

## **Modulation of miRNA expression in aged rat hippocampus by buttermilk and krill oil**

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### **Supporting Information**

Additional supporting information may be found in the online version of this article.

**Table S1.** miRNA sequencing results.

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**Table S7.** Sequence of miRNAs primers selected for validation.

**Table S8.** Sequence of genes primers selected for validation.

**Table S1.** miRNA sequencing results

	<b>miRNA</b>	<b>logFC</b>	<b>PValue</b>	<b>FDR</b>
<b>BMFC vs Control</b>	<b>miR-146a-3p</b>	-3.674	0.000	0.013
	<b>miR-3552</b>	-3.389	0.000	0.013
	<b>miR-181a-5p</b>	-2.760	0.001	0.039
	<b>miR-17-1-3p</b>	-2.168	0.001	0.032
	<b>miR-483-3p</b>	-2.083	0.001	0.032
	<b>miR-17-5p</b>	-1.756	0.001	0.033
	<b>miR-20a-5p</b>	-1.575	0.000	0.013
	<b>miR-339-5p</b>	-1.569	0.000	0.013
	<b>miR-99a-3p</b>	-1.501	0.001	0.032
	<b>miR-3102</b>	-1.397	0.000	0.008
	<b>miR-31a-5p</b>	-1.394	0.001	0.033
	<b>miR-582-5p</b>	-1.376	0.001	0.032
	<b>miR-3065-3p</b>	-1.261	0.001	0.035
	<b>miR-369-3p</b>	-1.255	0.002	0.044
	<b>miR-872-3p</b>	-1.238	0.001	0.039
	<b>miR-219a-5p</b>	-1.147	0.001	0.032
	<b>miR-674-5p</b>	-1.004	0.002	0.041
	<b>miR-450a-5p</b>	-0.970	0.001	0.032
	<b>miR-15b-5p</b>	-0.956	0.002	0.044
	<b>miR-181b-5p</b>	-0.932	0.001	0.040
	<b>let-7f-5p</b>	0.822	0.002	0.045
	<b>miR-148a-5p</b>	0.918	0.002	0.047
	<b>miR-195-3p</b>	1.023	0.002	0.042

	<b>miRNA</b>	<b>logFC</b>	<b>PValue</b>	<b>FDR</b>
	<b>miR-17-5p</b>	-2.747	0.001	0.008
	<b>miR-3583-3p</b>	-2.575	0.011	0.045
	<b>miR-362-3p</b>	-2.464	0.002	0.013
	<b>miR-17-1-3p</b>	-2.377	0.007	0.031
	<b>miR-20a-5p</b>	-2.305	0.000	0.000
	<b>miR-449a-5p</b>	-2.287	0.002	0.014
	<b>miR-369-3p</b>	-2.254	0.000	0.006
	<b>miR-362-5p</b>	-2.242	0.001	0.009
	<b>let-7f-2-3p</b>	-2.192	0.002	0.015
	<b>miR-339-5p</b>	-2.154	0.000	0.004
	<b>miR-764-3p</b>	-2.115	0.011	0.043
	<b>miR-107-3p</b>	-2.102	0.000	0.002
	<b>miR-194-5p</b>	-1.922	0.000	0.002
	<b>miR-770-5p</b>	-1.899	0.001	0.009
	<b>miR-144-5p</b>	-1.896	0.006	0.028
	<b>miR-93-3p</b>	-1.876	0.003	0.018
	<b>miR-374-5p</b>	-1.849	0.000	0.001
	<b>miR-582-5p</b>	-1.824	0.000	0.003
	<b>miR-3065-3p</b>	-1.797	0.000	0.002
	<b>miR-137-3p</b>	-1.771	0.000	0.002
	<b>miR-99a-3p</b>	-1.699	0.001	0.008
	<b>miR-106b-5p</b>	-1.696	0.000	0.001
	<b>miR-324-3p</b>	-1.689	0.000	0.005
	<b>miR-539-3p</b>	-1.671	0.000	0.006
	<b>miR-143-5p</b>	-1.592	0.001	0.008
	<b>miR-34c-3p</b>	-1.585	0.003	0.017
	<b>miR-30c-5p</b>	-1.546	0.001	0.007
	<b>miR-494-3p</b>	-1.543	0.000	0.006
	<b>miR-872-3p</b>	-1.522	0.001	0.009

