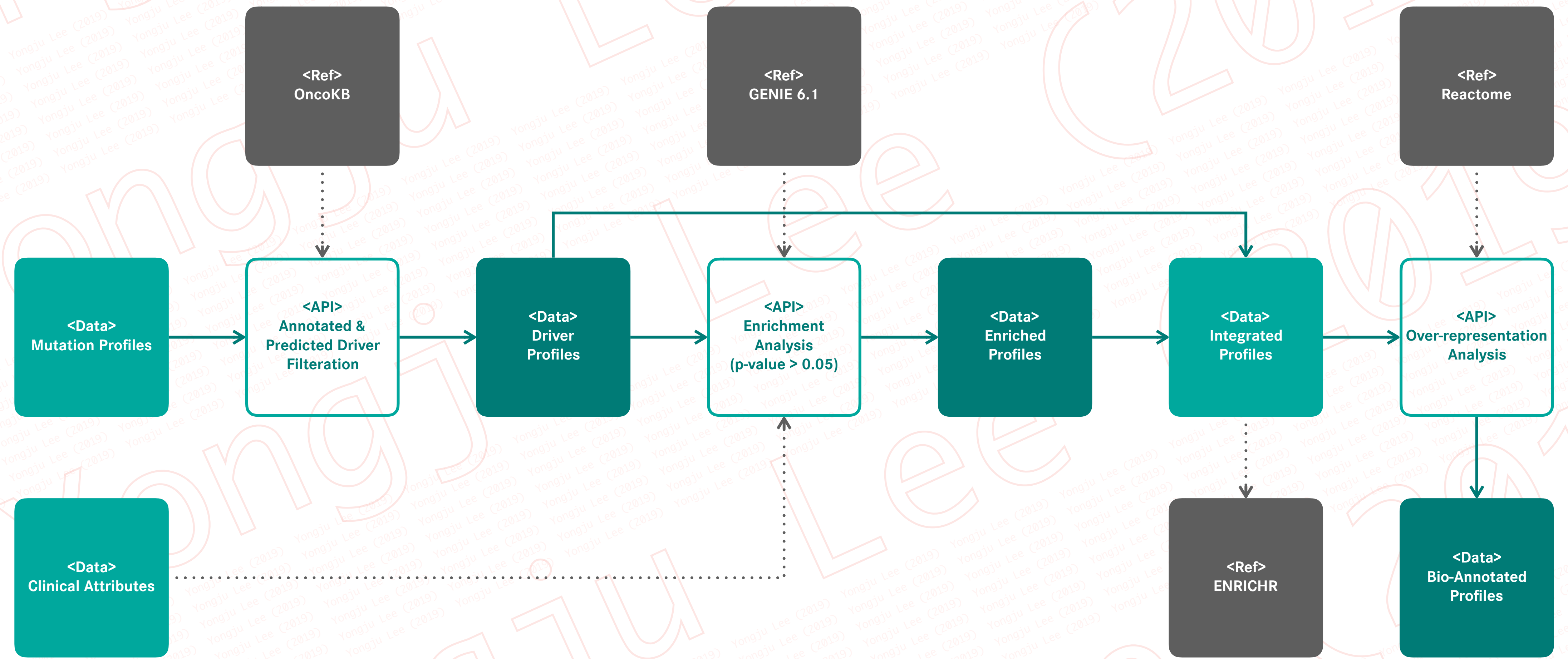
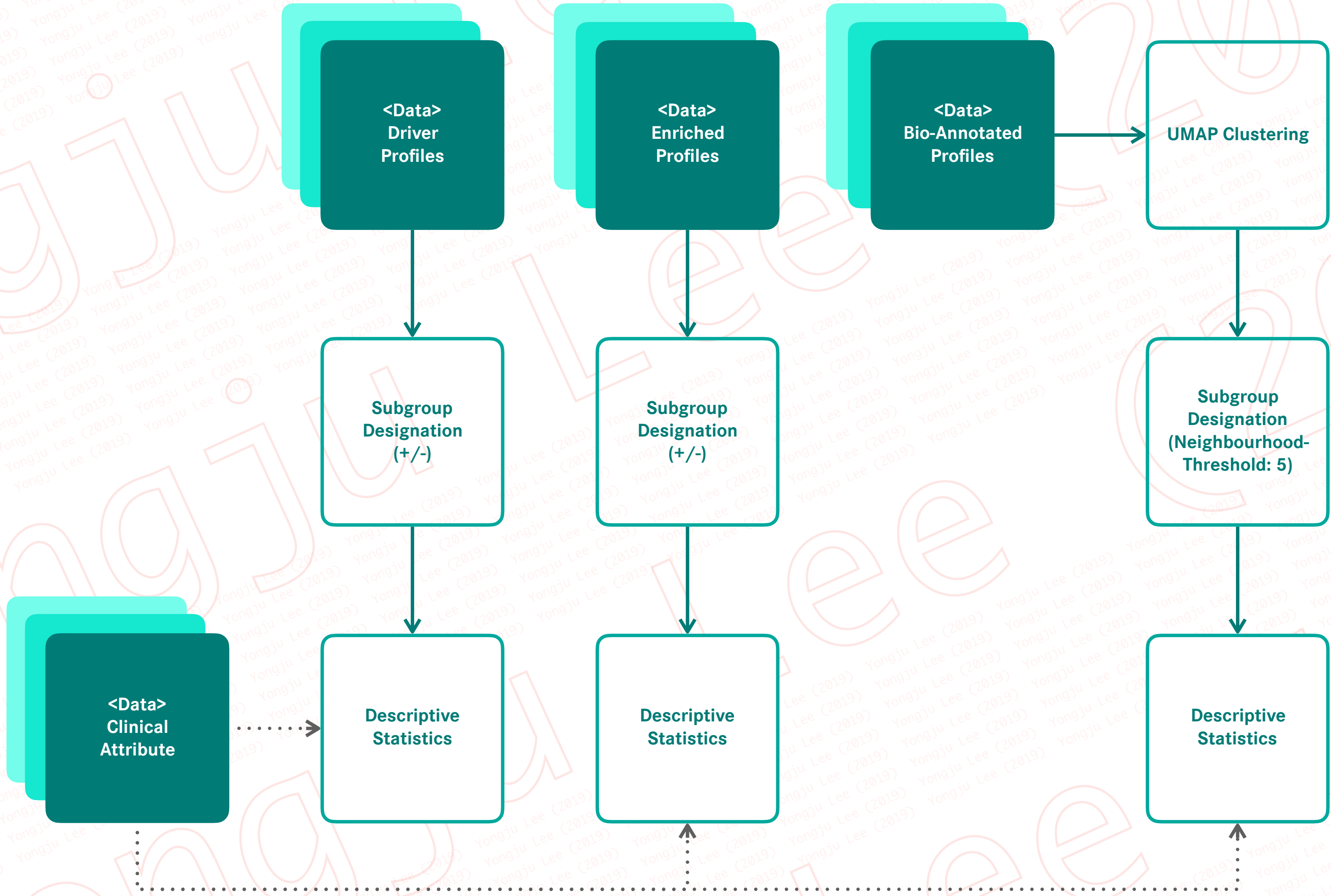


Patient-level

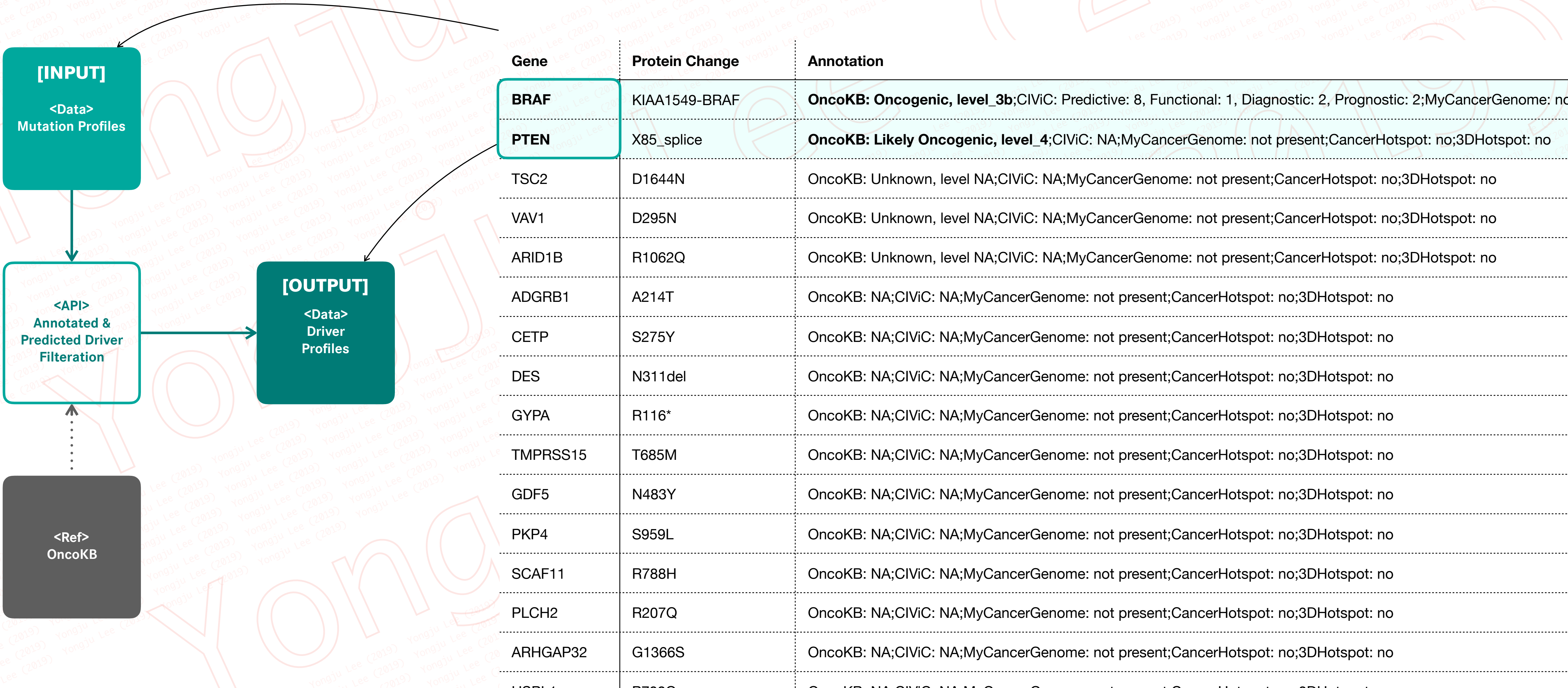


Study-level



1. 환자별 Mutation 리스트 & OncoKB Annotation

- [INPUT] 환자별 Mutation을 Gene과 해당 Gene의 Protein Change를 받아
- [FILTER] OncoKB 최신 Annotation dataset을 맵핑하여
- [OUTPUT] 1) Oncogenic이거나 2) Likely Oncogenic인 “Driver Gene”들만 추림



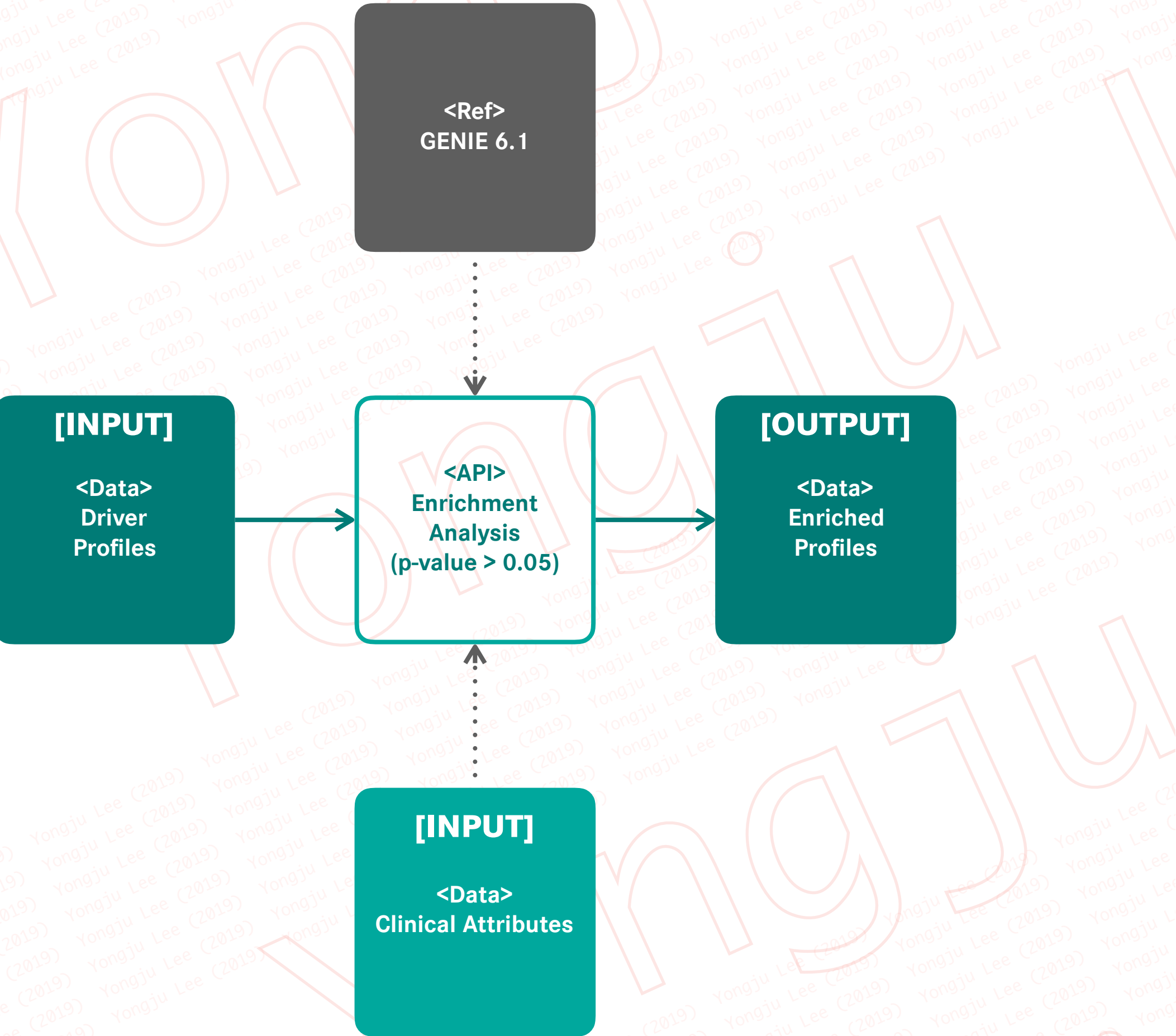
2. Enrichment Analysis

[INPUT] 1) 추려진 “Driver Gene”들과, 2) 환자의 a. Oncotree(i.e., BRCA, ICD, ILC, PRAD, etc.) 코드와, b. Sample type(i.e., Primary or Metastasis)를

[FILTER] CBioPortal Genie에서 동일한 a. Oncotree code와 b. Sample type 조건을 준 뒤,

“Driver Gene”당 “: MUT_DRIVER;” 스트링을 붙여 (OQL; Oncology Query Language) Enrichment 분석 (이때, Genomic profile은 Somatic mutations만)

[OUTPUT] 1) Patient-level Enrichment, 2) Enrichment group 은 Mutual Exclusivity로 지정한 뒤, 3) p-value < 0.05인 Gene symbol을 가져옴



[INPUT]

| | |
|------|-----------|
| BRAF | : DRIVER; |
| PTEN | : DRIVER; |

Selected Studies:

Modify

PRAD_Metastasis (1071 total samples)

Select Genomic Profiles:

☒ Somatic mutations ?

