

**Integration of copy number and transcriptomics provides risk stratification in prostate cancer:
a discovery and validation cohort study**

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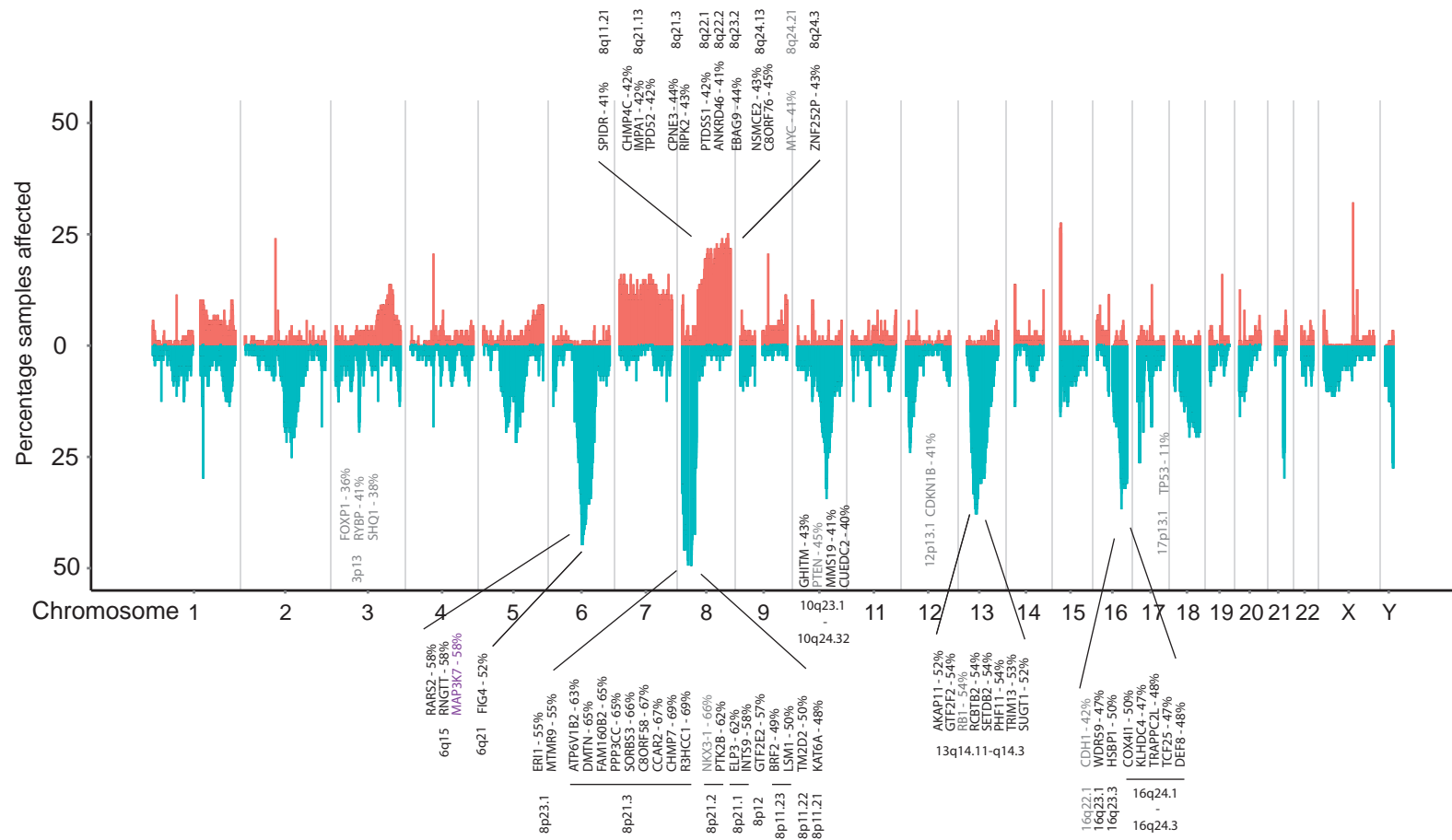
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Supplementary Figure 1-18

Supplementary Methods

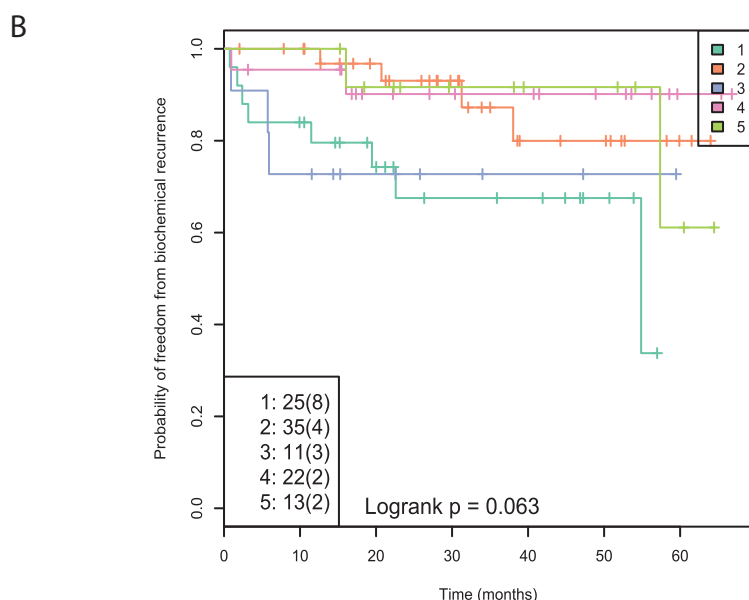
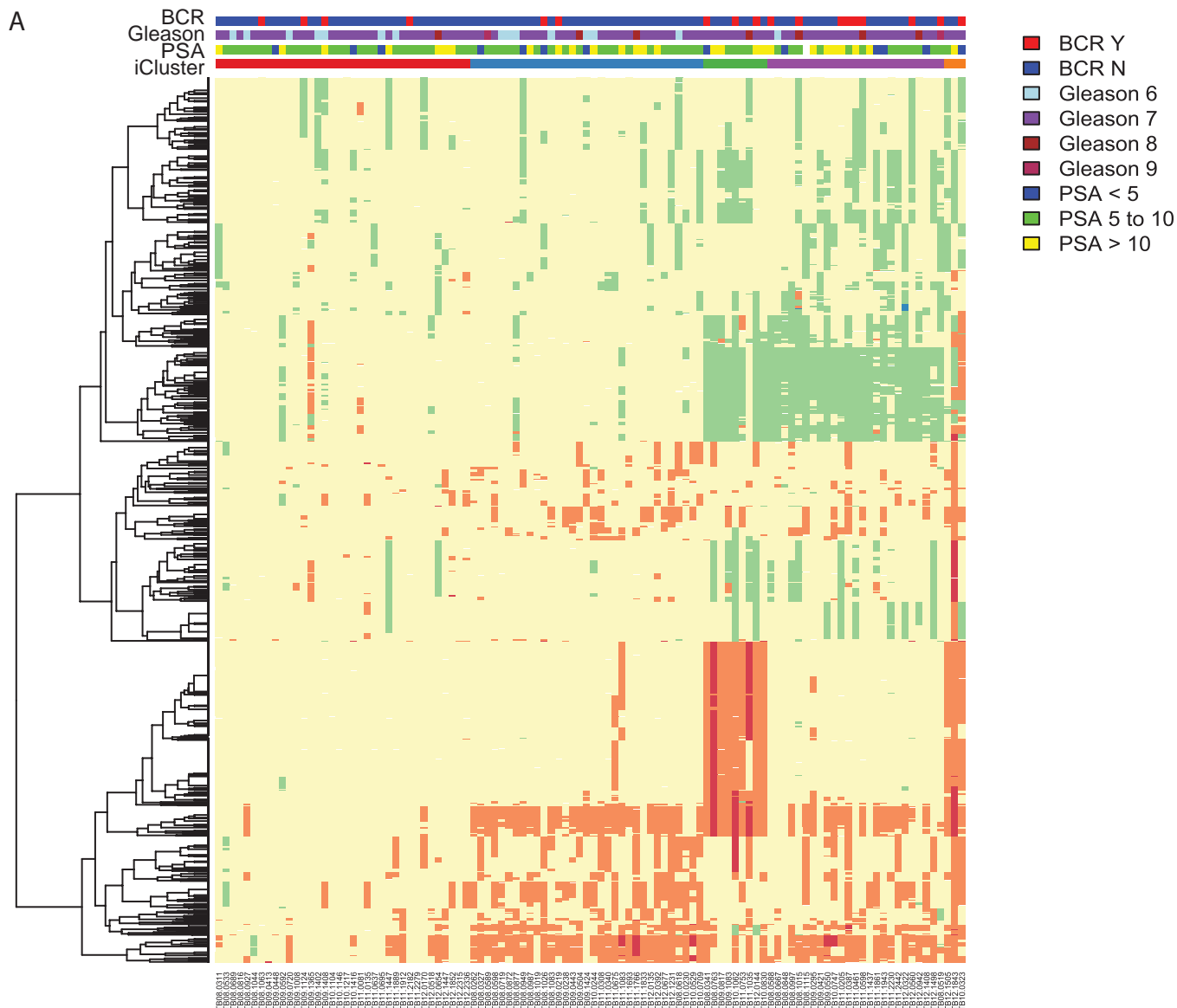
Supplementary Case Example

Supplementary Tables 1,3-12 (Supplementary Table 2 is a separate file)



Supplementary Figure 1. A genome-wide copy number profile of the Stockholm validation cohort.

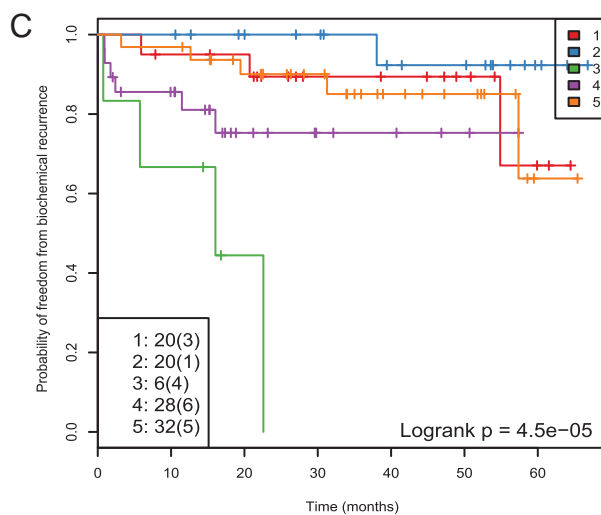
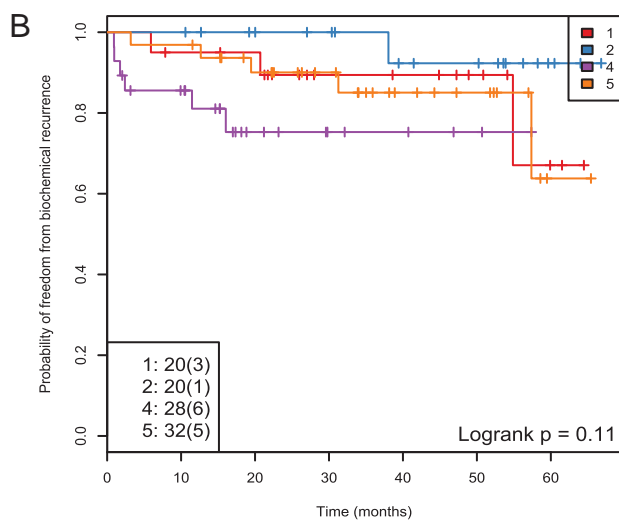
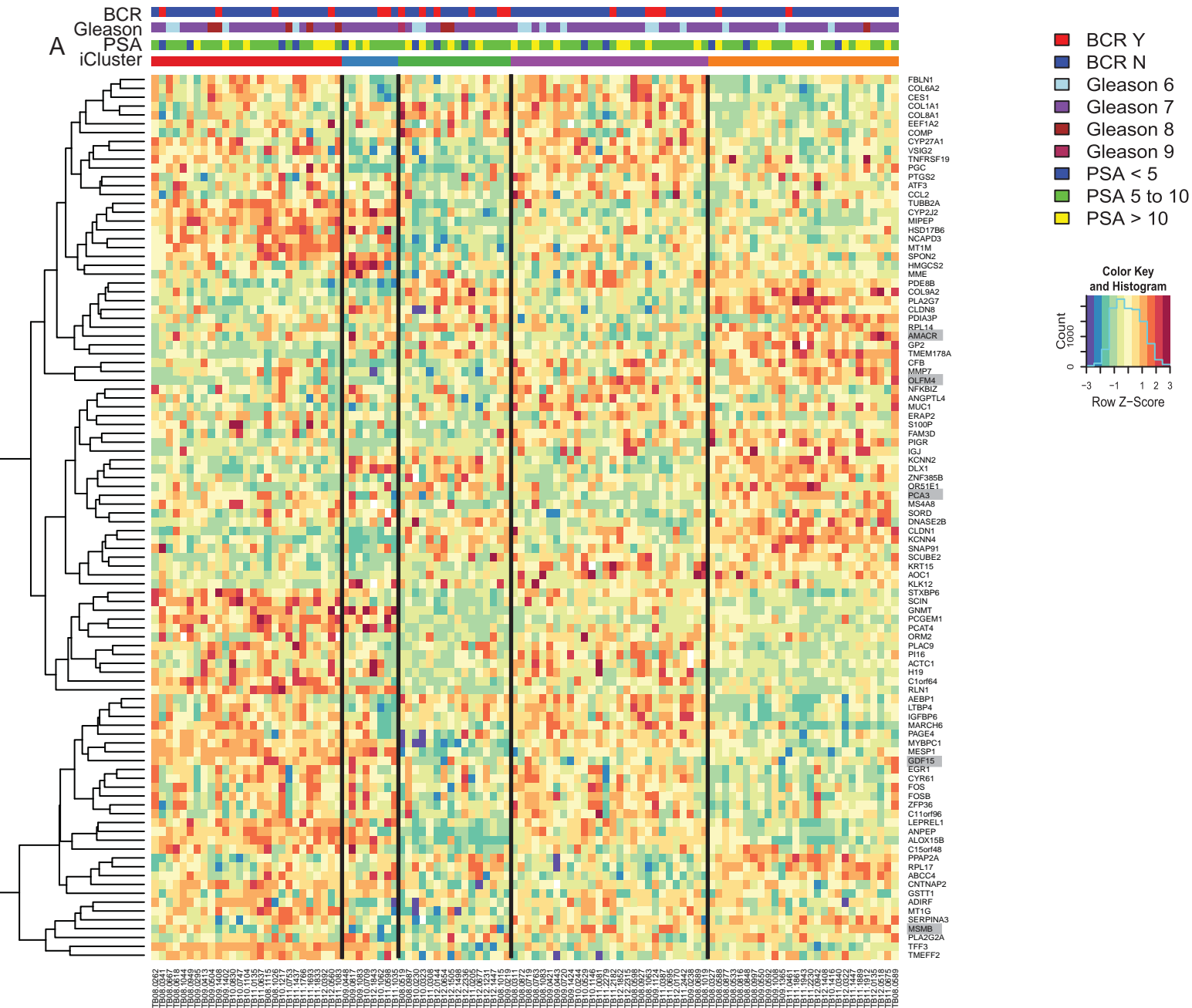
The percentage of samples containing copy number aberrations (CNA) at each locus is shown by gain/loss (red/blue); left hand y-axis. Established prostate cancer risk genes commonly disrupted by CNAs (from Williams et al. (2014) meta-analysis) are indicated in grey (gene name and frequency altered in this cohort are shown, see also Suppl. Table 3); only those affected in >10% samples are annotated. Novel CN changes identified in this cohort (>10% samples) also relevant in our 100-gene set are indicated in black type. The same general patterns of gains/losses identified in the discovery cohort are seen in this dataset, which was assayed on lower-density Affymetrix SNP6 arrays and analysed using ASCAT (van Loo et al., 2010) (see Methods), with no filtering of results. Chromosome ends are delineated by grey, vertical stripes. No gene expression data are available for this cohort, as no matched normal tissues were available.



Supplementary Figure 2. Clustering based on CN alone.

A. Genome-wide profiles identify five groups of patients with distinct CN profiles (see Methods). Gains are shown in pink (heterozygous; +1) and red (homozygous; +2), losses are indicated in green (heterozygous; -1) and blue (homozygous; -2). The distribution of clinical characteristics is shown. PSA = pre-operative PSA (ng/ml). Gleason = Prostatectomy Gleason Score. BCR = biochemical relapse.

B. Kaplan-Meier biochemical relapse curves using clusters generated above. These CN profiles are not prognostic (logrank $p=0.063$). Copy number data were analysed in OncoSNP (Yau et al., 2010); all rank calls were used to identify patient clusters to increase signal and ensure even small CNAs were captured (ranks3/4/5) and not just larger copy number alterations (ranks1/2).

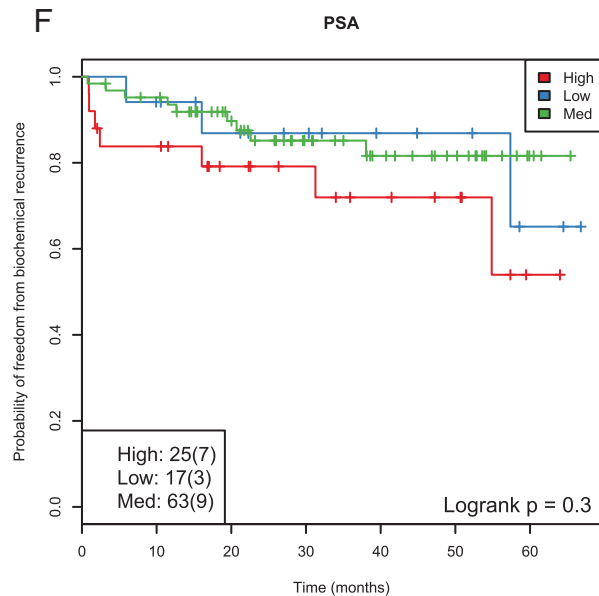
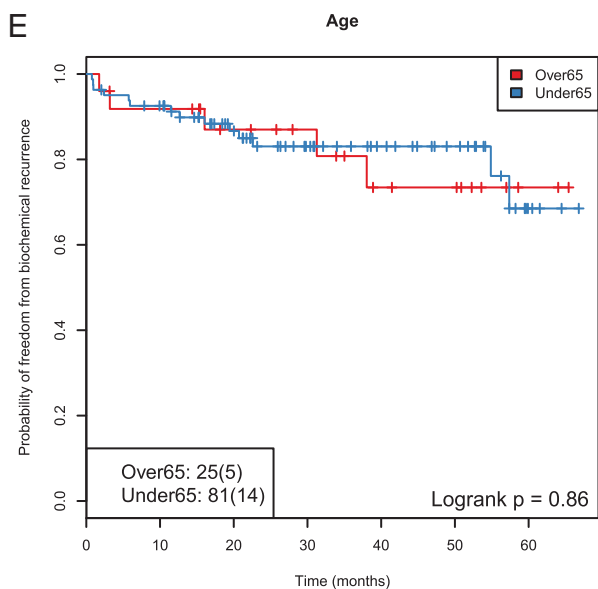
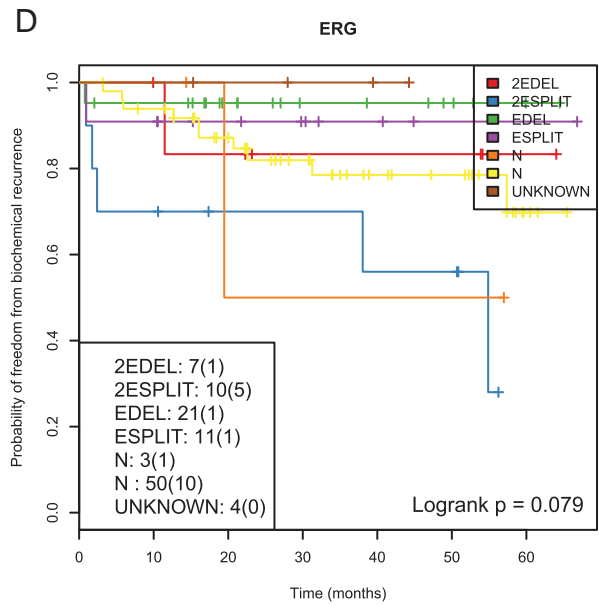
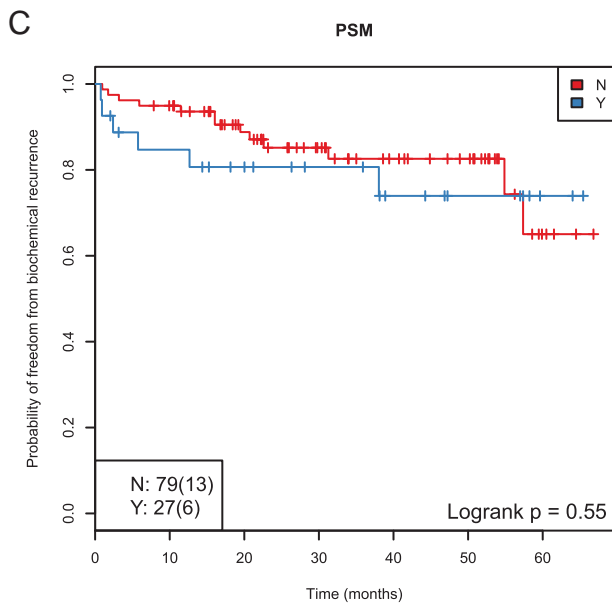
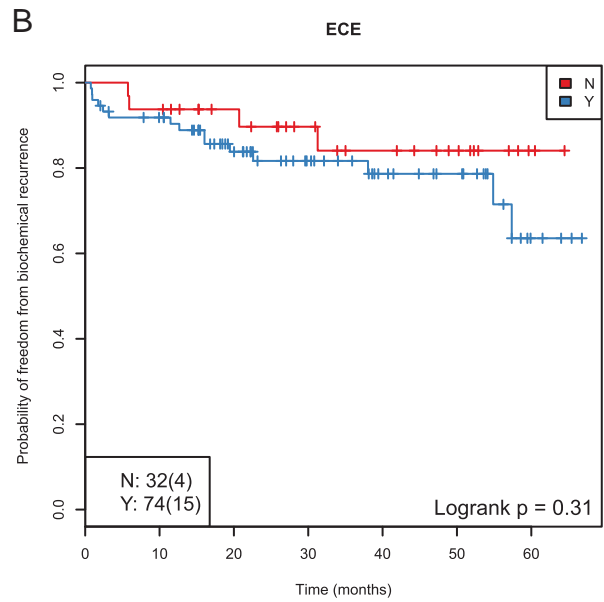
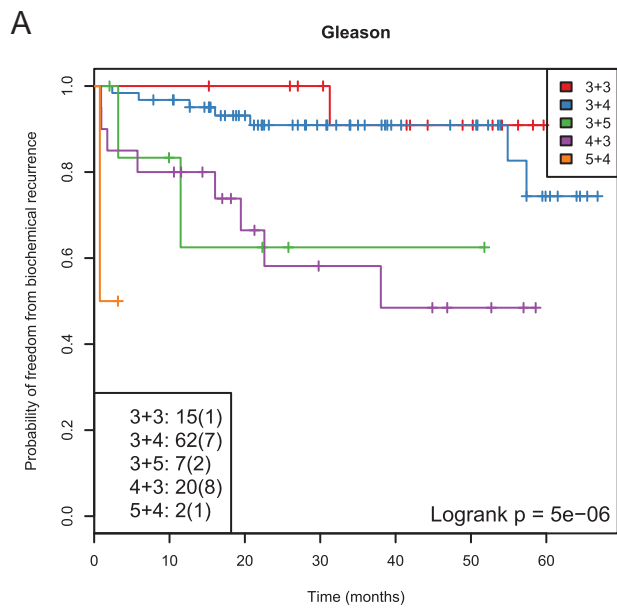


Supplementary Figure 3. Clustering based on Gene Expression (GEX) alone.

A. k-means clustering of the top 100 most variable inter-tumour probes (Gorlov et al., 2014) identified 5 clusters based on Illumina HT12 mRNA transcript data from primary prostate cancer tissue samples (see Methods). The distribution of clinical characteristics is shown. Relative expression of the 100 most variable genes (between tumours) are shown. This approach identified known prostate cancer risk genes AMACR, PCA3, GDF15, OLFM4 and MSMB, highlighted in grey. PSA = pre-operative PSA (ng/ml). Gleason = Prostatectomy Gleason Score. BCR = biochemical relapse.

