

Supplementary Tables

Supplementary Table 1: Genotyping Primers

| Gene name | Forward sequence (5' to 3') | Reverse sequence (3' to 5') |
|--------------------------|-------------------------------|-----------------------------|
| <i>Rbp1Cre transgene</i> | CAA GTG TGA GAG ACA GCA TTG | TCC TTA GCG CCG TAA ATC AA |
| <i>tdTomato wildtype</i> | AAG GGA GCT GCA GTG GAG TA | CCG AAA ATC TGT GGG AAG TC |
| <i>tdTomato mutant</i> | CTG TTC CTG TAC GGC ATG G | GGC ATT AAA GCA GCG TAT CC |
| <i>RPE65</i> | CAC TGT GGT CTC TGC TAT CTT C | GGT GCA GTT CCA CTT CAG TT |

Supplementary Table 2: Primary Antibodies

| Antibody | Concentration | Company, Catalog # |
|--------------------------------------|----------------------|-------------------------------|
| rat anti RFP (tdTomato) | 1:500 | Antibodies online, ABIN334653 |
| mouse anti glutamine synthetase (GS) | 1:200 | Millipore, MAB302 |
| rabbit anti Sox9 | 1:1000 | Millipore, AB5535 |
| rabbit anti GFAP | 1:1000 | Dako, Z033401-2 |
| goat anti Otx2 | 1:250 | R&D Systems, AF1979 |

Supplementary Table 3. Glial gene expression in normal and light damaged Müller glia (Microarray).

| Gene Symbol | Undamaged control log2 relative expression | 7 days after light damage log2 relative expression | Fold change light damage versus control |
|-------------|--|--|---|
| Glul | 13.84596 | 13.06892 | -0.78 |
| Rlbp1 | 12.38061 | 11.62174 | -0.76 |
| Aqp4 | 11.80212 | 11.42555 | -0.38 |
| Vim | 11.88869 | 13.19273 | 1.30 |
| Slc1a3 | 12.14886 | 12.14886 | 0.00 |
| Sox9 | 10.11897 | 10.73825 | 0.62 |
| Sox2 | 10.94028 | 11.58379 | 0.64 |
| Gfap | 6.282722 | 10.5111 | 4.23 |

Supplementary Table 4: Expression levels (decline) of the Müller glia (MG) miRNAs (cutoff 3000 counts) after light damage (LD), in normal adult wild type and Dicer-CKO MG (Nanostring).

| Probe Name | normal MG (counts) | MG Light damage (counts) | Dicer-CKO MG (counts) | % decline after LD | % decline after cKO | Accession # |
|-----------------|--------------------|--------------------------|-----------------------|--------------------|---------------------|--------------|
| mmu-miR-204 | 285706 | 51963 | 30438 | -82 | -89 | MIMAT0000237 |
| mmu-miR-125b-5p | 109596 | 19063 | 18202 | -83 | -83 | MIMAT0000136 |
| mmu-miR-9 | 75416 | 38870 | 18100 | -48 | -76 | MIMAT0000142 |
| mmu-miR-181a | 48829 | 19965 | 8300 | -59 | -83 | MIMAT0000210 |
| mmu-let-7c | 26178 | 9888 | 9984 | -62 | -62 | MIMAT0000523 |
| mmu-miR-99a | 19815 | 9626 | 2284 | -51 | -88 | MIMAT0000131 |
| mmu-let-7b | 18043 | 11425 | 3875 | -37 | -79 | MIMAT0000522 |
| mmu-let-7g | 15643 | 13871 | 6952 | -11 | -56 | MIMAT0000121 |
| mmu-miR-30c | 13955 | 7419 | 3296 | -47 | -76 | MIMAT0000514 |
| mmu-let-7d | 13294 | 6634 | 3557 | -50 | -73 | MIMAT0000383 |
| mmu-miR-135a | 12833 | 11626 | 4865 | -9 | -62 | MIMAT0000147 |
| mmu-let-7a | 8129 | 3648 | 2704 | -55 | -67 | MIMAT0000521 |
| mmu-miR-16 | 7260 | 2732 | 3358 | -62 | -54 | MIMAT0000527 |
| mmu-miR-125a-5p | 5639 | 1275 | 661 | -77 | -88 | MIMAT0000135 |
| mmu-miR-100 | 5626 | 1507 | 524 | -73 | -91 | MIMAT0000655 |
| mmu-miR-1944 | 5024 | 2622 | 3535 | -48 | -30 | MIMAT0009409 |

