

## Appendix

### **Targeting *miR-223* in neutrophils enhances the clearance of *Staphylococcus aureus* in infected wounds**

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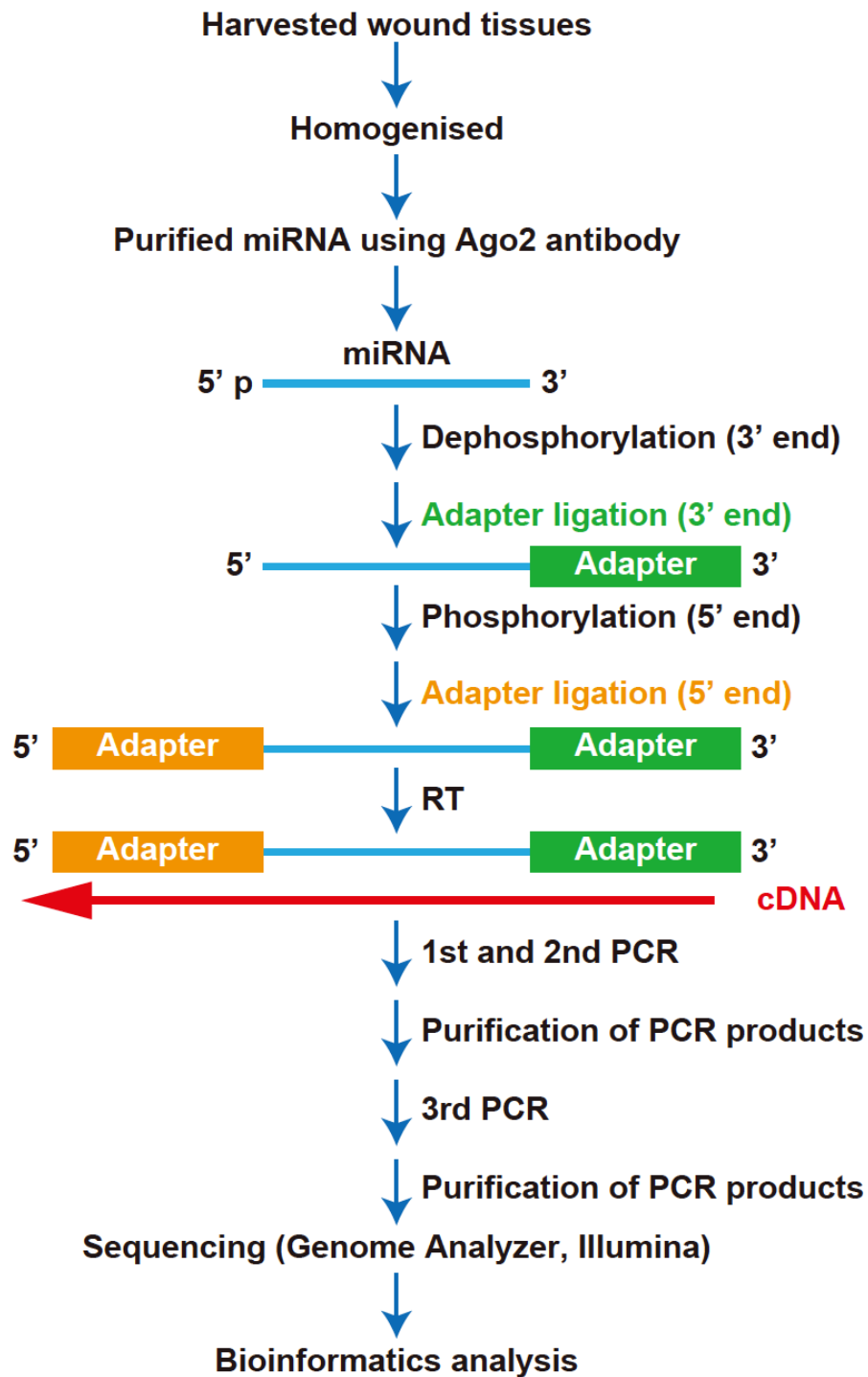
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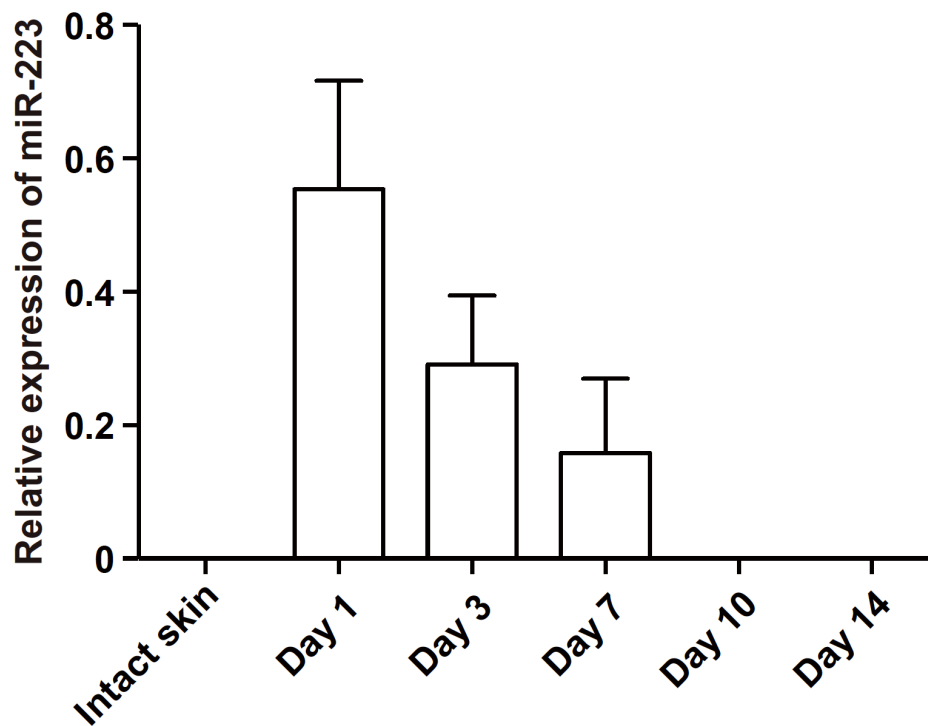
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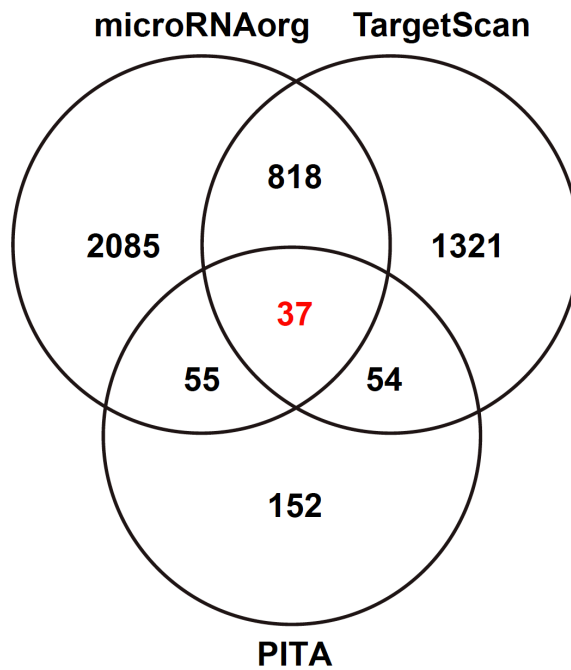
**Appendix Figure S1. Flowchart for the comprehensive identification of wound-induced Ago2 bound-miRNA complexes using NGS.**



