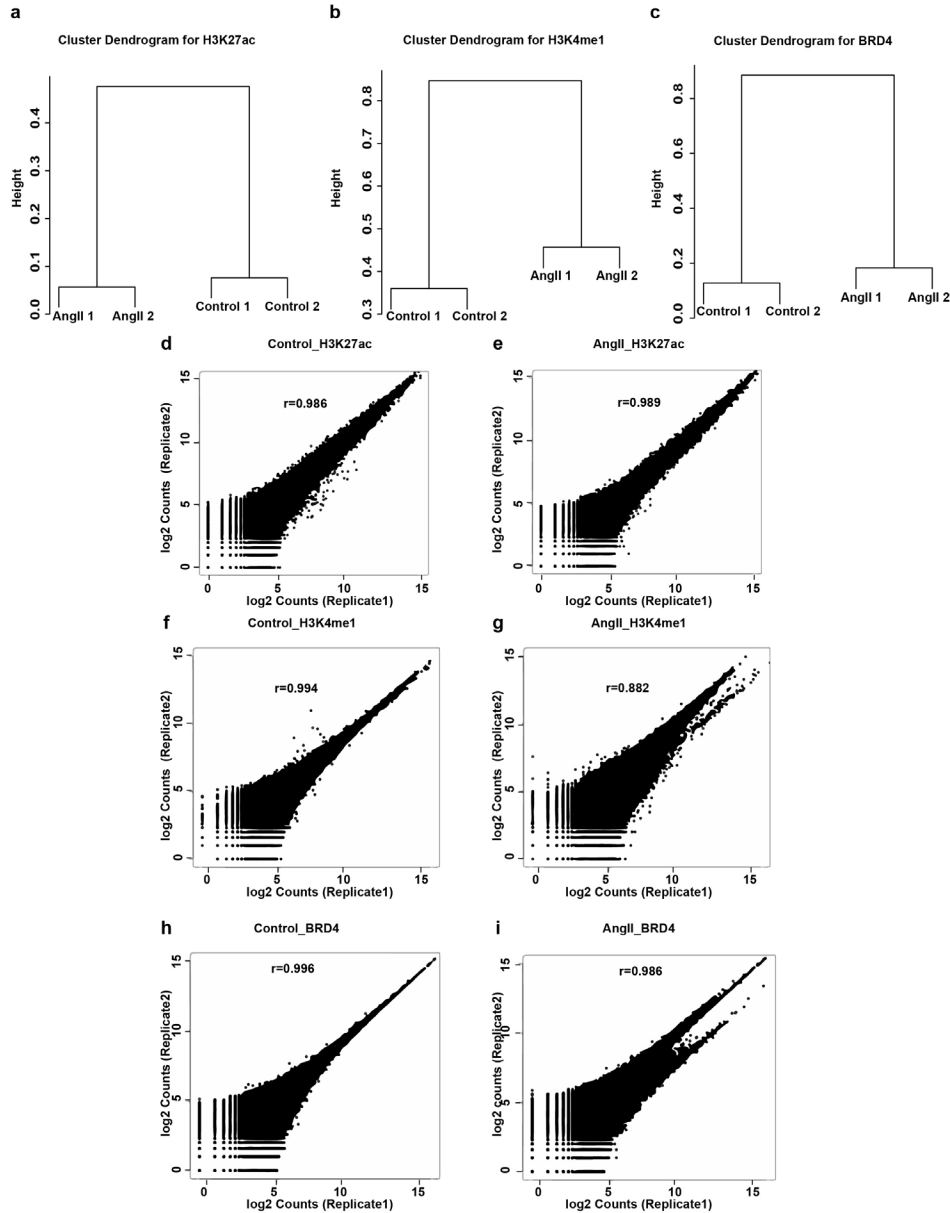
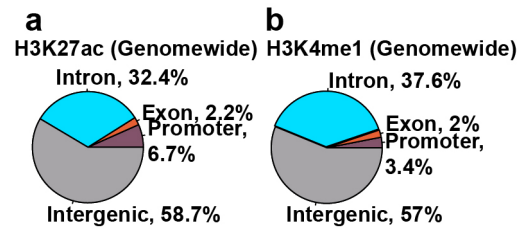


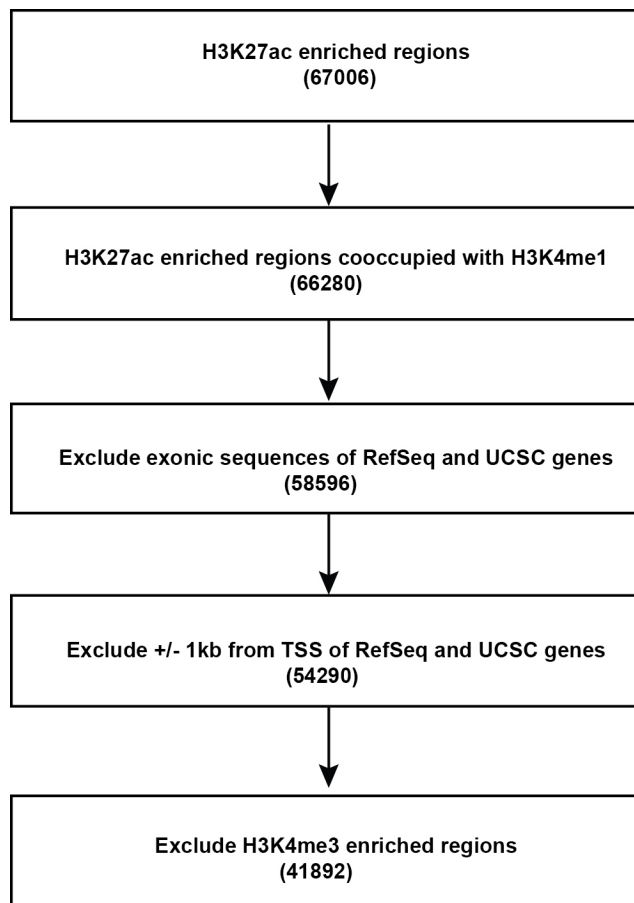
## Supplementary Figures and Tables



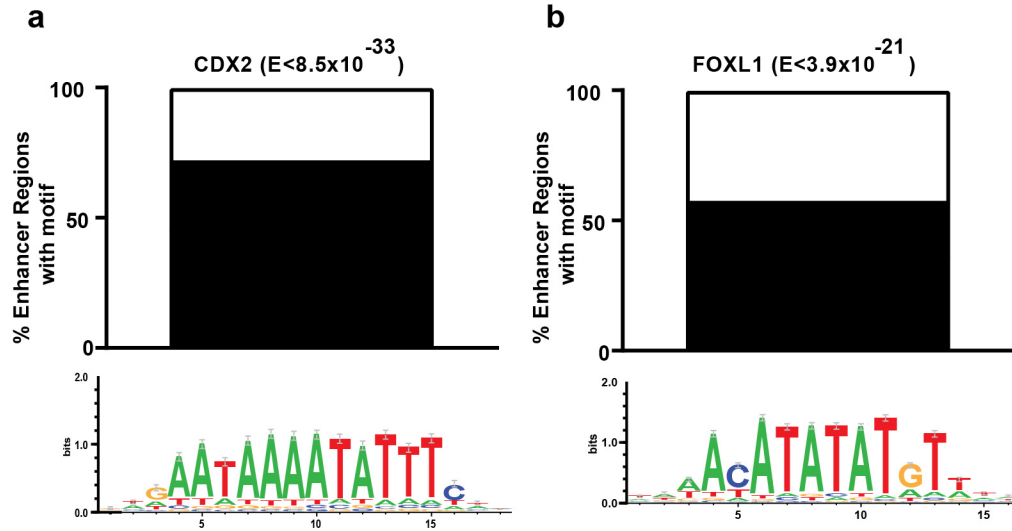
**Supplementary Figure 1. Correlation between ChIP-seq replicates.** (a) Cluster analysis of H3K27ac. (b) H3K4me1. (c) BRD4 ChIP-seq. Rat genome was divided into 566848 non-overlapping 5-kb windows. For each sample including both ChIP and input samples, coverage was calculated on each window and scaled to total number of reads across all the samples.  $\text{Log}_2(\text{ChIP}/\text{input})$  was then calculated for each window. The windows with enrichment levels ( $\text{log}_2(\text{ChIP}/\text{Input}) < 2$  in more than 2 samples) were excluded. The dissimilarities, defined as  $1 - \text{cor}$ , from the resulting data were finally calculated across the 4 samples (2 AngII and 2 controls) which were subjected to cluster analyses and shown as dendrograms. (d-i) Scatterplots of read counts ( $\text{log}_2$  counts) on identified peak regions for each replicate of H3K27ac, H3K4me1 and BRD4 ChIP-seq data for Control and AngII samples as indicated. Data shows very good correlation between the replicates (Pearson correlation coefficient ( $r$ )=0.882-0.996).