☐ Copy-number alterations ?

Select Patient/Case Set:

To build your own case set, try out our enhanced Study View.

All (1071)

×

▼

Enter Genes:

User-defined List

×

▼

BRAF : DRIVER;

PTEN : DRIVER;

Hint: Learn Onco Query Language (OQL) to write more powerful queries

All gene symbols are valid.

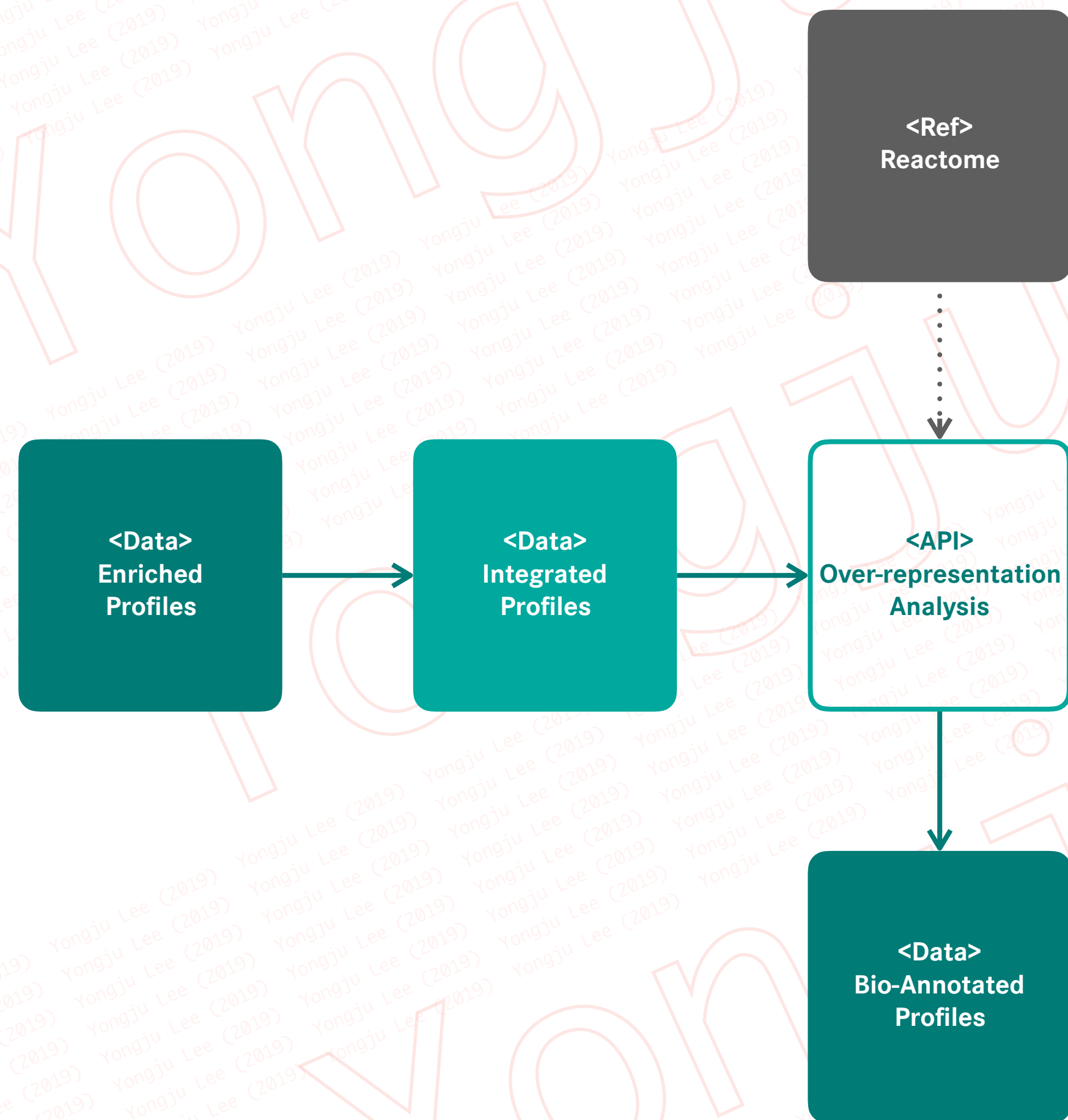
3. 환자별 Mutation 리스트 & OncoKB Annotation

[INPUT] 1) Mutated Gene 프로파일과, 2) Enriched Gene 프로파일 병합된 Integrated 프로파일(Gene symbol만)을

[FILTER] Reactome GSEA(Gene set enrichment analysis)로 분석 (이때, parameter 설정을 a. Project to human & b. Include interactors를 TRUE로 분석)


랭킹된 Pathway list에서 “Entities pValue”와 “Entities FDR”의 값을 $-\log_{10}$ 값으로 치환하여

[OUTPUT] 환자별 패스웨이 랭킹을 가져옴 (Bio-annotated ranked profile)




Step 2: Select your preferred options.

✓ Project to human

 All non-human identifiers are converted to their human equivalents (expand for more info...)

☒ Include interactors

 IntAct interactors are used to increase the analysis background (expand for more info...)

[illegible]

4. UMAP-based 클러스터링 Preparation

[INPUT] Bio-annotated ranked profile에서 a. -LOG10(Entities pValue) 버전과, b. -LOG10(Entities FDR) 버전을 나눠

[OUTPUT] 1) vector와 2) label 리스트 tsv 생성

| HSA_NAME | HSA_CODE | STDEV | AVG | CNT | 9115586 | 9115572 | 9115539 |
|--|---------------|-----------|-----------|-----|-------------------|-------------------|-------------------|
| Defective Mismatch Repair Associated With MLH1 | R-HSA-5545483 | ⚠ | 1.4055506 | 1 | 1.405550654493160 | | |
| Resolution of D-loop Structures through Synthesis-Dependent S | R-HSA-5693554 | 0.2039106 | 0.8533256 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| Resolution of D-loop Structures through Holliday Junction Interr | R-HSA-5693568 | 0.2020603 | 0.8524245 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| Resolution of D-Loop Structures | R-HSA-5693537 | 0.2014205 | 0.8518870 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| HDR through Single Strand Annealing (SSA) | R-HSA-5685938 | 0.3092483 | 0.8995658 | 19 | 1.229146916723810 | 0.672630280305592 | 0.879823343399318 |
| Processing of DNA double-strand break ends | R-HSA-5693607 | 0.2144646 | 0.7749681 | 17 | 1.229146916723810 | 0.630004274129010 | 0.816978853975654 |
| HDR through Homologous Recombination (HRR) | R-HSA-5685942 | 0.1993074 | 0.7821921 | 17 | 1.229146916723810 | 0.590856054495176 | 0.775590112742391 |
| Meiotic recombination | R-HSA-912446 | 0.2184479 | 0.7518827 | 17 | 1.229146916723810 | 0.585624564469399 | 0.770039850544110 |
| Transcriptional Regulation by E2F6 | R-HSA-8953750 | 0.2871504 | 0.7176407 | 15 | 1.229146916723810 | 0.515963152367605 | 0.695636053797126 |
| SUMO E3 ligases SUMOylate target proteins | R-HSA-3108232 | 0.4393138 | 0.4551416 | 23 | 1.229146916723810 | 0.264650651162941 | 0.516064715395482 |
| SUMOylation | R-HSA-2990846 | 0.4390480 | 0.4475852 | 23 | 1.229146916723810 | 0.254496116843433 | 0.502298482580464 |
| Metabolism of proteins | R-HSA-392499 | 0.4951786 | 0.2633776 | 24 | 1.229146916723810 | 0.066480159992483 | 0.044547689690695 |
| TP53 Regulates Transcription of DNA Repair Genes | R-HSA-6796648 | 2.0919406 | 3.6150411 | 17 | 1.229146916723810 | | 4.29997324972002E |
| G2/M DNA damage checkpoint | R-HSA-69473 | 0.4657978 | 1.1867209 | 16 | 1.229146916723810 | | 0.879823343399318 |
| G2/M Checkpoints | R-HSA-69481 | 0.3154110 | 1.1020147 | 15 | 1.229146916723810 | | 0.879823343399318 |
| Regulation of TP53 Activity through Phosphorylation | R-HSA-6804756 | 0.3010464 | 1.0730907 | 18 | 1.229146916723810 | | 0.879823343399318 |
| Homologous DNA Pairing and Strand Exchange | R-HSA-5693579 | 0.1820286 | 0.8502934 | 14 | 1.229146916723810 | | 0.879823343399318 |
| Presynaptic phase of homologous DNA pairing and strand excha | R-HSA-5693616 | 0.1821164 | 0.8442072 | 14 | 1.229146916723810 | | 0.879823343399318 |
| Meiotic synapsis | R-HSA-1221632 | 0.2493609 | 0.6870204 | 15 | 1.229146916723810 | | 0.700141795961886 |

| | | | | | | | |
|----|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
| 1 | 1.405550654 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1. |
| 2 | 0 | 0.754337392 | 0.754337392 | 0.754337392 | 0.67263028 | 0.630004274 | 0.59085605 |
| 3 | 0 | 0.879823343 | 0.879823343 | 0.879823343 | 0.879823343 | 0.816978854 | 0.77559011 |
| 4 | 0 | 0 | 0 | 0 | 0 | 0.084607889 | 0.079360377 |
| 5 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 6 | 0 | 0 | 0 | 1.403568034 | 0 | 0 | 1.414996791 |
| 7 | 0 | 0 | 0 | 0.884457004 | 0 | 0 | 0.646277146 |
| 8 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.68353998 |
| 9 | 0 | 0 | 0 | 0 | 0 | 1.025544325 | 1.366544469 |
| 10 | 0 | 0 | 0 | 0 | 0 | 0.891488505 | 0.891488505 |
| 11 | 0 | 0.904915122 | 0.904915122 | 0.904915122 | 0.894130778 | 0.84905199 | 0.90491512 |
| 12 | 0 | 0.808416197 | 0.808416197 | 0.808416197 | 0.808416197 | 0.668179673 | 0.62847941 |
| 13 | 0 | 0.892110401 | 0.892110401 | 0.887955178 | 0.812566628 | 0.711170702 | 0.81256662 |
| 14 | 0 | 1.252824627 | 1.237505999 | 1.232523412 | 1.81101513 | 1.075948279 | 1.02675944 |
| 15 | 0 | 0 | 0 | 0 | 0 | 0.750537141 | 0.252258512 |
| 16 | 0 | 0.649690695 | 0.649690695 | 0.649690695 | 0.649690695 | 0.595762688 | 0.64969069 |
| 17 | 0 | 0.600928448 | 0.600928448 | 0.600928448 | 0.600928448 | 0.579904013 | 0.54157377 |
| 18 | 0 | 0.836561138 | 0.836561138 | 0.836561138 | 0.832002844 | 0.787477293 | 0.74637333 |
| 19 | 0 | 0.957861341 | 0.957861341 | 0.957861341 | 0.957861341 | 0.922837747 | 0.88058653 |
| 20 | 0 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.64902855 |
| 21 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.68353998 |
| 22 | 0 | 0.679030141 | 0.679030141 | 0.679030141 | 0.679030141 | 0.612437282 | 0.67903014 |
| 23 | 0 | 0.816846642 | 0.816846642 | 0.816846642 | 0.73243996 | 0.689006864 | 0.64902855 |
| 24 | 0 | 0 | 0 | 0 | 0 | 0.101121417 | 0.00E+00 |
| 25 | 0 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.16656087 |

환자리스트

Pivot된 벡터

-LOG10(FDR)값

| |
|----------|
| 9115586 |
| 9115572 |
| 9115539 |
| 9115522 |
| 9115517 |
| 9115425 |
| 9115422 |
| 9115339 |
| 97115125 |
| 9115109 |
| 9115059 |
| 9115051 |
| 6115120 |
| 6115114 |
| 6115247 |
| 6115395 |
| 96115135 |
| 5115650 |
| 5115615 |
| 5115610 |
| 5115609 |
| 5115359 |
| 5115075 |
| 5115031 |
| 1115019 |