**Appendix Figure S2. Expression of *miR-223* in skin wound healing.**

Expression of *miR-223* in murine skin wound healing measured by qPCR relative to snoRNA202 (n = 4 - 6). Part of this results (intact, d 1, d 3, d7) are related to expression of *miR-223* corresponding to each time point in Figure 1. Data information: All values represent the mean ± SD.

Number of miR-223-3p target mRNAs



37 miR-223-3p target mRNAs

<i>Acvr2a</i>	<i>Il6</i>	<i>Slc39a1</i>	<i>Hapln2</i>	<i>Aldh1l2</i>
<i>Alcam</i>	<i>Nfia</i>	<i>Actr1a</i>	<i>Myh10</i>	<i>Hlf</i>
<i>Axin2</i>	<i>Ppp1r14b</i>	<i>Agtpbp1</i>	<i>Klhl1</i>	<i>Pkp4</i>
<i>Cd247</i>	<i>Rbbp6</i>	<i>Atp2b1</i>	<i>Slc16a6</i>	<i>Lrrc8d</i>
<i>Ebf3</i>	<i>Tgfr3</i>	<i>Tmem178</i>	<i>Galnt7</i>	<i>Med19</i>
<i>F3</i>	<i>Psmc6</i>	<i>Ndnf</i>	<i>Ube2q2</i>	
<i>Gpam</i>	<i>Pclo</i>	<i>Ttc4</i>	<i>Csrnp2</i>	
<i>Has3</i>	<i>Tnfrsf19</i>	<i>Tmed5</i>	<i>Frmd4a</i>	

