Supplementary Note 1. PCAWG data wrangling carried out in UCSC Xena.

PCAWG analysis working group primary results files (listed in the Data Availability Section) were downloaded and wrangled into two Xena cohorts: (i) the 'PCAWG donor-centric' cohort, in which all datasets use ICGC Donor IDs, such as DO217962 and (ii) the 'PCAWG specimen-centric' cohort, in which all datasets use ICGC Specimen IDs, such as SP117136.

Several steps were taken to wrangle all genomic and phenotypic datasets into Xena. For genomic datasets, we mapped data from aliquot IDs to donor IDs for the donor-centric cohort, and to specimen IDs for the specimen-centric cohort. Specific to the donor-centric cohort, data from normal specimens were removed, so the data represent only tumor profiles. When multiple specimens were available for the same donor, an average was taken and assigned to the donor.

For phenotype data, we mapped the specimen histology classifications back to the donor IDs after data from normal specimens were removed. We propagated donor clinical data to all specimen IDs belonging to the donor.

We extracted coding mutations from the consensus simple somatic mutation datasets and made the resulting dataset available on the open-access PCAWG hub (Consensus SNVs and indels - coding). The protected whole-genome consensus dataset containing simple somatic mutations was downloaded and wrangled into Xena-ready format. The resulting Xena-ready file was uploaded to GNOS (https://gtrepo-osdc-

tcga.annailabs.com/cghub/metadata/analysisFull/86a1f73a-fabf-4abe-a597-a47d84a8c980), in which the data remains under controlled access.

Supplementary Note 2. Recreating the panels in Figure 1.

ICGC Data Repository (1a)

Figure 1a shows a view of the PCAWG data files in the ICGC Data Repository. It can be accessed through https://dcc.icgc.org/repositories?filters=%7B%22file%22:%7B%22study%22:%7B%22is%22:%5 B%22PCAWG%22%5D%7D%7D%7D&files=%7B%22from%22:1%7D Shortened url: https://tinyurl.com/yburnde5

UCSC Xena Visual Spreadsheet (1b)

Figure 1b shows a view of the *ERG* fusion based on RNAseq and DNAseq data. It can be accessed through https://xenabrowser.pet/beatmap/2bookmark=29d8db43ebacda4f54e666d7666cf164

https://xenabrowser.net/heatmap/?bookmark=29d8db43ebacda4f54e666d7666cf164 Shortened url: https://tinyurl.com/y78adbl5

Chromothripsis Explorer (1c)

Figure 1c shows a Circos plot visualization of a donor (DO38283) for whom fusion of the oncogene *ERG* was detected. To reproduce Figure 1c; (i) visit

http://compbio.med.harvard.edu/chromothripsis/; (ii) click on the blue box containing the text 'CLICK HERE TO EXPAND' below the panel 'Find cases of interest quickly!'; (iii) type the ICGC donor ID for the tumor (e.g., DO38283 for the first result listed in Figure 1b) in the search bar below the column 'ICGC donor ID'. The table shows associated information for the donor, including the donor_unique_id (e.g., PRAD-US::f6381367-142c-45d0-92b3-c1727d1813ce); (iv) go to the panel 'Interactive circos plots reporting SNVs, indels, total CNV, minor CN (LOH), and SV calls' and select (e.g., 'PROST-ADENOCA') in the dropdown menu 'Cancer type' located on the left-hand side menu. Next, select the row corresponding to the donor of interest (e.g., PRAD-US::f6381367-142c-45d0-92b3-c1727d1813ce) in the dropdown menu 'Donor ID'; (v) finally, select the number of chromosomes to display in the Circos plot using the dropdown menu 'Chromosome selection for circos plot'.

Expression Atlas (1d)

The query in Figure 1d, showing PCAWG data for prostate adenocarcinoma together with adjacent normal tissue and normal prostate gland tissue from GTEx can be accessed through

https://www.ebi.ac.uk/gxa/experiments/E-MTAB-

5200/Results?specific=true&geneQuery=%255B%257B%2522value%2522%253A%2522Erg% 2522%252C%2522category%2522%253A%2522symbol%2522%257D%252C%257B%2522val ue%2522%253A%2522tmprss2%2522%252C%2522category%2522%253A%2522symbol%25 22%257D%252C%257B%2522value%2522%253A%2522SLC45A3%2522%252C%2522categ ory%2522%253A%2522symbol%2522%257D%255D&filterFactors=%257B%2522DISEASE%2 522%253A%255B%2522normal%2520-