벡터 순으로 나열된 환자 id

4. UMAP-based 클러스터링

[INPUT] Vector와 Label 리스트 tsv를,

[PROCESS] Neighborhood parameter를 최솟값이 5로 지정한뒤, UMAP clustering

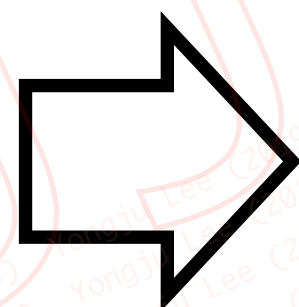
[OUTPUT] Cluster group별 리스트 가져오기

| | | | | | | | | |
|----|-------------|---------------|-------------|---------------|-------------|---------------|-------------|-----------|
| 1 | 1.405550654 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1 |
| 2 | 0 | 0.754337392 | 0.754337392 | 0.754337392 | 0.67263028 | 0.630004274 | 0.5908560 | 0 |
| 3 | 0 | 0.879823343 | 0.879823343 | 0.879823343 | 0.879823343 | 0.816978854 | 0.7755904 | 0 |
| 4 | 0 | 0 0 0 0 0 0 0 | 0 | 0.084607889 | 0.079360377 | 7.566E-05 | 5 | 1 |
| 5 | 0 | 0 0 0 0 0 0 0 | 0 | 0 0 0 0 0 0 0 | 0 | 0 0 0 0 0 0 0 | 0 | 0 |
| 6 | 0 | 0 0 0 0 0 0 0 | 0 | 0 | 0 | 1.414996791 | 1.414996791 | 1.414 |
| 7 | 0 | 0 0 0 0 0 0 0 | 0 | 0.884457004 | 0 | 0 | 0.646277146 | 0.3514808 |
| 8 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.6835399 | 0 |
| 9 | 0 | 0 0 0 0 0 0 0 | 1.025544325 | 1.366544469 | 1.025544325 | 1.0255443 | 1.0255443 | 0 |
| 10 | 0 | 0 0 0 0 0 0 0 | 0 | 0 | 0.891488505 | 0.891488505 | 0.597070818 | 0 |
| 11 | 0 | 0.904915122 | 0.904915122 | 0.904915122 | 0.894130778 | 0.84995199 | 0.9049151 | 0 |
| 12 | 0 | 0.808416197 | 0.808416197 | 0.808416197 | 0.808416197 | 0.668179673 | 0.6284794 | 0 |
| 13 | 0 | 0.892110401 | 0.892110401 | 0.887955178 | 0.812566628 | 0.71110702 | 0.8125666 | 0 |
| 14 | 0 | 1.252824627 | 1.237505999 | 1.232523412 | 1.81101513 | 1.075948279 | 1.0267594 | 0 |
| 15 | 0 | 0 0 0 0 0 0 0 | 0 | 0.750537141 | 0 | 0.25900116 | 0.252258512 | 0.026 |
| 16 | 0 | 0.649690695 | 0.649690695 | 0.649690695 | 0.649690695 | 0.595762688 | 0.6496906 | 0 |
| 17 | 0 | 0.600928448 | 0.600928448 | 0.600928448 | 0.600928448 | 0.579904013 | 0.5415737 | 0 |
| 18 | 0 | 0.836561138 | 0.836561138 | 0.836561138 | 0.832002844 | 0.78477293 | 0.7463733 | 0 |
| 19 | 0 | 0.957861341 | 0.957861341 | 0.957861341 | 0.957861341 | 0.922837747 | 0.8805865 | 0 |
| 20 | 0 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.6490285 | 0 |
| 21 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.6835399 | 0 |
| 22 | 0 | 0.679030141 | 0.679030141 | 0.679030141 | 0.679030141 | 0.612437282 | 0.6790301 | 0 |
| 23 | 0 | 0.816846642 | 0.816846642 | 0.816846642 | 0.73243996 | 0.689006864 | 0.6490285 | 0 |
| 24 | 0 | 0 0 0 0 0 0 0 | 0 | 0 0 0 0 0 0 0 | 0 | 0.101121417 | 0.00E+00 | 0 |
| 25 | 0 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.1665608 | 0 |

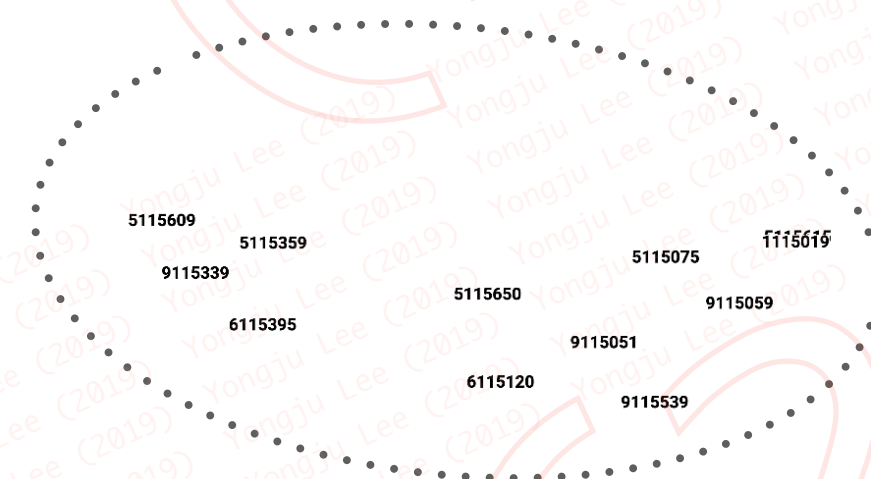
Pivot된 벡터

9115586
9115572
9115539
9115522
9115517
9115425
9115422
9115339
97115125
9115109
9115059
9115051
6115120
6115114
6115247
6115395
96115135
5115650
5115615
5115610
5115609
5115359
5115075
5115031
1115019

벡터 순으로 나열된 환자 id



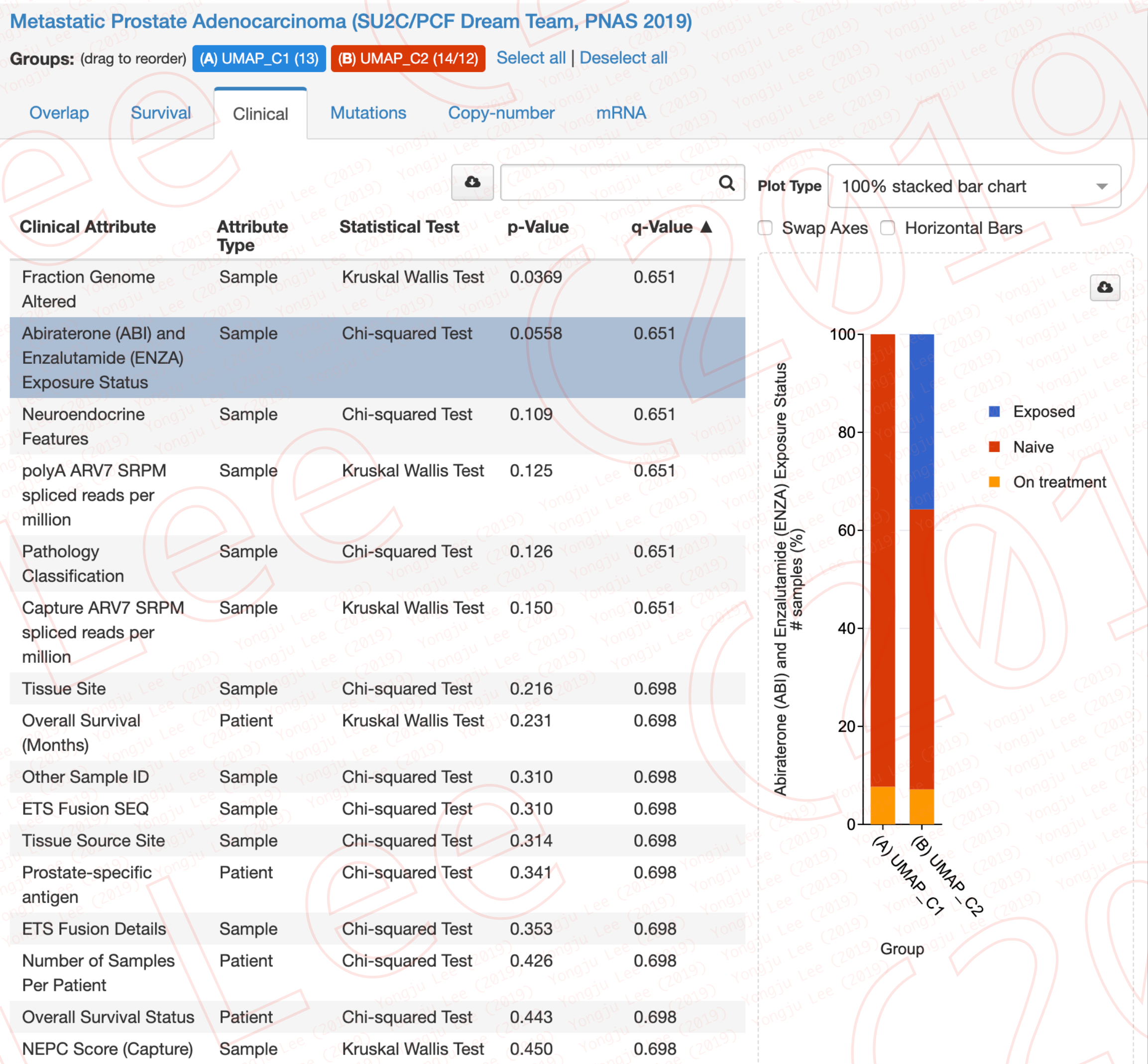
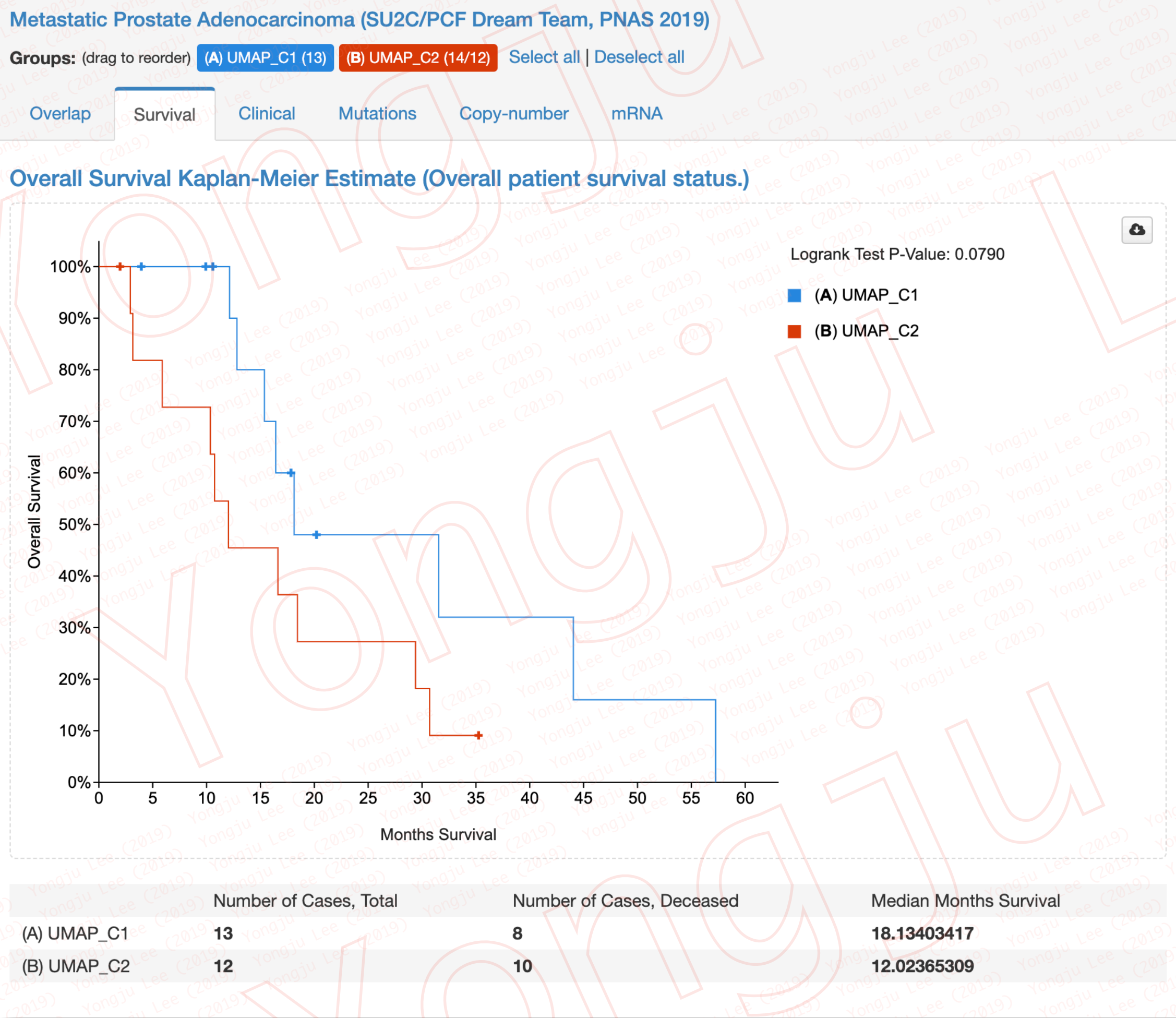
Cluster 1



Cluster 2

5. 클러스터 기반 Group Analysis

[INPUT] 클러스터된 환자들을 기준으로 그룹을 생성하여,
[OUTPUT] CBioPortal에서 제공되는 모든 탭의 Group Analysis결과 가져오기



SUMMIT TRIAL (2018)

Driver Only

With Enriched Entities (CO*s & ME*s)

