

## SUPPLEMENTARY TABLES

**Supplementary Table 1. Paired treatment naïve LN and PB CLL samples: patient characteristics.**

		Number	%
<b>Median age (years)</b>		56 (33-73)	
<b>Gender</b>			
	Male	10	59
	Female	7	41
<b>Rai Stage</b>			
	I	5	29.5
	II	4	23.5
	III	4	23.5
	IV	4	23.5
<b>IGHV mutation</b>			
	Un-mutated	10	59
	Mutated	4	23.5
<b>ZAP70</b>			
	+	9	53
	-	7	41
<b>CD38</b>			
	+	9	53
	-	8	47
<b>17p deletion</b>			
	Deleted	5	29.5
	Non deleted	12	70.5

**Supplementary Table 2. Clinical and laboratory parameters of CLL patients treated with ibrutinib.**

		<b>Number</b>	<b>%</b>
<b>Median age (years)</b>		66 (33 - 85)	
<b>Gender</b>			
Male		22	55
Female		18	45
<b>Median absolute lymphocytic count ALC (x10<sup>9</sup>/L)</b>			
ALC pre-treatment (min. - max.)		75.6 (0.5 - 402)	
ALC 28 days post-treatment (min. - max.)		87 (2.6 - 475.3)	
<b>Rai Stage</b>			
I		8	20
II		3	7.5
III		5	12.5
IV		24	60
<b>IGHV mutation</b>			
Unmutated		23	57.5
Mutated		17	42.5
<b>ZAP70</b>			
+		12	30
-		15	37.5
<b>CD38</b>			
+		15	37.5
-		25	62.5
<b>17p deletion</b>			
Deleted		23	57.5
Non deleted		17	42.5
<b>Nodal response to ibrutinib</b>			
Responders		25	62.5
Non-responders		13	32.5

**Supplementary Table 3. RT and PCR primers.**

Primer	Sequence
let-7a_F	5'- GGG CCT GAG GTA GTA GGT TGT ATA GTT -3'
let-7a-3p_F	5'- GGG GCC CCT ATA CAA TCT ACT GTC TTT C -3'
let-7g_F	5'- GGC CGT GAG GTA GTA GTT TGT ACA GTT -3'
miR-9_F	5'- GGG CCT CTT TGG TTA TCT AGC TGT ATG A -3'
miR-15a_F	5'- GTG CCT AGC AGC ACA TAA TGG TTT GTG -3'
miR-16_F	5'- GTG CCT AGC AGC ACG TAA ATA TTG GCG -3'
miR-17_F	5'- GGC AAA GTG CTT ACA GTG CAG GTA G -3'
miR-20a_F	5'- GGG CCT AAA GTG CTT ATA GTG CAG GTA G -3'
miR-21_F	5'- GGG CCC TAG CTT ATC AGA CTG ATG TTG A -3'
miR-22_F	5'- GGC AGT TCT TCAGTGGCAAGC TTT A-3'
miR-29a_F	5'- CCC GGA CTG ATT TCT TTT GGT GTT CAG -3'
miR-29c-3p_F	5'- GGC CTA GCA CCA TTT GAA ATC GGT TA -3'
miR-29c_F	5'- GGT GAC CGA TTT CTC CTG GTG TTC -3'
miR-34a_F	5'- GGT GGC AGT GTC TTA GCT GGT TGT -3'
miR-106b_F	5'- GTG CCT AAA GTG CTG ACA GTG CAG AT -3'
miR-146b-F	5'- GGC CTGAGAACTGAATTCCATAGGCT-3'
miR-150_F	5'- GTC TCC CAA CCC TTG TAC CAG TG -3'
miR-155_F	5'- GGG CCT TAA TGC TAA TCG TGA TAG GGG T -3'
miR-181b_F	5'- TGA CGA ACA TTC ATT GCT GTC GG -3'
miR-185_F	5'- GCT GGA GAG AAA GGC AGT TCC TGA -3'
miR-223_F	5'- GTG CCC GTG TAT TTG ACA AGC TGA GTT -3'
miR-650_F	5'- AGG AGG CAG CGC TCT CAG GAC -3'
RNU44_F	5' - GGC AAA TGC TGA CTG AAC ATG AAG GTC - 3'
Reverse transcription anchor primer	5'- CGA CTC GAT CCA GTC TCA GGG TCC GAG GTA TTC GAT CCT AAC CCT CTC CTC GGT ATC GAG TCG CAC TTT TTT TTT TTT V - 3'
PCR reverse primer	5' - GTC CGA GGT ATT CGA TCC TAA C - 3'
TaqMan probe	5' - 6-FAM/ TCT CCT CGG /ZEN/ TAT CGA GTC GCA CT /IABKFQ/ - 3'

