

## Original Article

# Identification of lncRNA-NR\_104160 as a biomarker and construction of a lncRNA-related ceRNA network for essential hypertension

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Received September 10, 2019; Accepted June 3, 2020; Epub October 15, 2020; Published October 30, 2020

**Abstract:** Objectives: To identify long noncoding RNAs (lncRNAs) and construct a competing endogenous RNA (ceRNA) network for essential hypertension. Methods: An RNA microarray and two-step quantitative real-time PCR were applied to identify differentially expressed RNAs (DE-RNAs), and a luciferase assay was performed to explore the binding relationship between RNAs. A generalized linear model and logistic regression model were used to analyze the associations between different RNAs and of RNAs with hypertension. Receiver operating characteristic curve analysis was executed to evaluate the diagnostic performance. Bioinformatics analysis was applied for network construction. Results: In total, 439 DE-RNAs (387 lncRNAs and 52 mRNAs) were identified in the microarray, and 71 'lncRNA-miRNA-mRNA' loops formed the ceRNA network. The first validation confirmed that five RNAs (NR\_104160, Inc-GPR63-8:1, Inc-Hprt1-9:1, ID1 and RSL24D1) were significantly upregulated in hypertensives ( $P < 0.05$ ). NR\_104160 was significantly associated with hypertension (OR = 2.863, 95% CI: 1.143-7.172;  $P = 0.025$ ) after adjusting for confounding factors. NR\_104160 was included in the hypertension diagnostic model, with an area under the curve of 0.852 (95% CI: 0.761-0.944). In the second validation, NR\_104160 showed a constant significant difference ( $P = 0.001$ ). An elevated expression level of NR\_104160 was associated with the expression of ID1 ( $\beta = 0.2235$ ,  $P = 0.005$ ). Luciferase assays showed hsa-miR-101-3p stimulation significantly inhibited the reporter gene activation ability of the NR\_104160 wild-type plasmid ( $P < 0.001$ ). Conclusions: Our study constructed a ceRNA network to provide hypotheses regarding the mechanism of hypertension development. IncRNA-NR\_104160 was identified as a hub element that participates in hypertension transcriptional regulation and as a potential biomarker.

**Keywords:** lncRNA, hypertension, ceRNA, network, biomarker

## Introduction

Essential hypertension is a chronic noncommunicable disease caused by genetic and environmental factors [1]. The prevalence of hypertension in China is 39.7% [2]. As a major risk factor for stroke, myocardial infarction, heart failure and end-stage renal disease [3], hypertension has become a serious public health problem that needs to be addressed. This highlights the urgent need to identify biomarkers that predict the key factors of hypertension occurrence and therapy.

Several studies have confirmed that long non-coding RNAs (lncRNAs) participate in the development of hypertension, acting as biomarkers, potential therapeutic targets or strong indicators of prognosis [4]. Researchers have reported that microvascular dysfunction induced by hypertension is aggravated by knockdown of lncRNA GAS5 [5], and the expression of GAS5 in hypertensives was significantly reduced. LncRNA AK098656 is a human vascular smooth muscle cell (VSMC)-dominant lncRNA that is increased in hypertensive patients [6]. However, the underlying molecular mechanism related to

lncRNAs in the context of hypertension remains largely unknown.

Recently, an abundance of studies have focused on the competing endogenous RNA (ceRNA) hypothesis of diseases, which claimed that RNA transcripts (including mRNA, pseudogenes, lncRNA and circular RNA) can competitively combine with microRNAs (miRNAs) via miRNA response elements (MREs) [7]. According to the ceRNA mechanism, a new network-based regulatory pattern has been recognized, thus forming a ‘ceRNA-miRNA-mRNA’ network. Growing evidence has shown that this novel regulatory mechanism underlying the crosstalk among lncRNAs, miRNAs and mRNAs plays a pivotal role in the pathophysiological processes of cardiovascular diseases (CVDs) [8, 9]. However, few studies have reported on the ceRNA network in hypertension.

Therefore, this study aimed to identify differentially expressed lncRNAs (DE-lncRNAs) in the context of hypertension and systematically analyze the ‘lncRNA-miRNA-mRNA’ ceRNA network involved in hypertension.

## Materials and methods

### *Patients and samples for microarray and quantitative real-time PCR*

All participants were selected from the System Epidemiology Study on Salt Sensitivity of Blood Pressure (EpiSS) cohort study, which was set up by our research group, and details can be found in the previously published protocol [10]. In brief, 20 subjects (10 hypertensive and 10 normotensive individuals) were involved in creating the microarray, and two-step validation was conducted in 70 subjects (42 hypertensive and 28 normotensive individuals) and 56 subjects (35 hypertensive and 21 normotensive individuals) by quantitative real-time PCR (*qRT-PCR*). All participants were 35–70 years old. Primary hypertensive patients with systolic blood pressure (SBP) in the range of 140–159 mmHg or diastolic blood pressure (DBP) 90–99 mmHg were enrolled in this study. In addition, individuals with CVDs, kidney disease, liver disease, malignant tumor or pregnancy or who had used antihypertensive drugs in the past month were excluded. This study was approved by the ethics committee of Capital Medical University, and all subjects provided written informed consent.

Blood samples were collected in the early morning after the participants had fasted for eight hours. Serum samples were isolated and used to perform biochemical examination, including the assessment of total cholesterol (TC), triglycerides (TG), high-density lipoprotein cholesterol (HDL-C), low-density lipoprotein cholesterol (LDL-C) and fasting blood glucose (FBG). Blood samples (2.5 mL) were collected in a PAXgene Blood RNA Tube (PreAnalytiX GmbH, Hombrechtikon, Switzerland) where they were maintained until RNA extraction was performed.

### *RNA isolation and quality control assay*

Total RNA was extracted and purified using a PAXgene Blood RNA Kit (cat. no. 762174, QIAGEN GmbH, Hilden, Germany). The RNA integrity number (RIN) of each RNA was assessed to evaluate RNA integration using an Agilent Bioanalyzer 2100 (Agilent Technologies, Santa Clara, CA, USA). RNA quantity was measured using a NanoDrop ND-1000 UV-VIS spectrophotometer (Nanodrop, ND1000) and an Agilent Bioanalyzer 2100 (Agilent Technologies, Santa Clara, CA, USA). If the RIN was more than seven and the 28 s/18 s ratio was more than 0.7, the RNA extract was considered adequate and was used for further analysis.

### *Microarray of expression profiles*

The microarray used in the current study was SBC human (4\*180K) ceRNA array v1.0, which contains 68,423 lncRNAs and 18,853 mRNAs. The microarray analysis was performed according to the standard procedure. First, cDNA was generated via reverse transcription of RNA obtained from samples using a Low-Input Quick Amp WT Labeling kit (One-Color, Agilent Technologies Santa Clara, USA). Then, the cDNA was hybridized to the SBC human (4\*180K) ceRNA array v1.0. Finally, the hybridized arrays were washed, fixed and scanned with an Agilent Microarray Scanner (Agilent Technologies, Santa Clara, CA, USA).

### *Significant differential gene analysis*

Raw data were normalized using a quantile normalization algorithm, and differential gene expression analysis of the microarray data was conducted using the Empirical Bayes method; both analyses were run using the limma package [11] in R software (<http://www.bioconductor.org/packages/2.14/bioc/html/limma.html>).

conductor.org/packages/release/bioc/html/lmma.html). In our study, DE-lncRNAs were identified by the following criteria: (i) Foldchange > 2 or < 0.5; (ii) P-value < 0.05; and (iii) Filtering genes satisfying the following two conditions [12]: Genes with an expression variability among 20 samples lower than the median of all the expression differences calculated for each RNA, and genes with a mean expression signal among 20 samples lower than the median of all the expression signals calculated for each RNA. The screening criteria for differentially expressed mRNAs (DE-mRNAs) were as follows: (i) Foldchange > 2 or < 0.5; (ii) P-value < 0.05. The heatmap of the top 10 upregulated and downregulated DE-lncRNAs and DE-mRNAs was drawn by the pheatmap package in R software.

#### *Prediction of lncRNA target genes*

We selected genes regulated by cis-regulation that were 10 kb upstream and downstream from the lncRNA as the target genes [13]. Trans-regulation [14] is the lncRNA-mediated transcriptional activation and expression regulation of coding genes on other chromosomes. Generally, the gene sequence of the corresponding species (human) in the database is used. First, the complementary or similar sequence is selected by BLAST. Second, the complementary energy between the two sequences is calculated by RNAPlex [15]. Third, we selected genes with energy  $\leq -30$  as the target genes regulated by trans-regulation.

#### *Functional annotation of targeted genes*

Gene Ontology (GO) enrichment and Kyoto Encyclopedia of Genes and Genomes (KEGG) pathway enrichment analyses for genes targeted by DE-lncRNAs were performed using the Database for Annotation, Visualization and Integrated Discovery [16, 17] (DAVID, <https://david.ncifcrf.gov/>).

#### *Coexpression network construction*

The lncRNA-mRNA coexpression network was constructed based on DE-lncRNAs and DE-mRNAs. We paired all DE-lncRNAs and DE-mRNAs, calculated the Pearson's correlation coefficient (PCC) for each pair, and then choose the significantly related pairs ( $PCC > 0.95$

or  $PCC < -0.95$ ) to construct the coexpression network using Cytoscape software version 3.4.0 [18].

#### *CeRNA network construction*

All lncRNAs and mRNAs in the ceRNA network were from the lncRNA-mRNA coexpression network. Each 'lncRNA-miRNA-mRNA' loop was built using the following steps: (i) Predicting the target miRNAs of the lncRNA by miRDB (<http://mirdb.org/miRDB/>); (ii) Predicting the target mRNAs of the abovementioned miRNAs by miRDB, TarBase (<http://diana.imis.athena-innovation.gr/>) and TargetScan ([http://www.targetscan.org/vert\\_71/](http://www.targetscan.org/vert_71/)); and (iii) Finding the intersection of the abovementioned targeted mRNAs and mRNAs that have a coexpression relationship with the lncRNA in step (i). Then, we combined these loops together and constructed a 'lncRNA-miRNA-mRNA' ceRNA network using Cytoscape software.

#### *Quantitative real-time PCR*

The expression of RNAs was detected by an ABI 7900HT Real-Time PCR System (Applied Biosystem, Foster City, CA, USA) using SYBR Green PCR Master Mix (ABI, 4368708), and glyceraldehyde 3-phosphate dehydrogenase (GAPDH) was used as the internal control. The relative expression level was determined with the  $2^{-\Delta\Delta Ct}$  method [19]. Each sample was performed in triplicate, and the average value was calculated. The primer sequences for qRT-PCR are summarized in [Table S1](#).

#### *Luciferase reporter assay*

HEK 293T cells were seeded into 24-well plates and cotransfected with plasmids (GV272-NR\_104160-WT, GV272-NR\_104160-Mut, GV272-ID1-3UTR-WT, GV272-ID1-3UTR-Mut) and miRNA mimics (hsa-miR-101-3p) using X-tremeGENE HP (cat. no. 06366236001, Roche, Basel, Switzerland). Luciferase activity was measured using the Dual-Luciferase<sup>®</sup> Reporter Assay System (cat. no. E2910, Promega, Madison, WI, USA) after 48 h of incubation according to the manufacturer's instructions. Independent experiments were performed in triplicate, and the average value was calculated. Relative luciferase activity was normalized to the Renilla luciferase internal control.

### Statistical analysis

All statistical analyses were performed using SPSS 20.0 (SPSS, Inc., Chicago, IL, USA), R software (version 3.4.4) and SAS (version 9.4). A *P*-value < 0.05 indicated a statistically significant difference, and all *P*-values were two-sided. Measurement data are presented as the mean ± standard deviation. Normally distributed data were analyzed using Student's *t*-tests. Nonnormally distributed data were analyzed using a Wilcoxon rank sum test. Enumeration data were compared using a Pearson  $\chi^2$  test. The differentially expressed RNAs in the microarray data were tested by the Bayesian adjusted *t*-statistics from the linear models for Microarray data (limma) package, and multiple testing correction based on the false discovery rate was performed. PCC between RNAs was calculated in the correlation analysis. Differential expression levels of RNAs between groups in qRT-PCR validation were analyzed using an independent two-sample *t*-test. The association between RNAs and hypertension was analyzed using an unconditional logistic regression model and adjusted for age, sex, TC, TG, BMI, HDL-C, LDL-C, FBG, smoking, family history of hypertension, and the consumption of sauce and poultry. Odds ratios (ORs) represent the risk of developing hypertension. Receiver operating characteristic (ROC) curve analysis was performed to determine the differences in the area under the curve (AUC). A generalized linear model (GLM) was conducted to explore the associations between the mRNA and lncRNA expression levels.

## Results

### Characteristics of subjects

We enrolled ten paired hypertensive patients and healthy control individuals for the microarray analysis and 70 (42 hypertensive and 28 normotensive individuals) and 56 subjects (35 hypertensive and 21 normotensive individuals) for separate qRT-PCR analyses. The characteristics of the subjects are summarized in **Tables 1** and **S2**. Variables in the microarray test were balanced between the two groups except for SBP (*P* = 0.030) and TC (*P* = 0.037). Among the samples in the qRT-PCR test, SBP, DBP, HDL-C, LDL-C and sauce consumption per month differed between the two groups in the first validation group (*P* < 0.05), while SBP, TC,

TG and HDL-C showed significant differences in the second validation group (*P* < 0.05).

### Overview of lncRNA and mRNA expression profiles

In total, 387 DE-lncRNAs and 52 DE-mRNAs were identified (**Figure 1A** and **1D**; **Table S3**). Of these, 355 DE-lncRNAs and 39 DE-mRNAs were upregulated, while 32 DE-lncRNAs and 13 DE-mRNAs were downregulated. The DE-lncRNAs were found to be widely distributed among all chromosomes, and chromosome (chr) 1 and chr3 were the top two chromosomes (**Figure 1B**). Among the 387 DE-lncRNAs, 170, 77, 59, 55, 14 and 12 DE-lncRNAs belong to the intergenic, exonic sense, intronic antisense, intronic sense, exonic antisense and bidirectional groups, respectively. For DE-mRNAs, there was no distribution on chr14, chr16, or chr21, with chr12 and chr19 being the top two chromosomes (**Figure 1E**). **Table 2** shows the information of the top 20 DE-lncRNAs and DE-mRNAs, and their heatmap results are shown in **Figure 1C** and **1F**.

### Functional annotation of targeted genes of DE-lncRNAs

A total of 311 target genes of DE-lncRNAs were identified by cis-regulation and 671 genes by trans-regulation (**Table S4**). Only 38 DE-lncRNAs are associated with trans-regulated genes, suggesting that one lncRNA is associated with many target genes via a trans-regulatory mechanism. The GO and KEGG pathway enrichment analyses suggested that these genes might play roles in some important GO terms ('response to estrogen', 'endothelial cell proliferation', etc.) and KEGG pathways ('HIF-1 signaling pathway', etc.) (**Figure 2**).

### Construction of the lncRNA-mRNA coexpression network

A total of 102 DE-lncRNAs, 12 DE-mRNAs (*ID1*, *CD69*, *SNRPF*, *GZF1*, *EGR3*, *PTS*, *RSL24D1*, *ERMN*, *CDADC1*, *SFN*, *C12orf79* and *RPL17*) and 195 pairs formed the coexpression network (**Figure 3** and **Table S5**). All the DE-lncRNAs and DE-mRNAs were upregulated in the hypertensive group compared to the control group, and each lncRNA-mRNA pair was positively correlated in the expression profile. The *RSL24D1* and *RPL17* genes were associated

**Table 1.** Clinical characteristics of study participants in the hypertensive and normotensive groups of microarray test

Variables	Hypertensive (N = 10)	Normotensive (N = 10)	Total (N = 20)	P-value
Gender (female, n (%))	6 (60)	6 (60)	12 (60)	1.000 <sup>‡</sup>
Age (years)	64.20 ± 1.55	64.90 ± 2.77	64.55 ± 2.21	0.494 <sup>*</sup>
SBP (mmHg)	143.30 ± 13.96	129.20 ± 12.69	136.25 ± 14.86	0.030 <sup>*</sup>
DBP (mmHg)	79.80 ± 7.001	74.10 ± 7.676	76.95 ± 7.72	0.100 <sup>*</sup>
BMI (kg/m <sup>2</sup> )	24.88 ± 2.70	24.19 ± 1.98	24.54 ± 2.33	0.523 <sup>*</sup>
TC (mmol/L)	3.45 ± 2.02	5.21 ± 1.38	4.33 ± 1.91	0.037 <sup>*</sup>
TG (mmol/L)	2.83 ± 1.99	1.48 ± 0.64	2.15 ± 1.60	0.067 <sup>*</sup>
HDL-C (mmol/L)	2.18 ± 1.38	2.77 ± 0.98	2.48 ± 1.20	0.218 <sup>†</sup>
LDL-C (mmol/L)	2.17 ± 1.11	1.95 ± 1.10	2.06 ± 1.08	0.326 <sup>†</sup>
FBG (mmol/L)	6.01 ± 1.17	5.70 ± 0.74	5.85 ± 0.97	0.545 <sup>†</sup>
Heart rate	73.60 ± 6.15	74.90 ± 7.37	74.25 ± 6.64	0.674 <sup>*</sup>
Smoke (yes, n (%))	5 (50)	3 (30)	8 (80)	0.650 <sup>‡</sup>
Family history of hypertension (yes, n (%))	6 (60)	4 (40)	10 (50)	0.656 <sup>‡</sup>
Salt consumption per month				1.000 <sup>‡</sup>
≤ 500 gram	5 (50)	4 (40)	9 (45)	
> 500 gram	5 (50)	6 (60)	11 (55)	
Pickle consumption per month				0.650 <sup>‡</sup>
≤ 250 gram	7 (70)	5 (50)	12 (60)	
> 250 gram	3 (30)	5 (50)	8 (40)	
Sauce consumption per month				1.000 <sup>‡</sup>
≤ 150 gram	7 (70)	7 (70)	14 (70)	
> 150 gram	3 (30)	3 (30)	6 (30)	
Poultry consumption per week				0.656 <sup>‡</sup>
≤ 3 times	6 (60)	4 (40)	10 (50)	
> 3 times	4 (40)	6 (60)	10 (50)	

BMI, body mass index; TC, total cholesterol; TG, triglycerides; HDL-C, high-density-lipoprotein-cholesterol; LDL-C, low-density-lipoprotein-cholesterol; FBG, fasting blood glucose; SBP, systolic blood pressure; DBP, diastolic blood pressure. P < 0.05 was considered statistically significant. \*, Statistical testing by independent-samples t test. †, Statistical testing by Wilcoxon rank sum test. ‡, Statistical testing by  $\chi^2$  test.

with 64 and 86 DE-lncRNAs, respectively, and could be the central nodes in the coexpression network.

