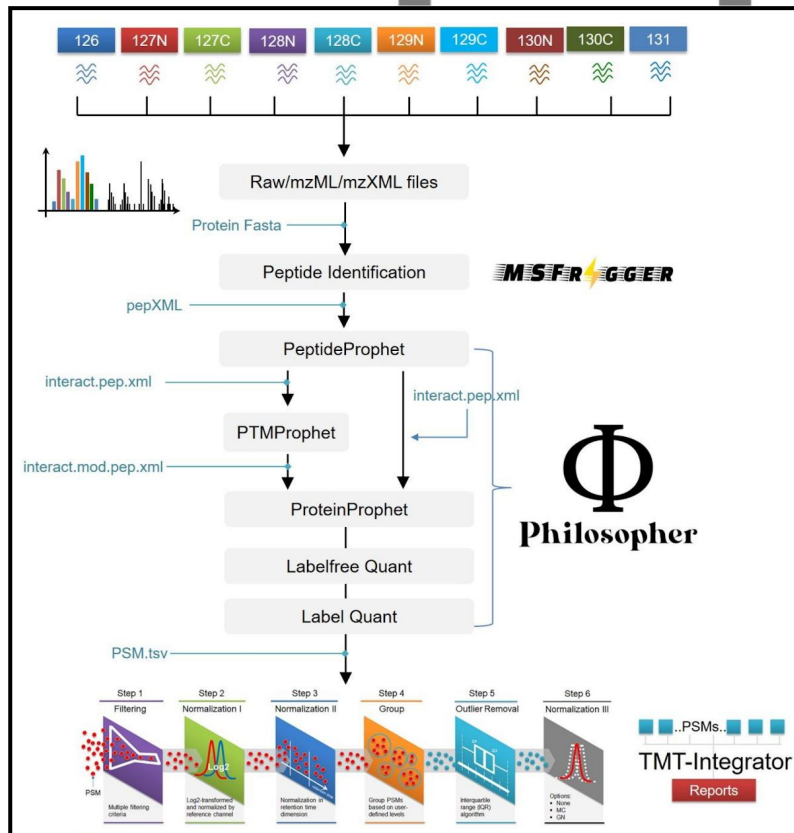


FragPipe Proteomics Pipeline



- Developed by Nesvizhskii lab at the University of Michigan
- **FragPipe**: a complete proteomics pipeline for comprehensive analysis of proteomics data which is powered by
 - **MSFragger**, an ultrafast peptide identification tool for mass spec-based proteomics
 - **Philosopher** toolkit, for post-processing MSFragger results
 - **TMT-Integrator**, a tools for integrating channel abundances from multiple TMT or iTRAQ-labeled samples and generating reports

<https://github.com/Nesvilab/FragPipe>



Find the tools you need in the Public Apps Gallery

A curated collection of **500+** bioinformatics tools & workflows

- Optimized for speed & cost in the cloud
- Fully parameterized & customizable
- Accessible via the GUI & API

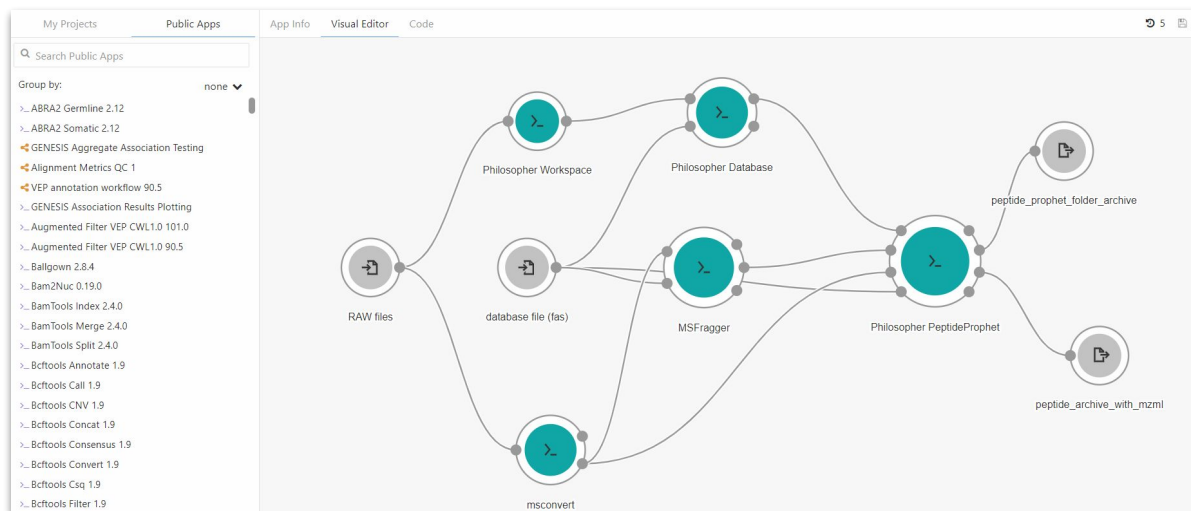
<https://cgc.sbgenomics.com/public/apps>