<b>BMFC+KOC vs Control</b>	<b>miR-98-3p</b>	-1.465	0.009	0.038
	<b>miR-31a-5p</b>	-1.457	0.000	0.004
	<b>miR-16-5p</b>	-1.453	0.000	0.002
	<b>miR-126a-5p</b>	-1.408	0.001	0.008
	<b>miR-708-5p</b>	-1.396	0.000	0.005
	<b>miR-410-3p</b>	-1.391	0.000	0.006
	<b>miR-1193-3p</b>	-1.344	0.008	0.033
	<b>miR-425-3p</b>	-1.326	0.002	0.015
	<b>miR-93-5p</b>	-1.272	0.000	0.004
	<b>miR-674-5p</b>	-1.266	0.001	0.008
	<b>miR-376b-3p</b>	-1.238	0.004	0.022
	<b>miR-361-5p</b>	-1.225	0.001	0.009
	<b>miR-181b-5p</b>	-1.222	0.001	0.009
	<b>miR-221-3p</b>	-1.216	0.001	0.008
	<b>miR-219a-5p</b>	-1.200	0.007	0.033
	<b>miR-450a-5p</b>	-1.181	0.001	0.008
	<b>miR-186-5p</b>	-1.177	0.000	0.006
	<b>miR-138-5p</b>	-1.166	0.001	0.009
	<b>miR-497-5p</b>	-1.146	0.007	0.029
	<b>miR-3589</b>	-1.126	0.003	0.019
	<b>let-7e-3p</b>	-1.079	0.001	0.009
	<b>miR-384-5p</b>	-1.073	0.004	0.022
	<b>miR-98-5p</b>	-1.071	0.001	0.009
	<b>miR-15b-5p</b>	-1.048	0.006	0.027
	<b>miR-384-3p</b>	-0.997	0.006	0.028
	<b>miR-874-3p</b>	-0.986	0.004	0.023
	<b>miR-379-3p</b>	-0.970	0.002	0.015
	<b>miR-23a-3p</b>	-0.892	0.006	0.028
	<b>miR-195-5p</b>	-0.891	0.006	0.028
	<b>miR-191a-5p</b>	-0.874	0.003	0.016
	<b>miR-140-5p</b>	-0.863	0.003	0.019
	<b>miR-421-3p</b>	-0.863	0.009	0.038
	<b>miR-26b-5p</b>	-0.848	0.004	0.022
	<b>miR-146a-5p</b>	-0.844	0.004	0.021
	<b>miR-29a-3p</b>	-0.835	0.007	0.033
	<b>miR-543-3p</b>	0.690	0.012	0.047
	<b>miR-455-5p</b>	0.763	0.012	0.047
	<b>miR-495</b>	0.810	0.006	0.028
	<b>let-7i-5p</b>	0.811	0.007	0.030
	<b>miR-541-5p</b>	0.822	0.009	0.038
	<b>miR-134-5p</b>	0.840	0.010	0.041
<b>miR-143-3p</b>	0.884	0.005	0.024	
<b>miR-135a-3p</b>	0.899	0.008	0.033	
<b>miR-125-1-3p</b>	0.899	0.009	0.038	
<b>let-7f-5p</b>	0.905	0.005	0.027	
<b>miR-139-5p</b>	0.923	0.001	0.009	
<b>miR-99a-5p</b>	0.940	0.004	0.021	
<b>miR-361-3p</b>	0.965	0.006	0.028	
<b>miR-185-3p</b>	0.969	0.009	0.038	
<b>miR-99b-5p</b>	0.973	0.010	0.038	
<b>miR-380-5p</b>	0.975	0.005	0.023	
<b>miR-6331</b>	0.976	0.004	0.021	
<b>miR-128-3p</b>	0.981	0.003	0.018	
<b>miR-30c-2-3p</b>	0.995	0.001	0.010	
<b>miR-1843-5p</b>	1.008	0.005	0.023	
<b>miR-106b-3p</b>	1.015	0.003	0.018	
<b>miR-192-5p</b>	1.042	0.001	0.010	
<b>miR-3559-3p</b>	1.060	0.009	0.038	