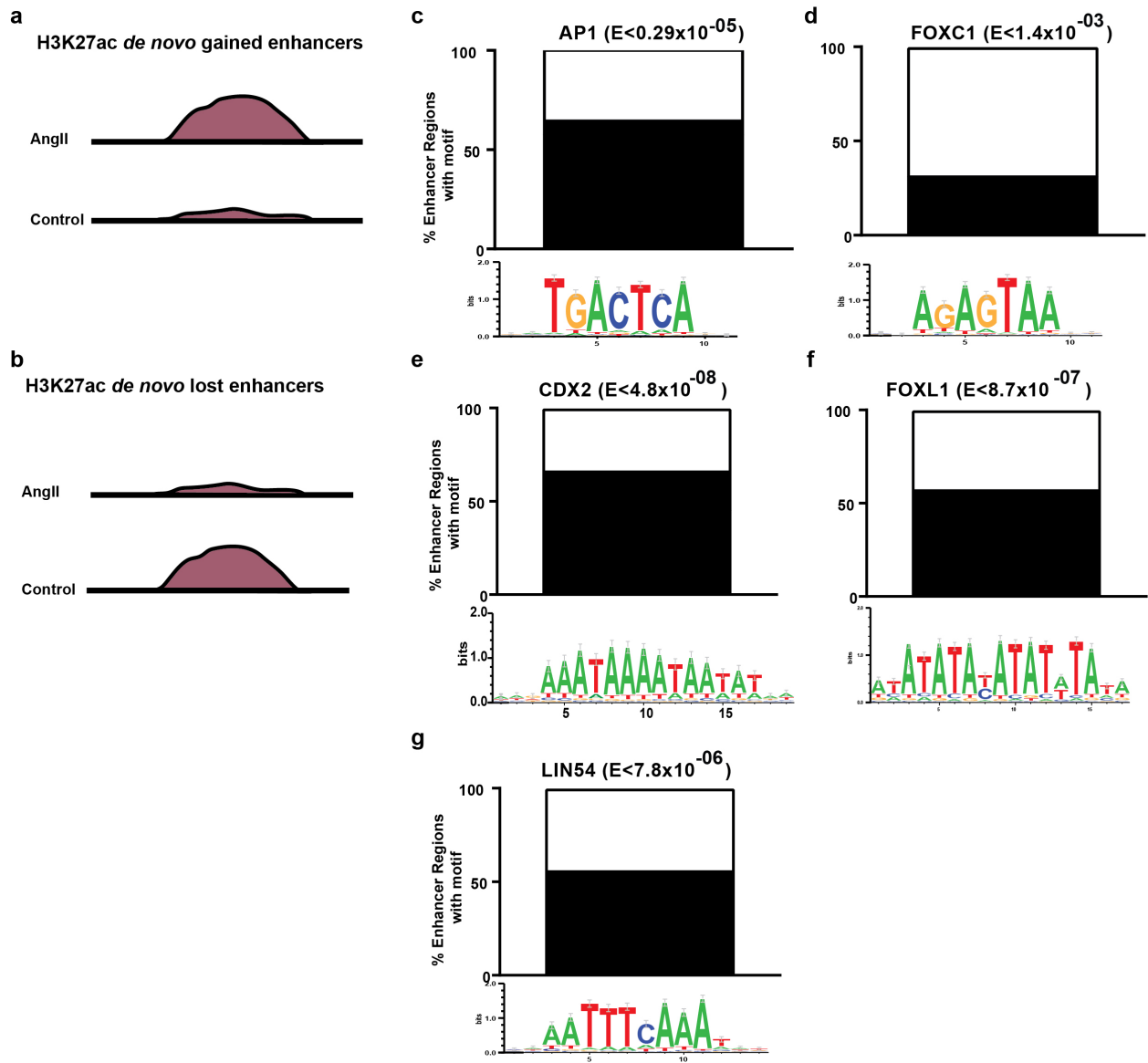
**Supplementary Figure 2. Genomic distribution of enhancer marks in VSMC.** Venn diagrams depicting (a) Distribution of H3K27ac and (b) H3K4me1 in the indicated genomic regions.



**Supplementary Figure 3. Strategy for enhancer identification.** Briefly, H3K27ac enriched regions were identified and filtered based on co-occupancy of H3K4me1. H3K27ac and H3K4me1 co-occupied regions were used to filter out exonic and promoter sequences of RefSeq and UCSC genes followed by removal of H3K4me3 enriched regions.

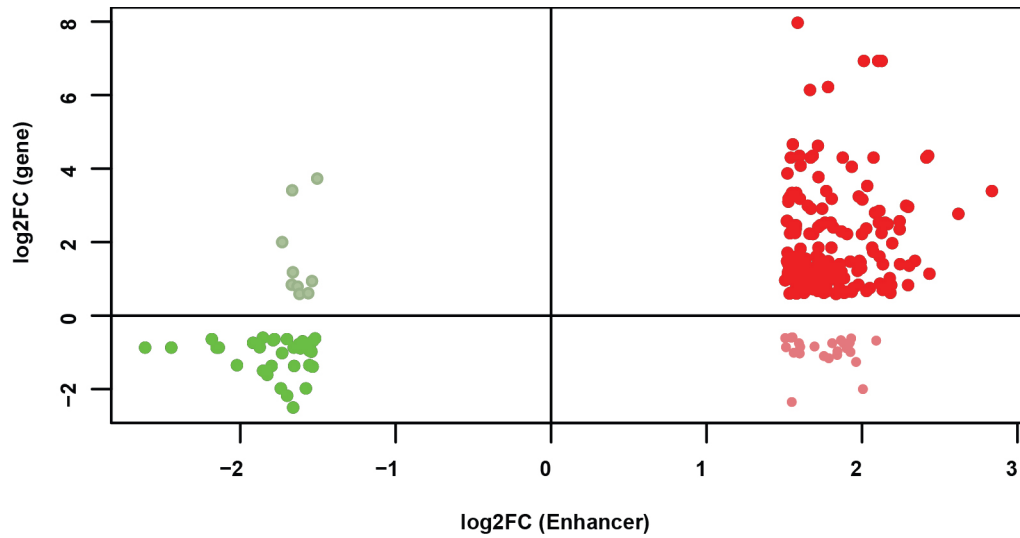


**Supplementary Figure 4. Motif discovery from AngII-downregulated enhancers. (a-b)** Enrichment of CDX2 and FOXL1 motifs in AngII-downregulated enhancers.

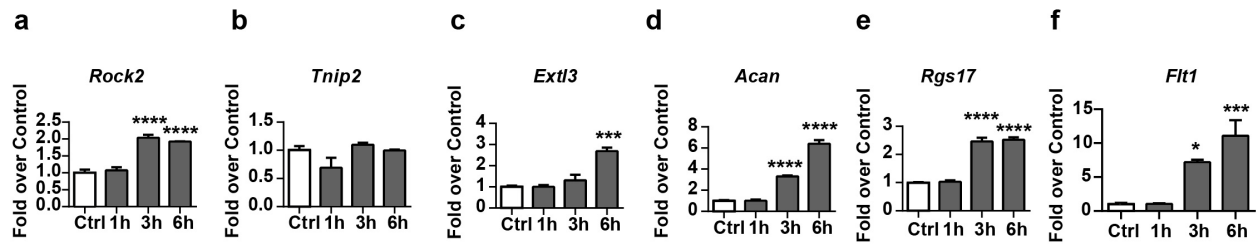


**Supplementary Figure 5. Enrichment of TF motifs on *de novo* gained and lost enhancers.**

**(a)** Schematic represents formation of *de novo* H3K27ac gained enhancers by AngII. **(b)** Schematic represents loss of H3K27ac enhancers by AngII. **(c-d)** Representative motifs of AP1 and FOXC1 are statistically enriched in H3K27ac-gained enhancers (where AngII treated H3K27ac ChIP enrichment signals are more than  $\log_2(1.5)$  and Control H3K27ac ChIP enrichment signals are less than  $\log_2(1.1)$ ) and shown as % of enhancers with individual motif. **(e-g)** Representative motifs of CDX2, FOXL1 and LIN54 enriched in H3K27ac lost enhancers (where AngII treated H3K27ac ChIP enrichment signals are less than  $\log_2(1.1)$  and Control H3K27ac ChIP enrichment signals are more than  $\log_2(1.5)$ ) and shown as % of enhancers with individual motif. E is Bonferroni adjusted *P* value.



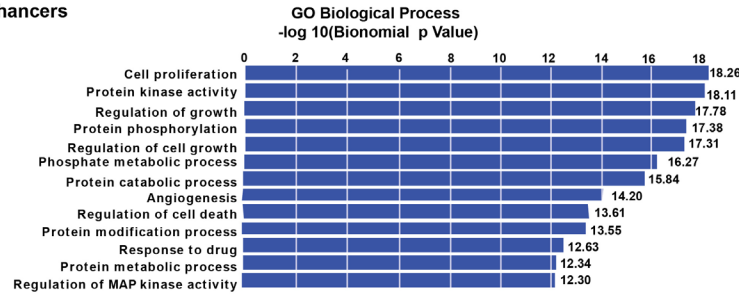
**Supplementary Figure 6. Correlation between differentially enriched enhancers and differentially regulated genes.** Scatter plot showing the log<sub>2</sub> fold change in expression of genes associated with differentially enriched H3K27ac enhancers ( $-1.5 > \log_2FC > 1.5$ ). Genes associated with upregulated enhancers are colored in shades of red and those associated with downregulated enhancers are colored in shades of green.