%2520blood%2520%28GTEx%29%2522%252C%2522normal%2522%252C%2522normal%25 20-%2520amygdala%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520Brodmann%2520%281909%29%2520area%252024%2520%28GTEx%29%2522%252C %2522normal%2520-

%2520Brodmann%2520%281909%29%2520area%25209%2520%28GTEx%29%2522%252C %2522normal%2520-

%2520C1%2520segment%2520of%2520cervical%2520spinal%2520cord%2520%28GTEx%29 %2522%252C%2522normal%2520-

%2520caudate%2520nucleus%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520cerebellar%2520hemisphere%2520%28GTEx%29%2522%252C%2522normal%2520%2520cerebellum%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520cerebral%2520cortex%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520hippocampus%2520proper%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520hypothalamus%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520nucleus%2520accumbens%2520%28GTEx%29%2522%252C%2522normal%2520%2520putamen%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520substantia%2520nigra%2520%28GTEx%29%2522%252C%2522normal%2520-%2520breast%2520%28GTEx%29%2522%252C%2522normal%2520-

%2520esophagogastric%2520junction%2520%28GTEx%29%2522%252C%2522normal%2520 -%2520esophagus%2520mucosa%2520%28GTEx%29%2522%252C%2522normal%2520-%2520esophagus%2520muscularis%2520mucosa%2520%28GTEx%29%2522%252C%2522n ormal%2520-

%2520cortex%2520of%2520kidney%2520%28GTEx%29%2522%252C%2522normal%2520-%2520sigmoid%2520colon%2520%28GTEx%29%2522%252C%2522normal%2520-%2520transverse%2520colon%2520%28GTEx%29%2522%252C%2522normal%2520-%2520liver%2520%28GTEx%29%2522%252C%2522normal%2520-%2520lung%2520%28GTEx%29%2522%252C%2522normal%2520-%2520minor%2520salivary%2520gland%2520%28GTEx%29%2522%252C%2522normal%252 0-%2520ovary%2520%28GTEx%29%2522%252C%2522normal%2520-%2520pancreas%2520%28GTEx%29%2522%252C%2522prostate%2520adenocarcinoma%25 22%252C%2522normal%2520-%2520prostate%2520gland%2520%28GTEx%29%2522%252C%2522normal%2520-%2520skeletal%2520muscle%2520tissue%2520%28GTEx%29%2522%252C%2522normal%2 520-%2520lower%2520leg%2520skin%2520%28GTEx%29%2522%252C%2522normal%2520-%2520suprapubic%2520skin%2520%28GTEx%29%2522%252C%2522normal%2520-%2520stomach%2520%28GTEx%29%2522%252C%2522normal%2520-%2520thyroid%2520gland%2520%28GTEx%29%2522%252C%2522normal%2520-%2520urinary%2520bladder%2520%28GTEx%29%2522%252C%2522normal%2520-%2520ectocervix%2520%28GTEx%29%2522%252C%2522normal%2520-%2520endocervix%2520%28GTEx%29%2522%252C%2522normal%2520-%2520uterus%2520%28GTEx%29%2522%255D%257D&cutoff=%257B%2522value%2522%2 53A30%257D&unit=%2522TPM%2522

Shortened url: https://tinyurl.com/y9fefymf

PCAWG-Scout on demand analysis (1e)

Exclusivity analysis of non-ERG fusion donors in PCAWG-Scout

To reproduce the exclusivity analysis, we first need to produce the list of samples without *ERG* fusions. That is done by generating the list of donors with *ERG* fusions and taking them out of the list of all donors with SV (somatic structural variant) data.

To generate the list of ERG fusion donors in Prost-AdenoCa: (1) go to the report for Prost-AdenoCa (http://pcawgscout.bsc.es/entity/Study/Prost-AdenoCa); (2) click on 'SV summary' button toward the bottom of the page; (3) in the resulting SV (structural variant) table, select fusions involving *ERG* by filtering the table (click on the 'filter' button at the bottom of the table and then, in the popup window, typing '*ERG*' in the field for 'Gene 1'); (4) after the table is filtered you can find the associated donors by clicking on the 'column' button at the bottom of the table and then, in the popup window, clicking 'save list' button next to the label 'Fusion donors'. The list will open in a pop-up window. Open the report in the main window using the button with a link-out icon on the popup window header bar. You can rename the list using the 'Edit' button on the sidebar of the report. Make it a favorite by clicking on the star icon at the top so it can be used for later analyses. To find the complement of the list: (1) go back to the report for Prost-AdenoCa (<u>http://pcawgscout.bsc.es/entity/Study/Prost-AdenoCa</u>) and click on the link for 'SV donors'; (2) from the sidebar, select the button 'Compare'; (3) Since the previous list was made a favorite, it now appears in the popup window, where you click the button 'Remove' to generate the complement donor list.