<b>miR-1843-5p</b>	1.092	0.004	0.022
<b>miR-99b-3p</b>	1.156	0.001	0.008
<b>miR-184</b>	1.180	0.003	0.019
<b>miR-30e-3p</b>	1.208	0.002	0.013
<b>miR-676</b>	1.227	0.000	0.003
<b>miR-344-1-3p</b>	1.236	0.000	0.001
<b>miR-673-3p</b>	1.264	0.000	0.004
<b>miR-770-3p</b>	1.302	0.000	0.005
<b>miR-493-3p</b>	1.320	0.002	0.015
<b>miR-370-3p</b>	1.367	0.001	0.009
<b>miR-485-5p</b>	1.383	0.001	0.010
<b>miR-379-5p</b>	1.393	0.000	0.002
<b>miR-3557-5p</b>	1.422	0.000	0.003
<b>miR-127-3p</b>	1.423	0.000	0.001
<b>miR-129-5p</b>	1.460	0.002	0.015
<b>miR-219-1-3p</b>	1.464	0.002	0.012
<b>miR-150-3p</b>	1.605	0.001	0.007
<b>miR-671</b>	1.615	0.000	0.004
<b>miR-381-3p</b>	1.639	0.000	0.002
<b>miR-674-3p</b>	1.665	0.000	0.001
<b>miR-148a-5p</b>	1.682	0.000	0.001
<b>miR-6329</b>	1.690	0.000	0.001
<b>miR-10a-5p</b>	1.697	0.002	0.012
<b>miR-27a-5p</b>	1.725	0.000	0.002
<b>miR-409b</b>	1.728	0.000	0.001
<b>miR-10b-5p</b>	1.748	0.002	0.012
<b>miR-195-3p</b>	1.829	0.000	0.001
<b>miR-708-3p</b>	1.935	0.000	0.000
<b>miR-6315</b>	1.984	0.000	0.003
<b>miR-151-3p</b>	2.087	0.000	0.004
<b>miR-148a-3p</b>	2.288	0.000	0.000
<b>miR-124-5p</b>	3.505	0.002	0.013

BMFC: buttermilk fat concentrate rich in phospho- and sphingolipids; KOC: krill oil concentrate rich in omega-3 fatty acids (eicosapentaenoic acid and docosahexaenoic acid) and phospholipids; BMFC+KOC: combination of BMFC and KOC.

**Table S2.** Selected miRNAs for validation by RT-qPCR.

	<b>miRNA</b>	<b>logFC</b>	<b>PValue</b>	<b>FDR</b>	<b>Function</b>
<b>BMFC</b>	<b>let-7f-5p</b>	0.822	0.002	0.045	Down-regulation in AD
	<b>miR-195-3p</b>	1.023	0.002	0.042	Role in determining dementia susceptibility
	<b>miR-148a-5p</b>	0.918	0.002	0.047	Inhibition induces hepatocellular tumorigenesis
<b>BMFC+KOC</b>	<b>miR-99a-5p</b>	0.940	0.004	0.021	Neuroprotective effect
	<b>miR-128-3p</b>	0.981	0.003	0.018	Critical for hippocampus-related contextual learning
	<b>miR-148a-3p</b>	2.288	0.000	0.000	Downregulation in AD
	<b>miR-29a-3p</b>	-0.835	0.007	0.033	Neuronal synapse formation and plasticity
	<b>miR-191a-5p</b>	-0.874	0.003	0.016	Target genes to apoptosis and cell death pathways
	<b>miR-381-3p</b>	1.639	0.000	0.002	Linked to insulin resistance or T2D
	<b>miR-379-5p</b>	1.393	0.000	0.002	Predispose to neuropathic pain identified
	<b>miR-30e-3p</b>	1.208	0.002	0.013	Linked to insulin resistance
	<b>miR-146a-5p</b>	-0.844	0.004	0.021	Associated with AD.
	<b>miR-370-3p</b>	1.367	0.001	0.009	Downregulated by Mild Stress in Rat Hippocampal Tissues
	<b>let-7f-5p</b>	0.905	0.005	0.027	Downregulation in AD
	<b>miR-770-3p</b>	1.302	0.000	0.005	Down-regulated in the hippocampus of Sprague-Dawley rats with temporal lobe epilepsy
	<b>miR-106b-3p</b>	1.015	0.003	0.018	Protect mitochondrial functions
	<b>miR-195-3p</b>	1.829	0.000	0.001	Downregulation in AD
	<b>miR-148a-5p</b>	1.682	0.000	0.001	Cancer

