

Supplementary Materials:

Method of transcriptome sequencing

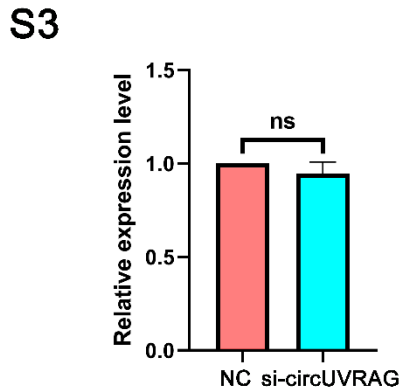
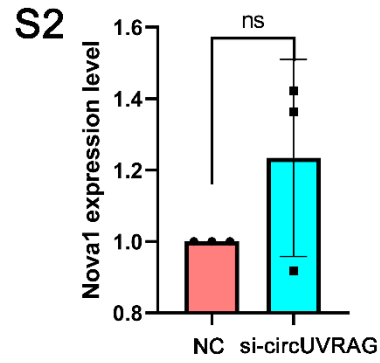
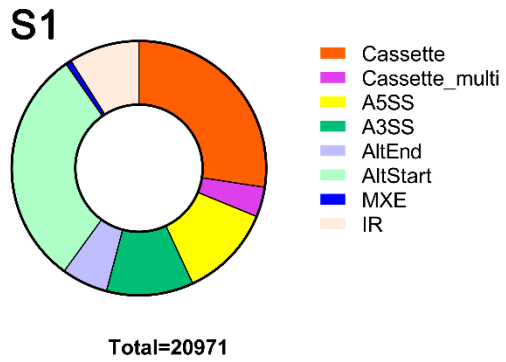
4 pairs of graft veins and control were obtained after one week of surgery. RNA from samples extracted by Trizol reagent (Invitrogen) with RIN>6.0 was utilized to construct rRNA depletion library (NEBNext Ultra Directional RNA Library Prep Kit) according to the manufacturer's instructions. Whole transcriptome sequencing data sequenced by HiSeq™ Sequencer was filtered (removing the adaptor sequences, reads with >5% ambiguous bases (noted as N) and low-quality reads containing more than 20 percent of bases with qualities of <20) and mapped to rat genome (Btgenome Version 5.0.1 NCBI) utilizing HISAT2 was used to calculate the gene count of mRNA and non-coding RNAs. NovelBrain Cloud Analysis Platform (www.novelbrain.com, NovelBioinformatics Ltd., Co.) was used for the bioinformatics analysis. DE genes, including mRNA, miRNA, and circRNA, in the vein graft and control were analyzed utilizing DESeq with the following criteria: fold change > 2, $p < 0.05$, and false discovery rate (FDR) < 0.05.

Supplementary Fig. S1. Our sequencing results predicted abundant alternative mRNA splicing events in graft vein. Cassette: cassette exon, i.e. Skipped exon; Cassette_multi: multiple adjacent Cassette exons; A5SS: Alternative 5' splice site; A3SS: Alternative 3' splice site; AltStart: Alternative start exon; AltEnd: Alternative end exon; MXE: Mutually exclusive exons; IR: intron retention

Supplementary Fig. S2. After knockdown of circUVRAG, the expression of NOVA1 mRNA did not change significantly, which indicated that NOVA1 was not regulated by circUVRAG(n=3).

Supplementary Fig. S3. Knockdown of circUVRAG had no significant effect on the VSMC proliferation detected with the BrdU assay (n=4). Colorimetric BrdU kit (Roche Diagnostics) was used to detect the proliferation of VSMCs. Enzyme linked immunosorbent assay plate reader (Bio-Rad 680) was used to measure the absorbance of culture medium with a wave length of 450 nm and 630 nm.

Supplementary Fig. S4. The sequence of circUVRAG in rat.



S4

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>chr1_164064815_164025518_-39297-Uvrag
AATTTTATAGGAGTGAAGTGATTAAGAATTC
CTTGAATCCAACGTGGAGGAGTCTTGACT
TTGGAATAATGCCGGACCGTCTGGATACAT
CTGTGTCTGCTTTGTGGTGAAGATTTGGG
GTGGAAAGGAAGAGGCCTATCAGCTGTTGA
TAGAGTGGAAAGTCTACTTGGATGGGCTGA
AGTACTTGGGT CAGCAGATCCATGCCCGCA
ACCAGAAT GAAATCATT TTTGGGCTGAACG
ATGGCTACTACGGT GCTCCATTT GAACACA
AGGCTCATCCAAATGCACAGAAGAACCTCC
TTCAGGTGGACCAGAACTGTGTTCCGAATT
CCTATGATGTGTTTTCTTTGCTGCG
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Supplementary Table S1. PCR primer sequence, siRNA sequence, and FISH probe sequence