As before you can rename the list and save it as a favorite. For your convenience the list is accessible as **Prost-AdenoCa ERG non-fusion donors**

(<u>http://pcawgscout.bsc.es/entity_list/Sample/Prost-AdenoCa ERG non-fusion donors</u>). You can now click on the 'Characteristic alterations' button at the bottom of the page to perform the exclusivity analysis. Doing so will start the on-demand analysis, which should complete in less than a minute. The result is a list of gene alterations that are enriched in the non-fusion donors, with associated statistical significance, as shown in **Figure 1d**

(http://pcawgscout.bsc.es/entity_list_action/Sample/characteristic_alterations/Prost-AdenoCa ERG non-fusion donors)

Annotation and 3D clustering of SPOP mutations in PCAWG-Scout

To reproduce the image with the SPOP/PTEN structure, search for SPOP in the search box at the top of the page and select **the first protein isoform**

(http://pcawgscout.bsc.es/entity/Protein:Ensembl Protein

ID/ENSP00000240327?organism=Hsa/feb2014) from the report sidebar. To show the protein report in the JMOL viewer, click the 'JMOL' tab and select the PDB 4o1v. To display the mutation density gradient overlaid, click the 'PCAWG' tab, then the 'Highlight' button, and go back to the JMOL' tab to show the gradient with respect to all PCAWG donors.

To focus on prostate samples, first click the 'PCAWG' tab, then click on 'filter', type 'Prost-AdenoCa' in the field 'histology_abbreviation', and click 'submit' to filter down to only those samples. Click 'Highlight' and then click on the 'JMOL' tab to view *SPOP* mutations only from prostate samples.

You can use the 'Sequence' tab to visualize the *SPOP* mutation clustering on a linear depiction of protein sequences (not included in figure). For further confirmation click the 'Protein feature incidence' to see a binomial distribution analysis that detects if a region is both significantly mutated and annotated as 'Important for binding substrate proteins'. The mutation 'F -> A' in residue 133 (the most frequent mutation in the cohort) is annotated as 'Strongly reduced affinity for substrate protein'.

Supplementary Note 3. Recreating Figure 5.

1- Select the CNS-tumors from the 'Meta-cohorts' tab.

PCAWG									
PanCancer Analysis of Whole Genomes									
Welcome to the PCAWG-Scout. This portal will help explore the data produced by the PCAWG project. It's organized as a collection of reports centered around entities and lists of entities: donors, cohorts, genes, etc. These reports are computed on-demand and cached, and some even take parameters. Please consult the help page for more information.									
You may also find tips on some parts of the site with additional information. These tips are green dots with a question mark inside appear at the top right part of a page section, and are revealed when hovering over that section; look, there is one down there! You can start by checking out some of the cohorts bellow or by searching for a gene of interest in the search area above.									
All abbreviations Meta cohor	s Histology tier:	Histology tier2 Histology tier	3 Histolog	gy tier4 Tumour histologica	l type				
Adenocarcinoma_tumors	NH	Breast_tumors	NH	CNS_tumors	NH	Carcinoma_tumors	NH		
Digestive_tract_tumors	NH	Female_reproductive_system_tum	NH	Glioma_tumors	NH	Hematopoietic_tumors	NH		
Kidney_tumors	NH	Lung_tumors	NH	Lymph_tumors	NH	Myeloid_tumors	NH		
Sarcoma_tumors	NH	Squamous_tumors	NH						

- 2- On the entity page for the cohort click, on the button for the 'Gene report' action and type '*IDH1*'.
- 3- In the gene report, click on the link that points to the 22 samples affected. Doing so will open the sample list report for the samples.