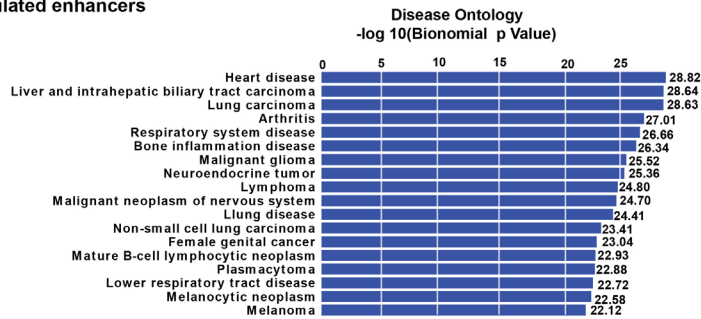
**Supplementary Figure 7. Enhancer-associated neighboring gene expression. (a-f)**

Expression of the indicated genes were validated in RVSMCs treated  $\pm$ AngII for 1-6 hrs *in vitro*. Mean $\pm$ SEM; \* $P$ <0.05; \*\*\* $P$ <0.001; \*\*\*\* $P$ <0.0001; vs. Ctrl;  $n$ =3, using one-way ANOVA, Dunnett's multiple comparisons test.

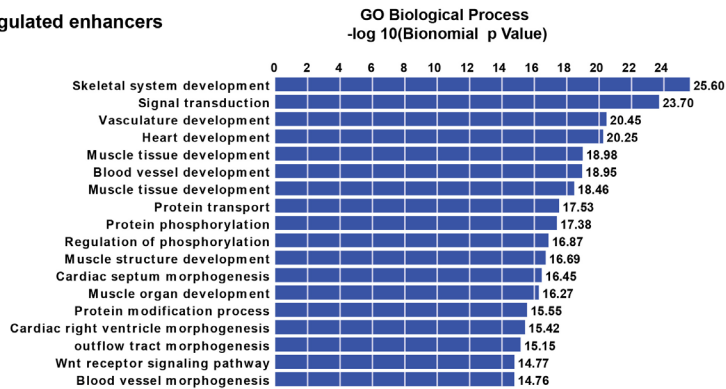
**a Upregulated enhancers**



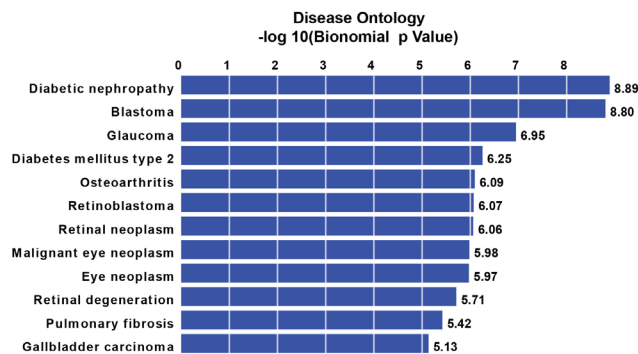
**b Upregulated enhancers**



**c Downregulated enhancers**

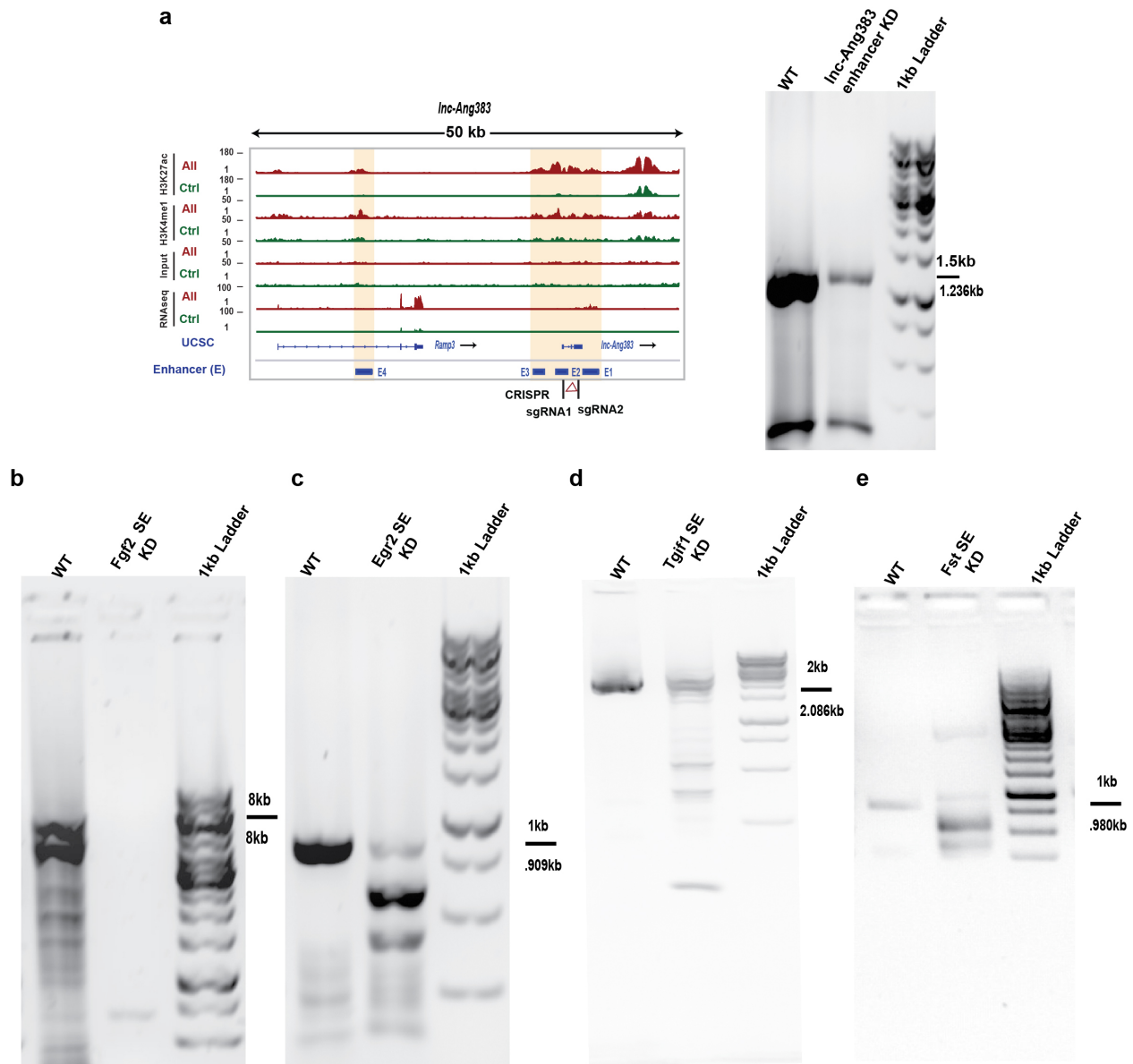


**d Downregulated enhancers**

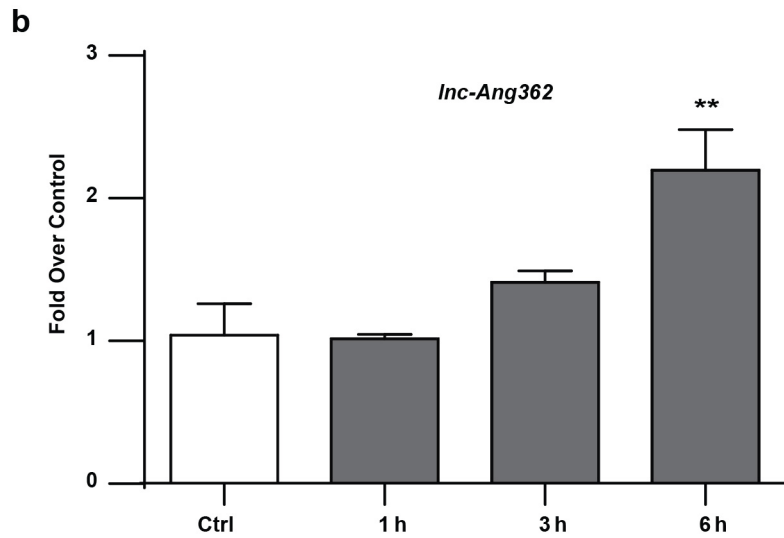
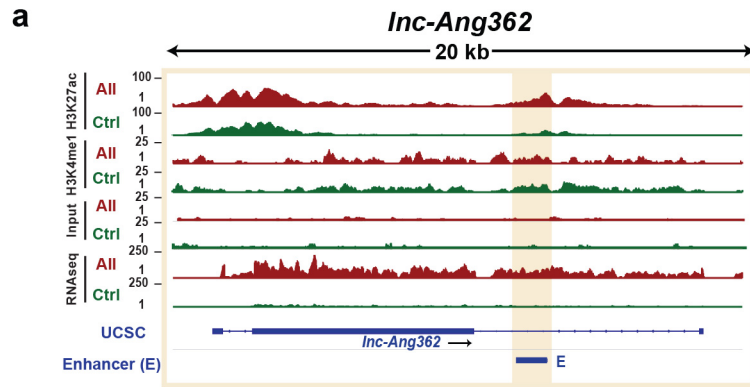


**Supplementary Figure 8. Functional annotation of differentially regulated enhancers. (a-d)** Identification of GO biological processes and Disease Ontology using GREAT tool. Upregulated enhancers are shown in panels (a) and (b), whereas downregulated enhancers are shown in panels (c) and (d). Values represent  $-\log_{10}(\text{Binomial } P \text{ value})$ .

