## Supplemental methods:

Expression Plasmids and sgRNA. The full-length human BRG1, FBXW7, GSK3 $\beta$ and PTEN cDNA was cloned into pMSCV-puro/neo (Clontech), pcDNA5 (Flag tag) (Invitrogen), and pcDNA3.1 (Myc or HA tag) (Invitrogen) to generate expression plasmids. To generate BRG1SA and BRG1-SD mutated PC3 cells, a DNA template in which serine (S) in BRG1 1417/1421 sites were substituted by alanine acid (A) or aspartic acid (D) and flanked by 500 bp homology arms was used. By CRISPR-Cas9 induced homology directed repair, the template was incorporated into the native locus resulting in endogenous SA or SD mutation in PC3 cells. The shRNA, sgRNA and siRNA sequences are listed in Supplemental Table 4.

Immunohistochemical and Western Blotting Antibodies. The antibodies used for western blot analysis and immunohistochemistry were pAKT Ser473 (Cell Signaling Technology; 4060, 1:1000 dilution), PTEN (Cell Signaling Technology; 9556, 1:1000 dilution), AKT (Cell Signaling Technology; 4691, 1:1000 dilution), pERK Thr202/Tyr204 (Cell Signaling Technology; 4370, 1:1000 dilution), FBXW7 (Bethyl, A301-721; 1:1000 dilution), pGSK3 $\beta$ S9 (Cell Signaling Technology; 9323, 1:1000 dilution), GSK3 $\beta$ (Cell Signaling Technology; 5558, 1:1000 dilution), AR N-20 (Santa Cruz Biotechnology; SC-816, 1:1000 dilution), SMA $\alpha$ (Sigma; A2547, 1:5000 dilution), P63 (Santa Cruz Biotechnology; SC-8431, 1:1000 dilution), c-Myc (Abcam; ab32072, 1:1000 dilution), and $\beta$-actin (Santa Cruz Biotechnology; SC-47778, 1:10000 dilution). Antibody specific to p-BRG1-1417/1421 was prepared commercially from immunizing rabbits at Shanghai Genomic Inc. (with 1:500 working dilution). Biotinylated secondary antibodies were purchased from Jackson Immunology. Staining was visualized with ABC Kit Vectastain Elite (Vector Laboratories) and DAB substrate (Vector Laboratories).

RNA isolation and real-time PCR. Total RNA was extracted using TRIzol followed by RNeasy Mini kit (Qiagen) clean up. First strand cDNA was synthesized using Superscript II (Invitrogen) and $2 \mu \mathrm{~g}$ of total RNA was used in each cDNA synthesis reaction. SYBR green Universal Master Mix reagents (Roche) and primer mixtures (Supplemental Table 4) were used for the RT-qPCR assay.

Immunoprecipitation and western blotting. Cells were lysed in $0.3 \%$ Nonidet P40 buffer ( $150 \mathrm{mM} \mathrm{NaCl}, 50 \mathrm{mM}$ Tris- $\mathrm{HCl}, \mathrm{pH} 7.5$ ) containing inhibitors ( 1 mM phenylmethylsulphonyl fluoride, $1 \mu \mathrm{~g} / \mathrm{ml}$ of aprotinin, $1 \mu \mathrm{~g} / \mathrm{ml}$ of leupeptin, $1 \mu \mathrm{~g} / \mathrm{ml}$ of pepstatin, 1 mM Na 3 VO 4 , and 1 mM NaF , all in their final concentrations). Cell debris were removed by centrifuging at $4^{\circ} \mathrm{C}$, $13,000 \mathrm{r} . \mathrm{p} . \mathrm{m}$. for 15 min , and lysates were incubated for 6 h at $4^{\circ} \mathrm{C}$ with anti-Flag M2 agarose (Sigma). The immunoprecipitates were washed three times with $0.3 \%$ Nonidet P40 buffer before boiled and analyzed by western blotting according to the standard methods. The following primary antibodies were commercially obtained: Flag (Sigma, ab1162, 1:10,000 working dilution), BRG1(Abcam, ab110641, 1;2000), HA (cell signaling, 3724, 1:1,000), FBXW7 (Bethyl, A301-721, 1:500), AKT (Cell signaling, 4691, 1:1,000), PTEN (Cell signaling, 9556, with 1:1,000 working dilution), c-Myc (Abcam, ab32072, 1:2,000), PhosphoERK (Cell signaling, 4370, 1:1,000) and Ub (Cell signaling, 3933, 1:1,000).

Mass spectrometry analysis. To identify BRG1 phosphorylation sites, 293T cells stably expressing Flag-BRG1 with or without GSK3 $\beta$ overexpression. Cell lysates were collected to perform Flag-IP and the band corresponding to Flag-BRG1 was excised and sent for mass spectrometry analysis by National Facility for Protein Science in Shanghai, China.