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|----------------|------|------|------|-----|-----|--------------|
| mmu-miR-22 | 4916 | 4438 | 1236 | -10 | -75 | MIMAT0000531 |
| mmu-miR-30d | 4676 | 1910 | 909 | -59 | -81 | MIMAT0000515 |
| mmu-let-7e | 3790 | 1521 | 984 | -60 | -74 | MIMAT0000524 |
| mmu-miR-335-5p | 3329 | 1702 | 1311 | -49 | -61 | MIMAT0000766 |
| mmu-miR-23a | 3327 | 1796 | 1515 | -46 | -54 | MIMAT0000532 |

Supplementary Table 5: Expression levels (increase) of the Müller glia (MG) miRNAs after light damage (LD, cutoff 5000 counts after LD), in normal adult wild type and Dicer-CKO MG (Nanostring).

| Probe Name | normal MG (counts) | MG Light damage (counts) | Dicer-CKO MG (counts) | Fold increase after LD | Fold increase cKO | Accession # |
|------------|--------------------|--------------------------|-----------------------|------------------------|-------------------|--------------|
| miR-720** | 23904 | 172827 | 10983 | 7.2 | 0.5 | MIMAT0003484 |
| miR-29a | 8716 | 10969 | 4052 | 1.3 | 0.5 | MIMAT0000535 |
| miR-29b | 1748 | 7515 | 2005 | 4.3 | 1.1 | MIMAT0000127 |
| miR-124 | 1354 | 5506 | 12106 | 4.1 | 8.9 | MIMAT0000134 |
| miR-1937a | 1094 | 6129 | 1298 | 5.6 | 1.2 | MIMAT0009401 |

** The sequence annotated as miR-720 is likely to be a fragment of a tRNA, and therefore is removed from the miRBase database ([Schopman et al., 2010](#)).

Supplementary Table 6: Expression levels of Argonaute2-bound miRNAs in whole mouse retinas (data set from the Natoli lab (Chu-Tan et al., 2020)).

| Probe Name | Log2CPM light damage | Probe Name | Log2CPM light damage |
|-------------|----------------------|------------|----------------------|
| miR-204-5p | 11.17327 | miR-1944 | - |
| miR-125b-5p | 13.76099 | miR-22-3p | 8.905651 |
| miR-9-5p | 4.631367 | miR-30d-5p | 7.862324 |
| miR-181a-5p | 12.86331 | let-7e-5p | 7.14469 |
| let-7c-5p | 9.809521 | miR-335-5p | 7.864721 |
| miR-99a-5p | 9.097142 | miR-23a-5p | 3.497926 |
| let-7b-5p | 9.04164 | miR-720 | - |
| let-7g-5p | 10.04702 | miR-29a-5p | 9.068966 |
| miR-30c-5p | 10.32144 | miR-29b-3p | 13.17489 |
| let-7d-5p | 8.755262 | miR-124-3p | 19.58201 |
| miR-135a-5p | 9.069131 | miR-1937a | - |
| let-7a-5p | 5.292606 | miR-183-5p | 14.30006 |
| miR-16-5p | 8.524132 | miR-182-3p | 7.636908 |
| miR-125a-5p | 13.37126 | miR-96-5p | 10.51531 |
| miR-100-5p | 6.124773 | miR-22-3p | 8.905651 |

Supplementary Table 7: Top 50 genes upregulated after light damage (Microarray).