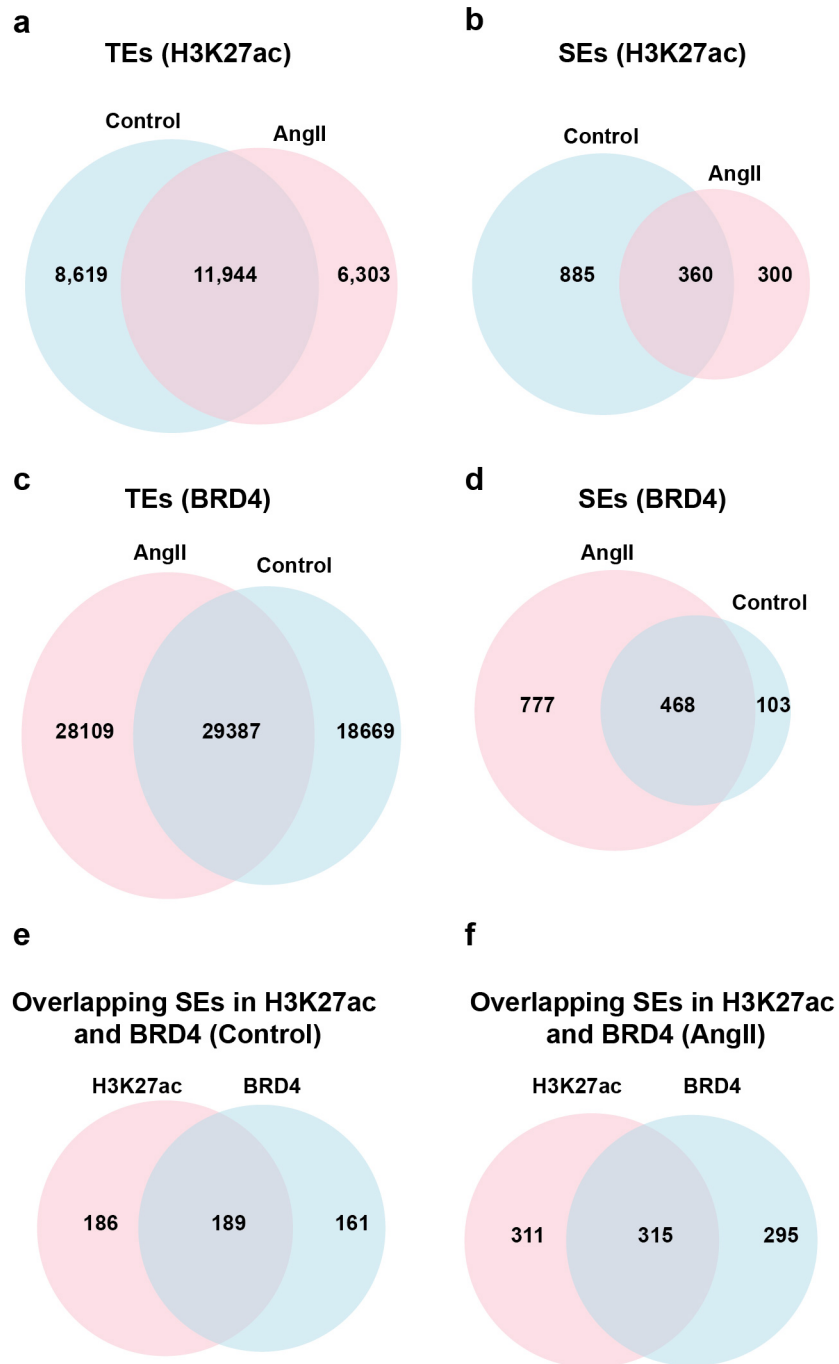




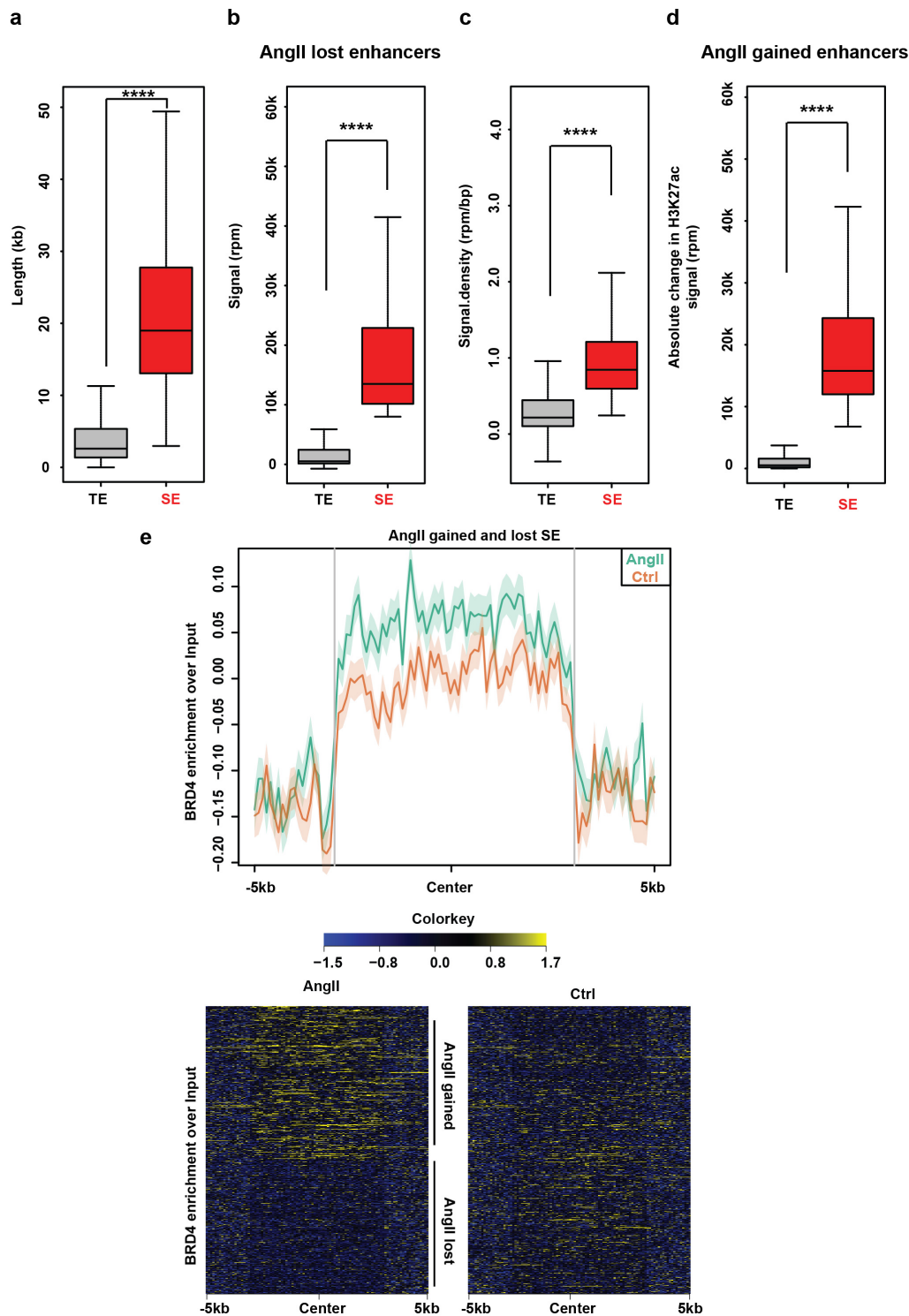
**Supplementary Figure 9. CRISPR-Cas9 mediated deletion of candidate enhancers/SEs. (a)** Genome browser tracks showing the relative position of the indicated enhancers (highlighted in orange) with respect to nearby indicated gene and the position of the sgRNAs (denoted by black lines) designed to delete the enhancer overlapping with lncRNA, *Inc-Ang383* using CRISPR-Cas9 editing in RVSMCs, and genotyping PCR result of *Inc-Ang383* enhancer deletion. **(b-e)** Genotyping PCR results to confirm deletion of SEs for *Fgf2* **(b)**, *Egr2* **(c)**, *Tgif1* **(d)**, and *Fst* **(e)**. The expected band size for each PCR is indicated on the right side of each figure.



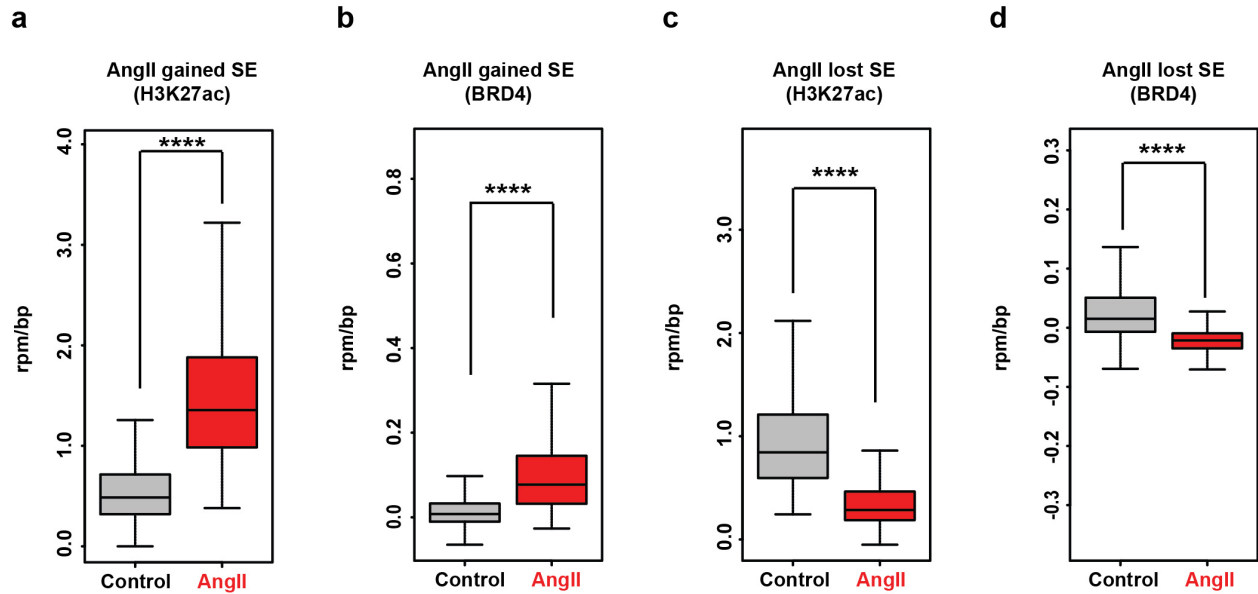
**Supplementary Figure 10. Genomic organization of *Inc-Ang362* overlapping enhancer and its regulation by AngII.** (a) Tracks show profiles of H3K27ac and H3K4me1 signal on RVSMC specific enhancer associated with *Inc-Ang362*. Each data track shown is on the same scale for both Control (Ctrl) and AngII. (b) Bars represent fold change in expression of *Inc-Ang362* upon treatment of RVSMCs with AngII for the indicated times. Mean+SEM; \*\* $P < 0.01$ ; vs. Ctrl;  $n = 3$ , using one-way ANOVA, Dunnett's multiple comparisons test.