In vitro ubiquitination assay. The procedure for in vitro ubiquitination assay was conducted according to the manufacturer's instructions. Flag-tagged BRG1, BRG1-SA, BRG1-SD and Flag-FBXW7 immunocomplexes were purified from 293 cells using Flag M2 beads (Sigma), and then eluted by incubating with a molar excess of Flag peptide. The FBXW7 immunocomplex was mixed with BRG1 substrate, and this mixture was added to a ubiquitin ligation reaction (Enzo Life Sciences). After the reactions, and the samples were submitted to immunoblotting with the anti-Ub antibody to examine ubiquitin ladder formation.

In vitro Kinase Assay. GSK3 $\beta$ was purchased from Abcam (ab63193). Briefly, $1 \mu \mathrm{~g}$ of purified Flag tagged proteins were incubated with GSK $3 \beta$ in the presence of $5 \mu \mathrm{Ci}[\gamma-32 \mathrm{P}]$ ATP and 200 $\mu \mathrm{M}$ cold ATP in the reaction buffer for $15-30 \mathrm{~min}$. The reaction was stopped by the addition of

SDS-containing lysis buffer and detected by autoradiography.

GST pull-down assay. BRG1 truncated protein was obtained from in-vitro translation (Promega). BL21 E. coli transformed with pGEX-GST-GSK3 $\beta$ plasmid was induced (or not induced) by isopropyl- $\beta$-D-thiogalactoside $(0.1 \mathrm{mM})$ at $20^{\circ} \mathrm{C}$ for 12 h . Protein was then purified through GST antibody-conjugated beads, and incubated with BRG1 protein. Beads were subsequently harvested through centrifugation and washed four times by $0.2 \% \mathrm{NP} 40$ buffer before boiled by $1 \times$ SDS-polyacrylamide gel electrophoresis loading buffer and subjected to western blotting.

Chromatin-immunoprecipitation assays. The ChIP assays were performed using Magnetic ChIP kit (Millipore). The procedure was as described in the kit provided by the manufacturer. BRG1 (Abcam, ab110641), H3K27ac (Cell signaling, 9733), H3K27me3 (Abcam, ab6002) were then used for immunoprecipitation. After reverse-crosslinking, the precipitated DNA was amplified by primers and quantified by the Step One Plus real-time-PCR machine. Primer sequences can be found in the Supplemental Table 4.

PTEN-WT 22RV-1

| Symbol | log2 beta score (Day 0 vs Day 45) | $p$ value | Symbol | $\log 2$ beta score (Day 0 vs Day 45) | $p$ value |
| :---: | :---: | :---: | :---: | :---: | :---: |
| BRD8 | 0.61494 | 0.001106 | H3F3A | 0.85445 | 0.001869 |
| DAXX | 0.8287 | 0.002277 | HIST2H2AC | 0.70635 | 0.007765 |
| DDX23 | 0.7485 | 0.004962 | INTS4 | 0.85057 | 0.001933 |
| DDX27 | 0.72716 | 0.006304 | KAT8 | 0.72222 | 0.00668 |
| DDX51 | 1.0182 | 0.000494 | KTI12 | 0.90983 | 0.001171 |
| DHX16 | 0.79607 | 0.00305 | LSM2 | 1.1627 | 0.00014 |
| DIDO1 | 0.81236 | 0.002674 | MCM4 | 0.80682 | 0.002792 |
| HDAC3 | 0.74673 | 0.005059 | NCAPD3 | 0.73538 | 0.005735 |
| INTS6 | 0.87064 | 0.001611 | PRMT5 | 0.72195 | 0.00668 |
| LSM7 | 0.75212 | 0.004801 | RFC2 | 0.69224 | 0.009043 |
| RAD54L | 0.77266 | 0.003888 | RUVBL1 | 0.95215 | 0.000752 |
| SETD1A | 0.87651 | 0.001536 | RUVBL2 | 1.0464 | 0.000354 |
| TERF2 | 0.90433 | 0.001257 | SMG6 | 0.89365 | 0.001343 |
| TPR | 0.76217 | 0.004339 | SNRNP200 | 1.3751 | $6.44 \mathrm{E}-05$ |
| ANKRD36 | 1.0025 | 0.000537 | SNRPB | 1.4705 | $5.37 \mathrm{E}-05$ |
| BANF1 | 0.77831 | 0.003695 | SNRPD2 | 0.95652 | 0.000709 |
| CHAF1A | 0.82799 | 0.002288 | SNRPE | 0.96633 | 0.000666 |
| CHAF1B | 1.0918 | 0.000226 | SNRPG | 0.89621 | 0.0013 |
| CHD4 | 0.6883 | 0.009322 | TCP1 | 0.93805 | 0.000827 |
| CHMP6 | 0.90345 | 0.001257 | TERF1 | 0.78264 | 0.003609 |
| DDX41 | 0.7632 | 0.004318 | TINF2 | 0.77808 | 0.003705 |
| DDX42 | 0.72112 | 0.006713 | XRCC6 | 1.1658 | 0.00014 |
| DDX49 | 1.2246 | 0.000118 |  |  |  |
| EIF4A3 | 0.85503 | 0.001858 |  |  |  |
| EP400 | 0.74755 | 0.005005 |  |  |  |