**Supplementary Table 4: Gene list of the nanoString nCounter assay.**

AKT1	DNMT3A	LDOC1	RUNX1	TP53
ARID1A	E2F1	MAP2K1	RUNX3	WT1
ARID1B	FBXW7	MRE11A	SAMHD1	XPO1
ARID2	FOXP1	NFKBIA	SF3B1	GUSB
ASXL1	GPS2	NOTCH1	SIRT1	HPRT1
ATM	HDAC1	PDCD1	SMAD4	TBP
BAX	IBTK	PHLPP	SOCS1	
CDKN1A	IDH1	PTEN	TCF21	
CYLD	IKBKB	RARRES3	TET2	
DAPK1	KLF4	RB1	TNFAIP3	

**Supplementary Table 5. Differential miRNA expression in circulating CLL cells from 38 patients compared to circulating B cells from 33 healthy controls.**

miRNA	Median (IQR) in CTRL	Median (IQR) in CLL	Fold	P value	Trend in CLL
<b>Let-7a</b>	129.6 (77.8-237)	767.6 (397-1080)	6	< 0.0001	Up
<b>Let7a-3p</b>	0.1 (0.04-0.2)	0.76 (0.3-2.4)	7	< 0.0001	Up
<b>Let-7g</b>	29.8 (18.8-50.7)	185.6 (60-342)	6	< 0.0001	Up
<b>miR-9</b>	0.02 (0.01-0.1)	0.13 (0.02-0.9)	6.5	0.002	Up
<b>miR-15a</b>	39.4 (13.7-196)	869.7 (146.6-8131)	22	< 0.0001	Up
<b>miR-16</b>	66 (33.6-195.7)	1303 (453.7-2239)	19	< 0.0001	Up
<b>miR-17</b>	0.6 (0.3-2)	6 (1.8-12.5)	10	< 0.0001	Up
<b>miR-20a</b>	2.2 (1.2-8)	15.5 (5.8-34.3)	8	< 0.0001	Up
<b>miR-21</b>	15.05 (6.8-30.2)	505.2 (202-655)	32	< 0.0001	Up
<b>miR-22</b>	0.02 (0.02-0.06)	0.2 (0.02-0.4)	15	0.0009	Up
<b>miR-29a</b>	0.1 (0.02-0.5)	1.6 (0.4-3.7)	16	< 0.0001	Up
<b>miR-29c-3p</b>	45.6 (26-83.5)	681 (313-1126)	15	< 0.0001	Up
<b>miR-29c</b>	0.1 (0.06-0.3)	0.8 (0.3-1.4)	8	< 0.0001	Up
<b>miR-34a</b>	0.5 (0.1-1.02)	43.7 (12.4-174)	87	< 0.0001	Up
<b>miR-106b</b>	1.3 (0.3-5.4)	9 (1.8-39)	7	0.0007	Up
<b>miR-146b</b>	14.4 (4.8-44.7)	127.4 (23-586.3)	9	< 0.0001	Up
<b>miR-150</b>	6.9 (3.9-21)	192 (30.6-336)	30	< 0.0001	Up
<b>miR-155</b>	10.6 (3.5-23.8)	307.4 (153.6-698)	29	< 0.0001	Up
<b>miR-181b</b>	0.66 (0.5-1.1)	0.24 (0.07-0.85)	-2.5	0.01	Down

<b>miR-185</b>	0.7 (0.2-1.5)	2.4 (1-3.7)	3	< 0.0001	Up
<b>miR-223</b>	0.03 (0.01-0.16)	0.17 (0.07-0.4)	5	0.0017	Up
<b>miR-650</b>	0.001 (0.0005-0.01)	0.085 (0.008-0.3)	85	< 0.0001	Up

Twenty-two miRNAs were assayed by TaqMan qPCR and with standard curves in PB CLL cells and circulating B-cells from healthy controls. The median and interquartile ranges of the miRNA quantity (fmoles/ng of total RNA) are shown. CTRL, healthy control; IQR, interquartile range. *P* values calculated by Student's T-test.