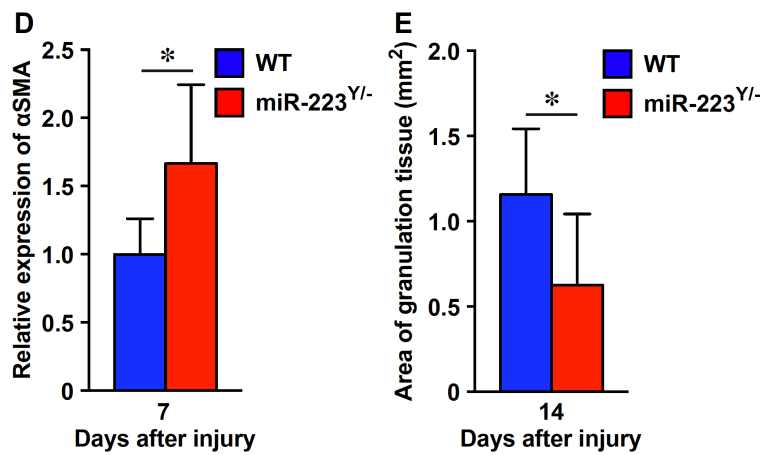
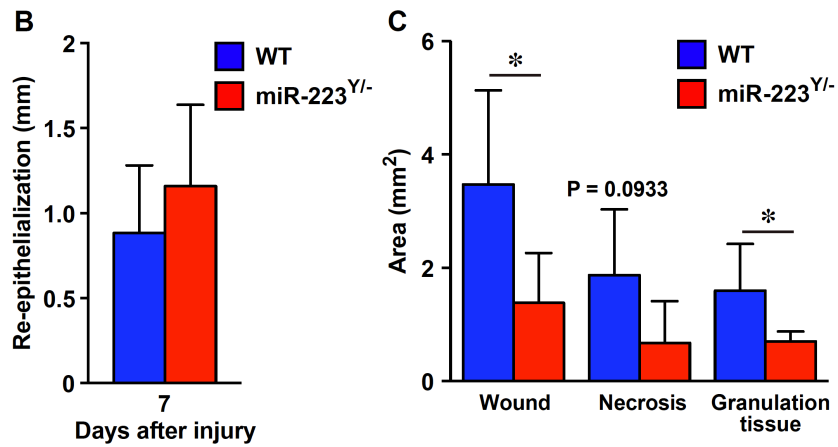
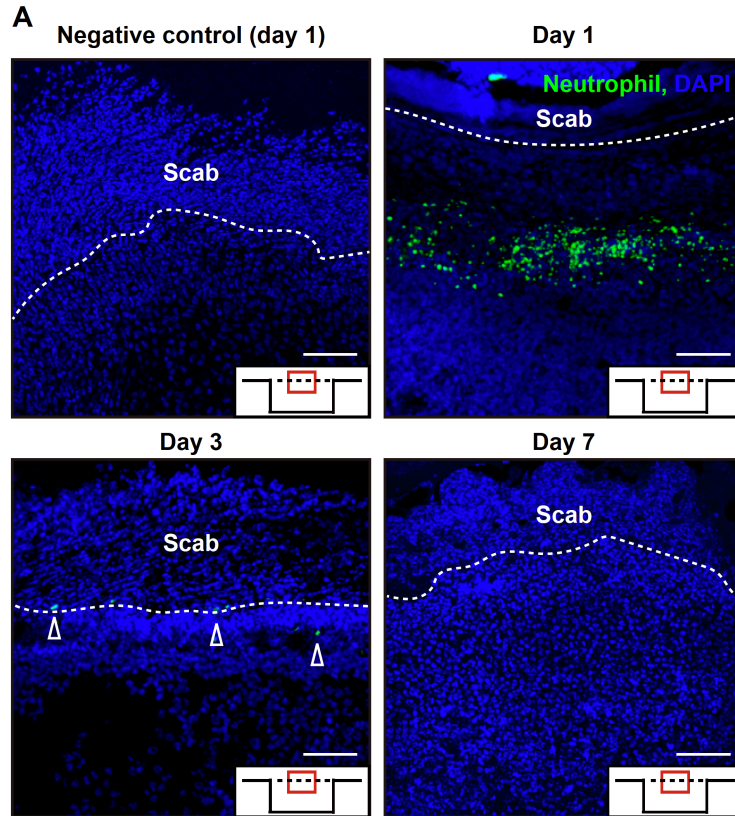


Screening of miR-223-3p target genes  
comparison 37 miRs with NGS data

Identification of miR-223-3p target mRNA 1 d wound sites

*Il6*, *Pclo*

Appendix Figure S3. Flowchart for the comprehensive identification of *miR-223* target mRNAs using Strand NGS and NGS data.



**Appendix Figure S4. *miR-223*<sup>Y/-</sup> neutrophils improve *S. aureus*-infected skin wound healing.**

**A.** Transplanted EGFP-expressing WT neutrophils in *S. aureus* skin wound sites. Representative images of EGFP-expressing WT neutrophils in *S. aureus* skin wound sites at various timepoints after EGFP-expressing neutrophil transplantation (n = 4). Scale bar, 50  $\mu$ m.

**B.** Measurement of the epithelial tongue at d 7 after injury in WT neutrophil-transplanted (WT) and *miR-223*<sup>Y/-</sup> neutrophil-transplanted (*miR-223*<sup>Y/-</sup>) wounds (n = 5).

**C.** Measurement of the total wound area, necrotic lesion area, and granulation tissue area at d 7 after injury in WT neutrophil-transplanted and *miR-223*<sup>Y/-</sup> neutrophil-transplanted wounds (n = 5).

**D.** Quantification of the expression of  $\alpha$ SMA at d 7 in WT neutrophil-transplanted and *miR-223*<sup>Y/-</sup> neutrophil-transplanted wounds (n = 6).

**E.** Measurement of the granulation tissue area at d 14 wound sites from WT neutrophil-transplanted and *miR-223*<sup>Y/-</sup> neutrophil-transplanted wounds (n = 6).

Data information: All values represent the mean  $\pm$  SD. Unpaired *t*-tests were used to generate *P*-values indicated in the Figure. \**P*<0.05.