PTEN-KD (shPTEN) 22RV-1

| Symbol | $\log 2$ beta score (Day 0 vs Day 45) | $p$ value | Symbol | $\log 2$ beta score (Day 0 vs Day 45) | $p$ value |
| :---: | :---: | :---: | :---: | :---: | :---: |
| ACTL6A | 0.72582 | 0.008839 | ANKRD36 | 0.84749 | 0.002943 |
| CENPT | 0.71975 | 0.009269 | BANF1 | 0.90995 | 0.001622 |
| DDX11 | 0.76006 | 0.00653 | CHAF1A | 0.77751 | 0.005617 |
| DDX18 | 0.83743 | 0.003233 | CHAF1B | 1.422 | $6.44 \mathrm{E}-05$ |
| DDX24 | 0.8656 | 0.00246 | CHD4 | 0.84712 | 0.002964 |
| DDX47 | 0.83538 | 0.003276 | CHMP6 | 0.89407 | 0.001923 |
| DDX55 | 0.94985 | 0.001149 | DDX41 | 0.97964 | 0.000827 |
| DHX37 | 0.93767 | 0.001278 | DDX42 | 0.80198 | 0.004543 |
| ELP3 | 0.85423 | 0.002792 | DDX49 | 1.1413 | 0.000247 |
| EP300 | 0.80391 | 0.004457 | EIF4A3 | 0.97152 | 0.00087 |
| HIST1H2AH | 0.823 | 0.00377 | EP400 | 0.96808 | 0.000902 |
| HIST1H2AI | 0.7758 | 0.005703 | H3F3A | 1.0506 | 0.000462 |
| HUWE1 | 0.77828 | 0.005574 | HIST2H2AC | 0.73011 | 0.008474 |
| JMJD6 | 0.61851 | 0.003952 | INTS4 | 0.82212 | 0.00377 |
| LSM12 | 0.87185 | 0.00232 | KAT8 | 0.72206 | 0.009118 |
| LSM6 | 0.74705 | 0.007282 | KTI12 | 2.0273 | $2.15 \mathrm{E}-05$ |
| PHF5A | 1.1137 | 0.000279 | LSM2 | 1.3152 | 8.59E-05 |
| RBBP5 | 0.73854 | 0.007829 | MCM4 | 1.0127 | 0.000677 |
| RTF1 | 0.71355 | 0.009945 | NCAPD3 | 0.85276 | 0.002835 |
| SAP18 | 1.33 | 7.52E-05 | PRMT5 | 0.80299 | 0.004479 |
| SF3B1 | 1.1258 | 0.000258 | RFC2 | 0.99659 | 0.000698 |
| SKIV2L2 | 0.91178 | 0.001568 | RUVBL1 | 1.1518 | 0.000226 |
| SMARCA4 | 1.048 | 0.000473 | RUVBL2 | 1.2143 | 0.000172 |
| SMARCA5 | 0.86202 | 0.002545 | SMG6 | 0.73655 | 0.007991 |
| SMARCB1 | 1.2251 | 0.000172 | SNRNP200 | 1.3738 | 7.52E-05 |
| SMARCE1 | 0.88691 | 0.00203 | SNRPB | 1.2148 | 0.000172 |
| SMC3 | 1.052 | 0.000462 | SNRPD2 | 1.0812 | 0.000344 |
| SNRPF | 0.98523 | 0.000784 | SNRPE | 1.192 | 0.000183 |
| SUZ12 | 0.83128 | 0.003383 | SNRPG | 1.1927 | 0.000183 |
| TADA2B | 0.89269 | 0.001923 | TCP1 | 0.75983 | 0.006541 |
| UPF1 | 0.96248 | 0.000999 | TERF1 | 1.1181 | 0.000269 |
| YEATS4 | 0.713 | 0.009956 | TINF2 | 0.97171 | 0.00087 |
|  |  |  | XRCC6 | 0.94623 | 0.001181 |

B


Supplemental Figure 1. An epigenome-wide based CRISPR-Cas9 screen identifies BRG1 as a synthetic lethal target in PTEN-deficient PCa cells. (A) Summary of the gene list showing the decreased sgRNA abundance in PTEN-WT and PTEN-KD (shPTEN) 22RV-1 cells. Genes in red denotes that their abundances are only decreased in PTEN knockdown cells. (B) The correlation between BRG1 expression and Gleason score or PSA levels in patients. Wilcoxon rank sum test was used to determine statistical significance (Asian radical prostatectomy cohort).