**Supplementary Table 6. Differential miRNA expression in PB CLL cells from 38 patients pre- and post-treatment (day 28) with ibrutinib.**

miRNA	Median in CLL patient samples pre-ibrutinib treatment (IQR) fmoles/ng total RNA	Median in CLL patient samples post-ibrutinib treatment (IQR) fmoles/ng total RNA	Fold change	p-value Wilcoxon n	Trend
<b>Let7a</b>	805.2 (421-1079.7)	782 (446.5-1090)	1	0.9	-
<b>Let7a-3p</b>	0.78 (0.36-2.4)	0.6 (0.3-2.4)	-1.3	0.25	-
<b>Let7g</b>	204.7 (76-342)	156.3 (94-321.2)	-1.3	0.5	-
<b>miR-9</b>	0.15 (0.04-0.9)	0.06 (0.01-0.3)	-2.5	0.2	-
<b>miR-15a</b>	1079 (281-8131)	550.4 (152.5-3033)	-2	0.08	-
<b>miR-16</b>	1315 (501.3-2239)	1279.8 (594-2219.3)	1	0.8	-
<b>miR-17</b>	6.4 (2.4-12.5)	5.6 (2.3-12.8)	-1.1	0.9	-
<b>miR-20a</b>	17.2 (7.3-34.3)	19.5 (10-37)	1.1	0.5	-
<b>miR-21</b>	530.9 (240.9-655)	324 (131.7-576)	-1.6	0.3	-
<b>miR-22</b>	0.2 (0.04-0.4)	0.06 (0.02-0.15)	-3.33	0.0008	Down
<b>miR-29a</b>	1.7 (0.7-3.7)	2 (0.6-5.5)	1.17	0.3	-
<b>miR-29c-3p</b>	688.7 (351.5-1126)	817.2 (291.1-1586.5)	1.18	0.59	-
<b>miR-29c</b>	0.8 (0.4-1.4)	0.7 (0.3-1.2)	-1.1	0.78	-
<b>miR-34a</b>	49.3 (14.6-174)	8.9 (3.4-45.3)	-5.56	0.01	Down
<b>miR-106b</b>	13 (2.1-39)	9.2 (3.7-22.2)	-1.4	0.67	-
<b>miR-146b</b>	151.5 (68.3-687.8)	27.6 (7.8-81.2)	-5.56	< 0.0001	Down
<b>miR-150</b>	200.3 (68.3-336)	227.4 (99-495)	1.1	0.45	-
<b>miR-155</b>	343.9 (163.6-698)	239 (122.5-469)	-1.4	0.08	-

<b>miR-181b</b>	0.26 (0.078-0.85)	0.1 (0.02-0.24)	-2.6	0.019	Down
<b>miR-185</b>	2.6 (1.3-3.7)	1.8 (0.9-2.6)	-1.4	0.079	-
<b>miR-223</b>	0.19 (0.1-0.4)	0.15 (0.05-0.3)	-1.3	0.1	-
<b>miR-650</b>	0.1 (0.009-0.3)	0.06 (0.006-0.2)	-1.4	0.9	-

The miRNA level was quantified by TaqMan qPCR and with standard curves. The median and interquartile range of the miRNA quantity (fmoles/ng of total RNA) are shown. Pre-tx, pre-treatment; post-tx, post-treatment; IQR, interquartile range. *P* values calculated by Wilcoxon paired test.