The screenshot displays the 'Public apps' interface. At the top, there is a search bar with the text 'Search workflows and tools', a category dropdown set to 'Proteomics', and a toolkit dropdown set to 'FragPipe'. A 'Reset search' button is also present. Below the search bar, three workflow cards are shown. Each card features a 'W' icon, a title, the 'NESVILAB' logo, the version 'FragPipe v15.0', a brief description, a 'PROTEOMICS' tag, and 'Copy' and 'Run' buttons. The first card is 'FragPipe: Convert - Identify - PeptideProphet', the second is 'FragPipe: Filter - Quant - Report', and the third is 'FragPipe: TMT Integrator and QC'.

Bring Your Own Tools & Tailor new Pipelines in the Platform with Web Composer

An intuitive and flexible software development kit for developing and porting custom tools to the platform

Conformance with community standards to ensure pipeline portability & reproducibility



COMMON
WORKFLOW
LANGUAGE



User Flow

Create a Project

Organizational unit
within the CGC

Find datasets of interest

Many ways to find and
bring in data:

- Data Browser
- Desktop
uploader
- Command line
uploader
- Volumes

Bring/Build tools or workflows

Tools, workflows, and
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Gallery
- Tools or
workflows
wrapped in
CWL
- R packages
- Python libraries

Analyze

Specify how an
analysis will be run

- Task page
- Notebooks in
RStudio or
JupyterLab



FragPipe Workflows

Dashboard Files **Apps** Tasks

PDC Workshop ⓘ





Search names and description 🔍

Category: All ▼

Toolkit: All ▼

CWL Version: All ▼

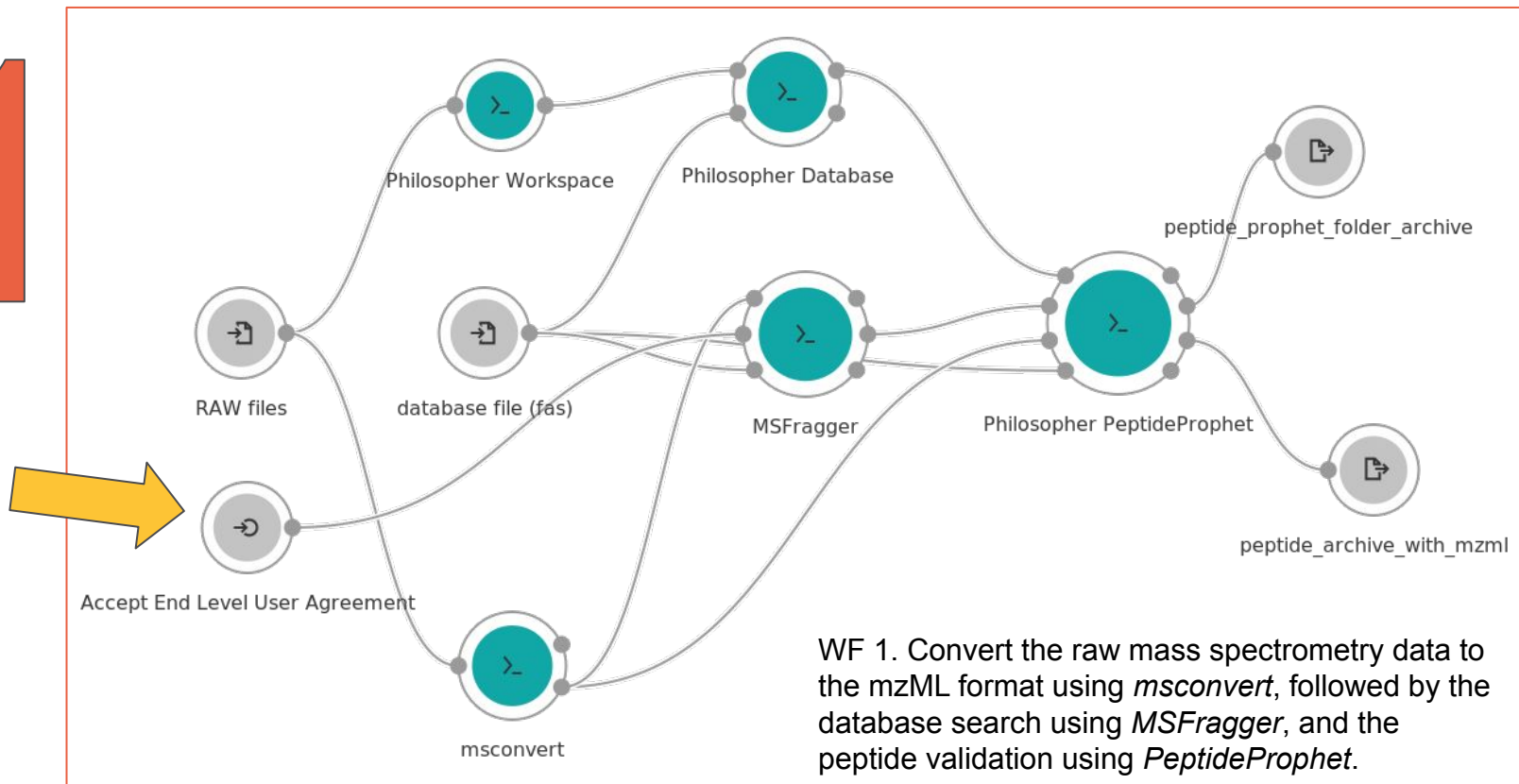
Status: Available ▼

▲ Name	Type
 01 FragPipe: Convert - Identify - Peptide Prophet The first step of the workflow consists of converting the raw mass spectrometry data to t...	Workflow
 02 FragPipe: Protein Prophet This workflow step takes the PeptideProphet output files from the first step containing the peptide validation and...	Workflow
 03 FragPipe: Filter - Quant - Report This workflow takes the PeptideProphet, and the ProteinProphet output files, and applies a stringent Fal...	Workflow
 04 FragPipe: TMT Integrator and QC This workflow step executes TMT-Integrator using the report tables generated by Philosopher. The pro...	Workflow