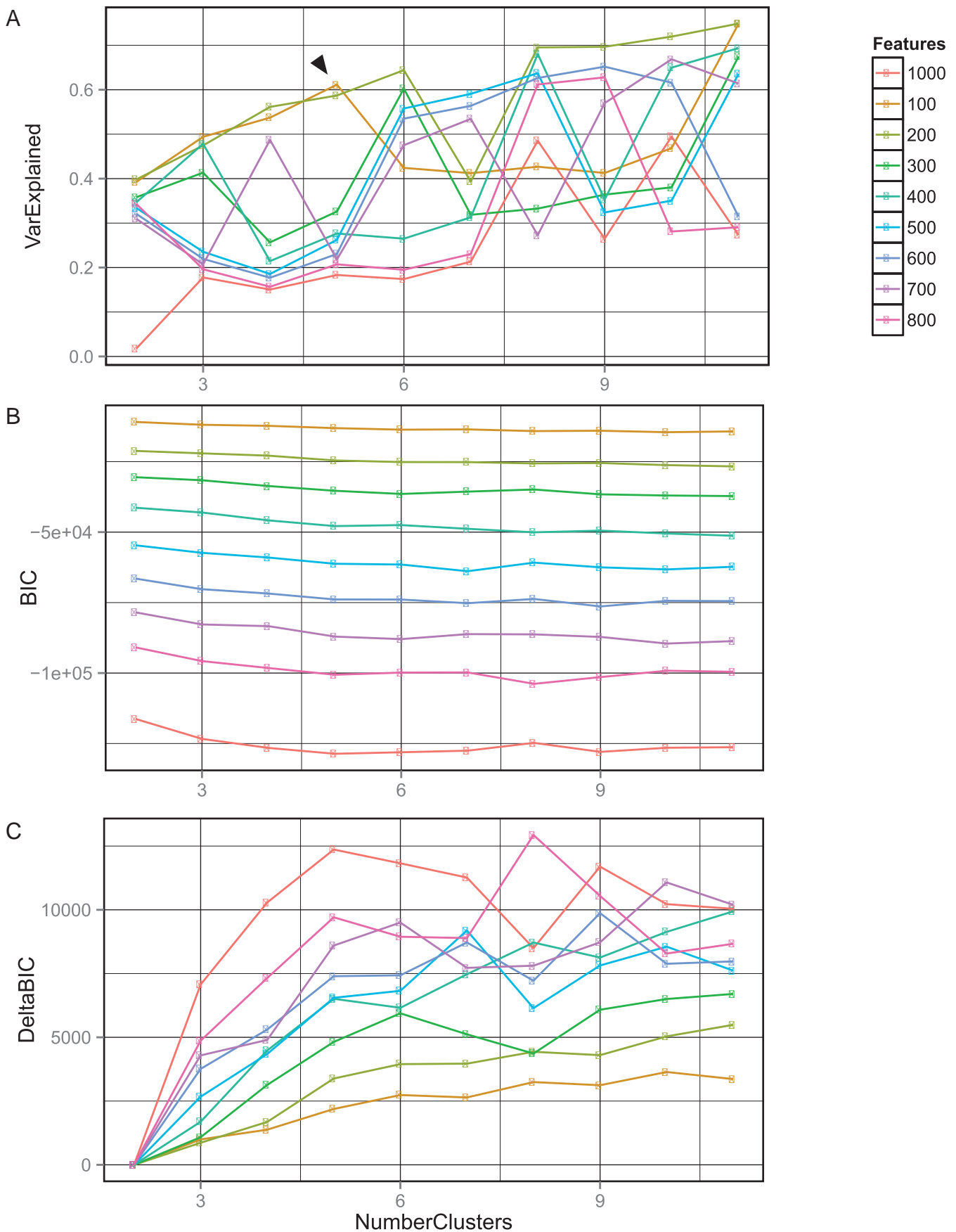
B. Kaplan-Meier biochemical relapse curves using clusters generated above. These GEX profiles alone are not prognostic (logrank $p=0.11$).

C. Transcript-only profiling was only statistically significant when a small group of outliers was included in the analysis.



Supplementary Figure 4. Relapse-free survival curves based on clinical characteristics.

- A.** Kaplan-Meier biochemical relapse curves split by Gleason score in the Discovery Cohort.
B. Extracapsular extension (ECE).
C. Positive surgical margin (PSM).
D. TMPRSS2-ERG gene fusion status (ERG).
E. Age.
F. Pre-operative PSA (low < 3 ng/ml; medium 3-10 ng/ml; high > 10 ng/ml).

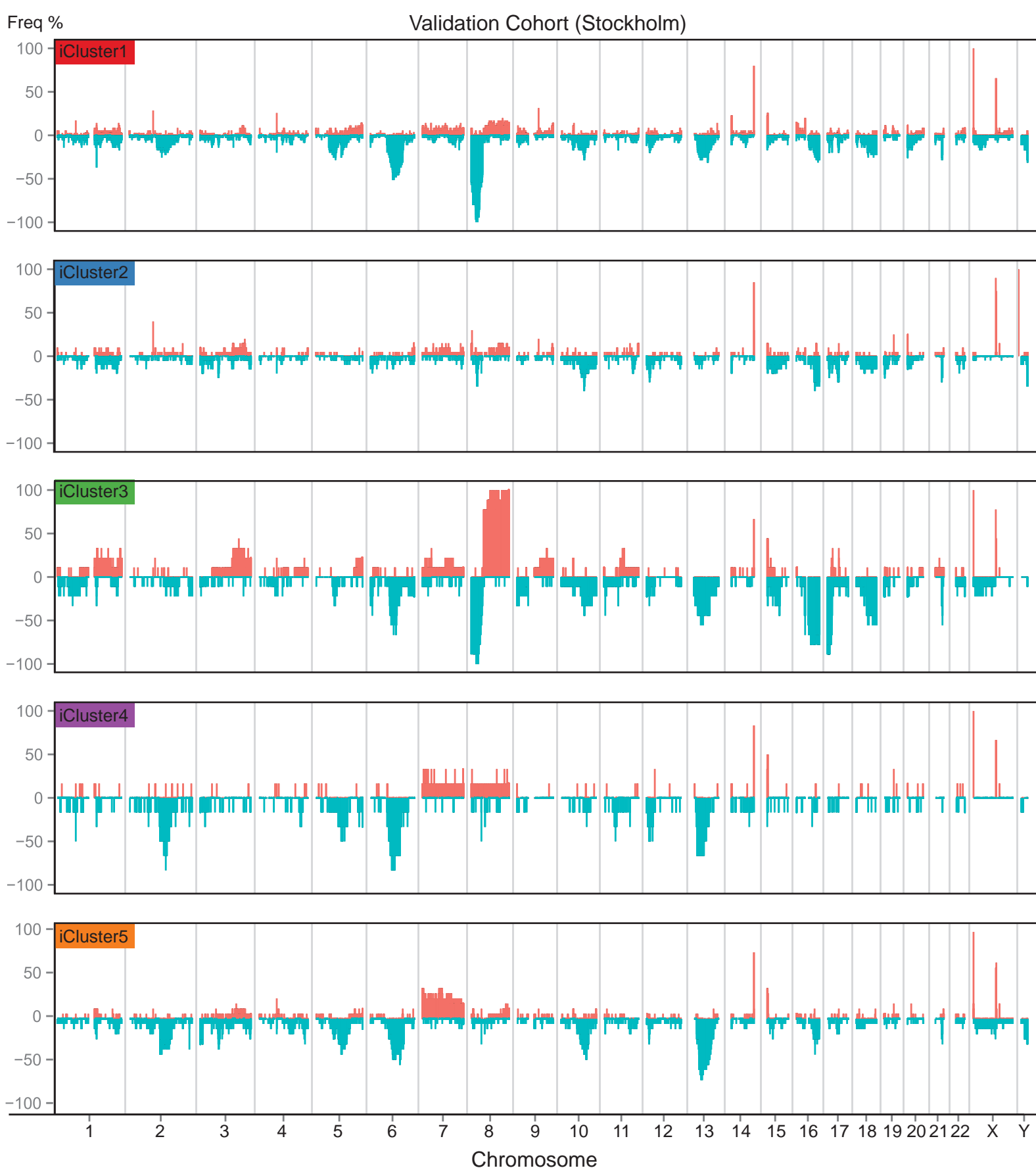


Supplementary Figure 5. Selection of integrated clusters in the Cambridge Discovery Cohort.

A. Variance plotted against number of clusters for 10 increasing size feature sets. The smallest number of features ($n=100$) consistently described the most variance for lower numbers of clusters. The inflection point at $n=5$ clusters was chosen as describing the most variance with largest numbers of men per cluster ($k+1$; $iCluster+ k=4$; see Methods & Supplementary Methods).

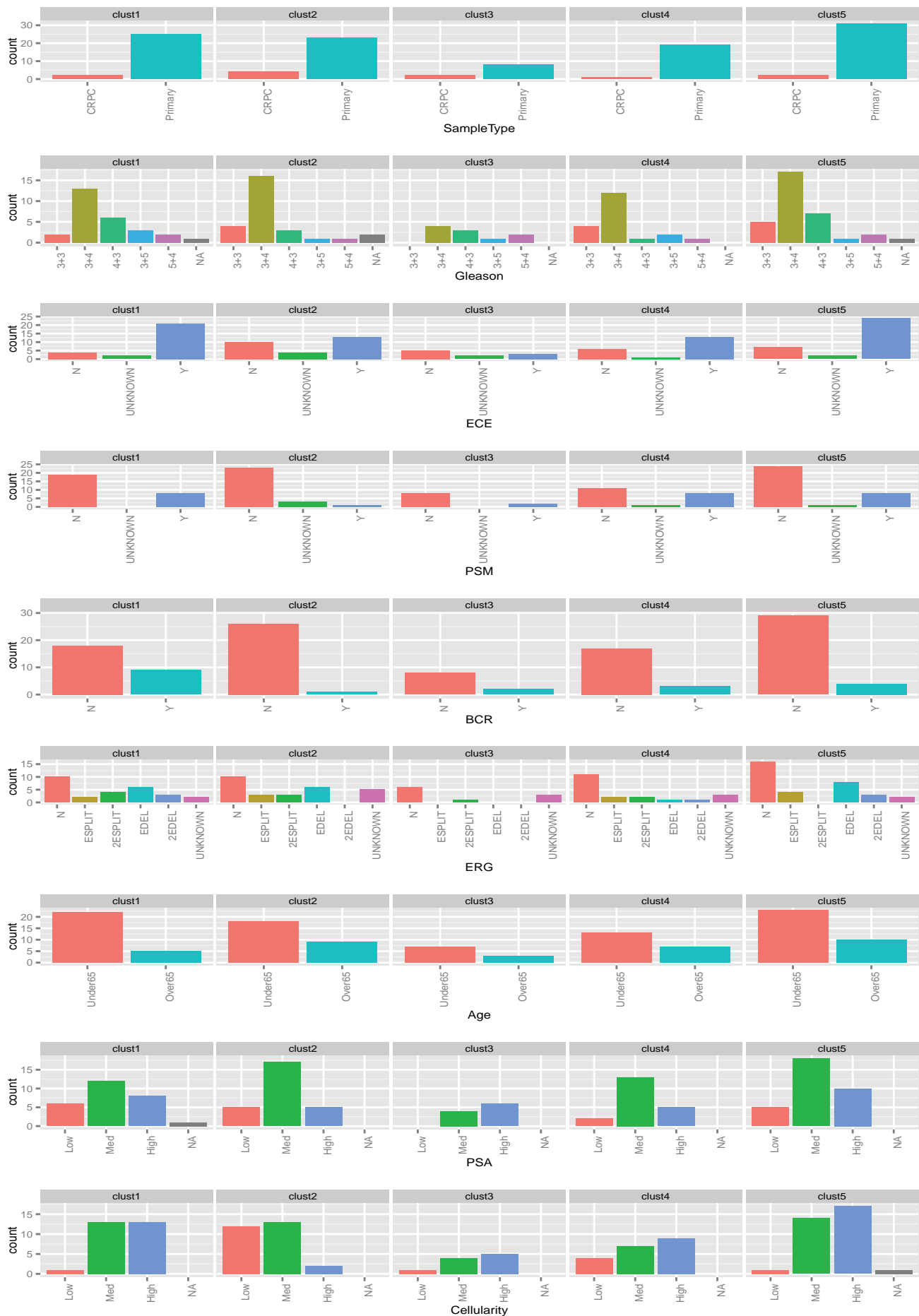
B. Optimal number of features was selected to minimize Bayesian information criterion (BIC) for a given k .

C. One over BIC was plotted (Delta BIC) to better display incremental differences in BIC. Clear inflection is seen in this figure at 5 clusters.

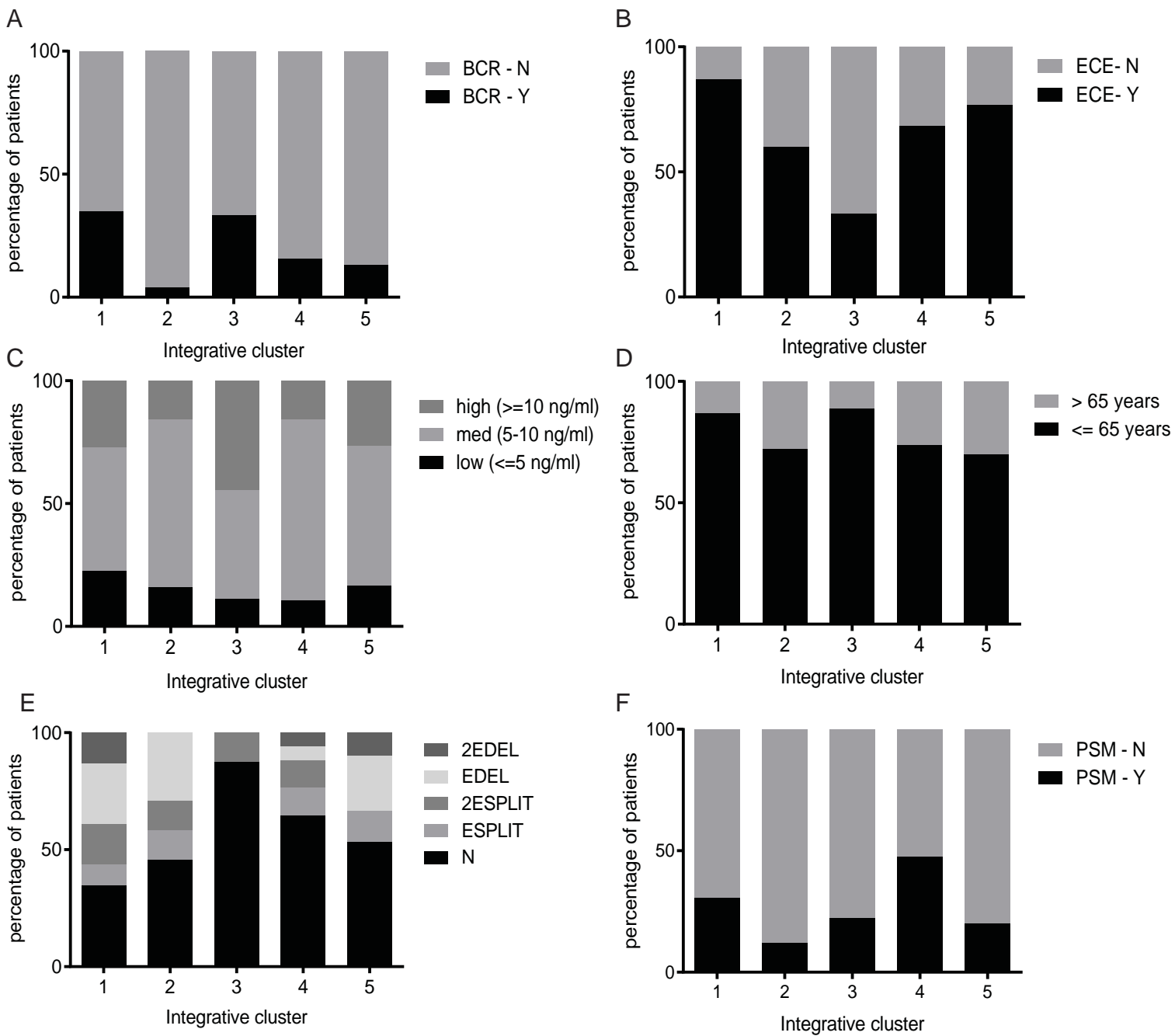


Supplementary Figure 6. Validation of integrative clusters (Stockholm Validation Cohort)

Genome-wide frequencies of somatic copy number alterations (CNAs) presented as a percentage of samples (left y-axis) in each Integrated Cluster (iCluster). Regions of copy number gain are indicated in red and regions of loss in blue. Subgroups were identified by integrated hierarchical clustering (as described in methods) of the discovery cohort (Figure 2; $n=125$; Illumina Omni2.5 array). For the validation cohort ($n=99$; Affymetrix SNP6 array) men were allocated to these same clusters based on the 100 CN and GEx features analysed using PAMr (Tibshirani et al., 2003; see Methods). Integrative clusters 1-5 correspond with survival analysis curves in Figure 3B.

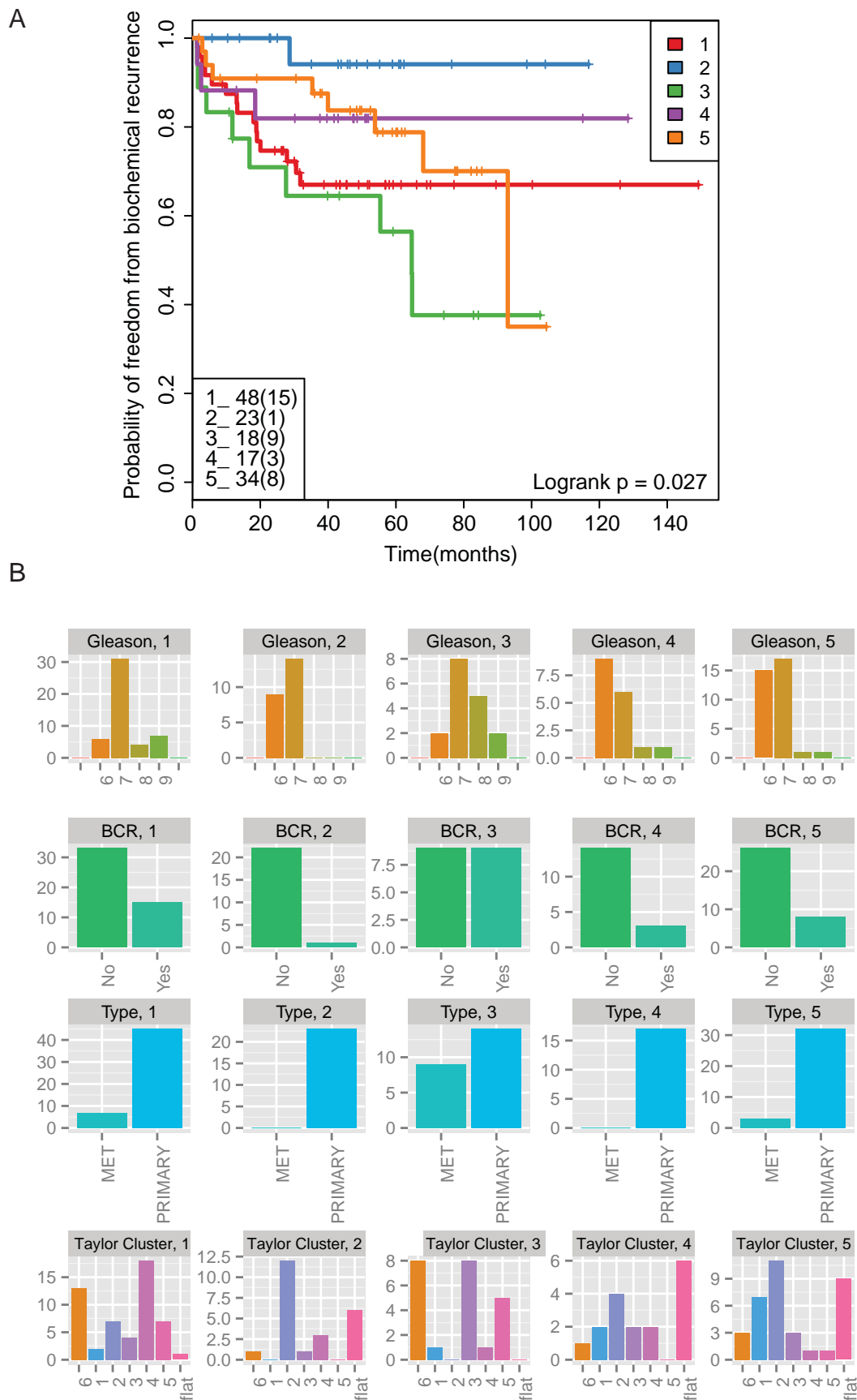


Supplementary Figure 7. Clinical Features of the integrative clusters (Cambridge Discovery Cohort). Distribution of samples between each sub-group is shown. CRPC samples are also included in the distribution (though not used in defining the clusters). Additional clinical information includes the presence of extra-capsular extension (ECE), positive surgical margin (PSM), biochemical relapse (BCR), TMPRSS2:ERG fusion status (ERG), age, PSA (low = <3ng/ml, med = 3-10ng/ml, high = >10ng/ml) and cellularity of the sample analysed.

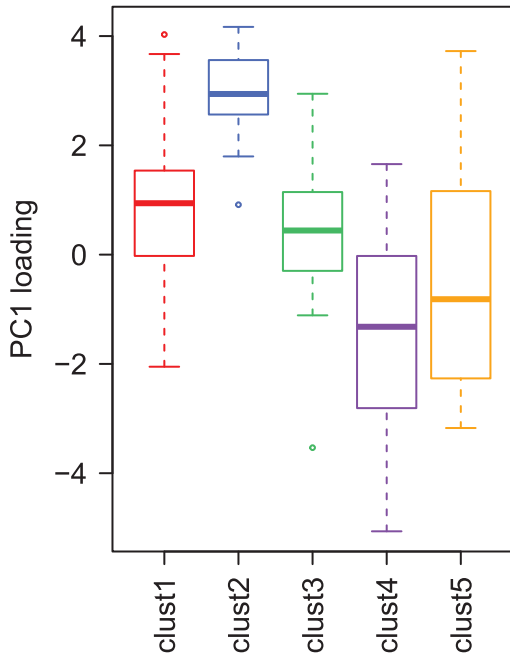
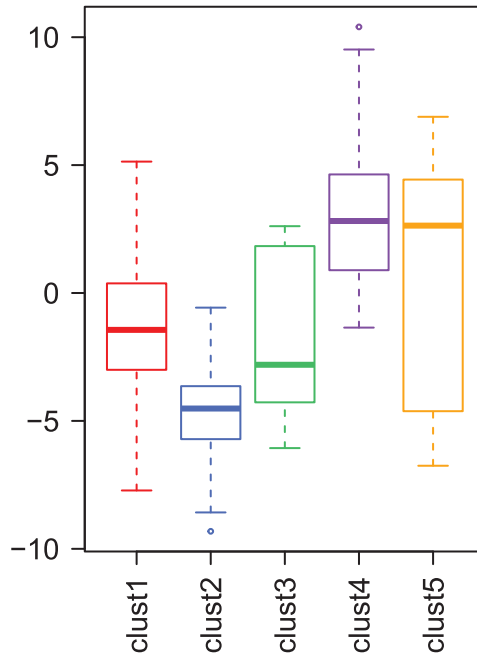
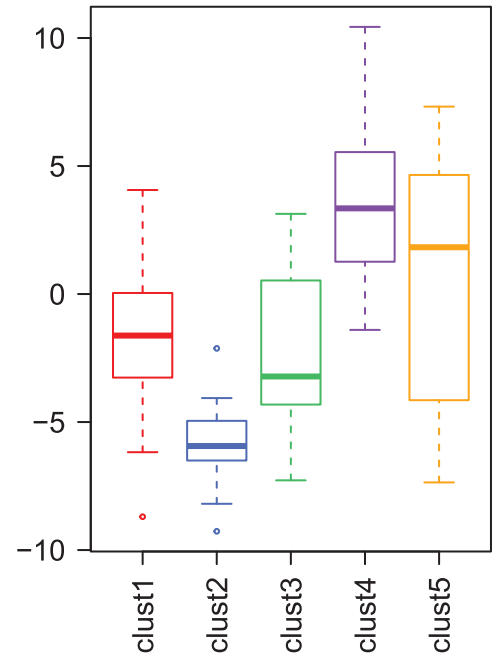
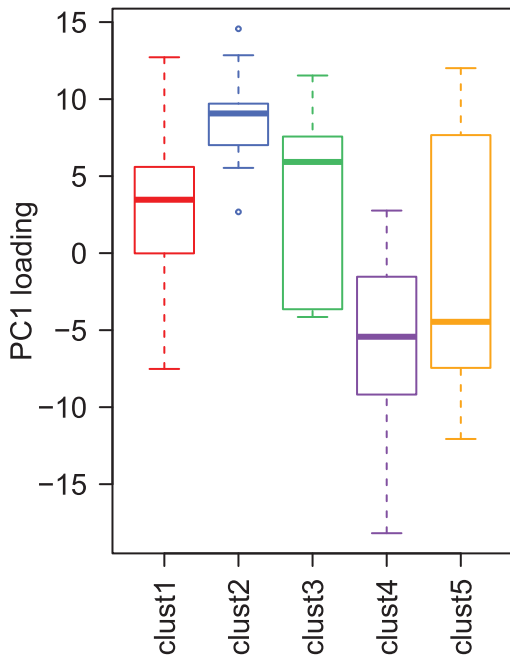
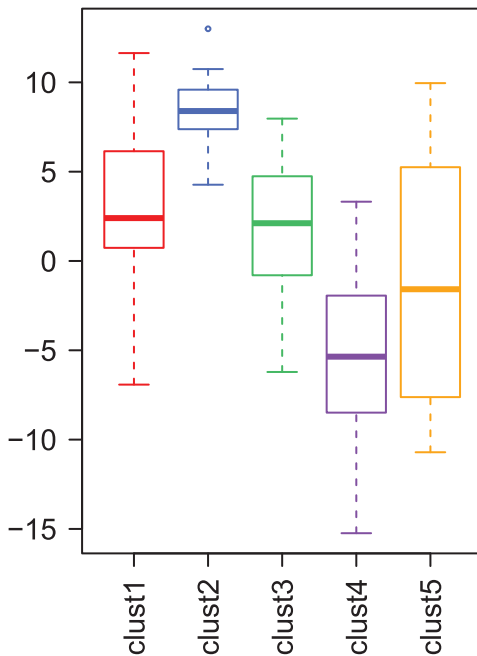
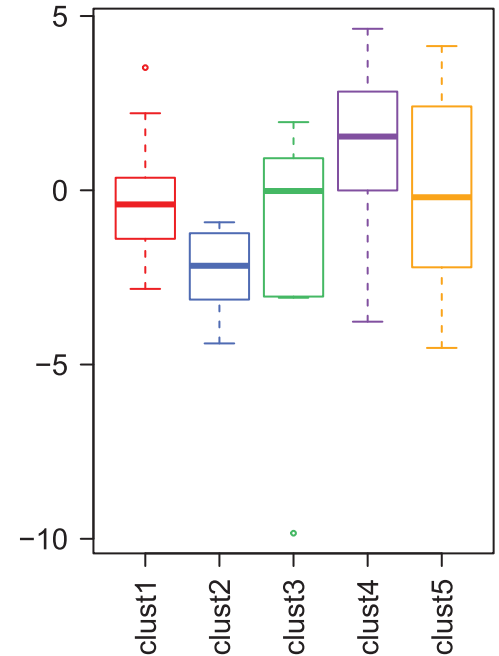


Supplementary Figure 8. Additional display of clinical features across integrative clusters (Cambridge Discovery Cohort).