With Enriched Entities (ME* only)

without
Over-representation*

| DRIVER_TP53+/- | | | | | DRIVER_PIK3CA+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value | Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Sample | Kruskal Wallis Test | 4.6E-03 | 0.0462 | Mutation Count | Sample | Kruskal Wallis Test | 0.0354 | 0.239 |
| Mutation Burden | Sample | Kruskal Wallis Test | 4.62E-03 | 0.0462 | Mutation Burden | Sample | Kruskal Wallis Test | 0.0357 | 0.239 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0143 | 0.0954 | Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0359 | 0.239 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0650 | 0.288 | Waterfall Group | Sample | Chi-squared Test | 0.0920 | 0.460 |
| Waterfall Group | Sample | Chi-squared Test | 0.0821 | 0.288 | Best objective response rate | Patient | Chi-squared Test | 0.189 | 0.696 |
| Coverage | Sample | Kruskal Wallis Test | 0.0864 | 0.288 | Purity | Sample | Kruskal Wallis Test | 0.234 | 0.696 |
| Bubble Group | Sample | Chi-squared Test | 0.159 | 0.453 | Bubble Group | Sample | Chi-squared Test | 0.244 | 0.696 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.252 | 0.554 | Qualifying Mutation | Sample | Chi-squared Test | 0.363 | 0.744 |
| Central | Sample | Chi-squared Test | 0.256 | 0.554 | Central | Sample | Chi-squared Test | 0.371 | 0.744 |
| Local Test | Sample | Chi-squared Test | 0.277 | 0.554 | Local Test | Sample | Chi-squared Test | 0.372 | 0.744 |
| Best objective response rate | Patient | Chi-squared Test | 0.392 | 0.712 | Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.514 | 0.801 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.732 | MSI Status | Sample | Chi-squared Test | 0.585 | 0.801 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.732 | Oncoprint | Sample | Chi-squared Test | 0.585 | 0.801 |
| Data | Sample | Chi-squared Test | 0.535 | 0.732 | Criteria | Sample | Chi-squared Test | 0.653 | 0.801 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.732 | Clonality | Sample | Chi-squared Test | 0.661 | 0.801 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.840 | IMPACT | Sample | Chi-squared Test | 0.661 | 0.801 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.840 | Coverage | Sample | Kruskal Wallis Test | 0.695 | 0.801 |
| Amplification Status | Sample | Chi-squared Test | 0.788 | 0.846 | Data | Sample | Chi-squared Test | 0.721 | 0.801 |
| Purity | Sample | Kruskal Wallis Test | 0.808 | 0.846 | ER_PR | Sample | Chi-squared Test | 0.836 | 0.880 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.846 | Amplification Status | Sample | Chi-squared Test | 0.954 | 0.954 |

N/A

with
Over-representation*

| 2 GROUP CLUSTERING (DRIVER) | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 1.46E-03 | 0.0293 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 9.20E-03 | 0.0642 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0127 | 0.0642 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0128 | 0.0642 |
| Coverage | Sample | Kruskal Wallis Test | 0.0451 | 0.181 |
| Waterfall Group | Sample | Chi-squared Test | 0.0567 | 0.189 |
| Bubble Group | Sample | Chi-squared Test | 0.125 | 0.338 |
| Best objective response rate | Patient | Chi-squared Test | 0.135 | 0.338 |
| Central | Sample | Chi-squared Test | 0.162 | 0.361 |
| Data | Sample | Chi-squared Test | 0.191 | 0.372 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.205 | 0.372 |
| IMPACT | Sample | Chi-squared Test | 0.356 | 0.594 |
| Local Test | Sample | Chi-squared Test | 0.509 | 0.783 |
| ER_PR | Sample | Chi-squared Test | 0.576 | 0.823 |
| Amplification Status | Sample | Chi-squared Test | 0.642 | 0.856 |
| Clonality | Sample | Chi-squared Test | 0.693 | 0.866 |
| Purity | Sample | Kruskal Wallis Test | 0.808 | 0.876 |
| MSI Status | Sample | Chi-squared Test | 0.833 | 0.876 |
| Oncoprint | Sample | Chi-squared Test | 0.833 | 0.876 |
| Criteria | Sample | Chi-squared Test | 0.933 | 0.933 |

| 2 GROUP CLUSTERING (ENR) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0191 | 0.138 |
| Central | Sample | Chi-squared Test | 0.0220 | 0.138 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0227 | 0.138 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0365 | 0.138 |
| Best objective response rate | Patient | Chi-squared Test | 0.0384 | 0.138 |
| Purity | Sample | Kruskal Wallis Test | 0.0415 | 0.138 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0662 | 0.189 |
| Local Test | Sample | Chi-squared Test | 0.174 | 0.399 |
| Amplification Status | Sample | Chi-squared Test | 0.179 | 0.399 |
| Data | Sample | Chi-squared Test | 0.253 | 0.496 |
| Waterfall Group | Sample | Chi-squared Test | 0.273 | 0.496 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.744 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.744 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.744 |
| Bubble Group | Sample | Chi-squared Test | 0.558 | 0.744 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.677 | 0.794 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.794 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.794 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.891 |
| Coverage | Sample | Kruskal Wallis Test | 0.903 | 0.903 |

| 3 GROUP CLUSTERING (ENR) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Central | Sample | Chi-squared Test | 0.0389 | 0.284 |
| Purity | Sample | Kruskal Wallis Test | 0.0498 | 0.284 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0524 | 0.284 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0639 | 0.284 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0745 | 0.284 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0933 | 0.284 |
| Amplification Status | Sample | Chi-squared Test | 0.0995 | 0.284 |
| Data | Sample | Chi-squared Test | 0.136 | 0.307 |
| Best objective response rate | Patient | Chi-squared Test | 0.143 | 0.307 |
| Clonality | Sample | Chi-squared Test | 0.153 | 0.307 |
| MSI Status | Sample | Chi-squared Test | 0.229 | 0.382 |
| Oncoprint | Sample | Chi-squared Test | 0.229 | 0.382 |
| Waterfall Group | Sample | Chi-squared Test | 0.249 | 0.382 |
| Local Test | Sample | Chi-squared Test | 0.267 | 0.382 |
| IMPACT | Sample | Chi-squared Test | 0.318 | 0.424 |
| Bubble Group | Sample | Chi-squared Test | 0.501 | 0.599 |
| ER_PR | Sample | Chi-squared Test | 0.509 | 0.599 |
| Coverage | Sample | Kruskal Wallis Test | 0.630 | 0.701 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.676 | 0.712 |
| Criteria | Sample | Chi-squared Test | 0.825 | 0.825 |

| 2 GROUP CLUSTERING (ME) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Patient | Kruskal Wallis Test | 0.0191 | 0.138 |
| Central | Sample | Chi-squared Test | 0.0220 | 0.138 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0227 | 0.138 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0365 | 0.138 |
| Best objective response rate | Patient | Chi-squared Test | 0.0384 | 0.138 |
| Purity | Sample | Kruskal Wallis Test | 0.0415 | 0.138 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0662 | 0.189 |
| Local Test | Sample | Chi-squared Test | 0.174 | 0.399 |
| Amplification Status | Sample | Chi-squared Test | 0.179 | 0.399 |
| Data | Sample | Chi-squared Test | 0.253 | 0.496 |
| Waterfall Group | Sample | Chi-squared Test | 0.273 | 0.496 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.744 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.744 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.744 |
| Bubble Group | Sample | Chi-squared Test | 0.558 | 0.744 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.677 | 0.794 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.794 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.794 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.891 |
| Coverage | Sample | Kruskal Wallis Test | 0.903 | 0.903 |

| 3 GROUP CLUSTERING (ME) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 6.3E-03 | 0.127 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0242 | 0.133 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0260 | 0.133 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0266 | 0.133 |
| Best objective response rate | Patient | Chi-squared Test | 0.0530 | 0.171 |
| Purity | Sample | Kruskal Wallis Test | 0.0599 | 0.171 |
| Coverage | Sample | Kruskal Wallis Test | 0.0603 | 0.171 |
| Central | Sample | Chi-squared Test | 0.0683 | 0.171 |
| Amplification Status | Sample | Chi-squared Test | 0.0995 | 0.221 |
| Waterfall Group | Sample | Chi-squared Test | 0.211 | 0.408 |
| ER_PR | Sample | Chi-squared Test | 0.241 | 0.408 |
| IMPACT | Sample | Chi-squared Test | 0.245 | 0.408 |
| Bubble Group | Sample | Chi-squared Test | 0.277 | 0.426 |
| Data | Sample | Chi-squared Test | 0.342 | 0.488 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.435 | 0.527 |
| MSI Status | Sample | Chi-squared Test | 0.448 | 0.527 |
| Oncoprint | Sample | Chi-squared Test | 0.448 | 0.527 |
| Local Test | Sample | Chi-squared Test | 0.494 | 0.548 |
| Criteria | Sample | Chi-squared Test | 0.825 | 0.869 |
| Clonality | Sample | Chi-squared Test | 0.949 | 0.949 |

- CO : Co-occurrent mutations
- ME : Mutually exclusive mutations
- Over-representation : Reactome based enrichment analysis

CCRCC (2019)

| DV_VHL+/- | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Sex | Patient | Chi-squared Test | 0.0350 | 0.484 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0561 | 0.484 |
| First Line Therapy | Patient | Chi-squared Test | 0.0781 | 0.484 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0815 | 0.484 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.0956 | 0.484 |
| Treatment Group | Patient | Chi-squared Test | 0.112 | 0.484 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.133 | 0.484 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.139 | 0.484 |
| Purity | Sample | Kruskal Wallis Test | 0.152 | 0.484 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.170 | 0.484 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.184 | 0.484 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.193 | 0.484 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.250 | 0.571 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.277 | 0.571 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.285 | 0.571 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.370 | 0.668 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.378 | 0.668 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.433 | 0.722 |
| Genome Doublings | Sample | Chi-squared Test | 0.498 | 0.784 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.571 | 0.784 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.576 | 0.784 |
| Age | Patient | Kruskal Wallis Test | 0.601 | 0.784 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.601 | 0.784 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.647 | 0.809 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.695 | 0.826 |
| RECIST | Patient | Chi-squared Test | 0.716 | 0.826 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.787 | 0.862 |
| Progression Free Censor | Patient | Chi-squared Test | 0.805 | 0.862 |
| All Mutations | Sample | Kruskal Wallis Test | 0.862 | 0.891 |
| Ploidy | Sample | Kruskal Wallis Test | 0.981 | 0.981 |

| ME_SMAD3+/- | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Treatment Group | Patient | Chi-squared Test | 0.0334 | 0.519 |
| First Line Therapy | Patient | Chi-squared Test | 0.0498 | 0.519 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0617 | 0.519 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.0956 | 0.519 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.102 | 0.519 |
| Sex | Patient | Chi-squared Test | 0.128 | 0.519 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.142 | 0.519 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.186 | 0.519 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.201 | 0.519 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.204 | 0.519 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.232 | 0.519 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.240 | 0.519 |
| Purity | Sample | Kruskal Wallis Test | 0.248 | 0.519 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.250 | 0.519 |
| Age | Patient | Kruskal Wallis Test | 0.266 | 0.519 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.277 | 0.519 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.307 | 0.541 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.543 | 0.763 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.560 | 0.763 |
| Genome Doublings | Sample | Chi-squared Test | 0.566 | 0.763 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.571 | 0.763 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.576 | 0.763 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.585 | 0.763 |
| Ploidy | Sample | Kruskal Wallis Test | 0.635 | 0.787 |
| Progression Free Censor | Patient | Chi-squared Test | 0.656 | 0.787 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.695 | 0.802 |
| All Mutations | Sample | Kruskal Wallis Test | 0.769 | 0.843 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.787 | 0.843 |
| RECIST | Patient | Chi-squared Test | 0.816 | 0.844 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.916 | 0.916 |

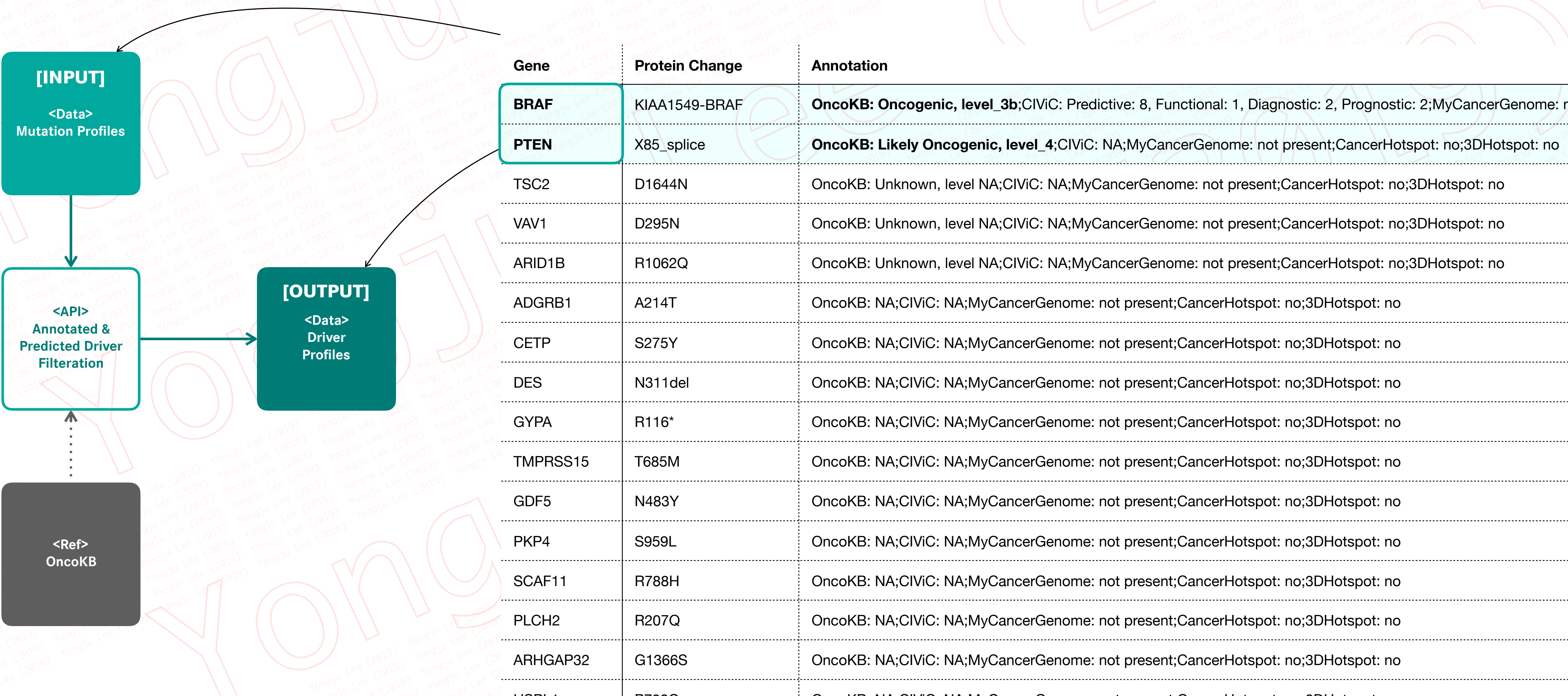
| DV_PBRM1+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 5.37E-03 | 0.161 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.0131 | 0.196 |
| RECIST | Patient | Chi-squared Test | 0.0275 | 0.275 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.0584 | 0.438 |
| Sex | Patient | Chi-squared Test | 0.0774 | 0.465 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.144 | 0.633 |
| Purity | Sample | Kruskal Wallis Test | 0.175 | 0.633 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.186 | 0.633 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.633 |
| Ploidy | Sample | Kruskal Wallis Test | 0.280 | 0.815 |
| Age | Patient | Kruskal Wallis Test | 0.327 | 0.815 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.815 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.393 | 0.815 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.815 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.408 | 0.815 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.454 | 0.851 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.501 | 0.885 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.552 | 0.891 |
| All Mutations | Sample | Kruskal Wallis Test | 0.564 | 0.891 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.645 | 0.903 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.659 | 0.903 |
| First Line Therapy | Patient | Chi-squared Test | 0.662 | 0.903 |
| Treatment Group | Patient | Chi-squared Test | 0.757 | 0.907 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.759 | 0.907 |
| Progression Free Censor | Patient | Chi-squared Test | 0.765 | 0.907 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.786 | 0.907 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.908 | 0.960 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.921 | 0.960 |
| Genome Doublings | Sample | Chi-squared Test | 0.948 | 0.960 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.960 | 0.960 |