BMFC: buttermilk fat concentrate rich in phospho- and sphingolipids; KOC: krill oil concentrate rich in omega-3 fatty acids (eicosapentaenoic acid and docosahexaenoic acid) and phospholipids; BMFC+KOC: combination of BMFC and KOC.

**Table S3** (see Excel file attached to this manuscript supporting information)

**Table S4.** Gene expression microarray results.

	<b>Gene</b>	<b>logFC</b>	<b>P.Value</b>	<b>Adj.P.Val</b>
<b>KOC</b>	<b>Kcnj13</b>	-3	0.0000	0.0315
	<b>Dpt</b>	-2.4	0.0000	0.0275
	<b>Tph1</b>	-2.1	0.0000	0.0000
	<b>Slc35f4</b>	-2	0.0001	0.0452
	<b>Exoc5</b>	-1.8	0.0000	0.0315
	<b>Barx1</b>	-1.7	0.0000	0.0005
	<b>Ippk</b>	-1.5	0.0000	0.0013
	<b>Lilrb3l</b>	-1.4	0.0000	0.0013
	<b>P4hb</b>	-1.3	0.0000	0.0008
	<b>Cenpn</b>	-1.3	0.0001	0.0438
	<b>---</b>	-1.2	0.0000	0.0049
	<b>Enpp2</b>	-1.2	0.0001	0.0380
	<b>Osr1</b>	-1.1	0.0000	0.0275
	<b>Ptgds</b>	-1.1	0.0000	0.0315
	<b>LOC654482</b>	-1.1	0.0001	0.0341
	<b>Slc6a20</b>	-1.1	0.0001	0.0380
	<b>Bcd2</b>	-1	0.0000	0.0226
	<b>Ch25h</b>	-1	0.0000	0.0293
	<b>Arhgef15</b>	-1	0.0001	0.0371
	<b>Sox21</b>	-1	0.0001	0.0452
	<b>Nutm2f</b>	-1	0.0001	0.0452
	<b>Rarres1</b>	-0.9	0.0000	0.0315
	<b>Car13</b>	-0.9	0.0001	0.0332
	<b>Id4</b>	-0.9	0.0001	0.0438
	<b>Efhc1</b>	-0.9	0.0002	0.0490
	<b>Ctsc</b>	-0.8	0.0001	0.0452
	<b>Aox4</b>	-0.8	0.0001	0.0452
	<b>Gpr135</b>	0.8	0.0001	0.0426
	<b>Pygo2</b>	0.8	0.0001	0.0452
	<b>Fam163b</b>	0.8	0.0001	0.0452
	<b>Ptpn4</b>	0.8	0.0001	0.0452
	<b>St6gal2</b>	0.8	0.0002	0.0495
	<b>Pak6</b>	0.9	0.0000	0.0315
<b>Arc</b>	0.9	0.0000	0.0315	
<b>Ramp3</b>	0.9	0.0001	0.0341	
<b>Cckbr</b>	0.9	0.0001	0.0371	
<b>N4bp2</b>	1	0.0000	0.0266	
<b>Yeats2</b>	1	0.0001	0.0438	
<b>BMFC</b>	<b>Kcnj13</b>	-2.8	2.53E-12	2.91E-08
	<b>Dpt</b>	-2.3	8.23E-11	4.74E-07
	<b>Zfp358</b>	-2.1	4.15E-09	6.81E-06
	<b>Cbr3</b>	-2	2.60E-09	4.99E-06