Supplemental Figure 2. BRG1 is important for PTEN-null PCa cells. (A) IB analysis of BRG1 knockdown efficiency in PCa cells as indicated. (B) Soft agar assays in parental and BRG1-KD cells. (C) IB of lysates (left) and cell growth measurements (right) in control and BRG1-KD (shBRG1) PC3 and LNCaP cells with or without PTEN overexpression (quantitative results shown are representative of 3 experiments, 2-way ANOVA followed by Tukey's multiple comparisons test). (D) Ki67 and C-Caspase 3 (C-C3) staining from xenografts derived from PTEN-KD and PTEN/BRG-KD 22RV-1 cells, and the quantitative results are shown in the right panel ( $\mathrm{n}=8$, two-tailed Student's t test). Scale bar: $100 \mu \mathrm{~m} .{ }^{* *} \mathrm{p}<0.01$.


Supplemental Figure 3. BRG1 loss inhibits prostate tumorigenesis elicited by Pten loss. (A) H\&E-stained sections of representative anterior prostate (AP), dorsal-lateral prostate (DLP) and ventral prostate (VP) of 6-month-old control and $\operatorname{Brg} 1^{\mathrm{PC}-/-}$ mice. Scale bar: $100 \mu \mathrm{~m}$. (B) IB analysis of BRG1 protein in the prostatic lysates from $\mathrm{Pten}^{\mathrm{PC}-/-}$ and $\mathrm{Pten}^{\mathrm{PC}-/-}$; $\mathrm{Brg} 1^{\mathrm{PC}-/-}$ mice. (C) Ki67 staining of prostate section from $\mathrm{Pten}{ }^{\mathrm{PC}-/-}$ and $\mathrm{Pten}{ }^{\mathrm{PC}-/-}$; $\mathrm{Brg} 1^{\mathrm{PC}-/-}$ mice, and the quantitative results are shown in the lower panel ( $\mathrm{n}=8$, two-tailed Student's t-test). Scale bar: $50 \mu \mathrm{~m}$. (D) RT-qPCR analysis of BRG1 and PTEN in the organoids derived from wild-type, Pen-null mice as indicated (two-tailed Student's t-test). (E) BRG1 knockdown efficiency in c-Myc overexpressed (Hi-Myc) organoids (two-tailed student's t-test). (F) Representative images and quantitation of organoid sizes from Pten; Tp53 null prostates (Pten ${ }^{\mathrm{PC}-/-}$; Tp53 ${ }^{\mathrm{PC}-/-}$ ) with or without BRG1 KD. BRG1 KD efficiency is shown on the bottom (two-tailed student's t-test). Scale bar: $400 \mu \mathrm{~m}$. (G) IB of lysates (top) and cell growth measurements (low) in control and BRG1-KD PC3 and 22RV-1 (shPTEN) cells with or without SPOP (F133V) overexpression (2-way ANOVA followed by Tukey's multiple comparisons test). *p $<$ $0.01, * * \mathrm{p}<0.01$. Data represents mean $\pm$ S.E.M of 3 independent experiments (D-G).


Supplemental Figure 4. PTEN loss stabilizes BRG1 in PCa cells. (A) IB analysis of BRG1 protein in the prostatic lysates from wild type and Pten ${ }^{\mathrm{PC}-/-}$ mice. Quantitative results shown are representative of 3 experiments, two-tailed student's t-test. (B) GST-pull down analysis to map the region in BRG1 responsible for the interaction with GSK3 $\beta$. (C) IB analysis of BRG1 stability in 293 cells with Flag tagged GSK3 $\beta$ or GSK3 $\beta-$ S9A overexpression.


Supplemental Figure 5. The phosphorylation of BRG1 in Serine 1417 and 1421 facilities FBXW7 binding and subsequent BRG1 degradation. (A) Spectrum results of peptide containing BRG1 S1417 and S1421 phosphorylation. The phosphorylation signal is stimulated upon GSK3 $\beta$ overexpression in 293 cells. (B) Specificity of phospho-S1417/1421-BRG1 antibody is determined by dot blot assay. PVDF membrane was spotted with indicated amounts of phospho-S1417/1421 or none phosphorylated peptide, and probed with p-S1417/1421-BRG1 antibody. (C) IB analysis of BRG1 in PC3 and LNCaP cell transfected with scramble or FBXW7, $\beta-\operatorname{TrCP}$ oligonucleotides. (D) Examination of exogenously expressed Flag tagged wild type, SA and SD BRG1 protein in 293 cells. (E) Sequencing validations of CRISPR BRG1-SA and BRG1-SD knock-in allele in PC3 cells. (F) WT, SA and SD cell lysates were subjected to IP with anti-BRG1 antibody and IB with anti-ubiquitin (anti-Ub). (G) Wild-type or SD PC3 cells were treated with $100 \mu \mathrm{~g} / \mathrm{ml}$ cycloheximide (CHX), and IB analysis of WCL at indicated time points.