| ME_EP300+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 3.29E-03 | 0.0593 |
| Response to Immunotherapy | Patient | Chi-squared Test | 4.0E-03 | 0.0593 |
| Sex | Patient | Chi-squared Test | 8.90E-03 | 0.0890 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0217 | 0.132 |
| RECIST | Patient | Chi-squared Test | 0.0220 | 0.132 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.0321 | 0.160 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0568 | 0.244 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.0921 | 0.345 |
| Age | Patient | Kruskal Wallis Test | 0.107 | 0.357 |
| First Line Therapy | Patient | Chi-squared Test | 0.120 | 0.359 |
| Treatment Group | Patient | Chi-squared Test | 0.161 | 0.438 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.475 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.310 | 0.716 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.769 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.769 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.410 | 0.769 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.447 | 0.782 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.469 | 0.782 |
| Purity | Sample | Kruskal Wallis Test | 0.497 | 0.784 |
| Progression Free Censor | Patient | Chi-squared Test | 0.568 | 0.845 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.592 | 0.845 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.620 | 0.845 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.691 | 0.901 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.721 | 0.901 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.781 | 0.937 |
| All Mutations | Sample | Kruskal Wallis Test | 0.842 | 0.953 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.858 | 0.953 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.889 | 0.953 |
| Genome Doublings | Sample | Chi-squared Test | 0.933 | 0.965 |
| Ploidy | Sample | Kruskal Wallis Test | 1.00 | 1.00 |

| UMAP_C1 vs C2 | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 2.84E-03 | 0.0515 |
| Response to Immunotherapy | Patient | Chi-squared Test | 3.43E-03 | 0.0515 |
| Sex | Patient | Chi-squared Test | 5.78E-03 | 0.0578 |
| RECIST | Patient | Chi-squared Test | 0.0171 | 0.128 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0367 | 0.220 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0497 | 0.249 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.0627 | 0.269 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.0921 | 0.345 |
| First Line Therapy | Patient | Chi-squared Test | 0.149 | 0.498 |
| Age | Patient | Kruskal Wallis Test | 0.189 | 0.518 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.518 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.231 | 0.577 |
| Treatment Group | Patient | Chi-squared Test | 0.281 | 0.648 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.724 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.387 | 0.724 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.724 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.410 | 0.724 |
| Purity | Sample | Kruskal Wallis Test | 0.451 | 0.741 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.469 | 0.741 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.608 | 0.856 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.608 | 0.856 |
| Progression Free Censor | Patient | Chi-squared Test | 0.628 | 0.856 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.676 | 0.880 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.704 | 0.880 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.775 | 0.894 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.819 | 0.894 |
| Ploidy | Sample | Kruskal Wallis Test | 0.851 | 0.894 |
| All Mutations | Sample | Kruskal Wallis Test | 0.879 | 0.894 |
| Genome Doublings | Sample | Chi-squared Test | 0.891 | 0.894 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.894 | 0.894 |

1. Retrieving patient's mutated gene profiles filtered by OncoKB annotations

[INPUT] Patient's mutated genes and their protein change retrieval
[FILTER] Identify gene-protein change from OncoKB for their status of oncogenesis (either 1. Oncogenic or 2. Likely oncogenic)
[OUTPUT] Patient's driver mutation profile



2. Enrichment Analysis

[INPUT] Based on extracted driver mutations and patient’s clinical attribute (inc. oncotree code and sample type of tumor sample),

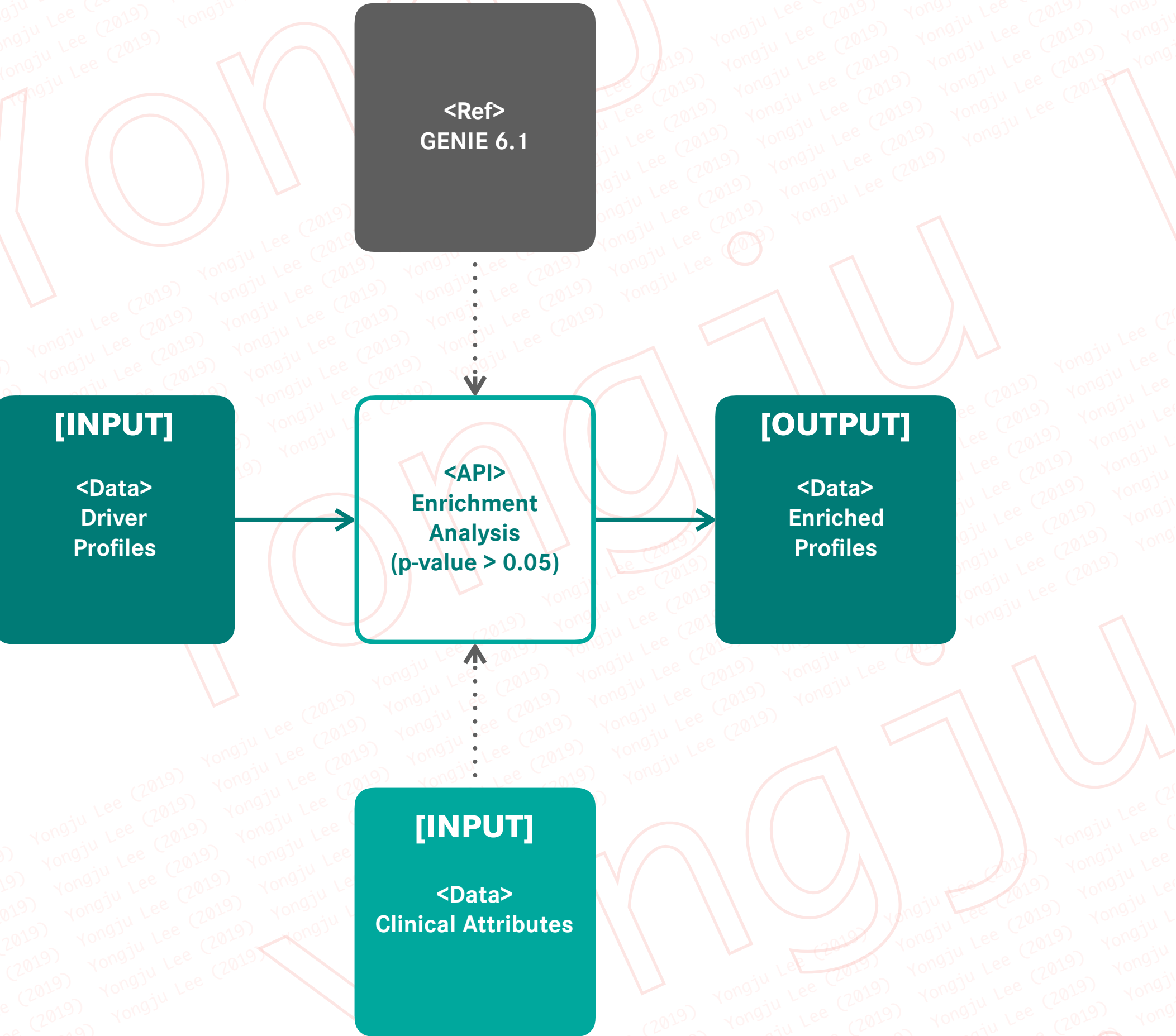
[FILTER]

A. Perform enrichment analysis with GENIE 6.1. CBioPortal Enrichment Tab with certain conditions applied.

***Conditions (1. Filter GENIE 6.1. patients with same clinical attribute; 2. Select somatic mutation profiles (not CNVs); 3. Enter driver genes with ‘: DRIVER;’)

B. Extract enriched genes with certain conditions applied.

1) Select patient-level enrichment, 2) Enrichment group : Mutual Exclusivity, 3) p-value cutoff : under 0.05



[INPUT]

| | |
|------|-----------|
| BRAF | : DRIVER; |
| PTEN | : DRIVER; |

Selected Studies:

Modify

PRAD_Metastasis (1071 total samples)

Select Genomic Profiles:

☒ Somatic mutations ?

☐ Copy-number alterations ?

Select Patient/Case Set:

To build your own case set, try out our enhanced Study View.

All (1071)

×

▼

Enter Genes:

User-defined List

×

▼

Hint: Learn Onco Query Language (OQL) to write more powerful queries

BRAF : DRIVER;
PTEN : DRIVER;

✔ All gene symbols are valid.

3. Pathway enrichment analysis

[INPUT]

Integrate both 1) mutated and 2) enriched profiles (GENE symbol only)

[FILTER]

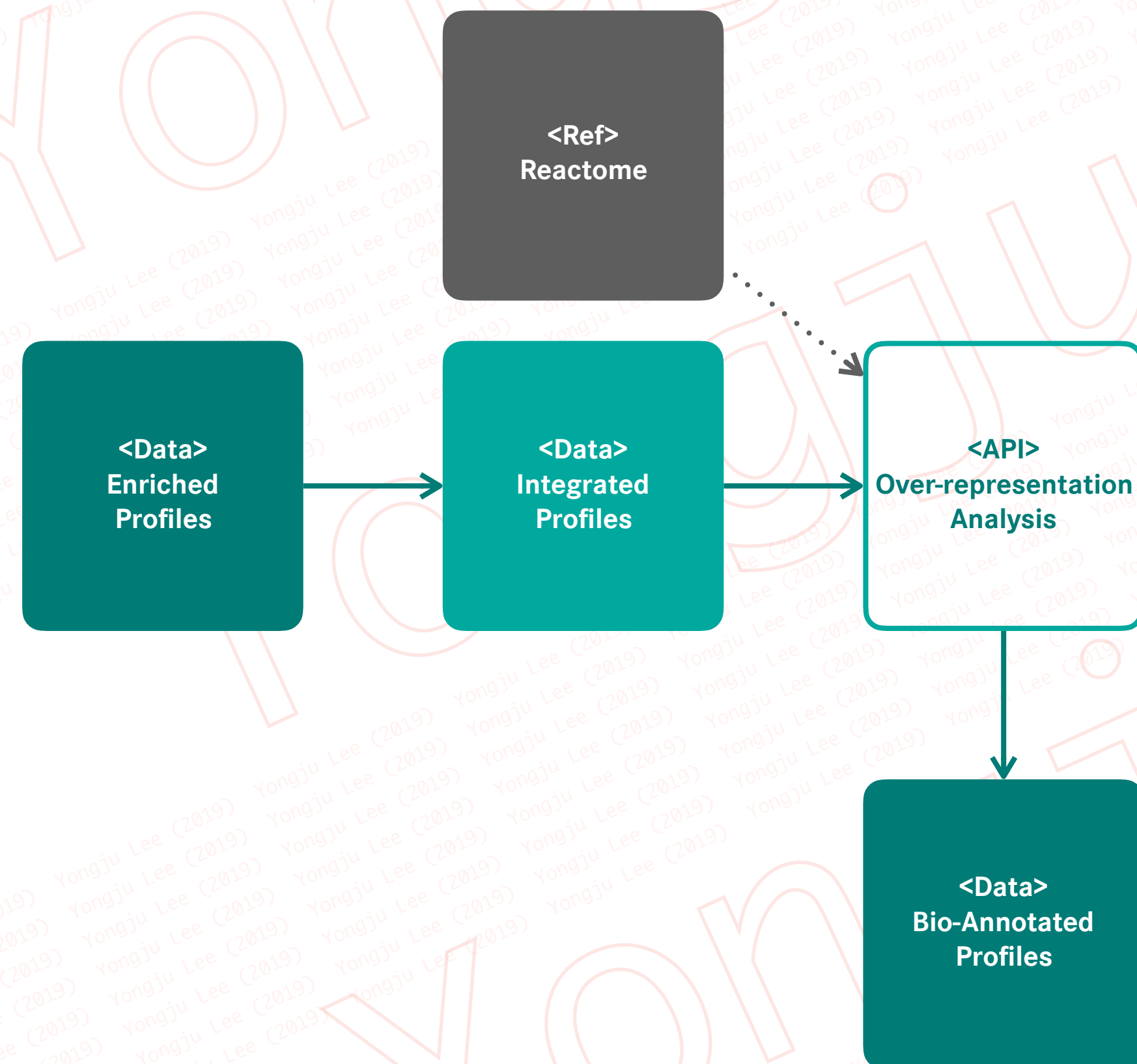
Perform pathway enrichment analysis with certain parameters applied.

A. Project to human = TRUE;

B. Include interactions = TRUE;

C. Apply $-\log_{10}()$ for both p-value column and FDR column

[OUTPUT] Patient specific ranked over-representation with p-value and FDR



Step 2: Select your preferred options.

- ✓ Project to human

 All non-human identifiers are converted to their human equivalents (expand for more info...)

☒ Include interactors

 IntAct interactors are used to increase the analysis background (expand for more info...)

[illegible]

4. UMAP-based clustering Preparation

[INPUT]

Prepare both 1) -log10(Entities p-value) and 2) -log10(Entities FDR).