- A.** Biochemical relapse (chi-sq p=0.0462).
- B.** Extra-capsular extension (chi-sq p=0.029).
- C.** Diagnostic PSA (Kruskal-Wallis p=0.2529).
- D.** Age (chi-sq p=0.5202).
- E.** TMPRSS2-ERG deletion status (K-W p=0.2751).
- F.** Positive surgical margins (chi-sq p=0.0912).

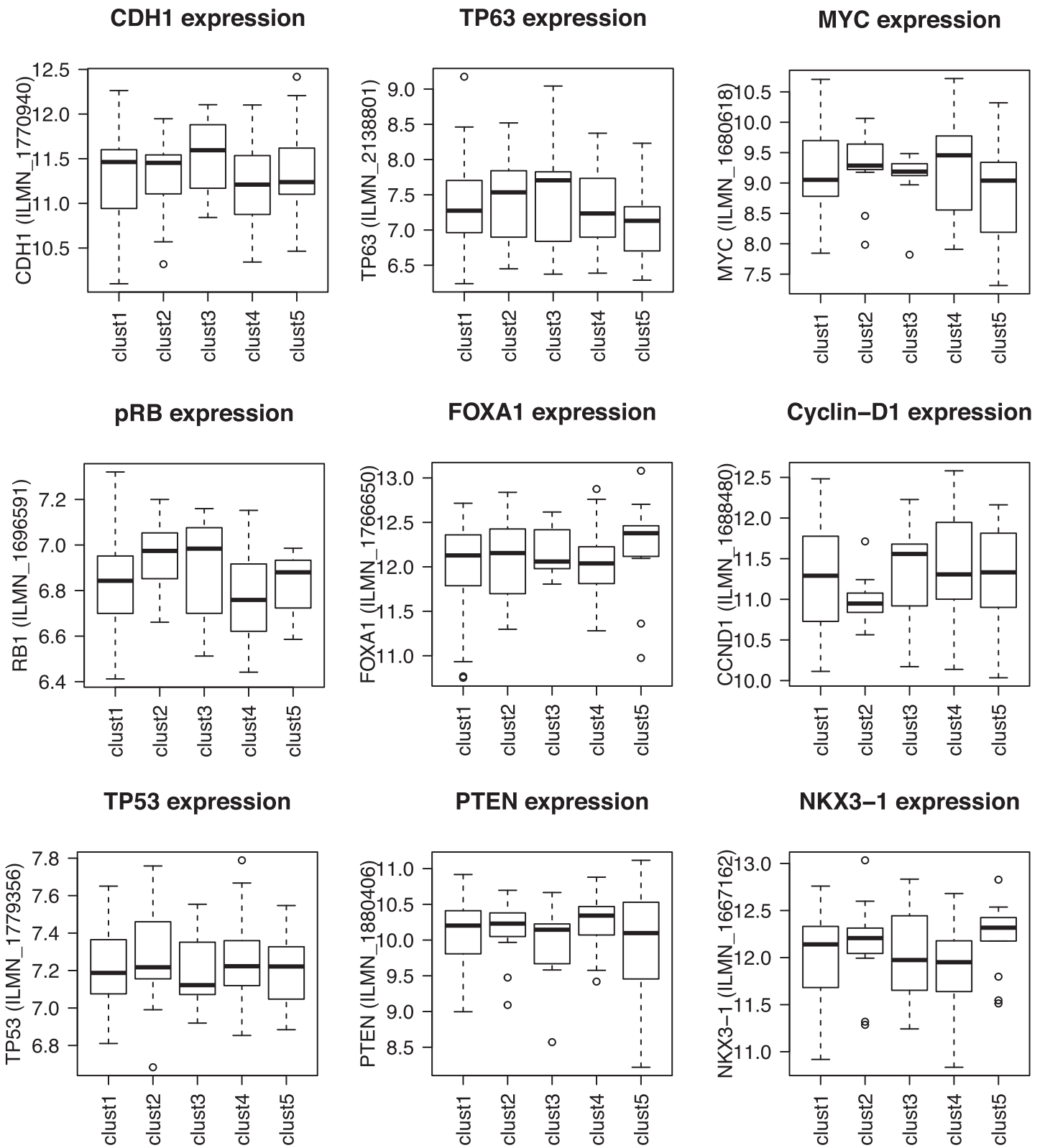


Supplementary Figure 9. Further validation of integrative clusters in predicting biochemical relapse.
A. The five molecular subtypes identified in the discovery cohort were assessed in this third validation set with 120-month follow-up ($p=0.027$; Taylor et al, 2010; see Supplementary Table 7 for clinical features of this cohort).
B. Clinical features of each subgroup are shown.

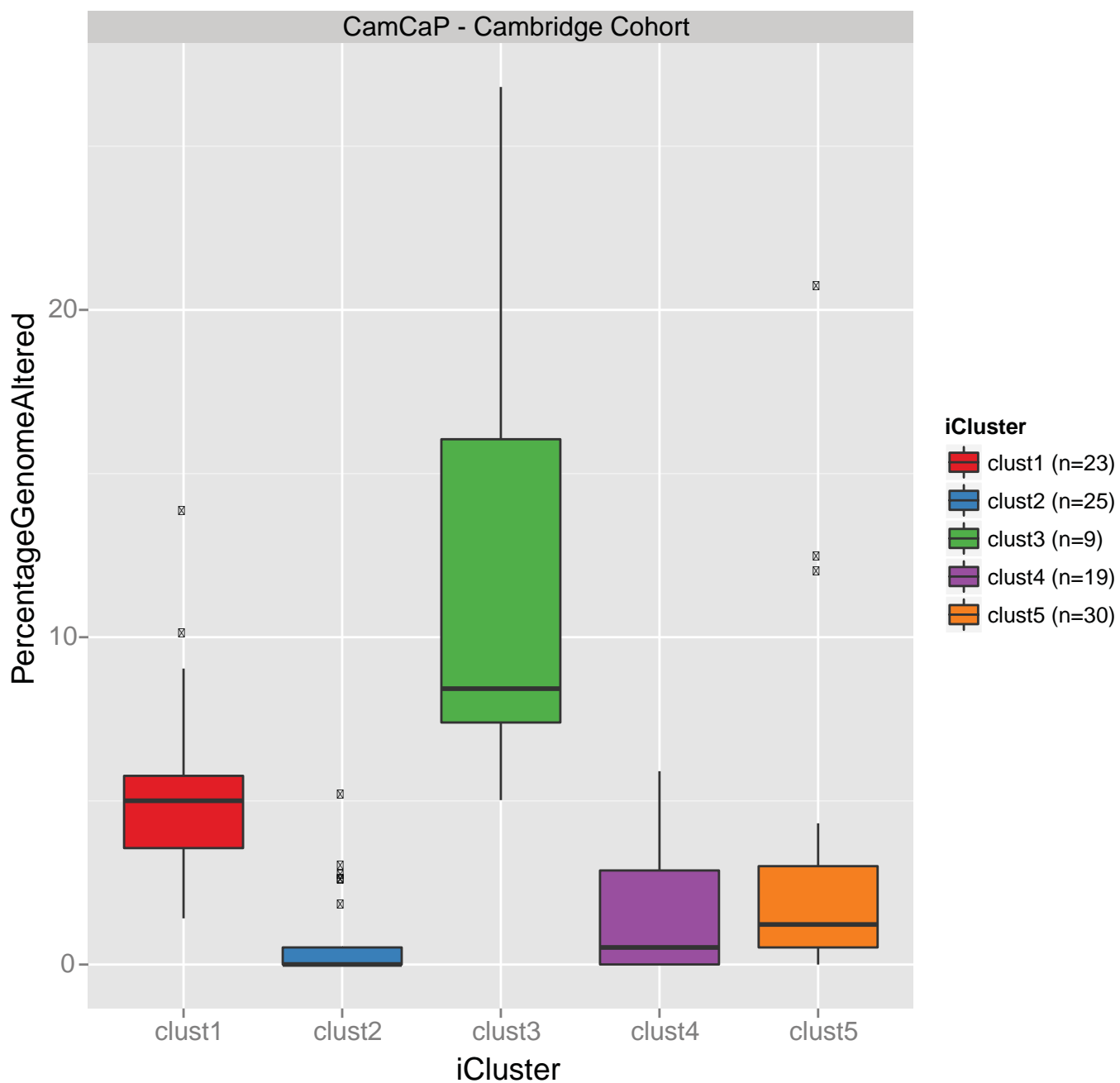
ERG gene set**Mendiratta AR gene set****AR cell line down-regulated****AR cell line up-regulated****AR CRPC castration down****HES6 gene set**

Supplementary Figure 10. Integrated Clusters showed distinct expression of ERG, AR and HES6 gene sets.

Principal components analyses (PCA) on the mRNA data from the Cambridge Discovery Cohort assessing the predominance of published gene sets in different integrated clusters (iCluster). Gene sets were identified from the following: ERG: Massie et al., (2010); Mendiratta et al. (2009), AR cell line up- and down-regulation: Massie et al., (2012); AR CRPC castration down (Sharma et al., 2013) and HES6 (Ramos-Montoya et al., 2014).

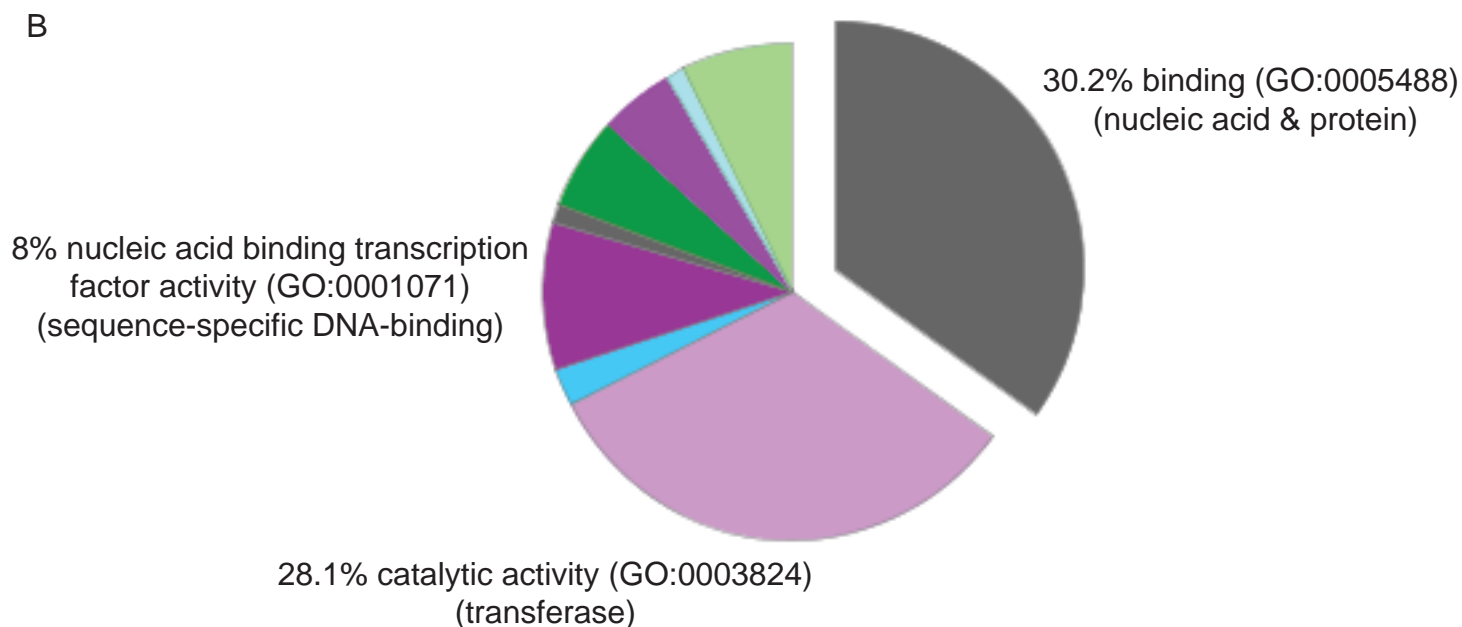
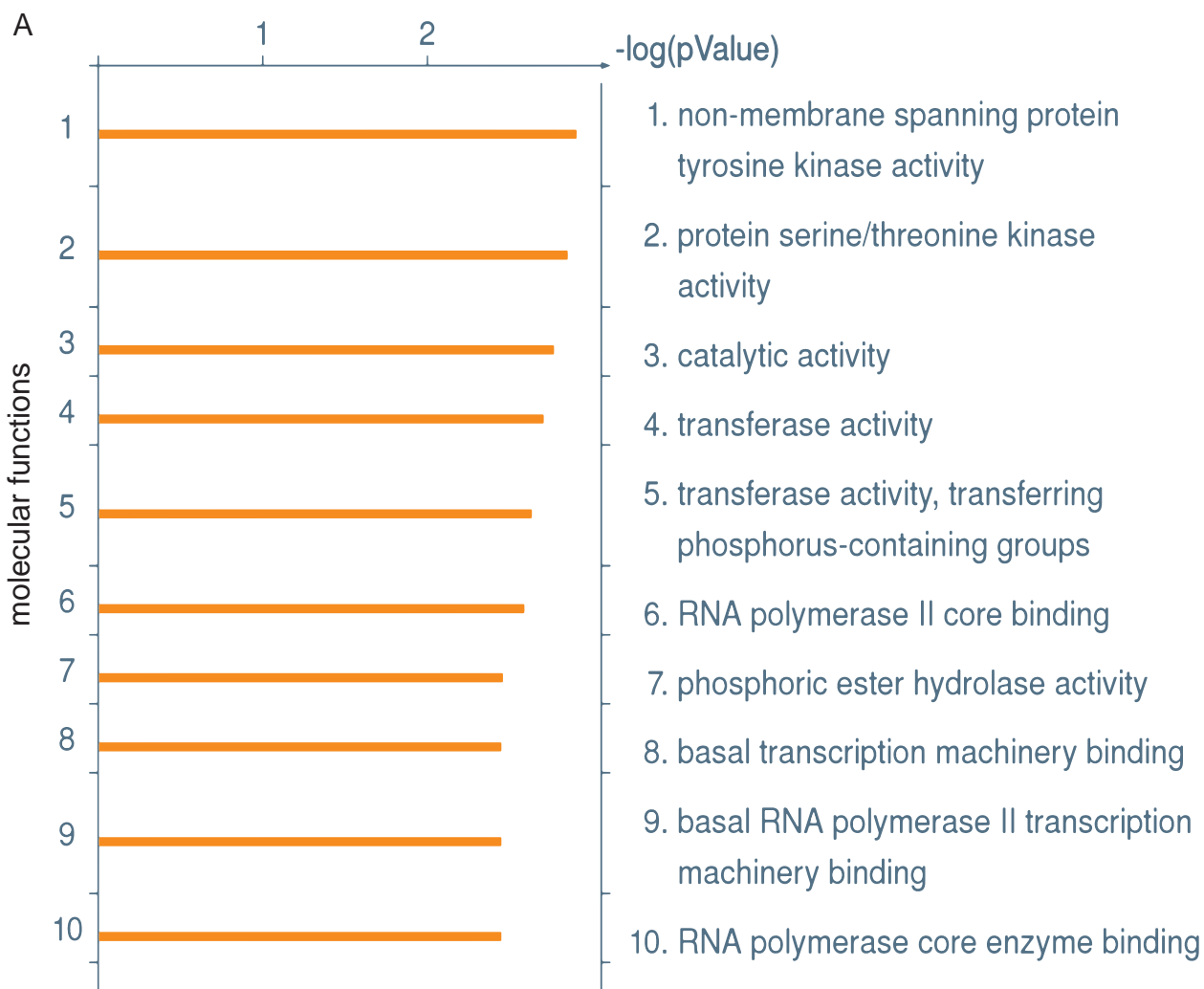


Supplementary Figure 11. Integrated Clusters showed distinct expression of individual genes.
 Relative expression levels of key prostate cancer genes in each cluster in the Cambridge Discovery Cohort



Supplementary Figure 12. Percentage Genome Affected (PGA).

The percentage of the genome affected by copy number change was calculated and mean value plotted for each cluster.

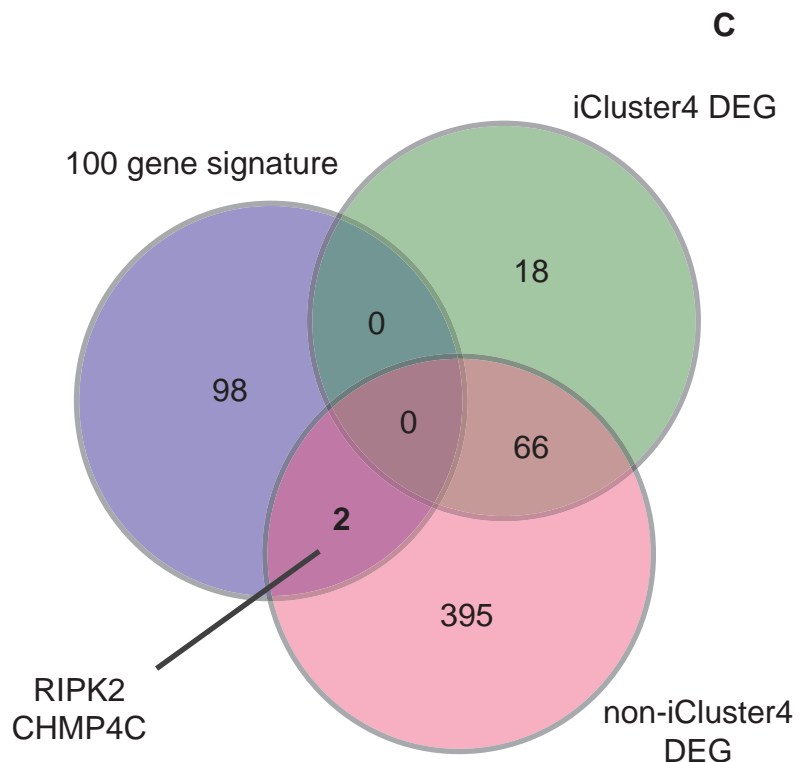
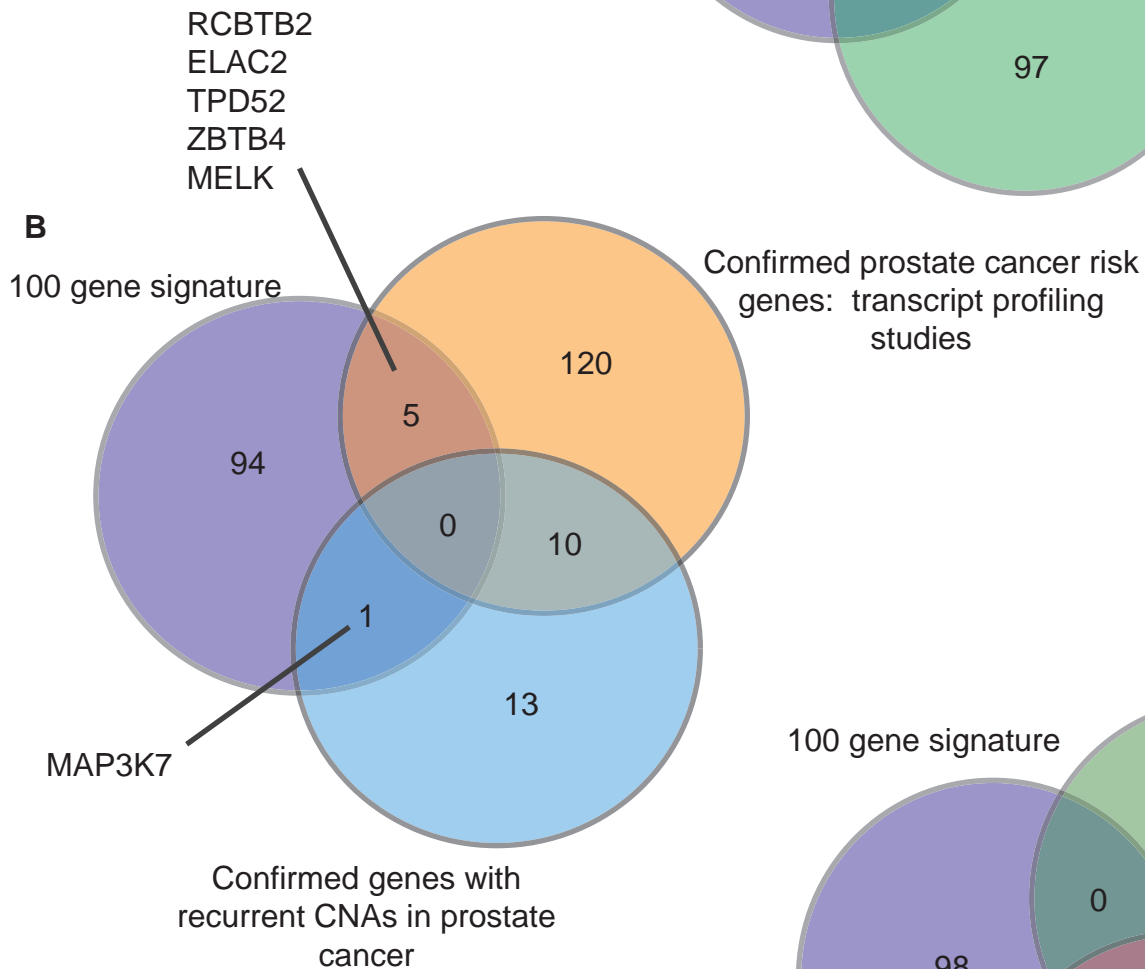
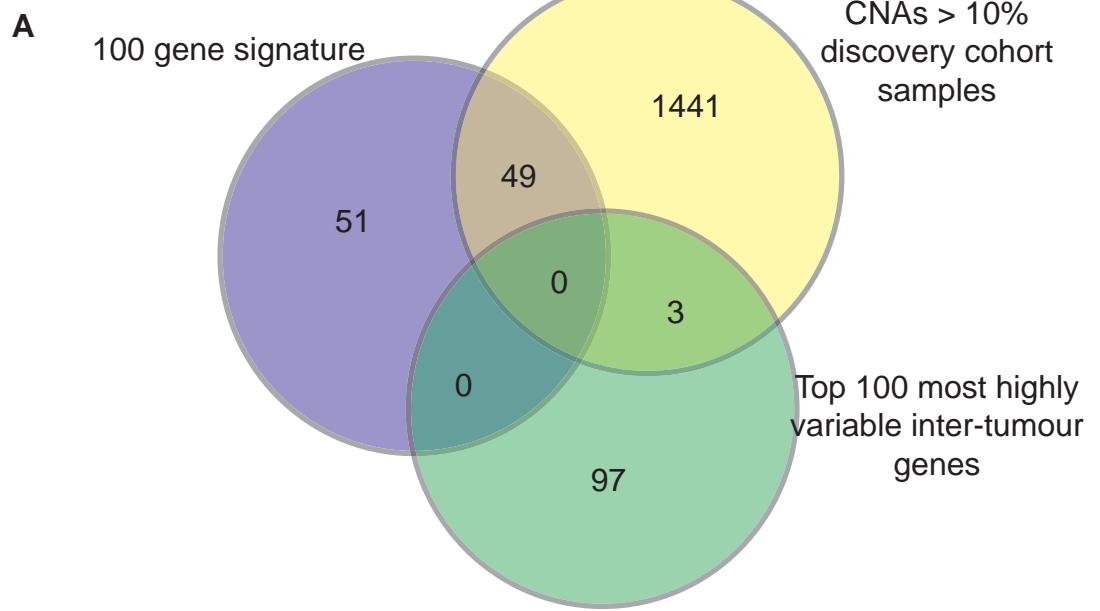


Supplementary Figure 13. Gene ontology pathway analysis of the 100-gene set

A. GeneGO MetaCore (Thomas Reuters) gene ontology analysis.

B. PantherDB (www.pantherdb.org) gene ontology analysis.

Both analyses identified enrichment of pathways relating to RNA and DNA processing – specifically sequence-specific transcription factor and nucleic acid binding – and the phosphorylation of proteins (particularly serine/threonine kinases) in this gene set.