C	Gene	e list report	Binomial signif	icance	Gene dan	nage bias	Sample gene in	cidence Gene report	PPI interfaces	Mutation	signatures Mu	tation signatures r	ature2013
		SV s	ummary										
met	ta=C	CNS_tum	ors										
Gene	report	t											
Report	alteratio	ons over a given ge	ne in the study me	ta=CNS_tum	nors								
Sample	IDH1 o	overview (287 tota	l samples)										
All sam 25	ples												
Sample 22 Sample 22	s affecte s damag	ed ged											
Sample 22 Sample 22 Samp	s affecte s damag ble	ed ged Ensembl Gene ID	overlapping	affected	broken	splicing	mutated_isoform	damaged_mutated_isoform	TSS promoter (1000 bp)	missing	compound_mutatio	n homozygous	CNV status
Sample 22 Sample 22 Sample DO1	s affecte s damag ole 1046	ed ged Ensembl Gene IDH1	overlapping true	affected true	broken true	splicing false	mutated_isoform true	damaged_mutated_isoform true	TSS promoter (1000 bp) false	missing false	compound_mutation	homozygous false	CNV status normal
Sample 22 Sample 22 Sample 22 DO1	s affecte s damag ole 1046 2352	Ensembl Gene IDH1 IDH1	overlapping true true	affected true true	broken true true	splicing false false	mutated_isoform true true	damaged_mutated_isoform true true	TSS promoter (1000 bp) false false	missing false false	compound_mutation false false	homozygous false false	CNV status normal
Sample 22 Sample 22 Sample 22 Sample 22 DO11 DO11	s affecte s damag ole 1046 2352 3575	Ensembl Gene DH1 IDH1 IDH1 IDH1	overlapping true true true	affected true true true	broken true true true	splicing false false false	mutated_isoform true true true	damaged_mutated_isoform true true true	TSS promoter (1000 bp) false false false	missing false false false	compound_mutation false false false	homozygous false false false	CNV status normal normal
Sample 22 Sample 22 Sample 22 DO1 DO1 DO1 DO1	s affecte s damag ole 1046 2352 3575 3971	Ensembl Gene IDH1 IDH1 IDH1 IDH1 IDH1	coverlapping true true true true	affected true true true true	broken true true true true	splicing false false false false	mutated_isoform true true true true	damaged_mutated_isoform true true true true	TSS promoter (1000 bp) false false false false	missing false false false false false	compound_mutation false false false false	homozygous false false false false false	CNV status normal normal normal

4- Click on the 'Differential expression' button and then submit the differential expression analysis. The first table shows all the genes and their corresponding statistics for differential expression between the 22 samples affected by *IDH1* mutations and the rest

AWGScout 🖈	Entities • Lists • Maps •			Workflows	 guest 	Help
CFH	-1.6569	-5.0729	-6.5507e-06	-0.00010188	-	
FUCA2				3	¢	
GCLC						
NFYA	Ensembl Gene ID	save map save list				
STPG1						
NIPAL3	ratio	save map				
LAS1L	too loo a					
ENPP4	t.values	save map				
SEMA3F	pyaluer					
CFTR	pleades	save map				
ANKIB1	adjusted by values					
CYP51A1		save map				
KRIT1						
RAD52						
« » all						
	Up-regu	lated genes			_	
Ensembl Gene ID	ratio	t.values	p.values	adjusted.p.values		

of the samples in the cohort. Negative values represent repression.

- 5- Click on the button 'column' on the table footer and then on the button 'Save map' next to the header 't.values'. After a short while, a window will open with a table listing the genes and their corresponding t-values. That is the map for use in coloring the genes. After a few seconds, you will be able to open the map in a different window by clicking on the top right button with a blue link-out icon. From the page accessed, you can rename the map (e.g., as 't-values for *IDH1* mutants'). After renaming the map a new page will open. Clicking on the star on the top-bar will turn it yellow. That change of color indicates that the map is placed in your top-bar drawer for favorite maps, under the sub-header 'Gene'.
- 6- Go back to the cohort entity report from step 2. Click on the link under the header 'Genotyped donors'. That will open the sample list report for all cohort donors and feature a table with the clinical information. From that table, use the 'column' button to extract the map between samples and the feature 'donor_survival_time'. As before, open the map into a separate window, rename it, and make sure it is a favorite by clicking on the top-bar star.
- 7- Go back to the cohort entity report from step 2. Click on the button 'Gene damage bias' to open a driver prediction report, which compares the predicted damage of the mutations found in a given protein with the predicted damage of all possible mutations. That highlights proteins harboring mutations significantly more damaging than expected at random. From the table, follow the usual procedure to extract the map between genes and p-values; except this time, before renaming the list, convert p-values to 'P-value'

scores', which are better suited to be used in plots.