[OUTPUT]

Generate both vector and label tsv.

| HSA_NAME | HSA_CODE | STDEV | AVG | CNT | 9115586 | 9115572 | 9115539 |
|--|---------------|-----------|-----------|-----|-------------------|-------------------|-------------------|
| Defective Mismatch Repair Associated With MLH1 | R-HSA-5545483 | ⚠ | 1.4055506 | 1 | 1.405550654493160 | | |
| Resolution of D-loop Structures through Synthesis-Dependent S | R-HSA-5693554 | 0.2039106 | 0.8533256 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| Resolution of D-loop Structures through Holliday Junction Interr | R-HSA-5693568 | 0.2020603 | 0.8524245 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| Resolution of D-Loop Structures | R-HSA-5693537 | 0.2014205 | 0.8518870 | 17 | 1.229146916723810 | 0.754337392352587 | 0.879823343399318 |
| HDR through Single Strand Annealing (SSA) | R-HSA-5685938 | 0.3092483 | 0.8995658 | 19 | 1.229146916723810 | 0.672630280305592 | 0.879823343399318 |
| Processing of DNA double-strand break ends | R-HSA-5693607 | 0.2144646 | 0.7749681 | 17 | 1.229146916723810 | 0.630004274129010 | 0.816978853975654 |
| HDR through Homologous Recombination (HRR) | R-HSA-5685942 | 0.1993074 | 0.7821921 | 17 | 1.229146916723810 | 0.590856054495176 | 0.775590112742391 |
| Meiotic recombination | R-HSA-912446 | 0.2184479 | 0.7518827 | 17 | 1.229146916723810 | 0.585624564469399 | 0.770039850544110 |
| Transcriptional Regulation by E2F6 | R-HSA-8953750 | 0.2871504 | 0.7176407 | 15 | 1.229146916723810 | 0.515963152367605 | 0.695636053797126 |
| SUMO E3 ligases SUMOylate target proteins | R-HSA-3108232 | 0.4393138 | 0.4551416 | 23 | 1.229146916723810 | 0.264650651162941 | 0.516064715395482 |
| SUMOylation | R-HSA-2990846 | 0.4390480 | 0.4475852 | 23 | 1.229146916723810 | 0.254496116843433 | 0.502298482580464 |
| Metabolism of proteins | R-HSA-392499 | 0.4951786 | 0.2633776 | 24 | 1.229146916723810 | 0.066480159992483 | 0.044547689690695 |
| TP53 Regulates Transcription of DNA Repair Genes | R-HSA-6796648 | 2.0919406 | 3.6150411 | 17 | 1.229146916723810 | | 4.29997324972002E |
| G2/M DNA damage checkpoint | R-HSA-69473 | 0.4657978 | 1.1867209 | 16 | 1.229146916723810 | | 0.879823343399318 |
| G2/M Checkpoints | R-HSA-69481 | 0.3154110 | 1.1020147 | 15 | 1.229146916723810 | | 0.879823343399318 |
| Regulation of TP53 Activity through Phosphorylation | R-HSA-6804756 | 0.3010464 | 1.0730907 | 18 | 1.229146916723810 | | 0.879823343399318 |
| Homologous DNA Pairing and Strand Exchange | R-HSA-5693579 | 0.1820286 | 0.8502934 | 14 | 1.229146916723810 | | 0.879823343399318 |
| Presynaptic phase of homologous DNA pairing and strand excha | R-HSA-5693616 | 0.1821164 | 0.8442072 | 14 | 1.229146916723810 | | 0.879823343399318 |
| Meiotic synapsis | R-HSA-1221632 | 0.2493609 | 0.6870204 | 15 | 1.229146916723810 | | 0.700141795961886 |

환자리스트

| | | | | | | | |
|----|-------------|-------------|-------------|-------------|-------------|-------------|-------------|
| 1 | 1.405550654 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 |
| 2 | 0 | 0.754337392 | 0.754337392 | 0.754337392 | 0.67263028 | 0.630004274 | 0.59085605 |
| 3 | 0 | 0.879823343 | 0.879823343 | 0.879823343 | 0.879823343 | 0.816978854 | 0.77559011 |
| 4 | 0 | 0 | 0 | 0 | 0 | 0.084607889 | 0.079360377 |
| 5 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 6 | 0 | 0 | 0 | 1.403568034 | 0 | 0 | 1.414996791 |
| 7 | 0 | 0 | 0 | 0.884457004 | 0 | 0 | 0.646277146 |
| 8 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.68353998 |
| 9 | 0 | 0 | 0 | 0 | 0 | 1.025544325 | 1.366544469 |
| 10 | 0 | 0 | 0 | 0 | 0 | 0.891488505 | 0.891488505 |
| 11 | 0 | 0.904915122 | 0.904915122 | 0.904915122 | 0.894130778 | 0.84905199 | 0.90491512 |
| 12 | 0 | 0.808416197 | 0.808416197 | 0.808416197 | 0.808416197 | 0.668179673 | 0.62847941 |
| 13 | 0 | 0.892110401 | 0.892110401 | 0.887955178 | 0.812566628 | 0.711170702 | 0.81256662 |
| 14 | 0 | 1.252824627 | 1.237505999 | 1.232523412 | 1.81101513 | 1.075948279 | 1.02675944 |
| 15 | 0 | 0 | 0 | 0 | 0 | 0.750537141 | 0.252258512 |
| 16 | 0 | 0.649690695 | 0.649690695 | 0.649690695 | 0.649690695 | 0.595762688 | 0.64969069 |
| 17 | 0 | 0.600928448 | 0.600928448 | 0.600928448 | 0.600928448 | 0.579904013 | 0.54157377 |
| 18 | 0 | 0.836561138 | 0.836561138 | 0.836561138 | 0.832002844 | 0.787477293 | 0.74637333 |
| 19 | 0 | 0.957861341 | 0.957861341 | 0.957861341 | 0.957861341 | 0.922837747 | 0.88058653 |
| 20 | 0 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.64902855 |
| 21 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.68353998 |
| 22 | 0 | 0.679030141 | 0.679030141 | 0.679030141 | 0.679030141 | 0.612437282 | 0.67903014 |
| 23 | 0 | 0.816846642 | 0.816846642 | 0.816846642 | 0.73243996 | 0.689006864 | 0.64902855 |
| 24 | 0 | 0 | 0 | 0 | 0 | 0.101121417 | 0.00E+00 |
| 25 | 0 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.16656087 |

Pivot된 벡터

-LOG10(FDR)값

| |
|----------|
| 9115586 |
| 9115572 |
| 9115539 |
| 9115522 |
| 9115517 |
| 9115425 |
| 9115422 |
| 9115339 |
| 97115125 |
| 9115109 |
| 9115059 |
| 9115051 |
| 6115120 |
| 6115114 |
| 6115247 |
| 6115395 |
| 96115135 |
| 5115650 |
| 5115615 |
| 5115610 |
| 5115609 |
| 5115359 |
| 5115075 |
| 5115031 |
| 1115019 |

벡터 순으로 나열된 환자 id

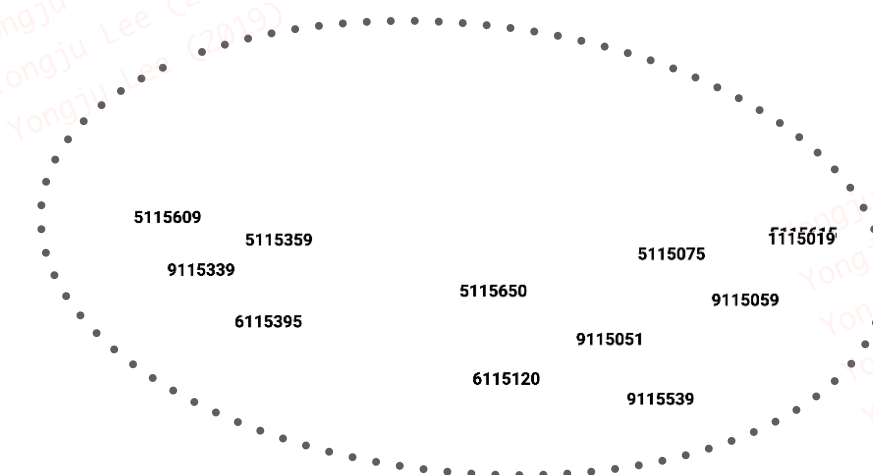
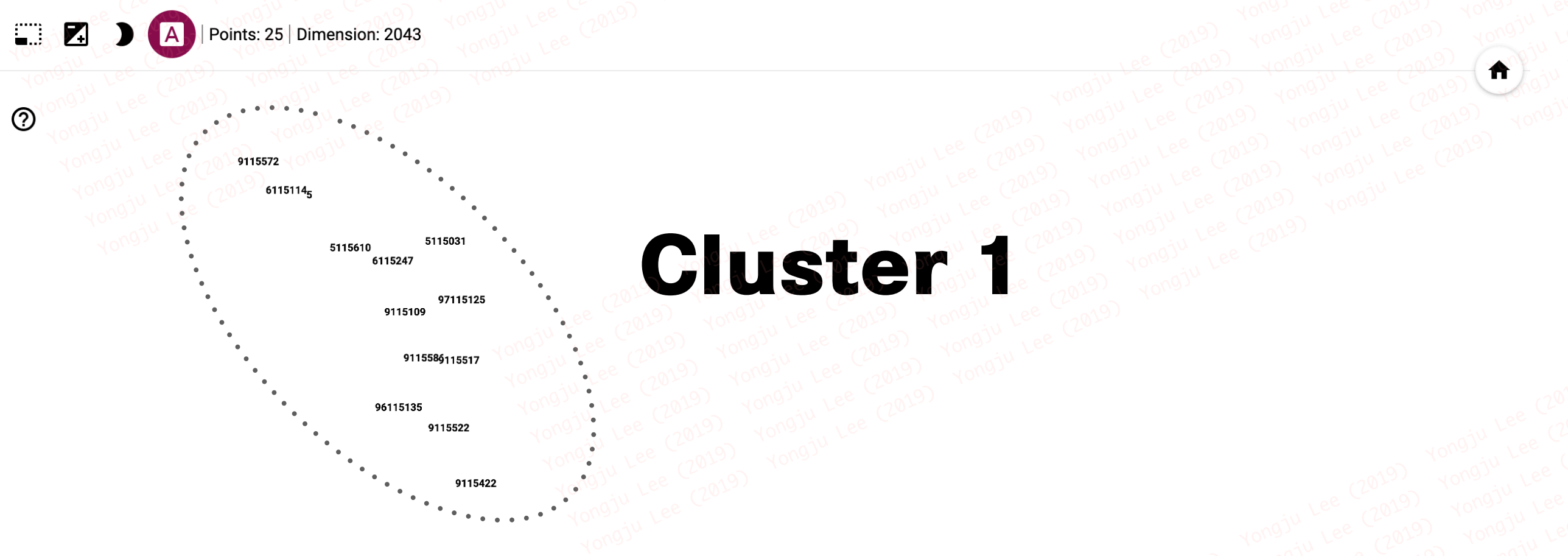
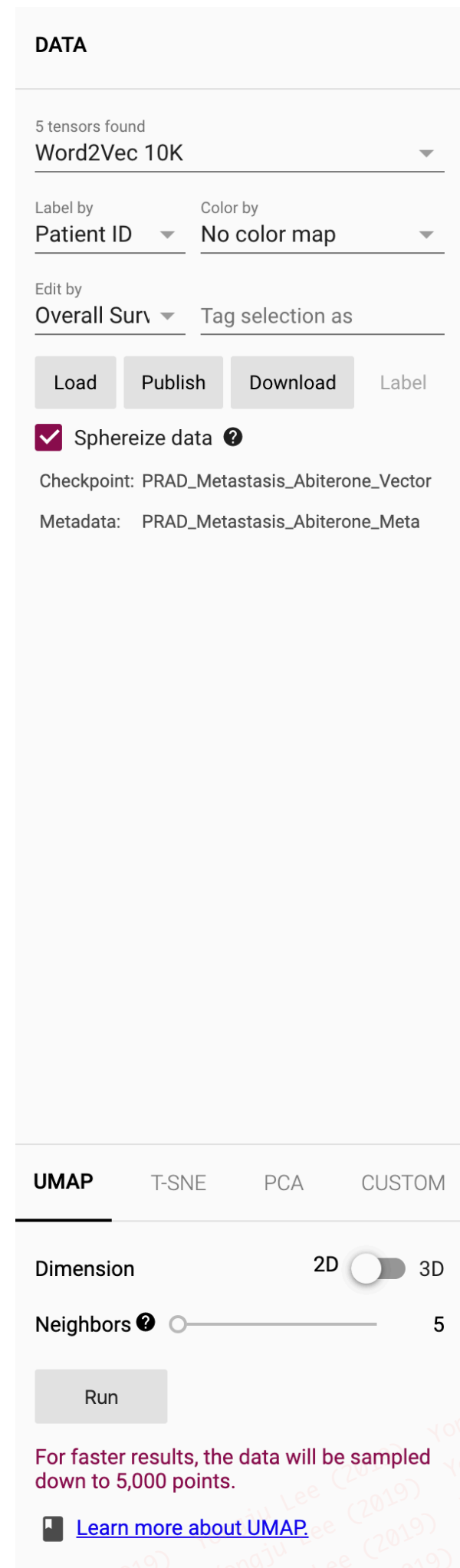
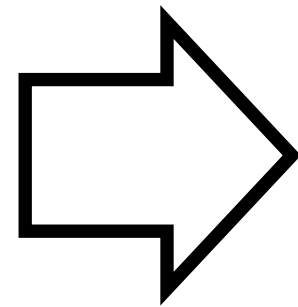
4. UMAP-based clustering

| | | | | | | | | |
|----|-------------|-------------|-------------|-------------|-------------|--------------|-------------|-------------|
| 1 | 1.405550654 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1.229146917 | 1 |
| 2 | 0 | 0.754337392 | 0.754337392 | 0.754337392 | 0.67263028 | 0.630004274 | 0.5908560 | |
| 3 | 0 | 0.879823343 | 0.879823343 | 0.879823343 | 0.879823343 | 0.816978854 | 0.7755901 | |
| 4 | 0 | 0 | 0 | 0 | 0 | 0.08460789 | 0.079360377 | 7.566E-05 |
| 5 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 6 | 0 | 0 | 0 | 0 | 1.403568034 | 0 | 0 | 0 |
| 7 | 0 | 0 | 0 | 0.884457004 | 0 | 0 | 0.646277146 | 0.366462845 |
| 8 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.6835399 | |
| 9 | 0 | 0 | 0 | 0 | 0 | 1.025544325 | 1.366544469 | 1.025544325 |
| 10 | 0 | 0 | 0 | 0 | 0 | 0 | 0.891488505 | 0.597070818 |
| 11 | 0 | 0.904915122 | 0.904915122 | 0.904915122 | 0.894130778 | 0.84905199 | 0.9049151 | |
| 12 | 0 | 0.808416197 | 0.808416197 | 0.808416197 | 0.808416197 | 0.668179673 | 0.6284794 | |
| 13 | 0 | 0.892110401 | 0.892110401 | 0.887955178 | 0.812566628 | 0.711170702 | 0.8125666 | |
| 14 | 0 | 1.252824627 | 1.237505999 | 1.232523412 | 1.81101513 | 1.075948279 | 1.026594 | |
| 15 | 0 | 0 | 0 | 0 | 0.750537141 | 0.0.25900116 | 0.252258512 | 0.026 |
| 16 | 0 | 0.649690695 | 0.649690695 | 0.649690695 | 0.649690695 | 0.595762688 | 0.6496906 | |
| 17 | 0 | 0.600928448 | 0.600928448 | 0.600928448 | 0.600928448 | 0.579904013 | 0.5145737 | |
| 18 | 0 | 0.836561138 | 0.836561138 | 0.836561138 | 0.832002844 | 0.787477293 | 0.7463733 | |
| 19 | 0 | 0.957861341 | 0.957861341 | 0.957861341 | 0.957861341 | 0.922837747 | 0.8805865 | |
| 20 | 0 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.685993135 | 0.6490285 | |
| 21 | 0 | 0.683539988 | 0.683539988 | 0.683539988 | 0.683539988 | 0.564794148 | 0.6835399 | |
| 22 | 0 | 0.679030141 | 0.679030141 | 0.679030141 | 0.679030141 | 0.612437282 | 0.6790301 | |
| 23 | 0 | 0.816846642 | 0.816846642 | 0.816846642 | 0.73243996 | 0.689006804 | 0.6490285 | |
| 24 | 0 | 0 | 0 | 0 | 0 | 0 | 0.101121417 | 0.00E+00 |
| 25 | b | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.190970032 | 1.1665608 | |