Supplementary Figure 14. Comparison of 100-gene integrative gene set with known CN and gene expression loci.

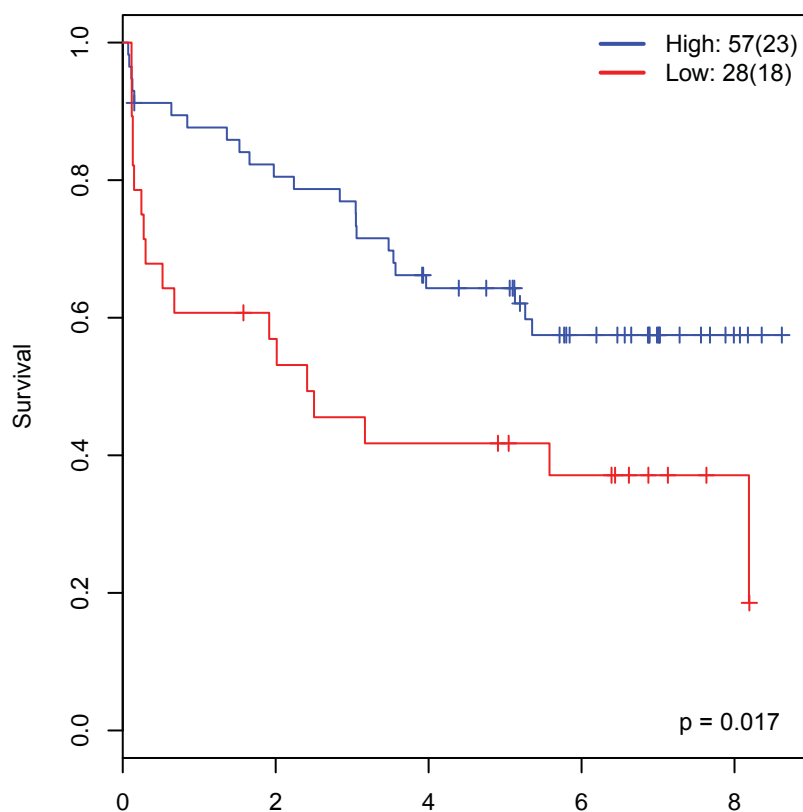
A. Within the Cambridge Discovery Cohort, only 3.3% (49/1490) of all CN changes (>10 % samples tested) were found in the 100-gene set. None of the top 100 most variable inter-tumour genes was included.

B. Only six risk genes from the 100-gene set have previously been described according to published meta-analyses – CN changes in MAP3K7 and gene expression changes in RCBTB2, ELAC2, TPD52, ZBTB4 and MELK. Our eQTL-based gene set therefore identifies 94 targets that would not have been identified using copy number or transcript profiling data alone.

C. Comparison of differentially expressed genes between iCluster4 and non-iCluster 4 identified CHMP4C and RIPK2 in the 100-gene set that could separate iCluster4 from non-iCluster4 using expression levels alone.

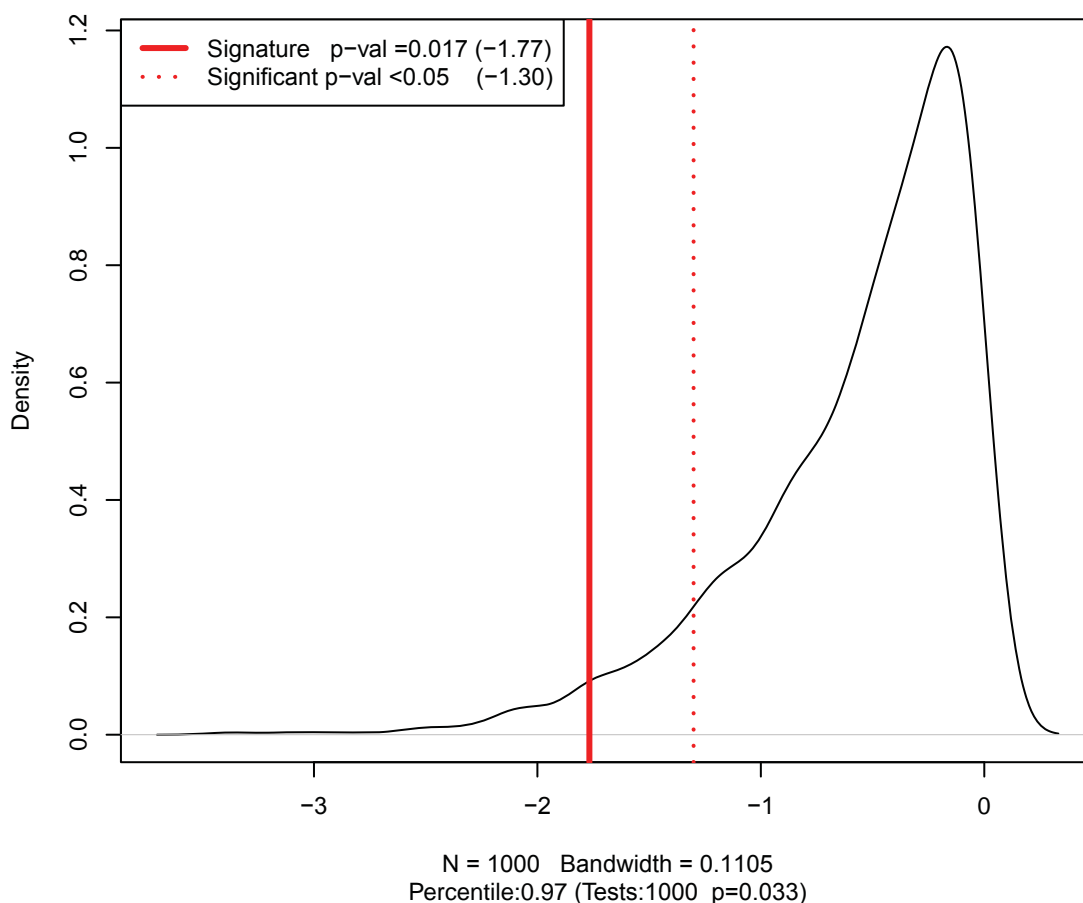
A

Survival: Validation Set



B

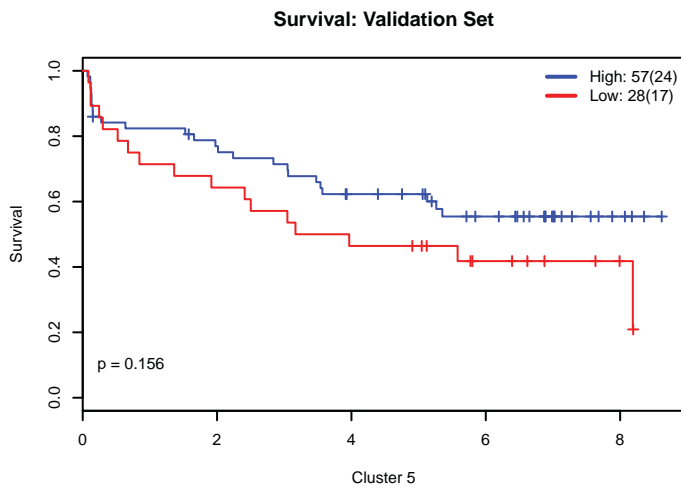
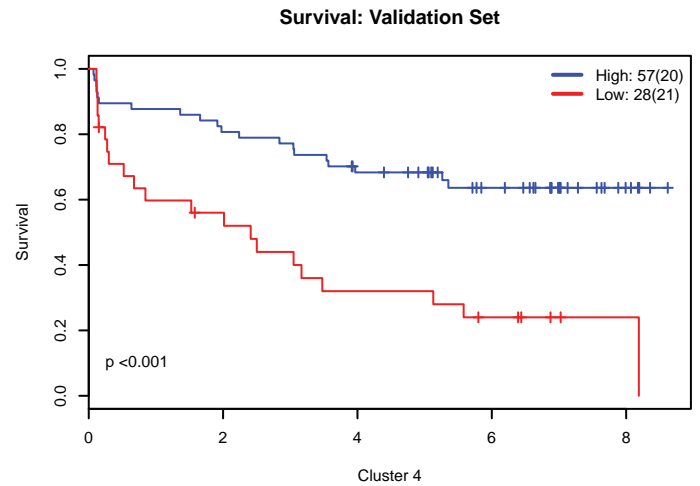
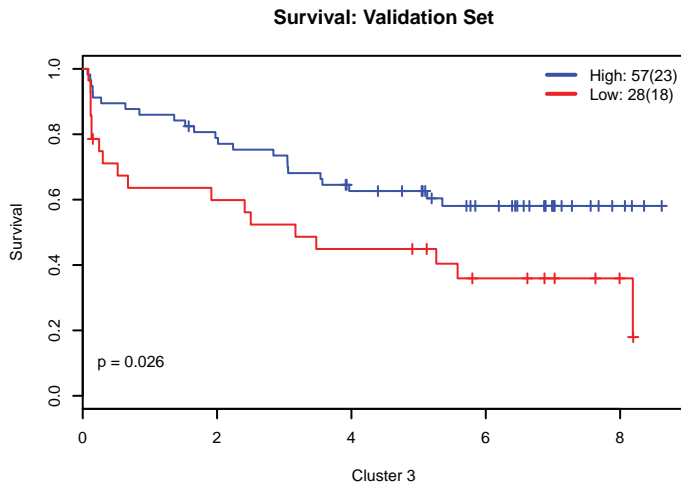
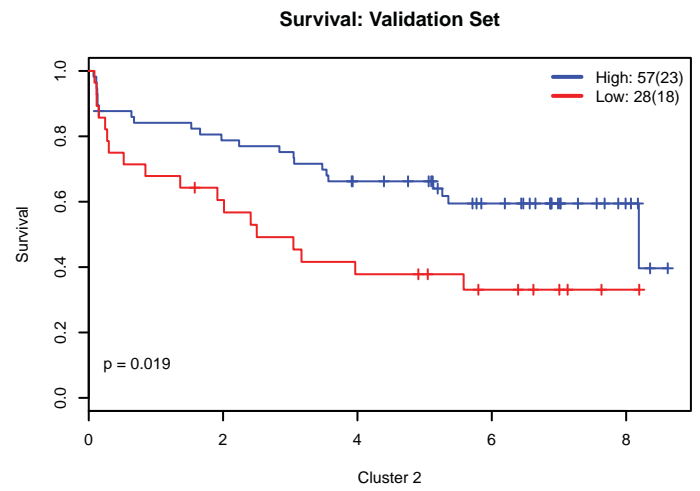
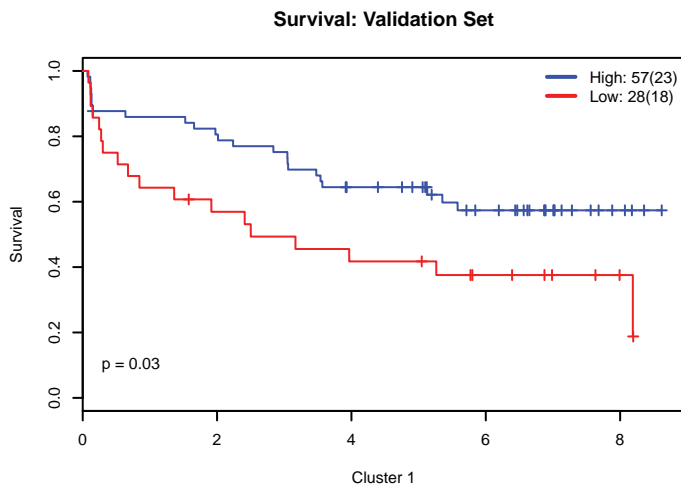
Survival: Random Signatures



Supplementary Figure 15. Performance of 100-gene set as predictive signature in Stockholm validation cohort.

A. Kaplan-Meier plot of biochemical relapse-free survival in the Stockholm validation cohort over 96 months. Patient data were divided into two categories by calculating the mean value over all features for each sample; the 'high' enriched group comprised those samples >33rd percentile, while the 'low' enriched group consisted of samples \leq 33rd percentile. Log-rank p -value = 0.017.

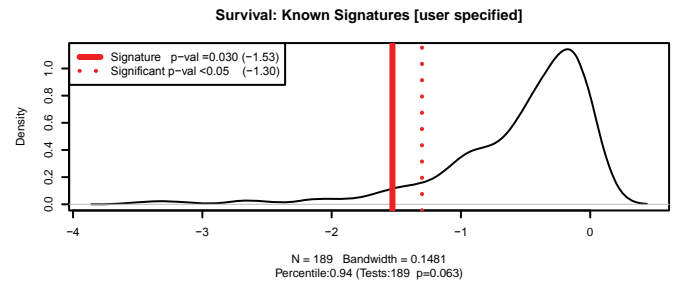
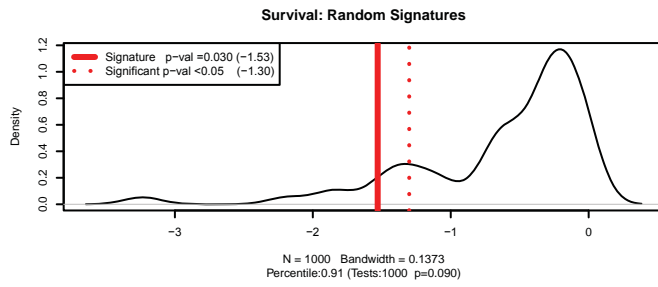
B. The 100-gene set was compared to 1 000 random signatures of the same size. The solid red vertical line shows the performance of the Cambridge signature, while the dotted red vertical line shows $p = 0.05$.



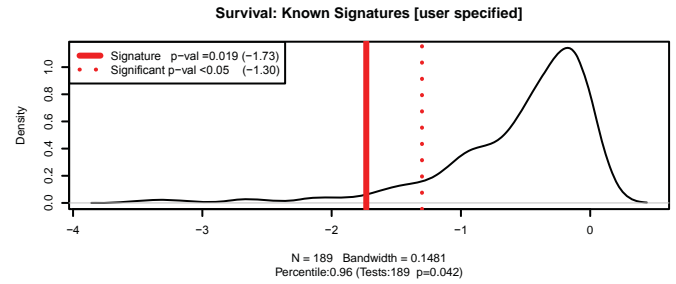
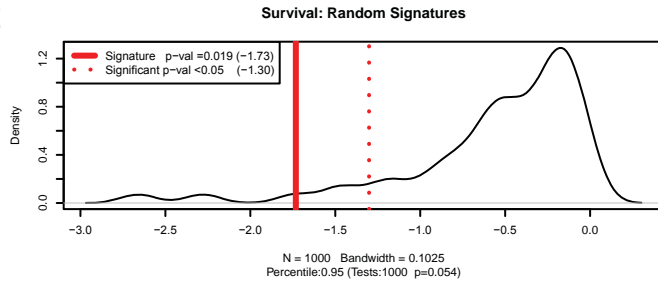
Supplementary Figure 16. Relapse-free survival curves for cluster-specific signatures

Kaplan Meier plots for each of the five clusters, using the 50-feature signatures from rankings computed by PAMR (Tibshirani et al., (2003), applied to the Stockholm validation set with no a priori clustering (see Suppl. Methods).

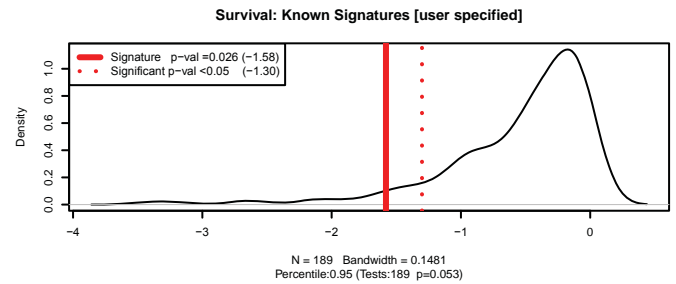
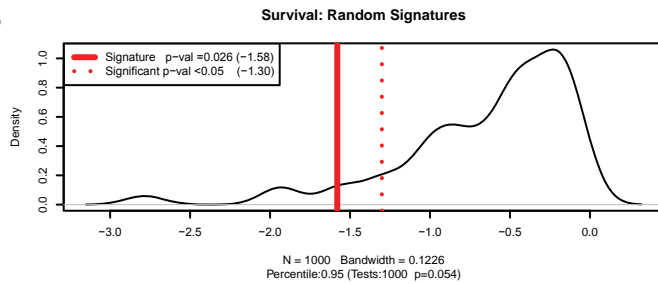
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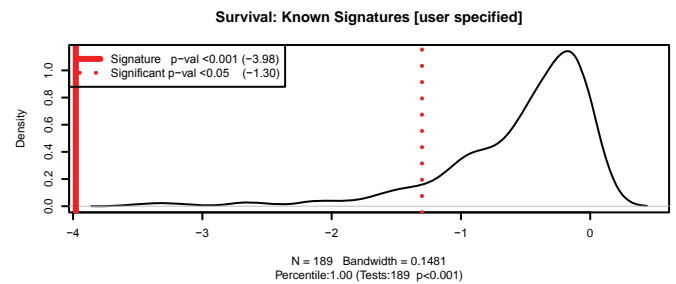
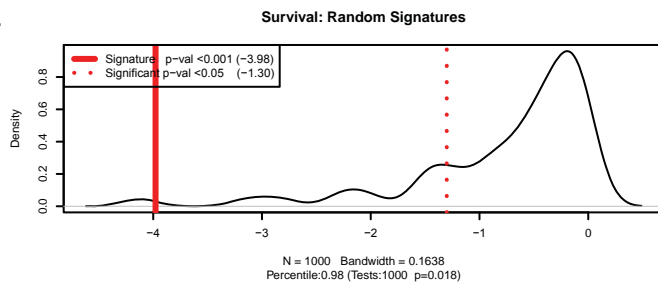
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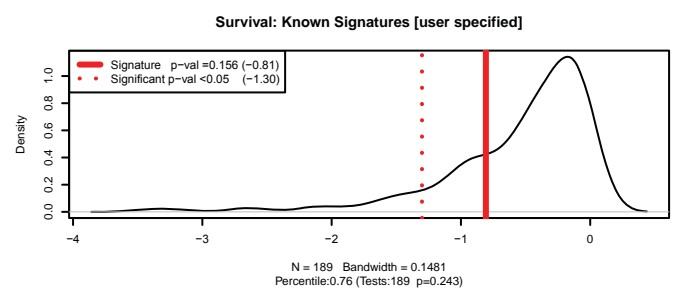
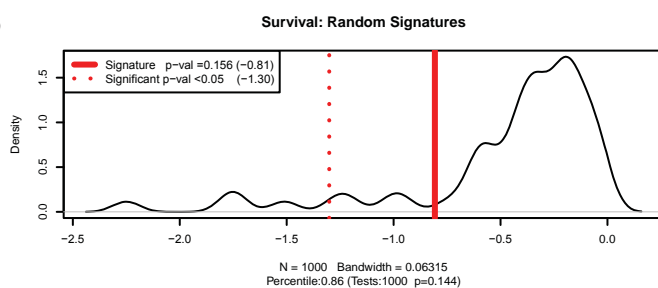
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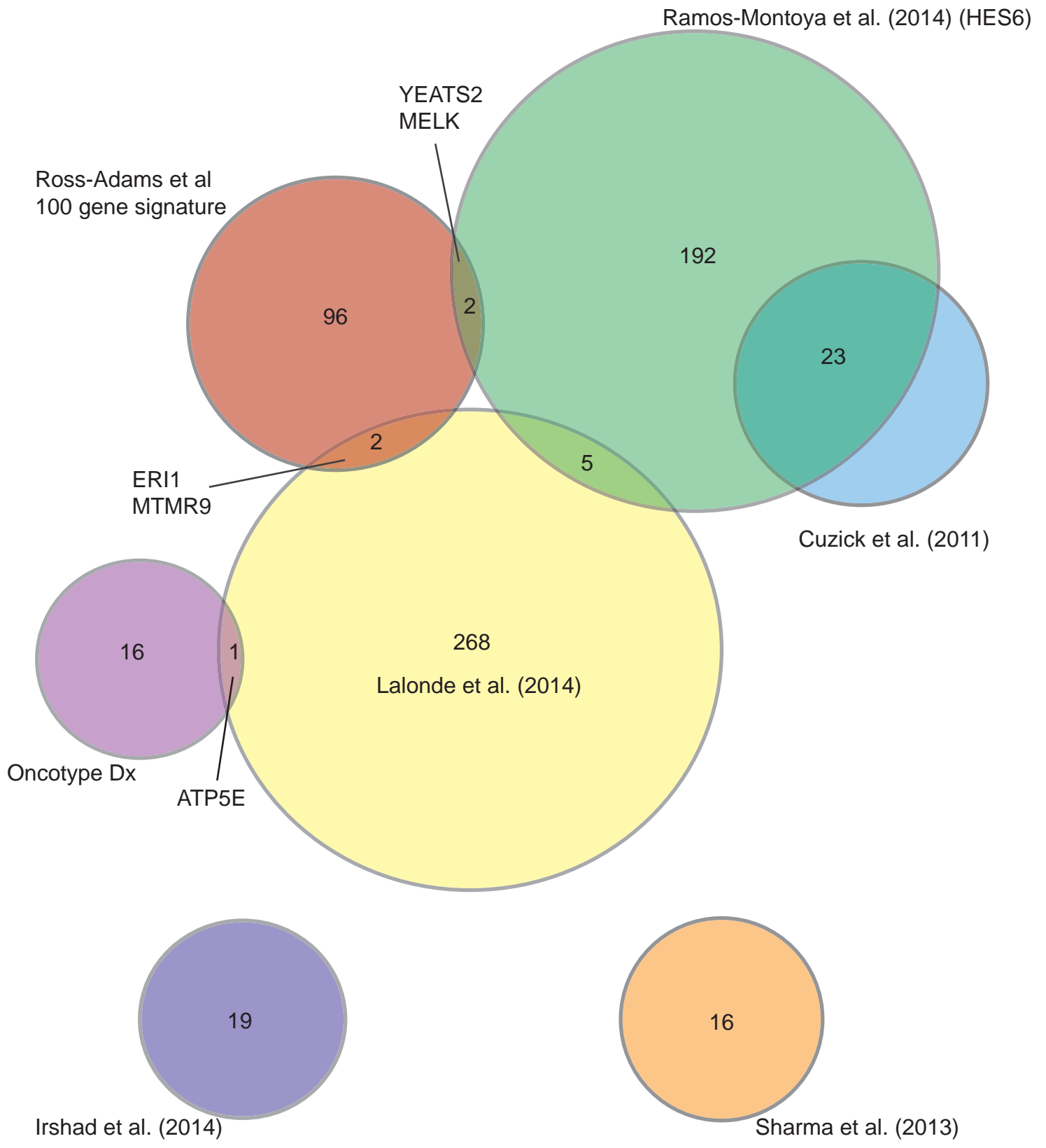
Cluster 4



Cluster 5



Supplementary Figure 17. Cluster-specific refinement of 100-gene set improves prediction of relapse. Subsets of the 100-gene set were identified that best describes each iCluster (see Suppl. Table 11 for gene lists, and Suppl. Methods). These were tested for power to predict relapse in the Stockholm validation set, against 1 000 random signatures of similar size (left column; $p < 0.001$), and 189 oncological signatures (right column; $p < 0.001$) (Subramanian et al., 2005). In each case, the solid red line shows the performance of each Cluster-specific gene signature, as compared to dotted red line $p = 0.05$, against the background/null distribution of the gene signatures tested.