**Supplementary Table 7. Differential miRNA expression in PB CLL cells from 38 patients pre- and post-treatment (day 28) with ibrutinib: Responders and Non-Responders.**

miRNA	Responders (N=24)	Non-Responders (N=12)	Responders Pre-treatment compared to post-ibrutinib p-value	Non- Responders Pre-treatment compared to post-ibrutinib p-value	Responder Pre-treatment compared to Non-Responder Pre-treatment p-value
	Median change in CLL cells post- ibrutinib treatment compared to pre- treatment (IQR)	Median change in CLL cells post- ibrutinib treatment compared to pre- treatment (IQR)			
	Percent Change	Percent Change			
<b>Let7a</b>	4.36% (-38.11%-136.23%)	-14.4% (-54.4%-44.43%)	0.65	0.53	0.73
<b>Let7a-3p</b>	-0.18% (-63.17%-67.95%)	-13.3% (-84.8%-71.4%)	0.58	0.27	0.63
<b>Let7g</b>	-34.62% (-60.35% - 149.8%)	0.63% (-43.42%-185.9%)	0.3	0.8	0.89
<b>miR-9</b>	-60.24% (-92.25% -160.5%)	-88.82% (-94.1% - 29.6%)	0.58	0.11	0.48
<b>miR-15a</b>	-57.41% (-92.34%-58.30%)	6.13% (-47.15%-777.0%)	0.08	0.94	0.39
<b>miR-16</b>	10.64% (-60.89%-588.72%)	-30.66% (-59.66%-33.57%)	0.84	0.2	0.65
<b>miR-17</b>	4.29% (-66.41%-209.4%)	-31.75% (-59.16 – 151.5%)	0.88	0.70	0.90
<b>miR-20a</b>	55.89% (-58.59% - 437.4%)	50.77% (-38.64% - 77.88%)	0.66	0.5	0.78
<b>miR-21</b>	-13.11% (-70.92% - 98.68%)	-66.26% (-76.46 – 9.94%)	0.90	0.16	0.54
<b>miR-22</b>	-59.44% (-92.1% - 25.3%)	-39.09% (-92.89% - -4.27%)	<b>0.02*</b>	<b>0.015*</b>	0.20
<b>miR-29a</b>	151.1% (-66.67% - 558.2%)	-4.29% (-65.52% - 123.7%)	0.4	0.3	0.95
<b>miR-29c-3p</b>	36.94% (-43.49% -113.29%)	-6.08% (-54.51%-149.11%)	0.22	0.8	0.90
<b>miR-29c</b>	23.30% (-57.91% - 98.1%)	17.5% (-46.23 – 77.1%)	0.95	0.70	0.76
<b>miR-34a</b>	-70.17% (-90.36% - 31.22%)	-71.82% (-93.71% - 11.23%)	<b>0.007**</b>	0.13	0.88
<b>miR-106b</b>	76.46%	-24.47%	0.50	0.80	0.17

	(-85.90% - 1140.9%)	(-90.03 – 219.6%)			
<b>miR-146b</b>	-87.49% (-925.39 – -58.81%)	-87.8% (-93.65 - -75.1%)	<b>0.001***</b>	<b>0.02*</b>	0.41
<b>miR-150</b>	85.8% (-56.26% - 379.4%)	2.37% (-40.78% - 165.0%)	0.25	1	0.22
<b>miR-155</b>	-14.10% (-64.37% - 137.15%)	-53.79% (-74.38% - -36.9%)	0.60	<b>0.028*</b>	0.70
<b>miR-181b</b>	-76.19% (-96.27% - 10.94%)	5.34% (-92.64% - 79.02%)	<b>0.024*</b>	0.48	0.72
<b>miR-185</b>	-39.68% (-77.86% - 8.39%)	-47.13% (-60.1% - 106.6%)	<b>0.049*</b>	0.81	0.87
<b>miR-223</b>	-51.1% (-88.00% - 191.4%)	-56.79% (-76.84% - -21.7%)	0.53	0.08	<b>0.007**</b>
<b>miR-650</b>	-19.15% (-96.77% - 2647.51%)	-4.29% (-65.52% - 123.7%)	0.98	0.9	0.67