#### Construction of the ceRNA network

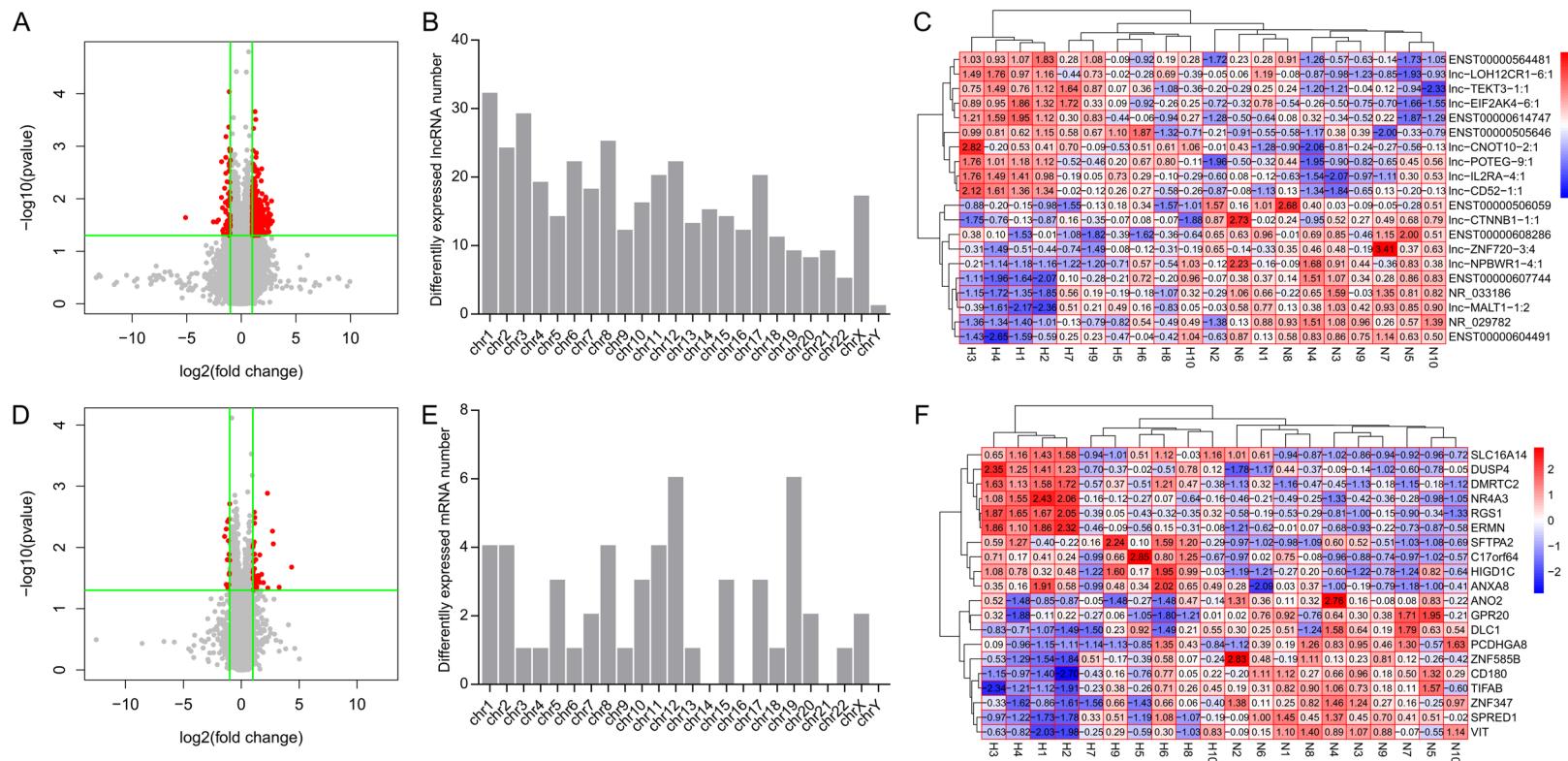
In total, 485 lncRNA-miRNA interactions (62 lncRNAs and 381 miRNAs) were identified in miRDB (Table S6), and 54 miRNAs of the above pairs were associated with the aforementioned 12 DE-mRNAs in the coexpression network, thus forming the ceRNA network, which was composed of 10 mRNAs, 30 lncRNAs, and 54 target miRNAs, including 71 'lncRNA-miRNA-mRNA' ceRNA circular pathways (Figure 4 and Table S7). In the ceRNA network, NR\_104160 was associated with seven ceRNA circular pathways, making it the top lncRNA. RSL24D1 was involved with 44 ceRNA loops, making it the top mRNA. Furthermore, five miRNAs (hsa-miR-4426, hsa-miR-335-3p, hsa-miR-4662b,

hsa-miR-4647 and hsa-miR-103a-2-5p) were related to three ceRNA pathways, making them the top miRNAs.

#### The results of selected RNA relative expression levels in the first qRT-PCR

To analyze the association between hypertension and ceRNA pathways, we decided to validate NR\_104160 and ID1 by qRT-PCR, which both interacted with hsa-miR-101-3p. Moreover, we reviewed the literature on the aforementioned five miRNAs and decided to identify RNAs (lnc-GPR63-8:1, lnc-HPRT1-9:1, lnc-MDC1-1:1 and RSL24D1) that all interacted with hsa-miR-103a-2-5p. The results showed that RNA expression was upregulated in hypertensive patients compared with that in normotensive patients, which was in line with the microarray analysis. Furthermore, NR\_104160, lnc-GPR63-8:1, lnc-HPRT1-9:1, ID1 and

## lncRNA-NR\_104160 and ceRNA network for hypertension

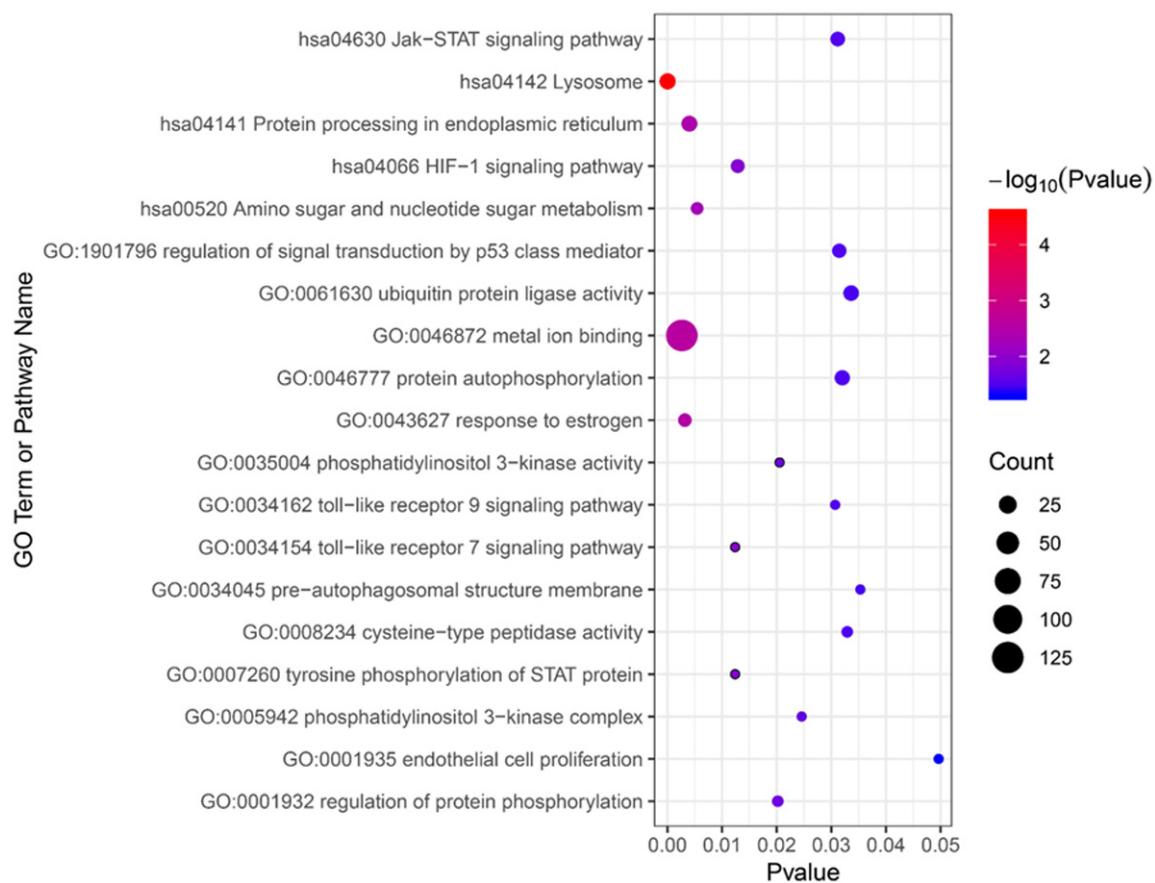


**Figure 1.** Overview of lncRNA and mRNA expression profiles. A. Volcano plot of differentially expressed lncRNAs. The vertical green lines correspond to twofold increased and decreased expression, and the horizontal green line represents  $P = 0.05$ . The red points represent differentially expressed lncRNAs between hypertensive and normotensive individuals. B. Chromosomal distribution of differentially expressed lncRNAs. C. Heatmap of the top 20 differentially expressed lncRNAs. The red and blue colors indicate high and low expression of the 20 dysregulated RNAs among samples, respectively. The values in the heatmap represent the levels of RNAs in each group. H: hypertensive samples; N: normal samples. D. Volcano plot of differentially expressed mRNAs. The vertical green lines correspond to twofold increased and decreased expression, and the horizontal green line represents  $P = 0.05$ . The red points represent differentially expressed mRNAs between hypertensive and normotensive individuals. E. Chromosomal distribution of differentially expressed mRNAs. F. Heatmap of the top 20 differentially expressed mRNAs. The red and blue colors indicate high and low expression, respectively, of the 20 dysregulated RNAs among samples. The values in the heatmap represent the levels of RNAs in each group. H: hypertensive samples; N: normal samples.

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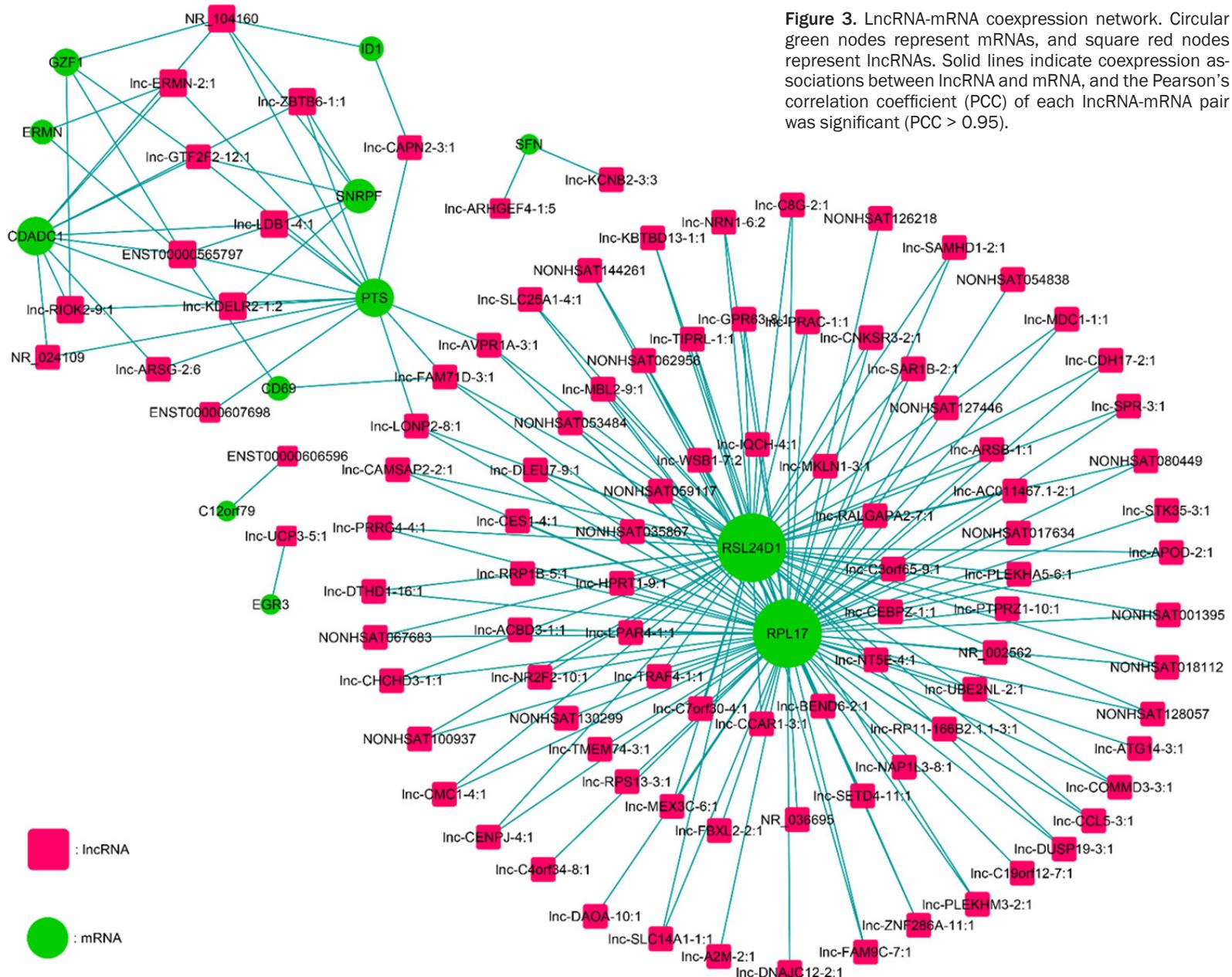
**Table 2.** Top 10 dysregulated lncRNAs and mRNAs

	lncRNA	P-value	Fold change	mRNA	P-value	Fold change
Upregulated	lnc-EIF2AK4-6:1	0.00269	3.66	DMRTC2	0.00130	4.78
	lnc-IL2RA-4:1	0.00300	2.43	SFTPA2	0.00265	2.28
	lnc-CD52-1:1	0.00457	3.55	HIGD1C	0.00325	2.24
	ENST00000614747	0.00556	2.47	ANXA8	0.00402	2.21
	lnc-CNOT10-2:1	0.00558	2.16	RGS1	0.00541	6.49
	ENST00000505646	0.00638	2.16	SLC16A14	0.00816	2.17
	ENST00000564481	0.00718	2.84	NR4A3	0.00875	6.80
	lnc-LOH12CR1-6:1	0.00723	2.20	ERMN	0.00918	2.04
	lnc-POTEG-9:1	0.00751	2.68	DUSP4	0.00982	2.01
	lnc-TEKT3-1:1	0.00779	2.20	C17orf64	0.01282	2.20
Downregulated	ENST00000608286	0.00043	0.45	ZNF347	0.00194	0.50
	NR_033186	0.00184	0.50	SPRED1	0.00350	0.47
	NR_029782	0.00197	0.29	CD180	0.00379	0.44
	ENST00000506059	0.00475	0.41	VIT	0.00501	0.41
	lnc-NPBWR1-4:1	0.00631	0.35	GPR20	0.00661	0.37
	lnc-MALT1-1:2	0.00751	0.48	ANO2	0.00777	0.46
	lnc-CTNNB1-1:1	0.00800	0.44	PCDHGA8	0.01295	0.48
	lnc-ZNF720-3:4	0.00944	0.28	DLC1	0.01589	0.43
	ENST00000607744	0.00957	0.50	TIFAB	0.01628	0.50
	ENST00000604491	0.00987	0.48	ZNF585B	0.02631	0.49



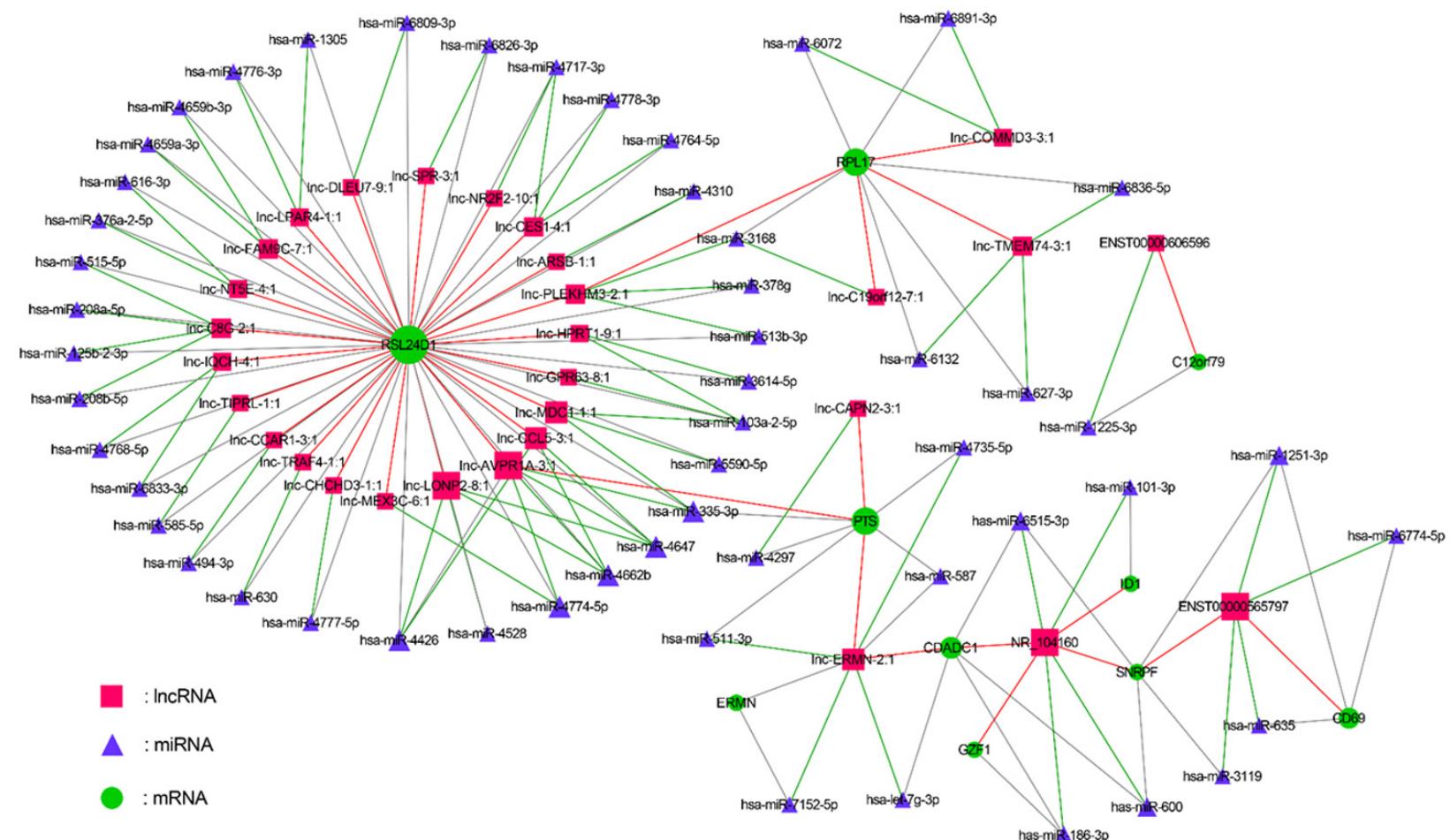
**Figure 2.** The GO Term and KEGG Pathway enrichment analyses of genes targeted by differentially expressed lncRNAs. Count represents the gene number in the GO term or KEGG pathway.

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**Figure 3.** LncRNA-mRNA coexpression network. Circular green nodes represent mRNAs, and square red nodes represent lncRNAs. Solid lines indicate coexpression associations between lncRNA and mRNA, and the Pearson's correlation coefficient (PCC) of each lncRNA-mRNA pair was significant ( $PCC > 0.95$ ).

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**Figure 4.** LncRNA-miRNA-mRNA ceRNA network. In this network, the triangle, square and circular nodes represent miRNA, lncRNAs and mRNAs, respectively. Red edges indicate the coexpression relationship between a lncRNA and mRNA, and the Pearson's correlation coefficient of each lncRNA-mRNA pair was  $> 0.95$ . The green and gray edges represent predicted lncRNA-miRNA and miRNA-mRNA associations, respectively, predicted by bioinformatic software.

*RSL24D1* were significantly upregulated in the hypertension group compared to their expression levels in the control group ( $P = 0.012, 0.024, 0.035, 0.028$  and  $0.039$ , respectively), but the  $P$ -value of *Inc-MDC1-1:1* was  $0.074$ . **Figure 5** illustrates the relative expression levels of five differentially expressed RNAs.

#### The results of multiple logistic regression analysis

The associations between hypertensive status and  $\log_2$ -transformed relative expression levels of RNAs (NR\_104160, Inc-GPR63-8:1, Inc-HPRT1-9:1, RSL24D1, ID1) were assessed. After adjusting for confounding factors, including TG, HDL-C, and sauce and poultry consumption, only NR\_104160 was significantly associated with the risk of hypertension (OR = 2.863, 95% CI: 1.143-7.172;  $P = 0.025$ ). The final model was logit ( $P = \text{hypertension}$ ) =  $1.913 + 1.052 \times \text{NR\_104160} + 1.415 \times \text{TG} - 0.815 \times \text{HDL-C} + 1.234 \times \text{poultry} + 1.917 \times \text{sauce}$  (**Table 3**).

#### The diagnostic performance by ROC analysis

Predicted probabilities based on a multiple logistic regression model were examined by ROC curve analysis. We evaluated the diagnostic value of NR\_104160 and five RNA combinations in human peripheral blood samples for hypertension (**Figure 6**). After adjusting for TG, HDL-C, and sauce and poultry consumption, the AUCs of NR\_104160 and the combination of five RNAs were 0.852 (95% CI: 0.761-0.944;  $P < 0.001$ ) and 0.821 (95% CI: 0.721-0.922;  $P < 0.001$ ), the sensitivity values were 73.8% (95% CI: 58.0%-86.1%) and 78.6% (95% CI: 63.2-89.7%), and the specificity values were 89.3% (95% CI: 71.8-97.7%) and 78.6% (95% CI: 59.1-91.7%), respectively.