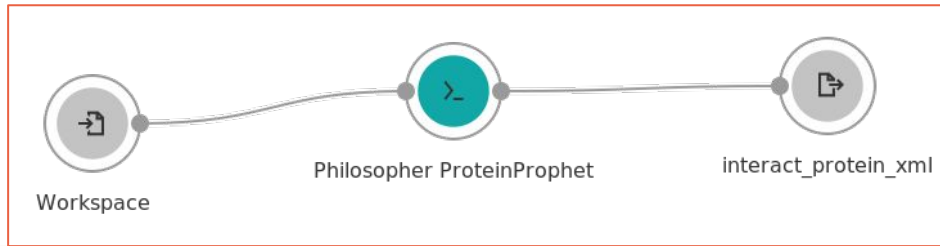


FragPipe - 4 Workflows

1

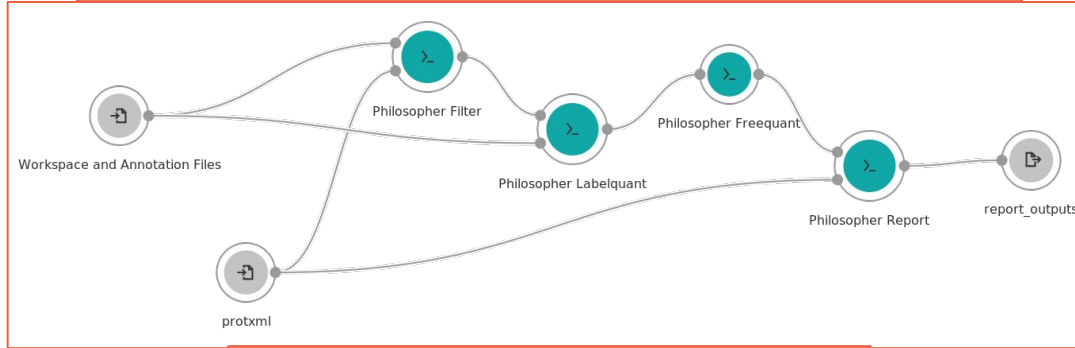


2



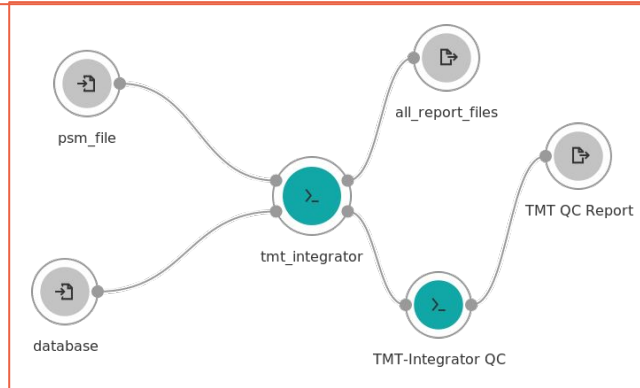
WF 2. Takes the PeptideProphet output files from the first step containing the peptide validation and calculates the protein inference using ProteinProphet.

3



WF 3. takes the PeptideProphet, and the ProteinProphet output files, and applies a stringent False Discovery Rate (FDR) filtering. Peptide and proteins are filtered individually at 1% FDR. The high-quality PSMs, peptides, and proteins are then quantified using a label-free algorithm that uses the apex peak intensity as a measurement. Finally, the isobaric tags are quantified and annotated with the correct sample labels.

4



WF 4. Executes TMT-Integrator using the report tables generated by Philosopher. The program applies a series of statistical filters, and high-quality thresholds to filter the data. Summary report tables are created containing peptides, proteins, genes, and phosphosites (only for phospho-enriched data sets).



Features to control cloud costs



Monitor analysis at each step in **macro** or micro levels

Dashboard Files Apps **Tasks** sailakss - FragPipe Proteomics Pipeline Tutorial ? Interactive Analysis Notes

COMPLETED **FragPipe: Convert - Identify - PeptideProphet run - 04-21-21 18:07:35: Plex o...** ? [Get support](#) [View stats & logs](#) [Edit and rerun](#)

Executed on Apr. 21, 2021 13:12 by sailakss

Spot Instances: **On** ? | Memoization (WorkReuse): **Off** ? **Price: \$0.31** ? Refund [View refunds](#) **Duration: 1 hour, 3 minutes** ?

App: FragPipe: Convert - Identify - PeptideProphet - Revision: 2

Inputs

- RAW files ? ?
 - CPTAC-3 22CPTAC_CCRCC_W_JHU_20180625_LUMOS_f...
 - CPTAC-3 22CPTAC_CCRCC_W_JHU_20180625_LUMOS_f...
 - CPTAC-3 22CPTAC_CCRCC_W_JHU_20180625_LUMOS_f...
 - CPTAC-3 22CPTAC_CCRCC_W_JHU_20180625_LUMOS_f...
 - CPTAC-3 22CPTAC_CCRCC_W_JHU_20180625_LUMOS_f......and 20 more items
- database file (fas) ? ?
 - 2020-03-30-decoys-reviewed-contam-UP000005640.fas

App Settings

Show non-default ?

- MSFragger (#msfragger)
 - Accept End Level User Agreement True

Outputs

- peptide_archive_with_mzml ?
 - 22CPTAC_CCRCC_Proteome_JHU_20180625.for_filter.tar.gz
- peptide_prophet_folder_archive ?
 - 22CPTAC_CCRCC_Proteome_JHU_20180625.for_protein.tar.gz