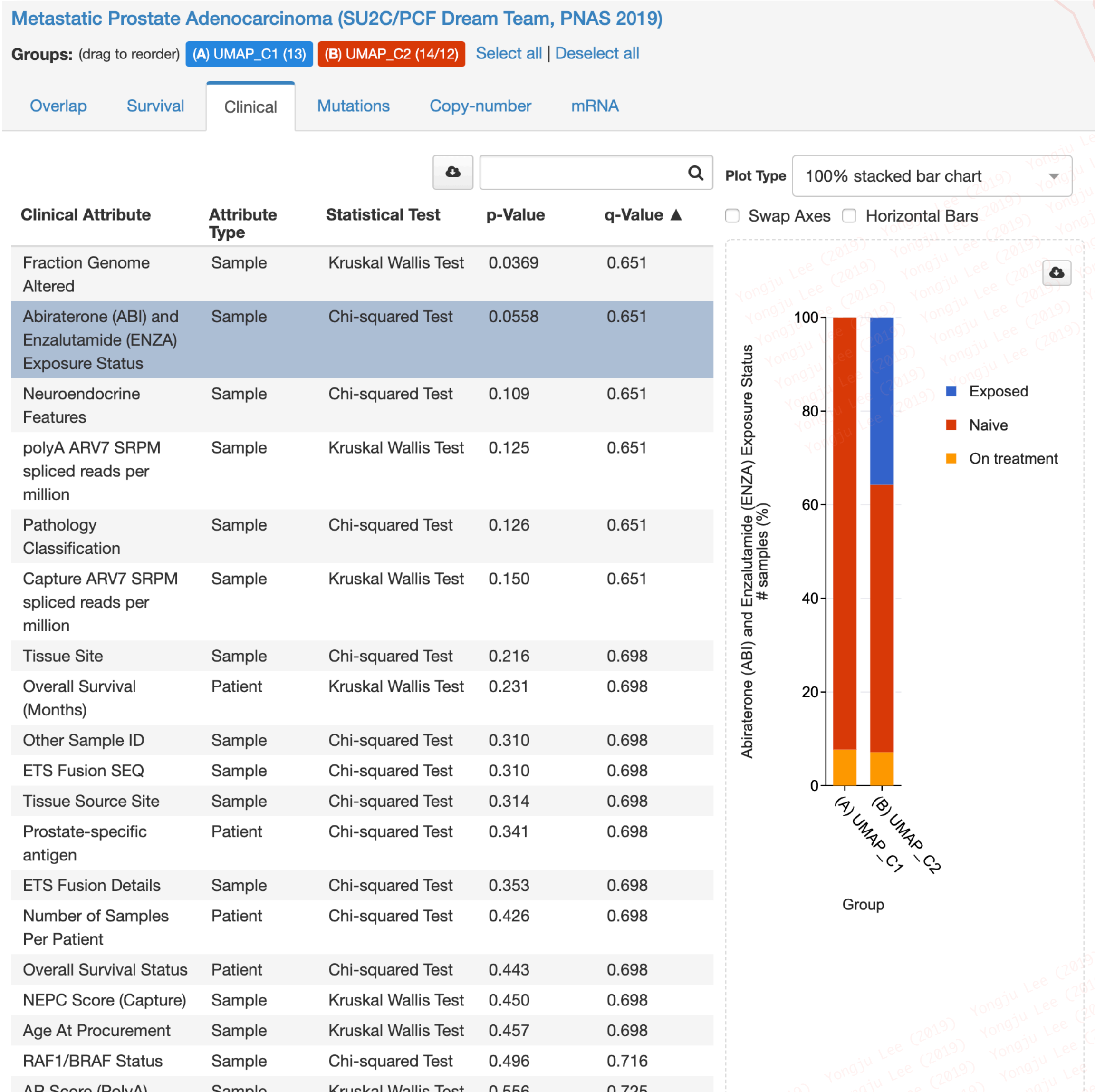
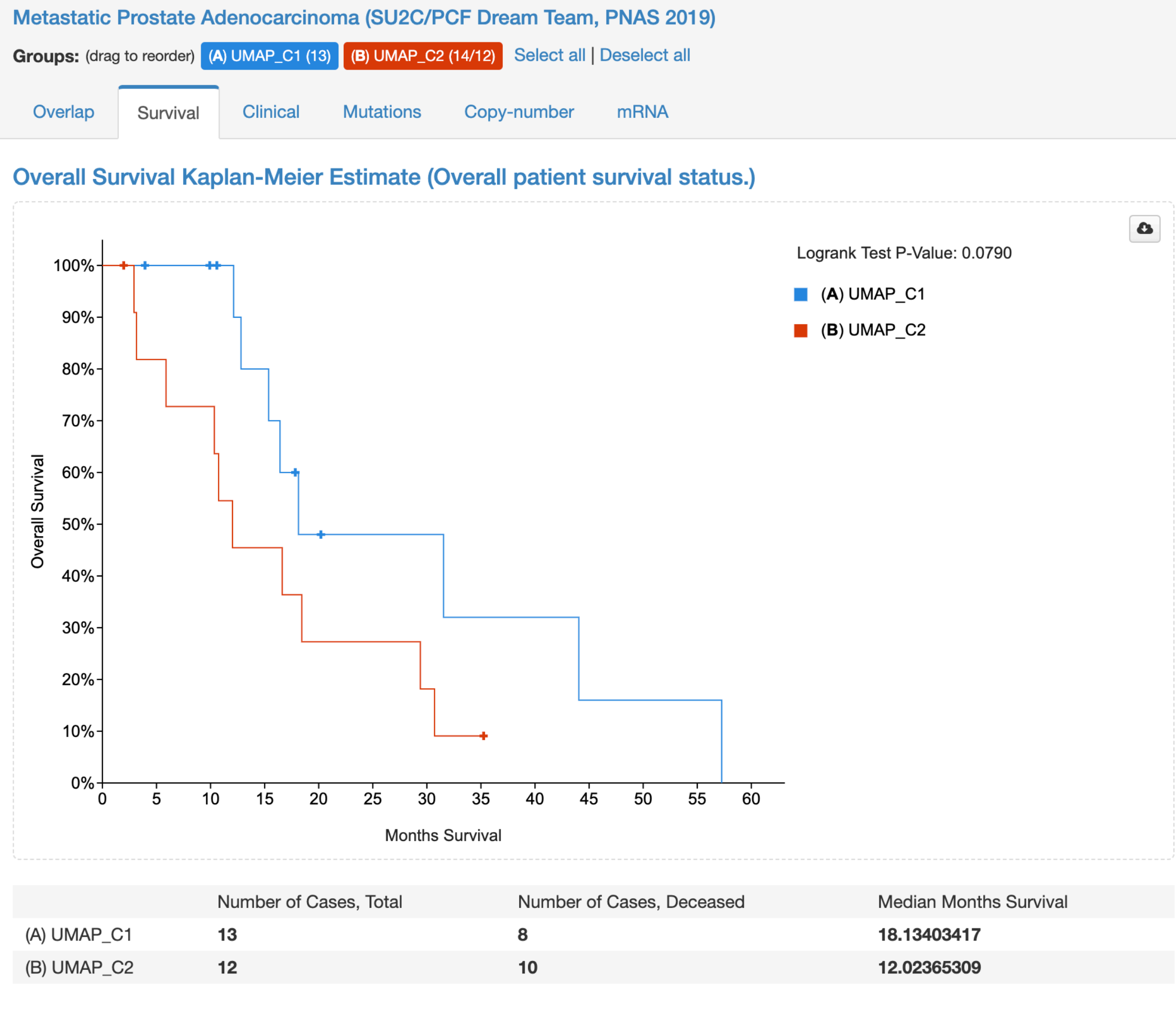
Pivot된 벡터

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6115395
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5115359
5115075
5115031
1115019

벡터 순으로 나열된 환자 id



5. Cluster based group Analysis



SUMMIT TRIAL (2018)

Driver Only

With Enriched Entities (CO*s & ME*s)

With Enriched Entities (ME* only)

without
Over-representation*

| DRIVER_TP53+/- | | | | | DRIVER_PIK3CA+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value | Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Sample | Kruskal Wallis Test | 4.6E-03 | 0.0462 | Mutation Count | Sample | Kruskal Wallis Test | 0.0354 | 0.239 |
| Mutation Burden | Sample | Kruskal Wallis Test | 4.62E-03 | 0.0462 | Mutation Burden | Sample | Kruskal Wallis Test | 0.0357 | 0.239 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0143 | 0.0954 | Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0359 | 0.239 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0650 | 0.288 | Waterfall Group | Sample | Chi-squared Test | 0.0920 | 0.460 |
| Waterfall Group | Sample | Chi-squared Test | 0.0821 | 0.288 | Best objective response rate | Patient | Chi-squared Test | 0.189 | 0.696 |
| Coverage | Sample | Kruskal Wallis Test | 0.0864 | 0.288 | Purity | Sample | Kruskal Wallis Test | 0.234 | 0.696 |
| Bubble Group | Sample | Chi-squared Test | 0.159 | 0.453 | Bubble Group | Sample | Chi-squared Test | 0.244 | 0.696 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.252 | 0.554 | Qualifying Mutation | Sample | Chi-squared Test | 0.363 | 0.744 |
| Central | Sample | Chi-squared Test | 0.256 | 0.554 | Central | Sample | Chi-squared Test | 0.371 | 0.744 |
| Local Test | Sample | Chi-squared Test | 0.277 | 0.554 | Local Test | Sample | Chi-squared Test | 0.372 | 0.744 |
| Best objective response rate | Patient | Chi-squared Test | 0.392 | 0.712 | Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.514 | 0.801 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.732 | MSI Status | Sample | Chi-squared Test | 0.585 | 0.801 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.732 | Oncoprint | Sample | Chi-squared Test | 0.585 | 0.801 |
| Data | Sample | Chi-squared Test | 0.535 | 0.732 | Criteria | Sample | Chi-squared Test | 0.653 | 0.801 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.732 | Clonality | Sample | Chi-squared Test | 0.661 | 0.801 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.840 | IMPACT | Sample | Chi-squared Test | 0.661 | 0.801 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.840 | Coverage | Sample | Kruskal Wallis Test | 0.695 | 0.801 |
| Amplification Status | Sample | Chi-squared Test | 0.788 | 0.846 | Data | Sample | Chi-squared Test | 0.721 | 0.801 |
| Purity | Sample | Kruskal Wallis Test | 0.808 | 0.846 | ER_PR | Sample | Chi-squared Test | 0.836 | 0.880 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.846 | Amplification Status | Sample | Chi-squared Test | 0.954 | 0.954 |