**Appendix Table S1. Number of annotated reads at various time points in skin wound sites.**

Category	Intact	Day 1	Day 3	Day 7	Day 10	Day 14
total_reads	2,448,759	2,172,903	1,499,496	2,179,215	2,031,016	2,338,879
filtered_out_by_trim	92,478	135,833	105,836	74,749	86,501	87,952
filtered_pass	2,356,281	2,037,070	1,393,660	2,104,466	1,944,515	2,250,927
genome_aligned	2,334,669	2,019,622	1,386,056	2,094,861	1,909,111	2,239,740
target_miRNA	1,920,358 (82.3%)	1,776,639 (88.0%)	1,219,212 (88.0%)	1,924,139 (91.9%)	1,462,271 (76.6%)	2,038,726 (91.0%)
other_miRNA	6,679 (0.3%)	5,210 (0.3%)	4,135 (0.3%)	5,187 (0.2%)	6,209 (0.3%)	5,845 (0.3%)
piRNA	51,457 (2.2%)	18,437 (0.9%)	13,974 (1.0%)	10,887 (0.5%)	51,115 (2.7%)	11,480 (0.5%)
rRNA	12,620 (0.5%)	2,354 (0.1%)	2,302 (0.2%)	2,838 (0.1%)	11,169 (0.6%)	3,259 (0.1%)
tRNA	757 (0.03%)	254 (0.01%)	28 (0.0002%)	371 (0.02%)	330 (0.02%)	485 (0.02%)
snRNA	3,443 (0.1%)	6,197 (0.3%)	5,628 (0.4%)	1,313 (0.1%)	5,108 (0.3%)	1,465 (0.1%)
snoRNA	3,177 (0.1%)	2,245 (0.1%)	996 (0.1%)	815 (0.04%)	2,218 (0.1%)	933 (0.04%)
scRNA	0	0	0	0	0	0
miscRNA	2,120 (0.1%)	606 (0.03%)	368 (0.03%)	514 (0.02%)	1,665 (0.1%)	723 (0.03%)
RefSeq transcripts	193,247 (8.3%)	84,105 (4.2%)	43,293 (3.1%)	63,332 (3.0%)	229,026 (12.0%)	79,483 (3.5%)
Only Genome	140,811 (6.0%)	123,575 (6.1%)	96,120 (6.9%)	85,465 (4.1%)	140,000 (7.3%)	97,341 (4.3%)

Abbreviations: piRNA, Piwi-interacting RNA; rRNA, ribosomal RNA; tRNA, transfer RNA; snRNA, small nuclear RNA; snoRNA, small nucleolar RNA; scRNA, small cytoplasmic RNA; miscRNA, miscellaneous other RNA.

\*Values in parentheses show the percentage of similarity for each genome aligned sample.

**Appendix Table S2. Top 9 candidate inflammation-related miRNA reads at various time points in skin wound sites.**

miRNA	Intact	Day 1	Day 3	Day 7	Day 10	Day 14	Day 1 / Intact
mmu-miR-147	6	442	96	8	12	8	73.67
mmu-miR-223	3,957	138,618	29,135	7,251	2,968	5,883	35.03
mmu-miR-129-3p	19	582	64	47	47	54	30.63
mmu-miR-139-5p	575	7,693	1,331	1,051	859	1,114	13.38
mmu-miR-21*	256	2,751	1,721	76	67	34	10.75
mmu-miR-340-5p	250	1,672	1,331	462	179	439	6.69
mmu-miR-142-3p	1,872	12,270	6,540	5,416	2,617	4,324	6.55
mmu-miR-142-5p	2,493	15,944	8,980	5,492	3,146	3,254	6.40
mmu-miR-486	1,834	8,266	3,040	5,010	4,851	4,243	4.51

Abbreviation: mmu, *Mus musculus*. Note: Cutoff value > 4.5 data from Dataset EV1.



**Appendix Table S3. High expression of miRNA reads at various time points in skin wound sites.**

**Intact skin vs. day 3**

miRNA	Intact	Day 1	Day 3	Day 7	Day 10	Day 14	Day 3 / Intact
mmu-miR-147	6	442	96	8	12	8	16.00
mmu-miR-223	3957	138618	29135	7251	2968	5883	7.36
mmu-miR-21*	256	2751	1721	76	67	34	6.72
mmu-miR-450b-5p	2	13	12	33	8	20	6.00
mmu-miR-18b	3	7	16	2	3	3	5.33
mmu-miR-340-5p	250	1672	1331	462	179	439	5.32
mmu-miR-21	46385	108302	218138	400610	195053	267573	4.70
mmu-miR-363	7	57	32	38	14	27	4.57

Note: Cutoff value > 4.5 data from Dataset EV1.

**Intact skin vs. day 7**

miRNA	Intact	Day 1	Day 3	Day 7	Day 10	Day 14	Day 7 / Intact
mmu-miR-592	2	5	8	54	66	24	27.00
mmu-miR-155	23	18	23	409	302	307	17.78
mmu-miR-450b-5p	2	13	12	33	8	20	16.50
mmu-miR-335-3p	9	3	2	146	36	46	16.22
mmu-miR-297a	1	1	2	13	9	6	13.00
mmu-miR-214*	84	109	104	900	611	973	10.71
mmu-miR-672	4	0	0	42	19	9	10.50
mmu-miR-29a*	14	25	17	134	123	226	9.57
mmu-miR-450a-5p	210	207	179	1913	581	1177	9.11
mmu-miR-1188	4	4	8	36	24	10	9.00
mmu-miR-21	46385	108302	218138	400610	195053	267573	8.64
mmu-miR-574-5p	4	6	2	29	30	45	7.25
mmu-miR-455*	19	7	11	136	87	130	7.16
mmu-miR-137	1	5	2	7	3	9	7.00

mmu-miR-294	1	1	2	7	1	8	7.00
mmu-miR-499	4	4	6	27	4	14	6.75
mmu-miR-27a*	2	13	0	12	6	9	6.00
mmu-miR-875-5p	1	0	0	6	1	5	6.00
mmu-miR-126-5p	1244	632	498	7372	6015	8831	5.93
mmu-miR-136	1197	1034	522	7008	4893	7569	5.85
mmu-miR-679	3	5	3	17	10	9	5.67
mmu-miR-667	60	56	55	329	299	224	5.48
mmu-miR-34b-3p	33	81	113	180	103	83	5.45
mmu-miR-363	7	57	32	38	14	27	5.43
mmu-miR-467c	5	16	4	23	19	16	4.60
mmu-miR-467d	5	0	2	23	19	20	4.60

Note: Cutoff value > 4.5 data from Dataset EV1.