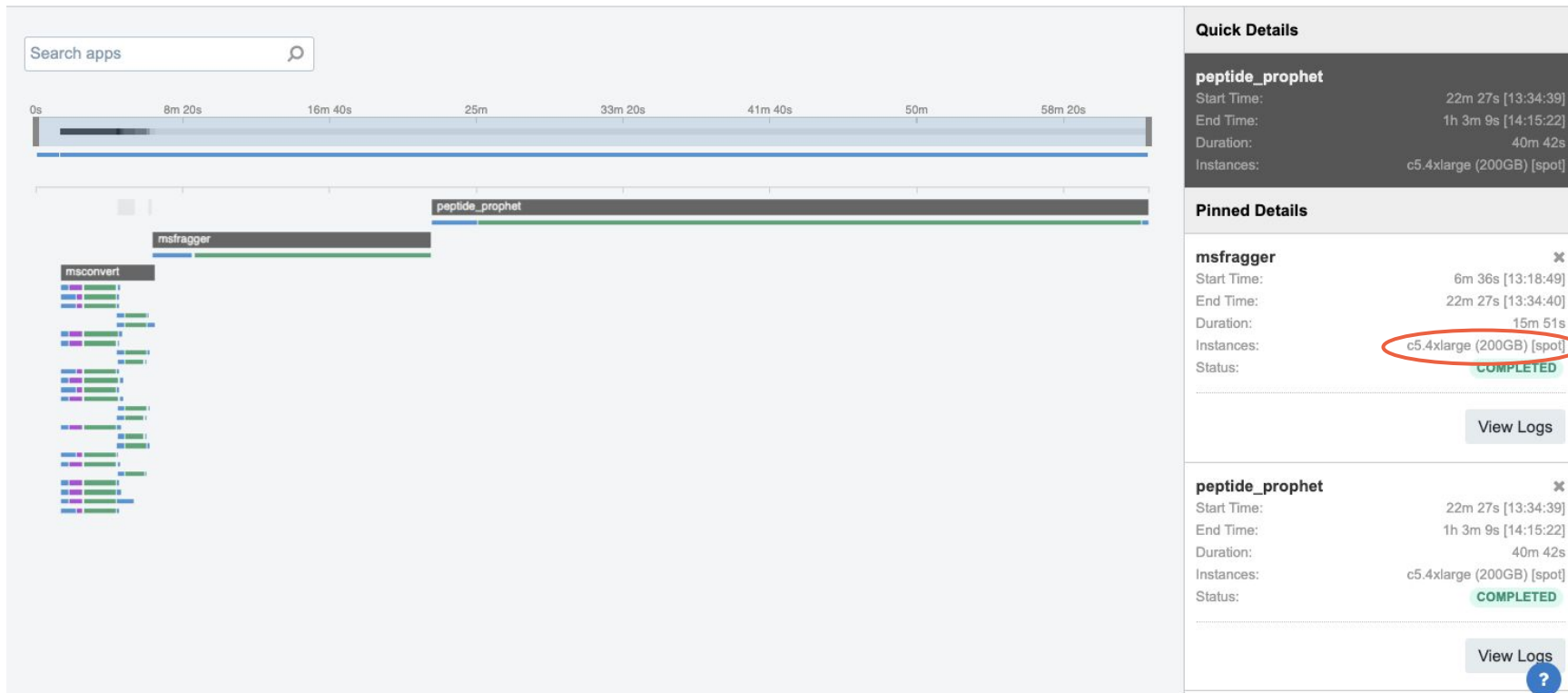


Monitor analysis at each step in macro or **micro** levels

COMPLETED Tasks / FragPipe: Convert - Identify - PeptideProphet run - 04-21-21 18:07:35: Plex or dataset name: 22CPTAC_CCRCC_Proteome_JHU_20180625 / Stats

Instance metrics

[View task logs](#)



Scale Up with **batching** by PDC file metadata

App: 01 FragPipe: Convert - Identify - Peptide Prophet - Revision: 2

Task Inputs Execution Settings

Inputs

Batching On

RAW files * Change selection

Batch by: File metadata

This task will be batched by file metadata (Plex or dataset name) and this will create 21 groups.

- ▶ 01cptac_ccrcc_proteome_jhu_20171007 (25 items) ✕
- ▶ 02cptac_ccrcc_proteome_jhu_20171003 (25 items) ✕
- ▶ 03cptac_ccrcc_proteome_jhu_20171022 (25 items) ✕
- ▶ 04cptac_ccrcc_proteome_jhu_20171026 (25 items) ✕
- ▶ 05cptac_ccrcc_proteome_jhu_20171030 (25 items) ✕
- ▶ 06cptac_ccrcc_proteome_jhu_20171120 (25 items) ✕
- ▶ 07cptac_ccrcc_proteome_jhu_20171127 (25 items) ✕
- ▶ 08cptac_ccrcc_proteome_jhu_20171205 (25 items) ✕
- ▶ 09cptac_ccrcc_proteome_jhu_20171215 (25 items) ✕
- ▶ 10cptac_ccrcc_proteome_jhu_20180119 (25 items) ✕
- ▶ 11cptac_ccrcc_proteome_jhu_20180126 (25 items) ✕