							C.
Ensembl G	iene ID Map)					
Map gene-p.v	alue in gene da	mage blas li	n meta=Cl	NS tumors	5		
Rename	Pvalue score	QQ-plot	Name	Ranks	Rank products	Compare plot	
	Map gene-p.value	e in gene damag	e bias in met	a=CNS tumo	rs		
Ensembl Gene	D		p.va	lue			
CCDC132			0.26	246			
ABCB5			0.84	609			
CREBBP			5.58	9e-05			
CACNA1G			0.28	514			
CRAMP1L			0.33	208			
PKD1			0.94	296			
TENIM1			0.15	094			

- 8- We have gathered all of the information needed to annotate the network. The network can be found on the cohort report from step 2. Open the 'Aesthetics' menu to map visual characteristics to different values.
- 9- Select 'nodes', the 'color' aesthetic, and select the map from step 5. That will color a gradient from the most repressed in purple to the most over-expressed in gold, going through gray, which represents no dysregulation.
- 10- Perform a similar procedure with the aesthetic 'borderWidth' and the survival time map from step 6. That will highlight donors with longer survival times.
- 11- Repeat the procedure with the aesthetic 'size' and the map from step 7 to highlight genes that are significantly damaged.
- 12- If you click on the edges of the graph, a window will open describing the nature of the relationship between the sample and the gene, e.g. whether the mutation is a validated driver or not. To visualize that information, select the element 'edges' and the aesthetic 'color', and assign to it the field 'Validated'. On the map select the option 'Use textarea'.

In the text area, put the line 'validated green' and, below that, the line 'predicted orange'.								
×	Edges	Entities	Aesthetics	Help				

	Acstrictics	Theip	
Map entity aesthetic			
elem (default: nodes)			
edges			•
aesthetic (default: opacity)			
color			•
field (default: id)			
Validated			
map			
Use textarea			•
map_tsv			
validated green			
predicted ordinge			
Submit Query			

13- You can use the contextual menu to help organize they layout of the nodes, for instance using the 'surround with first neighbors' to organize the samples around the main driver genes.

Supplementary Figure 1. PCAWG-scout uses PanDrugs for prediction of recommended therapies.

PanDrugs is a web-based tool (http://www.pandrugs.org) to guide the selection of therapies from the results of genome-wide studies in cancers. It allows identification of actionable molecular alterations and prioritization of drugs by calculating gene-drug scores (GScore and DScore, respectively). Those scores take into account: i) the relevance in cancer of the affected gene and the specific variant type; ii) the target pathway context; iii) the drug approval status (FDA, clinical trial, or experimental small molecule inhibitor); and iv) manually curated pharmacological information retrieved from the literature. PanDrugs GScore indicates the biological relevance in cancer of the gene affected, the functional impact, and the clinical actionability of the specific mutation, integrating evidence from public resources. The DScore indicates the suitability of the drug according to the genes and their susceptibility to be targeted, reflecting the strength of evidence for the gene-drug association. That measure can be used to assist in clinical decision-making. Additionally, it incorporates manually curated information about the drug approval status and its usage in cancer therapies or clinical studies in this field.

The current version of PanDrugs integrates data from 24 sources, supporting more than 56,000 drug-target associations obtained from approximately 4800 genes and 9000 unique compounds.

- [1] Piñeiro-Yáñez E. et al. (2018) Genome Medicine. 10:41
- [2] Wagner AH. et al. (2016) Nucleic Acids Res. Jan 4;44(D1):D1036-44
- [3] Van Allen EM. et al. (2014) Nat Med. Jun;20(6):682-8
- [4] Basu A. et al. (2013) Cell. Aug 29;154(5):1151-61
- [5] Iorio F. et al. (2016) Cell. Jul 28;166(3):740-54