The miRNA level was quantified by TaqMan qPCR and with standard curves. The median and interquartile range of the miRNA percentage change from pre-treatment to post-treatment are shown. Pre-tx, pre-treatment; post-tx, post-treatment; IQR, interquartile range. *P* values calculated by Wilcoxon paired test. Significant p-values are bolded and marked with asterisks (\* p-value < 0.05 \*\* p-value < 0.01 \*\*\* p-value < 0.001)

**Supplementary Table 8. Differential miRNA expression in PB CLL cells from 38 patients pre- and post-treatment (day 28) with ibrutinib: Absolute values of miRNA in CLLs that were found to be significant for comparisons in responders and non-responders.**

miRNA	Responders		Non-Responders			
	Median in CLL patient samples pre-ibrutinib treatment (IQR) fmoles/ng total RNA	Median in CLL patient samples post-ibrutinib treatment (IQR) fmoles/ng total RNA	P Value	Median in CLL patient samples pre-ibrutinib treatment (IQR) fmoles/ng total RNA	Median in CLL patient samples post-ibrutinib treatment (IQR) fmoles/ng total RNA	P Value
miR-22	0.119 (0.03 – 0.42)	0.045 (0.013 – 0.15)	0.02	0.282 (0.15 – 0.49)	0.115 (0.018 – 0.18)	0.015
miR-34a	48.64 (13.1 – 175.65)	9.00 (3.70 – 32.75)	0.007	52.17 (22.74 – 92.26)	7.54 (3.41 – 45.32)	0.13
miR-146b	202.65 (45.71 - 725.67)	41.32 (10.95 - 95.78)	0.001	127.41 (80.79 – 203.58)	12.53 (7.32 – 31.1)	0.02
miR-155	265.95 (163.6 – 846.9)	310.79 (152.3 – 496.5)	0.6	402.00 (204.4 – 652.9)	198.93 (97.53 – 445.25)	0.028
miR-181b	0.289 (0.09 – 1.49)	0.09 (0.02 – 0.26)	0.024	0.21 (0.09 – 0.75)	0.1 (0.03 – 0.18)	0.48
miR-185	3.28 (0.89 - 3.86)	1.88 (0.66 – 2.60)	0.049	2.45 (1.69 – 3.00)	1.65 (0.97 – 3.79)	0.81
miR-223	0.14 (0.07 – 0.28)	0.14 (0.034 – 0.29)	0.53	0.42 (0.31-1.74)	0.22 (0.91 – 0.56)	0.08

The miRNA level was quantified by TaqMan qPCR in duplicate and with standard curves. The median and interquartile range of the miRNA quantity (fmoles/ng of total RNA) are shown. Pre-tx, pre-treatment; post-tx, post-treatment; IQR, interquartile range.

**Supplementary Table 9. mRNA counts of putative tumor suppressor targets of miR-22, miR-34a, miR-146b, and miR-181b in PB CLL cells from 38 patients pre- and post-treatment (day 28) with ibrutinib.**

mRNA	Avg. (SD) pre-tx	Avg. (SD) post-tx	Fold change	P value	Trend in post-tx
<b>ARID1B</b>	578 ( $\pm 325$ )	787 ( $\pm 377$ )	1.4	0.011	Up
<b>ARID2</b>	490 ( $\pm 260$ )	635 ( $\pm 275$ )	1.3	0.027	Up
<b>ATM</b>	274 ( $\pm 130$ )	332 ( $\pm 209$ )	1.2	0.046	Up
<b>CYLD</b>	2132 ( $\pm 1124$ )	3014 ( $\pm 1298$ )	1.4	0.033	Up
<b>FOXP1</b>	1001 ( $\pm 950$ )	1561 ( $\pm 1304$ )	1.6	0.023	Up
<b>HDAC1</b>	740 ( $\pm 303$ )	981 ( $\pm 410$ )	1.3	0.012	Up
<b>IBTK</b>	141 ( $\pm 159$ )	172 ( $\pm 133$ )	1.2	0.04	Up
<b>PTEN</b>	1090 ( $\pm 406$ )	1272 ( $\pm 668$ )	1.2	0.05	Up
<b>SMAD4</b>	834 ( $\pm 402$ )	1159 ( $\pm 588$ )	1.4	0.003	Up

mRNA counts were obtained from nanoString nCounter assay. Avg., average; SD, standard deviation; pre-tx, pre-treatment; post-tx, post-treatment. *P* values calculated by Wilcoxon paired test.