Supplementary Figure 18. Published gene signature overlap.
 Intersects compared between our 100 gene-set and other published gene signatures as shown.

SUPPLEMENTARY METHODS

Integrative clustering

The GenomicRanges and GenomicFeatures infrastructure in Bioconductor was used to define a set of gene-centric coordinates for hg19, and to perform overlaps between the copy number and expression data.¹ For each gene in the TxDb.Hsapiens.UCSC.hg19.knownGene database, we computed a single copy-number and expression value for each sample. For the Cambridge cohort, copy number values were taken as the OncoSNP² Rank 1 call with most-aberrant log-ratio that overlapped the gene. For Stockholm, the longest segment overlapping the gene was used in order to remove short, spurious calls from the data. In both cohorts, gene-centric expression data were derived by first considering Illumina expression probes that overlapped the coordinates of each gene. In the case of multiple overlapping probes, the probe with the highest inter-quartile range (IQR) was used. An ANOVA test was used to compute a p-value, testing the association between expression and copy-number for each gene. All genes were then ranked according to this p-value; lowest (i.e. strongest evidence for association) to highest. The top N ranked genes were then taken forward for integrative analysis; where increasing values of N were used in order to determine to optimal number of features (see below).

Integrative analysis (cluster identification) was performed using copy number and gene expression data from the same 125 PC tissue samples in iCluster+³, with seed set to zero to ensure that the same parameters were generated each time where iCluster+ uses kNN (nearest neighbour) method. All features were included, and used to generate diagnostic plots in both discovery and validation cohorts. To increase stability, penalty parameters (lambda) were set to zero for all (no penalty). Variable selection was done after cluster generation, to reduce variability in results. Since the percentage variation explained naturally increases with an increase in number of clusters (and a reduction in cluster size), we selected the optimal k (5 clusters= $k+1$; $k=4$) at the point beyond which additional sample divisions provided no significant improvement in variation explained – see Suppl. Fig 5. In-group proportions (IGP) were used to determine optimal number of $k=4$ (5 clusters), where IGP is defined as the proportion of data in a group whose nearest neighbours are classified to the same group (Pearson's centred correlation).

The stability of iCluster+ was confirmed by considering the *distribution* of explained variation for each k after running 100 different analyses with 100 features each and a range of $k=1$ to 10. There was no value for k after which the explained variation became more unstable; clusters were reliably reproduced. After training of the classifier using primary tumours of the Cambridge cohort with iCluster+³, pamr⁴ in Bioconductor was used to predict clusters for the Cambridge CRPC samples, as well as in the Stockholm and Taylor et al., (2010)⁵ validation cohorts.

iCluster comparison

A core set of 106 samples pairs were used to accurately compare partitioning of samples based on either CN or mRNA or CN+mRNA data (integrative). We specified the iClusters to be 'true', and applied the recommended Adjusted Rand Index (ARI)⁶ using fpc in Bioconductor to measure the relative accuracy of alternative clustering, where identical clustering gives ARI=1, and ARI=0 implies completely random re-clustering. Since there is no absolute threshold for similarity/divergence (but 0.35 would not occur by chance; Suppl. Table 6), and outputs must be considered relative to the set 'true' value, we also calculated the Variation of Information Index (VII)⁷ in Bioconductor, where VII=0 for identical clustering and VII>0 for more disparate groupings; there is no maximum value.

Cambridge prostate cancer (CamCaP) gene signature analysis

This section describes an analysis of the genes and features identified as driving our clustering, treated as a prognostic gene signature. The goal is to identify the ability of these genes to stratify patients into outcome groups, and to determine how unique they are in predicting outcome for prostate cancer patients.

Genes, features, training set

We identified 100 genes for use in deriving tumour subtype clusters. For every sample, each of these genes has two features: a gene expression value and a copy number value. The genes were chosen based on a correlation between the expression value and the copy number values in the Cambridge (discovery) cohort samples (i.e. eQTL), here known as the *training/discovery set*.

Validation Sets

Evaluation of the performance of the 100-gene set was performed using the Stockholm patient set, consisting of 103 samples, 99 with both copy number and expression and of which 85 samples had relapse information.

Patient categories

Patients were divided into two categories using the 100-gene set by calculating the mean expression over all the features associated with genes in the signature for each sample, and selecting a “high” enriched group consisting of those above the 33rd percentile, and a “low” enriched group at or below the 33rd percentile.

Prognostic power of the gene signature

The baseline test of the gene signature involved survival analyses on the validation data to determine the ability of the gene signature to separate a class of patients with worse outcome (as measured by time to relapse) from other, better performing patients. For this, we generated Kaplan-Meier plots, and computed associated p-values using a chi-squared test. For this test, the p-value was computed as $p=0.017$ (Suppl. Fig 14A).

Specificity of the gene signature

It has been shown that most oncogenic gene signatures are highly predictive of prognosis in breast cancer, even if they were derived using data for a different cancer⁸. We therefore performed a more rigorous set of tests to determine whether our gene signature had prognostic power beyond other possible gene sets. For this we used SigCheck in Bioconductor⁹. This package performs tests against randomly selected gene signatures, as well as sets of previously published signatures with discriminatory power value in various oncogenic contexts.

a. Compared to random sets of genes

We tested the p-value of our signature against a null distribution of random signatures of similar size (number of genes). This distribution is constructed by generating 1 000 random samples of 1 000 genes, and repeating the survival analysis for each set. An empirical p-value was computed by counting the number of random gene signatures with p-values equal to or lower than that computed to our gene signature. Our gene signature performed at the 97th percentile, for an empirical p-value computed at $p=0.033$. Suppl. Fig 14B shows how the signature compares to the background distribution of randomly selected 100-gene signatures.

b. Compared to known gene signatures

The second set of tests is against previously published gene signatures. We use two sets of cancer signatures.

1. A set of 189 oncogenic signatures in MSigDB¹⁰.
Our signature performed at the 96th percentile, for an empirical p-value of $p=0.042$.
2. A set of six signatures specific to prostate cancer:
 - Sharma (2013): 16 genes relating to CRPC¹¹
 - Cuzick (2011): 31 genes relating to prognosis in primary PC¹²
 - Irshad (2013): 19 genes relating to indolent PC¹³
 - HES6 (2014): A set of 222 genes associated with HES6 & CRPC¹⁴
 - Lalonde (2014): A set of 276 genes (corresponding to 100 loci) relating to chromosome instability, and predicting <5 year relapse (intended to be used with copy number data only)¹⁵
 - OncoTypeDx: A set of 17 genes on commercial Oncotype Dx prognostic arrays

Not only did our signature outperform all six of these, none of the previously identified prostate-cancer specific signatures achieved significant results on the Stockholm dataset, mean gene expression:

<u>Signature</u>	<u>p-value</u>
SHARMA	0.174
CUZICK	0.263
IRSHAD	0.853
HES6	0.189
LALONDE	0.495
ONCOTYPEDX	0.732

Cluster-specific signatures

In our final analysis of signature performance, we looked at the features *within* our 100 gene set that were most important for defining our five clusters, yielding five distinct prognostic signatures (one for each cluster). We wanted to see if any of these five clusters had particularly interesting prognostic power.

We obtained a signature for each cluster by ranking the importance of features on a classification task, separating the samples in that cluster from those in other clusters. This was repeated for each cluster. As we had 100 genes and two features for each gene (expression and copy number), we had a total of 200 features from which to select a signature. For this analysis, we generated signatures comprised of the genes corresponding to the 50 highest-ranked *features* for each cluster, selected using rankings computed by the PAMR package in Bioconductor⁴. These signatures were then each applied to the Stockholm validation set (with no *a priori* clustering) (Suppl. Fig 16). When both the copy number and expression features for the same gene were used, these mapped to only one gene in the signature, so there could be as few as 25 genes in a signature.

Four out of five of the clusters exhibited significant ability to separate a poor outcome class in the Stockholm validation set, as shown in Suppl. Fig 15 and Table 3. In addition, it showed how none of the five previously identified signatures or the commercially available OncoType Dx gene signature, was able to separate a distinct survival group using mean expression values. We next compared the performance of the cluster signatures to 1 000

signatures of equivalent size (number of genes), as well as 189 previously identified oncogenic signatures (see Suppl. Fig 16).

The best performing cluster signature is that obtained for iCluster4 (purple curve; figure 3). On the survival task, we calculated a p-value of 0.0001. When compared to random signatures, iCluster4 signature performed in the 98th percentile for an empirical p-value of $p=0.018$. This signature also out-performed all 189 oncogenic signatures from MSigDB (Suppl. Fig 16 & Suppl. Table 9). These are presented in PAMR-ranked order of importance in discriminating the cluster. Further investigation showed that not all 50 genes are required to generate good survival performance in the Stockholm validation set, with even lower p-values obtained for as few as 15-20 genes. However, we strictly applied the same criteria for all five signatures in the discovery cohort, and did not select a lower cut-off that performed better in the validation set, since performance in a validation set cannot be used to set any parameters.

Supplementary Methods - References

1. Lawrence M, Huber W, Pagès H, et al. Software for Computing and Annotating Genomic Ranges. *PLoS Comput Biol* 2013; **9**(8): e1003118.
2. Yau C, Mouradov D, Jorissen RN, et al. A statistical approach for detecting genomic aberrations in heterogeneous tumor samples from single nucleotide polymorphism genotyping data. *Genome biology* 2010; **11**(9): R92.
3. Mo Q, Wang S, Seshan VE, et al. Pattern discovery and cancer gene identification in integrated cancer genomic data. *Proceedings of the National Academy of Sciences of the United States of America* 2013; **110**(11): 4245-50.
4. Tibshirani R, Hastie T, Narasimhan B, Chu G. Class Prediction by Nearest Shrunken Centroids, with Applications to DNA Microarrays. 2003: 104-17.
5. Taylor BS, Schultz N, Hieronymus H, et al. Integrative genomic profiling of human prostate cancer. *Cancer cell* 2010; **18**(1): 11-22.
6. Hubert L, Arabie P. Comparing partitions. *Journal of Classification* 1985; **2**(1): 193-218.
7. Meilă M. Comparing clusterings—an information based distance. *Journal of Multivariate Analysis* 2007; **98**(5): 873-95.
8. Venet D, Dumont JE, Detours V. Most Random Gene Expression Signatures Are Significantly Associated with Breast Cancer Outcome. *PLoS Comput Biol* 2011; **7**(10): e1002240.
9. Norden JS, Rory. SigCheck: Check a gene signature's classification performance against random signatures, permuted data, and known signatures. R package version 1.0.2. ed; 2014.
10. Subramanian A, Tamayo P, Mootha VK, et al. Gene set enrichment analysis: A knowledge-based approach for interpreting genome-wide expression profiles. *Proceedings of the National Academy of Sciences* 2005; **102**(43): 15545-50.
11. Sharma NL, Massie CE, Ramos-Montoya A, et al. The androgen receptor induces a distinct transcriptional program in castration-resistant prostate cancer in man. *Cancer cell* 2013; **23**(1): 35-47.
12. Cuzick J, Swanson GP, Fisher G, et al. Prognostic value of an RNA expression signature derived from cell cycle proliferation genes in patients with prostate cancer: a retrospective study. *The Lancet Oncology* 2011; **12**(3): 245-55.
13. Irshad S, Bansal M, Castillo-Martin M, et al. A molecular signature predictive of indolent prostate cancer. *Science translational medicine* 2013; **5**(202): 202ra122.
14. Ramos-Montoya A, Lamb AD, Russell R, et al. HES6 drives a critical AR transcriptional programme to induce castration-resistant prostate cancer through activation of an E2F1-mediated cell cycle network. *EMBO Mol Med* 2014; **6**(5): 651-61.
15. Lalonde E, Ishkanian AS, Sykes J, et al. Tumour genomic and microenvironmental heterogeneity for integrated prediction of 5-year biochemical recurrence of prostate cancer: a retrospective cohort study. *The Lancet Oncology* 2014; **15**(13): 1521-32.

CASE EXAMPLE

Integration of copy number and transcriptomics provides risk stratification in prostate cancer: a discovery and validation cohort study

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Men who have clinically low risk disease at diagnosis but who turn out to have unexpectedly aggressive disease are a problem for the prostate cancer medical community. There seem to be two principal reasons for this. The first is sampling error. Prostate biopsies cannot sample the whole prostate gland with complete accuracy and therefore may miss the dominant tumour nodule. But the second is that our current methods in disease risk stratification are inadequate. In our study we have proposed a molecular classification for men with prostate cancer that will improve such stratification.

We present here a case where such molecular knowledge, derived from information discovered in our study, would have changed clinical management.

Case

This man was diagnosed with prostate cancer in 2008 and his key clinical parameters are outlined below.

	Diagnostic Biopsy	Radical Prostatectomy
Gleason Score	3+3=6; 3/10 cores; 5%	3+4=7
PSA	4.3	No change
TNM	T1cN0Mx	T2aN0Mx
ECE	N/A	No
SMS	N/A	Negative

On the basis of these details he would be considered to have low risk prostate cancer according to D'Amico criteria at biopsy. He was upgraded and upstaged at prostatectomy but would still have been considered to be at relatively low risk of relapse (see Figure 1). If the Memorial Sloane Kettering nomogram is used (<https://www.mskcc.org/nomograms/prostate/pre-op>) then he would be considered to have a probability of progression-free survival at 5 years of 94% based on the diagnostic biopsy and 91% based on the prostatectomy specimen.

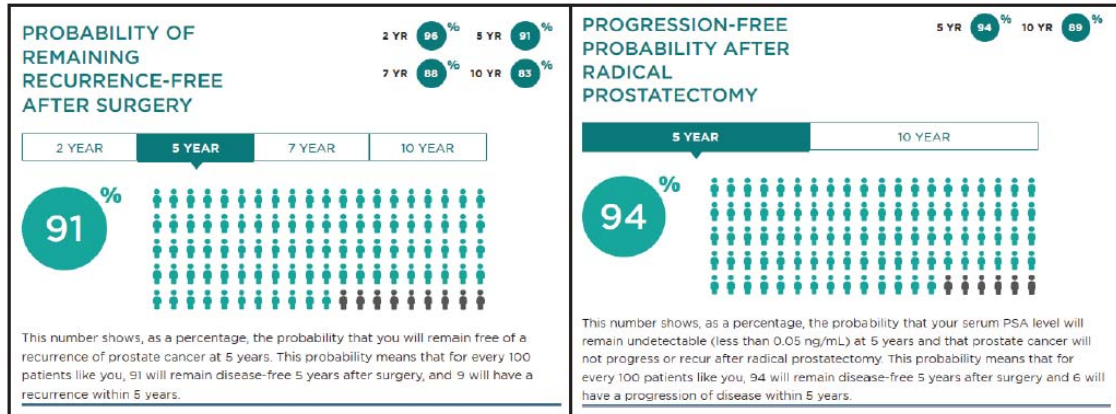


Figure 1. Memorial Sloan Kettering prostate cancer risk nomogram.

Patient details input to <https://www.mskcc.org/nomograms/prostate> for both pre-operative and post-operative outcomes.

On integrated genomic analysis, this man was found to be in iCluster3, our highest risk subgroup. So, although his clinical data suggested he was at low risk of relapse he is actually, according to our data, between 30 and 78% likely to relapse. In the event, this man had adjuvant / early salvage radiotherapy at 7 months and this seems to have been effective as his PSA has remained steady since (Figure 2) suggesting that his relapse was due to recurrence from unidentified locally advanced disease (local equivalent of micrometastatic disease). We propose that genomic information contained within this study could be used at the time of biopsy to advocate surgery with wider margins, or after prostatectomy to advocate early adjuvant radiotherapy or even neoadjuvant treatment with an appropriate small molecule.

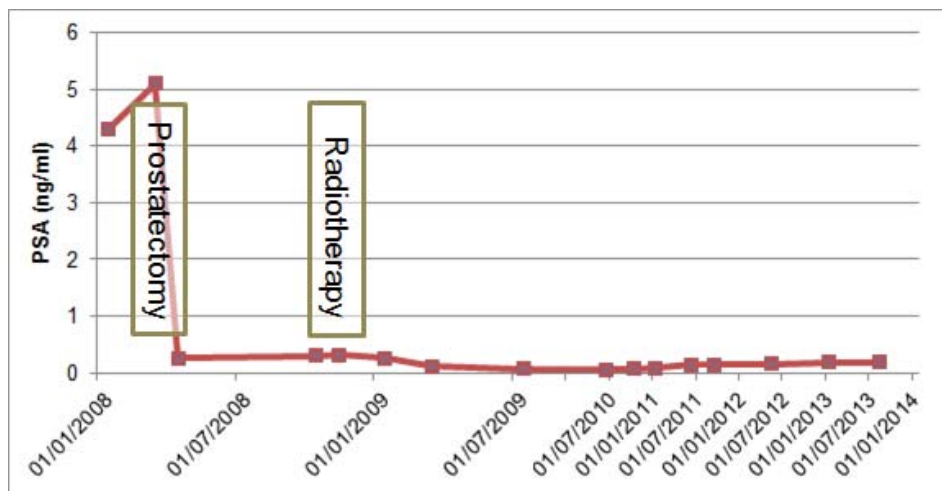


Figure 2. PSA kinetics from diagnosis to most recent recorded follow up.

In addition to signature profiling another area in which this information could be used is in immunohistochemistry of biopsy or prostatectomy specimens. In order to help investigate this we generated a tissue microarray (TMA) containing tumour cores for every man in the Cambridge cohort undergoing prostatectomy. These specimens were taken from regions of the prostate paraffin megablock directly adjacent to the frozen section from which nucleic acids were extracted. A representative H&E stain of one of the four TMA blocks is shown (Figure 3A) as well as of the man

described here. A core demonstrating his prostate cancer is presented (Figure 3B) as well as our first attempt at staining for a protein coded by a member of the 100-gene set presented in this study, MELK. This staining needs optimising and it is hoped that further antibody selection for genes discriminating the high-risk subgroups will yield clinically useful diagnostic information.

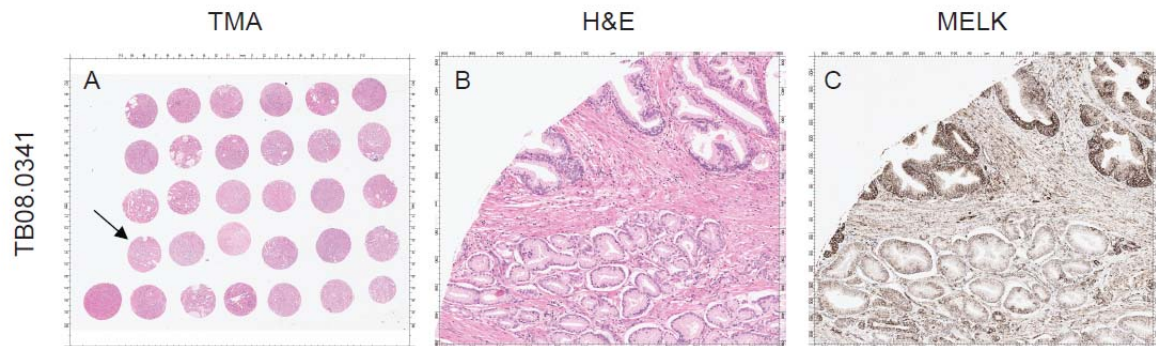


Figure 3. Histology from CamCaP robotic prostatectomy TMA

A. Screenshot of relevant block of TMA (one of four blocks). Each 3mm core is taken from a section of megablock neighbouring the region from which nucleic acids were extracted for integrated analysis. Arrow indicates the man in this case example.

B. Haematoxylin & Eosin stain at 10x magnification of region of infiltrating cancer. Scale as shown.

C. MELK immunohistochemistry (IHC, Sigma Atlas #HPA017214). MELK is one of the 100-gene set of discriminating genes. Staining is poor here. Further work-up of relevant antibodies is needed to identify members of the gene-set that are accurate in identifying members of high-risk subgroups before an IHC/histological approach could be used.