<b>Mfrp</b>	-2	1.51E-08	2.17E-05
<b>Tmem72</b>	-1.9	5.70E-08	7.29E-05
<b>Sostdc1</b>	-1.8	4.66E-10	1.34E-06
<b>Barx1</b>	-1.8	4.56E-10	1.34E-06
<b>Ippk</b>	-1.6	2.29E-09	4.99E-06
<b>Slco1a5</b>	-1.6	1.05E-07	0.00011518
<b>Tph1</b>	-1.6	2.45E-06	0.0011723
<b>Ahsg</b>	-1.5	3.48E-07	0.00026687
<b>Slc35f4</b>	-1.4	1.10E-07	0.00011518
<b>Exoc5</b>	-1.4	1.75E-07	0.00016792
<b>Slc4a5</b>	-1.4	2.40E-07	0.00020184
<b>Slc13a4</b>	-1.4	4.28E-07	0.00029527
<b>Prlr</b>	-1.4	1.45E-05	0.00476127
<b>Ict1</b>	-1.3	2.46E-07	0.00020184
<b>C7</b>	-1.3	4.88E-07	0.00029527
<b>Prr32</b>	-1.3	6.79E-07	0.00039051
<b>Zfp24</b>	-1.3	6.74E-06	0.0025839
<b>Ptgds</b>	-1.2	4.47E-07	0.00029527
<b>---</b>	-1.2	4.70E-07	0.00029527
<b>Slc6a20</b>	-1.2	7.25E-07	0.00039735
<b>Aqp1</b>	-1.2	7.68E-06	0.00276169
<b>Cldn2</b>	-1.2	5.98E-05	0.01465844
<b>Myog</b>	-1.1	1.60E-06	0.00079918
<b>Ctsc</b>	-1.1	3.03E-06	0.00134359
<b>Enpp2</b>	-1.1	3.04E-06	0.00134359
<b>Ch25h</b>	-1.1	3.43E-06	0.00146244
<b>Abca4</b>	-1.1	4.89E-06	0.00200889
<b>Folr1</b>	-1.1	2.45E-05	0.00721838
<b>Cbr1</b>	-1.1	5.20E-05	0.01329863
<b>Emp3</b>	-1	6.23E-06	0.00247225
<b>Bicd2</b>	-1	1.24E-05	0.00430572
<b>LOC654482</b>	-1	1.31E-05	0.0044311
<b>Mospd1</b>	-1	1.67E-05	0.00533843
<b>Lepr</b>	-1	1.77E-05	0.00551727
<b>P4hb</b>	-1	2.09E-05	0.00632722
<b>Atp11c</b>	-1	3.81E-05	0.01086025
<b>Cenpn</b>	-1	4.38E-05	0.01184456
<b>Igf2</b>	-1	6.44E-05	0.01544012
<b>Rpl9</b>	-1	8.19E-05	0.01884002
<b>Sst</b>	-1	8.66E-05	0.01952348
<b>Tmem27</b>	-1	0.000134	0.02854155
<b>Elovl7</b>	-0.9	3.87E-05	0.01086025
<b>Pla2g5</b>	-0.9	4.43E-05	0.01184456
<b>Ranbp3l</b>	-0.9	4.77E-05	0.01247554
<b>Hmgb2l1</b>	-0.9	5.99E-05	0.01465844