App Settings

Edit parameters Show editable ▼

- ▶ **Philosopher PeptideProphet** (#peptide_prophet)
 - ▼ **MSFragger** (#msfragger)
 - add_A_alanine
 - add_B_user_amino_acid
 - add_C_cysteine
 - add_Cterm_peptide
 - add_Cterm_protein
 - add_D_aspartic_acid

Outputs

- peptide_archive_with_mzml *No value*
- peptide_prophet_folder_archive *No value*



Review Outputs without downloading

Dashboard **Files** Apps Tasks **PDC Virtual Workshop**

Files

gene_report.html [Edit Metadata](#) [Copy](#)

1.9 MIB (1,970,530 bytes) · Produced on April 17, 2021 10:36 (Eastern Daylight Time), by 04 FragPipe: TMT Integrator and QC run - 04-17-21 01:22:03 · Modified on April 17, 2021 10:36 (1)

Metadata Raw View **Preview**

The preview area displays a dendrogram at the top, showing hierarchical clustering of samples. Below the dendrogram is a heatmap with a blue color scale, representing gene expression levels across the samples. The heatmap is divided into two main clusters by the dendrogram's branching structure.

Metrics of FragPipe Cloud vs. Local

Analysis of ccRCC whole cell lysate samples @Michigan on local server (non-cloud)

Computation:

- local server
- 56 cores (Xeon(R) CPU, 2.60GHz)
- 500GB RAM

Time for analysis:

27 hours, and a total of 16 GB RAM

Analysis of ccRCC samples on the CGC

Time for accessing the data:

Less than a minute; no need for downloads

Time for analysis on CGC:

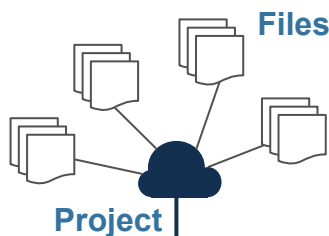
4.5 hours total. Up to 23 aws nodes in parallel @ 8 vCPU 16GB RAM, 48 aws nodes cumulative

Cost of analysis on CGC:

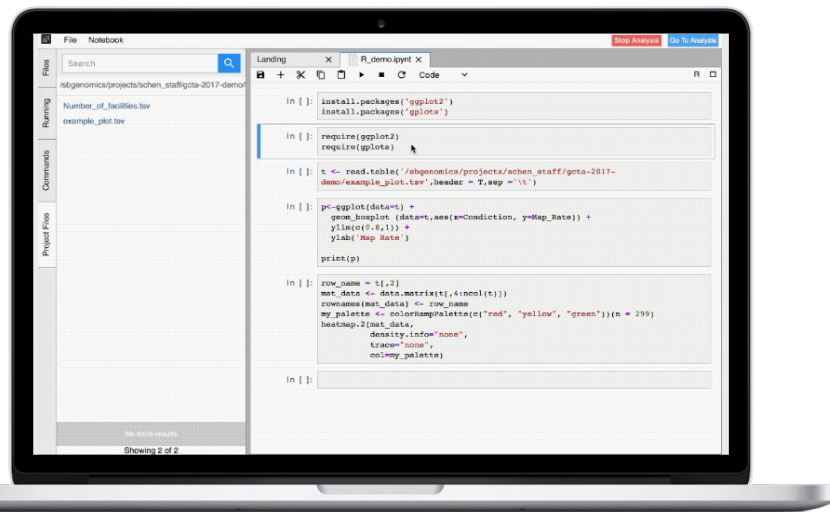
\$0.49 per TMT plex, ~\$12 for total analysis