#### NR\_104160 is a potential biomarker for hypertension

NR\_104160 was further tested by qRT-PCR on another 56 individuals and showed a significantly higher expression level in hypertensives than normotensives ( $P = 0.001$ ) (**Figure 7A**). The NR\_104160-related diagnostic model was externally verified in these samples, and the predicted probabilities were used to perform ROC curve analysis (**Figure 7B**). The AUC was 0.769 (95% CI: 0.641-0.898;  $P < 0.001$ ), with

sensitivity and specificity values of 68.6% (95% CI: 50.7%-83.2%) and 86.7% (95% CI: 63.7%-97.0%), respectively.

#### NR\_104160 serves as an miRNA sponge for hsa-miR-101-3p and the association between NR\_104160 and ID1

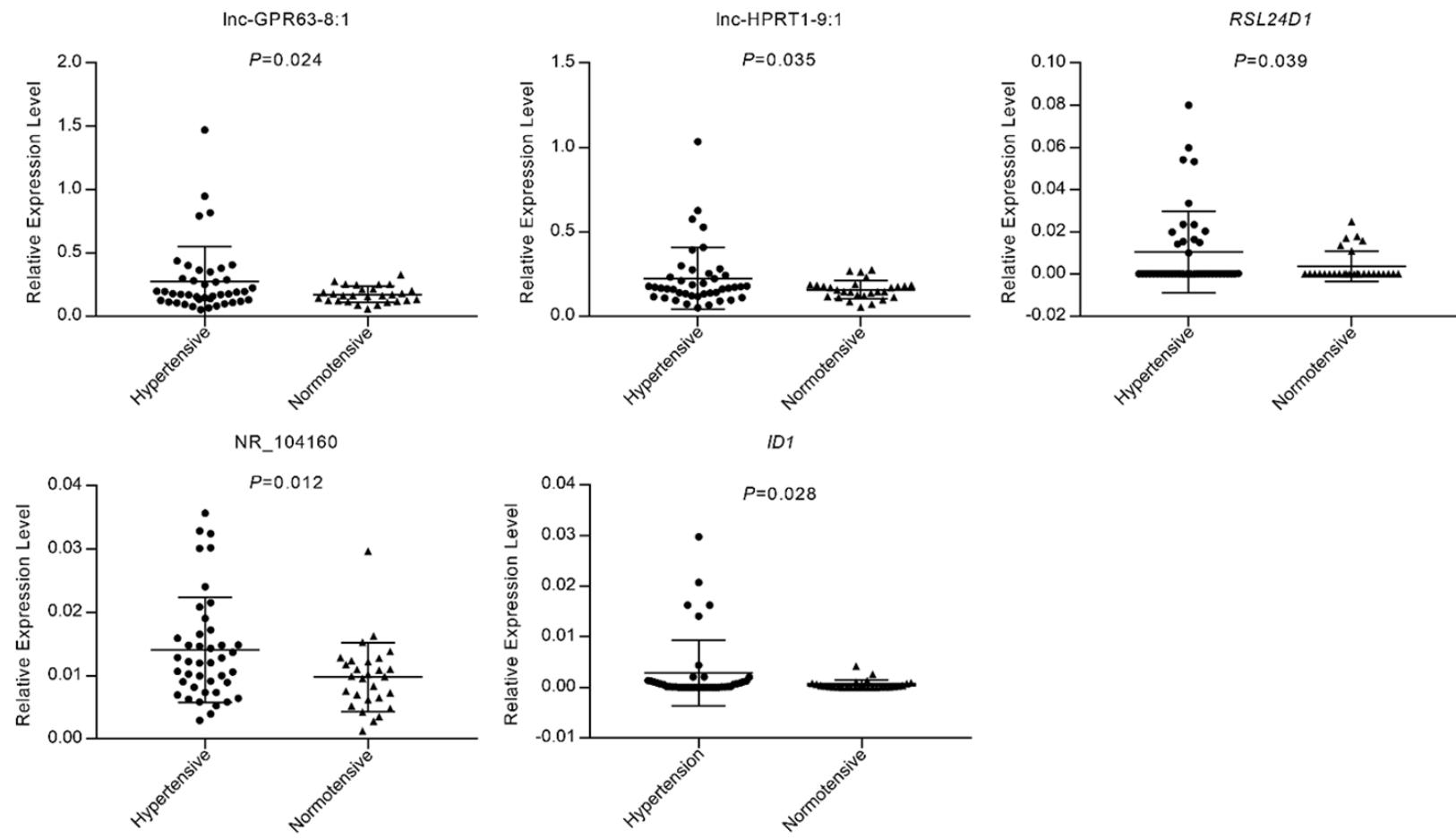
NR\_104160 is located at 10p11.22 (chr10: 32345112-32347218), and its associated gene symbol is *EPC1*. Bioinformatics analysis revealed that hsa-miR-101-3p was targeted by both NR\_104160 in miRDB and *ID1* in TarBase v8. Luciferase assays were applied to determine whether miR-101-3p directly targets NR\_104160 and *ID1*. Potential binding sites of miR-101-3p were identified within the NR\_104160 and *ID1* sequences (**Figure 8A**). The results illustrated that miR-101-3p mimics significantly inhibited the luciferase activity of GV272-NR\_104160-WT ( $P < 0.001$ ) but did not affect the luciferase activity of GV272-NR\_104160-Mut (**Figure 8B**). These data suggest that NR\_104160 may serve as a sponge for miR-101-3p. However, *ID1* was not a binding target of miR-101-3p (**Figure 8C**).

According to the correlation analysis, the PCC of *ID1* and NR\_104160 in the first qRT-PCR test was 0.565. **Table 4** illustrates the results of GLM after adjusting for age, sex, MAP, TC, TG, HDL-C, and LDL-C. The results showed that an elevated expression level of NR\_104160 increased the expression of *ID1* ( $\beta = 0.2235$ ,  $P = 0.005$ ).

#### Discussion

Our results identified 387 DE-lncRNAs and 52 DE-mRNAs by microarray and verified that five RNAs (Inc-GPR63-8:1, Inc-HPRT1-9:1, NR\_104160, *ID1* and *RSL24D1*) were significantly upregulated in hypertensive patients compared with their expression levels in controls. NR\_104160 was significantly associated with the risk of hypertension, and in the hypertension diagnostic model, the AUC was 0.852 (95% CI: 0.761-0.944). Luciferase assays indicated that NR\_104160 serves as a sponge for miR-101-3p. Our study constructed a ceRNA network to provide a hypothesis for the understanding of the mechanism of hypertension. lncRNA-NR\_104160 was identified as a hub element that participates in hypertension tran-

## lncRNA-NR\_104160 and ceRNA network for hypertension



**Figure 5.** Relative expression levels of five RNAs according to qRT-PCR. Independent two-sample t-test was analyzed to compare the expression of RNAs (Inc-GPR63-8:1, Inc-HPRT1-9:1, RSL24D1, NR\_104160 and ID1) between 42 hypertensives and 28 normotensives. The long and short lines represent the mean and standard deviation, respectively, of the RNA relative expression level.

**Table 3.** Analysis of the association between RNAs and hypertension depending on unconditional logistic regression analysis

Variables	$\beta$	P-value	OR	95% CI	
				Lower	Upper
NR_104160	1.052	0.025	2.863	1.143	7.172
TG	1.415	0.018	4.118	1.272	13.326
HDL-C	-0.815	0.009	0.443	0.240	0.815
Poultry	1.234	0.007	3.435	1.407	8.384
Sauce	1.917	0.044	6.799	1.049	44.055
Constant	1.913	0.519	6.775		

TG, triglycerides; HDL-C, high-density-lipoprotein-cholesterol; Poultry, poultry consumption per week; Sauce, sauce consumption per month. Statistical testing by stepwise regression method.  $P < 0.05$  was considered statistically significant.

scriptional regulation and acts as a potential biomarker.

Recently, an increasing number of functional lncRNAs have been reported in hypertension studies [20], such as lncRNA-Sone [21], Inc-Ang362 [22], and CDKN2B-AS1 [23]. However, few studies using microarrays have focused on lncRNAs in the context of human hypertension. Cai J [6] explored the lncRNA expression profile using Human lncRNA Array v2.0 ( $8 \times 60K$ , Arraystar) and demonstrated that lncRNA-AK098656 was strongly upregulated in the plasma of hypertensive patients. In the current study, lncRNA-NR\_104160 was highly expressed in hypertensive individuals compared with its expression in normotensive individuals, and the risk of hypertension increased by 1.863 times. Furthermore, the ROC curve analysis revealed that the AUC of NR\_104160 was equal to 0.852 and demonstrated 73.8% sensitivity and 89.3% specificity. This model was externally verified in 56 individuals in the second step of the validation, with an AUC of 0.769 (95% CI: 0.641-0.898). These results suggested that NR\_104160 was significantly and independently associated with hypertension.

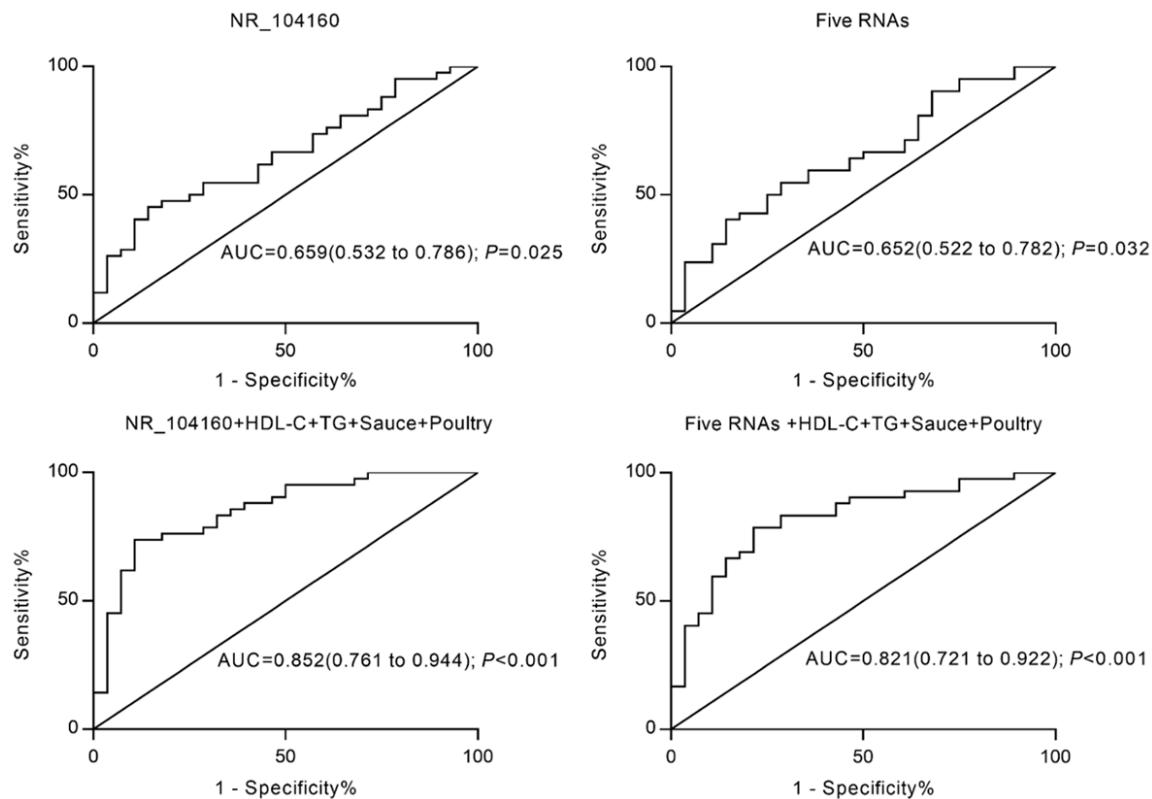
Our results revealed a relatively new biomarker of NR\_104160 that was coexpressed with five DE-mRNAs (*ID1*, *PTS*, *SNRPF*, *GZF1*, *CDADC1*). In general, the high PCC between these RNAs represented the high possibility of similar functions. The gene *PTS* encodes the enzyme that is greatly associated with the biosynthesis of tetrahydrobiopterin (BH4). BH4 is the cofactor

of nitric oxide synthase (eNOS), and many studies have focused on the mechanisms of eNOS dysfunction in hypertension because restoring eNOS function might be a potential novel therapeutic strategy to treat hypertension [24]. Researchers have found that the upregulation of the BH4 pathway might ameliorate the hypertension-related decline in the reendothelialization capacity of endothelial progenitor cells [25]. In addition, studies demonstrated that antihypertensive treatment can restore BH4, e.g., triple therapy with reserpine + hydrochlorothiazide + hydralazine or oral BH4 [26]. These data suggest that *PTS* might be related to hypertension. Few hypertension-related studies have assessed *SNRPF*, *GZF1*, and *CDADC1*. Researchers have found that *SNRPF* expression is downregulated by ubiquitin C-terminal hydrolase-L5, resulting in the inhibition of the migration and invasion of glioma cells [27]. *GZF1* mutations could cause the expansion of genetic heterogeneity in Larsen syndrome [28].

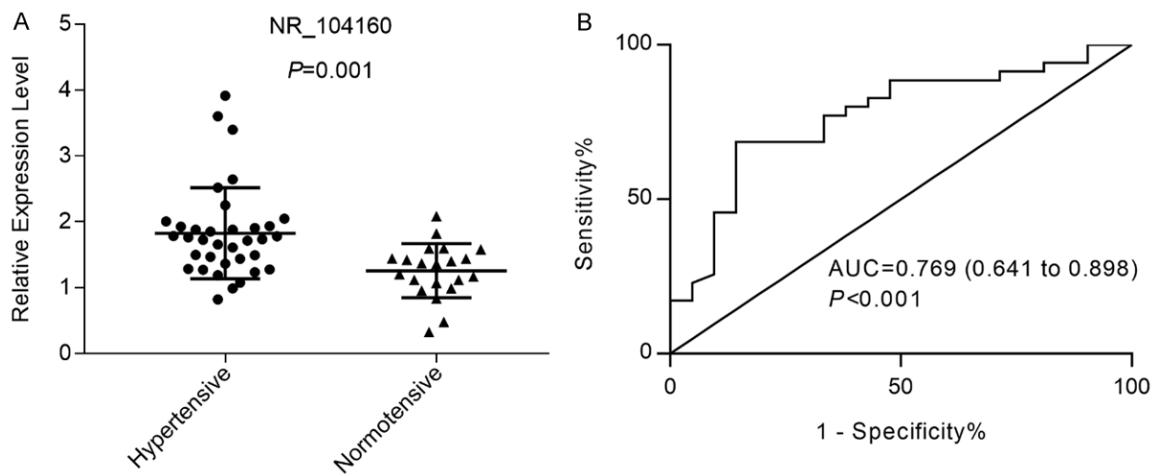
LncRNAs can regulate coding RNAs in the development of diseases, such as pulmonary hypertension [29], and the heart against myocardial I/R injury [30], by acting as miRNA sponges [31, 32]. In the novel regulatory mechanism, miRNAs are viewed as messengers and MREs as the letters of an 'RNA language'; thus, RNA-RNA crosstalk appears. We constructed a ceRNA network for hypertension that included 30 lncRNAs, 10 mRNAs, 54 miRNAs and 71 'lncRNA-miRNA-mRNA' loops. Strikingly, lncRNAs and mRNAs in each ceRNA loop interacted with the same miRNAs and had a coexpression relationship between themselves.

Within the ceRNA network, NR\_104160 was involved in seven ceRNA pathways, and NR\_104160→hsa-miR-101-3p→*ID1* was one of them. The gene symbol of NR\_104160 is *EPC1*, and it might be named lncRNA-EPC1 according to the current default rules. Thus, we referred to the literature on *EPC1*, hsa-miR-101-3p and *ID1*. *EPC1* was reported to be associated with many biological functions and diseases, such as endometrial stromal sarcoma [33] and aberrant spermatid development [34]. A previous study revealed that *ID1* may be associated with familial pulmonary arterial hypertension [35] and is relevant to the proliferation of pulmonary artery smooth muscle

## lncRNA-NR\_104160 and ceRNA network for hypertension



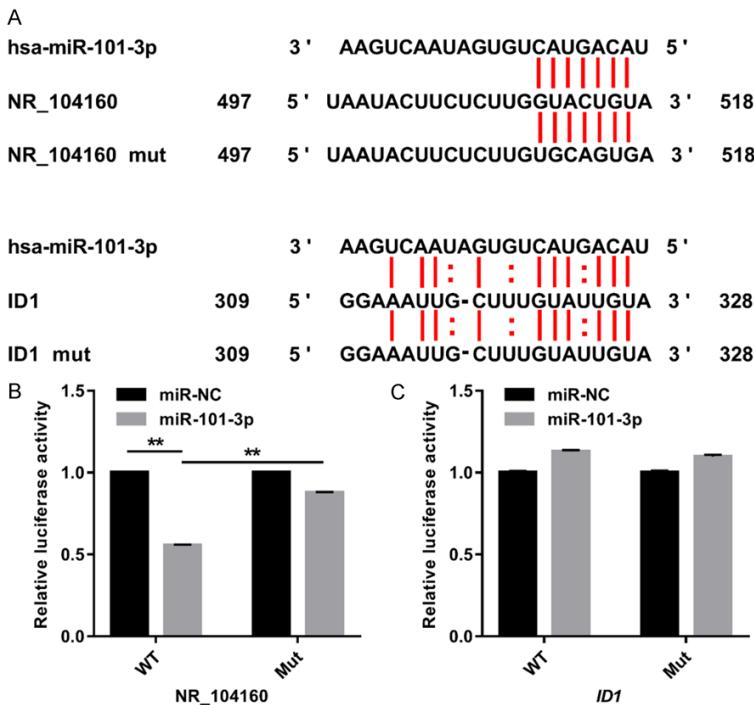
**Figure 6.** ROC analysis of NR\_104160 and the combination model of five RNAs. AUC, area under the curve; HDL-C, high-density lipoprotein cholesterol; TG, triglycerides; Sauce, sauce consumption per month; Poultry, poultry consumption per week; Five RNAs, Inc-GPR63-8:1 + Inc-HPRT1-9:1 + NR\_104160 + ID1 + RSL24D1.



**Figure 7.** The second qRT-PCR validation of NR\_104160. A. Relative expression levels of NR\_104160. Independent two-sample t-tests were performed to compare the expression levels between 35 hypertensive patients and 21 normotensive patients. The long and short lines represent the mean and standard deviation, respectively. B. ROC analysis of the NR\_104160-related model. AUC, area under the curve.

cells [36], endothelial proliferation and angiogenesis [37]. Hsa-miR-101-3p is involved in ceRNA pathways in various diseases, such as

the PTAR/miR-101-3p/ZEB1 pathway in ovarian cancer [38], the SNHG12/miR-101-3p/FOXP1 axis in glioma [39], and the LINC00052/miR-



**Figure 8.** The potential relationships between lncRNA-NR\_104160, hsa-miR-101-3p and ID1. A. Hsa-miR-101-3p binding sequences in lncRNA-NR\_104160 or ID1 and mutant sites in lncRNA-NR\_104160 or ID1. B. Luciferase assay showed that the luciferase activity of lncRNA-NR\_104160-WT (wild type) was inhibited by hsa-miR-101-3p mimics but that the luciferase activity of lncRNA-NR\_104160-mut (mutant) was not. \*\*,  $P < 0.001$ . C. Luciferase assay showed that the luciferase activities of ID1-WT and ID1-mut were not inhibited by hsa-miR-101-3p mimics.

**Table 4.** Analysis of the influencing factors of mRNA expression depending on Generalized linear model

Dependent	Variables	$\beta$	95% CI		P-value
			Lower	Upper	
<i>ID1</i>	NR_104160	0.2235	0.0663	0.3806	0.005
	Age	0.0001	-0.0002	0.0003	0.602
	Gender	0.0007	-0.0012	0.0027	0.456
	MAP	< 0.0001	-0.0001	0.0001	0.376
	TC	-0.0014	-0.0029	0.0001	0.072
	TG	0.0014	0.0004	0.0025	0.009
	HDL-C	0.0016	< -0.0001	0.0033	0.056
	LDL-C	0.0010	-0.0007	0.0027	0.237
	Constant	-0.0109	-0.0288	0.0070	0.224

TC, total cholesterol; TG, triglycerides; HDL-C, high-density-lipoprotein-cholesterol; LDL-C, low-density-lipoprotein-cholesterol; MAP, mean arterial pressure.

101-3p/SOX9 axis in hepatocellular carcinoma [40]. Hence, we hypothesized that NR\_104160 might regulate *ID1* by binding to hsa-miR-101-3p.