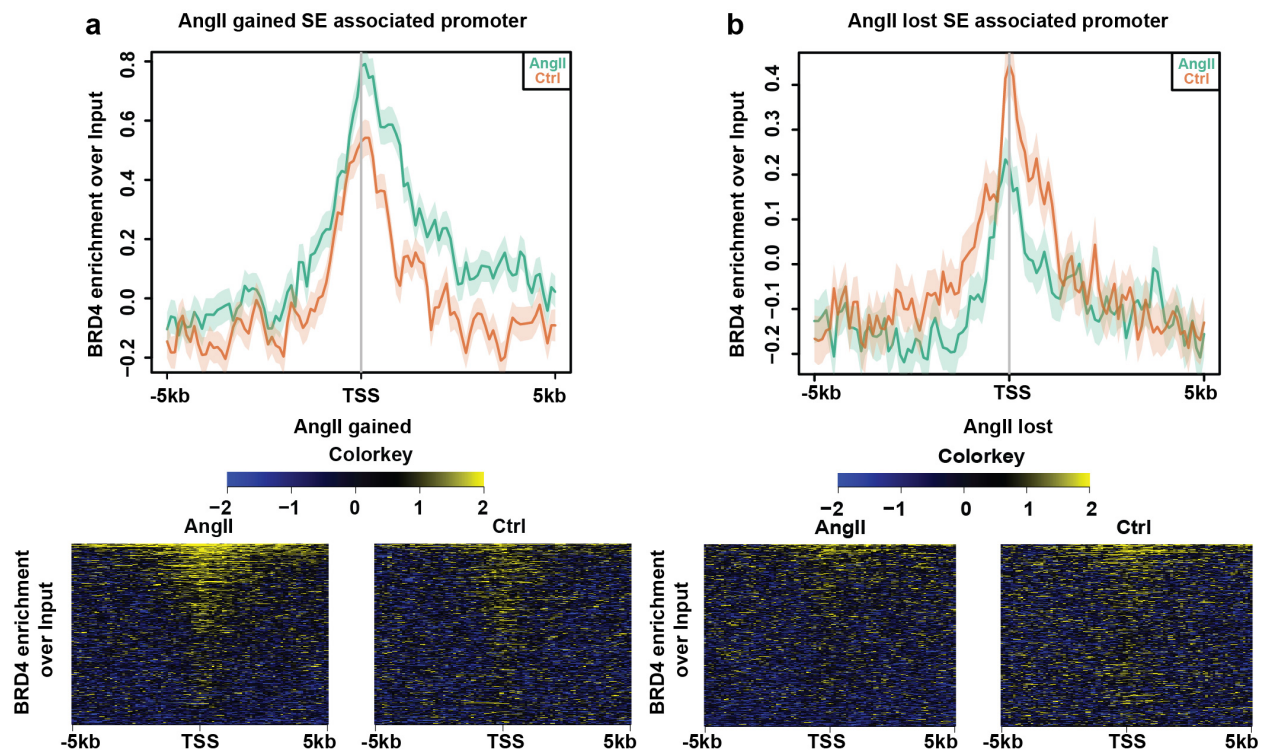
**Supplementary Figure 11. Comparison of H3K27ac and BRD4 specific TEs and SEs.** (a) Venn diagram showing the overlap between H3K27ac TEs and (b) SEs identified in Control and AngII conditions. (c) Overlap between BRD4 TEs and (d) SEs identified in Control and AngII conditions. (e) Overlap between H3K27ac and BRD4 SEs identified in Control (untreated) and (f) AngII treated RVSMCs.



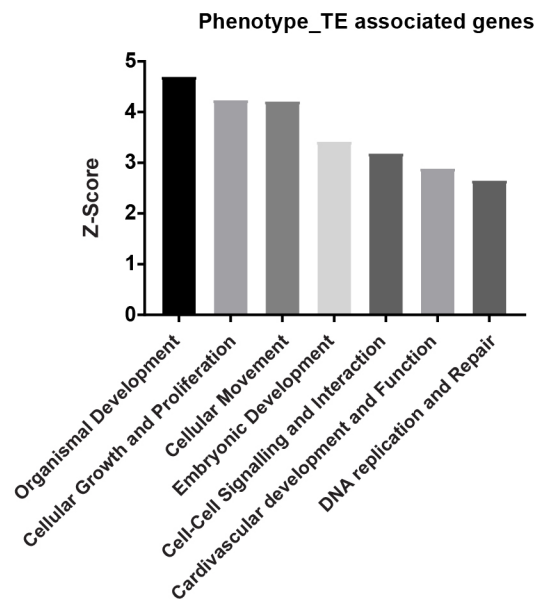
**Supplementary Figure 12. Characterization of AngII gained and lost SEs.** (a) Boxplots show median enhancer length (kb), (b) signal (rpm), and (c) density (rpm/bp) in AngII-lost TEs and SEs. (d) Boxplot shows the absolute change in H3K27ac signal in response to AngII treatment measured at AngII gained TEs and SEs. For (a-d), \*\*\*\* $p < 0.0001$  using unpaired two-tailed t test. (e) Average profile and heatmaps of BRD4 enrichment over SEs gained and lost upon AngII treatment.



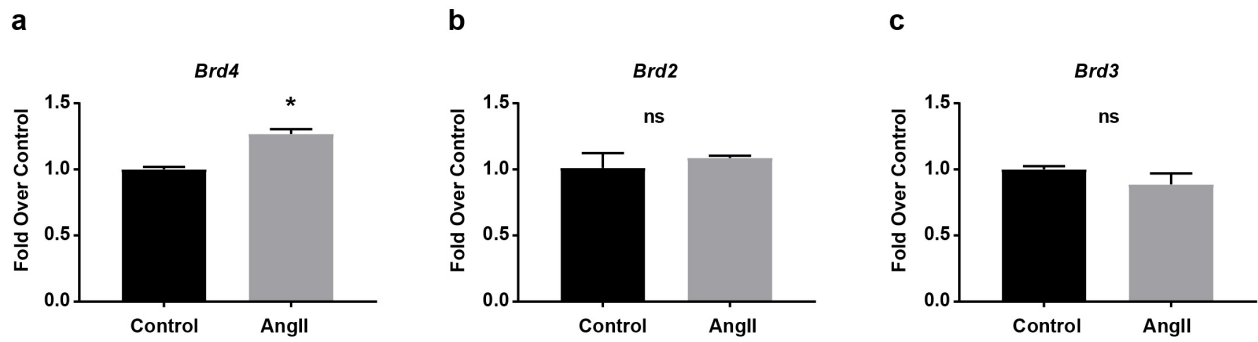
**Supplementary Figure 13. AngII-induces co-ordinate changes in H3K27ac and BRD4 occupancy over SEs. (a)** Boxplot of H3K27ac and **(b)** BRD4 signal density (rpm/bp) at AngII gained SEs. **(c)** Boxplot of H3K27ac and **(d)** BRD4 signal density (rpm/bp) at AngII lost SEs. \*\*\*\* $P < 0.0001$ , using unpaired-two tailed t test.