Supplementary Table 1. Available prostate cancer datasets with both mRNA and copy number data

Dataset	Platform Gene expression	Platform CN	# primary tumours	# advanced tumours
Varambally <i>et al.</i> (2005)	Affymetrix U133 2.0	N/A	7	6
Tomlins <i>et al.</i> (2005)	not reported	N/A	22	9
Taylor <i>et al.</i> (2010)	Affymetrix Human Exon 1.0 ST	Agilent 244K	109	19
Markert <i>et al.</i> (2011)	Illumina Custom Array (6,144 genes)	N/A	281	
Barbieri <i>et al.</i> (2012)	Illumina RNA-seq	Affymetrix SNP6	112 (CN)	-
			22 (mRNA & CN)	
Grasso <i>et al.</i> (2012)	Exome-seq (Illumina GAIIx)	Agilent 105K or 244K	59	32
Baca <i>et al.</i> (2013)*	Exome-seq (Illumina GAIIx)	WGS (Illumina GAIIx) & Affymetrix SNP 6	55	2
TCGA - provisional‡	Illumina RNA-seq	Affymetrix SNP6	171	-

* data not available

‡ data only available after analysis completed

Supplementary Table 3. Frequencies of copy number variations (CNVs) in known prostate cancer risk genes in discovery and validation cohorts

genes tagging region*	chr	% other primary PC cohorts*	% Cambridge discovery cohort	% Stockholm validation cohort
NKX3-1	8p	55.7%	39.6%	44.7%
RB1	13q	44.9%	23.6%	42.7%
RYBP	3p13	19.8%	19.8%	32.0%
PTEN	10q	26.4%	17.9%	37.9%
CDKN1B	12p	23.8%	17.0%	29.1%
SHQ1	3p13	19.8%	15.1%	30.1%
MAP3K7	6q	40.8%	15.1%	45.6%
FOXP1	3p13	19.8%	15.1%	29.1%
MYC	8q24	20.9%	13.2%	36.9%
TP53	17p	28.4%	11.3%	32.0%
BRCA2	13q	44.9%	9.4%	32.0%
BCL2	18q	25.5%	7.5%	35.9%
MCM7	7p	14.1%	7.5%	33.0%
CDH1	16q	44.3%	7.5%	31.1%
CHD1	5q	35.9%	6.6%	37.9%
APC	5q	35.9%	6.6%	36.9%
ETV4	17q	17.4%	6.6%	30.1%
ETV5	3q	9.7%	6.6%	29.1%
SMAD4	18q	25.5%	5.7%	35.9%
CXCR4	2q	22.9%	4.7%	37.9%
EGFR	7p	14.1%	4.7%	31.1%
PIK3CA	3q	9.7%	4.7%	30.1%
BRCA1	17q	17.4%	3.8%	27.2%
BRAF	7p	14.1%	3.8%	31.1%
ERG	21q	25.2%	2.8%	15.5%
ETV1	7p	14.1%	1.9%	32.0%
TMPRSS2	21q	25.1%	0.0%	19.4%

* From Williams et al. (2014) meta-analysis of recurrent CNVs in PC

Cambridge cohort OMNI2.5M data were analysed in OncoSNP (Yau et al., 2010); all rank calls 1-5 counted
Stockholm SNP6 data were analysed in ASCAT (van Loo et al., 2010); no filtering on calls

Supplementary Table 4. All genes with CN altered in $\geq 10\%$ Cambridge samples tested (OMNI2.5M; Onc

chr	chrStart	chrEnd	gene	strand	CNA type
chr8	47752507	47767407	LINC00293	+	gain
chr8	48100929	48104439	LOC100287846	+	gain
chr8	48173488	48678652	SPIDR	+	gain
chr8	48872762	48890719	MCM4	+	gain
chr8	48920994	48974454	UBE2V2	+	gain
chr8	49966894	49988642	C8orf22	+	gain
chr8	50822348	51705427	SNTG1	+	gain
chr8	53852467	53853454	NPBWR1	+	gain
chr8	54764367	54871863	RGS20	+	gain
chr8	55047780	55061074	MRPL15	+	gain
chr8	55370494	55373456	SOX17	+	gain
chr8	55528626	55682531	RP1	+	gain
chr8	56015016	56438710	XKR4	+	gain
chr8	56685790	56738005	TGS1	+	gain
chr8	56792385	56925006	LYN	+	gain
chr8	57124314	57131176	CHCHD7	+	gain
chr8	58131647	58145388	LOC100507651	+	gain
chr8	58173784	58179170	LOC286177	+	gain
chr8	58192101	58197290	LINC00588	+	gain
chr8	58907112	59062277	FAM110B	+	gain
chr8	59323822	59364060	UBXN2B	+	gain
chr8	59465727	59495419	SDCBP	+	gain
chr8	61429468	61536203	RAB2A	+	gain
chr8	61591323	61780586	CHD7	+	gain
chr8	62200524	62414204	CLVS1	+	gain
chr8	62627346	62627418	MIR4470	+	gain
chr8	63161500	63903628	NKAIN3	+	gain
chr8	63890419	63897460	UG0898H09	+	gain
chr8	64081111	64125346	YTHDF3	+	gain
chr8	64681987	64698054	LINC01289	+	gain
chr8	65285774	65295842	MIR124-2HG	+	gain
chr8	65291705	65291814	MIR124-2	+	gain
chr8	65492794	65496191	BHLHE22	+	gain
chr8	66556887	66622798	MTFR1	+	gain
chr8	66933790	67012755	DNAJC5B	+	gain
chr8	67039277	67087718	TRIM55	+	gain
chr8	67104348	67109554	LINC00967	+	gain
chr8	67341262	67342968	RRS1	+	gain
chr8	67344717	67381044	ADHFE1	+	gain
chr8	67405490	67430759	C8orf46	+	gain
chr8	67579786	67593377	C8orf44	+	gain
chr8	67579786	67774257	C8orf44-SGK3	+	gain
chr8	67624652	67774257	SGK3	+	gain
chr8	67782983	67834283	MCMD2	+	gain
chr8	67976602	68108849	CSPP1	+	gain
chr8	68864602	69143897	PREX2	+	gain
chr8	69242956	69731258	C8orf34	+	gain
chr8	70378858	70573147	SULF1	+	gain

chr8	71520811	71575513	LOC286190	+	gain
chr8	71581599	71648177	XKR9	+	gain
chr8	72755357	72968547	LOC100132891	+	gain
chr8	73449625	73850584	KCNB2	+	gain
chr8	73921096	73959987	TERF1	+	gain
chr8	74206836	74237520	RDH10	+	gain
chr8	74332308	74353753	STAU2-AS1	+	gain
chr8	74888376	74895018	TMEM70	+	gain
chr8	74903563	74941307	LY96	+	gain
chr8	75262617	75279335	GDAP1	+	gain
chr8	75460777	75460852	MIR5681A	+	gain
chr8	75512100	75670587	FLJ39080	+	gain
chr8	75617927	75617982	MIR2052	+	gain
chr8	75736771	75767264	PI15	+	gain
chr8	75896707	75946793	CRISPLD1	+	gain
chr8	76320182	76479061	HNF4G	+	gain
chr8	77593514	77779521	ZFHX4	+	gain
chr8	79428335	79517502	PKIA	+	gain
chr8	79578281	79631997	ZC2HC1A	+	gain
chr8	80523048	80578410	STMN2	+	gain
chr8	81153623	81153708	MIR5708	+	gain
chr8	81397853	81438500	ZBTB10	+	gain
chr8	82192717	82197012	FABP5	+	gain
chr8	82644687	82671748	CHMP4C	+	gain
chr8	85095452	85834078	RALYL	+	gain
chr8	86019322	86058314	LRRCC1	+	gain
chr8	86089618	86126753	E2F5	+	gain
chr8	86132844	86352138	CA13	+	gain
chr8	86351055	86361267	CA3	+	gain
chr8	86376130	86393721	CA2	+	gain
chr8	87111138	87166454	ATP6VOD2	+	gain
chr8	87354993	87480178	WWP1	+	gain
chr8	87526655	87573726	CPNE3	+	gain
chr8	87878675	88394955	CNBD1	+	gain
chr8	90769974	90803292	RIPK2	+	gain
chr8	90914095	90940095	OSGIN2	+	gain
chr8	91013579	91064227	DECR1	+	gain
chr8	91233715	91400187	LINC00534	+	gain
chr8	91803920	91971630	NECAB1	+	gain
chr8	92082423	92099323	OTUD6B	+	gain
chr8	92114846	92231464	LRRC69	+	gain
chr8	92217712	92217786	MIR4661	+	gain
chr8	92221721	92410378	SLC26A7	+	gain
chr8	94712772	94740671	FAM92A1	+	gain
chr8	94752338	94753047	RBM12B-AS1	+	gain
chr8	94767071	94830347	TMEM67	+	gain
chr8	94929082	94938296	PDP1	+	gain
chr8	95653363	95719694	ESRP1	+	gain
chr8	95732102	95806076	DPY19L4	+	gain
chr8	95835517	95892721	INTS8	+	gain

chr8	95970235	96088396	NDUFAF6	+	gain
chr8	96085141	96085221	MIR3150A	+	gain
chr8	96145948	96168913	PLEKHF2	+	gain
chr8	96281063	96822371	C8orf37-AS1	+	gain
chr8	97274166	97346774	PTDSS1	+	gain
chr8	97505881	97624037	SDC2	+	gain
chr8	97657498	98155722	CPQ	+	gain
chr8	98656406	98742488	MTDH	+	gain
chr8	98787808	98864830	LAPTM4B	+	gain
chr8	98881310	99048946	MATN2	+	gain
chr8	99076749	99105838	ERICH5	+	gain
chr8	99129520	99172069	POP1	+	gain
chr8	99439249	99443023	KCNS2	+	gain
chr8	99956630	99964326	OSR2	+	gain
chr8	100025493	100889814	VPS13B	+	gain
chr8	101162838	101166230	POLR2K	+	gain
chr8	101170262	101254132	SPAG1	+	gain
chr8	101394990	101395073	MIR4471	+	gain
chr8	102064281	102088479	FLJ42969	+	gain
chr8	102381120	102381823	NACAP1	+	gain
chr8	102504667	102681952	GRHL2	+	gain
chr8	103137659	103137743	MIR5680	+	gain
chr8	103563847	103573245	ODF1	+	gain
chr8	104033247	104085285	ATP6V1C1	+	gain
chr8	104152920	104242533	BAALC	+	gain
chr8	104166841	104166917	MIR3151	+	gain
chr8	104310660	104345094	FZD6	+	gain
chr8	104383742	104395232	CTHRC1	+	gain
chr8	104426941	104455680	DCAF13	+	gain
chr8	104512975	105266656	RIMS2	+	gain
chr8	105352023	105368917	DCSTAMP	+	gain
chr8	106331146	106816767	ZFPM2	+	gain
chr8	107282405	107764921	OXR1	+	gain
chr8	109455852	109499136	EMC2	+	gain
chr8	110099652	110131812	TRHR	+	gain
chr8	110346551	110358189	ENY2	+	gain
chr8	110374705	110543500	PKHD1L1	+	gain
chr8	110551928	110577391	EBAG9	+	gain
chr8	113655721	113655812	MIR2053	+	gain
chr8	117778741	117786921	UTP23	+	gain
chr8	117886662	117889107	RAD21-AS1	+	gain
chr8	117950463	117956239	AARD	+	gain
chr8	117962511	118188953	SLC30A8	+	gain
chr8	118532964	118552501	MED30	+	gain
chr8	119633239	119738306	SAMD12-AS1	+	gain
chr8	120079423	120119202	COLEC10	+	gain
chr8	120220609	120257914	MAL2	+	gain
chr8	120428551	120436678	NOV	+	gain
chr8	120885899	121063157	DEPTOR	+	gain
chr8	121137346	121384273	COL14A1	+	gain

chr8	121457637	121535875	MTBP	+	gain
chr8	122651585	122657564	HAS2-AS1	+	gain
chr8	123793900	123986755	ZHX2	+	gain
chr8	124084919	124164392	TBC1D31	+	gain
chr8	124191286	124222318	FAM83A	+	gain
chr8	124428964	124454260	WDYHV1	+	gain
chr8	124780881	124827690	FAM91A1	+	gain
chr8	124864226	125132302	FER1L6	+	gain
chr8	125463047	125465266	TRMT12	+	gain
chr8	125487007	125500859	RNF139	+	gain
chr8	125551342	125580751	NDUFB9	+	gain
chr8	125951883	125963337	LINC00964	+	gain
chr8	125985538	125991630	ZNF572	+	gain
chr8	126010719	126034525	SQLE	+	gain
chr8	126104082	126379367	NSMCE2	+	gain
chr8	126442562	126450644	TRIB1	+	gain
chr8	128025398	128033259	PCAT1	+	gain
chr8	128427856	128429441	POU5F1B	+	gain
chr8	128747764	128753680	MYC	+	gain
chr8	128806802	129113499	PVT1	+	gain
chr8	128808207	128808274	MIR1204	+	gain
chr8	128972878	128972941	MIR1205	+	gain
chr8	129021143	129021202	MIR1206	+	gain
chr8	129061397	129061484	MIR1207	+	gain
chr8	129162361	129162434	MIR1208	+	gain
chr8	132916355	133025886	EFR3A	+	gain
chr8	133787603	133861052	PHF20L1	+	gain
chr8	133879204	134147143	TG	+	gain
chr8	134203281	134243932	WISP1	+	gain
chr8	135610313	135612932	ZFAT-AS1	+	gain
chr8	136246373	136311962	LOC286094	+	gain
chr8	136469715	136659848	KHDRBS3	+	gain
chr8	141521396	141527252	CHRA1	+	gain
chr8	142138719	142205906	DENND3	+	gain
chr8	142350647	142354720	LINC01300	+	gain
chr8	142432006	142441620	PTP4A3	+	gain
chr8	143257699	143257779	MIR4472-1	+	gain
chr8	143279716	143290364	LINC00051	+	gain
chr8	143545376	143626368	BAI1	+	gain
chr8	143751725	143764145	PSCA	+	gain
chr8	143781528	143785584	LY6K	+	gain
chr8	143808620	143818350	THEM6	+	gain
chr8	143916216	143928262	GML	+	gain
chr8	144077244	144079080	CDC42P3	+	gain
chr8	144099901	144103827	LY6E	+	gain
chr8	144120678	144135720	C8orf31	+	gain
chr8	144295067	144299044	GPIHBP1	+	gain
chr8	144328990	144359101	ZFP41	+	gain
chr8	144349606	144359101	GLI4	+	gain
chr8	144373558	144382120	ZNF696	+	gain

chr8	144451024	144466390	RHPN1	+	gain
chr8	144635556	144645231	GSDMD	+	gain
chr8	144680073	144682485	TIGD5	+	gain
chr8	144718182	144735900	ZNF623	+	gain
chr8	144766621	144794977	ZNF707	+	gain
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chr8	144798506	144804633	MAPK15	+	gain
chr8	144816309	144828507	FAM83H-AS1	+	gain
chr8	145064225	145067583	GRINA	+	gain
chr8	145086581	145102015	SPATC1	+	gain
chr8	145133521	145135551	EXOSC4	+	gain
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chr8	145149941	145152428	CYC1	+	gain
chr8	145159304	145162515	MAF1	+	gain
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chr8	145515269	145538385	HSF1	+	gain
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chr8	145729464	145732555	GPT	+	gain
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chr8	56980738	56987140	RPS20	-	gain
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chr8	57025500	57026541	MOS	-	gain
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chr8	87484577	87521009	RMDN1	-	gain
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chr8	100548863	100548958	MIR599	-	gain
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chr8	146065828	146126846	COMMD5	-	gain
chr8	146102335	146126846	ZNF250	-	gain
chr8	146155743	146176274	ZNF16	-	loss
chr8	146198974	146228285	ZNF252P	-	loss
chr3	71803200	71804328	GPR27	+	loss
chr3	72200407	72223490	LINC00870	+	loss
chr3	71003864	71633140	FOXP1	-	loss
chr3	71591120	71591240	MIR1284	-	loss
chr3	71728439	71803924	EIF4E3	-	loss
chr3	71820805	71834357	PROK2	-	loss
chr3	72423743	72495774	RYBP	-	loss
chr3	72798427	72897598	SHQ1	-	loss
chr6	80816343	81055987	BCKDHB	+	loss
chr6	83072922	83077133	TPBG	+	loss
chr6	83777384	83881065	DOPEY1	+	loss
chr6	83903031	83906256	RWDD2A	+	loss
chr6	84222193	84235421	PRSS35	+	loss
chr6	84562984	84567234	RIPPLY2	+	loss
chr6	84569369	84670146	CYB5R4	+	loss
chr6	84743419	84800605	MRAP2	+	loss
chr6	86159301	86205509	NT5E	+	loss
chr6	87647023	87726397	HTR1E	+	loss
chr6	87862550	87973406	ZNF292	+	loss
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chr6	88054570	88075181	C6orf163	+	loss
chr6	88091719	88109459	C6orf164	+	loss
chr6	88117689	88205858	C6orf165	+	loss
chr6	88146443	88222057	SLC35A1	+	loss
chr6	88299784	88377172	ORC3	+	loss
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chr8	21964382	21967456	NUDT18	-	loss
chr8	21971931	21988565	HR	-	loss
chr8	21995532	21999448	REEP4	-	loss
chr8	22004342	22014344	LGI3	-	loss
chr8	22077215	22089851	PHYHIP	-	loss
chr8	22102474	22102556	MIR320A	-	loss
chr8	22477946	22526661	BIN3	-	loss
chr8	22497883	22499722	BIN3-IT1	-	loss
chr8	22545173	22550815	EGR3	-	loss
chr8	22570764	22785421	PEBP4	-	loss
chr8	22877647	22926700	TNFRSF10B	-	loss
chr8	22993100	23021543	TNFRSF10D	-	loss
chr8	23048969	23082680	TNFRSF10A	-	loss
chr8	23154409	23261722	LOXL2	-	loss
chr8	23243441	23315244	ENTPD4	-	loss
chr8	23536205	23540450	NKX3-1	-	loss
chr8	23559963	23564111	NKX2-6	-	loss
chr8	23699433	23712320	STC1	-	loss
chr8	24808468	24814383	NEFL	-	loss
chr8	25276773	25282556	GNRH1	-	loss
chr8	25285363	25315984	KCTD9	-	loss
chr8	25699245	25902640	EBF2	-	loss
chr8	26362195	26371483	PNMA2	-	loss
chr8	26605666	26722922	ADRA1A	-	loss
chr8	27093813	27115903	STMN4	-	loss
chr8	27142403	27168834	TRIM35	-	loss
chr8	27317277	27336813	CHRNA2	-	loss
chr8	27454433	27472328	CLU	-	loss
chr8	27559189	27559284	MIR3622B	-	loss
chr8	27590832	27630170	CCDC25	-	loss
chr8	27667137	27695572	PBK	-	loss
chr8	27727398	27850369	SCARA5	-	loss
chr8	27743555	27743633	MIR4287	-	loss
chr8	27879480	27941388	NUGGC	-	loss
chr8	28203101	28347835	ZNF395	-	loss
chr8	28285925	28347835	FBXO16	-	loss

chr8	28362632	28362699	MIR4288	-	loss
chr8	28625174	28747698	INTS9	-	loss
chr8	28924794	29120610	KIF13B	-	loss
chr8	29190578	29208267	DUSP4	-	loss
chr8	29578775	29605625	LINC00589	-	loss
chr8	29814787	29814864	MIR3148	-	loss
chr8	29920257	30108213	MIR54802	-	loss
chr8	29920630	29940649	SARAF	-	loss
chr8	29989186	30002200	MBOAT4	-	loss
chr8	30239634	30242917	RBPMS-AS1	-	loss
chr8	30436030	30515738	GTF2E2	-	loss
chr8	30535579	30585486	GSR	-	loss
chr8	30643125	30670352	PPP2CB	-	loss
chr8	30689059	30706533	TEX15	-	loss
chr8	30853320	30891231	PURG	-	loss
chr8	33228343	33330664	FUT10	-	loss
chr8	33330903	33370703	TTI2	-	loss
chr8	33405272	33424643	RNF122	-	loss
chr8	33448853	33457439	DUSP26	-	loss
chr8	37604073	37605564	LOC728024	-	loss
chr8	37701397	37707431	BRF2	-	loss
chr8	37716464	37757015	RAB11FIP1	-	loss
chr8	37791799	37797647	GOT1L1	-	loss
chr8	37820513	37824184	ADRB3	-	loss
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chr8	38020838	38034248	LSM1	-	loss
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chr8	38132560	38239790	WHSC1L1	-	loss
chr8	38268655	38326352	FGFR1	-	loss
chr8	38368351	38386180	C8orf86	-	loss
chr8	38457692	38458775	RNF5P1	-	loss
chr8	38846326	38854041	TM2D2	-	loss
chr8	39601254	39695808	ADAM2	-	loss
chr8	40388110	40755343	ZMAT4	-	loss
chr8	41119475	41166990	SFRP1	-	loss
chr8	41503828	41643385	NKX6-3	-	loss
chr8	41510743	41754280	ANK1	-	loss
chr8	41517958	41518026	MIR486-1	-	loss
chr8	41786996	41909505	KAT6A	-	loss
chr8	42032235	42065194	PLAT	-	loss
chr8	42231585	42234674	DKK4	-	loss
chr8	42273979	42397356	SLC20A2	-	loss
chr8	42607779	42623929	CHRNA6	-	loss
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chr8	42704779	42751866	RNF170	-	loss
chr8	42751339	42751418	MIR4469	-	loss
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chr10	88426550	88495824	LDB3	+	loss
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chr10	88718287	88723017	SNCG	+	loss

chr10	88728187	88730666	ADIRF	+	loss
chr10	88730497	88784487	AGAP11	+	loss
chr10	88780045	88784487	FAM25A	+	loss
chr10	88854952	88951222	FAM35A	+	loss
chr10	88985204	88994733	NUTM2A	+	loss
chr10	89102167	89103331	LOC439994	+	loss
chr10	89117476	89130452	NUTM2D	+	loss
chr10	89263637	89263711	MIR4678	+	loss
chr10	89264222	89313218	MINPP1	+	loss
chr10	89419475	89507462	PAPSS2	+	loss
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chr10	89623194	89728532	PTEN	+	loss
chr10	90346518	90366733	LIPJ	+	loss
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chr10	90484300	90512513	LIPK	+	loss
chr10	90521162	90537999	LIPN	+	loss
chr10	90562486	90580303	LIPM	+	loss
chr10	90639943	90734910	STAMBPL1	+	loss
chr10	90729552	90775542	FAS	+	loss
chr10	90823092	90823167	MIR4679-1	+	loss
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chr10	88809958	88854776	GLUD1	-	loss
chr10	88965544	89102315	NUTM2A-AS1	-	loss
chr10	89156330	89167457	LINC00864	-	loss
chr10	89512874	89577917	ATAD1	-	loss
chr10	89618917	89623194	KLLN	-	loss
chr10	89892056	90343082	RNLS	-	loss
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chr10	90694830	90751147	ACTA2	-	loss
chr10	90751180	90752732	FAS-AS1	-	loss
chr10	90823091	90823168	MIR4679-2	-	loss
chr10	90965693	90967071	CH25H	-	loss
chr10	90973325	91174382	LIPA	-	loss
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chr12	11323779	11328619	LOC100129361	+	loss
chr12	11700963	11717335	LINC01252	+	loss
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chr12	12764830	12798041	CREBL2	+	loss
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chr12	12878850	12944399	APOLD1	+	loss
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chr12	12917582	12917677	MIR613	+	loss
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chr12	13043955	13066600	GPRC5A	+	loss
chr12	13068762	13068852	MIR614	+	loss
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chr12	14927269	14930936	H2AFJ	+	loss
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chr12	15125955	15134799	PDE6H	+	loss
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chr12	10851675	10875953	YBX3	-	loss
chr12	10954130	10955226	TAS2R7	-	loss
chr12	10958649	10959579	TAS2R8	-	loss
chr12	10961692	10962767	TAS2R9	-	loss
chr12	10977944	10978868	TAS2R10	-	loss
chr12	10998447	11002075	PRR4	-	loss
chr12	10998447	11324224	PRH1-PRR4	-	loss
chr12	11033559	11036874	PRH1	-	loss
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chr12	11060524	11062161	TAS2R13	-	loss
chr12	11090852	11091806	TAS2R14	-	loss
chr12	11138511	11139511	TAS2R50	-	loss
chr12	11148560	11150474	TAS2R20	-	loss
chr12	11174270	11175170	TAS2R19	-	loss
chr12	11182985	11184006	TAS2R31	-	loss
chr12	11213963	11214893	TAS2R46	-	loss
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chr12	11338598	11339543	TAS2R42	-	loss
chr12	11418856	11422641	PRB3	-	loss
chr12	11504756	11508524	PRB1	-	loss
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chr12	12268960	12419811	LRP6	-	loss
chr12	12482217	12503169	MANSC1	-	loss
chr12	12508341	12510001	LOH12CR2	-	loss
chr12	12626215	12715448	DUSP16	-	loss
chr12	12813994	12849121	GPR19	-	loss
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chr12	13127798	13153243	HEBP1	-	loss
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chr12	13523604	13529679	C12orf36	-	loss
chr12	13714409	14133022	GRIN2B	-	loss
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chr12	14765567	14849519	GUCY2C	-	loss
chr12	14923653	14924065	HIST4H4	-	loss
chr12	14939411	14956401	WBP11	-	loss
chr12	14957583	14967116	SMCO3	-	loss
chr12	14982244	14996413	ART4	-	loss
chr12	15034114	15038853	MGP	-	loss
chr12	15066975	15091463	ERP27	-	loss
chr12	15094949	15114562	ARHGDIB	-	loss
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chr13	36920567	36943872	SPG20OS	+	loss
chr13	37005656	37017019	CCNA1	+	loss

chr13	37248048	37271975	SERTM1	+	loss
chr13	37393338	37403740	RFXAP	+	loss
chr13	37574677	37583751	EXOSC8	+	loss
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chr13	38923941	38937143	UFM1	+	loss
chr13	39141860	39153653	LINC00366	+	loss
chr13	39261172	39461267	FREM2	+	loss
chr13	39612447	39624244	NHLRC3	+	loss
chr13	40229763	40365802	COG6	+	loss
chr13	40755945	40763167	LINC00332	+	loss
chr13	41363546	41386596	SLC25A15	+	loss
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chr13	41635696	41658139	WBP4	+	loss
chr13	41885340	41951166	NAA16	+	loss
chr13	42016699	42017902	OR7E37P	+	loss
chr13	42031541	42045013	RGCC	+	loss
chr13	42535304	42555701	VWA8-AS1	+	loss
chr13	42614171	42830716	DGKH	+	loss
chr13	42846288	42897403	AKAP11	+	loss
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chr13	44947977	44971850	SERP2	+	loss
chr13	45563686	45607743	GPALPP1	+	loss
chr13	45694630	45858239	GTF2F2	+	loss
chr13	45915479	45965618	TPT1-AS1	+	loss
chr13	46039029	46110833	COG3	+	loss
chr13	46276445	46288693	SPERT	+	loss
chr13	46626982	46675482	CPB2-AS1	+	loss
chr13	47127295	47327175	LRCH1	+	loss
chr13	48611702	48621282	NUDT15	+	loss
chr13	48651272	48654129	MED4-AS1	+	loss
chr13	48807273	48836232	ITM2B	+	loss
chr13	48877882	49056026	RB1	+	loss
chr13	49280952	49283498	CYSLTR2	+	loss
chr13	49550047	49783915	FND3A	+	loss
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chr13	50018428	50069139	SETDB2	+	loss
chr13	50069800	50103117	PHF11	+	loss
chr13	50202434	50208008	ARL11	+	loss
chr13	50464544	50467516	CTAGE10P	+	loss
chr13	50571142	50592603	TRIM13	+	loss
chr13	50589389	50595058	KCNRG	+	loss
chr13	50656413	51102670	DLEU1	+	loss
chr13	50746153	50747751	ST13P4	+	loss
chr13	51381990	51424041	DLEU7-AS1	+	loss
chr13	51483813	51544596	RNASEH2B	+	loss