Supplemental Figure 6. BRG1 modulated chromatin configurations in PTEN-deficient PCa cells to initiate pro-tumorigenic transcriptome. (A) Unsupervised cluster analysis of differentially expressed genes in control, BRG1-KD, PTEN-KD and PTEN; BRG1-KD 22RV-1 cells. (B) KEGG-DEGs relationship network between PTEN-KD and PTEN; BRG1-KD cells. (C) The genomic distribution of BRG1 intervals in PTEN-KD 22RV-1 cells. (D) KEGG pathway enrichment analysis of the overlapping genes (PTEN-dependent BRG1 signature), which harbor BRG1 peaks and the changes of DEGs and OCRs. (E) The activity of PTEN-dependent BRG1 signature in PTEN-WT and PTEN-deleted PCa tumors (TCGA dataset, 1-way ANOVA followed by Tukey's multiple comparisons test). (F) IB analysis of c-Myc and p-ERK in the prostatic lysates from Pten ${ }^{\mathrm{PC}-/-}$, and $\mathrm{Pten}^{\mathrm{PC}-}$ ${ }^{\prime-}$; $\operatorname{Brg} 1^{\mathrm{PC}-/-}$ mice. Blots images are derived from replicate samples run on parallel gels. (G) Correlations (by Pearson's) between the BRG1 transcriptome (GSE115619), MAPK signature (GSE4739), and Myc overexpressed signature (GSE10954) within PCa specimens (TCGA, $n=374$ ) are shown. Yellow, high-signature scoring in prostate tumor specimens; blue, low-signature scoring. $* * \mathrm{p}<0.01$.


Supplemental Figure 7. Treatment of PFI-3 specifically inhibits the growth of PTEN-deficient PCa cells. (A) MTT analysis of effect for PFI-3 treatment ( 500 nM ) in wild-type 22RV-1 (left), PTEN-KD (shPTEN) 22RV-1 (middle) or PC3 (right) cells with or without BRG1 knockdown (shBRG1) (data from 3 independent experiments, two-way ANOVA followed by Tukey's multiple comparisons test). (B) Ki67 and C-C3 staining of mice prostatic sections from vehicle and PFI-3 treated Pten ${ }^{\text {PC }-\checkmark}$ mice. Scale bar: $50 \mu \mathrm{~m}$. (C-D) Body weight changes (C) and daily food intake (D) in Pten ${ }^{\text {PC- }-}$ mice treated with vehicle or PFI-3 as indicated ( $\mathrm{n}=5$, two-tailed Student's t -test). **p $<0.01$.


Supplemental Figure 8. Analysis of BRG1 and PTEN in PCa patients and the other tumor type of cancer cells.
(A) Genomic alterations of BRG1 and PTEN genes in prostate cancer datasets. (B) MTT analysis of control and BRMdepleted PC3 and 22RV-1 cell. BRM knockdown efficiency is verified by RT-qPCR analysis (right, 1-way ANOVA followed by Tukey's multiple comparisons test). (C) MTT analysis of glioma cells with or without BRG1 KD (shBRG1) as indicated (right, ). BRG1 KD efficiency is shown by western blotting (left). (D) IB of lysates (left) and cell growth measurements (right) in control and BRG1-KD (shBRG1) melanoma (A375) and breast cancer cells (MCF7) with or without PTEN KD (shPTEN). $\mathrm{n}=3$ independent experiments, 2-way ANOVA followed by Tukey's multiple comparisons test (B left, C and D). ${ }^{*} \mathrm{p}<0.05,{ }^{* *} \mathrm{p}<0.01$.

Supplemental Table 1. The gene list of chromatin regulators library.