**Supplementary Figure 14. BRD4 enrichment mirrors H3K27ac enrichment on SE regions. (a) Average profile and heatmaps of BRD4 enrichment over promoters of AngII gained and (b) AngII lost SE-associated genes.**

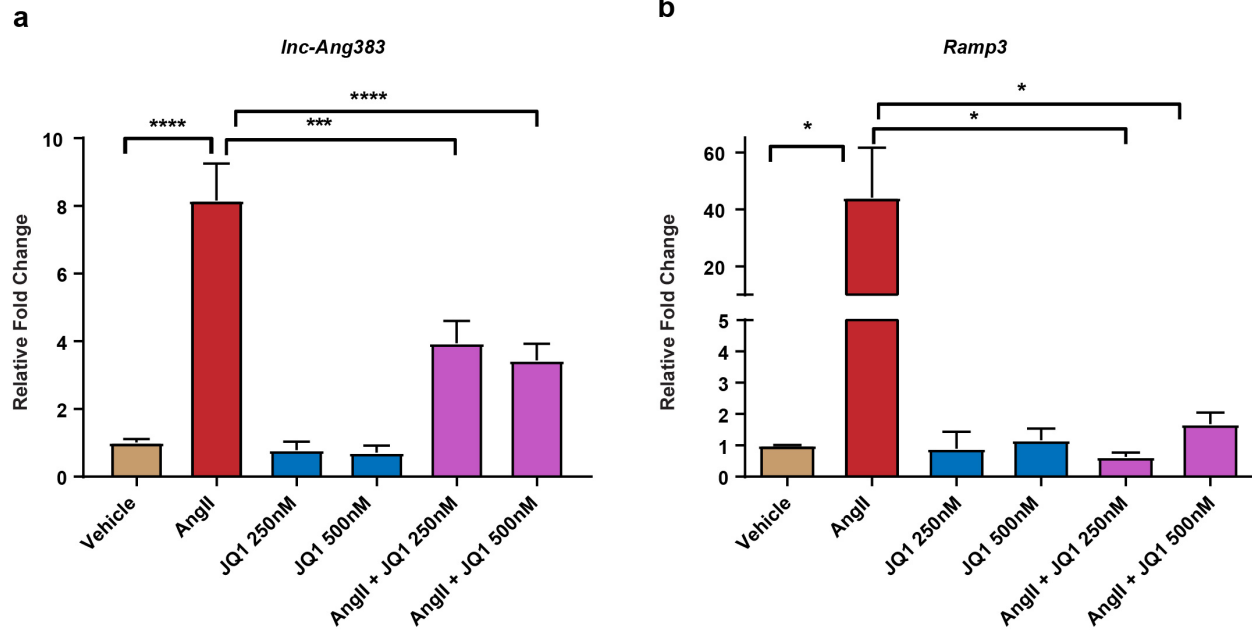


**Supplementary Figure 15. Phenotypes associated with TEs.** IPA Pathway analysis of genes associated with TEs.

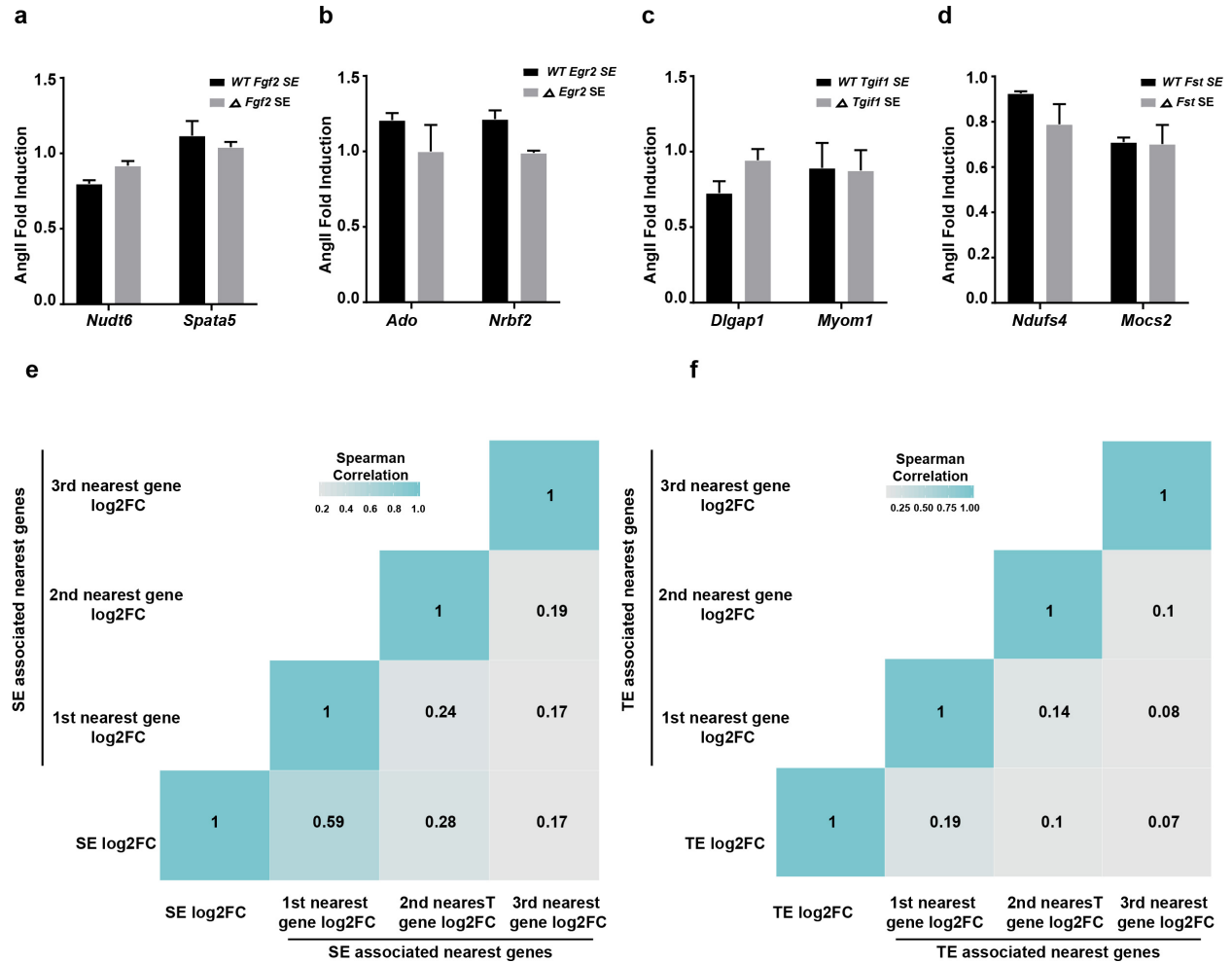


**Supplementary Figure 16. Expression of BET family proteins in VSMCs.** (a-c) *Brd4*, *Brd2* and *Brd3* expression were quantified by RT-qPCRs in RVSMCs treated  $\pm$ AngII for 3 hrs *in vitro*. Mean+SEM; \* $P$ <0.05; vs. Control;  $n$ =3, using unpaired-two tailed t test.





**Supplementary Figure 17. JQ1 attenuates AngII-induced expression of *IncAng383* and *Ramp3*.** (a-b) Bars represent the expression of SE-associated genes measured by RT-qPCR in RVSMCs pre-treated with vehicle (DMSO) or the BRD4 inhibitor JQ1 (250 and 500 nM) and then treated *in vitro* with AngII as indicated. Gene expression was normalized with *Ppia* and represented as relative fold change with respect to Vehicle. Mean+SEM; \* $P < 0.05$ ; \*\*\* $P < 0.001$ ; \*\*\*\* $P < 0.0001$ ;  $n = 3$ , using one-way ANOVA, Tukey's multiple comparisons test.



**Supplementary Figure 18. SEs regulate the first nearest AngII-regulated genes. (a-d)** Bar graphs show the AngII-induced expression of indicated genes in WT and SE deleted ( $\Delta$ SE) RVSMCs. Gene expression was normalized to *Ppia* and expressed as AngII response vs. respective Controls. Mean  $\pm$ SEM,  $n=3$ . Data is not significant, using unpaired two-tailed t tests. **(e)** Correlation heatmaps representing Spearman correlation co-efficients between log2 fold change in H3K27ac occupancy at SEs and **(f)** TEs and the expression changes of 1<sup>st</sup>, 2<sup>nd</sup> and 3<sup>rd</sup> nearest gene mapped within  $\pm$  500 kb from the enhancer centre. For **(e-f)**, all differentially regulated enhancers (SEs/TEs) that had 1<sup>st</sup>, 2<sup>nd</sup> and 3<sup>rd</sup> nearest genes mapped within  $\pm$  500 kb from the enhancer centre were considered regardless of whether the nearest genes were AngII-regulated or not.

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**qRT-PCR expression****Refers to Figure 1 G-J**

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<b>Gene</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
<i>Jun</i>	agggttaagacaccgcttg	ggagaacctctgtcgctg
<i>fos</i>	ctccgtttctctctcttcag	tccttacggactccccac
<i>Ets1</i>	ctctcaactggcaggaagatg	gctcgtcaggtaggattcag
<i>Ets2</i>	acggatgggaattcaagctc	tgcccgatgtcttgtaagt
<i>Ppia</i>	ccgctgtctctttcgcc	gctgtctttggaactttgtctg

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**Refers to Figure 2**

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<b>Gene</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
<i>Esm1</i>	cttcccttcttccagtatgc	gttccggtctccaatctcttc
<i>Spry2</i>	atcagatcttggcgggtgtg	tgtgtaccttgctgttgag
<i>Agtr1a</i>	ggcagtcattctggattctttg	acacaacctcccagaaaag
<i>mEsm1</i> *	ggaagatgtctggactttccc	cgttcccttctccaatctcttc
<i>mSpry2</i> *	atcaggctcttggcagtggtg	agaggattcaaggagagagg
<i>mAgtr1a</i> *	gctggcattttgtctggataac	gcttttctgggttgagttgg
<i>mPpia</i> *	atggtcaaccccaccgtg	ttcttgctgtctttggaactttgtc

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**Refers to Figure 3**

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<b>Gene</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
<i>Fgf7</i>	tgaatactatcttgcctgaacaaag	acgaacatttcccctccg
<i>Slc29a2</i>	tggtgaagggtgatttgagc	tgaggaagagggtgctgtag
<i>Serpine1</i>	cacaagcactacaaaaggtaag	gccgaaccacaaagagaaaag
<i>CD44</i>	cttcagctctccagtaagt	ctccagtcattagacaaccttc
<i>Egr3</i>	attacactcagatggctacagc	ctggaaagagcccgaataagag