Powerful, collaborative, & reproducible interactive analysis

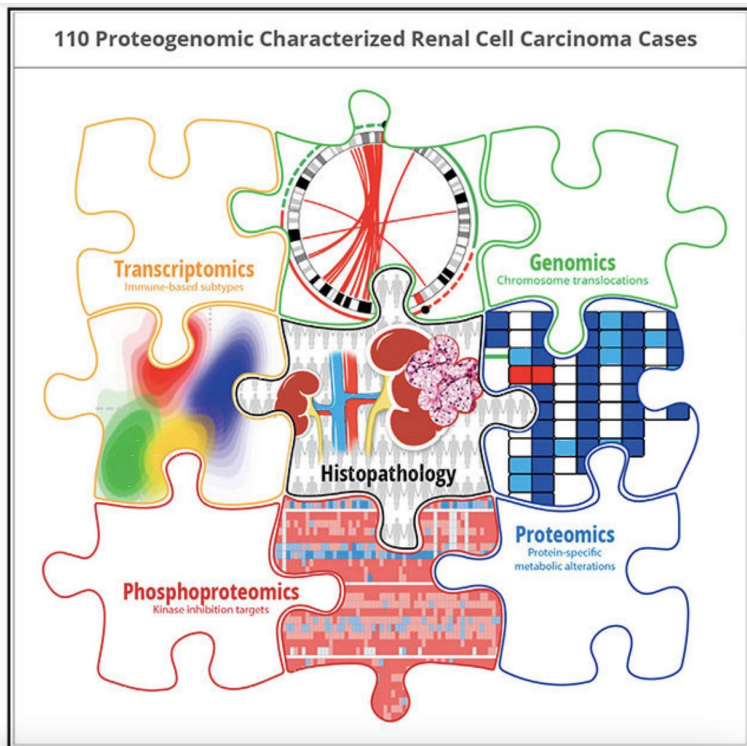
Users create interactive analysis sessions within a project - all files are available and over 50 instances can be used (*c3.xlarge* to *x1.32xlarge* on AWS)



Instance



Multi-omic data is critical for cancer research



Cancer is a complex disease!

Many research questions go beyond genomics data!

Different modalities should be examined to comprehensively understanding the full picture of a research question

As a researcher, it is essential to focus on the data. The CGC provides all tools for a multi omics analysis, so you can spend more time interpreting the results, and not configuring programs.

Clark et al., Integrated Proteogenomic Characterization of Clear Cell Renal Cell Carcinoma, Cell. 2019 Oct 31; 179(4): 964–983.e31. DOI:<https://doi.org/10.1016/j.cell.2019.10.007>





Ongoing Projects

Adding new resources and capabilities to CGC
to support broader areas of cancer research



Epigenetic mechanisms for transcription regulation in cancer

Overall aim:

- Enable secondary analysis of epigenomics data on CGC, focusing on ATACseq and ChIPseq experiments
- Develop multi-omics analysis for data coming from different omics experiments (RNAseq, ATACseq, ChIPseq, WGBS, RRBS, proteomics)
- Analyze data from several publicly available repositories in order to characterize epigenetic markers in cancer.

Deliverables (publicly available on CGC soon):

- Workflows for ATACseq and ChIPseq analysis, based on ENCODE's specification
- Data Cruncher Interactive Analysis combining ATACseq, ChIPseq, WGBS, RNAseq and proteomics data
- Workflow for multi-omics analysis as a one-click solution



Multi-omics and imaging machine learning analysis

Overall aim:

Create a predictive model from both genomics and image features combined with the available clinical data in order to **predict therapy response among patients with cancer**.

- Build machine learning tools for processing multi-omics and imaging data from dbGAP/TCGA dataset.
- Use existing deep learning algorithms and libraries and adapt them for execution on CGC platform.

Deliverables (publicly available on CGC soon):

- Data Cruncher Interactive Analysis
- Tools and workflows for deep learning models adapted for imaging data, utilizing GPU instances and Tensorflow Python deep learning library.

Support and Resources

CGC Monthly Webinar Series

- Learn about CGC platform features that you can use in your projects.
- 4th Wednesday of each month at 2pm ET
- Upcoming webinar info, slides and recordings are available at:

<https://www.cancergenomicscloud.org/webinars>

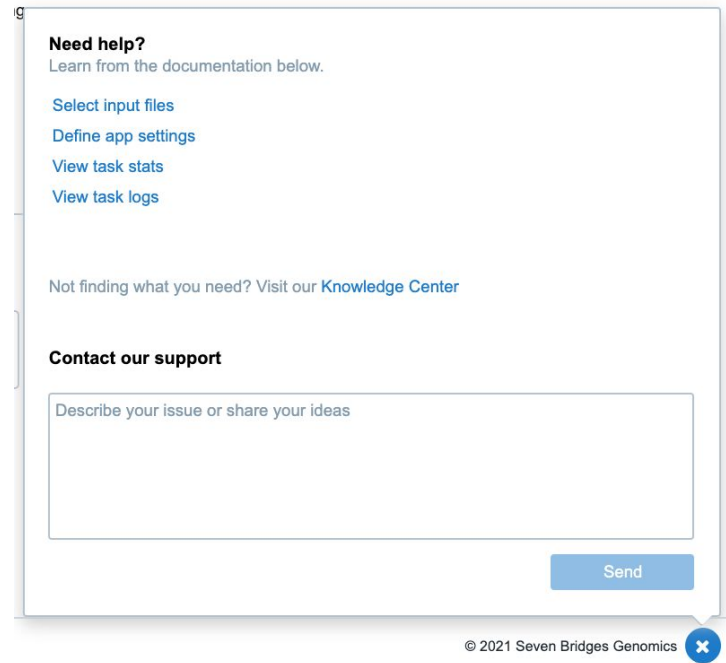
CGC Knowledge Center

<https://docs.cancergenomicscloud.org/>

Contact CGC Support: cgc@sevenbridges.com

Office Hours: Every week on Thursdays

<https://www.cancergenomicscloud.org/officehours>



Need help?
Learn from the documentation below.

- [Select input files](#)
- [Define app settings](#)
- [View task stats](#)
- [View task logs](#)

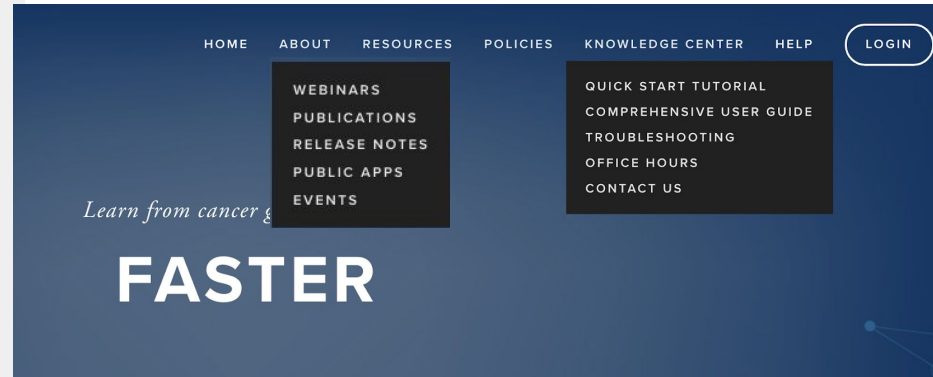
Not finding what you need? Visit our [Knowledge Center](#)

Contact our support

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OFFICE HOURS
CONTACT US

Learn from cancer g

FASTER

In Summary



Data Access



Immediately access petabytes of **Open and Controlled** TCGA, CPTAC, and other omics datasets

Bring your own private cohorts alongside public data.



Tools and Workflows



Standard bioinformatics pipelines

Bring your own analysis tools directly to the platform

Connect multiple tools together using our interactive custom workflow builder



Collaborate on the cloud



Collaborate with other researchers around the world in a secure workspace
Access to high-throughput, cost-effective cloud computing resources and storage on demand and at cost.



6000 users

>80 countries

1,600,000+ computational tasks

1400+ years of total compute time

66,800+ workflows

500+ public apps

R

Interactive Analysis



The ability to perform custom, interactive analysis and visualization on the platform using Python, RStudio.



Support & Resources



Access comprehensive online documentation and training resources; Technical support from a team of >200 expert scientists, bioinformaticians, and engineers.



Towards best practice in cancer mutation detection with whole-genome and whole-exome sequencing

May 26, 2021

2 pm EDT/11 am PDT/7 pm GMT



Dr. Wenming Xiao

Lead Bioinformatics Scientist

Office of New Drug, Office of Oncological Diseases, Center for
Drug Evaluation and Research,
Food and Drug Administration

Acknowledgements

CGC Team

Manisha Ray

Jelena Radenkovic

Milos Stanojevic

Milos Trboljevac

Marko Tasic

Ana Stelkic

Dave Roberson

Sai Lakshmi Subramanian

Jack DiGiovanna

Brandi Davis-Dusenbery

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Guo Ci Teo

Fengchao Yu



PDC Team

Paul Rudnick

Rajesh Thangudu



NATIONAL
CANCER
INSTITUTE

The Seven Bridges Cancer Research Data Commons Cloud Resource has been funded in whole or in part with Federal funds from the National Cancer Institute, National Institutes of Health, Contract No. HHSN261201400008C and ID/IQ Agreement No. 17X146 under Contract No. HHSN261201500003I and 75N91019D00024.

Questions?