| $\overline{A C D}$ | CCDC67 | DDX6 | H1FOO | JMJD6 | MYSM1 | PRMT8 | SKIV2L2 | TNIP2 |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| ACIN1 | CCDC96 | DDX60 | H2AFB1 | JMY | N6AMT1 | PYGO1 | SMARCA1 | TNKS |
| ACTL6A | CDYL | DFFB | H2AFV | KAT2A | N6AMT2 | PYGO2 | SMARCA2 | TNKS2 |
| ACTL6B | CDYL2 | DHX15 | H3F3A | KAT2B | NAP1L1 | RABGAP1L | SMARCA4 | TOX4 |
| AEBP2 | CENPA | DHX16 | HAT1 | KAT5 | NAP1L2 | RAD18 | SMARCA5 | TPR |
| AICDA | CENPE | DHX29 | HDAC1 | KAT6B | NAP1L3 | RAD50 | SMARCAD1 | TRAF5 |
| AKAP8 | CENPT | DHX30 | HDAC10 | KAT7 | NAP1L5 | RAD54B | SMARCAL1 | TRAF7 |
| ALKBH8 | CEP63 | DHX32 | HDAC11 | KAT8 | NAT10 | RAD54L | SMARCB1 | TRDMT1 |
| ANKRD26 | CEP72 | DHX35 | HDAC2 | KDM1A | NCAPD2 | RAD54L2 | SMARCC1 | TRIM27 |
| ANKRD36 | CETN2 | DHX37 | HDAC3 | KDM1B | NCAPD3 | RAG1 | SMARCC2 | TRMT11 |
| APITD1 | CHAF1A | DHX38 | HDAC4 | KDM2A | NCOA1 | RAG2 | SMARCD1 | TROVE2 |
| AQR | CHAF1B | DHX40 | HDAC5 | KDM3A | NCOR1 | RAI1 | SMARCD2 | TSPYL2 |
| ARID1A | CHD1 | DHX57 | HDAC6 | KDM3B | NINL | RBBP5 | SMARCD3 | TSPYL4 |
| ARID1B | CHD1L | DHX9 | HDAC7 | KDM4A | NOVA2 | RBBP7 | SMARCE1 | TTF2 |
| ARID2 | CHD2 | DICER1 | HDAC8 | KDM4C | NPM2 | RBX1 | SMC1B | UBE2A |
| ARID3C | CHD3 | DIDO1 | HDAC9 | KDM4D | NSD1 | RCBTB1 | SMC3 | UBE2B |
| ARID4A | CHD4 | DNA2 | HELB | KDM5B | NSMCE1 | RECQL | SMC5 | UBR2 |
| ASCC3 | CHD5 | DNAJC18 | HELQ | KDM5C | NUMA1 | RECQL5 | SMC6 | UHRF2 |
| ASH1L | CHD6 | DNAJC2 | HELZ | KDM6A | ODF2L | RFC1 | SMG6 | UPF1 |
| ASH2L | CHD7 | DNMT3A | HFM1 | KDM8 | OFD1 | RFC2 | SMYD1 | USP22 |
| ASXL3 | CHD8 | DNMT3B | HIF1AN | KLHDC3 | PARP1 | RILPL1 | SMYD2 | UTP3 |
| ATG14 | CHD9 | DNMT3L | HIST1H1A | KTI12 | PBRM1 | RING1 | SMYD3 | UTY |
| ATRX | CHMP2A | DOT1L | HIST1H1B | L3MBTL1 | PCGF1 | RNF168 | SMYD4 | WDR11 |
| ATXN2 | CHMP2B | DPF1 | HIST1H1D | L3MBTL2 | PCGF2 | RNF2 | SMYD5 | WDR5 |
| ATXN7L2 | CHMP4B | DPF2 | HIST1H1E | LIN28B | PCGF3 | RNF20 | SNRNP200 | WHSC1 |
| BAG6 | CHMP4C | DPF3 | HIST1H1T | LMNA | PCGF5 | RNF40 | SNRPB | WRN |
| BANF1 | CHMP5 | DQX1 | HIST1H2AA | LMNB1 | PCGF6 | RNF6 | SNRPD2 | WRNIP1 |
| BAP1 | CHMP6 | DZIP3 | HIST1H2AB | LMNB2 | PCNT | RNF8 | SNRPD3 | XRCC5 |
| BAZ2A | CHRAC1 | EDC3 | HIST1H2AC | LRRC45 | PHC1 | RPTOR | SNRPE | XRCC6 |
| BAZ2B | CNBP | EHMT1 | HIST1H2AD | LRRFIP2 | PHF1 | RSF1 | SNRPF | YEATS4 |
| BICD1 | COQ5 | EHMT2 | HIST1H2AG | LSM1 | PHF11 | RTEL1 | SNRPG | YLPM1 |
| BLM | CREBBP | EIF4A3 | HIST1H2AH | LSM12 | PHF12 | RTF1 | SNX29 | YTHDC2 |
| BLOC1S1 | DAXX | ELP3 | HIST1H2AI | LSM14A | PHF13 | RUVBL1 | SPAST | ZCCHC11 |
| BMI1 | DCLRE1C | EP300 | HIST1H2AK | LSM2 | PHF14 | RUVBL2 | SPOP | ZCCHC4 |
| BRCA1 | DDX1 | EP400 | HIST2H2AC | LSM6 | PHF2 | SAP18 | SUDS3 | ZMYND11 |
| BRCC3 | DDX10 | EPC2 | HIST3H2A | LSM7 | PHF20L1 | SBNO1 | SUPT7L | ZMYND8 |
| BRD1 | DDX11 | ERCC1 | HLCS | LUZP2 | PHF21B | SET | SUPV3L1 | ZRANB3 |
| BRD8 | DDX18 | ERCC2 | HMGA2 | MBD1 | PHF23 | SETD1A | SUV39H1 |  |
| BRIP1 | DDX19A | ERCC3 | HMGB3 | MBD3 | PHF3 | SETD1B | SUV39H2 |  |
| BRPF1 | DDX19B | ERCC4 | HMGN2 | MCM4 | PHF5A | SETD2 | SUV420H1 |  |
| BRPF3 | DDX20 | ERCC6 | HMGN5 | MECP2 | PHF6 | SETD3 | SUV420H2 |  |
| BRWD1 | DDX23 | ERCC6L | HP1BP3 | MEN1 | PHF7 | SETD5 | SUZ12 |  |
| BRWD3 | DDX24 | EXOSC10 | HR | METTL5 | PHF8 | SETD7 | SVEP1 |  |
| CALCOCO1 | DDX25 | EZH1 | HUWE1 | MGMT | PHLDB1 | SETD8 | SYCE1 |  |
| CARM1 | DDX26B | EZH2 | IFT74 | MIER2 | PHLDB3 | SETDB1 | TADA2A |  |
| CASC4 | DDX27 | FAM117A | IGHMBP2 | MKRN1 | PHRF1 | SETDB2 | TADA2B |  |
| CBX1 | DDX28 | FAM81A | IKBKG | MLH3 | PIBF1 | SETMAR | TAF3 |  |
| CBX2 | DDX3Y | FANCG | ING1 | MORF4L1 | PIF1 | SETX | TCP1 |  |
| CBX4 | DDX4 | FANCL | ING2 | MORF4L2 | PINX1 | SF3B1 | TDG |  |
| CBX6 | DDX41 | FANCM | ING3 | MOV10 | PLA2G4B | SHPRH | TDRD3 |  |
| CBX7 | DDX42 | G3BP1 | ING4 | MOV10L1 | PLEKHA5 | SHROOM4 | TDRD9 |  |
| CBX8 | DDX46 | GATAD1 | ING5 | MPHOSPH8 | PLEKHA7 | SIPA1 | TERF1 |  |
| CCDC122 | DDX47 | GATAD2A | INO80 | MSL3 | POC5 | SIRT1 | TERF2 |  |
| CCDC125 | DDX49 | GATAD2B | INTS12 | MTA1 | POLQ | SIRT2 | TERF2IP |  |
| CCDC146 | DDX51 | GCC2 | INTS4 | MTA2 | PRDM9 | SIRT3 | TERT |  |
| CCDC151 | DDX52 | GMCL1 | INTS6 | MTA3 | PRMT1 | SIRT4 | TES |  |
| CCDC160 | DDX54 | GMNN | IQCE | MTF2 | PRMT3 | SIRT5 | TET1 |  |
| CCDC39 | DDX55 | GRIPAP1 | JAKMIP2 | MYEOV2 | PRMT5 | SIRT6 | TFPT |  |
| CCDC40 | DDX58 | GTF3C4 | JARID2 | MYOCD | PRMT6 | SIRT7 | TINF2 |  |
| CCDC6 | DDX59 | H1F0 | JMJD4 | MYPOP | PRMT7 | SKIV2L | TMEM38B |  |