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chr13	51915167	51938418	SERPINE3	+	loss
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chr13	52158483	52336171	WDFY2	+	loss
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chr13	53029494	53050763	CKAP2	+	loss
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chr13	53191604	53217919	HNRNPA1L2	+	loss
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chr13	55748588	55748683	MIR5007	+	loss
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chr13	57741330	57744352	PRR20E	+	loss
chr13	58205788	58303065	PCDH17	+	loss
chr13	60586884	60618490	DIAPH3-AS1	+	loss
chr13	60718831	60727639	DIAPH3-AS2	+	loss
chr13	60970590	61148013	TDRD3	+	loss
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chr13	67399300	67489163	PCDH9-AS2	+	loss
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chr13	71589272	71742549	LINC00348	+	loss
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chr13	77502584	77503224	BTF3P11	+	loss
chr13	77522693	77532776	IRG1	+	loss
chr13	77566058	77576652	CLN5	+	loss
chr13	77649648	77661911	MYCBP2-AS1	+	loss
chr13	78109808	78219398	SCEL	+	loss
chr13	78271988	78338377	SLAIN1	+	loss
chr13	78493823	79191460	RNF219-AS1	+	loss
chr13	78587017	78627730	LINC00446	+	loss
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chr13	36742344	36788752	SOHLH2	-	loss
chr13	36742344	36871992	CCDC169-SOHLH	-	loss
chr13	36801178	36871992	CCDC169	-	loss
chr13	36875774	36944317	SPG20	-	loss
chr13	37418967	37494409	SMAD9	-	loss

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chr13	37677396	37679801	CSNK1A1L	-	loss
chr13	38136718	38183563	POSTN	-	loss
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chr13	38624953	38717369	LINC00571	-	loss
chr13	39540061	39564996	STOML3	-	loss
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chr13	41048130	41240734	FOXO1	-	loss
chr13	41301963	41302011	MIR320D1	-	loss
chr13	41303431	41345347	MRPS31	-	loss
chr13	41371120	41495910	TPTE2P5	-	loss
chr13	41486024	41495910	SUGT1P3	-	loss
chr13	41506054	41593508	ELF1	-	loss
chr13	41701708	41706936	KBTD6	-	loss
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chr13	41790515	41837742	MTRF1	-	loss
chr13	42140960	42535221	VWA8	-	loss
chr13	42142421	42142531	MIR5006	-	loss
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chr13	44410488	44453826	CCDC122	-	loss
chr13	44720605	44732358	SMIM2-IT1	-	loss
chr13	45006278	45150701	TSC22D1	-	loss
chr13	45373639	45383766	LINC00330	-	loss
chr13	45513383	45563613	NUFIP1	-	loss
chr13	45766987	45775175	KCTD4	-	loss
chr13	45911303	45915297	TPT1	-	loss
chr13	45911614	45911744	SNORA31	-	loss
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chr13	46115431	46189874	ERICH6B	-	loss
chr13	46354415	46425846	SIAH3	-	loss
chr13	46529804	46626894	ZC3H13	-	loss
chr13	46627321	46679211	CPB2	-	loss
chr13	46700057	46756459	LCP1	-	loss
chr13	46870579	46871979	LINC00563	-	loss
chr13	46916136	47012325	KIAA0226L	-	loss
chr13	47345390	47371367	ESD	-	loss
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chr13	48516790	48612125	SUCLA2	-	loss
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chr13	49882785	50018221	CAB39L	-	loss

chr13	50106081	50159719	RCBTB1	-	loss
chr13	50234809	50265623	EBPL	-	loss
chr13	50273442	50367057	KPNA3	-	loss
chr13	50486841	50510625	SPRYD7	-	loss
chr13	50556687	50699677	DLEU2	-	loss
chr13	50570550	50570637	MIR3613	-	loss
chr13	50623108	50623337	MIR16-1	-	loss
chr13	51286758	51417885	DLEU7	-	loss
chr13	51456514	51484848	RNASEH2B-AS1	-	loss
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chr13	51658150	51746524	LINC00371	-	loss
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chr13	52741844	52865680	MRPS31P5	-	loss
chr13	52951302	52980629	THSD1	-	loss
chr13	52986736	53024763	VPS36	-	loss
chr13	53277399	53313947	LECT1	-	loss
chr13	53418108	53422775	PCDH8	-	loss
chr13	54886106	54886183	MIR1297	-	loss
chr13	60239720	60738119	DIAPH3	-	loss
chr13	61773931	61774014	MIR3169	-	loss
chr13	61983818	62002079	PCDH20	-	loss
chr13	66876965	67804468	PCDH9	-	loss
chr13	69435416	69459457	LINC00550	-	loss
chr13	70274724	70682625	KLHL1	-	loss
chr13	72012097	72441330	DACH1	-	loss
chr13	73282494	73301938	MZT1	-	loss
chr13	73329539	73356344	DIS3	-	loss
chr13	74260148	74708394	KLF12	-	loss
chr13	75811888	75814517	CTAGE11P	-	loss
chr13	75858808	76056250	TBC1D4	-	loss
chr13	76099349	76111991	COMMD6	-	loss
chr13	77454303	77460540	KCTD12	-	loss
chr13	77579388	77601331	FBXL3	-	loss
chr13	77618791	77901177	MYCBP2	-	loss
chr13	78272146	78272251	MIR3665	-	loss
chr13	78469615	78549664	EDNRB	-	loss
chr13	79173229	79177695	POU4F1	-	loss
chr13	79188420	79234700	RNF219	-	loss
chr13	79361453	79414185	LINC00331	-	loss
chr16	74330672	74340186	PSMD7	+	loss
chr16	75032914	75144892	ZNRF1	+	loss
chr16	75182420	75206132	ZFP1	+	loss
chr16	75252883	75258822	CTRB1	+	loss
chr16	75600248	75611779	GABARAPL2	+	loss

chr16	75681634	75691341	TERF2IP	+	loss
chr16	76311175	76593135	CNTNAP4	+	loss
chr16	76902832	76902916	MIR4719	+	loss
chr16	77224835	77233543	MON1B	+	loss
chr16	77233348	77246976	SYCE1L	+	loss
chr16	77756388	77776157	NUDT7	+	loss
chr16	77822482	78014001	VAT1L	+	loss
chr16	78056442	78066001	CLEC3A	+	loss
chr16	78133326	79246564	WVOX	+	loss
chr16	80574853	80584541	DYNLRB2	+	loss
chr16	81040102	81066709	CENPN	+	loss
chr16	81069457	81080951	ATMIN	+	loss
chr16	81272295	81324747	BCO1	+	loss
chr16	81348570	81413803	GAN	+	loss
chr16	81418622	81418698	MIR4720	+	loss
chr16	81478774	81745367	CMIP	+	loss
chr16	81698958	81700879	LOC100129617	+	loss
chr16	81812898	81991899	PLCG2	+	loss
chr16	82068857	82132139	HSD17B2	+	loss
chr16	82660398	83830215	CDH13	+	loss
chr16	83541950	83542013	MIR3182	+	loss
chr16	83841507	83846607	HSBP1	+	loss
chr16	83932729	83949787	MLYCD	+	loss
chr16	83982671	83999937	OSGIN1	+	loss
chr16	84002236	84036379	NECAB2	+	loss
chr16	84178864	84211524	DNAAF1	+	loss
chr16	84224722	84230772	ADAD2	+	loss
chr16	84328400	84363450	WFDC1	+	loss
chr16	84402132	84497793	ATP2C2	+	loss
chr16	84682130	84695916	KLHL36	+	loss
chr16	84733554	84813527	USP10	+	loss
chr16	84853586	84943116	CRISPLD2	+	loss
chr16	85061374	85127828	KIAA0513	+	loss
chr16	85170755	85183049	LOC400548	+	loss
chr16	85316563	85321685	LINC00311	+	loss
chr16	85645028	85709812	GSE1	+	loss
chr16	85833172	85840607	COX4I1	+	loss
chr16	85932773	85956211	IRF8	+	loss
chr16	86320036	86326995	LOC146513	+	loss
chr16	86544132	86548070	FOXF1	+	loss
chr16	86588925	86590905	FLJ30679	+	loss
chr16	86600856	86602537	FOXC2	+	loss
chr16	86612114	86615304	FOXL1	+	loss
chr16	87425800	87438380	MAP1LC3B	+	loss
chr16	87635440	87731761	JPH3	+	loss
chr16	87984237	88110924	BANP	+	loss
chr16	88493878	88507165	ZNF469	+	loss
chr16	88520013	88601574	ZFPM1	+	loss
chr16	88535325	88535439	MIR5189	+	loss
chr16	88636788	88698372	ZC3H18	+	loss

chr16	88705000	88706882	IL17C	+	loss
chr16	88729780	88753594	SNAI3-AS1	+	loss
chr16	88772890	88781786	CTU2	+	loss
chr16	88870185	88875666	CDT1	+	loss
chr16	88922627	88927591	TRAPPC2L	+	loss
chr16	89160216	89222171	ACSF3	+	loss
chr16	89225627	89230083	LINC00304	+	loss
chr16	89232790	89235555	LOC400558	+	loss
chr16	89238162	89261900	CDH15	+	loss
chr16	89284110	89295965	ZNF778	+	loss
chr16	89387540	89391518	LOC100287036	+	loss
chr16	89574804	89624174	SPG7	+	loss
chr16	89627064	89633237	RPL13	+	loss
chr16	89627837	89627909	SNORD68	+	loss
chr16	89642175	89663654	CPNE7	+	loss
chr16	89679715	89704839	DPEP1	+	loss
chr16	89724151	89736866	SPATA33	+	loss
chr16	89753075	89762772	CDK10	+	loss
chr16	89778263	89784573	VPS9D1-AS1	+	loss
chr16	89786807	89807332	ZNF276	+	loss
chr16	89894906	89937727	SPIRE2	+	loss
chr16	89939993	89977792	TCF25	+	loss
chr16	89984286	89987385	MC1R	+	loss
chr16	89985258	90002505	TUBB3	+	loss
chr16	90015138	90034468	DEF8	+	loss
chr16	90038987	90067195	AFG3L1P	+	loss
chr16	90086068	90111379	GAS8	+	loss
chr16	74366303	74402153	LOC283922	-	loss
chr16	74442528	74455649	CLEC18B	-	loss
chr16	74481325	74641042	GLG1	-	loss
chr16	74907470	75019017	WDR59	-	loss
chr16	75145757	75150670	LDHD	-	loss
chr16	75237993	75241072	CTRB2	-	loss
chr16	75262927	75301951	BCAR1	-	loss
chr16	75327607	75467387	CFDP1	-	loss
chr16	75477141	75498584	TMEM170A	-	loss
chr16	75507021	75528926	CHST6	-	loss
chr16	75562427	75579393	CHST5	-	loss
chr16	75572014	75590184	TMEM231	-	loss
chr16	75632246	75657221	ADAT1	-	loss
chr16	75661621	75682541	KARS	-	loss
chr16	77316024	77469011	ADAMTS18	-	loss
chr16	79627744	79634622	MAF	-	loss
chr16	80637675	80838175	CDYL2	-	loss
chr16	81009698	81040502	CMC2	-	loss
chr16	81087101	81110872	C16orf46	-	loss
chr16	81115551	81129980	GCSH	-	loss
chr16	81134483	81253975	PKD1L2	-	loss
chr16	82031250	82045093	SDR42E1	-	loss
chr16	82181766	82203829	MPHOSPH6	-	loss

chr16	84043388	84075762	SLC38A8	-	loss
chr16	84087368	84150517	MBTPS1	-	loss
chr16	84155743	84178800	HSDL1	-	loss
chr16	84211452	84220676	TAF1C	-	loss
chr16	84254740	84273356	KCNG4	-	loss
chr16	84509965	84538288	TLDC1	-	loss
chr16	84599203	84651669	COTL1	-	loss
chr16	85008066	85045141	ZDHHC7	-	loss
chr16	85131964	85146114	FAM92B	-	loss
chr16	85339831	85339931	MIR5093	-	loss
chr16	85711279	85722588	GINS2	-	loss
chr16	85741123	85784689	C16orf74	-	loss
chr16	85775226	85775306	MIR1910	-	loss
chr16	85812230	85833148	EMC8	-	loss
chr16	86365455	86379285	LINC00917	-	loss
chr16	86508130	86542466	FENDRR	-	loss
chr16	86563781	86588841	MTHFSD	-	loss
chr16	87336403	87351026	C16orf95	-	loss
chr16	87362941	87425708	FBXO31	-	loss
chr16	87439851	87525460	ZCCHC14	-	loss
chr16	87736728	87799598	KLHDC4	-	loss
chr16	87863628	87903100	SLC7A5	-	loss
chr16	87921624	87970112	CA5A	-	loss
chr16	88709696	88717492	CYBA	-	loss
chr16	88718347	88729495	MVD	-	loss
chr16	88744089	88752882	SNAI3	-	loss
chr16	88762902	88772829	RNF166	-	loss
chr16	88781745	88851372	PIEZO1	-	loss
chr16	88782685	88782745	MIR4722	-	loss
chr16	88875876	88878342	APRT	-	loss
chr16	88880141	88923374	GALNS	-	loss
chr16	88928035	88933068	PABPN1L	-	loss
chr16	88941262	89043504	CBFA2T3	-	loss
chr16	89262168	89266529	SLC22A31	-	loss
chr16	89334028	89556969	ANKRD11	-	loss
chr16	89710838	89724193	CHMP1A	-	loss
chr16	89762764	89768121	SPATA2L	-	loss
chr16	89773540	89787394	VPS9D1	-	loss
chr16	89803958	89883065	FANCA	-	loss
chr16	90036182	90039240	CENPBD1	-	loss
chr16	90071278	90086328	DBNDD1	-	loss
chr16	90095315	90096309	C16orf3	-	loss
chr16	90106168	90114191	URAHP	-	loss
chr16	90122973	90142338	PRDM7	-	loss
chr21	40177230	40196878	ETS2	+	loss
chr21	40687632	40695144	BRWD1-AS1	+	loss
chr21	40752212	40786214	WRB	+	loss
chr21	40817796	40887433	SH3BGR	+	loss
chr21	40928368	41034815	B3GALT5	+	loss
chr21	41117333	41174023	IGSF5	+	loss

chr21	41239346	41301322	PCP4	+	loss
chr21	41755009	41757285	DSCAM-AS1	+	loss
chr21	42539483	42539556	MIR3197	+	loss
chr21	42539727	42648524	BACE2	+	loss
chr21	42676176	42729654	FAM3B	+	loss
chr21	42733949	42780869	MX2	+	loss
chr21	42792519	42831141	MX1	+	loss
chr21	39739182	40033704	ERG	-	loss
chr21	40110878	40145401	LINC00114	-	loss
chr21	40547371	40555440	PSMG1	-	loss
chr21	40557403	40685712	BRWD1	-	loss
chr21	40714240	40721047	HMGN1	-	loss
chr21	40777769	40817670	LCA5L	-	loss
chr21	40969074	40984749	C21orf88	-	loss
chr21	41384342	42219039	DSCAM	-	loss
chr21	41584278	41584358	MIR4760	-	loss
chr21	41990432	42002693	DSCAM-IT1	-	loss
chr21	42513426	42519991	LINC00323	-	loss
chr21	42547157	42557166	PLAC4	-	loss
chrY	14774297	14804153	TTY15	+	loss
chrY	14813159	14972768	USP9Y	+	loss
chrY	15016018	15032390	DDX3Y	+	loss
chrY	15815446	15817902	TMSB4Y	+	loss
chrY	16634487	16955848	NLGN4Y	+	loss
chrY	21729235	21769056	TXLNGY	+	loss
chrY	22737610	22755040	EIF1AY	+	loss
chrY	22917953	22942918	RPS4Y2	+	loss
chrY	24455005	24462352	RBMY2FP	+	loss
chrY	24455005	24564028	RBMY1F	+	loss
chrY	24636543	24660784	PRY	+	loss
chrY	25365580	27053187	DAZ2	+	loss
chrY	14517914	14533389	GYG2P1	-	loss
chrY	15346460	15592550	UTY	-	loss
chrY	16905521	16915913	NLGN4Y-AS1	-	loss
chrY	21094584	21239302	TTY14	-	loss
chrY	21152525	21154705	CD24	-	loss
chrY	21617316	21665039	BCORP1	-	loss
chrY	21867300	21906825	KDM5D	-	loss
chrY	22627553	22681114	TTY10	-	loss
chrY	23557033	23563448	RBMY2EP	-	loss
chrY	23745485	23756552	TTY13	-	loss
chrY	24026222	24329089	RBMY1E	-	loss
chrY	24217902	24242154	PRY2	-	loss
chrY	24442944	24445023	TTY5	-	loss
chrY	25275501	25345254	DAZ1	-	loss

oSNP)

Supplementary Table 5. Top 100 genes with the most highly variable inter-tumour expression

Gene ID	Ensembl ID	chromosome
ABCC4	ENSG00000125257	chr13q32.1
ACTC1	ENSG00000159251	chr15q14
ADIRF	ENSG00000148671	chr10q23.2
AEBP1	ENSG00000106624	chr7p13
ALOX15B	ENSG00000179593	chr17p13.1
AMACR	ENSG00000242110	chr5p13.2
ANGPTL4	ENSG00000167772	chr19p13.2
ANPEP	ENSG00000166825	chr15q26.1
AOC1	ENSG00000002726	chr7q36.1
ATF3	ENSG00000162772	chr1q32.3
C11orf96	ENSG00000187479	chr11p11.2
C15orf48	ENSG00000166920	chr15q21.1
C1orf64	ENSG00000183888	chr1p36.13
CCL2	ENSG00000108691	chr17q12
CES1	ENSG00000262243	chr16q12.2
CFB	ENSG00000239754	chr6p21.33
CLDN1	ENSG00000163347	chr3q28
CLDN8	ENSG00000156284	chr21q22.11
CNTNAP2	ENSG00000174469	chr7q35
COL1A1	ENSG00000108821	chr17q21.33
COL6A2	ENSG00000142173	chr21q22.3
COL8A1	ENSG00000144810	chr3q12.1
COL9A2	ENSG00000049089	chr1p34.2
COMP	ENSG00000105664	chr19p13.11
CYP27A1	ENSG00000135929	chr2q35
CYP2J2	ENSG00000134716	chr1p32.1
CYR61	ENSG00000142871	chr1p22.3
DLX1	ENSG00000144355	chr2q31.1
DNASE2B	ENSG00000137976	chr1p31.1
EEF1A2	ENSG00000101210	chr20q13.33
EGR1	ENSG00000120738	chr5q31.2
ERAP2	ENSG00000164308	chr5q15
FAM3D	ENSG00000198643	chr3p14.2
FBLN1	ENSG00000077942	chr22q13.31
FOS	ENSG00000170345	chr14q24.3
FOSB	ENSG00000125740	chr19q13.32
GDF15	ENSG00000130513	chr19p13.11
GNMT	ENSG00000124713	chr6p21.1
GP2	ENSG00000169347	chr16p12.3
GSTT1	ENSG00000277656	chr22q11.23
H19	ENSG00000130600	chr11p15.5
HMGCS2	ENSG00000134240	chr1p12
HSD17B6	ENSG00000025423	chr12q13.3
IGFBP6	ENSG00000167779	chr12q13.13
IGJ	ENSG00000132465	chr4q13.3
KCNN2	ENSG00000080709	chr5q22.3
KCNN4	ENSG00000104783	chr19q13.31
KLK12	ENSG00000186474	chr19q13.41

KRT15	ENSG00000171346	chr17q21.2
LTBP4	ENSG00000090006	chr19q13.2
MARCH6	ENSG00000145495	chr5p15.2
MESP1	ENSG00000166823	chr15q26.1
MIPEP	ENSG00000027001	chr13q12.12
MME	ENSG00000196549	chr3q25.2
MMP7	ENSG00000137673	chr11q22.2
MS4A8	ENSG00000166959	chr11q12.2
MSMB	ENSG00000263639	chr10q11.22
MT1G	ENSG00000125144	chr16q13
MT1M	ENSG00000205364	chr16q13
MUC1	ENSG00000185499	chr1q22
MYBPC1	ENSG00000196091	chr12q23.2
NCAPD3	ENSG00000151503	chr11q25
NFKBIZ	ENSG00000144802	chr3q12.3
OLFM4	ENSG00000102837	chr13q14.3
OR51E1	ENSG00000180785	chr11p15.4
ORM2	ENSG00000228278	chr9q32
PAGE4	ENSG00000101951	chrXp11.23
PCA3	ENSG00000225937	chr9q21.2
PCAT4	ENSG00000251321	chr4q21.21
PCGEM1	ENSG00000227418	chr2q32.3
PDE8B	ENSG00000113231	chr5q13.3
PGC	ENSG00000096088	chr6p21.1
PI16	ENSG00000164530	chr6p21.2
PIGR	ENSG00000162896	chr1q32.1
PLA2G2A	ENSG00000188257	chr1p36.13
PLA2G7	ENSG00000146070	chr6p12.3
PLAC9	ENSG00000189129	chr10q22.3
PPAP2A	ENSG00000067113	chr5q11.2
PTGS2	ENSG00000073756	chr1q31.1
RLN1	ENSG00000107018	chr9p24.1
RPL14	ENSG00000188846	chr3p22.1
RPL17	ENSG00000265681	chr18q21.1
S100P	ENSG00000163993	chr4p16.1
SCIN	ENSG00000006747	chr7p21.3
SCUBE2	ENSG00000175356	chr11p15.4
SERPINA3	ENSG00000196136	chr14q32.13
SNAP91	ENSG00000065609	chr6q14.2
SORD	ENSG00000140263	chr15q21.1
SPON2	ENSG00000159674	chr4p16.3
STXBP6	ENSG00000168952	chr14q12
TFF3	ENSG00000160180	chr21q22.3
TMEFF2	ENSG00000144339	chr2q32.3
TMEM178A	ENSG00000152154	chr2p22.1
TNFRSF19	ENSG00000127863	chr13q12.12
TUBB2A	ENSG00000137267	chr6p25.2
VSIG2	ENSG00000019102	chr11q24.2
ZFP36	ENSG00000128016	chr19q13.2
ZNF385B	ENSG00000144331	chr2q31.3

LEPREL1	ENSG00000090530	chr3q29
PDIA3P	ENSG00000180867	chr1q21.1

Supplementary Table 6. Discriminating 100 gene set, derived from gene expression and copy number feature data*