### Intact skin vs. day 10

miRNA	Intact	Day 1	Day 3	Day 7	Day 10	Day 14	Day 10 / Intact
mmu-miR-592	2	5	8	54	66	24	33.00
mmu-miR-155	23	18	23	409	302	307	13.13
mmu-miR-297a	1	1	2	13	9	6	9.00
mmu-miR-29a*	14	25	17	134	123	226	8.79
mmu-miR-574-5p	4	6	2	29	30	45	7.50
mmu-miR-214*	84	109	104	900	611	973	7.27
mmu-miR-1934	1	10	1	4	7	5	7.00
mmu-miR-206	3585	8831	5407	15022	23863	14210	6.66
mmu-miR-1188	4	4	8	36	24	10	6.00
mmu-miR-466f-3p	2	22	3	4	12	5	6.00
mmu-miR-384-5p	1	0	0	3	5	2	5.00
mmu-miR-701	2	0	5	7	10	7	5.00
mmu-miR-667	60	56	55	329	299	224	4.98

mmu-miR-126-5p	1244	632	498	7372	6015	8831	4.84
mmu-miR-672	4	0	0	42	19	9	4.75
mmu-miR-455*	19	7	11	136	87	130	4.58
mmu-miR-297c	2	8	1	4	9	9	4.50

Note: Cutoff value > 4.5 data from Dataset EV1.

### Intact skin vs. day 14

miRNA	Intact	Day 1	Day 3	Day 7	Day 10	Day 14	Day 14 / Intact
mmu-miR-29a*	14	25	17	134	123	226	16.14
mmu-miR-155	23	18	23	409	302	307	13.35
mmu-miR-592	2	5	8	54	66	24	12.00
mmu-miR-214*	84	109	104	900	611	973	11.58
mmu-miR-574-5p	4	6	2	29	30	45	11.25
mmu-miR-450b-5p	2	13	12	33	8	20	10.00
mmu-miR-137	1	5	2	7	3	9	9.00
mmu-miR-294	1	1	2	7	1	8	8.00
mmu-miR-126-5p	1244	632	498	7372	6015	8831	7.10
mmu-miR-190b	1	1	0	1	3	7	7.00
mmu-miR-455*	19	7	11	136	87	130	6.84
mmu-miR-136	1197	1034	522	7008	4893	7569	6.32
mmu-miR-297a	1	1	2	13	9	6	6.00
mmu-miR-21	46385	108302	218138	400610	195053	267573	5.77
mmu-miR-450a-5p	210	207	179	1913	581	1177	5.60
mmu-miR-9	12	12	10	44	24	65	5.42
mmu-miR-124	9	19	13	23	39	46	5.11
mmu-miR-335-3p	9	3	2	146	36	46	5.11
mmu-miR-146a	1394	2833	3633	5372	4318	7019	5.04
mmu-miR-1934	1	10	1	4	7	5	5.00
mmu-miR-875-5p	1	0	0	6	1	5	5.00

mmu-miR-27a*	2	13	0	12	6	9	4.50
mmu-miR-297c	2	8	1	4	9	9	4.50

Note: Cutoff value > 4.5 data from Dataset EV1.

**Appendix Table S4. The profile of each human sample.**

Sex	Age (years)	Etiology	Biopsy site
M	67	Epidermal cyst	Face
M	75	Cellulitis with microabscess	Face
M	60	Inflammatory granulation tissue with microabscess	Abdominal
F	32	Cystic lesion with inflammatory granulation tissue and fibrosis	Inguinal
F	79	Abscess	Face
F	71	Epidermal cyst	Neck
M	28	Epidermal cyst with microabscess and inflammatory granulation tissue	Face
M	82	Necrotizing lesion with neutrophil and eosinophil infiltration	Head
F	56	Epidermal cyst with multiple abscess and chronic inflammation and psoriasiform dermatitis	Hip
F	12	Abscess forming granulomatous inflammation	Eyelid
F	23	Cystic lesion with inflammatory granulation tissue, foreign body reaction, and granuloma	Arm
F	33	Pustular dermatitis with erosion, ulcer, and gram positive cocci	Arm
F	47	Dermal neutrophilic infiltration	Thing
F	75	Abscess	Leg
M	62	Cutaneous ulcer	Foot

Abbreviations: M, Male; F, Female

**Appendix Table S5. List of antibodies.**

<b>Primary antibody</b>	<b>Company</b>	<b>Catalog No.</b>	<b>Species</b>	<b>Dilution</b>
$\alpha$ smooth muscle actin	Abcam	ab5694	Rabbit	1:500
F4/80	Abcam	ab6640	Rat	1:400
IL6	Abcam	ab6672	Rabbit	1:400
Neutrophil (Ly-6G and Ly-6C)	Abcam	ab2557	Rat	1:400
Alexa Fluor 568	ThermoFisher	A-11011	Goat	1:1000
DIG	Roche Diagnostics	11093274910	Sheep	1:800
Ago2	Wako Pure Chemical Industries	292-67301	Mouse	5 $\mu$ g/IP
C/EBP $\alpha$	GeneTex	GTX100674	Rabbit	5 $\mu$ g/IP
Histone H3	Cell Signaling Technology	4620	Rabbit	5 $\mu$ g/IP
Normal rabbit IgG	Cell Signaling Technology	2729	Rabbit	5 $\mu$ g/IP

Abbreviation: IP, immunoprecipitation.