Therefore, luciferase reporter assays were applied to determine whether miR-101-3p directly targets NR\_104160 and *ID1*. The results illustrated that NR\_104160 may serve as a sponge for miR-101-3p. Although *ID1* was not a binding target of miR-101-3p, our GLM results revealed that the expression level of *ID1* was associated with NR\_104160 ( $\beta = 0.2235$ ,  $P = 0.005$ ), which suggested that the relationship between NR\_104160 and *ID1* may be very complicated.

Some strengths and limitations of the current study should be acknowledged. First, we performed two-step qRT-PCR validations for NR\_104160, and the results showed consistent significantly higher expression in hypertensive individuals than in controls. Second, NR\_104160 was found to be significantly and independently associated with the risk of hypertension, and the robustness of the diagnostic model was verified externally. Third, the construction of a ceRNA network might provide a potential basis for revealing the pathogenesis of hypertension, and we found that NR\_104160 could serve as a sponge for miR-101-3p. Fourth, we used bioinformatics technology, which can aid in the understanding of the genes and networks that are responsible for diseases. The limitations are as follows: the number of samples limited the statistical power of the microarray analyses. This stu-

dy only considered one lncRNA regulatory mechanism, but lncRNA-related regulation is very complicated. The result of the luciferase reporter assay suggested that *ID1* was not a

binding target of miR-101-3p, and the relationship between NR\_104160 and *ID1* may be very complicated and require further exploration. However, this study aimed to identify lncRNAs as potential biomarkers and construct a ceRNA network for hypertension.

In conclusion, our study verified that NR\_104160 was significantly upregulated in the human peripheral blood of hypertensive patients compared to its expression in normotensive patients. Moreover, NR\_104160 was identified as a potential biomarker for hypertension. Our results suggest that systematically analyzing the ‘lncRNA-miRNA-mRNA’ ceRNA network could help to elucidate the pathogenesis of hypertension and the RNA-RNA crosstalk among lncRNAs, miRNAs and mRNAs.

### Acknowledgements

This study was supported by the National Key Research and Development Program of China (No. 2016YFC0900600/2016YFC0900603), the Natural Science Foundation of China (No. 81973121 & 81373076), and the Beijing Natural Science Foundation (No. 7172023).

### Disclosure of conflict of interest

None.

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**Table S1.** The primer sequences for qRT-PCR

NO.	Gene	Primer sequence
1	lnc-TIPRL-1:1	F: AGGCAGACTTGAGGGGTTCTC R: CCACCATATCCCTGCTGAC
	lnc-TIPRL-1:1	
2	lnc-MDC1-1:1	F: GAAGGAATTCCCAGAGCTGAA R: TTGCCTTTGAATCATCTTTGG
	lnc-MDC1-1:1	
3	lnc-GPR63-8:1	F: GTTGGTAGGGTCGGAAAGC R: GACAACATGCTCCCCACTGA
	lnc-GPR63-8:1	
4	lnc-HPRT1-9:1	F: GCGATCTTGAGGACTGGTC R: TGTTCTGCTGTGCTTGTC
	lnc-HPRT1-9:1	
5	NR_104160	F: AAACAGGGTGCAGTGTATTGACA R: CCACAACAAATTGAGCACACAA
	NR_104160	
6	RSL24D1	F: CTTTCTAGCCCAGCATTGA R: TTAAAAGGCCGCTGGGATTCTC
	RSL24D1	
7	ID1	F: TTTCTGAGGAAATTGCTTGTATTGT R: CAGCCCCACAGAACTATTGTAAAA
	ID1	
8	GAPDH	F: TGACTTCAACAGCGACACCCA R: CACCCTGTTGCTGTAGCCAAA
	GAPDH	

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**Table S2.** Clinical characteristics of study participants in the hypertensive and normotensive groups of qRT-PCR

Variables	First quantitative real-time PCR				Second quantitative real-time PCR			
	Hypertensive (N = 42)	Normotensive (N = 28)	Total (N = 70)	P-value	Hypertensive (N = 35)	Normotensive (N = 21)	Total (N = 56)	P-value
Gender (female, n (%))	28 (66.67)	18 (64.29)	46 (65.71)	0.837 <sup>‡</sup>	25 (71.43)	10 (47.62)	35 (62.50)	0.075 <sup>‡</sup>
Age (years)	63.67 ± 4.14	63.18 ± 4.44	63.47 ± 4.24	0.640 <sup>*</sup>	63.06 ± 6.51	62.84 ± 6.23	63.11 ± 5.98	0.739 <sup>*</sup>
SBP (mmHg)	151.19 ± 16.83	127.5 ± 10.62	141.71 ± 18.68	< 0.001 <sup>*</sup>	146.09 ± 17.88	129.38 ± 12.16	139.82 ± 17.83	< 0.001 <sup>*</sup>
DBP (mmHg)	79.60 ± 8.64	74.21 ± 8.75	77.44 ± 9.02	0.013 <sup>*</sup>	78.17 ± 14.58	76.07 ± 8.66	77.38 ± 12.64	0.552 <sup>*</sup>
BMI (kg/m <sup>2</sup> )	25.49 ± 3.24	23.92 ± 3.37	24.87 ± 3.36	0.054 <sup>*</sup>	26.03 ± 3.54	24.15 ± 3.43	25.32 ± 3.59	0.057 <sup>*</sup>
TC (mmol/L)	4.64 ± 1.53	4.93 ± 0.97	4.76 ± 1.33	0.476 <sup>†</sup>	3.37 ± 1.80	4.42 ± 1.31	3.76 ± 1.70	0.049 <sup>†</sup>
TG (mmol/L)	1.85 ± 1.25	1.38 ± 0.57	1.66 ± 1.05	0.275 <sup>†</sup>	3.44 ± 1.93	1.97 ± 1.68	2.90 ± 1.96	0.011 <sup>†</sup>
HDL-C (mmol/L)	2.14 ± 1.09	2.91 ± 0.81	2.45 ± 1.05	0.001 <sup>†</sup>	2.09 ± 1.00	2.75 ± 0.85	2.34 ± 0.99	0.012 <sup>†</sup>
LDL-C (mmol/L)	2.23 ± 1.18	1.61 ± 0.73	1.98 ± 1.06	< 0.001 <sup>†</sup>	1.78 ± 1.07	1.62 ± 1.08	1.72 ± 1.07	0.741 <sup>†</sup>
FBG (mmol/L)	5.78 ± 1.15	5.63 ± 0.88	5.72 ± 1.04	0.806 <sup>†</sup>	5.97 ± 1.54	5.50 ± 0.89	5.80 ± 1.34	0.397 <sup>†</sup>
Heart rate	76.29 ± 10.09	72.96 ± 9.83	74.96 ± 10.05	0.177 <sup>*</sup>	73.54 ± 10.94	72.38 ± 10.93	73.11 ± 10.85	0.702 <sup>*</sup>
Smoke (yes, n (%))	10 (23.81)	6 (21.43)	16 (22.86)	0.816 <sup>‡</sup>	4 (11.43)	6 (28.57)	10 (17.86)	0.105 <sup>‡</sup>
Family history of hypertension (yes, n (%))	30 (71.43)	15 (53.57)	45 (64.29)	0.127 <sup>‡</sup>	26 (74.29)	11 (52.38)	37 (66.07)	0.094 <sup>‡</sup>
Salt consumption per month				0.762 <sup>‡</sup>				0.213 <sup>‡</sup>
≤ 500 gram	15 (35.71)	11 (39.29)	26 (37.14)		14 (40.00)	12 (57.14)	26 (46.43)	
> 500 gram	27 (64.29)	17 (60.71)	44 (62.86)		21 (60.00)	9 (42.86)	30 (53.57)	
Pickle consumption per month				0.917 <sup>‡</sup>				0.068 <sup>‡</sup>
≤ 250 gram	28 (66.67)	19 (67.86)	47 (67.14)		20 (57.14)	17 (80.95)	37 (66.07)	
> 250 gram	14 (33.33)	9 (32.14)	23 (32.86)		15 (42.86)	4 (19.05)	19 (33.93)	
Sauce consumption per month				0.048 <sup>‡</sup>				0.215 <sup>‡</sup>
≤ 150 gram	27 (64.28)	24 (85.71)	51 (72.86)		21 (60.00)	16 (76.19)	37 (66.07)	
> 150 gram	15 (35.71)	4 (14.29)	19 (27.14)		14 (40.00)	5 (23.81)	19 (33.93)	
Poultry consumption per week				0.202 <sup>‡</sup>				0.183 <sup>‡</sup>
≤ 3 times	16 (38.10)	15 (53.57)	31 (44.29)		23 (65.71)	10 (57.62)	34 (60.71)	
> 3 times	26 (61.90)	13 (46.43)	39 (55.71)		12 (34.29)	11 (52.38)	22 (39.21)	

BMI, body mass index; TC, total cholesterol; TG, triglycerides; HDL-C, high-density-lipoprotein-cholesterol; LDL-C, low-density-lipoprotein-cholesterol; FBG, fasting blood glucose; SBP, systolic blood pressure; DBP, diastolic blood pressure. P < 0.05 was considered statistically significant. \*, Statistical testing by independent-samples t test. †, Statistical testing by Wilcoxon rank sum test. ‡, Statistical testing by  $\chi^2$  test.

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**Table S3-1.** All DE-lncRNAs between hypertensive and normotensive groups

DE-lncRNA	P-value	Fold Changes	DE-lncRNA	P-value	Fold Changes
ENST00000608286	0.000431	0.451954	lnc-SPR-3:1	0.029937	2.149257
NR_033186	0.001842	0.499125	lnc-CHCHD3-1:1	0.029962	2.241278
NR_029782	0.001968	0.286406	lnc-IRAK1BP1-6:1	0.029968	3.191759
lnc-EIF2AK4-6:1	0.002691	3.656553	lnc-UNC5C-2:1	0.029976	2.492030
lnc-IL2RA-4:1	0.003001	2.427779	lnc-RCSD1-2:1	0.029980	2.461092
lnc-CD52-1:1	0.004570	3.553413	lnc-WDR36-1:1	0.029981	0.490155
ENST00000506059	0.004747	0.406730	lnc-BMP2K-4:1	0.030193	2.161710
ENST00000614747	0.005561	2.468473	lnc-NR4A2-8:1	0.030258	2.232988
lnc-CNOT10-2:1	0.005582	2.164794	lnc-KDELR2-1:2	0.030267	2.570226
lnc-NPBWR1-4:1	0.006309	0.347671	lnc-UBE2NL-2:1	0.030268	3.054562
ENST00000505646	0.006378	2.161718	lnc-PLIN1-1:1	0.030318	4.660597
ENST00000564481	0.007182	2.844158	lnc-MEX3C-6:1	0.030330	2.263003
lnc-LOH12CR1-6:1	0.007230	2.199606	lnc-C4orf34-8:1	0.030545	3.133566
lnc-POTEG-9:1	0.007507	2.683584	lnc-TRAPPC8-4:1	0.030573	3.099935
lnc-MALT1-1:2	0.007510	0.476829	lnc-CNKSRS3-2:1	0.030724	3.241606
lnc-TEKT3-1:1	0.007793	2.199088	lnc-PSEN2-3:1	0.030799	3.969550
NR_024109	0.007942	2.033839	lnc-NDUFS8-6:1	0.030854	3.317164
lnc-CTNNB1-1:1	0.008001	0.441364	lnc-TMEM185B-7:1	0.030934	3.235290
lnc-FKBP1C-2:1	0.008001	3.720573	NONHSAT083013	0.031080	2.519326
NR_027064	0.008194	2.413124	lnc-DPYSL2-1:2	0.031187	0.369112
lnc-SLITRK2-2:1	0.008466	2.135440	lnc-TRAF4-1:1	0.031297	2.922623
lnc-ACTR8-2:2	0.008950	2.010220	lnc-HSP90AA1-12:1	0.031488	2.965284
lnc-GPR125-5:1	0.009259	2.249563	lnc-DPY19L4-2:1	0.031507	3.277730
lnc-GLI2-8:1	0.009269	4.732693	lnc-INSM2-3:1	0.031540	4.369118
lnc-ZNF720-3:4	0.009444	0.284125	lnc-GJA1-5:1	0.031616	4.414721
ENST00000607744	0.009565	0.498854	lnc-C21orf91-7:1	0.031910	2.057613
ENST00000604491	0.009868	0.483914	lnc-HNRPDL-1:1	0.032011	2.246524
lnc-XRCC2-11:1	0.010028	4.751639	lnc-SLC36A4-4:1	0.032026	5.435425
lnc-KCNB2-3:3	0.010433	3.170159	lnc-AF165138.7.1-6:1	0.032115	3.129283
lnc-ARHGEF4-1:5	0.010667	2.284153	lnc-UCP3-5:1	0.032162	2.459402
lnc-WDR60-10:1	0.010714	4.353585	ENST00000418620	0.032233	2.865087
lnc-ITPR1-1:1	0.010979	2.810121	lnc-MKLN1-3:1	0.032452	3.154286
lnc-MTFR1-4:1	0.011047	6.836786	lnc-SEMA6A-9:1	0.032508	2.967021
ENST00000562459	0.011098	0.413284	lnc-RRP1B-5:1	0.032558	2.988848
lnc-ABCC11-2:1	0.011194	2.555013	lnc-FAM133A-1:1	0.032572	2.822855
lnc-PSMB4-2:1	0.011320	2.635221	lnc-RUNX1T1-8:1	0.032599	5.508733
lnc-NCDN-4:1	0.011472	3.118238	lnc-SAMHD1-2:1	0.032674	2.944065
lnc-FAM150A-4:1	0.011629	3.014977	lnc-HAS2-4:1	0.033028	3.272611
lnc-TMEM14C-3:1	0.011791	0.363498	lnc-TAAR8-2:1	0.033103	3.198109
lnc-HELT-4:1	0.012120	3.399447	NONHSAT035867	0.033243	2.275326
lnc-EREG-1:1	0.012501	5.222298	lnc-SLAIN1-10:1	0.033278	2.633741
lnc-RNF152-10:1	0.012682	2.833123	lnc-KRAS-3:1	0.033320	2.120663
NONHSAT055372	0.012769	2.364184	lnc-RP11-1396013.13.1-3:1	0.033426	3.399453
lnc-PSMA8-3:1	0.013580	3.586623	NONHSAT080449	0.033499	2.284116
lnc-RNF146-3:2	0.013682	0.450543	lnc-CORO6-2:1	0.033600	2.033234
ENST00000588182	0.013795	2.483066	NR_105043	0.033635	0.497849
lnc-ARL14-1:1	0.013947	2.152969	lnc-FGF9-1:2	0.033740	2.012303

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NONHSAT082239	0.014164	2.734858	lnc-GTF2F2-12:1	0.033829	3.110996
lnc-C19orf40-2:1	0.014524	3.582016	lnc-ZNF485-5:1	0.034382	2.888447
lnc-GSG2-2:1	0.014790	2.298855	lnc-SLC25A1-4:1	0.034402	3.591687
lnc-IFNGR1-1:1	0.014800	2.017367	NONHSAT067683	0.034453	2.106330
lnc-KIAA0754-4:1	0.014866	0.303337	lnc-TCEANC-3:1	0.034475	3.222334
ENST00000580686	0.015090	2.006726	lnc-HEATR2-2:3	0.034536	2.122075
lnc-NHSL2-5:1	0.015458	5.364915	NONHSAT069830	0.034658	2.336815
lnc-CNTNAP1-2:1	0.015672	2.185384	ENST00000623366	0.034730	2.247730
NR_028484	0.015842	2.466656	NONHSAT055039	0.034745	2.569516
lnc-NIPBL-5:1	0.015956	2.068993	lnc-XRCC6BP1-7:1	0.034967	2.403190
lnc-GSTM4-3:1	0.016001	4.558921	lnc-DLEU7-9:1	0.035358	3.739677
lnc-CSPG4-4:1	0.016239	2.832767	lnc-NXPH1-3:1	0.035573	2.622335
ENST00000562878	0.016451	3.780205	ENST00000469216	0.035583	2.991286
lnc-C9orf93-4:1	0.016462	4.292720	lnc-SAR1B-2:1	0.035584	2.945021
lnc-COA5-1:2	0.016706	2.281758	ENST00000607698	0.035631	2.236715
NONHSAT037842	0.016826	2.494149	lnc-FAM9C-7:1	0.035635	2.208227
NR_003319	0.017015	2.055342	NR_002564	0.035655	2.022936
lnc-SLAIN1-6:1	0.017050	2.347503	lnc-LPHN3-9:1	0.035655	2.887387
NONHSAT014088	0.017060	3.315290	lnc-API5-11:1	0.035713	5.360648
lnc-NR2C1-2:5	0.017601	3.187805	lnc-CNTROB-1:1	0.035773	3.034125
NR_003326	0.017605	2.237539	lnc-CREB3L4-5:1	0.035829	2.486738
ENST00000618697	0.018278	2.811176	lnc-PHYHIP-3:1	0.036129	2.411950
lnc-AC011467.1-2:1	0.018886	3.873938	lnc-CCL5-3:1	0.036142	2.169636
lnc-AC024940.1-3:1	0.018950	5.721876	lnc-PKP4-4:1	0.036273	3.574996
ENST00000520185	0.019191	3.153546	lnc-ERN1-2:1	0.036506	2.060532
lnc-CAPN2-3:1	0.019274	3.059190	NONHSAT007752	0.036575	2.068952
NONHSAT023699	0.019354	3.654491	lnc-PLEKHA5-6:1	0.036903	3.173178
ENST00000457602	0.019451	2.808044	lnc-APOD-2:1	0.036948	3.037368
lnc-ATP7A-1:1	0.019482	2.117638	ENST00000513581	0.037019	2.052258
NONHSAT054838	0.019612	2.348478	lnc-RP11-144F15.1.1-1:1	0.037149	0.416976
ENST00000607849	0.020036	2.749882	lnc-WNT7B-2:1	0.037222	2.334190
lnc-RP4-811H24.6.1-3:1	0.020089	2.203762	NONHSAT128057	0.037247	2.046945
lnc-PRDM10-2:1	0.020139	6.452253	lnc-TIPRL1-1:1	0.037326	2.473408
ENST00000522057	0.020262	0.447348	lnc-TLK1-3:1	0.037420	0.405312
ENST00000440714	0.020451	2.693063	NONHSAT144261	0.037427	2.086151
lnc-C8G-2:1	0.020470	2.017927	lnc-SLC25A17-2:1	0.037491	2.112698
lnc-XRCC2-5:1	0.020536	2.252912	ENST00000516179	0.037776	3.698214
lnc-C20orf196-5:1	0.020537	2.206133	NONHSAT097497	0.037930	3.233874
lnc-ERMN-2:1	0.020549	2.316518	lnc-GCA-6:1	0.038066	0.322882
lnc-MMADHC-5:1	0.020674	2.714973	lnc-CCAR1-3:1	0.038094	2.796137
ENST00000556386	0.020962	0.492902	NONHSAT101519	0.038116	2.296229
lnc-C11orf63-3:1	0.021055	5.800559	lnc-CPB2-1:1	0.038135	2.716338
lnc-GFM1-3:1	0.021398	2.138727	lnc-ZBTB6-1:1	0.038167	2.522111
NONHSAT018112	0.021474	2.380851	lnc-CEBPZ-1:1	0.038312	2.498023
ENST00000620896	0.021615	4.355321	NONHSAT036908	0.038429	2.323269
lnc-EIF2B1-5:1	0.021669	2.252508	lnc-SCAF11-3:1	0.038451	2.155351
lnc-DAOA-10:1	0.021737	2.116635	lnc-GPR63-8:1	0.038508	2.298733
lnc-CSGALNACT1-4:1	0.021804	2.754306	lnc-A2M-2:1	0.038565	2.059797
lnc-EGR1-1:1	0.021885	2.784704	lnc-PTPRZ1-10:1	0.038600	3.074862