	<b>Trim34</b>	-0.9	6.94E-05	0.01628543
	<b>Capn7</b>	-0.9	0.00011415	0.02525007
	<b>Slc2a12</b>	-0.9	0.00013173	0.02854155
	<b>C1qtnf3</b>	-0.9	0.00014208	0.02971226
	<b>Mdk</b>	-0.9	0.00014941	0.03068729
	<b>Slc22a8</b>	-0.8	0.00021362	0.04310616
	<b>Spata20</b>	-0.8	0.00022686	0.04498921
	<b>Arc</b>	1	7.18E-06	0.00266579
	<b>Adss</b>	1.3	1.03E-06	0.00053942
<b>BMFC+KOC</b>	<b>Zfp358</b>	-3.6	0.0000	0.0000
	<b>Kcnj13</b>	-3.9	0.0000	0.0000
	<b>Dpt</b>	-3.7	0.0000	0.0000
	<b>Zfp24</b>	-3.5	0.0000	0.0000
	<b>Cbr3</b>	-3.4	0.0000	0.0000
	<b>Mfrp</b>	-3.2	0.0000	0.0000
	<b>Sostdc1</b>	-2.9	0.0000	0.0000
	<b>Tmem72</b>	-2.9	0.0000	0.0000
	<b>Prlr</b>	-2.6	0.0000	0.0000
	<b>Ahsg</b>	-2.4	0.0000	0.0000
	<b>Exoc5</b>	-2.4	0.0000	0.0000
	<b>Cldn2</b>	-2.4	0.0000	0.0001
	<b>Slc35f4</b>	-2.2	0.0000	0.0000
	<b>Slc4a5</b>	-2.1	0.0000	0.0000
	<b>Folr1</b>	-2	0.0000	0.0000
	<b>Prr32</b>	-2	0.0000	0.0000
	<b>Slco1a5</b>	-2	0.0000	0.0000
	<b>Ict1</b>	-1.9	0.0000	0.0000
	<b>Tmem27</b>	-1.8	0.0000	0.0010
	<b>Sst</b>	-1.6	0.0000	0.0011
	<b>Cenpn</b>	-1.5	0.0000	0.0002
	<b>Aqp1</b>	-1.5	0.0000	0.0007
	<b>Abca4</b>	-1.5	0.0000	0.0007
	<b>Cbr1</b>	-1.5	0.0000	0.0022
	<b>Slc13a4</b>	-1.5	0.0000	0.0117
	<b>Htr2c</b>	-1.4	0.0000	0.0008
	<b>Capn7</b>	-1.3	0.0000	0.0014
	<b>Itpril1</b>	-1.2	0.0000	0.0015
	<b>Glycam1</b>	-1.2	0.0000	0.0015
	<b>Pla2g5</b>	-1.2	0.0000	0.0017
	<b>LOC654482</b>	-1.2	0.0000	0.0022
	<b>Rpl9</b>	-1.2	0.0000	0.0063
	<b>Enpp2</b>	-1.1	0.0000	0.0021
	<b>Spata20</b>	-1.1	0.0000	0.0044
	<b>Itgb6</b>	-1.1	0.0000	0.0057
	<b>Atp11c</b>	-1.1	0.0000	0.0070



<b>Scgb1c1</b>	-1.1	0.0000	0.0077
<b>Aimp2</b>	-1.1	0.0000	0.0098
<b>Mdk</b>	-1.1	0.0001	0.0231
<b>Slc2a12</b>	-1.1	0.0003	0.0475
<b>Efhc1</b>	-1	0.0000	0.0075
<b>Tbc1d10a</b>	-1	0.0000	0.0088
<b>Creg1</b>	-1	0.0001	0.0177
<b>Diaph3</b>	-1	0.0001	0.0239
<b>Narfl</b>	-0.9	0.0000	0.0082
<b>Wdr63</b>	-0.9	0.0000	0.0100
<b>RGD1564074</b>	-0.9	0.0000	0.0111
<b>Glipr2</b>	-0.9	0.0000	0.0117
<b>Fbxo45</b>	-0.9	0.0000	0.0119
<b>LOC100909960</b>	-0.9	0.0001	0.0134
<b>Lama2</b>	-0.9	0.0001	0.0163
<b>Cars2</b>	-0.9	0.0001	0.0264
<b>Olr109</b>	-0.9	0.0001	0.0289
<b>Slc4a2</b>	-0.9	0.0002	0.0316
<b>Wfs1</b>	-0.9	0.0002	0.0365
<b>Muc1911</b>	-0.8	0.0001	0.0239
<b>Igfbpl1</b>	-0.8	0.0002	0.0383
<b>Mast3</b>	-0.8	0.0002	0.0384
<b>---</b>	-0.8	0.0002	0.0415
<b>Fam92b</b>	-0.8	0.0003	0.0473
<b>Six3</b>	-0.8	0.0003	0.0486
<b>LOC100911887</b>	0.8	0.0001	0.0276
<b>Samd15</b>	0.8	0.0002	0.0365
<b>Tnks</b>	0.8	0.0003	0.0465
<b>Sptssb</b>	0.8	0.0003	0.0469
<b>Satb2</b>	0.8	0.0003	0.0489
<b>ATP6</b>	0.8	0.0003	0.0489
<b>MGC116197</b>	0.9	0.0002	0.0342
<b>LOC102550585</b>	1	0.0000	0.0068
<b>Cckbr</b>	1	0.0000	0.0070
<b>LOC102555217</b>	1	0.0000	0.0082
<b>Npas4</b>	1	0.0002	0.0351