N/A

with
Over-representation*

| 2 GROUP CLUSTERING (DRIVER) | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 1.46E-03 | 0.0293 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 9.20E-03 | 0.0642 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0127 | 0.0642 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0128 | 0.0642 |
| Coverage | Sample | Kruskal Wallis Test | 0.0451 | 0.181 |
| Waterfall Group | Sample | Chi-squared Test | 0.0567 | 0.189 |
| Bubble Group | Sample | Chi-squared Test | 0.125 | 0.338 |
| Best objective response rate | Patient | Chi-squared Test | 0.135 | 0.338 |
| Central | Sample | Chi-squared Test | 0.162 | 0.361 |
| Data | Sample | Chi-squared Test | 0.191 | 0.372 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.205 | 0.372 |
| IMPACT | Sample | Chi-squared Test | 0.356 | 0.594 |
| Local Test | Sample | Chi-squared Test | 0.509 | 0.783 |
| ER_PR | Sample | Chi-squared Test | 0.576 | 0.823 |
| Amplification Status | Sample | Chi-squared Test | 0.642 | 0.856 |
| Clonality | Sample | Chi-squared Test | 0.693 | 0.866 |
| Purity | Sample | Kruskal Wallis Test | 0.808 | 0.876 |
| MSI Status | Sample | Chi-squared Test | 0.833 | 0.876 |
| Oncoprint | Sample | Chi-squared Test | 0.833 | 0.876 |
| Criteria | Sample | Chi-squared Test | 0.933 | 0.933 |

| 2 GROUP CLUSTERING (ENR) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0191 | 0.138 |
| Central | Sample | Chi-squared Test | 0.0220 | 0.138 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0227 | 0.138 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0365 | 0.138 |
| Best objective response rate | Patient | Chi-squared Test | 0.0384 | 0.138 |
| Purity | Sample | Kruskal Wallis Test | 0.0415 | 0.138 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0662 | 0.189 |
| Local Test | Sample | Chi-squared Test | 0.174 | 0.399 |
| Amplification Status | Sample | Chi-squared Test | 0.179 | 0.399 |
| Data | Sample | Chi-squared Test | 0.253 | 0.496 |
| Waterfall Group | Sample | Chi-squared Test | 0.273 | 0.496 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.744 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.744 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.744 |
| Bubble Group | Sample | Chi-squared Test | 0.558 | 0.744 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.677 | 0.794 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.794 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.794 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.891 |
| Coverage | Sample | Kruskal Wallis Test | 0.903 | 0.903 |

| 3 GROUP CLUSTERING (ENR) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Central | Sample | Chi-squared Test | 0.0389 | 0.284 |
| Purity | Sample | Kruskal Wallis Test | 0.0498 | 0.284 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0524 | 0.284 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0639 | 0.284 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0745 | 0.284 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0933 | 0.284 |
| Amplification Status | Sample | Chi-squared Test | 0.0995 | 0.284 |
| Data | Sample | Chi-squared Test | 0.136 | 0.307 |
| Best objective response rate | Patient | Chi-squared Test | 0.143 | 0.307 |
| Clonality | Sample | Chi-squared Test | 0.153 | 0.307 |
| MSI Status | Sample | Chi-squared Test | 0.229 | 0.382 |
| Oncoprint | Sample | Chi-squared Test | 0.229 | 0.382 |
| Waterfall Group | Sample | Chi-squared Test | 0.249 | 0.382 |
| Local Test | Sample | Chi-squared Test | 0.267 | 0.382 |
| IMPACT | Sample | Chi-squared Test | 0.318 | 0.424 |
| Bubble Group | Sample | Chi-squared Test | 0.501 | 0.599 |
| ER_PR | Sample | Chi-squared Test | 0.509 | 0.599 |
| Coverage | Sample | Kruskal Wallis Test | 0.630 | 0.701 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.676 | 0.712 |
| Criteria | Sample | Chi-squared Test | 0.825 | 0.825 |

| 2 GROUP CLUSTERING (ME) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Mutation Count | Patient | Kruskal Wallis Test | 0.0191 | 0.138 |
| Central | Sample | Chi-squared Test | 0.0220 | 0.138 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0227 | 0.138 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0365 | 0.138 |
| Best objective response rate | Patient | Chi-squared Test | 0.0384 | 0.138 |
| Purity | Sample | Kruskal Wallis Test | 0.0415 | 0.138 |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 0.0662 | 0.189 |
| Local Test | Sample | Chi-squared Test | 0.174 | 0.399 |
| Amplification Status | Sample | Chi-squared Test | 0.179 | 0.399 |
| Data | Sample | Chi-squared Test | 0.253 | 0.496 |
| Waterfall Group | Sample | Chi-squared Test | 0.273 | 0.496 |
| Clonality | Sample | Chi-squared Test | 0.494 | 0.744 |
| IMPACT | Sample | Chi-squared Test | 0.494 | 0.744 |
| Criteria | Sample | Chi-squared Test | 0.549 | 0.744 |
| Bubble Group | Sample | Chi-squared Test | 0.558 | 0.744 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.677 | 0.794 |
| MSI Status | Sample | Chi-squared Test | 0.714 | 0.794 |
| Oncoprint | Sample | Chi-squared Test | 0.714 | 0.794 |
| ER_PR | Sample | Chi-squared Test | 0.846 | 0.891 |
| Coverage | Sample | Kruskal Wallis Test | 0.903 | 0.903 |

| 3 GROUP CLUSTERING (ME) | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Duration of Therapy Days | Patient | Kruskal Wallis Test | 6.3E-03 | 0.127 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.0242 | 0.133 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0260 | 0.133 |
| Mutation Burden | Sample | Kruskal Wallis Test | 0.0266 | 0.133 |
| Best objective response rate | Patient | Chi-squared Test | 0.0530 | 0.171 |
| Purity | Sample | Kruskal Wallis Test | 0.0599 | 0.171 |
| Coverage | Sample | Kruskal Wallis Test | 0.0603 | 0.171 |
| Central | Sample | Chi-squared Test | 0.0683 | 0.171 |
| Amplification Status | Sample | Chi-squared Test | 0.0995 | 0.221 |
| Waterfall Group | Sample | Chi-squared Test | 0.211 | 0.408 |
| ER_PR | Sample | Chi-squared Test | 0.241 | 0.408 |
| IMPACT | Sample | Chi-squared Test | 0.245 | 0.408 |
| Bubble Group | Sample | Chi-squared Test | 0.277 | 0.426 |
| Data | Sample | Chi-squared Test | 0.342 | 0.488 |
| Qualifying Mutation | Sample | Chi-squared Test | 0.435 | 0.527 |
| MSI Status | Sample | Chi-squared Test | 0.448 | 0.527 |
| Oncoprint | Sample | Chi-squared Test | 0.448 | 0.527 |
| Local Test | Sample | Chi-squared Test | 0.494 | 0.548 |
| Criteria | Sample | Chi-squared Test | 0.825 | 0.869 |
| Clonality | Sample | Chi-squared Test | 0.949 | 0.949 |

- CO : Co-occurrent mutations
- ME : Mutually exclusive mutations
- Over-representation : Reactome based enrichment analysis

CCRCC (2019)

| DV_VHL+/- | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Sex | Patient | Chi-squared Test | 0.0350 | 0.484 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0561 | 0.484 |
| First Line Therapy | Patient | Chi-squared Test | 0.0781 | 0.484 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.0815 | 0.484 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.0956 | 0.484 |
| Treatment Group | Patient | Chi-squared Test | 0.112 | 0.484 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.133 | 0.484 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.139 | 0.484 |
| Purity | Sample | Kruskal Wallis Test | 0.152 | 0.484 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.170 | 0.484 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.184 | 0.484 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.193 | 0.484 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.250 | 0.571 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.277 | 0.571 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.285 | 0.571 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.370 | 0.668 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.378 | 0.668 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.433 | 0.722 |
| Genome Doublings | Sample | Chi-squared Test | 0.498 | 0.784 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.571 | 0.784 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.576 | 0.784 |
| Age | Patient | Kruskal Wallis Test | 0.601 | 0.784 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.601 | 0.784 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.647 | 0.809 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.695 | 0.826 |
| RECIST | Patient | Chi-squared Test | 0.716 | 0.826 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.787 | 0.862 |
| Progression Free Censor | Patient | Chi-squared Test | 0.805 | 0.862 |
| All Mutations | Sample | Kruskal Wallis Test | 0.862 | 0.891 |
| Ploidy | Sample | Kruskal Wallis Test | 0.981 | 0.981 |

| ME_SMAD3+/- | | | | |
|---------------------------------|----------------|---------------------|---------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Treatment Group | Patient | Chi-squared Test | 0.0334 | 0.519 |
| First Line Therapy | Patient | Chi-squared Test | 0.0498 | 0.519 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0617 | 0.519 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.0956 | 0.519 |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 0.102 | 0.519 |
| Sex | Patient | Chi-squared Test | 0.128 | 0.519 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.142 | 0.519 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.186 | 0.519 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.201 | 0.519 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.204 | 0.519 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.232 | 0.519 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.240 | 0.519 |
| Purity | Sample | Kruskal Wallis Test | 0.248 | 0.519 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.250 | 0.519 |
| Age | Patient | Kruskal Wallis Test | 0.266 | 0.519 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.277 | 0.519 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.307 | 0.541 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.543 | 0.763 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.560 | 0.763 |
| Genome Doublings | Sample | Chi-squared Test | 0.566 | 0.763 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.571 | 0.763 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.576 | 0.763 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.585 | 0.763 |
| Ploidy | Sample | Kruskal Wallis Test | 0.635 | 0.787 |
| Progression Free Censor | Patient | Chi-squared Test | 0.656 | 0.787 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.695 | 0.802 |
| All Mutations | Sample | Kruskal Wallis Test | 0.769 | 0.843 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.787 | 0.843 |
| RECIST | Patient | Chi-squared Test | 0.816 | 0.844 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.916 | 0.916 |

| DV_PBRM1+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 5.37E-03 | 0.161 |
| Response to Immunotherapy | Patient | Chi-squared Test | 0.0131 | 0.196 |
| RECIST | Patient | Chi-squared Test | 0.0275 | 0.275 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.0584 | 0.438 |
| Sex | Patient | Chi-squared Test | 0.0774 | 0.465 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.144 | 0.633 |
| Purity | Sample | Kruskal Wallis Test | 0.175 | 0.633 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.186 | 0.633 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.633 |
| Ploidy | Sample | Kruskal Wallis Test | 0.280 | 0.815 |
| Age | Patient | Kruskal Wallis Test | 0.327 | 0.815 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.815 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.393 | 0.815 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.815 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.408 | 0.815 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.454 | 0.851 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.501 | 0.885 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.552 | 0.891 |
| All Mutations | Sample | Kruskal Wallis Test | 0.564 | 0.891 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.645 | 0.903 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.659 | 0.903 |
| First Line Therapy | Patient | Chi-squared Test | 0.662 | 0.903 |
| Treatment Group | Patient | Chi-squared Test | 0.757 | 0.907 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.759 | 0.907 |
| Progression Free Censor | Patient | Chi-squared Test | 0.765 | 0.907 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.786 | 0.907 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.908 | 0.960 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.921 | 0.960 |
| Genome Doublings | Sample | Chi-squared Test | 0.948 | 0.960 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.960 | 0.960 |

| ME_EP300+/- | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 3.29E-03 | 0.0593 |
| Response to Immunotherapy | Patient | Chi-squared Test | 4.0E-03 | 0.0593 |
| Sex | Patient | Chi-squared Test | 8.90E-03 | 0.0890 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0217 | 0.132 |
| RECIST | Patient | Chi-squared Test | 0.0220 | 0.132 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.0321 | 0.160 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0568 | 0.244 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.0921 | 0.345 |
| Age | Patient | Kruskal Wallis Test | 0.107 | 0.357 |
| First Line Therapy | Patient | Chi-squared Test | 0.120 | 0.359 |
| Treatment Group | Patient | Chi-squared Test | 0.161 | 0.438 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.475 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.310 | 0.716 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.769 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.769 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.410 | 0.769 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.447 | 0.782 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.469 | 0.782 |
| Purity | Sample | Kruskal Wallis Test | 0.497 | 0.784 |
| Progression Free Censor | Patient | Chi-squared Test | 0.568 | 0.845 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.592 | 0.845 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.620 | 0.845 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.691 | 0.901 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.721 | 0.901 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.781 | 0.937 |
| All Mutations | Sample | Kruskal Wallis Test | 0.842 | 0.953 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.858 | 0.953 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.889 | 0.953 |
| Genome Doublings | Sample | Chi-squared Test | 0.933 | 0.965 |
| Ploidy | Sample | Kruskal Wallis Test | 1.00 | 1.00 |

| UMAP_C1 vs C2 | | | | |
|---------------------------------|----------------|---------------------|----------|---------|
| Clinical Attribute | Attribute Type | Statistical Test | p-Value | q-Value |
| Progress Free Survival (Months) | Patient | Kruskal Wallis Test | 2.84E-03 | 0.0515 |
| Response to Immunotherapy | Patient | Chi-squared Test | 3.43E-03 | 0.0515 |
| Sex | Patient | Chi-squared Test | 5.78E-03 | 0.0578 |
| RECIST | Patient | Chi-squared Test | 0.0171 | 0.128 |
| NonSynonymous Uneval | Sample | Kruskal Wallis Test | 0.0367 | 0.220 |
| FrameShift Indels | Sample | Kruskal Wallis Test | 0.0497 | 0.249 |
| Overall Survival (Months) | Patient | Kruskal Wallis Test | 0.0627 | 0.269 |
| PDL1 Positive 1% | Patient | Kruskal Wallis Test | 0.0921 | 0.345 |
| First Line Therapy | Patient | Chi-squared Test | 0.149 | 0.498 |
| Age | Patient | Kruskal Wallis Test | 0.189 | 0.518 |
| Cell Membrane PDL1 2%+ | Patient | Kruskal Wallis Test | 0.190 | 0.518 |
| Mutations Uneval | Sample | Kruskal Wallis Test | 0.231 | 0.577 |
| Treatment Group | Patient | Chi-squared Test | 0.281 | 0.648 |
| Cell Membrane PDL1 3%+ | Patient | Kruskal Wallis Test | 0.376 | 0.724 |
| Overall Survival Censor | Patient | Chi-squared Test | 0.387 | 0.724 |
| PDL1 Positive 5% | Patient | Kruskal Wallis Test | 0.398 | 0.724 |
| Cell Membrane PDL1 0% | Patient | Kruskal Wallis Test | 0.410 | 0.724 |
| Purity | Sample | Kruskal Wallis Test | 0.451 | 0.741 |
| Cell Membrane PDL1 1%+ | Patient | Kruskal Wallis Test | 0.469 | 0.741 |
| Nonsynonymous Mutation Count | Sample | Kruskal Wallis Test | 0.608 | 0.856 |
| Nonsynonymous Clonal | Sample | Kruskal Wallis Test | 0.608 | 0.856 |
| Progression Free Censor | Patient | Chi-squared Test | 0.628 | 0.856 |
| Mutations Clonal | Sample | Kruskal Wallis Test | 0.676 | 0.880 |
| Mutation Count | Sample | Kruskal Wallis Test | 0.704 | 0.880 |
| All Synonymous Mutations | Sample | Kruskal Wallis Test | 0.775 | 0.894 |
| Mutations Subclonal | Sample | Kruskal Wallis Test | 0.819 | 0.894 |
| Ploidy | Sample | Kruskal Wallis Test | 0.851 | 0.894 |
| All Mutations | Sample | Kruskal Wallis Test | 0.879 | 0.894 |
| Genome Doublings | Sample | Chi-squared Test | 0.891 | 0.894 |
| NonSynonymous Subclonal | Sample | Kruskal Wallis Test | 0.894 | 0.894 |