| Gene | Undamaged control log2 relative expression | Light Damage log2 relative expression | Fold change light damage versus control |
|-------------|---|--|--|
| Myc | 5.546965 | 10.58033 | 5.03 |
| Ednrb | 7.38606 | 12.32283 | 4.94 |
| Serpina3n | 7.027112 | 11.8461 | 4.82 |
| Cxcl10 | 5.769135 | 10.57737 | 4.81 |
| Lcn2 | 7.960248 | 12.71421 | 4.75 |
| Gfap | 6.282722 | 10.5111 | 4.23 |
| Timp1 | 6.105936 | 10.3307 | 4.22 |
| Gadd45b | 7.56354 | 11.64912 | 4.09 |
| Ifi203 | 4.298881 | 8.316713 | 4.02 |
| Anxa2 | 5.3885 | 9.285599 | 3.90 |
| Steap4 | 6.341545 | 10.20562 | 3.86 |
| Mt2 | 8.264072 | 12.11129 | 3.85 |
| Klf6 | 8.065225 | 11.78304 | 3.72 |
| Ccl2 | 6.402283 | 10.11604 | 3.71 |
| S100a6 | 6.641229 | 10.29619 | 3.65 |
| Tubb6 | 6.196977 | 9.784472 | 3.59 |
| Gbp2 | 6.841433 | 10.37846 | 3.54 |

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|---------|----------|----------|------|
| Hmox1 | 6.483181 | 10.01682 | 3.53 |
| Acsl5 | 5.492249 | 9.016914 | 3.52 |
| Emp1 | 6.499097 | 10.00783 | 3.51 |
| Ch11 | 7.436919 | 10.9089 | 3.47 |
| Maff | 5.317573 | 8.760696 | 3.44 |
| Penk | 6.833021 | 10.2276 | 3.39 |
| Akap12 | 7.17374 | 10.51307 | 3.34 |
| Rnd3 | 7.763508 | 11.06458 | 3.30 |
| Il1rap | 6.830734 | 10.12385 | 3.29 |
| Nupr1 | 7.160269 | 10.37722 | 3.22 |
| Bace2 | 6.047527 | 9.25269 | 3.21 |
| Rsad2 | 6.317559 | 9.501802 | 3.18 |
| Ptgs2 | 5.797745 | 8.972202 | 3.17 |
| Aldh1a3 | 6.87675 | 10.044 | 3.17 |
| Atf3 | 8.045691 | 11.1855 | 3.14 |
| Cdh13 | 7.377244 | 10.51443 | 3.14 |
| Lgals1 | 7.972818 | 11.10459 | 3.13 |
| Fosl1 | 6.344384 | 9.40595 | 3.06 |
| Tagln2 | 5.956065 | 8.991003 | 3.03 |
| Phlda1 | 6.358819 | 9.392575 | 3.03 |
| Gap43 | 8.421201 | 11.44139 | 3.02 |

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|---------|----------|----------|------|
| Ifitm3 | 8.424528 | 11.38966 | 2.97 |
| Lif | 6.396519 | 9.344319 | 2.95 |
| Slc7a11 | 7.8706 | 10.81352 | 2.94 |
| Osmr | 8.618423 | 11.55899 | 2.94 |
| Ahnak2 | 5.90714 | 8.836987 | 2.93 |
| Cd44 | 7.835063 | 10.75019 | 2.92 |
| Layn | 5.905998 | 8.788326 | 2.88 |
| Kitl | 5.588583 | 8.422109 | 2.83 |
| Nes | 6.868566 | 9.689172 | 2.82 |
| Trp53 | 6.800496 | 9.614729 | 2.81 |
| Hbegf | 7.970807 | 10.77976 | 2.81 |
| Gas5 | 7.477266 | 10.28387 | 2.81 |
| Egr2 | 7.160723 | 9.940983 | 2.78 |