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lnc-LDB1-4:1	0.022076	2.266362	lnc-C2orf57-3:1	0.038703	3.093121
lnc-RIOK2-9:1	0.022275	2.295135	lnc-DPPA4-3:1	0.038898	2.208222
ENST00000486671	0.022392	0.488640	NONHSAT062956	0.038927	2.120931
lnc-NDUFB4-3:1	0.022459	5.001966	lnc-NPIPL2-5:1	0.039054	2.182963
lnc-TNFRSF11A-1:1	0.022544	4.141308	lnc-PRAC-1:1	0.039085	2.238885
NONHSAT117148	0.022834	0.029067	lnc-ZMYND8-6:1	0.039155	3.492289
NONHSAT126218	0.022998	2.281553	lnc-CMC1-4:1	0.039284	3.469623
lnc-TMSB15A-4:1	0.023142	2.936737	lnc-MOK-6:1	0.039415	2.039396
NONHSAT127886	0.023160	4.047682	lnc-CDH17-2:1	0.039494	2.614143
lnc-ZNF366-13:1	0.023319	2.269892	ENST00000516415	0.039614	3.495111
lnc-GP5-4:1	0.023343	2.568824	lnc-TMEM202-4:1	0.039682	2.298466
lnc-AC023632.1-3:1	0.023438	5.825887	lnc-SLC25A26-6:1	0.039883	3.238884
lnc-ZNF292-1:1	0.023504	3.040714	ENST00000619883	0.040129	2.017658
lnc-RALGAPA2-7:1	0.023527	2.401887	lnc-SNX14-5:1	0.040182	3.950922
lnc-C7orf30-4:1	0.023586	3.099895	ENST00000609669	0.040244	2.298269
lnc-ADD3-5:1	0.023669	3.831370	lnc-NXPH2-11:1	0.040282	2.443955
lnc-TAS1R1-1:1	0.023671	4.650636	lnc-PLEKHM3-2:1	0.040292	2.240296
lnc-SEL1L-12:1	0.023680	4.131716	lnc-FBXO10-1:1	0.040456	2.259429
lnc-STX7-1:1	0.023691	2.371396	lnc-NT5E-4:1	0.040664	2.560400
lnc-SMC4-2:1	0.023745	2.502074	lnc-PON3-1:1	0.040670	0.307763
lnc-LPAR1-2:1	0.023811	2.397395	lnc-ACAT1-2:1	0.040989	3.059496
ENST00000432229	0.023891	4.875449	lnc-GPR19-2:1	0.041076	2.262905
lnc-CD40LG-4:1	0.023948	2.138773	lnc-IL20RA-2:1	0.041154	3.912937
lnc-FANCB-2:1	0.023994	2.493826	ENST00000565797	0.041229	2.161403
lnc-CAMP-3:1	0.024049	2.761479	lnc-FAM71D-3:1	0.041316	2.273365
lnc-VPS13C-3:1	0.024080	2.315048	NR_046400	0.041377	3.314155
lnc-DLD-2:1	0.024095	3.265361	lnc-ETV6-5:1	0.041393	2.025817
NONHSAT001395	0.024147	3.443800	lnc-NOS2-6:1	0.041898	2.534273
lnc-PSMA6-3:1	0.024153	4.842954	lnc-SLC14A1-1:1	0.041905	2.130369
lnc-CDH13-4:1	0.024223	2.417426	NONHSAT100937	0.042076	2.366031
lnc-DENND5B-2:1	0.024256	5.941084	lnc-RPS27-2:2	0.042202	2.938509
lnc-ATG14-3:1	0.024596	4.650949	NONHSAT053484	0.042323	2.018450
lnc-SFT2D2-3:1	0.024654	3.993808	lnc-RPL7A-2:1	0.042364	2.197304
NR_104160	0.025050	2.397668	lnc-CENPJ-4:1	0.042422	3.089729
lnc-TAOK3-11:1	0.025193	3.647123	lnc-ZKSCAN2-4:1	0.042664	2.042803
lnc-IFT74-6:1	0.025394	2.916980	lnc-FAM126A-1:1	0.042732	5.815825
lnc-CC2D2A-1:1	0.025426	0.234307	lnc-TMEM183A-2:1	0.042736	2.777235
lnc-VANGL1-3:1	0.025667	2.019714	lnc-MDC1-1:1	0.042821	2.315465
NONHSAT004372	0.025673	2.575596	ENST00000515939	0.043099	2.617044
lnc-RGS13-2:2	0.025673	3.580933	lnc-RPL36A-1:1	0.043108	3.133511
lnc-AC012123.1.1-4:1	0.025757	4.067355	lnc-MBL2-9:1	0.043200	2.849667
lnc-SIRT5-4:2	0.025865	2.481641	lnc-CD82-3:1	0.043634	2.187287
ENST00000563775	0.025868	0.455186	lnc-FBXL2-2:1	0.043716	2.139475
lnc-RP11-439E19.8.1-1:1	0.025948	2.443394	lnc-PRDX6-4:1	0.043902	4.837367
lnc-KATNAL2-10:1	0.025968	2.710647	lnc-NRN1-6:2	0.043947	3.215138
lnc-NPY5R-5:1	0.026096	3.325271	NR_029663	0.043988	2.225406
lnc-BORA-14:1	0.026121	3.183868	lnc-ALS2CR8-2:1	0.044054	2.014730
lnc-GALNT12-2:1	0.026131	3.003232	ENST00000517226	0.044094	2.500426
lnc-C10orf68-16:1	0.026166	3.963789	lnc-DTHD1-16:1	0.044149	2.875734

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NONHSAT130299	0.026220	2.243392	lnc-WSB1-7:2	0.044329	3.056061
lnc-TPRG1-1:1	0.026246	2.128522	NR_028287	0.044415	2.220527
lnc-SLC36A4-7:1	0.026323	4.331721	lnc-RP11-32K4.2.1-3:1	0.044501	3.962538
lnc-C7orf41-2:1	0.026356	3.284729	ENST00000539313	0.044646	2.238493
lnc-FAM82A1-8:1	0.026435	3.770875	lnc-ADAMTS7-2:1	0.044664	2.180753
NONHSAT127446	0.026492	3.308516	lnc-DUSP19-3:1	0.044704	2.342081
lnc-OR6J1-1:1	0.026586	4.849661	lnc-PTPN13-1:1	0.044755	3.610328
lnc-ACTL6A-1:1	0.026680	2.917185	ENST00000608498	0.044916	0.419118
NR_125398	0.026682	2.149512	lnc-TFDP2-9:1	0.044919	3.345986
NONHSAT113353	0.026731	5.714059	lnc-LETMD1-1:1	0.045215	2.509107
lnc-CNOT7-5:1	0.026778	5.887736	lnc-C3orf65-9:1	0.045545	3.148205
NONHSAT078561	0.026787	7.189788	lnc-PLBD1-1:3	0.045730	2.433243
lnc-GRIA3-2:1	0.026812	2.423567	lnc-ARSG-2:6	0.045782	2.319127
lnc-RNF222-1:1	0.027005	2.224587	lnc-CES1-4:1	0.045810	2.797083
lnc-LTA4H-2:1	0.027083	2.262939	lnc-NR2F2-10:1	0.045881	2.529856
lnc-LONP2-8:1	0.027114	2.016856	ENST00000412944	0.045981	0.323891
NR_003327	0.027121	2.951477	lnc-C3orf72-4:2	0.046103	2.685749
ENST00000517272	0.027323	3.226000	lnc-RP11-166B2.1.1-3:1	0.046171	2.168401
ENST00000603472	0.027359	3.022812	lnc-PRRG4-4:1	0.046265	3.035505
ENST00000413347	0.027542	0.211350	lnc-RP11-24B21.1.1-2:7	0.046327	2.100781
lnc-DYRK1A-1:1	0.027543	0.169983	lnc-KBTBD13-1:1	0.046483	2.141858
lnc-DTX1-2:1	0.027592	3.653670	lnc-ARSB-1:1	0.046819	2.560501
lnc-AC016586.1-1:4	0.027715	2.007466	lnc-SUPT6H-2:7	0.046823	2.069449
lnc-ACBD3-1:1	0.027924	4.281501	lnc-C19orf12-7:1	0.046990	2.778798
lnc-GTF2A1-4:1	0.027953	3.111798	lnc-ACBD6-2:1	0.047054	2.514431
ENST00000516592	0.027980	4.373258	lnc-COMMD3-3:1	0.047063	2.503576
lnc-STK35-3:1	0.028143	2.317525	NR_002562	0.047112	2.053924
lnc-RP11-849H4.2.1-2:1	0.028536	4.199084	lnc-NDFIP2-21:1	0.047219	2.111534
lnc-TUFM-4:1	0.028538	2.813072	lnc-MUC20-7:2	0.047226	2.128360
lnc-RBMY1J-7:1	0.028578	2.727599	lnc-CXCR3-12:1	0.047232	4.464054
lnc-CYP7B1-9:1	0.028702	2.930639	lnc-CYTL1-5:2	0.047503	2.901246
lnc-SMARCAD1-1:1	0.028779	3.654037	lnc-RNF144B-5:1	0.047593	2.016126
lnc-TMEM74-3:1	0.028832	2.465546	lnc-IQCH-4:1	0.048068	2.173871
NR_036695	0.028848	2.776613	lnc-C5orf22-8:1	0.048284	3.203795
NONHSAT017634	0.028876	2.028523	lnc-CNTN4-3:1	0.048311	2.025867
lnc-NAP1L3-8:1	0.028947	3.242532	NONHSAT059117	0.048379	2.403889
lnc-HPRT1-9:1	0.029041	2.730397	lnc-LPAR4-1:1	0.048691	2.107284
lnc-KCNB1-1:2	0.029166	5.540174	lnc-JAK1-8:1	0.049024	3.100057
NONHSAT137858	0.029188	2.286954	lnc-LINC00470-3:1	0.049145	2.468314
lnc-SNX10-4:1	0.029264	3.962636	ENST00000516699	0.049186	2.252165
lnc-SETD4-11:1	0.029358	5.268727	lnc-MRP63-7:2	0.049218	2.257006
lnc-BTG1-3:1	0.029364	4.188679	lnc-C11orf82-9:1	0.049221	2.226389
lnc-EBF3-11:1	0.029501	2.170030	NR_045117	0.049409	0.459001
lnc-RPS13-3:1	0.029547	5.254605	lnc-CTNNA3-8:1	0.049423	2.616655
ENST00000606596	0.029629	3.836024	lnc-ZNF286A-11:1	0.049549	2.501332
lnc-SH3RF2-4:1	0.029730	2.693729	lnc-CAMSAP2-2:1	0.049577	3.010308
lnc-DNAJC12-2:1	0.029822	3.449972	lnc-IL6-10:1	0.049733	2.217231
lnc-AVPR1A-3:1	0.029871	2.560073	lnc-BEND6-2:1	0.049876	2.067030
lnc-TNP1-7:1	0.029881	6.443609			

DE-lncRNAs, differentially expressed lncRNAs.

## lncRNA-NR\_104160 and ceRNA network for hypertension

**Table S3-2.** All DE-mRNAs between hypertensive and normotensive groups

DEG	P-value	FC	DEG	P-value	FC
DMRTC2	0.001304	4.781866	PERP	0.025326	2.020140
ZNF347	0.001943	0.498468	ZNF585B	0.026308	0.488404
SFTPA2	0.002651	2.180156	CTAG1B	0.027474	3.382239
HIGD1C	0.003248	2.243524	FFAR4	0.027566	0.483000
SPRED1	0.003497	0.471448	KCNK7	0.028835	2.692425
CD180	0.003790	0.441027	CDADC1	0.029385	2.201790
ANXA8	0.004021	2.207667	RPL17	0.029567	2.531104
VIT	0.005005	0.407097	NWD1	0.030331	2.816975
RGS1	0.005414	6.486876	SERINC2	0.030338	2.362981
GPR20	0.006608	0.368496	KLHDC7A	0.030688	2.047292
ANO2	0.007765	0.461345	PTS	0.031218	2.250719
SLC16A14	0.008164	2.174513	TOB1	0.032654	2.019600
NR4A3	0.008749	6.799959	RSL24D1	0.034190	2.221835
ERMN	0.009180	2.044825	SNRPF	0.034645	2.757363
DUSP4	0.009823	2.009596	GPR87	0.037195	3.032198
C17orf64	0.012822	2.203667	CTAG2	0.037550	3.759738
PCDHGA8	0.012950	0.478581	EGR3	0.038435	2.320014
LACTB	0.013003	2.127301	SEC14L2	0.038820	2.171246
CD69	0.013229	3.024099	CNTN1	0.040303	0.405808
SFN	0.015809	2.376723	C17orf107	0.043330	2.506023
DLC1	0.015886	0.432207	AREG	0.044621	9.667943
TIFAB	0.016275	0.499804	SIGLEC14	0.045134	0.443825
C12orf79	0.017032	2.105432	ZNF98	0.045337	2.516863
ID1	0.020953	20.331717	NR4A2	0.045953	4.892434
CALN1	0.022189	2.198163	VWDE	0.047126	2.261225
MS4A3	0.022953	2.077624	GZF1	0.049491	2.059196

DE-mRNAs, differentially expressed mRNAs.

**Table S4-1.** Cis Targeted Genes of 387 DE-lncRNAs identified by microarray

DE-lncRNA	Target Genes	N	DE-lncRNA	Target Genes	N
NR_029782	NR6A1; MIR181B-2	2	Inc-SPR-3:1	EXOC6B	1
Inc-EIF2AK4-6:1	THBS1	1	Inc-CHCHD3-1:1	EXOC4	1
Inc-IL2RA-4:1	IL2RA; Os07g0271000	2	Inc-BMP2K-4:1	BMP2K	1
Inc-CD52-1:1	AIM1L; UBXN11	2	Inc-KDELR2-1:2	KDELR2; KDELR	2
ENST00000614747	SLC18A2	1	Inc-PLIN1-1:1	PEX11A; PLIN1	2
ENST00000505646	UGT2B11; UGT1B4	2	Inc-C4orf34-8:1	UBE2K	1
ENST00000564481	AP3M2	1	Inc-TRAPP8-4:1	RNF125; F12L6.25	2
Inc-LOH12CR1-6:1	LRP6	1	Inc-TMEM185B-7:1	PTPN4	1
NR_024109	TYW1B; SBDS-1; CTNNB1-B	3	Inc-TRAF4-1:1	FAM222B	1
Inc-ACTR8-2:2	SELK	1	Inc-HSP90AA1-12:1	DYNC1H1	1
Inc-GPR125-5:1	KCNIP4	1	Inc-DPY19L4-2:1	ESRP1	1
Inc-ZNF720-3:4	ZNF720; GM10037	2	Inc-C21orf91-7:1	BTG3	1
ENST00000607744	LRRKIP2	1	Inc-HNRPDL-1:1	TMEM150C	1
ENST00000604491	FOXP1	1	Inc-AF165138.7.1-6:1	SAMSN1	1
Inc-KCNB2-3:3	TERF1	1	ENST00000418620	SP3	1
Inc-ARHGEF4-1:5	PLEKHB2	1	Inc-MKLN1-3:1	PODXL	1
Inc-ITPR1-1:1	BHLHE40	1	Inc-RRP1B-5:1	HSF2BP	1
ENST00000562459	GPR20	1	Inc-SAMHD1-2:1	RBL1	1

# lncRNA-NR\_104160 and ceRNA network for hypertension

lnc-PSMB4-2:1	POGZ	1	NONHSAT035867	RNF212B	1
lnc-NCDN-4:1	ZMYM4	1	NONHSAT080449	CASS4	1
lnc-FAM150A-4:1	ST18	1	lnc-CORO6-2:1	SSH2	1
lnc-RNF152-10:1	CDH20	1	lnc-GTF2F2-12:1	COG3	1
NONHSAT055372	RGS9	1	lnc-SLC25A1-4:1	CLTCL1; CHC	2
lnc-PSMA8-3:1	SS18	1	NONHSAT067683	ZNF813; VMN1R82	2
lnc-RNF146-3:2	ECHDC1	1	lnc-TCEANC-3:1	EGFL6	1
lnc-C19orf40-2:1	CEP89; FAAP24	2	lnc-HEATR2-2:3	PRKAR1B; KIN-2	2
lnc-GSG2-2:1	GSG2	1	ENST00000623366	HSPA4	1
lnc-IFNGR1-1:1	IFNGR1	1	NONHSAT055039	RPS6KB1	1
lnc-KIAA0754-4:1	MACF1	1	lnc-SAR1B-2:1	DDX46; CG6227	2
lnc-CNTNAP1-2:1	EZH1	1	ENST00000607698	RSRP1	1
NR_028484	ADORA2A; MGC82114	2	lnc-FAM9C-7:1	FRMPD4	1
lnc-NIPBL-5:1	C5orf42; 2410089E03Rik	2	NR_002564	SLC3A2	1
lnc-GSTM4-3:1	GSTM4; RPL7A	2	lnc-CNTROB-1:1	CNTROB	1
lnc-CSPG4-4:1	NRG4	1	lnc-CREB3L4-5:1	GATA2B	1
lnc-COA5-1:2	MGAT4A; MGATB4B	2	lnc-PHYHIP-3:1	SLC39A14	1
NONHSAT037842	NEK9	1	lnc-CCL5-3:1	HEATR9; RPL24	2
lnc-AC011467.1-2:1	LOC100193910	1	ENST00000520185	IDO1	1
lnc-NR2C1-2:5	NR2C1	1	lnc-PKP4-4:1	CCDC148	1
ENST00000618697	IGF1R	1	lnc-ERN1-2:1	ERN1	1
lnc-CAPN2-3:1	TP53BP2	1	lnc-PLEKHA5-6:1	AEBP2	1
ENST00000457602	ARHGAP25	1	lnc-APOD-2:1	APOD	1
lnc-ATP7A-1:1	PGK1; TAF9B; CG9961	3	ENST00000513581	SEC31A; THAP9	2
ENST00000607849	VGLL4	1	lnc-RP11-144F15.1.1-1:1	POLR3B; RPLLL128	2
lnc-RP4-811H24.6.1-3:1	BSDC1	1	lnc-TIPRL-1:1	TIPRL; RPL3402	2
lnc-PRDM10-2:1	APLP2	1	lnc-TLK1-3:1	GORASP2; GRASP65	2
ENST00000522057	PPP1R3B	1	lnc-SLC25A17-2:1	MKL1; ADSL	2
lnc-C8G-2:1	TRAF2; RABL6	2	ENST00000516179	SOD2	1
lnc-XRCC2-5:1	RL36A	1	lnc-C11orf63-3:1	LOC103435013	1
lnc-DAOA-10:1	RPL-35	1	lnc-TNFRSF11A-1:1	RPL17	1
NONHSAT117148	MAMU-DPA	1	lnc-TMSB15A-4:1	GPRASP1	1
lnc-MMADHC-5:1	LYPD6B	1	lnc-CCAR1-3:1	TET1	1
NONHSAT018112	FAR1	1	NONHSAT101519	IL6ST	1
ENST00000620896	SRGN	1	lnc-CPB2-1:1	ZC3H13	1
lnc-EIF2B1-5:1	ATP6VOA2	1	lnc-ZBTB6-1:1	RC3H2	1
lnc-CSGALNACT1-4:1	PSD3	1	lnc-CEBPZ-1:1	SULT6B1	1
lnc-EGR1-1:1	EGR1	1	NONHSAT036908	GNPNAT1; MYEOV2	2
lnc-LDB1-4:1	C10orf76	1	lnc-A2M-2:1	PZP; A2M	2
lnc-RIOK2-9:1	CHD1	1	NONHSAT062956	JUND	1
NONHSAT126218	SFRP1	1	lnc-NPIPL2-5:1	GLG1	1
NONHSAT127886	LAPTM4B; RPL39	2	lnc-CMC1-4:1	ZCWPW2	1
lnc-AC023632.1-3:1	LOC100306335	1	ENST00000432229	FARP1	1
lnc-ZNF366-13:1	TNP01	1	lnc-MOK-6:1	HSP90AA1	1
lnc-ZNF292-1:1	HTR1E; RPL7L1	2	ENST00000516415	B4GALT1	1
lnc-RALGAPA2-7:1	CFAP61	1	lnc-TMEM202-4:1	SENP8	1
lnc-C7orf30-4:1	IGF2BP3	1	ENST00000619883	RHN01; FOXM1	2
lnc-TAS1R1-1:1	NOL9	1	lnc-NXPH2-11:1	LRP1B	1
lnc-STX7-1:1	STX7	1	lnc-PLEKHM3-2:1	PLEKHM3	1
ENST00000609669	UAP1	1	lnc-PON3-1:1	PON2	1
lnc-NOS2-6:1	NOS2	1	NONHSAT100937	RPL17	1
lnc-CENPJ-4:1	OS08G0156800	1	lnc-FAM126A-1:1	FAM126A	1
lnc-SMC4-2:1	IFT80	1	lnc-FBXO10-1:1	POLR1E	1