BMFC: buttermilk fat concentrate rich in phospho- and sphingolipids; KOC: krill oil concentrate rich in omega-3 fatty acids (eicosapentaenoic acid and docosahexaenoic acid) and phospholipids; BMFC+KOC: combination of BMFC and KOC.

**Table S5** (see Excel file attached to this manuscript supporting information)

**Table S6.** Differently expressed genes from microarray results selected for validation by RT-qPCR.

<b>Gene</b>	<b>miR Interaction</b>	<b>Study group</b>	<b>Gene Function</b>	<b>miRNA Function</b>
<b>Arc</b>	7f-5p	KOC	Neuronal plasticity and memory	Downregulation in AD
<b>Cbr3</b>	191a-5p	BMFC	Parkinson's disease	Target genes to apoptosis and cell death pathways
	106b	BMFC+KOC	Parkinson's disease.	Protect mitochondrial functions
<b>tshz3</b>	148a-5p	BMFC+KOC	Reduced expression of this gene and consequent caspase upregulation may be correlated with progression of Alzheimer's disease	Inhibition induces hepatocellular tumorigenesis
	195-3p			Downregulation in AD
	30e-3p			Linked to insulin resistance
	381-3p			
<b>pla2g5</b>	106b	BMFC+KOC	Increased in several neurological and neurodegenerative disorders	Protect mitochondrial functions
	148a-3p			Downregulation in AD
	195-3p			Downregulation in AD
<b>htr2c</b>	195-3p	BMFC+KOC	Signaling of neurotransmitters	Downregulation in AD
	379-5p			Predispose to neuropathic pain identified
<b>Mrps23</b>	379-5p	KOC	Mitochondrial ribosomal proteins	Predispose to neuropathic pain identified
<b>Capn7</b>	128-3p	BMFC	Calpains are ubiquitous, have been implicated in neurodegenerative processes,	Critical for hippocampus-related contextual learning
	370-3p	KOC		Downregulated by Mild Stress in Rat Hippocampal Tissues
	128-3p	BMFC+KOC		Critical for hippocampus-related contextual learning
	370-3p			Downregulated by Mild Stress in Rat Hippocampal Tissues

	381-3p			Linked to insulin resistance or T2D
<b>Sst</b>	106b	BMFC+KOC	Neurotransmission	Protect mitochondrial functions
<b>Satb2</b>	128-3p	BMFC+KOC	Memory	Critical for hippocampus-related contextual learning
	148a-5p			Inhibition induces hepatocellular tumorigenesis
	381-3p			Linked to insulin resistance or T2D
<b>Ip6k1</b>	148a-5p	BMFC+KOC	Neurotransmission	Inhibition induces hepatocellular tumorigenesis
	770-3p			Down-regulated in the hippocampus of Sprague-Dawley rats with temporal lobe epilepsy
<b>Tph1</b>	379-5p	BMFC & KOC	biosynthesis of serotonin	Identified in primary sensory neurons that are associated with neuropathic pain

Arc: activity-regulated cytoskeleton-associated protein; Cbr3: carbonyl reductase 3; Tshz3: teashirt zinc finger homeobox 3; Pla2g5: phospholipase A2, group V; Htr2c: 5-hydroxytryptamine receptor 2C; Mrps23: mitochondrial ribosomal protein S23; Capn7: calpain 7; Sst: somatostatin; Satb2: SATB homeobox 2; Ip6k1: inositol hexakisphosphate kinase 1; Tph1: tryptophan hydroxylase 1; BMFC: buttermilk fat concentrate rich in phospho- and sphingolipids; KOC: krill oil concentrate rich in omega-3 fatty acids (eicosapentaenoic acid and docosahexaenoic acid) and phospholipids; BMFC+KOC: combination of BMFC and KOC.