Chromosome	start position^	end position	Associated gene	% CNAs in cohort
chr8p21.3	23119071	23119120	CHMP7	37%
chr8p21.3	23153663	23153712	R3HCC1	37%
chr8p21.3	22432879	22432928	SORBS3	36%
chr8p21.3	22477693	22477742	CCAR2	36%
chr8p21.3	22461341	22461390	C8orf58	36%
chr8p21.3	21961474	21961523	FAM160B2	36%
chr8p21.3	21939833	21939882	DMTN	36%
chr8p21.3	22398199	22398248	PPP3CC	36%
chr8p21.3	20078945	20078994	ATP6V1B2	35%
chr8p23.1	11185430	11185479	MTMR9	35%
chr8p23.1	8890610	8890659	ERI1	34%
chr8p21.2	27316655	27316704	PTK2B	34%
chr8p12	30436174	30436223	GTF2E2	33%
chr8p21.1	28048518	28048567	ELP3	33%
chr8p21.1	28625279	28625328	INTS9	33%
chr8p11.22	38846564	38846613	TM2D2	25%
chr8p11.21	41788540	41788589	KAT6A	25%
chr8p11.23	38020937	38020986	LSM1	25%
chr8p11.23	37701676	37701725	BRF2	25%
chr13q14.2	49063443	49063492	RCBTB2	21%
chr13q14.13	45858012	45858061	GTF2F2	20%
chr13q14.2	50065906	50065955	SETDB2	20%
chr13q14.2	50591839	50591888	TRIM13	20%
chr13q14.2	50102892	50102941	PHF11	20%
chr13q14.3	53262319	53262368	SUGT1	19%
chr13q14.11	42897015	42897064	AKAP11	18%
chr16q24.1	85840509	85840558	COX4I1	16%
chr16q24.2	87741499	87741548	KLHDC4	16%
chr6q15	88228353	88228402	RARS2	15%
chr16q24.3	89977633	89977682	TCF25	15%
chr16q24.3	88927406	88927455	TRAPPC2L	15%
chr16q24.3	90034280	90034329	DEF8	15%
chr16q23.3	83846402	83846451	HSBP1	15%
chr6q15	89320073	89320122	RNGTT	14%
chr6q15	91225641	91225690	MAP3K7	13%
chr8q21.3	90802376	90802425	RIPK2	13%
chr8q24.13	124232447	124232496	C8orf76	13%
chr8q24.13	126379143	126379192	NSMCE2	13%
chr6q21	110146366	110146415	FIG4	12%
chr8q11.21	48648011	48648060	SPIDR	12%
chr8q21.13	82570430	82570479	IMPA1	12%
chr8q21.3	87573314	87573363	CPNE3	12%
chr8q22.2	101533288	101533337	ANKRD46	12%
chr8q23.2	110576909	110576958	EBAG9	12%
chr8q21.13	80947140	80947189	TPD52	12%
chr8q21.13	82671480	82671529	CHMP4C	12%
chr8q22.1	97346438	97346487	PTDSS1	12%
chr8q24.3	146199160	146199209	ZNF252P	11%

chr16q23.1	74907697	74907746	WDR59	11%
chr17p13.1	7529756	7529805	SAT2	9%
chr17p13.1	7363017	7363066	ZBTB4	9%
chr17p13.1	7491180	7491229	MPDU1	9%
chr17p13.2	4862717	4862766	SPAG7	9%
chr17p13.2	4840686	4840735	SLC25A11	9%
chr10q23.1	85912689	85912738	GHITM	8%
chr18q23	77806076	77806125	RBFA	8%
chr18q23	77662567	77662616	PQLC1	8%
chr6q22.31	125597068	125597117	HDDC2	7%
chr12p13.31	6840857	6840906	COPS7A	7%
chr16q22.1	70286329	70286378	AARS	7%
chr16q22.1	70514063	70514112	FUK	7%
chr17p13.2	4060604	4060653	CYB5D2	7%
chr17p13.2	3564717	3564766	CTNS	7%
chr17p13.2	4067363	4067412	ANKFY1	7%
chr18q21.1	43846636	43846685	C18orf25	7%
chr17p12	12895998	12896047	ELAC2	7%
chr18q21.1	47808738	47808787	CXXC1	6%
chr18q21.33	59711985	59712034	PIGN	6%
chr7q32.1	127231472	127231521	FSCN3	6%
chr7q32.1	127231472	127231521	ARF5	6%
chr18q21.1	44633811	44633860	HDHD2	6%
chr17p12	14111849	14111898	COX10	6%
chr10q24.32	104183242	104183291	CUEDC2	5%
chr10q24.1	99218137	99218186	MMS19	5%
chr18p11.22	9537980	9538029	RALBP1	5%
chr18p11.21	12329202	12329251	AFG3L2	5%
chr2q14.3	128015187	128015236	ERCC3	4%
chr2q14.3	128461058	128461107	SFT2D3	4%
chr2q24.1	158593153	158593202	ACVR1	4%
chr3q27.1	184052784	184052833	EIF4G1	4%
chr3q27.1	183530057	183530106	YEATS2	4%
chr13q34	115038036	115038085	CDC16	3%
chr3q26.2	169557364	169557413	LRRC31	3%
chr1p21.2	101007343	101007392	GPR88	2%
chr3q22.1	131083043	131083092	NUDT16P1	2%
chr9q33.3	129647594	129647643	ZBTB34	2%
chr9q33.3	128321925	128321974	MAPKAP1	2%
chr3q22.1	129023910	129023959	HMCES	2%
chr3q22.1	128631728	128631777	ACAD9	2%
chr12q13.3	57924185	57924234	DCTN2	2%
chr20q13.33	61962106	61962155	COL20A1	1%
chr2p21	47747953	47748002	KCNK12	1%
chr9q22.33	101611248	101611297	GALNT12	1%
chr9q31.1	102648734	102648783	RP11-60I3.5	1%
chr9p13.2	36677336	36677385	MELK	1%
chr9q31.1	104087030	104087079	LPPR1	1%
chr12q12	46316836	46316885	SCAF11	1%
chr20q11.21	30422380	30422429	MYLK2	1%

chr17q23.2	62016057	62016106	SCN4A	1%
chr12q12	46119857	46119906	LINC00938	1%

^ mapped to hg19

* Only OncoSNP (Yau et al., 2010) calls rank1+2 considered - see Methods.

Supplementary Table 7. Measures of cluster accuracy, compared to integrative clustering

	iCluster vs Gx alone (T variability)	iCluster vs Gx (fold change)	iCluster vs CN alone	Gx (T variability) vs Gx (fold change)
Adjusted Rand Index (Hubert 1985)	0.13	0.08	0.35	0.47
Variation of Information Index (Meilă 2007)	2.4	2.7	1.7	1.5

Gx = gene expression

CN = copy number

iCluster = integrative clustering, performed in iCluster+ (Mo et al., 2013).

T variability - tumour variability (see Gorlov et al. (2012) and Gorlov et al. (2014))

Supplementary Table 8. Summary of clinical characteristics of 2nd validation cohort (from Taylor et al., 2010).

	Primary tumour		Metastases	
	n = 181	%	n = 37	%
Age (years)				
Mean	60.9		60	
Range	37.3 - 83		41-82	
Pre-operative PSA (ng/ml)				
< 4	31	25%	4	13%
4 - 10	105	84%	6	19%
> 10	44	35%	22	69%
Biopsy Gleason Grade				
5	2	1%	-	-
6	101	56%	2	6%
7	61	34%	16	46%
8	11	6%	8	23%
9	6	3%	9	25%
Clinical stage				
cT1c	95	52%	8	22%
cT2c	76	42%	12	33%
cT3	9	5%	3	8%
pT4	-	-	1	3%
not available	-	-	9	25%
Ethnicity				
Black	29	16%	2	6%
Asian	4	2%	-	-
White Hispanic	-	0%	2	6%
White non-Hispanic	142	79%	32	89%
unknown	6	3%	-	-

*Only samples with aCGH and mRNA were used in this analysis: 109 primary tumours and 19 metastases.

Genes.SYMBOL	Coef.clust1 - Benign	t.clust1 - Benign	p.value.adj.clust1 - Benign	Genes.IlluminaID
AMACR	2.598	11.51	0	ILMN_1759670
HPN	2.191	14.38	0	ILMN_2358760
HPN	2.175	14.13	0	ILMN_1687235
PCA3	2.142	9.97	0	ILMN_3239648
AGR2	2.053	8.15	0	ILMN_3307841
PLA2G7	1.942	9.23	0	ILMN_1701195
ABCC4	1.931	9.58	0	ILMN_2194009
DLX1	1.832	12.24	0	ILMN_1672094
TRPM4	1.781	10.06	0	ILMN_1679401
SPON2	1.766	6.74	0	ILMN_1676099
ABCC4	1.689	8.93	0	ILMN_1788457
GDF15	1.591	5.29	4.00E-06	ILMN_2188862
TSPAN13	1.554	11.12	0	ILMN_1669881
THBS4	1.527	7.17	0	ILMN_1736078
RAP1GAP	1.5	11.04	0	ILMN_1776519
LEPREL1	-1.507	-5.76	0	ILMN_1657373
SMTN	-1.524	-8.75	0	ILMN_1785618
TCEAL2	-1.529	-10.94	0	ILMN_1765310
BNIP1	-1.53	-8.72	0	ILMN_1699989
PARM1	-1.532	-11.61	0	ILMN_1656560
ZNF827	-1.548	-9.72	0	ILMN_1727574
SCGB3A1	-1.55	-7.79	0	ILMN_1679666
PDK4	-1.566	-8.08	0	ILMN_1684982
SLC2A5	-1.611	-8.46	0	ILMN_1671337
PCP4	-1.615	-8.76	0	ILMN_1682326
FOS	-1.619	-5.05	1.00E-05	ILMN_1669523
KIAA1210	-1.628	-8.33	0	ILMN_1732962
COL17A1	-1.641	-10.68	0	ILMN_1651282
GCNT2	-1.67	-9.22	0	ILMN_1680390
FLRT3	-1.686	-10.56	0	ILMN_1805665
KIAA0353	-1.697	-10.38	0	ILMN_1712075
ANPEP	-1.736	-4.98	1.30E-05	ILMN_1763837
DES	-1.741	-9.68	0	ILMN_1698995
WFDC2	-1.817	-10.51	0	ILMN_3307693
OLFM4	-1.875	-3.96	0.000619	ILMN_2116877
MSMB	-1.884	-5.87	0	ILMN_2376313
MME	-1.904	-7.65	0	ILMN_1678170
MSMB	-1.917	-6.13	0	ILMN_1790906
PRDM8	-2.037	-9.68	0	ILMN_1802082
KRT15	-2.079	-10.15	0	ILMN_1770612
TP63	-2.259	-11.43	0	ILMN_3305055

Genes.SYMBOL	Coef.clust2 - Benign	t.clust2 - Benign	p.value.adj.clust2 - Benign	Genes.IlluminaID
HPN	1.637	10.98		0 ILMN_1687235
PCA3	1.567	7.53		0 ILMN_3239648
HPN	1.556	10.54		0 ILMN_2358760

Genes.SYMBOL	Coef.clust3 - Benign	t.clust3 - Benign	p.value.adj.clust3 - Benign	Genes.IlluminaID
KLK12	3.189	9.89	0	ILMN_1725912
AMACR	2.879	8.63	0	ILMN_1759670
SPON2	2.642	6.82	0	ILMN_1676099
FGFRL1	2.267	13.52	0	ILMN_1795865
TRPM4	2.221	8.49	0	ILMN_1679401
HPN	2.115	9.3	0	ILMN_1687235
CYP2J2	2.113	7.26	0	ILMN_1758731
GDF15	2.11	4.75	6.40E-05	ILMN_2188862
UNC5A	2.107	7.84	0	ILMN_1712913
HPN	2.033	9.03	0	ILMN_2358760
DLX1	2.016	9.11	0	ILMN_1672094
MYL2	2.009	3.92	0.001179	ILMN_2113807
PCA3	1.918	6.04	0	ILMN_3239648
MS4A8	1.913	7.28	0	ILMN_1808249
TRIB1	1.885	6.49	0	ILMN_1803811
MYL1	1.879	4.25	0.000392	ILMN_1700860
MYH7	1.833	3.91	0.001212	ILMN_1669714
KLK12	1.816	8.93	0	ILMN_2411511
TMSB15A	1.794	5.31	7.00E-06	ILMN_1681737
TFF3	1.718	2.97	0.018336	ILMN_1811387
TSPAN13	1.717	8.31	0	ILMN_1669881
CLDN8	1.717	6.21	0	ILMN_2072568
C15orf48	1.684	3.19	0.010379	ILMN_1805410
MB	1.664	4.96	2.90E-05	ILMN_1666109
HOXC4	1.657	7.89	0	ILMN_1719975
RAB17	1.65	8.56	0	ILMN_2052373
ELF5	1.614	8.42	0	ILMN_1813270
ABCC4	1.59	5.69	1.00E-06	ILMN_1788457
ABCC4	1.569	5.27	8.00E-06	ILMN_2194009
FLJ22184	1.554	8	0	ILMN_2387471
TSPAN13	1.534	8.25	0	ILMN_2130525
GNMT	1.53	4.03	0.000811	ILMN_1736238
CLDN8	1.525	6.53	0	ILMN_1746676
SLN	1.503	3.67	0.002579	ILMN_1778595
JAM3	-1.51	-7.46	0	ILMN_1769575
EDNRA	-1.513	-7.99	0	ILMN_1796629
LIMS2	-1.515	-8.06	0	ILMN_1760493
KRT13	-1.516	-3.11	0.012789	ILMN_1721218
ITGA5	-1.522	-5.53	3.00E-06	ILMN_1792679
MSRB3	-1.525	-8.08	0	ILMN_1676088
DPYSL3	-1.528	-8.35	0	ILMN_1679262
RARRES1	-1.528	-4.76	6.00E-05	ILMN_1800091
TGM4	-1.531	-2.66	0.038728	ILMN_1669363
CALD1	-1.531	-8.51	0	ILMN_1730487
HSPB8	-1.532	-6.91	0	ILMN_1791280
SLC14A1	-1.54	-8.21	0	ILMN_1805561
ALDH1A2	-1.541	-8.03	0	ILMN_1748538
PTGS2	-1.55	-4.58	0.000119	ILMN_2054297
HSPB6	-1.553	-5.85	1.00E-06	ILMN_1721283
TNFRSF19	-1.556	-4.92	3.40E-05	ILMN_1704154

ID1	-1.559	-5.24	9.00E-06	ILMN_1664861
CFB	-1.56	-3.26	0.00843	ILMN_1774287
SCGB3A1	-1.565	-5.32	7.00E-06	ILMN_1679666
LAMB3	-1.57	-8.11	0	ILMN_1715684
CCND2	-1.577	-7.02	0	ILMN_2067656
DLK2	-1.577	-7.88	0	ILMN_1676728
CPED1	-1.577	-8.66	0	ILMN_1677038
ANXA1	-1.58	-6.84	0	ILMN_2184184
C2orf40	-1.587	-5.28	8.00E-06	ILMN_1676822
PAGE4	-1.588	-4.88	3.90E-05	ILMN_1782754
AHNAK2	-1.589	-7.75	0	ILMN_3243156
ROR2	-1.594	-10.63	0	ILMN_1694426
ATP2B4	-1.595	-9.19	0	ILMN_1680579
PDE8B	-1.602	-6.17	0	ILMN_2301722
TGFB111	-1.625	-7.83	0	ILMN_2389876
MYH11	-1.641	-10.54	0	ILMN_1660086
LGALS1	-1.671	-9.33	0	ILMN_1723978
STOM	-1.676	-9.43	0	ILMN_1766657
TPM1	-1.679	-7.33	0	ILMN_2360710
CLIP3	-1.683	-7.12	0	ILMN_1789733
TP63	-1.683	-9.34	0	ILMN_2138801
ISL1	-1.687	-8.34	0	ILMN_1763390
BNIPL	-1.689	-6.51	0	ILMN_1699989
TPM1	-1.694	-7.37	0	ILMN_1716687
GPR124	-1.696	-8.19	0	ILMN_1773059
CCK	-1.707	-5.62	2.00E-06	ILMN_1712943
KRT5	-1.707	-7.45	0	ILMN_1801632
FLNA	-1.709	-6.57	0	ILMN_1687335
AHNAK2	-1.714	-7.72	0	ILMN_1657111
C12orf75	-1.735	-8.36	0	ILMN_1716382
PTRF	-1.746	-9.01	0	ILMN_1757552
TMEM158	-1.759	-6.33	0	ILMN_1792455
FLNC	-1.769	-6.11	0	ILMN_1715748
CFD	-1.771	-5.19	1.10E-05	ILMN_1777190
SORBS1	-1.78	-7.52	0	ILMN_1749792
ATP2B4	-1.799	-9.21	0	ILMN_2367753
SLC2A5	-1.801	-6.4	0	ILMN_1671337
GPX3	-1.809	-8.07	0	ILMN_1726666
FBLN1	-1.814	-6.14	0	ILMN_1700541
VSIG2	-1.82	-4.94	3.10E-05	ILMN_1666536
KCNAB1	-1.824	-6.4	0	ILMN_1744968
LDHB	-1.827	-8.48	0	ILMN_1728132
MEIS2	-1.836	-9.99	0	ILMN_1695945
COL17A1	-1.84	-8.11	0	ILMN_1651282
ADIRF	-1.842	-6.55	0	ILMN_1680110
PDK4	-1.864	-6.51	0	ILMN_1684982
CES1	-1.9	-6.8	0	ILMN_2359945
AOC1	-1.907	-4.57	0.000123	ILMN_1731433
TCEAL2	-1.938	-9.38	0	ILMN_1765310
KCNMB1	-1.944	-8.8	0	ILMN_1652065

CALD1	-2.014	-6.65	0 ILMN_1717990
CSRP1	-2.06	-10.58	0 ILMN_1811921
CAV1	-2.061	-10.35	0 ILMN_2149226
MYL9	-2.111	-9.1	0 ILMN_1675062
MYLK	-2.125	-8.92	0 ILMN_2364768
PNCK	-2.129	-9.57	0 ILMN_1697189
TAGLN	-2.13	-9.2	0 ILMN_2400935
FLRT3	-2.184	-9.26	0 ILMN_1805665
DES	-2.208	-8.31	0 ILMN_1698995
SMTN	-2.212	-8.59	0 ILMN_1785618
CNN1	-2.216	-9.2	0 ILMN_1810054
KIAA1210	-2.232	-7.73	0 ILMN_1732962
PGM5	-2.297	-9.52	0 ILMN_1709590
PGM5	-2.311	-9.7	0 ILMN_1781388
PGM5	-2.396	-9.33	0 ILMN_2271149
PRDM8	-2.403	-7.73	0 ILMN_1802082
KRT15	-2.48	-8.19	0 ILMN_1770612
TP63	-2.554	-8.75	0 ILMN_3305055
WFDC2	-2.555	-10	0 ILMN_3307693
OLFM4	-2.566	-3.67	0.002634 ILMN_2116877
MSMB	-2.578	-6.71	0 ILMN_1699243
MYLK	-2.634	-9.52	0 ILMN_1691476
PCP4	-2.649	-9.73	0 ILMN_1682326
SYNM	-2.65	-10.97	0 ILMN_1712075
MSMB	-3.216	-6.78	0 ILMN_2376313
MSMB	-3.226	-6.98	0 ILMN_1790906

Genes.SYMBOL	Coef.clust4 - Benign	t.clust4 - Benign	p.value.adj.clust4 - Benign	Genes.IlluminaID
TRPM4	2.136	11.2		0 ILMN_1679401
TFF3	1.892	4.49	0.000126	ILMN_1811387
HPN	1.765	10.65		0 ILMN_1687235
FLJ22184	1.712	12.09		0 ILMN_2387471
HPN	1.688	10.29		0 ILMN_2358760
C1orf64	1.63	5.02	1.60E-05	ILMN_2066088
GDF15	1.506	4.65	6.90E-05	ILMN_2188862
PRDM8	-1.51	-6.66		0 ILMN_1802082
MSMB	-1.548	-5.53	2.00E-06	ILMN_1699243
MSMB	-1.674	-4.97	2.00E-05	ILMN_1790906
TP63	-1.698	-7.98		0 ILMN_3305055
OLFM4	-1.742	-3.42	0.004484	ILMN_2116877
MSMB	-1.748	-5.05	1.50E-05	ILMN_2376313

Genes.SYMBOL	Coef.clust5 - Benign	t.clust5 - Benign	p.value.adj.clust5 - Benign	Genes.IlluminaID
AMACR	2.596	12.68		0 ILMN_1759670
HPN	2.131	15.27		0 ILMN_1687235
HPN	2.106	15.25		0 ILMN_2358760
PCA3	2.081	10.68		0 ILMN_3239648
ABCC4	1.924	10.52		0 ILMN_2194009
TRPM4	1.907	11.88		0 ILMN_1679401
AGR2	1.889	8.27		0 ILMN_3307841
C1orf64	1.749	6.4		0 ILMN_2066088
THBS4	1.74	9		0 ILMN_1736078
GDF15	1.642	6.02		0 ILMN_2188862
COMP	1.596	5.66	1.00E-06	ILMN_1677636
FLJ22184	1.513	12.69		0 ILMN_2387471
RAB17	1.508	12.74		0 ILMN_2052373
COL9A2	1.507	8.05		0 ILMN_1685122
SLC2A5	-1.504	-8.71		0 ILMN_1671337
PDK4	-1.506	-8.57		0 ILMN_1684982
MSMB	-1.526	-5.38	3.00E-06	ILMN_1790906
DES	-1.555	-9.54		0 ILMN_1698995
PRDM8	-1.665	-8.73		0 ILMN_1802082
MME	-1.688	-7.48		0 ILMN_1678170
KRT15	-1.837	-9.88		0 ILMN_1770612
TP63	-1.849	-10.31		0 ILMN_3305055

Supplementary Table 10. Refined gene signature for each patient subtype

iCluster1	iCluster2	iCluster3	iCluster4	iCluster5
ATP6V1B2	AKAP11	AKAP11	ACVR1	AFG3L2
BRF2	ANKRD46	ANKRD46	ANKRD46	ANKRD46
C8orf58	ATP6V1B2	ARF5	C8orf76	ATP6V1B2
CCAR2	C18orf25	ATP6V1B2	CCAR2	BRF2
CHMP7	C8orf58	C8orf58	CHMP4C	C18orf25
COPS7A	CCAR2	C8orf76	CHMP7	C8orf58
CTNS	CDC16	CCAR2	COX10	CCAR2
CXXC1	CHMP4C	CHMP4C	COX4I1	CHMP7
DMTN	CHMP7	COL20A1	CPNE3	CUEDC2
ELP3	CUEDC2	COX4I1	CTNS	CXXC1
ERI1	DCTN2	CPNE3	CXXC1	DMTN
FAM160	DMTN	DMTN	CYB5D2	EBAG9
FAM160B2	ELP3	EBAG9	DEF8	ELAC2
GHITM	ERI1	ELAC2	DMTN	ELP3
GTF2E2	FAM160B2	ELP3	EBAG9	ERI1
INTS9	GHITM	ERI1	ELAC2	FAM160B2
KAT6A	GTF2E2	FSCN3	ELP3	FIG4
KLHDC4	GTF2F2	GTF2E2	ERCC3	GPR88
LSM1	HDDC2	HSBP1	FAM160B2	GTF2E2
MAPKAP1	HDHD2	IMPA1	FUK	HDHD2
MTMR9	HSBP1	INTS9	GHITM	INTS9
NUDT16P1	INTS9	MAPKAP1	GTF2F2	KAT6A
PPP3CC	KCNK12	NSMCE2	IMPA1	LINC00938
PQLC1	LSM1	PPP3CC	INTS9	LSM1
PTK2B	MAP3K7	PTDSS1	KAT6A	MAP3K7
R3HCC1	MAPKAP1	PTK2B	KLHDC4	MAPKAP1
SAT2	MTMR9	R3HCC1	MAPKAP1	MELK
SCAF11	PHF11	RIPK2	MELK	MMS19
SORBS3	PIGN	RP11-60I3.5	MMS19	MTMR9
TM2D2	PPP3CC	SORBS3	MPDU1	PPP3CC
TPD52	PTDSS1	SPIDR	MTMR9	PTDSS1
ZBTB4	PTK2B	TCF25	NSMCE2	PTK2B
	R3HCC1	TPD52	NUDT16P1	R3HCC1
	RARS2	ZBTB34	PIGN	RALBP1
	RCBTB2	ZBTB4	PQLC1	RARS2
	RNGTT	ZNF252P	PTK2B	RNGTT
	SETDB2		R3HCC1	SCAF11
	SORBS3		RBFA	SCN4A
	SPAG7		RCBTB2	SETDB2
	SUGT1		RIPK2	SLC25A11
	TM2D2		SAT2	SORBS3
	TRIM13		SCAF11	TM2D2
	WDR59		SFT2D3	TPD52
	ZBTB4		SLC25A11	ZBTB34
			SORBS3	
			TCF25	
			TRAPPC2L	
			TRIM13	
			YEATS2	
			ZBTB4	

Supplementary Table 11. Published molecular signatures of primary prostate cancer

Study	Signature type	# genes	genes/pathways identified
Cuzick et al. (2011)	RNA	31	cell cycle proliferation
Irshad et al. (2013)	RNA	19	B2M, CAT, CDKN1A, CFH, CLIC4, CLU, CTSH, CXCL1, FGFR1, GPX3, IGF1, ITM2A, LGALS3, MECP2, MSN, NFE2L2, PMP22, SERPING1, TXNIP
Sharma et al. (2013)	DNA	16	PECI, TNFSF10, ABHD2, XRCC3, MAD1L1, SEC61A1, GFM1, TSPAN13, STIL, EIF2B5, TM42F1, NDUFB11, SLC26A2, AGR2, NT5DC3, TRMT12
Ramos-Montoya et al. (2014)	RNA	222	E2F1-mediated cell cycle
Lalonde et al. (2014)	DNA	100	lipid metabolism genes enriched
Oncotype Dx prostate cancer assay	RNA	17	Genomic Prostate Score (GPS) given by expression levels of BGN, COL1A1, SFRP4, TX2, AZGP1, FAM13C, KLK2, SRD5A2, FLNC, GSN, GSTM2, TPM2, ARF1, ATP5E, CLTC, GPS1, PGK1

Supplementary Table 12. Comparison of prognostic power between our 100-gene signature and published PC signatures, based on p-value.

	Performance in Stockholm (all features)	Performance in Taylor et al. (2010) (gene expression)
Cambridge-derived novel 100-gene signature	8.96E-06	0.0213
Sharma et al. (2013)	0.1086	0.0076
Cuzick et al. (2011)	0.7471	0.4076
Irshad et al. (2014)	0.7146	0.7727
HES6/Ramos-Montoya et al. (2014)	0.0260	0.1384
Lalonde et al (2014)*	0.0891	0.7691
OncotypeDX	0.7766	0.3798

* Only copy number data available