Supplementary Table 8. Gene expression (Microarray) of genes reported to be expressed in light damaged retinas or MG of light damaged retinas

| Gene Symbol | Description | Undamaged control log2 relative expression | Light Damage log2 relative expression | Fold change light damage versus control |
|--------------------|--|---|--|--|
| Bcl2 | B cell leukemia/lymphoma 2 | 6.964325 | 7.039897 | 0.1 |
| Ctsd | cathepsin D | 11.26174 | 11.74302 | 0.5 |
| Kcnj10 | potassium inwardly-rectifying channel, subfamily J, member 10 | 10.01217 | 10.01217 | 0.0 |
| Ngfr p75 | nerve growth factor receptor (TNFR superfamily, member 16) | 9.180977 | 10.22241 | 1.0 |
| Fgf2 | fibroblast growth factor 2 | 7.965034 | 9.540471 | 1.6 |
| Ntf3 | neurotrophin 3 | 6.209656 | 6.530434 | 0.3 |
| Ntrk3 | neurotrophic tyrosine kinase, receptor, type 3 | 6.347559 | 5.990314 | -0.4 |
| Pcna | proliferating cell nuclear antigen | 10.28271 | 10.64208 | 0.4 |
| Mki67 | antigen identified by monoclonal antibody Ki 67 | 4.799019 | 4.799019 | 0.0 |
| Ccnd1 | cyclin D1 | 9.439362 | 11.09856 | 1.7 |
| Cntf | ciliary neurotrophic factor receptor | 10.44431 | 10.44431 | 0.0 |
| Gdnf | glial cell line derived neurotrophic factor | 6.911917 | 7.776315 | 0.9 |
| Aqp1 | aquaporin 1 | 9.807625 | 7.814171 | -2.0 |
| Fos | FBJ osteosarcoma oncogene | 11.96496 | 12.49366 | 0.5 |
| Nfkb1 | nuclear factor of kappa light polypeptide gene enhancer in B cells 1, p105 | 8.410229 | 9.486856 | 1.1 |

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|-------|--|----------|----------|------|
| Tnf | tumor necrosis factor | 6.7052 | 6.7052 | 0.0 |
| Ccl2 | chemokine (C-C motif) ligand 2 | 6.402283 | 10.11604 | 3.7 |
| Crb1 | crumbs family member 1, photoreceptor morphogenesis associated | 11.92124 | 10.88318 | -1.0 |
| Ace2 | angiotensin I converting enzyme (peptidyl-dipeptidase A) 2 | 4.17806 | 4.17806 | 0.0 |
| Ncan | neurocan | 7.059061 | 7.436745 | 0.4 |
| Vcan | versican | 5.366831 | 7.601713 | 2.2 |
| Bcan | brevican | 6.602848 | 6.602848 | 0.0 |
| Cd44 | CD44 antigen | 7.835063 | 10.75019 | 2.9 |
| Rac1 | RAS-related C3 botulinum substrate 1 | 10.58856 | 11.14308 | 0.6 |
| Rac2 | RAS-related C3 botulinum substrate 2 | 6.715644 | 6.64817 | -0.1 |
| Rac3 | RAS-related C3 botulinum substrate 3 | 6.947626 | 7.163431 | 0.2 |
| Rhoa | ras homolog gene family, member A | 11.11479 | 11.76252 | 0.6 |
| Rhob | ras homolog gene family, member B | 9.169951 | 10.23754 | 0.6 |
| Rhoc | ras homolog gene family, member C | 8.911922 | 9.589165 | 1.1 |
| Gpx1 | glutathione peroxidase 1 | 7.744192 | 8.224051 | 0.5 |
| Il6st | interleukin 6 signal transducer | 10.98721 | 11.81941 | 0.8 |
| Lifr | leukemia inhibitory factor receptor | 9.5482 | 9.000968 | -0.5 |
| Cntfr | ciliary neurotrophic factor receptor | 10.44431 | 10.44431 | 0.0 |
| Stat3 | signal transducer and activator of transcription 3 | 9.444607 | 11.28477 | 1.8 |