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<i>Vcl</i>	aggagaaacagaaaagagggc	ctgtaactcctaacctttcacatg
<i>Clu</i>	tcttgcaactctcacactg	agctttacagttcccggatg
<i>Foxp1</i>	gttttgtgcgagtagagaacg	cattgagaggtgtgcagtagg

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**Refers to Supplementary Figure 7**

<b>Gene</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
<i>Rock2</i>	cacaagaccacaaaagcagc	gagtaggttctccactttctgc
<i>Tnip</i>	ctgatgagaccaccaactg	ggatccctgcttgtatgctc
<i>Extl3</i>	accccacaatgcatcac	ctgcccctggaatgagatagag
<i>Acan</i>	aaccttgcctccaatgactc	ctcggtaaaagtccagtggtg
<i>Rgs17</i>	Cccacacaccacaaaatg	gagaccaggacaagacttcac
<i>Flt1</i>	cctcagcctaccatcaagtg	gccctcgattctgtttcctatg

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**ChIP Primers**

**Refers to Figure 3**

<b>Enhancer proximal to</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
Neg Ctrl	agcttaaagtgtctccagca	atatcaggtcctttgtgaagt
Spry2	actttctgacagagctggatt	tgaggtgtctcctggaatc
FGF7	tccttctccattgggtgatt	ggtacagagacagcagataga
Slc29a2	ggtcctcgacaagaacagaatag	taggaggaggaaggctcaaa
Serpine1	cttactggaagcgaggtgtag	gagacagcacttagaggaatgg
Sgk	ccagactacctgcctattaatctc	cctgagctgagtgatgtaagg
CD44	gcagggatgacacatcagaa	gttggctgtggtgtctcttta
Ezr	cctagtcaggttctgtctgtct	tgttctgtgtccctgtcttac
Esm1A	acatacaggcactcacagaac	gcatggcaactcagaatcac
Esm1B	agcaccaggaaccaactattc	ctcttgagagttcccttctaac
Esm1C	gaaccagaacatcaggaagtc	cagcgtctggcaagtagtaac

Egr3	ctgtgtatctggcatacgatctg	ctccgtgtgtgagtatgtgtatag
Cebpγ	ctgaagtggaaggactgaactg	gtgggtgggcagaatgagaaa
Jun	tggatggatggcacagttac	agagagagagagagagagagaga
Foxp en1	tgtctcttcagggaacataac	aagctgttgggttccttctatc
Foxp en2	ctccttaccttccctttctgttt	ggcccttggctcactgftatg
Ets2	cagtgagttctgcctactctg	caagggtggagacagggttatg
Agtr1aUA	cgactgggtgccttgtatct	ctagggaccagactgaataaac
Agtr1aUB	caggtgccactggatacaaa	accctcacctagatcagtaagt
Agtr1aDA	ctgacagactactctgcaatcc	ctgccttgggatctctcataaa
Agtr1aDB	caagtgccatacctactactg	ggctaaccctcgtcttctatg
Ago2	tggcaggtccttattgtctatc	gggtctcctagagatggacttac
Vcl	aacaggcaggagcaaaaca	gtacgtgtaggttccgtgtag
Clu	cttagctccattccctgacaaa	tctccttccagccctaaa
mEsm1A*	ttgccaccttgaggactattg	gcctgccttctctgtaaat
mEsm1B*	cccgtgggtgaatgagatatt	agcgtagggtgagacaaatg

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### Luciferase Constructs

Refers to Figure 4

Construct	gBLOCK sequence (5'-3')
Ccl2-Prom	ggaggcctaactggccgtacctagatctgctagcctcaggatatacagcggccgctggcaaaatgtctgggagtcaccaaagc agagaatgccatattcaacaagcctgataaccaaggactcagtggactaattggcagtcctatccagatccaaggctccttgagc caggggcaagctaggatgctcccaggatcttctcccttaggactttaggttcttggccacttctcttatttcagtgaaagcagatc cactcattgacacttgtggtcacagtctagcacgactgctcccttctcttttctcctcctgcgcagcttcattgctcccagtagtg gctggaaaaacaccaaattccaatccgcgggttctccttctacttctggaacatccaaggctcggcacttactcagcagattca aacttccacttccatcactcatcgaggatgatgctgctccttggcaccaccacctgcctgactccacccttggcttacaataaaa ggctgaggcagagccgctagaaatgcagagacacagacagaggccagccagaaaccagccaactctcactgaaaagctccc
Spry2-enh	ggaggcctaactggccctgaagttagggctcagtcctgctccaggtctgaactgtcttcccttggcagccttcttggcacacattaaa atgacagccacattccctcactactccctcagcgcctctgtagcagggaggtctgcttaattaggcttacaggcagactgactg ggtaattcttgatcttgagagaagtactttgtattgtaggctgttggtatcaccctggccttcccttctctattgccagcagtgaagtc caaaaaatctaaaggccttgcctaatggaccttgaggcaggggactagacaccaagaccttaagtagataatgttatcagcttc

ttccaagacttagcacetaagtctcagattatcagcttgaaagctcgaggaccatTTTTctctctctcatctgtggcatcacaattta  
caacaataaggtgcacggtaagagtttagtcagtgacgagtagttgcaccctggcctgaaagcttgggttttagcagtactatgga  
tccaacctaggctctcccacatgctagggccaccgctctacctaggagctacagcagttctactgtctcagacagttctgtttcaca  
gcgtagggcaaaagtctcagaaggcctcctactgggctcatcgggtgacttcacagatgataagtaagaagtcagttctgacag  
agctgggtatttcagtgttgfttaaggtttactcagtagctaagagtgcaacaaagtgaaccaatcaggctttagcccaacttgc  
ggccagattccaggagacacctcagctaaagccaagtgtcgaggcatggctgtgtcatctctggagtcagtccttacctgcg  
gccgcaaaaggaaaa

Esm1A-enh ggaggcctaactggcccacctccactcgtcaagtctgaagtgtttgaattgtgggtgtaattctgtatgcataaagtttccgctctg  
tacaattattcagttatggaacacagagactttgacagtgttctccagtcactgccctcaaaggagtgcctagaagctcgggtttct  
tgttctctgttaacccagtcgaccactgacattagccatcatgagcatctgtgtatcgtaccagaaaggaatgtctcaagctcagg  
tgacagaaatgaatcacatacaggcactcacagaacaaaggaacagtattttctcaataataaagacaatccgggaggtcttgc  
atcttcagggcagtgattctgagttgccatgccaggcctctagatgtcctttgttgaatctgtgtgaaactcccagactttgcaac  
tcttgcgtccttttgtgcttatttagatttctattgccatgataaaatacctgacaaaaagcatctaaaggcgaggagaggggttattc  
acctcatagctttagctcctgaagggaagtgcagcaggaattgggtggaacctggaggaggcacctgggagcagggactgaag  
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ccaggagtgccactgccaggagtgccactgccccagtgactgggctctctctatcaattagcaatcaaatgccccaggctt  
gcctatagacgagcctctaagtctgataatgcttttaaaaaatcaattcctgttcagatttaactctactgtgtcaaatcaacaaca  
caacagcagcagtagcagcagcagccaccacctaaccagaacaacatgatttaaaaccagcatcacatggaaaccaccagtc  
agttgctggcttgagtgaatctgggactcctgggaccaaaagacacattgacctccatgcttgacaactgaagatagggaaat  
atatcctggagagggcacttcatgaccagccagtgaaagtaagacacctagatcttagaggctagaggtgctctttctacaggta  
gcatctttcaaggagttactccatatccttgagcctacagtggagcggccgcaaaaggaaaa

Enhancer	Forward Primer (5'-3')	Reverse Primer (5'-3')
Esm1B-enh	ggaggcctaactggcctaccctcctgaatctatg	ctagctagccatagctcccctagtcc
Esm1C-enh	ggaggcctaactggcctaaaatccaagtggctttag	ctagctagcagaggtagcatgaaaacc
Egr3-enh	ggaggcctaactggccttcttctgtccttgg	ctagctagccatctagttcagcaaaagag

**qRT-PCR expression Inc-Ang184 and**

**IncAng383 CRISPR Phenotype**

Refers to Figure 5

Gene	Forward Primer (5'-3')	Reverse Primer (5'-3')
<i>Ccl2</i>	gcagctcagcagaggtagtgg	cggtatagggtctgagaagattaccg
<i>Serpine1</i>	gcagctctctgtagcacaagcc	cggcctctgttgattgtgccg

<i>IL6</i>	ggataccaccacacaacagaccag	cgatgagttttctgacagtgcacatc
<i>lnc-Ang383</i>	atgtccacactgtatgctc	ctgctccacttctctcactatg
<i>lnc-Ang184</i>	cagacctagtggcagtttc	cgctataacctcaacacttcc
<i>Ramp3</i>	gcacctctccctctgttg	tccagattcctgtctcattg

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**Dicer-substrate short-interfering RNAs (DsiRNAs)**

**Refers to Figure 5H**

<b>Name</b>	<b>Sense (5'-3')</b>	<b>Anti-sense (5'-3')</b>
Dsi-lnc-Ang383	gaacuaauagcuaaggcuuuccaaguu	cuuggaaagccuuagcuauuagutc

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**Refers to Supplementary Figure 10**

<b>Gene</b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
<i>lnc-Ang362</i>	gtcccattctgttcctttg	cacactcaccttctattccc