Supplemental Table 2. The clinical information of Asian radical prostate cohort.

| Variables | All patients |
| :---: | :---: |
| Numbers | 128 |
| Age at diagnosis, yr. | 61-71 |
| Year of surgery | 2006-2010 |
| No. of biochemical recurrence, $\mathrm{n}(\%)$ | 47 (39.8) |
| Preoperative PSA, ng/mL | 16.0 (10.4-31.6) |
| Pathologic Gleason score, n (\%) |  |
| $\leq 6$ | 29 (24.6) |
| 7 | 53 (44.9) |
| 8 | 21 (17.8) |
| $\geq 9$ | 15 (12.7) |
| Adverse pathologic events, n (\%) |  |
| Seminal vesicle invasion | 13 (11.0) |
| Lymph node invasion | 4 (3.4) |
| Positive surgical margins | 6 (5.1) |

Supplemental Table 3. Summary of the clinical information of GSE21032 and TCGA.

| Variables | GSE21032 | TCGA |
| :---: | :---: | :---: |
| Numbers | 179 | 426 |
| Normal sample | 29 | 52 |
| Tumor sample | 150 | 374 |
| Samples with clinical follow up, n | 140 | 366 |
| No. of biochemical recurrence, n (\%) | 36 (25.7\%) | 60 (16.4\%) |
| Preoperative PSA, ng/mL | 12.1 (1.15-506) |  |
|  | $\leq 678$ (55.7\%) |  |
| Pathologic Gleason score, n | 749 (35.0\%) |  |
|  | $\geq 813$ (9.3\%) |  |
| Seminal vesicle invasion | 20 (14.3\%) |  |
| Lymph node invasion | 12 (8.6\%) |  |