## lncRNA-NR\_104160 and ceRNA network for hypertension

lnc-LPAR1-2:1	MUSK	1	lnc-ACAT1-2:1	CUL5	1
lnc-CD40LG-4:1	ARHGEF6	1	lnc-GPR19-2:1	CREBL2	1
lnc-CAMP-3:1	MAP4; RPS27B	2	ENST00000565797	ABHD17A; LEP	2
lnc-DLD-2:1	THAP5; DNAJB9	2	lnc-FAM71D-3:1	GPHN; GPHNA	2
NONHSAT001395	EIF4G3	1	NR_046400	DIP2A	1
lnc-PSMA6-3:1	KIAA0391; AT1G33120	2	lnc-RPS27-2:2	NUP210L; RAB13	2
lnc-DENND5B-2:1	DENND5B; LOC100276690	2	NONHSAT053484	IKZF3	1
lnc-CC2D2A-1:1	C1QTNF7	1	ENST00000563775	C25H16orf59	1
lnc-ATG14-3:1	FBXO34	1	lnc-RPL7A-2:1	SURF6; RPL21A	2
NR_104160	EPC1	1	lnc-TMEM183A-2:1	TMEM183A	1
lnc-IFT74-6:1	TEK	1	lnc-MDC1-1:1	NRM; RPL7L1	2
NONHSAT004372	LRRC8C	1	ENST00000515939	UBE2R2	1
lnc-RGS13-2:2	RGS1	1	lnc-RPL36A-1:1	TIMM8A	1
lnc-AC012123.1.1-4:1	KLHL14	1	lnc-FBXL2-2:1	CLASP2; COX6C-PS1	2
lnc-SIRT5-4:2	PHACTR1	1	NR_029663	SMC4; IFT80; MIR15B	3
lnc-RP11-439E19.8.1-1:1	SCCPDH; LOC101866277	2	lnc-ALS2CR8-2:1	CARF	1
lnc-KATNAL2-10:1	ST8SIA5	1	NR_028287	ZNF774; GRL1	2
lnc-BORA-14:1	DACH1; DACHD	2	ENST00000539313	TSG101	1
lnc-TPRG1-1:1	LPP	1	lnc-ADAMTS7-2:1	MORF4L1	1
lnc-FAM82A1-8:1	RPL7	1	ENST00000603472	SH3D19	1
lnc-SLC36A4-7:1	FAT3	1	lnc-DUSP19-3:1	DNAJC10; DUSP19	2
ENST00000413347	SNAP47	1	lnc-AC016586.1.1:4	EBI3	1
lnc-PRDX6-4:1	RPL26	1	lnc-RP11-32K4.2.1-3:1	COX6C	1
ENST00000608498	RGD1306941	1	lnc-NR2F2-10:1	GM29683	1
lnc-TFDP2-9:1	TPT1	1	lnc-C3orf72-4:2	PIK3CB	1
lnc-SUPT6H-2:7	SPAG5	1	lnc-ACBD6-2:1	QSOX1	1
lnc-DYRK1A-1:1	DYRK1A	1	lnc-NDUFS8-6:1	UNC93B1	1
lnc-UCP3-5:1	CACD3	1	lnc-RP11-1396013.13.1-3:1	OLR1151	1
NR_105043	FBXL13	1	lnc-ZNF485-5:1	ZGC:173624	1
lnc-GCA-6:1	KCNH7	1	lnc-GPR63-8:1	T21H19.50	1
lnc-C7orf41-2:1	MTURN	1	lnc-PTPN13-1:1	C4orf36	1
lnc-CXCR3-12:1	MGC152340	1	NR_045117	WASH1	1
lnc-CYTL1-5:2	CG32944	1	lnc-CTNNAA3-8:1	LOC100194643	1
lnc-BEND6-2:1	DST	1	lnc-NHSL2-5:1	MGC152340	1
NR_125398	GATA2	1	lnc-LETMD1-1:1	TFCP2; LOC101866801	2
lnc-CNOT7-5:1	ZDHHC2	1	lnc-C3orf65-9:1	SENP2	1
lnc-RNF222-1:1	MYH10	1	lnc-PLBD1-1:3	ATF7IP	1
lnc-LONP2-8:1	N4BP1; LOC100160790	2	lnc-ARSG-2:6	PRKAR1A	1
lnc-DTX1-2:1	DTX1; OAS3	2	lnc-RP11-166B2.1.1-3:1	GSPT1	1
lnc-ACBD3-1:1	ACBD3	1	lnc-PRRG4-4:1	CCDC73	1
lnc-GTF2A1-4:1	STON2; LOC100284180	2	lnc-KBTBD13-1:1	UBAP1L; RPL17	2
lnc-RP11-849H4.2.1-2:1	RNF121	1	lnc-C19orf12-7:1	URI1; LOC100162964	2
lnc-TUFM-4:1	ATXN2L	1	NR_002562	SLC3A2	1
NR_036695	AGOS_ACL076W	1	lnc-WDR36-1:1	CAMK4	1
lnc-MEX3C-6:1	LOC105018348	1	lnc-MBL2-9:1	RPL31	1
lnc-RBMY1J-7:1	PRY2; RBMY	2	lnc-IQCH-4:1	AAGAB; RPS2402	2
NONHSAT017634	STIM1	1	lnc-C5orf22-8:1	PDZD2	1
lnc-HPRT1-9:1	PLAC1	1	lnc-LPAR4-1:1	LPAR4	1
lnc-KCNB1-1:2	KCNB1	1	lnc-JAK1-8:1	DNAJC6	1
lnc-EBF3-11:1	GLRX3	1	ENST00000516699	GPC5; GPC5A	2
lnc-RPS13-3:1	PIK3C2A	1	lnc-MRP63-7:2	SAP18	1
ENST00000606596	SARAF	1	lnc-ZNF286A-11:1	CDRT4	1
lnc-SH3RF2-4:1	LARS; RPL35A	2	lnc-CAMSAP2-2:1	GPR25	1

## lncRNA-NR\_104160 and ceRNA network for hypertension

lnc-DNAJC12-2:1	<i>HERC4</i>	1	lnc-TEKT3-1:1	<i>PMP22</i>	1
lnc-TMEM14C-3:1	<i>MAK</i>	1	lnc-EREG-1:1	<i>AREG</i>	1
ENST00000588182	<i>CYP4F2</i>	1	NONHSAT082239	<i>BACE2</i>	1
lnc-NPBWR1-4:1	<i>RB1CC1</i>	1	lnc-ABCC11-2:1	<i>LONP2</i>	1

DE-lncRNAs, differentially expressed lncRNAs.

**Table S4-2.** Trans Targeted Genes of 387 DE-lncRNAs identified by microarray

DE-lncRNA	Target Genes	N
NR_033186	ADAT1; ANGEL2; ANKRD60; ARPP19; ARSB; ATP6V0A2; C4orf26; CACUL1; CHMP1B; CHP1; CLUAP1; CNNM3; CXorf56; CYP51A1; DNAL1; EIF5A2; ERCC4; FAM117B; FCAR; GPATCH2L; GPI; IKZF3; LY75; MAGEA10; MDM4; MON2; MRPL44; NICN1; NMNAT1; PDXK; PGAM5; PNMA2; POU5F1; PPM1K; RASGRP4; RBM34; RBMS2; SCAMP4; SERPINB1; SFMBT2; SIX4; SLC16A4; SPRED1; STRIP2; SV2C; TRPV1; TTC28; USP33; XPNPEP3; ZNF555; ZNF713	51
lnc-CNOT10-2:1	ACPP; BAG1; BLZF1; CFLAR; CIAO1; CORO2A; CPT2; DCAF10; DTD2; EAF1; EXOG; IL4R; LPIN1; MARVELD3; MFSD11; MRPS25; NRIP2; ORAI2; PECAM1; RAD23B; SEMA5A; SLC35E1; SLC6A11; TCEANC2; TOP3A; VPS53; WDR73; ZC2HC1C; ZKSCAN1; ZNF43; ZNF841	31
lnc-LOH12CR1-6:1	BRCA1; CC2D2B	2
lnc-ARHGEF4-1:5	AKAP5; ATG14; C8orf44-SGK3; CDC157; CCL5; CENPBD1; CTNS; CYB5D2; CYCS; CYP20A1; DDX51; DESI1; DPP9; GLTP; HOOK3; ITIH5; KLHL28; MAP3K9; OGFOD1; OPRK1; PAK3; PDP2; PNPT1; RAD1; RBL1; SCAF11; SGK3; SLC31A1; SLC35E3; TAF8; URB1; WDR31; XIAP; ZBTB43; ZNF260; ZNF333; ZNF426; ZNF665	38
lnc-PSMB4-2:1	ADAT1; AHR; ALDH1B1; ALDH6A1; ALG1; ANKRD40; AP1S3; AP4E1; APOBEC3D; ARSA; ATP6V1A; BROX; C1orf210; CA5B; CDKAL1; CHMP1B; CLUAP1; CRISPLD2; CRX; CXorf56; CYB5R3; CYP51A1; DENND6A; DESI1; EMP2; FAM114A1; FAM117B; FAM234B; FBXW2; FOXK1; GFOD2; GGA2; GNE; GOLGA3; GPANK1; GSDMA; HOOK3; IBA57; IFNLR1; IKZF3; INMT; IRGQ; JOSD1; LAMP3; LMX1B; LY75; MAP3K13; MBOAT1; MEFV; MID1P1; MTX3; NCOA6; NDUFV3; NF2; NLRC3; NOL9; NT5DC3; NUAK2; NXN; OLFML2A; OPRM1; OR7D2; OSBPL2; PACS2; PDDC1; PDE12; PIAS2; PIK3C2B; POU5F1; PSTPIP2; R3HDM2; RBM34; RBMS2; RIPPLY3; RNFT; SCA1; SCAMP4; SERPINB9; SHROOM1; SIRT5; SLC15A1; SLC25A15; SLC30A6; SLC35F1; SLC48A1; SMIM14; SMU1; SNX22; SPAST; SPECC1; SPRED1; SS18; STK17B; STON2; STRIP2; STYX; SYNRG; TAF8; TANGO2; TAPBP; TATDN3; TBC1D15; TLCD2; TMEM236; TMEM56; TMX4; TOR1AIP2; TRAF3IP1; TRAPP2; TRPV1; UGGT1; WAC; WDR31; WDR55; WDR76; XIAP; ZC3H12B; ZCCHC8; ZNF250; ZNF260; ZNF426; ZNF445; ZNF451; ZNF543; ZNF550; ZNF555; ZNF699; ZNF765; ZNRF3; ZSCAN29	130
lnc-GSG2-2:1	ABHD11; AFAP1L1; ALDH6A1; AP4E1; ARPP19; CCL5; CHRNB1; CLN8; CLUAP1; DENND6A; FAM117B; GCLM; GLTP; GSDMA; GSTM3; HILPDA; IKZF3; LAMP2; LRRC27; MLANA; NF2; NXPE3; OPA3; ORAI2; PER2; PNPO; RAB36; RARS2; RLIM; RNF125; SCA1; SIRT5; SMCR8; SMIM14; SPAST; STAT2; STK17B; TMEM120B; TOP3A; TRIM16; WDR55; WDR76; ZKSCAN1; ZNF490; ZNF681; ZNF765; ZSCAN29	47
lnc-WDR60-10:1	BRCA1; CC2D2B	2
lnc-SLC25A26-6:1	BRCA1; CC2D2B	2
lnc-FBXO10-1:1	BRCA1; CC2D2B; TCF4	3
lnc-ADAMTS7-2:1	BRCA1	1
lnc-NDFIP2-21:1	BRCA1; CC2D2B	2
lnc-MUC20-7:2	BRCA1; CC2D2B	2
lnc-RNF144B-5:1	BRCA1; CC2D2B	2
NR_028484	ADORA2A	1
lnc-C8G-2:1	ACSL6; AKIP1; APOL1; APOL2; APPL1; C11orf58; C17orf75; C3orf70; CASP2; CASP8; CBX5; CCDC122; CDH6; CHST5; CPT1A; CRX; CXADR; CYB5R3; DSC3; DTD2; DTX3L; ERO1A; FBXL18; FBXL20; FBXO44; FOXK1; FUT1; GPR83; GREB1; HOMEZ; HOOK3; HYKK; IL17RD; INMT; IRAK4; JAK3; KIAA0825; KIAA1143; KIAA1210; KIAA1456; LEPROT; LIN54; LIX1L; LLGL1; LPP; LRRC28; LYZ; MALL; MAPK1IP1L; MAVS; MBOAT1; MPI; MPRIP; MTFFMT; NANOG; NEK2; NR6A1; NRIP3; NRXN3; OPRM1; PAPOLA; PCDH49; PDE7A; PEX13; PHKG2; PIGR; PLEKHG2; POLA2; PPIL6; PSPH; PTCHD4; PXMP4; RAB21; RBMS2; RRP15; SCARF1; SCN1M; SDE2; SERINC3; SERPINB9; SHROOM1; SIX4; SKA1; SKP2; SLC35F1; SLU7; SMAD2; SPATA5; SSR3; ST8SIA4; STEAP2; TADA3; TAPBP; TBC1D24; TBCCD1; TERF2; TMEM41B; TNFSF15; TOX4; TRAF3IP2; TRIM59; TTC39C; TVP23C; ULBP3; ULK2; UTP11; VHL; VIPR2; XPNPEP3; YES1; ZNF114; ZNF124; ZNF2; ZNF286A; ZNF286B; ZNF426; ZNF454; ZNF461; ZNF490; ZNF543; ZNF548; ZNF557; ZNF714; ZNF718; ZNF780B; ZNF793; ZNF805; ZSCAN22; ZSWIM1	129
ENST00000620896	AAK1; ADARB2; ANGPT4; ANKS4B; AP4S1; ATAD3C; ATP6V1C1; C14orf178; C15orf52; CCL28; CDCP1; CELF3; CLCC1; CLDN11; CLEC19A; CYP20A1; DERL2; DNAL1; DTD2; DTX3L; EEF2K; EIF2AK2; FAM156A; FAM227A; GBP6; GSDMA; H6PD; HUNK; HUS1; HYKK; IKZF3; IL17RA; ITPR1PL2; KCNE4; KIAA1324; LITAF; LRPAP1; LRRC10; LY75; MDM4; ME2; MPP7; MRPL30; MYL10; NOS3; NSL1; NXPE3; OGFLR1; OLFML1; P2RY2; PDE6A; PER2; PHACTR4; PLEKHH2; PM20D2; PRRG4; RBBP4; RBMS2; RNF135; S1PR2; SCA1; SHCBP1; SIRT3; SKA1; SLC24A2; SLC26A2; SLC35F6; SLC6A11; SMU1; SNX27; SPN; STAP2; STYX; TINAGL1; TLCD2; TMEM241; TMT1; TOR1AIP2; TRIP11; TRMT2B; TTC22; UVSSA; ZADH2; ZBTB40; ZC3HAV1; ZFP14; ZNF234; ZNF544	88

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Inc-TUFM-4:1	ADAMTSL2; ALDH6A1; APOL1; ARPP19; ATL3; BNIP1; CEBPZOS; CHMP1B; CHP1; COX15; COX6B2; CXADR; CYP20A1; CYP4V2; EIF5A2; EXPH5; FAM114A1; FBXO45; FOXK1; GREB1; GTF2E1; HECA; KBTBD12; KIN; KLHL30; MBOAT1; ME2; MPZL3; MS4A10; MYLK3; NOL9; NR6A1; PIK3C2B; PLEKHA5; PNPO; PPM1K; RAB22A; RAB33B; RDH10; REEP3; RLIM; SCARF1; SCNM1; SCUBE3; SGTB; SIX4; SKP2; SLC16A4; SPAST; TANGO2; TERF2; TGFBRAP1; TOR1AIP2; TOX4; TRAPPC2; ZNF426	56
Inc-BTG1-3:1	H6PD; ICMT; MTX3; RBMS2; REL; SLFN13	6
Inc-NR4A2-8:1	ALDH6A1; ATP6V1A; ATXN3; CEP104; CLUAP1; CXorf56; DSC3; ERCC4; FOXK1; GNG4; HAVCR2; HSD17B13; IFNLR1; MANSC1; MED18; MTFMT; MYEF2; NOL9; OGFR1L1; ORAI2; OSBP2; PDE12; PSTPIP2; RAB22A; RASGRP4; RBM34; SALL4; SAR1B; SGTB; SIRT5; SLC16A4; SLC25A32; SPAST; SPRED3; SS18; TANGO2; TIMM10B; TOP3A; VENTX; VPS33A; ZNF264; ZNF426; ZNF713	43
Inc-GPR19-2:1	BRCA1; CC2D2B	2
Inc-RPL7A-2:1	BRCA1; CC2D2B	2
Inc-RPL36A-1:1	BRCA1	1
Inc-ADD3-5:1	BRCA1; CC2D2B	2
Inc-STX7-1:1	BRCA1; CC2D2B	2
Inc-CDH13-4:1	BRCA1; CC2D2B	2
Inc-ZNF485-5:1	BRCA1	1
Inc-XRCC6BP1-7:1	CC2D2B	1
Inc-CNTROB-1:1	BRCA1	1
Inc-HNRPDL-1:1	ACSL6; ALDH5A1; ARIH20S; ATAD3C; ATF6; BCDIN3D; BRIP1; C17orf75; CAMK1D; CCL5; CD82; CD99; COA7; CPT1A; DENND2A; DIS3L; DNAJC21; ENTPD4; F2R; FAHD1; FBXL18; GBP4; GGPS1; GNE; GSTM3; ICA1L; IDO1; IDS; IRF1; KCNA7; KIAA1324; KREMEN1; LILRB3; LRRC27; LRPC58; MANEAL; MEAF6; MGAT4A; MRPS10; PCDH11Y; PDE6A; PIGX; PLEKH2; PPP1R12B; RAB3B; RNF14; RNF41; RUNDC1; SHROOM1; SHROOM4; SLC26A2; SLC43A2; SPATA5; SVOP; SYNPO2L; TADA2B; TBC1D15; TNFAIP8L1; TONS1; TPM3; UBN2; UGGT1; WDR45; XIAP; XPNPEP3; ZBTB24; ZC3H12D; ZFP82; ZMAT3; ZNF114; ZNF124; ZNF669	72
Inc-CORO6-2:1	BRCA1; CC2D2B; TCF4	3
Inc-RP11-144F15.1.1-1:1	ADAT1; ALDH6A1; ATG14; BROX; BVES; C17orf75; CA5B; CD99; CYP20A1; CYP4V2; CYP51A1; DNAL1; EMP2; FAM118A; FOXK1; GFPT1; GNG4; GREB1; HAVCR2; HDAC2; HOMEZ; HYKK; ICE2; JAKMIP2; LPP; MED18; MTFMT; MYLK3; NMT2; NOL9; NT5DC3; NUDT19; ORAI2; PAFAH1B2; PDDC1; PHC3; PIK3C2B; PTGIS; RAB21; RBMS2; RHD; RHOF; SAR1B; SCIMP; SH3BP2; SLC16A4; SLC1A2; SLC4A8; SLC9A7; SNTB2; SSR1; TAF8; TOR4A; TOX4; TRAF3IP2; TRIM16; VENTX; XPNPEP3; YES1; ZC3HAV1; ZNF426; ZNF43; ZNF445	63
Inc-WNT7B-2:1	ACER3; ADAT1; AFMID; AP3S2; AP4S1; ARCN1; ARGFX; ARSB; AS3MT; ASB6; ATL3; ATP1B4; ATP6VOA2; BLOC1S2; C15orf38-AP3S2; C19orf12; C9orf3; CA5B; CCDC141; CCNDBP1; CD99; CDCP1; CEP41; CHMP1B; CHP1; CLCC1; CRX; DES1; DISC1; EEF2K; EIF5A2; EVI5; FAAP20; FAM114A1; FBXL18; FLCN; FNX; GNE; GPI; GSMDA; GTPBP3; H6PD; IFNLR1; IKZF3; IL11; IL17RD; JAKMIP2; KCNJ11; KIF18B; KIF1B; LRRN4CL; MAP7D3; MAVS; MCTS1; MDM4; METTL8; MRO; NATD1; NDUFV3; NMNAT1; NMUR1; NQO1; OLFML2A; OPA3; P2RY2; PDDC1; PGPEP1; PHACTR4; PHC3; PIK3CD; POU5F1; PRRG4; RASGRP4; RBMS2; RHD; RPL15; S1PR2; SCA1; SEPN1; SFMBT2; SFXN2; SHISA9; SKP2; SLC14A2; SLC4A8; SLC5A5; SPAG9; SPATA5; SVOP; TFPD2; TGFBRAP1; TIMM50; TOX4; TRAF3IP2; TTC28; USP33; USP9X; XPNPEP3; ZBTB8A; ZNF280C; ZNF426; ZNF621; ZNF677; ZNF713; ZNF829	105
Inc-TLK1-3:1	GORASP2	1
Inc-C2orf57-3:1	BRCA1; CC2D2B	2
Inc-TMEM202-4:1	ABHD11; ADAT1; AP1S3; AP3M1; ARHGEF39; ATCAY; BCDIN3D; BHMT2; BORCS7; C11orf63; C6orf89; CARF; CCDC142; CCL5; CCR6; CD99; CEP19; COX18; CPM; CRX; CWC25; CXorf38; DBT; DCAF10; DMC1; DNAJC21; DNAL1; DNPEP; DPP9; DSG2; DSTYK; ECE1; EEF2K; ELMOD1; FBXL18; FKBP14; GLTP; GNE; GPR155; GPR82; GRM6; GSTM3; HCAR1; ICA1L; IDS; IKZF3; IL13RA1; IL17RD; IRF1; ITPR1L2; KCNJ3; KDELC2; KIAA0355; KREMEN1; LAIR1; LAMP2; LRRC58; LRRN4CL; MANEAL; MAPK13; MARVELD3; MBOAT2; METTL21A; METTL2A; METTL2B; MREG; NMT2; NUBPL; ORAI2; PARK2; PDDC1; PDE4C; PDE6A; PDE7A; PGPEP1; PHACTR4; RBBP5; RHBTL2; RPL7L1; RRP7A; SCAF11; SEC14L4; SGSM1; SHPK; SHROOM1; SLC25A15; SLC44A4; SLC6A4; SMIM14; SNIP1; SPATS2; SRSF10; SYNPO2L; TIMM50; TLR7; TMEM168; TMEM236; TMEM56; TNFSF15; TONS1; TPM3; TRAF3IP2; TRIM65; TRIM66; TSNAX; TPPAL; TUBGCP4; UGGT1; UPK1B; VHL; XIAP; ZBTB43; ZBTB8A; ZHX3; ZKSCAN1; ZMAT3; ZNF106; ZNF34; ZNF655; ZNF749; ZNFX1	121
Inc-CNTN4-3:1	BRCA1; CC2D2B	2
Inc-TRAPPC8-4:1	HK1	1
Inc-GJA1-5:1	HK1	1

DE-IncRNAs, differentially expressed IncRNAs.