**Appendix Table S6. List of primers.**

Gene symbol	Sequences (5'-3')
Rps18	(F) 5'-TTCTGGCCAACGGTCTAGACAAC-3' (R) 5'-CCAGTGGTCTTGGTGTGCTGA-3'
Il6 (M)	(F) 5'-CAACGATGATGCACTTGCAGA-3' (R) 5'-CTCCAGGTAGCTATGGTACTCCAGA-3'
Il6ra	(F) 5'- CCTGAGACTCAAGCAGAAATGG-3' (R) 5'-AGAAGGAAGGTCGGCTTCAGT-3'
Il6st	(F) 5'- CCGTGTGGTTACATCTACCCT-3' (R) 5'-CGTGGTTCTGTTGATGACAGTG-3'
Il1b	(F) 5'-TCCAGGATGAGGACATGAGCAC-3' (R) 5'-GAACGTCACACACCAGCAGGTTA-3'
B2M	(F) 5'-CGGGCATTCTGAAGCTGA-3' (R) 5'-GGATGGATGAAACCCAGACACATAG-3'
IL6 (H)	(F) 5'-AAGCCAGAGCTGTGCAGATGAGTA-3' (R) 5'-TGTCTGCAGCCACTGGTTC-3'
CEBPA	(F) 5'-TCGTGGGTCAGCTCTGAGGA-3' (R) 5'-GCAATGCTGAAGGCATACAGTACAA-3'
RUNX1	(F) 5'-CACTGGCGCTGCAACAAGA-3' (R) 5'-CCAGCCATCACAGTGACCAGA-3'
SPI1	(F) 5'-GCCCTATGACACGGATCTATACCAA-3' (R) 5'-TCTCGGCGAAGCTCTCGAA-3'
ChIP-PCR ChIP-qPCR	(F) 5'-GCCCTCTTTGTTGATGTGTC-3' (R) 5'-GGCAGCTATTAAAGTGCCCT-3'

Abbreviations: F; forward, R; reverse, M; *Mus musculus*, H; *Homo sapiens*.

**Appendix Table S7. Statistical analysis.**

**Fig 1. *miR-147***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	****	< 0.0001	Tukey's multiple comparisons test
Intact vs. d 3	***	0.0005	Tukey's multiple comparisons test
d 1 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test
d 3 vs. d 7	***	0.0005	Tukey's multiple comparisons test

**Fig 1. *miR-223***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	****	< 0.0001	Tukey's multiple comparisons test
Intact vs. d 3	**	0.0011	Tukey's multiple comparisons test
d 1 vs. d 3	**	0.0029	Tukey's multiple comparisons test
d 1 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test

**Fig 1. *miR-129-3p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	*	0.0461	Tukey's multiple comparisons test
Intact vs. d 3	****	< 0.0001	Tukey's multiple comparisons test
d 1 vs. d 3	***	0.0001	Tukey's multiple comparisons test
d 3 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test



**Fig 1. miR-139-5p**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	****	< 0.0001	Tukey's multiple comparisons test
Intact vs. d 3	***	0.0005	Tukey's multiple comparisons test
Intact vs. d 7	*	0.0252	Tukey's multiple comparisons test
d 1 vs. d 3	***	0.0009	Tukey's multiple comparisons test
d 1 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test

**Fig 1. miR-21\***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	**	0.0056	Tukey's multiple comparisons test
d 1 vs. d 3	*	0.0440	Tukey's multiple comparisons test
d 1 vs. d 7	*	0.0470	Tukey's multiple comparisons test

**Fig 1. miR-340-5p**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	**	0.0070	Tukey's multiple comparisons test
Intact vs. d 3	****	0.0005	Tukey's multiple comparisons test
d 1 vs. d 3	***	0.0001	Tukey's multiple comparisons test
d 3 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test

**Fig 1. *miR-142-3p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	*	0.0486	Tukey's multiple comparisons test
Intact vs. d 3	**	0.0027	Tukey's multiple comparisons test
d 3 vs. d 7	**	0.0080	Tukey's multiple comparisons test

**Fig 1. *miR-142-5p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. d 1	****	< 0.0001	Tukey's multiple comparisons test
Intact vs. d 3	****	< 0.0001	Tukey's multiple comparisons test
d 1 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test
d 3 vs. d 7	****	< 0.0001	Tukey's multiple comparisons test

**Fig 2A. Expression of *miR-223***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. 12 hrs	****	< 0.0001	Tukey's multiple comparison test
0.5hr vs. 12 hrs	****	< 0.0001	Tukey's multiple comparison test
3 hrs vs. 12 hrs	**	0.0062	Tukey's multiple comparison test
12 hrs vs. 24 hrs	***	0.0003	Tukey's multiple comparison test

**Fig 2A. Expression of *miR-142-3p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Intact vs. 12 hrs	**	0.0032	Tukey's multiple comparison test
0.5hr vs. 3 hrs	*	0.0500	Tukey's multiple comparison test
0.5hr vs. 12 hrs	**	0.0032	Tukey's multiple comparison test