**Supplemental Table 10. Differential gene expression between CLL cells in PB and in LN.**

mRNA	Median (IQR) in PB	Median (IQR) in LN	Fold change	P value	Trend
<b>ARID1B</b>	7 (4.6-8.3)	5 (3.3-6.2)	-1.4	0.01	Down
<b>ARID2</b>	20.9 (18.4-27.3)	22.8 (15.6-32.3)	1.1	0.36	-
<b>ATM</b>	31.1 (25.6-36.3)	45.2 (19.9-56.7)	1.3	0.11	-
<b>CYLD</b>	2.5 (1.7-3.5)	1.6 (1.3-2)	-1.6	0.04	Down
<b>FOXP1</b>	68.6 (52-72)	49.4 (35-74)	-1.4	0.006	Down
<b>HDAC1</b>	68.6 (52-72)	49.4 (35-74)	-1.4	0.4	-
<b>IBTK</b>	7 (4.6-8.3)	5 (3.3-6.2)	-1.4	0.06	-
<b>PTEN</b>	2.5 (1.7-3.5)	1.6 (1.3-2)	-1.5	0.22	-
<b>SMAD4</b>	68.6 (52-72)	49.4 (35-74)	-1.4	0.33	-

Gene expression was determined by TaqMan qPCR performed in triplicate. RNA from 17 paired and treatment naive PB and LN CLL cells was used. The median and interquartile range of the relative mRNA expression are shown. P values calculated by Wilcoxon paired test.

**Supplementary Table 11. Differential gene expression in cell lines treated with Ibrutinib**

mRNA	Cell Line	Percent Change Untreated to Treated	P value
<b>ARID1b</b>	MEC1	56.69%	0.0005
<b>ARID2</b>	MEC1	35.21%	0.0028
<b>ATM</b>	MEC1	35.48%	0.039
<b>CYLD</b>	MEC1	19.49%	0.0014
<b>FOXP1</b>	MEC1	199.22%	0.0055
<b>HDAC1</b>	MEC1	65.82%	0.033
<b>IBTK</b>	MEC1	45.42%	0.0037
<b>PTEN</b>	MEC1	20.57%	0.066
<b>SMAD4</b>	MEC1	12.89%	0.0038
<b>ARID1b</b>	SP53	175.97%	0.001
<b>ARID2</b>	SP53	155.97%	0.0009
<b>ATM</b>	SP53	290.76%	0.0005
<b>CYLD</b>	SP53	65.35%	0.01
<b>FOXP1</b>	SP53	164.80%	0.0002
<b>HDAC1</b>	SP53	76.92%	0.002
<b>IBTK</b>	SP53	157.62%	0.006
<b>PTEN</b>	SP53	180.27%	0.0001
<b>SMAD4</b>	SP53	118.89%	0.002

Gene expression was determined by TaqMan qPCR performed in triplicate. RNA from MEC1 and SP53 cell lines treated with of Ibrutinib for 48 hours was compared to untreated cells. Percent change in expression levels is shown from untreated cells to treated cells with the associated p-value. P values calculated by Student's t-test.

**Supplementary Table 12. Differential gene expression in cell lines transfected with pre-miRNA and anti-miRNA constructs.**