**Table S5.** lncRNA-mRNA co-expression pairs

lncRNA	mRNA	PCC	lncRNA	mRNA	PCC
NR_104160	ID1	0.951	lnc-CNKS3-2:1	RSL24D1	0.975
ENST00000565797	CD69	0.951	NONHSAT062956	RPL17	0.975
lnc-LDB1-4:1	SNRPF	0.951	lnc-HPRT1-9:1	RPL17	0.975
ENST00000565797	GZF1	0.952	NONHSAT053484	RSL24D1	0.976

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lnc-UCP3-5:1	<i>EGR3</i>	0.952	lnc-IQCH-4:1	<i>RSL24D1</i>	0.976
lnc-ZBTB6-1:1	<i>SNRPF</i>	0.952	lnc-FAM71D-3:1	<i>RSL24D1</i>	0.976
lnc-LONP2-8:1	<i>PTS</i>	0.952	NONHSAT035867	<i>RPL17</i>	0.976
lnc-AVPR1A-3:1	<i>PTS</i>	0.952	lnc-DUSP19-3:1	<i>RPL17</i>	0.976
NONHSAT018112	<i>RSL24D1</i>	0.952	lnc-NR2F2-10:1	<i>RPL17</i>	0.976
lnc-GTF2F2-12:1	<i>SNRPF</i>	0.953	lnc-RIOK2-9:1	<i>CDADC1</i>	0.977
lnc-SAMHD1-2:1	<i>RSL24D1</i>	0.953	lnc-LPAR4-1:1	<i>RSL24D1</i>	0.977
lnc-C7orf30-4:1	<i>RSL24D1</i>	0.953	lnc-CCAR1-3:1	<i>RSL24D1</i>	0.977
lnc-DAOA-10:1	<i>RPL17</i>	0.953	lnc-SAR1B-2:1	<i>RSL24D1</i>	0.977
lnc-DNAJC12-2:1	<i>RPL17</i>	0.953	lnc-SLC14A1-1:1	<i>RPL17</i>	0.977
lnc-SPR-3:1	<i>RSL24D1</i>	0.954	lnc-KBTBD13-1:1	<i>RPL17</i>	0.977
lnc-CDH17-2:1	<i>RSL24D1</i>	0.954	NR_104160	<i>PTS</i>	0.978
lnc-HPRT1-9:1	<i>RSL24D1</i>	0.954	NONHSAT144261	<i>RSL24D1</i>	0.978
lnc-SETD4-11:1	<i>RPL17</i>	0.954	lnc-CMC1-4:1	<i>RSL24D1</i>	0.978
lnc-ERMN-2:1	<i>PTS</i>	0.955	lnc-IQCH-4:1	<i>RPL17</i>	0.978
lnc-CAPN2-3:1	<i>PTS</i>	0.955	lnc-STK35-3:1	<i>RPL17</i>	0.978
lnc-TIPRL-1:1	<i>RSL24D1</i>	0.955	lnc-COMMD3-3:1	<i>RPL17</i>	0.978
ENST00000565797	<i>ERMN</i>	0.956	lnc-DUSP19-3:1	<i>RSL24D1</i>	0.979
lnc-RIOK2-9:1	<i>GZF1</i>	0.956	lnc-APOD-2:1	<i>RSL24D1</i>	0.979
lnc-GTF2F2-12:1	<i>GZF1</i>	0.956	NONHSAT067683	<i>RPL17</i>	0.979
lnc-KDELR2-1:2	<i>CDADC1</i>	0.956	lnc-PLEKHM3-2:1	<i>RPL17</i>	0.979
lnc-GTF2F2-12:1	<i>CDADC1</i>	0.956	lnc-CHCHD3-1:1	<i>RPL17</i>	0.979
NONHSAT054838	<i>RPL17</i>	0.956	lnc-CCL5-3:1	<i>RSL24D1</i>	0.98
lnc-C4orf34-8:1	<i>RPL17</i>	0.956	lnc-CEBPZ-1:1	<i>RSL24D1</i>	0.98
lnc-CAPN2-3:1	<i>ID1</i>	0.957	lnc-AVPR1A-3:1	<i>RSL24D1</i>	0.981
ENST00000565797	<i>SNRPF</i>	0.957	lnc-TRAF4-1:1	<i>RSL24D1</i>	0.981
NR_024109	<i>PTS</i>	0.957	lnc-CAMSAP2-2:1	<i>RSL24D1</i>	0.981
lnc-AC011467.1-2:1	<i>RSL24D1</i>	0.957	lnc-ZNF286A-11:1	<i>RPL17</i>	0.981
NONHSAT130299	<i>RPL17</i>	0.957	lnc-C7orf30-4:1	<i>RPL17</i>	0.981
lnc-RPS13-3:1	<i>RPL17</i>	0.957	NONHSAT100937	<i>RSL24D1</i>	0.982
lnc-KCNB2-3:3	<i>SFN</i>	0.958	lnc-CENPJ-4:1	<i>RSL24D1</i>	0.982
lnc-ZNF286A-11:1	<i>RSL24D1</i>	0.958	lnc-C3orf65-9:1	<i>RSL24D1</i>	0.982
NR_002562	<i>RPL17</i>	0.958	lnc-MEX3C-6:1	<i>RPL17</i>	0.982
lnc-TIPRL-1:1	<i>RPL17</i>	0.958	NONHSAT018112	<i>RPL17</i>	0.982
lnc-NAP1L3-8:1	<i>RPL17</i>	0.958	lnc-SAMHD1-2:1	<i>RPL17</i>	0.982
NONHSAT127446	<i>RPL17</i>	0.958	lnc-ERMN-2:1	<i>CDADC1</i>	0.983
lnc-LDB1-4:1	<i>CDADC1</i>	0.959	lnc-PRRG4-4:1	<i>RSL24D1</i>	0.983
lnc-FAM71D-3:1	<i>CD69</i>	0.959	lnc-WSB1-7:2	<i>RSL24D1</i>	0.983
lnc-ARSG-2:6	<i>PTS</i>	0.959	lnc-PTPRZ1-10:1	<i>RSL24D1</i>	0.983
lnc-RP11-166B2.1.1-3:1	<i>RPL17</i>	0.959	NONHSAT128057	<i>RPL17</i>	0.983
NONHSAT126218	<i>RPL17</i>	0.959	lnc-CCL5-3:1	<i>RPL17</i>	0.983
lnc-TMEM74-3:1	<i>RPL17</i>	0.959	lnc-MEX3C-6:1	<i>RSL24D1</i>	0.984
ENST00000607698	<i>PTS</i>	0.961	lnc-COMMD3-3:1	<i>RSL24D1</i>	0.984
lnc-FAM71D-3:1	<i>PTS</i>	0.961	lnc-NT5E-4:1	<i>RSL24D1</i>	0.984
NONHSAT035867	<i>RSL24D1</i>	0.961	lnc-CES1-4:1	<i>RSL24D1</i>	0.984
lnc-A2M-2:1	<i>RPL17</i>	0.961	lnc-C8G-2:1	<i>RPL17</i>	0.984
lnc-FAM71D-3:1	<i>RPL17</i>	0.961	NONHSAT059117	<i>RPL17</i>	0.984
lnc-C19orf12-7:1	<i>RPL17</i>	0.961	lnc-ARSB-1:1	<i>RPL17</i>	0.984
lnc-ZBTB6-1:1	<i>CDADC1</i>	0.963	lnc-AC011467.1-2:1	<i>RPL17</i>	0.984
lnc-BEND6-2:1	<i>RPL17</i>	0.963	lnc-PLEKHM3-2:1	<i>RSL24D1</i>	0.985

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lnc-SPR-3:1	<i>RPL17</i>	0.963	lnc-MBL2-9:1	<i>RSL24D1</i>	0.985
lnc-ARHGEF4-1:5	<i>SFN</i>	0.964	lnc-NT5E-4:1	<i>RPL17</i>	0.985
lnc-KDELR2-1:2	<i>PTS</i>	0.964	lnc-CES1-4:1	<i>RPL17</i>	0.985
NR_024109	<i>CDADC1</i>	0.965	lnc-CAMSAP2-2:1	<i>RPL17</i>	0.985
NONHSAT001395	<i>RSL24D1</i>	0.965	lnc-NR2F2-10:1	<i>RSL24D1</i>	0.986
lnc-KDELR2-1:2	<i>SNRPF</i>	0.966	lnc-ARSB-1:1	<i>RSL24D1</i>	0.986
lnc-RIOK2-9:1	<i>PTS</i>	0.966	lnc-NRN1-6:2	<i>RSL24D1</i>	0.986
lnc-PLEKHA5-6:1	<i>RSL24D1</i>	0.966	lnc-GPR63-8:1	<i>RPL17</i>	0.986
lnc-DLEU7-9:1	<i>RSL24D1</i>	0.966	lnc-CEBPZ-1:1	<i>RPL17</i>	0.986
NR_104160	<i>SNRPF</i>	0.967	lnc-CCAR1-3:1	<i>RPL17</i>	0.986
ENST00000565797	<i>PTS</i>	0.967	lnc-WSB1-7:2	<i>RPL17</i>	0.986
lnc-LONP2-8:1	<i>RSL24D1</i>	0.967	lnc-C3orf65-9:1	<i>RPL17</i>	0.986
lnc-CHCHD3-1:1	<i>RSL24D1</i>	0.967	lnc-KBTBD13-1:1	<i>RSL24D1</i>	0.987
NONHSAT080449	<i>RSL24D1</i>	0.967	lnc-DTHD1-16:1	<i>RSL24D1</i>	0.987
NONHSAT067683	<i>RSL24D1</i>	0.968	NONHSAT144261	<i>RPL17</i>	0.987
NR_104160	<i>GZF1</i>	0.969	NONHSAT100937	<i>RPL17</i>	0.987
ENST00000606596	<i>C12orf79</i>	0.969	lnc-PRRG4-4:1	<i>RPL17</i>	0.987
lnc-ARSG-2:6	<i>CDADC1</i>	0.969	lnc-CENPJ-4:1	<i>RPL17</i>	0.987
lnc-MKLN1-3:1	<i>RSL24D1</i>	0.969	lnc-MBL2-9:1	<i>RPL17</i>	0.988
lnc-CDH17-2:1	<i>RPL17</i>	0.969	lnc-FAM9C-7:1	<i>RSL24D1</i>	0.989
NR_036695	<i>RPL17</i>	0.969	lnc-LPAR4-1:1	<i>RPL17</i>	0.989
lnc-ATG14-3:1	<i>RPL17</i>	0.969	lnc-RALGAPA2-7:1	<i>RPL17</i>	0.989
lnc-ZBTB6-1:1	<i>PTS</i>	0.97	lnc-DTHD1-16:1	<i>RPL17</i>	0.989
lnc-MDC1-1:1	<i>RSL24D1</i>	0.97	lnc-TRAF4-1:1	<i>RPL17</i>	0.989
lnc-SLC25A1-4:1	<i>RSL24D1</i>	0.97	lnc-SAR1B-2:1	<i>RPL17</i>	0.989
lnc-FBXL2-2:1	<i>RPL17</i>	0.97	lnc-RRP1B-5:1	<i>RPL17</i>	0.989
lnc-ERMN-2:1	<i>ERMN</i>	0.971	lnc-C8G-2:1	<i>RSL24D1</i>	0.994
ENST00000565797	<i>CDADC1</i>	0.971	lnc-FAM9C-7:1	<i>RPL17</i>	0.994
lnc-GTF2F2-12:1	<i>PTS</i>	0.971	lnc-PTPRZ1-10:1	<i>RPL17</i>	0.994
NONHSAT128057	<i>RSL24D1</i>	0.971	NONHSAT053484	<i>RPL17</i>	0.993
lnc-UBE2NL-2:1	<i>RPL17</i>	0.971	lnc-MDC1-1:1	<i>RPL17</i>	0.993
lnc-LDB1-4:1	<i>PTS</i>	0.972	NONHSAT001395	<i>RPL17</i>	0.993
lnc-LONP2-8:1	<i>RPL17</i>	0.972	lnc-CMC1-4:1	<i>RPL17</i>	0.993
lnc-PRAC-1:1	<i>RPL17</i>	0.972	lnc-MKLN1-3:1	<i>RPL17</i>	0.992
lnc-ACBD3-1:1	<i>RPL17</i>	0.972	lnc-CNKS3-2:1	<i>RPL17</i>	0.992
lnc-SLC14A1-1:1	<i>RSL24D1</i>	0.973	lnc-GPR63-8:1	<i>RSL24D1</i>	0.991
lnc-PRAC-1:1	<i>RSL24D1</i>	0.973	lnc-DLEU7-9:1	<i>RPL17</i>	0.991
lnc-RALGAPA2-7:1	<i>RSL24D1</i>	0.973	NONHSAT080449	<i>RPL17</i>	0.99
NONHSAT017634	<i>RPL17</i>	0.973	lnc-AVPR1A-3:1	<i>RPL17</i>	0.99
NR_104160	<i>CDADC1</i>	0.974	lnc-APOD-2:1	<i>RPL17</i>	0.99
NONHSAT059117	<i>RSL24D1</i>	0.974	lnc-PLEKHA5-6:1	<i>RPL17</i>	0.99
lnc-RRP1B-5:1	<i>RSL24D1</i>	0.974	lnc-NRN1-6:2	<i>RPL17</i>	0.99
lnc-BEND6-2:1	<i>RSL24D1</i>	0.975	lnc-SLC25A1-4:1	<i>RPL17</i>	0.99
NONHSAT062956	<i>RSL24D1</i>	0.975			

PCC, Pearson's correlation coefficient.

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**Table S6.** lncRNA-targeted miRNAs predicted by miRDB

lncRNA	Targeted miRNAs	Pairs
NR_104160	hsa-miR-101-3p; hsa-miR-3692-3p; hsa-miR-6515-3p; hsa-miR-186-3p; hsa-miR-196a-5p; hsa-miR-600	6
lnc-DAOA-10:1	hsa-miR-4443; hsa-miR-4291; hsa-miR-3166	3
lnc-DNAJC12-2:1	hsa-miR-7106-3p; hsa-miR-4529-5p; hsa-miR-5581-3p; hsa-miR-4724-5p; hsa-miR-6854-3p; hsa-miR-5196-3p; hsa-miR-3182; hsa-miR-3679-5p; hsa-miR-1185-5p; hsa-miR-3158-5p; hsa-miR-1255b-2-3p; hsa-miR-4730	12
lnc-SETD4-11:1	hsa-miR-4268; hsa-miR-548ba; hsa-miR-3173-5p	3
lnc-RPS13-3:1	hsa-miR-150-3p; hsa-miR-3173-5p; hsa-miR-362-3p; hsa-miR-329-3p; hsa-miR-6828-3p	5
lnc-TIPRL-1:1	hsa-miR-585-5p; hsa-miR-3170; hsa-miR-7850-5p; hsa-miR-4520-2-3p; hsa-miR-939-3p; hsa-miR-6866-5p	6
lnc-NAP1L3-8:1	hsa-miR-3622b-3p; hsa-miR-3622a-3p; hsa-miR-1299; hsa-miR-4303; hsa-miR-4279; hsa-miR-3925-3p; hsa-miR-4661-5p; hsa-miR-103a-2-5p; hsa-miR-3119	9
lnc-TMEM74-3:1	hsa-miR-6836-5p; hsa-miR-6132; hsa-miR-4283; hsa-miR-4317; hsa-miR-5002-3p; hsa-miR-24-3p; hsa-miR-873-5p; hsa-miR-138-1-3p; hsa-miR-627-3p	9
lnc-A2M-2:1	hsa-miR-5096; hsa-miR-582-3p; hsa-miR-4276; hsa-miR-636; hsa-miR-329-5p; hsa-miR-1206; hsa-miR-3910; hsa-miR-3074-5p; hsa-miR-4660; hsa-miR-7153-3p; hsa-miR-4530; hsa-miR-26b-3p	12
lnc-C19orf12-7:1	hsa-miR-1275; hsa-miR-93-3p; hsa-miR-4294; hsa-miR-3168; hsa-miR-6723-5p; hsa-miR-5706; hsa-miR-4782-5p; hsa-miR-671-5p; hsa-miR-4709-5p; hsa-miR-1227-5p; hsa-miR-4682	11
lnc-BEND6-2:1	hsa-miR-330-5p; hsa-miR-548ah-5p; hsa-miR-326; hsa-miR-3609; hsa-miR-939-3p; hsa-miR-3944-5p; hsa-miR-6797-3p; hsa-miR-6076; hsa-miR-766-3p; hsa-miR-4677-5p; hsa-miR-30e-3p; hsa-miR-30d-3p; hsa-miR-2116-5p; hsa-miR-3686; hsa-miR-3201	15
lnc-SPR-3:1	hsa-miR-6738-3p; hsa-miR-6887-3p; hsa-miR-6826-3p; hsa-miR-4267; hsa-miR-7977	5
lnc-CDH17-2:1	hsa-miR-3173-5p; hsa-miR-4511; hsa-miR-362-3p; hsa-miR-329-3p; hsa-miR-6828-3p; hsa-miR-504-5p; hsa-miR-4725-5p	7
lnc-ATG14-3:1	hsa-miR-8075; hsa-miR-4310; hsa-miR-3173-5p; hsa-miR-7157-5p; hsa-miR-6126; hsa-miR-1267	6
lnc-ACBD3-1:1	hsa-miR-1238-3p; hsa-miR-362-3p; hsa-miR-329-3p; hsa-miR-651-3p; hsa-miR-4639-3p	5
lnc-HPRT1-9:1	hsa-miR-873-5p; hsa-miR-4314; hsa-miR-3614-5p; hsa-miR-1227-3p; hsa-miR-4319; hsa-miR-103a-2-5p; hsa-miR-6885-3p; hsa-miR-4279; hsa-miR-661; hsa-miR-6736-3p; hsa-miR-4646-5p; hsa-miR-204-3p	12
lnc-DUSP19-3:1	hsa-miR-544a; hsa-miR-501-5p; hsa-miR-422a; hsa-miR-378i; hsa-miR-378h; hsa-miR-378f; hsa-miR-378e; hsa-miR-378d; hsa-miR-378c; hsa-miR-378b; hsa-miR-378a-3p	11
lnc-NR2F2-10:1	hsa-miR-520f-3p; hsa-miR-943; hsa-miR-4717-3p; hsa-miR-4534	4
lnc-SLC14A1-1:1	hsa-miR-431-5p; hsa-miR-6819-5p; hsa-miR-6812-5p; hsa-miR-6737-5p; hsa-miR-4441; hsa-miR-7108-3p; hsa-miR-1200; hsa-miR-1290; hsa-miR-4324; hsa-miR-504-3p	10
lnc-KBTBD13-1:1	hsa-miR-548ah-5p; hsa-miR-3609; hsa-miR-4677-5p; hsa-miR-4286; hsa-miR-2116-5p; hsa-miR-3686; hsa-miR-3942-3p	7
lnc-IQCH-4:1	hsa-miR-4420; hsa-miR-181b-3p; hsa-miR-181b-2-3p; hsa-miR-4532; hsa-miR-4297; hsa-miR-6873-3p; hsa-miR-6833-3p; hsa-miR-4768-5p; hsa-miR-4715-3p	9
lnc-STK35-3:1	hsa-miR-4306; hsa-miR-4644; hsa-miR-185-5p; hsa-miR-103a-2-5p	4
lnc-COMMD3-3:1	hsa-miR-764; hsa-miR-6766-5p; hsa-miR-6756-5p; hsa-miR-552-3p; hsa-miR-6891-3p; hsa-miR-6072	6
lnc-PLEKHM3-2:1	hsa-miR-4709-5p; hsa-miR-6867-5p; hsa-miR-505-3p; hsa-miR-1227-5p; hsa-miR-378g; hsa-miR-513b-3p; hsa-miR-3168	7
lnc-CHCHD3-1:1	hsa-miR-548b-3p; hsa-miR-4456; hsa-miR-6081; hsa-miR-4641; hsa-miR-4267; hsa-miR-571; hsa-miR-4777-5p; hsa-miR-6845-3p	8
lnc-ZNF286A-11:1	hsa-miR-1227-5p; hsa-miR-409-3p; hsa-miR-671-5p; hsa-miR-7107-5p; hsa-miR-1234-3p	5
lnc-SAMHD1-2:1	hsa-miR-578; hsa-miR-544a; hsa-miR-4423-5p; hsa-miR-744-3p; hsa-miR-7106-3p; hsa-miR-127-5p	6
lnc-C8G-2:1	hsa-miR-6830-5p; hsa-miR-3674; hsa-miR-4328; hsa-miR-5095; hsa-miR-6511a-5p; hsa-miR-1910-3p; hsa-miR-5003-5p; hsa-miR-874-5p; hsa-miR-4763-5p; hsa-miR-515-5p; hsa-miR-645; hsa-miR-208b-5p; hsa-miR-208a-5p; hsa-miR-125b-2-3p; hsa-miR-7160-5p; hsa-miR-4306	16
lnc-ARSB-1:1	hsa-miR-4529-5p; hsa-miR-3622b-5p; hsa-miR-4310; hsa-miR-4288; hsa-miR-346; hsa-miR-6740-3p; hsa-miR-3679-5p; hsa-miR-1185-5p; hsa-miR-659-3p; hsa-miR-6806-3p; hsa-miR-3928-5p	11
lnc-NT5E-4:1	hsa-miR-616-3p; hsa-miR-504-3p; hsa-miR-1279; hsa-miR-597-3p; hsa-miR-376a-2-5p	5
lnc-CES1-4:1	hsa-miR-4778-3p; hsa-miR-4717-3p; hsa-miR-4323; hsa-miR-4764-5p; hsa-miR-5196-3p; hsa-miR-152-3p; hsa-miR-148b-3p; hsa-miR-148a-3p	8
lnc-CAMSAP2-2:1	hsa-miR-3173-5p; hsa-miR-6753-5p; hsa-miR-939-3p; hsa-miR-4511; hsa-miR-6828-3p	5
lnc-GPR63-8:1	hsa-miR-4319; hsa-miR-298; hsa-miR-103a-2-5p; hsa-miR-1299; hsa-miR-4765; hsa-miR-4279; hsa-miR-647	7
lnc-CEBPZ-1:1	hsa-miR-3186-3p; hsa-miR-4520-3p; hsa-miR-549a	3
lnc-CCAR1-3:1	hsa-miR-4288; hsa-miR-1297; hsa-miR-346; hsa-miR-10b-3p; hsa-miR-591; hsa-miR-4465; hsa-miR-26b-5p; hsa-miR-26a-5p; hsa-miR-31-5p; hsa-miR-4529-5p; hsa-miR-544a; hsa-miR-632; hsa-miR-942-3p; hsa-miR-4299; hsa-miR-494-3p	15
lnc-WSB1-7:2	hsa-miR-4268; hsa-miR-6753-5p; hsa-miR-939-3p; hsa-miR-4511	4
lnc-PRRG4-4:1	hsa-miR-4443; hsa-miR-6753-5p; hsa-miR-4641	3
lnc-CENPJ-4:1	hsa-miR-3173-5p; hsa-miR-939-3p	2