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**CRISPR-sgRNAs**

**Refers to Figure 5 (I-M) and 9 (A-D)**

<b>Enhancer sgRNA<sup>#</sup></b>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
lnc-Ang383SG	caccgactgtttattcgttctctac	aaccgtagagaacgaataaacagtc
lnc-Ang383ASG	caccgcttgattccagcaccgaag	aacccttcggtgctggaatcaaagc
Egr2SG	caccgccccttcccggcggtct	aaccagacacgccgggaagggcgc
Egr2ASG	caccggagcgcacggtatgcgggc	aaccgcccgcacccgatgcgctcc
Fgf2SG	caccgcggtctactgcaagaacgg	aaccccgctcttgtagtagaccgc
Fgf2ASG	caccgtctgggtccataaggctacc	aaccggtagccttatggaccagac
FstSG	caccgagactgcacttccaageta	aacctagcttgagagtgcagtcgc
FstASG	caccgatgcgccagaggttcggct	aaccagccgaacctctgggcgcatc

Tgif1SG	caccgatgetttaatcccatcactc	aaccgagtgatgggattaaagcatc
Tgif1ASG	caccgttcataatacccgaactgta	aacctacagtcgggtattatggaac

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### CRISPR Genotyping

Refers to Supplementary Figure 9

Gene	Forward Primer (5'-3')	Reverse Primer (5'-3')
<i>Fst</i>	ggagcatctgctaagcgtaaa	ctccgacagataattgggaagg
<i>Tgif1</i>	gcattgatacatgccagcaaa	gtgcatgctatgttcgcataat
<i>lnc-Ang383</i>	tacagaagaagacagcacacac	agctaaggctttccaagttaatg
<i>Fgf2</i>	agtccggctgcactagg	ggagggataatgctggagtttg
<i>Egr2</i>	ggtagagagaatggagcagaag	gtagcgtgggtcaagaaaga

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### qRT-PCR expression

Refers to Figure 8 A-N and Supplementary Figure 16 A-D and 18 A-D

SE associated Genes

Gene	Forward Primer (5'-3')	Reverse Primer (5'-3')
<i>Fst</i>	gagtgaagagataggggaagc	agcgagtgtgcatgaag
<i>Egr2</i>	gaaggctggtttctaggcg	aatggtgaactgggagggc
<i>Tgif1</i>	tgacgatgaaaggccaagg	cgaaatgatttacacgagcagg
<i>Fgf2</i>	accacacgtcaactacag	ggcgtcaaagaagaacactc
<i>Flt1</i>	cctcagcctaccatcaagtg	gccctcgattctgttctctatg
<i>Foxp1</i>	gttttgtgcgagtagagaacg	cattgagaggtgtgcagtagg
<i>Brd4</i>	ttgatgcttgagttgtgtttgg	catgatagtccaggcaaaggg
<i>Brd2</i>	gaagcgggaactagagaagc	gaactagaagcactgagacgg
<i>Brd3</i>	tcctctttcgatttgctgc	cccagacgagattgagattgac
<i>Nudt6</i>	ggtgctaacaatcaatacggc	ctctgggcttctgcttcc



<i>Spata5</i>	ccagtgctccttaccagttg	cagcttcacacccacattc
<i>Ado</i>	gcctggacgatggattactg	gcagtgacaaaaccaggaatc
<i>Nrbf2</i>	gggtctatggaagtgatggaag	tcagttcatggcttcagag
<i>Dlgap1</i>	gccacactttgaatgctacac	agaccgaaggagaaaatgcc
<i>Myom1</i>	tgattacggctattcccatgg	accatgtagttgcctgacag
<i>Ndufs4</i>	gatcccaggcattcatcgtag	aactacaggtacacttggcag
<i>Mocs2</i>	tgccaagaagctctcagtg	tgccatgggtagatatgcttc

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**ChIP**

**Refers to Figure 8 O-P**

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<i>SE Regions</i>	<b>Forward Primer (5'-3')</b>	<b>Reverse Primer (5'-3')</b>
Tgif1	agtaaggttgagtcgggatttg	tgccctccacaaagtgaataa
Fgf2	gcagcagtgttcttgcctttg	gtgaggtgggcttgtttact
Foxp1	atgtggaacctctccactttac	ccagtcctacagcattgat

**Supplementary Table 1. List of oligonucleotide sequences used in this study.** \* *m* denotes mouse genes. # SG denotes sense guide RNA and ASG denotes anti-sense guide RNA.

<b>GWAS chromosome coordinates</b>	<b>GWAS ID</b>	<b>PubMed ID</b>	<b>Authors (et al.)</b>	<b>Journal</b>	<b>Trait</b>	<b>Genes</b>	<b>hg19 co-ordinates</b>	<b>Rat enhancer co-ordinates</b>
chr9_22115958_22115959	rs2383207	20622881	Gretarsdottir S	Nat Genet	Abdominal aortic aneurysm	CKDN2B, CDKN2A	chr9_22114267_22116382	chr5_109036837_109039175
chr5_39397131_39397132	rs11959928	20383146	Kottgen A	Nat Genet	Chronic kidney disease	DAB2, C9	chr5_39397114_39401158	chr2_55707758_55711091
chr1_15095147_150951477	rs267734	20383146	Kottgen A	Nat Genet	Chronic kidney disease	SETDB1, LASS2, ANXA9, PRUNE, FAM63A, BNIPL	chr1_15094951_150952124	chr2_190219397_190221430
chr21_3559382_35593827	rs28451064	26343387	Nikpay M	Nat Genet	Coronary artery disease	KCNE2	chr21_3559306_35594753	chr11_32159399_32162443
chr9_2211528_22115286	rs944797	21971053	Takeuchi F	Eur J Hum	Coronary heart disease	CDKN2A, CDKN2B	chr9_22114267_22116382	chr5_109036837_109039175

				Genet				
chr2_62713532_62713533	rs6545946	22412388	Kenny EE	PLoS Genet	Crohn's disease	TMEM17, EHBP1, CPAMD8, AK3	chr2_62712712_62716292	chr14_10319761_1_103200563
chr1_83220655_83220656	rs11582609	26305897	Iyengar SK	PLoS Genet	Diabetic kidney disease	intergenic	chr1_83217751_83221886	chr2_246266690_246271012
chr20_10965997_10965998	rs1887320	25249183	Lu X	Hum Mol Genet	Diastolic blood pressure	JAG1	chr20_10957868_10966395	chr3_125541543_125548196
chr9_22115104_22115105	rs10965235	20601957	Uno S	Nat Genet	Endometriosis	CDKN2B AS	chr9_22114267_22116382	chr5_109036837_109039175
chr1_8150637_8150638	rs72634258	26192919	Liu JZ	Nat Genet	Inflammatory bowel disease	NR	chr1_8149972_8152812	chr5_167965360_167968601
chr9_22114468_22114469	rs10733376	25256182	Foroud T	Stroke	Intracranial aneurysm	CDKN2B AS, ANRIL	chr9_22114267_22116382	chr5_109036837_109039175
chr21_35593826_35593827	rs28451064	26343387	Nikpay M	Nat Genet	Myocardial infarction	KCNE2	chr21_35593064_35594753	chr11_32159399_32162443

chr13_102145559_102145560	rs1335587	23251661	Comuzzie AG	PLoS One	Obesity-related traits	ITGBL1	chr13_102123151_102146733	chr15_108803525_108814397
chr10_44458713_44458714	rs898549	23251661	Comuzzie AG	PLoS One	Obesity-related traits	C10orf136	chr10_44457081_44461770	chr4_153840182_153845918
chr14_94972241_94972242	rs11627075	23251661	Comuzzie AG	PLoS One	Obesity-related traits	SERPINA12	chr14_94972049_94977481	chr6_128102375_128105662
chr5_16913854_16913855	rs17614462	23251661	Comuzzie AG	PLoS One	Obesity-related traits	MYO10	chr5_16901265_16914607	chr2_77201740_77210481
chr5_131430117_131430118	rs657075	24390342	Okada Y	Nature	Rheumatoid arthritis	CSF2, IL3	chr5_131429842_131432784	chr10_39641675_39644527
chr5_131430117_131430118	rs657075	22446963	Okada Y	Nat Genet	Rheumatoid arthritis	CSF2	chr5_131429842_131432784	chr10_39641675_39644527
chr10_63785088_63785089	rs10821944	22446963	Okada Y	Nat Genet	Rheumatoid arthritis	ARID5B	chr10_63784082_63790216	chr20_21238868_21245054

chr1_20550929 7_205509298	rs1815022 28	26606652	Alarcon- Riquelme ME	Arthritis Rheumat ol	Systemic lupus erythematosus	DSTYK	chr1_20550622 9_205515389	chr13_44982874 _44986847
chr20_1096599 7_10965998	rs1887320	25249183	Lu X	Hum Mol Genet	Systolic blood pressure	JAG1	chr20_1095786 8_10966395	chr3_125541543 _125548196
chr2_12640740 _12640741	rs1534422	19430480	Barrett JC	Nat Genet	Type 1 diabetes	intergenic	chr2_12629402 _12645547	chr6_39702585_ 39708724
chr22_3058172 1_30581722	rs5753037	19430480	Barrett JC	Nat Genet	Type 1 diabetes	intergenic	chr22_3058136 9_30583153	chr14_84958839 _84960817
chr10_7145228 4_71452285	rs2812533	24509480	Mahajan A	Nat Genet	Type 2 diabetes	C10orf35	chr10_7144747 6_71452665	chr20_29456788 _29457999

**Supplementary Table 2: GWAS SNPs associated with AngII regulated enhancers. GWAS SNPs present in human sequences orthologous to rat enhancers were identified. Only diabetes and CVD associated SNPs are shown.**