Pre-miR				Anti-miR							
mRNA	Transfection	MEC-1		SP53		Transfection	MEC-1		SP53		
		% Change	p-value	% Change	p-value		% Change	p-value	% Change	p-value	
<b>ARID1b</b>	pre-miR 34a	-23.67%	0.007	-49.02%	0.014	anti-miR 34a	0.69%	0.91	-9.02%	0.86	
<b>ATM</b>	pre-miR 34a	1.89%	0.75	-19.30%	0.067	anti-miR 34a	28.62%	0.28	178.04%	0.003	
<b>CYLD</b>	pre-miR 34a	-52.39%	0.0004	-10.26%	0.31	anti-miR 34a	-34.15%	0.066	-3.92%	0.83	
<b>FOXP1</b>	pre-miR 34a	-19.23%	0.10	-1.21%	0.75	anti-miR 34a	-14.99%	0.53	898.78%	0.011	
<b>HDAC1</b>	pre-miR 34a	-20.06%	0.03	-27.21%	0.001	anti-miR 34a	39.56%	0.03	922.51%	0.001	
<b>IBKT</b>	pre-miR 34a	-35.81%	0.02	-14.88%	0.087	anti-miR 34a	-3.66%	0.52	10.68%	0.59	
<b>PTEN</b>	pre-miR 34a	-30.63%	0.065	-18.63%	0.25	anti-miR 34a	-8.25%	0.71	-4.54%	0.78	
<b>SMAD4</b>	pre-miR 34a	-16.37%	0.029	-24.57%	0.06	anti-miR 34a	265.02%	0.002	354.67%	0.023	
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<b>ARID1b</b>	pre-miR 146b	-46.84%	0.0002	-34.79%	0.066	anti-miR 146b	16.21%	0.15	175.97%	0.001	
<b>ATM</b>	pre-miR 146b	-36.18%	0.03	-24.32%	0.033	anti-miR 146b	21.67%	0.08	412.87%	0.003	
<b>CYLD</b>	pre-miR 146b	-47.34%	0.01	-19.99%	0.12	anti-miR 146b	-38.44%	0.071	-8.15%	0.64	
<b>FOXP1</b>	pre-miR 146b	-50.98%	0.0002	13.62%	0.06	anti-miR 146b	8.21%	0.33	8.63%	0.22	
<b>HDAC1</b>	pre-miR 146b	-40.83%	0.004	-29.26%	0.02	anti-miR 146b	46.65%	0.009	880.40%	0.001	
<b>IBKT</b>	pre-miR 146b	-12.83%	0.23	-25.19%	0.03	anti-miR 146b	-4.06%	0.41	-3.42%	0.81	
<b>PTEN</b>	pre-miR 146b	-9.15%	0.48	-28.97%	0.13	anti-miR 146b	-6.68%	0.38	-33.34%	0.083	
<b>SMAD4</b>	pre-miR 146b	-37.43%	0.0014	-28.88%	0.026	anti-miR 146b	847.99%	0.0005	369.78%	0.00001	

Gene expression was determined by TaqMan qPCR performed in triplicate. RNA from MEC1 and SP53 cell lines transfected with pre-miRNA constructs or anti-miRNA constructs was compared to those treated with scramble control oligos. Percent change in expression levels is shown from untreated cells to treated cells with the associated p-value. P values calculated by Student's t-test

**Supplementary Table 13. Differential gene expression of proliferation markers in cell lines transfected with anti-miRNA constructs.**

Marker	Cell/Transfection	Percent Change	P-Value
KI67	MEC1 anti-miR 34a	-7.49%	0.231
TOP2a	MEC1 anti-miR 34a	-17.81%	0.152
TPX	MEC1 anti-miR 34a	-28.86%	0.02
KI67	MEC1 anti-miR 146b	-18.34%	0.38
TOP2a	MEC1 anti-miR 146b	-23.54%	0.31
TPX	MEC1 anti-miR 146b	-23.01%	0.049
KI67	SP53 anti-miR 34a	-15.63%	0.035
TOP2a	SP53 anti-miR 34a	-30.34%	0.0225
TPX	SP53 anti-miR 34a	-34.09%	0.0001
KI67	SP53 anti-miR 146b	-30.55%	0.003
TOP2a	SP53 anti-miR 146b	-27.13%	0.026
TPX	SP53 anti-miR 146b	-2.55%	0.69

Gene expression was determined by TaqMan qPCR performed in triplicate. RNA from MEC1 and SP53 cell lines transfected with anti-miRNA constructs was compared to those treated with scramble control oligos. Percent change in expression levels is shown from untreated cells to treated cells with the associated p-value. *P* values calculated by Student's t-test