**Fig 2B. Expression of *miR-223***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
WT vs. <i>PU.1</i> <sup>+/-</sup>	****	< 0.0001	Tukey's multiple comparison test
WT vs. <i>PU.1</i> <sup>-/-</sup>	****	< 0.0001	Tukey's multiple comparison test
<i>PU.1</i> <sup>+/-</sup> vs. <i>PU.1</i> <sup>-/-</sup>	***	0.0004	Tukey's multiple comparison test

**Fig 2B. Expression of *miR-142-3p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
WT vs. <i>PU.1</i> <sup>+/-</sup>	****	< 0.0001	Tukey's multiple comparison test
WT vs. <i>PU.1</i> <sup>-/-</sup>	****	< 0.0001	Tukey's multiple comparison test
<i>PU.1</i> <sup>+/-</sup> vs. <i>PU.1</i> <sup>-/-</sup>	***	0.0004	Tukey's multiple comparison test

**Fig 2B. Expression of *miR-142-5p***

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
WT vs. <i>PU.1</i> <sup>+/-</sup>	****	< 0.0001	Tukey's multiple comparison test
WT vs. <i>PU.1</i> <sup>-/-</sup>	****	< 0.0001	Tukey's multiple comparison test
<i>PU.1</i> <sup>+/-</sup> vs. <i>PU.1</i> <sup>-/-</sup>	**	0.0024	Tukey's multiple comparison test

**Fig 2B. Expression of *miR-139-5p***

Group compared	Summary	P value	Statistical test
WT vs. <i>PU.1</i> <sup>-/-</sup>	**	0.0064	Tukey's multiple comparison test
<i>PU.1</i> <sup>+/-</sup> vs. <i>PU.1</i> <sup>-/-</sup>	**	0.0028	Tukey's multiple comparison test

**Fig 3F. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
3 d	***	0.0003	Sidak's multiple comparisons test
7 d	*	0.0376	Sidak's multiple comparisons test

**Fig 3H. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
3 d	*	0.0115	Unpaired Student's <i>t</i> -tests

**Fig 3J. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
7 d	*	0.0206	Unpaired Student's <i>t</i> -tests
14 d	*	0.0246	Unpaired Student's <i>t</i> -tests

**Fig 4B. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
3 h	*	0.0445	Unpaired Student's <i>t</i> -tests
3 d	*	0.0445	Sidak's multiple comparisons test

**Fig 4D. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
1 d	*	0.0322	Unpaired Student's <i>t</i> -tests
3 d	*	0.0025	Unpaired Student's <i>t</i> -tests

**Fig 4E. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
1 d	*	0.0285	Unpaired Student's <i>t</i> -tests
3 d	*	0.0101	Unpaired Student's <i>t</i> -tests

**Fig 4G. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
17 min	*	0.0356	Mann Whitney test
45 min	*	0.0356	Mann Whitney test
46 min	*	0.0415	Mann Whitney test
47 min	*	0.0259	Mann Whitney test
48 min	*	0.0167	Mann Whitney test
49 min	*	0.0156	Mann Whitney test
50 min	*	0.0151	Mann Whitney test
51 min	*	0.0173	Mann Whitney test
52 min	*	0.0146	Mann Whitney test
53 min	*	0.0162	Mann Whitney test
54 min	*	0.0141	Mann Whitney test

55 min	*	0.0173	Mann Whitney test
56 min	*	0.0146	Mann Whitney test
57 min	*	0.0156	Mann Whitney test
58 min	*	0.0131	Mann Whitney test
59 min	*	0.0151	Mann Whitney test
60 min	*	0.0113	Mann Whitney test

**Fig 5B. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
<i>Il6</i>	*	0.0245	Unpaired Student's <i>t</i> -tests

**Fig 5H. *miR-223* mimic vs. each sample**

Group compared	Summary	P value	Statistical test
vs. control	**	0.0093	Dunnett's multiple comparisons test
vs. mutation 2	*	0.0316	Dunnett's multiple comparisons test
vs. mutation 3	**	0.0064	Dunnett's multiple comparisons test

**Fig 6B. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
3 d	*	0.0467	Sidak's multiple comparisons test
7 d	**	0.0011	Sidak's multiple comparisons test

**Fig 6C. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
3d	*	0.0403	Unpaired Student's <i>t</i> -tests

**Fig 6E. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
1d	*	0.0403	Unpaired t test with Welch's correction

**Fig 6F. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
<i>Il6</i>	*	0.0394	Unpaired test
<i>Il6ra</i>	**	0.0013	Unpaired Student's <i>t</i> -tests
<i>Il1β</i>	*	0.0371	Unpaired t test with Welch's correction

**Fig 6H. 3d**

Group compared	Summary	P value	Statistical test
WT control vs. WT-neutrophil	-	0.4992	Tukey's multiple comparisons test
WT control vs. KO-neutrophil	*	0.0124	Tukey's multiple comparisons test
WT neutrophil vs. KO-neutrophil	-	0.1961	Tukey's multiple comparisons test

**Fig 6H. 7d**

Group compared	Summary	P value	Statistical test
WT control vs. WT-neutrophil	-	0.7993	Tukey's multiple comparisons test
WT control vs. KO-neutrophil	**	0.0023	Tukey's multiple comparisons test
WT neutrophil vs. KO-neutrophil	***	0.0002	Tukey's multiple comparisons test