a) Overview of the drug assignment distribution for 2494 donors for 37 different tumor types. Each bar represents the percentage of patients with a suggested therapy for each tumor type based on simple somatic mutations detected. Different colors correspond to different approval status for the drugs as indicated in the legend. Only high impact alterations (those with a PanDrugs GScore greater or equal to 0.6) are considered. b) Example of a therapy suggestion based on evidence from an affected gene. Donor DO13132 with Glioblastoma (CNS-GBM) has an EGFR p.Gly719Cys missense mutation, which leads to carcinogenic processes of cell growth and proliferation. The mutation confers sensitivity to EGFR inhibitors such as Erlotinib, one of the proposed therapies. c) Example of a therapy suggestion against the use of a conventional therapy based on evidence from an affected gene. EGFR inhibitor Cetuximab is a standard therapy for the treatment of colorectal cancer, but KRAS mutations have shown to be a predictor of resistance to cetuximab therapy. In the DO44094 case, the missense p.Gly12Val mutation indicates resistance to Cetuximab. d) Example of an indirect therapy suggestion based on evidence from a pathway membership: Melanoma patient DO220845 has the p.Val600Glu alteration in BRAF. The alteration suggests use of BRAF inhibitors such as Vemurafenib. It also suggests MEK inhibitors, for example Trametinib, according to the downstream position of those gene in relation to BRAF in the MAPK signaling pathway.



a) Percentage of donors with treatments suggested by PanDrugs

Supplementary Figure 2. View protected data using a local Xena Hub.

The Xena platform has two components: the web-based Xena Browser and the back-end Xena Hubs. The Xena Browser empowers biologists to explore data across multiple Xena Hubs. The back-end Xena Hubs host the genomics data, are configured to be public or private, and can be installed on laptops, public servers, behind a firewall, or in the cloud. Xena Browser simultaneously connects to any number of Xena Hubs, with integration occurring in the browser, enabling data to be distributed across multiple Xena Hubs. To view the controlled-access non-coding simple mutations, download the file that contains those mutations (https://gtrepo-osdc-tcga.annailabs.com/cghub/metadata/analysisFull/86a1f73a-fabf-4abe-a597-a47d84a8c980). Only authorized users can download the protected data. The file is pre-formatted to be imported directly it into a local Xena hub on a user's laptop. Once it is loaded, the UCSC Xena Browser will connect to both the local hub and the public PCAWG hub concurrently, while still keeping the protected data private. The arrangement allows users to visualize the protected whole-genome simple mutation data alongside the open-access PCAWG data. More information about using Xena private data hubs can be found at http://xena.ucsc.edu/private-hubs/.



Supplementary Figure 3. Secured PCAWG-Scout installation to support controlled-access data.

To protect controlled access data, in particular genomic mutations, the PCAWG-Scout is configured so that controlled data is on only a siloed machine behind a firewall. Approved analyses that do not compromise the security of the data are made available by configuring a remote workflow access file on the Rbbt installation. Approved analyses can be accessed via a web-browser, javascript plotting utilities, or command-line tools. The PCAWG-Scout machine does not hold the controlled access data, and the silo is not directly accessible from outside, keeping the data secure. Any remote installation of the PCAWG-Scout can request such analyses and will relay them to the silo. The procedure enables the system to be extended by the general research community without requiring all researchers to have access to the controlled data.



Supplementary Figure 4. Circos plots for the other 7 tumors with ERG fusion for all chromosomes (left side of figures) and for chromosome 21 (right side of figures).



Donor PRAD-US::59207bcd-f127-481f-b576-8716da7d7d97 (DO36372)

Donor PRAD-US::6845076e-e940-4437-8618-81db4a447544 (DO36345)





Donor PRAD-US::87959988-bb25-47ee-afc1-9af92592bbe7 (DO36273)

Donor PRAD-US::aa252f78-ebf3-42f9-86af-bb6e57345706 (DO36359)





Donor PRAD-US::be5411dc-3caa-4b25-9b94-ce463e8e52b4 (DO36265)

Donor PRAD-US::e5e4d3b6-8e58-4dcf-8f67-a00e656a8e27 (DO36335)





Donor PRAD-US::f6381367-142c-45d0-92b3-c1727d1813ce (DO36283)

Supplementary Figure 5. Prostate adenocarcinoma ERG structural variants

A UCSC Xena Visual Spreadsheet showing that 106 of the 195 prostate adenocarcinoma tumors showed the *ERG* fusion that joins an external piece of DNA from the 5' direction, replacing the 5' end of *ERG* with another piece of DNA (promoter fusion). The 195 prostate tumors were from four different projects: PRAD-CA, PRAD-US, PRAD-UK and EOPC-DE. Eighty-nine of the 195 prostate tumors (46%) did not show that type of fusion . Structural variants in *ERG* were detected in three samples, but they were not the 5' joining type (DO52498, DO50430, and DO51087). For structural variants (column B), the grey-colored lines represent the external DNA segments found to be fused to *ERG*, and the short black tick marks indicate the breakpoints in *ERG*. The breakpoints are clustered at the beginning of the coding regions. A live view can be accessed here:

https://xenabrowser.net/heatmap/?bookmark=4d07f983ccd528022a931703aa1abe47.