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Inc-MBL2-9:1	hsa-miR-140-5p	1
Inc-LPAR4-1:1	hsa-miR-1283; hsa-miR-4709-5p; hsa-miR-3679-5p; hsa-miR-1185-5p; hsa-miR-1305; hsa-miR-4717-5p; hsa-miR-6768-3p; hsa-miR-330-3p; hsa-miR-4776-3p; hsa-miR-6878-3p; hsa-miR-1297; hsa-miR-3193	12
Inc-DTHD1-16:1	hsa-miR-4691-3p; hsa-miR-4520-3p; hsa-miR-892a	3
Inc-TRAF4-1:1	hsa-miR-630; hsa-miR-216a-5p; hsa-miR-4469	3
Inc-SAR1B-2:1	hsa-miR-6753-5p; hsa-miR-7850-5p	2
Inc-FAM9C-7:1	hsa-miR-7114-5p; hsa-miR-3944-5p; hsa-miR-452-3p; hsa-miR-4659b-3p; hsa-miR-4659a-3p; hsa-miR-3177-3p; hsa-miR-1537-5p; hsa-miR-4677-5p; hsa-miR-8066; hsa-miR-324-5p; hsa-miR-5704; hsa-miR-2116-5p	12
Inc-MDC1-1:1	hsa-miR-103a-2-5p; hsa-miR-326; hsa-miR-1283; hsa-miR-181a-2-3p; hsa-miR-6509-3p; hsa-miR-5590-5p; hsa-miR-3679-5p; hsa-miR-1185-5p; hsa-miR-3665; hsa-miR-335-3p; hsa-miR-5588-5p; hsa-miR-5001-3p; hsa-miR-4799-3p; hsa-miR-4720-5p; hsa-miR-4695-3p	15
Inc-CMC1-4:1	hsa-miR-4448; hsa-miR-6844; hsa-miR-5006-5p; hsa-miR-4261; hsa-miR-345-5p; hsa-miR-652-5p; hsa-miR-6828-3p; hsa-miR-6893-3p; hsa-miR-370-3p; hsa-miR-5003-5p; hsa-miR-140-5p	11
Inc-CNKS3R3-2:1	hsa-miR-5000-3p; hsa-miR-764; hsa-miR-4663; hsa-miR-4745-3p; hsa-miR-1538; hsa-miR-943; hsa-miR-4436a; hsa-miR-1226-5p	8
Inc-DLEU7-9:1	hsa-miR-548m; hsa-miR-23b-5p; hsa-miR-23a-5p; hsa-miR-939-3p; hsa-miR-6809-3p	5
Inc-APOD-2:1	hsa-miR-500b-3p; hsa-miR-6718-5p; hsa-miR-203b-5p; hsa-miR-6865-3p; hsa-miR-1909-5p; hsa-miR-550a-5p; hsa-miR-1271-3p; hsa-miR-5587-5p	8
Inc-NRN1-6:2	hsa-miR-3173-5p; hsa-miR-939-3p; hsa-miR-6753-5p; hsa-miR-4750-3p; hsa-miR-4256; hsa-miR-4511	6
Inc-SLC25A1-4:1	hsa-miR-4268; hsa-miR-657	2
Inc-ARHGEF4-1:5	hsa-miR-22-5p; hsa-miR-616-5p; hsa-miR-373-5p; hsa-miR-371b-5p; hsa-miR-6827-3p; hsa-miR-340-3p; hsa-miR-3613-3p; hsa-miR-5192; hsa-miR-6780a-5p; hsa-miR-6779-5p; hsa-miR-3689c; hsa-miR-3689b-3p; hsa-miR-3689a-3p; hsa-miR-30b-3p; hsa-miR-1273h-5p	15
Inc-ERMN-2:1	hsa-miR-568; hsa-miR-3065-5p; hsa-miR-7159-5p; hsa-miR-7152-5p; hsa-miR-6728-3p; hsa-miR-8069; hsa-miR-6822-3p; hsa-miR-511-3p; hsa-miR-760; hsa-miR-7844-5p; hsa-miR-6760-5p; hsa-miR-708-3p; hsa-miR-6817-3P; hsa-miR-587; hsa-miR-149-5p; hsa-let-7g-3p; hsa-let-7a-2-3p; hsa-miR-1972; hsa-miR-4735-5p	19
ENST00000606596	hsa-miR-1908-5p; hsa-miR-1225-3p; hsa-miR-369-3p; hsa-miR-4492	4
ENST00000607698	hsa-miR-324-5p; hsa-miR-2355-5p; hsa-miR-6077; hsa-miR-548ax; hsa-miR-548ao-5p	5
ENST00000565797	hsa-miR-3119; hsa-miR-6774-5p; hsa-miR-635; hsa-miR-3130-3p; hsa-miR-1225-3p; hsa-miR-650; hsa-miR-3612; hsa-miR-6883-5p; hsa-miR-1251-3p	9
Inc-CAPN2-3:1	hsa-miR-4643; hsa-miR-4308; hsa-miR-518c-5p; hsa-miR-4254; hsa-miR-548al; hsa-miR-4445-3p; hsa-miR-4297; hsa-miR-1193; hsa-miR-4293; hsa-miR-1281	10
Inc-LONP2-8:1	hsa-miR-4426; hsa-miR-6894-3p; hsa-miR-6773-5p; hsa-miR-6724-5p; hsa-miR-6853-3p; hsa-miR-4662b; hsa-miR-4647; hsa-miR-561-3p; hsa-miR-3977; hsa-miR-4528; hsa-miR-7155-3p; hsa-miR-3136-3p; hsa-miR-4330	13
Inc-AVPR1A-3:1	hsa-miR-296-5p; hsa-miR-4426; hsa-miR-4763-5p; hsa-miR-335-3p; hsa-miR-561-3p; hsa-miR-4774-5p; hsa-miR-4662b; hsa-miR-4647	8
Inc-MEX3C-6:1	hsa-miR-4283; hsa-miR-1204; hsa-miR-1273h-3p; hsa-miR-6894-3p; hsa-miR-4774-5p; hsa-miR-6853-3p; hsa-miR-561-3p; hsa-miR-24-3p; hsa-miR-3166; hsa-miR-296-5p; hsa-miR-1304-3p; hsa-miR-4440; hsa-miR-6515-3p	13
Inc-CCL5-3:1	hsa-miR-4426; hsa-miR-449c-3p; hsa-miR-7155-5p; hsa-miR-4754; hsa-miR-6894-3p; hsa-miR-7159-3p; hsa-miR-6515-3p; hsa-miR-4283; hsa-miR-5787; hsa-miR-4505; hsa-miR-4662b; hsa-miR-4647; hsa-miR-3200-3p	13
Inc-FBXL2-2:1	hsa-miR-4279; hsa-miR-3120-3p; hsa-miR-3614-5p; hsa-miR-4435; hsa-miR-6772-3p; hsa-miR-1273h-3p	6

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**Table S7-1.** lncRNA-miRNA-mRNA ceRNA pathways in the lncRNA-miRNA-mRNA network

N	lncRNA	miRNA	mRNA	N	lncRNA	miRNA	mRNA
1	ENST00000565797	hsa-miR-3119	SNRPF	37	lnc-FAM9C-7:1	hsa-miR-4659a-3p	RSL24D1
2	ENST00000565797	hsa-miR-1251-3p	SNRPF	38	lnc-GPR63-8:1	hsa-miR-103a-2-5p	RSL24D1
3	ENST00000565797	hsa-miR-1251-3p	CD69	39	lnc-HPRT1-9:1	hsa-miR-3614-5p	RSL24D1
4	ENST00000565797	hsa-miR-6774-5p	CD69	40	lnc-HPRT1-9:1	hsa-miR-103a-2-5p	RSL24D1
5	ENST00000565797	hsa-miR-635	CD69	41	lnc-IQCH-4:1	hsa-miR-6833-3p	RSL24D1
6	ENST00000606596	hsa-miR-1225-3p	C12orf79	42	lnc-IQCH-4:1	hsa-miR-4768-5p	RSL24D1
7	lnc-ARSB-1:1	hsa-miR-4310	RSL24D1	43	lnc-LONP2-8:1	hsa-miR-4426	RSL24D1
8	lnc-AVPR1A-3:1	hsa-miR-4426	RSL24D1	44	lnc-LONP2-8:1	hsa-miR-4662b	RSL24D1
9	lnc-AVPR1A-3:1	hsa-miR-335-3p	RSL24D1	45	lnc-LONP2-8:1	hsa-miR-4647	RSL24D1
10	lnc-AVPR1A-3:1	hsa-miR-335-3p	PTS	46	lnc-LONP2-8:1	hsa-miR-4528	RSL24D1
11	lnc-AVPR1A-3:1	hsa-miR-4774-5p	RSL24D1	47	lnc-LPAR4-1:1	hsa-miR-1305	RSL24D1
12	lnc-AVPR1A-3:1	hsa-miR-4662b	RSL24D1	48	lnc-LPAR4-1:1	hsa-miR-4776-3p	RSL24D1
13	lnc-AVPR1A-3:1	hsa-miR-4647	RSL24D1	49	lnc-MDC1-1:1	hsa-miR-103a-2-5p	RSL24D1
14	lnc-C19orf12-7:1	hsa-miR-3168	RPL17	50	lnc-MDC1-1:1	hsa-miR-5590-5p	RSL24D1
15	lnc-C8G-2:1	hsa-miR-515-5p	RSL24D1	51	lnc-MDC1-1:1	hsa-miR-335-3p	RSL24D1
16	lnc-C8G-2:1	hsa-miR-208b-5p	RSL24D1	52	lnc-MEX3C-6:1	hsa-miR-4774-5p	RSL24D1
17	lnc-C8G-2:1	hsa-miR-208a-5p	RSL24D1	53	lnc-NR2F2-10:1	hsa-miR-4717-3p	RSL24D1
18	lnc-C8G-2:1	hsa-miR-125b-2-3p	RSL24D1	54	lnc-NT5E-4:1	hsa-miR-616-3p	RSL24D1
19	lnc-CAPN2-3:1	hsa-miR-4297	PTS	55	lnc-NT5E-4:1	hsa-miR-376a-2-5p	RSL24D1
20	lnc-CCAR1-3:1	hsa-miR-494-3p	RSL24D1	56	lnc-PLEKHM3-2:1	hsa-miR-3168	RPL17
21	lnc-CCL5-3:1	hsa-miR-4426	RSL24D1	57	lnc-PLEKHM3-2:1	hsa-miR-378g	RSL24D1
22	lnc-CCL5-3:1	hsa-miR-4662b	RSL24D1	58	lnc-PLEKHM3-2:1	hsa-miR-513b-3p	RSL24D1
23	lnc-CCL5-3:1	hsa-miR-4647	RSL24D1	59	lnc-SPR-3:1	hsa-miR-6826-3p	RSL24D1
24	lnc-CES1-4:1	hsa-miR-4778-3p	RSL24D1	60	lnc-TIPRL-1:1	hsa-miR-585-5p	RSL24D1
25	lnc-CES1-4:1	hsa-miR-4717-3p	RSL24D1	61	lnc-TMEM74-3:1	hsa-miR-6836-5p	RPL17
26	lnc-CES1-4:1	hsa-miR-4764-5p	RSL24D1	62	lnc-TMEM74-3:1	hsa-miR-6132	RPL17
27	lnc-CHCHD3-1:1	hsa-miR-4777-5p	RSL24D1	63	lnc-TMEM74-3:1	hsa-miR-627-3p	RPL17
28	lnc-COMMD3-3:1	hsa-miR-6891-3p	RPL17	64	lnc-TRAF4-1:1	hsa-miR-630	RSL24D1
29	lnc-COMMD3-3:1	hsa-miR-6072	RPL17	65	NR_104160	hsa-miR-6515-3p	CDADC1
30	lnc-DLEU7-9:1	hsa-miR-6809-3p	RSL24D1	66	NR_104160	hsa-miR-6515-3p	SNRPF
31	lnc-ERMN-2:1	hsa-miR-7152-5p	ERMN	67	NR_104160	hsa-miR-186-3p	GZF1
31	lnc-ERMN-2:1	hsa-miR-511-3p	PTS	68	NR_104160	hsa-miR-186-3p	CDADC1
33	lnc-ERMN-2:1	hsa-miR-587	PTS	69	NR_104160	hsa-miR-600	SNRPF
34	lnc-ERMN-2:1	hsa-miR-4735-5p	PTS	70	NR_104160	hsa-miR-600	CDADC1
35	lnc-ERMN-2:1	hsa-let-7g-3p	CDADC1	71	NR_104160	hsa-miR-101-3p	ID1
36	lnc-FAM9C-7:1	hsa-miR-4659b-3p	RSL24D1				

**Table S7-2.** RNAs and the number of their associated ceRNA pathways

lncRNA	Pathways	miRNA	Pathways	mRNA	Pathways
NR_104160	7	hsa-miR-4426	3	RSL24D1	44
lnc-AVPR1A-3:1	6	hsa-miR-335-3p	3	RPL17	7
ENST00000565797	5	hsa-miR-4662b	3	PTS	5
lnc-ERMN-2:1	5	hsa-miR-4647	3	CDADC1	4
lnc-LONP2-8:1	4	hsa-miR-103a-2-5p	3	SNRPF	4
lnc-C8G-2:1	4	hsa-miR-1251-3p	2	CD69	3
lnc-CCL5-3:1	3	hsa-miR-4774-5p	2	C12orf79	1
lnc-CES1-4:1	3	hsa-miR-3168	2	ID1	1

## lncRNA-NR\_104160 and ceRNA network for hypertension

lnc-MDC1-1:1	3	hsa-miR-4717-3p	2	<i>ERMN</i>	1
lnc-PLEKHM3-2:1	3	hsa-miR-6515-3p	2	<i>GZF1</i>	1
lnc-TMEM74-3:1	3	hsa-miR-186-3p	2		
lnc-COMMD3-3:1	2	hsa-miR-600	2		
lnc-FAM9C-7:1	2	hsa-miR-6774-5p	1		
lnc-HPRT1-9:1	2	hsa-miR-635	1		
lnc-IQCH-4:1	2	hsa-miR-1225-3p	1		
lnc-LPAR4-1:1	2	hsa-miR-4310	1		
lnc-NT5E-4:1	2	hsa-miR-3119	1		
ENST00000606596	1	hsa-miR-515-5p	1		
lnc-ARSB-1:1	1	hsa-miR-208b-5p	1		
lnc-C19orf12-7:1	1	hsa-miR-208a-5p	1		
lnc-CAPN2-3:1	1	hsa-miR-125b-2-3p	1		
lnc-CCAR1-3:1	1	hsa-miR-4297	1		
lnc-CHCHD3-1:1	1	hsa-miR-494-3p	1		
lnc-DLEU7-9:1	1	hsa-miR-4778-3p	1		
lnc-GPR63-8:1	1	hsa-miR-4764-5p	1		
lnc-MEX3C-6:1	1	hsa-miR-4777-5p	1		
lnc-NR2F2-10:1	1	hsa-miR-6891-3p	1		
lnc-SPR-3:1	1	hsa-miR-6072	1		
lnc-TIPRL-1:1	1	hsa-miR-6809-3p	1		
lnc-TRAF4-1:1	1	hsa-miR-7152-5p	1		
		hsa-miR-511-3p	1		
		hsa-miR-587	1		
		hsa-miR-4735-5p	1		
		hsa-let-7g-3p	1		
		hsa-miR-4659b-3p	1		
		hsa-miR-4659a-3p	1		
		hsa-miR-3614-5p	1		
		hsa-miR-6833-3p	1		
		hsa-miR-4768-5p	1		
		hsa-miR-4528	1		
		hsa-miR-1305	1		
		hsa-miR-4776-3p	1		
		hsa-miR-5590-5p	1		
		hsa-miR-616-3p	1		
		hsa-miR-376a-2-5p	1		
		hsa-miR-378g	1		
		hsa-miR-513b-3p	1		
		hsa-miR-6826-3p	1		
		hsa-miR-585-5p	1		
		hsa-miR-6836-5p	1		
		hsa-miR-6132	1		
		hsa-miR-627-3p	1		
		hsa-miR-630	1		
		hsa-miR-101-3p	1		