

Supplementary Table S1 | Main, extended data, and supplementary data figures statistics

Experiment	Measure	Statistical Test	Comparison	Statistic (F or t or q or U)	° of freedom	p	*	Fig.	
EEG/EMG recordings	Total time per day in Wake	Unpaired t-test, two-tailed	2:Ctrl vs. 2:AD	2.185	15	0.0452	*	1e	
	Total time per day in NREM			2.197	15	0.0441	*		
	Total time per day in REM			0.446	15	0.6622	-		
	Mean bout duration per day in Wake			0.518	15	0.6119	-		
	Mean bout duration per day in NREM			3.040	15	0.0083	**		
	Mean bout duration per day in REM			2.393	15	0.0303	*		
	Total time per day in Wake		12:Ctrl vs. 12:AD	1.624	14	0.1267	-	1g	
	Total time per day in NREM			3.414	14	0.0042	**		
	Total time per day in REM			1.204	14	0.2487	-		
	Mean bout duration per day in Wake			1.738	14	0.1041	-		
	Mean bout duration per day in NREM			1.487	14	0.1592	-		
	Mean bout duration per day in REM			0.118	14	0.9081	-		
	Delta power (AUC) during NREM		2:Ctrl vs. 2:AD	0.047	15	0.9628	-	1j	
	-		12:Ctrl vs. 12:AD	3.665	14	0.0025	**	1l	
-	Simple linear regression	Hippocampal plaque burden vs. Average NREM time per day	9.220	1, 6	0.0229	*	1o		
-		Cortical plaque burden vs. Average NREM time per day	4.972	1, 6	0.0673	-	1p		
EPM: Sundown (males)	Total open arm time	Two-way ANOVA	Age	49.300	2, 88	<0.0001	****	2d	
			Genotype	9.775	1, 88	0.0024	**		
			Age x Genotype	3.859	2, 88	0.0247	*		
		Tukey's multiple comparisons tests	2:Ctrl vs. 2:AD	0.368	88	0.9998	-		
			2:Ctrl vs. 12:Ctrl	3.029	88	0.2759	-		
			2:Ctrl vs. 12:AD	4.344	88	0.0327	*		
			2:Ctrl vs. 24:Ctrl	6.750	88	0.0001	***		
			2:Ctrl vs. 24:AD	13.770	88	<0.0001	****		
			2:AD vs. 12:Ctrl	2.642	88	0.4286	-		
			2:AD vs. 12:AD	3.977	88	0.0649	-		
			2:AD vs. 24:Ctrl	6.428	88	0.0002	***		
			2:AD vs. 24:AD	13.400	88	<0.0001	****		
			12:Ctrl vs. 12:AD	1.550	88	0.8816	-		
			12:Ctrl vs. 24:Ctrl	4.403	88	0.0291	*		
			12:Ctrl vs. 24:AD	11.480	88	<0.0001	****		
			12:AD vs. 24:Ctrl	2.940	88	0.3078	-		
12:AD vs. 24:AD	9.423	88	<0.0001	****					
24:Ctrl vs. 24:AD	5.325	88	0.0040	**					
EPM: Sundown (females)		Two-way ANOVA	Age	22.550	2, 70	<0.0001	****	2e	
			Genotype	14.920	1, 70	0.0002	***		
			Age x Genotype	5.487	2, 70	0.0061	**		
		Tukey's multiple comparisons tests	2:Ctrl vs. 2:AD	0.291	70	0.8379	-		
			12:Ctrl vs. 12:AD	3.471	70	0.0166	*		
			24:Ctrl vs. 24:AD	5.780	70	0.0001	***		