**Table S7.** Sequence of miRNAs primers selected for validation.

<b>miRNA</b>	<b>sequence</b>
<i>rno-let-7f-5p</i>	TGAGGTAGTAGATTGTATAGTT
<i>rno-miR-106b-3p</i>	CCGCACTGTGGGTA CT TGCTGC
<i>rno-miR-128-3p</i>	TCACAGTGAACCGGTCTCTTT
<i>rno-miR-146a-5p</i>	TGAGAACTGAATTCCATGGGTT
<i>rno-miR-148a-3p</i>	TCAGTGCACTACAGAACTTTG
<i>rno-miR-148a-5p</i>	AAAGTTCTGAGACACTCTGACTC
<i>rno-miR-191a-5p</i>	CAACGGAATCCCAA AAGCAGCTG
<i>rno-miR-195-3p</i>	CCAATATTGGCTGTGCTGCTCCA
<i>rno-miR-29a-3p</i>	TAGCACCATCTGAAATCGGTTA
<i>rno-miR-30e-3p</i>	CTTTCAGTCGGATGTTTACAGC
<i>rno-miR-370-3p</i>	GCCTGCTGGGGTGGAACCTGGT
<i>rno-miR-381-3p</i>	TATACAAGGGCAAGCTCT
<i>rno-miR-770-3p</i>	GTGGGCCTGACGTGGAG
<i>rno-miR-99a-5p</i>	AACCCGTAGATCCGATCTTGTG
<i>rno-miR-379-5p</i>	TGGTAGACTATGGAACGTAGG

**Table S8.** Sequence of genes primers selected for validation.

<b>Gene</b>	<b>Name</b>	<b>Sequence</b>
<b>Arc</b>	rno-Arc-FW	AAAGCAGCAGCAAGATGGTT
	rno-Arc-REV	GAGTCTTGCCTCCTGTCCTG
<b>Cbr3</b>	rno-Cbr3-FW	GCTGCCATAATGAAACCACA
	rno-Cbr3-REV	GTCTGGCCAACCTTCTCTCT
<b>Tshz3</b>	rno-Tshz3-FW	GCAGCACAGCCATTATCACG
	rno-Tshz3-REV	GGCCAGACTGTTGCTCATCT
<b>Pla2g5</b>	rno-Pla2g5-FW	CTTGGGCTGCCAGCATAAAC
	rno-Pla2g5-REV	GCAGCCGTAGAAGCCATAGT
<b>Htr2c</b>	rno-Htr2c-FW	CGGACGGGGTACAAAACCTGG
	rno-Htr2c-REV	AATCCAGACGGGGCACAAAT
<b>Mrps23</b>	rno-Mrps23-FW	CACATGGTGTGGTTCCTCGG
	rno-Mrps23-REV	CTTAGCCCAACCCGTGACAT
<b>Capn7</b>	rno-Capn7-FW	AGTTCTCCTCGCAGTGCCTC
	rno-Capn7-REV	TTGGGCAGCTTCCTTGAGTA
<b>Sst</b>	rno-Sst-FW	ACCCAGACTCCGTCAGTTT
	rno-Sst -REV	CCAGGGCATCGTTCTCTGTC
<b>Satb2</b>	rno-Satb2-FW	AAAACCTCGACACCGACAAC
	rno-Satb2-REV	CCAACGAAGCAGTTCACAGA
<b>Ip6k1</b>	rno-Ip6k1-FW	ACCAAGGCTGCATCATTTTGAC
	rno-Ip6k1-REV	AAACACACATTGCGTTGGGG

Arc: activity-regulated cytoskeleton-associated protein; Cbr3: carbonyl reductase 3; Tshz3: teashirt zinc finger homeobox 3; Pla2g5: phospholipase A2, group V; Htr2c: 5-hydroxytryptamine receptor 2C; Mrps23: mitochondrial ribosomal protein S23; Capn7: calpain 7; Sst: somatostatin; Satb2: SATB homeobox 2; Ip6k1: inositol hexakisphosphate kinase 1; Tph1: tryptophan hydroxylase 1; FW: Forward; REV: Reverse.