PCR primer sequence, siRNA sequence, and FISH probe sequence		
Primer sequence		
circUVRAG	Forward	TGGACCAGAACTGTGTTTCGCA
	Reverse	GCCTCTTCCTTTCCACCCCA
uvrag-mRNA	Forward	GGCTACTACGGTGCTCCATT
	Reverse	CATTGCTTGTCGATGTGAGTCT
uvrag-premRNA	Forward	GGCTGAACGATGGCTACTACG
	Reverse	CCGACTTCTCAGAGTGCTTGTG
uvrag for gDNA	Forward	GGAATAATGCCGGACCGTCT
	Reverse	ACAGCTGATAGGCCTCTTCCT
nova1	Forward	GCAGCACCAAGAGGACCAACAC
	Reverse	CTTGATCGTGGCTCCCGTTTCC
gapdh	Forward	TGAACTTGCCGTGGGTAGAG
	Reverse	GATGGTGAAGGTCGGTGTGA
siRNA sequence		
Negative control	Forward	UUCUUCGAACGUGUCACGUTT
	Reverse	ACGUGACACGUUCGGAGAATT
siRNA1 for circUVRAG	Forward	UUCUUUGCUGCGAAUUUUUATT
	Reverse	UAAAAUUCGCAGCAAAGAATT
siRNA2 for circUVRAG	Forward	UUGCUGCGAAUUUUUAUAGGTT
	Reverse	CCUAUAAAAUUCGCAGCAATT

siRNA1 for nova1	Forward	GCUGCUCAGUAAUUAAUUATT
	Reverse	UAAUUAUUACUGAGCAGCTT
siRNA2 for nova1	Forward	GGUCCUCAUACCUAGUUUATT
	Reverse	AUAACUAGGUAUGAGGACCTT
siRNA3 for nova1	Forward	GCUGCAACCAAUGGAUACUTT
	Reverse	AGUAUCCAUUGGUUGCAGCTT
Fish probe sequence		
Negative control	sense (5'-3')	TGCTTTGCACGGTAACGCCTGTTTT
circUVRAG probe mix	sense (5'-3')	GGCCCCAAATCAGAGTCCAT
	sense (5'-3')	CCGGGGAAGGAAGGGAAGAG
	sense (5'-3')	CCCCAAATCAGAGTCCATGG

Supplementary Table S2. The homologous circUVRAG in humans and mice

Position (genome browser link)	Strand	circRNA ID	Genomic length	Spliced length	Best transcript
chr7:10624782 1-106266815	-	mmu_circ_0013695	18994	315	ENSMUST000 00037968
chr7:10621430 9-106247983	-	mmu_circ_0001601	33674	320	NM_178635
chr11:7557279 0-75672593	+	hsa_circ_0023635	99803	464	NM_003369
chr11:7557279 0-75623083	+	hsa_circ_0023634	50293	358	NM_003369
chr11:7556292 7-75694557	+	hsa_circ_0023633	131630	709	NM_003369
chr11:7556292 7-75623083	+	hsa_circ_0096459	60156	476	NM_003369
chr11:7556292 7-75599947	+	hsa_circ_0140972	37020	390	NM_003369
chr11:7556292 7-75599947	+	hsa_circ_0096458	37020	390	NM_003369

Supplementary Table S3. The top 10 RNA Binding Proteins predicted by catrapid.

Protein	Transcript	Z-score	Discriminative Power	Interaction Strength	Motif
sp P63155 C RNL1_RAT	chr1_164064815_ _1	0.91	0.96	0.88	NO
sp Q496Z9 T RM1L_RAT	chr1_164064815_ _1	0.81	0.96	0.8	NO
sp Q99PF5 F UBP2_RAT	chr1_164064815_ _1	0.78	0.95	0.79	NO
sp Q641Y8 D DX1_RAT	chr1_164064815_ _1	0.71	0.95	0.85	NO
sp Q7TP47 HNRPQ_RA T	chr1_164064815_ _1	0.64	0.93	0.87	NO
sp P51400 R ED1_RAT	chr1_164064815_ _1	0.62	0.92	0.73	NO
sp B2RYD2 ESRP1_RAT	chr1_164064815_ _1	0.57	0.91	0.85	NO
sp P13383 N UCL_RAT	chr1_164064815_ _1	0.51	0.9	0.68	NO
sp Q5XI28 R AVR1_RAT	chr1_164064815_ _1	0.48	0.89	0.78	NO
sp P68101 IIF 2A_RAT	chr1_164064815_ _1	0.45	0.88	0.97	NO