**Fig 7B.**

Group compared	Summary	P value	Statistical test
Control ODN vs. <i>miR-223</i> AS ODN	****	< 0.0001	Unpaired Student's <i>t</i> -tests

**Fig 7D. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
6 h	**	0.0083	Unpaired t test with Welch's correction
1 d	****	< 0.0001	Unpaired Student's <i>t</i> -tests

**Fig 7E.**

Group compared	Summary	P value	Statistical test
Control ODN vs. <i>miR-223</i> AS ODN	*	0.0307	Unpaired Student's <i>t</i> -tests

**Fig 7G. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
3 d	***	0.0002	Sidak's multiple comparisons test
7 d	**	0.0064	Sidak's multiple comparisons test

**Fig 7H. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
Wound	*	0.0217	Unpaired t test with Welch's correction
Necrosis	*	0.0309	Unpaired t test with Welch's correction

**Fig 7I.**

Group compared	Summary	P value	Statistical test
Control ODN vs. <i>miR-223</i> AS ODN	**	0.0010	Unpaired Student's <i>t</i> -tests

**Fig 7J.**

Group compared	Summary	P value	Statistical test
Control ODN vs. <i>miR-223</i> AS ODN	***	0.0005	Unpaired Student's <i>t</i> -tests



**Fig 8A. Expression of *miR-223* in HL-60 cells**

Group compared	Summary	P value	Statistical test
Control vs. 2 d	*	0.0499	Holm-Sidak's multiple comparisons test
Control vs. 3 d	*	0.0137	Holm-Sidak's multiple comparisons test
Control vs. 4 d	****	< 0.0001	Holm-Sidak's multiple comparisons test
Control vs. 5 d	****	< 0.0001	Holm-Sidak's multiple comparisons test

**Fig 8B. *miR-223* AS ODN *miR-223* expression in dHL60**

Group compared	Summary	P value	Statistical test
Control vs. <i>miR-223</i> AS ODN	*	0.0385	Unpaired t test with Welch's correction
Control ODN vs. <i>miR-223</i> AS ODN	*	0.0384	Unpaired t test with Welch's correction

**Fig 8C. Non PGN vs. PGN**

Group compared	Summary	P value	Statistical test
Control	*	0.0176	Unpaired Student's <i>t</i> -tests
<i>miR-223</i> AS ODN	**	0.0056	Unpaired Student's <i>t</i> -tests

**Fig 8D. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
Non PGN	**	0.0056	Unpaired Student's <i>t</i> -tests
PGN	**	0.0096	Unpaired Student's <i>t</i> -tests

**Fig 8E. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
Non PGN	**	0.0089	Unpaired Student's <i>t</i> -tests
6 hrs	*	0.0120	Unpaired Student's <i>t</i> -tests
12 hrs	*	0.0116	Unpaired Student's <i>t</i> -tests

**Fig 8G. Relative binding activity**

Group compared	Summary	P value	Statistical test
Control vs. PGN	*	0.0361	Unpaired Student's <i>t</i> -tests

**Fig EV1C. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
d 7	*	0.0206	Unpaired <i>t</i> test
d 14	*	0.0246	Unpaired <i>t</i> test

**Fig EV1E.**

Group compared	Summary	P value	Statistical test
WT vs. <i>miR-223</i> <sup>Y/-</sup>	**	0.0013	Unpaired <i>t</i> test

**Fig EV2B. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
d 3	**	0.0044	Unpaired <i>t</i> test
d 7	*	0.0154	Unpaired <i>t</i> test

**Fig EV2C.**

Group compared	Summary	P value	Statistical test
WT vs. <i>miR-223</i> <sup>Y/-</sup>	*	0.0478	Unpaired t test

**Fig EV3C. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
d 3	*	0.0206	Unpaired t test
d 7	*	0.0367	Unpaired t test

**Fig EV3D. WT vs. *miR-223*<sup>Y/-</sup>**

Group compared	Summary	P value	Statistical test
Wound	*	0.0467	Unpaired t test
Necrosis	*	0.0434	Unpaired t test

**Fig EV3F.**

Group compared	Summary	P value	Statistical test
WT vs. <i>miR-223</i> <sup>Y/-</sup>	*	0.0470	Unpaired t test

**Fig EV4A. Control ODN vs. *miR-223* AS ODN**

Group compared	Summary	P value	Statistical test
d 7	***	0.0004	Sidak's multiple comparisons test

**Fig EV4B. Pluronic gel vs. PB gel**

Group compared	Summary	P value	Statistical test
d 3	**	0.0012	Sidak's multiple comparisons test

**Fig EV5A. Non PGN vs. PGN**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
<i>CEBPA</i>	**	0.0089	Unpaired t test

**Appendix Fig S4C. WT neutrophil vs. *miR-223*<sup>Y/-</sup> neutrophil**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
Wound	*	0.0467	Unpaired t test
Necrosis	-	0.0933	Unpaired t test
Granulation tissue	*	0.0441	Unpaired t test

**Appendix Fig S4D.**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
WT neutrophil vs. <i>miR-223</i> <sup>Y/-</sup> neutrophil	*	0.0282	Unpaired t test

**Appendix Fig S4E.**

<b>Group compared</b>	<b>Summary</b>	<b>P value</b>	<b>Statistical test</b>
WT neutrophil vs. <i>miR-223</i> <sup>Y/-</sup> neutrophil	*	0.0446	Unpaired t test