| | | | | |
|--------|---|----------|----------|-----|
| Lif | leukemia inhibitory factor | 6.396519 | 9.344319 | 2.9 |
| Mt1 | metallothionein 1 | 10.30838 | 12.41494 | 2.1 |
| Mt2 | metallothionein 2 | 8.264072 | 12.11129 | 3.8 |
| Cp | ceruloplasmin | 12.69996 | 13.53588 | 0.8 |
| Lcn2 | lipocalin 2 | 7.960248 | 12.71421 | 4.8 |
| Chi311 | Chitinase 3 Like 1 | 9.969642 | 10.76591 | 3.7 |
| Bag3 | BCL2-associated athanogene 3 | 8.030416 | 9.648602 | 0.8 |
| Bag1 | BCL2-associated athanogene 1 | 8.359955 | 8.655867 | 1.6 |
| S100a1 | S100 calcium binding protein A1 | 8.86525 | 10.03048 | 0.3 |
| Clic4 | chloride intracellular channel 4 (mitochondrial) | 9.118465 | 10.49558 | 1.2 |
| Clic1 | chloride intracellular channel 1 | 7.760585 | 10.41751 | 1.4 |
| Hdac11 | histone deacetylase 11 | 8.182041 | 7.539239 | 2.7 |

Supplementary Table 9: Fold change of 21 genes in Müller glia after LD and Dicer deletion. *genes with >2.7 (LD) and >1.4 fold increase (cKO) are in bold.

| Gene | Undamaged control log2 relative expression (Microarray) | Light Damage log2 relative expression (Microarray) | Fold change light damage versus control | Wild type (RNA-Seq) log2CPM | Dicer-cKO (RNA-Seq) log2CPM | Fold change cKO versus wild type |
|----------------|---|--|---|-----------------------------|-----------------------------|----------------------------------|
| Maff | 5.31757 | 8.7607 | 3.44 | 6.18847 | 8.76958 | 2.58 |
| Atf3 | 8.04569 | 11.1855 | 3.14 | 6.83319 | 8.98196 | 2.15 |
| Irf1 | 7.89435 | 9.26844 | 1.37 | 6.65689 | 8.79824 | 2.14 |
| Arc | 7.99967 | 9.42568 | 1.43 | 6.93241 | 8.75483 | 1.82 |
| Egr2 | 7.16072 | 9.94098 | 2.78 | 7.02422 | 8.8231 | 1.80 |
| Rrs1 | 7.41351 | 8.87402 | 1.46 | 4.62288 | 6.37159 | 1.75 |
| Gadd45b | 7.56354 | 11.6491 | 4.09 | 4.78525 | 6.4322 | 1.65 |
| Fosl2 | 7.10696 | 8.63265 | 1.53 | 5.19051 | 6.54333 | 1.35 |
| Csrnp1 | 7.04901 | 8.99495 | 1.95 | 6.24975 | 7.43475 | 1.19 |
| Cebpb | 7.8281 | 9.37884 | 1.55 | 4.73147 | 5.78673 | 1.06 |
| Lmna | 7.01591 | 8.57848 | 1.56 | 4.51125 | 5.56106 | 1.05 |
| C4b | 7.77481 | 9.44723 | 1.67 | 4.94628 | 5.9242 | 0.98 |
| Epha2 | 7.82134 | 9.26089 | 1.44 | 5.8779 | 6.7657 | 0.89 |
| Mt2 | 8.26407 | 12.1113 | 3.85 | 6.93133 | 7.81361 | 0.88 |
| Nfkbia | 8.25089 | 10.232 | 1.98 | 5.69714 | 6.54235 | 0.85 |
| Gfap | 6.28272 | 10.5111 | 4.23 | 4.37759 | 5.20687 | 0.83 |

| | | | | | | |
|-------|---------|---------|------|---------|---------|------|
| H2-D1 | 8.34902 | 9.89489 | 1.55 | 5.82285 | 6.62164 | 0.80 |
| Myh9 | 8.34219 | 9.65764 | 1.32 | 6.46277 | 7.24567 | 0.78 |
| Hbegf | 8.97967 | 9.60301 | 2.81 | 4.24778 | 5.01641 | 0.77 |
| Nr4a1 | 7.97081 | 10.7798 | 1.49 | 9.49231 | 10.2245 | 0.73 |
| Sbno2 | 6.99263 | 9.6866 | 2.69 | 5.40557 | 6.11851 | 0.71 |