Supplementary Table 1. Code availability.

Resource	Open source code availability
ICGC Data Portal	https://github.com/icgc-dcc/dcc-portal
PCAWG-Scout	http://mikisvaz.github.io/rbbt/; https://github.com/Rbbt- Workflows; https://github.com/Rbbt-Apps/PCAWGScout
UCSC Xena Browser	https://github.com/ucscXena/ucsc-xena-client
Expression Atlas	https://github.com/gxa/atlas
Chromothripsis Explorer	https://github.com/parklab/ShatterSeek

Supplementary Table 2. Embeddable javascript modules.

Javascript Module	Utility	Open source code availability
OncoGrid	Generate OncoGrids and related tracks	https://github.com/oncojs/oncogrid
Xena Visual Spreadsheet	Generate visual spreadsheet	https://github.com/ucscXena/ucsc-xena-client
Kaplan-Meier	Kaplan-meier estimator and log- rank test	https://github.com/ucscXena/kaplan-meier
static-interval-tree	Fast overlapping interval queries in javascript	https://github.com/ucscXena/static-interval- tree
Expression Atlas Widget: Heatmap & Anatomogram	View tissue- specific results on a heatmap and human figure	https://github.com/gxa/atlas-heatmap

Supplementary Table 3. List of PCAWG primary results supported by online visualization resources.

Table of primary results generated by PCAWG analysis working groups available for visualization by UCSC Xena, Expression Atlas, PCAWG-Scout, and Chromothripsis Explorer. Each primary result is referenced by corresponding synapse IDs. Synapse folder ID is the identifier for the synapse landing page for each type of primary results. The landing page typically includes a summary written by the analysis working group to briefly describe the bioinformatics methods used and a list of results generated. Because there are often multiple versions of the same results files (such as fpkm vs fpkm-uq gene expression estimations, or simple mutations from all specimens or aggregated by donors), synapse identifiers in the remaining columns point to the actual data file ingested by each online resource. The data snapshot was taken as of Feb 10, 2017.

Data	Synapse page ID	UCSC Xena	Expression Atlas	PCAWG- Scout	Chromo- thripsis Explorer
Consensus SNVs and indels	<u>syn7118450</u>	<u>syn7364923</u> syn7364924		<u>syn7364923</u>	<u>syn7357330</u>
Consensus SVs	<u>syn5964535</u>	<u>syn7596712</u>		<u>syn7596712</u>	<u>syn7596712</u>
Consensus copy number	<u>syn8042880</u>	<u>syn8042988</u>		<u>syn8042992</u>	<u>syn8042880</u>
Gene expression	<u>syn3104297</u>	<u>syn5553991</u>	<u>syn5553983</u> <u>syn5553985</u>	<u>syn5553991</u>	
GTEx gene expression derived using the PCAWG RNA-seq SOP	<u>syn8105922</u>		<u>syn8105922</u>		
RNAseq gene fusion	<u>syn7221157</u>	<u>syn7221157</u>			

RNAseq alternative promoter usage	<u>syn3354819</u>	<u>syn10332949</u>			
small RNA-Seq (miRNA) analyses	<u>syn5842981</u>	<u>syn5878064</u> <u>syn5878067</u>			
Patient-centric driver catalogue	<u>syn7250534</u>	<u>syn11050201</u>		<u>syn7328242</u>	
Integrated driver calls	<u>syn7359546</u>			<u>syn8035740</u>	
APOBEC mutagenesis analysis	<u>syn7437205</u>	<u>syn7511424</u>			
Tumour subtype and histology information	<u>syn4974831</u>	<u>syn10389164</u>	<u>syn7253569</u>	<u>syn7253569</u>	<u>syn4974831</u> (version 9)
Donor clinical data	<u>syn4974831</u>	<u>syn10389158</u>		<u>syn7772065</u>	<u>syn4974831</u> (version 9)
Consensus purity and ploidy					<u>syn8272483</u>