## Supplemental Table 4. Oligonucleotides. The RT-qPCR, ChIP-qPCR, shRNA , sgRNA and siRNA primer sequences are listed.

| qRT-PCR |  |  |
| :---: | :---: | :---: |
| BRG1 | AGCGATGACGTCTCTGAGGT | GTACAGGGACACCAGCCACT |
| PTEN | TTGGCGGTGTCATAATGTCT | GCAGAAAGACTTGAAGGCGTA |
| ETV1 | CTGAACCCTGTAACTCCTTTCC | AGACATCTGGCGTTGGTACATA |
| NCOA2 | GCAGTGCTTCGCTGTCTCT | TTCATGGGAACTCTTCTTGCC |
| WEE1 | GACGAAGATGATTGGGCATCC | TGGACTGGAGATCCTTGTTACA |
| ERBB2 | TGGCCTGTGCCCACTATAAG | AGGAGAGGTCAGGTTTCACAC |
| FGFR3 | CCCAAATGGGAGCTGTCTCG | CCCGGTCCTTGTCAATGCC |
| KRAS | ACAGAGAGTGGAGGATGCTTT | TTTCACACAGCCAGGAGTCTT |
| DOT1L | CGTGTATTGTTCGTTACCTGGA | TTCAGTAGTGGTCTGGTCTTGT |
| BMI1 | CTGCCGGTCTACGATAAACATC | AGCTTGAGATCCGGGATTTCT |
| SIN3a | GGTGGAGGATGCGCTATCTTA | GGGTGTCGATGCTCTGAGATTT |
| c-Myc | TCCCTCCACTCGGAAGGAC | CTGGTGCATTTTCGGTTGTTG |
| ChIP-qPCR |  |  |
| c-Myc | AACCAGGTAAGCACCGAAGTCCA | TTCATAAGGCAGAAATCTCGAAAGG |
| ETV1 | ACAGCAACTTTAATGAGGCAAGA | TAACAGATAAGGCAGTCAGGAAT |
| KRAS | AGCCTTGCTTCTGCTCTGCGGGTTT | CCAGCCTTCCCTGCTGCATTTGG |
| BMI1 | AGATCGGGGCGAGACAATGGGGATG | GACGCCGCTGTCAATGGGCAACC |
| shRNA |  |  |
| shBRG1-3'UTR-1 | CCGGCCATATTTATACAGCAGAGAACTCGAGTTCTCTGCTGTATAAATATGGTTTTTG |  |
|  | AATTCAAAAACCATATTTATACAGCAG | GTTTCTCTGCTGTATAAATATGG |
| shBRG1-3'UTR-2 | CCGGGGCATAGGCCTTAGCAGTAACCTCGAGGTTACTGCTAAGGCCTATGCCTTTTTG |  |
|  | AATTCAAAAAGGCATAGGCCTTAGCAGTAACCTCGAGGTTACTGCTAAGGCCTATGCC |  |
| shGSK3ß-1 | CCGGGATGAATTACGGGACCCAAATCTCGAGATTTGGGTCCCGTAATTCATCTTTTTG |  |
|  | AATTCAAAAAGATGAATTACGGGACCCAAATCTCGAGATTTGGGTCCCGTAATTCATC |  |
| shGSK3ß-2 | CCGGCCAATGTTTCGTATATCTGTTCTCGAGAACAGATATACGAAACATTGGTTTTTG |  |
|  | AATTCAAAAACCAATGTTTCGTATATCTGTTCTCGAGAACAGATATACGAAACATTGG |  |
| shFBXW7-1 | CCGGGGCAACAACGACGCCGAATTACTCGAGTAATTCGGCGTCGTTGTTGCCTTTTTG |  |
|  | AATTCAAAAAGGCAACAACGACGCCGAATTACTCGAGTAATTCGGCGTCGTTGTTGCC |  |
| shFBXW7-2 | CCGGGGCATACTAATAGAGTCTATTCTCGAGAATAGACTCTATTAGTATGCCTTTTTG |  |
|  | AATTCAAAAAGGCATACTAATAGAGTCTATTCTCGAGAATAGACTCTATTAGTATGCC |  |
| shPTEN-1 | CCGGGCAGATAATGACAAGGAATATTACTCGAGTAATATTCCTTGTCATTATCTGCTTTTTG |  |
|  | AATTCAAAAAGCAGATAATGACAAGGAATATTACTCGAGTAATATTCCTTGTCATTATCTGC |  |
| shPTEN-2 | CCGGGGTGAAGATATATTCCTCCAATACTCGAGTATTGGAGGAATATATCTTCACCTTTTTG |  |
|  | AATTCAAAAAGGTGAAGATATATTCC | GAGTATTGGAGGAATATATCTTCACC |
| sgRNA |  |  |
| BRG1-sgRNA | CACCGCACGCTGGAGGAGATCGAAG |  |
| BRG1-mutant-Template | AGTTCAAGACTGCAGTGAGCTATGAT |  |
| siRNA |  |  |
| siAKT-1 | GCUAUUGUGAAGGAGGGUUTT |  |
| siAKT-2 | GGCCCAACACCUUCAUCAUTT |  |
| siPTEN-1 | GCAGAUAAUGACAAGGAAUAUUATT |  |
| siPTEN-2 | GGUGAAGAUAUAUUCCUCCAAUATT |  |
| siBRG1-1 | UCUCCGUCAGUGAGUCGCUdTdT |  |
| siBRG1-2 | UCUCUAGGUCGUUGAGGCUdTdT |  |
| siFBXW7-1 | GGGCAUACUAAUAGAGUCUAUUCAUTT |  |
| siFBXW7-2 | AGUUGGCACUCUAUGUGCUUUCAUUTT |  |
| si $\beta$-TRCP-1 | GCGUUGUAUUCGAUUUGAUAATT |  |
| si $\beta$-TRCP-2 | ACUUGCCCAGGACCCAUUAAATT |  |