			2:Ctrl vs. 12:Ctrl	1.081	70	0.7260	-		
			2:Ctrl vs. 24:Ctrl	3.442	70	0.0455	-		
			12:Ctrl vs. 24:Ctrl	2.621	70	0.1600	-		
			2:AD vs. 12:AD	4.556	70	0.0054	**		
2:AD vs. 24:AD	9.933	70	<0.0001	****					
12:AD vs. 24:AD	5.974	70	0.0002	***					
OF: Sundown (males)	Distance traveled	Two-way ANOVA	Age	2.095	2, 49	0.1340	-	2h	
			Genotype	21.270	1, 49	<0.0001	****		
			Age x Genotype	1.860	2, 49	0.1665	-		
		Tukey's multiple comparisons tests (conducted based on trends observed and a priori interest)	2:Ctrl vs. 2:AD	1.796	49	0.2101	-		
			12:Ctrl vs. 12:AD	3.805	49	0.0097	**		
			24:Ctrl vs. 24:AD	6.334	49	<0.0001	****		
			2:Ctrl vs. 12:Ctrl	1.544	49	0.5235	-		
	2:Ctrl vs. 24:Ctrl	1.023	49	0.7509	-				
	12:Ctrl vs. 24:Ctrl	0.626	49	0.8978	-				
	2:AD vs. 12:AD	0.385	49	0.9600	-				
	2:AD vs. 24:AD	2.486	49	0.1946	-				
	12:AD vs. 24:AD	3.249	49	0.0655	-				
	Rearing No.	Two-way ANOVA	Age	4.571	2, 39	0.0165	*		2i
			Genotype	3.428	1, 39	0.0717	-		
Age x Genotype			2.179	2, 39	0.1267	-			
Tukey's multiple comparisons tests (conducted based on trends observed and a priori interest)		2:Ctrl vs. 2:AD	0.494	39	0.7288	-			
		12:Ctrl vs. 12:AD	1.484	39	0.3006	-			
		24:Ctrl vs. 24:AD	4.112	39	0.0060	**			
		2:Ctrl vs. 12:Ctrl	0.029	39	0.9998	-			
2:Ctrl vs. 24:Ctrl	0.887	39	0.8062	-					
12:Ctrl vs. 24:Ctrl	0.937	39	0.7864	-					
2:AD vs. 12:AD	1.894	39	0.3826	-					
2:AD vs. 24:AD	4.824	39	0.0042	**					
12:AD vs. 24:AD	2.559	39	0.1799	-					
OF: Sundown (females)	Distance traveled	Two-way ANOVA	Age	7.067	2, 59	0.0018	**	2j	
			Genotype	7.535	1, 59	0.0080	**		
			Age x Genotype	7.666	2, 59	0.0011	**		
		Tukey's multiple comparisons tests	2:Ctrl vs. 2:AD	0.372	59	0.9998	-		
			2:Ctrl vs. 12:Ctrl	1.984	59	0.7253	-		
			2:Ctrl vs. 12:AD	1.791	59	0.8018	-		
			2:Ctrl vs. 24:Ctrl	0.944	59	0.9848	-		
			2:Ctrl vs. 24:AD	6.177	59	0.0007	***		
			2:AD vs. 12:Ctrl	1.763	59	0.8120	-		
			2:AD vs. 12:AD	1.546	59	0.8821	-		
			2:AD vs. 24:Ctrl	0.596	59	0.9982	-		
			2:AD vs. 24:AD	6.384	59	0.0004	***		
			12:Ctrl vs. 12:AD	0.152	59	>0.9999	-		
	12:Ctrl vs. 24:Ctrl	1.390	59	0.9217	-				
	12:Ctrl vs. 24:AD	5.990	59	0.0011	**				
	12:AD vs. 24:Ctrl	1.136	59	0.9658	-				
	12:AD vs. 24:AD	5.740	59	0.0020	**				
	24:Ctrl vs. 24:AD	6.731	59	0.0002	***				
	Two-way ANOVA	Age	15.280	2, 49	<0.0001	****			
		Genotype	1.716	1, 49	0.1964	-			
Age x Genotype		4.451	2, 49	0.0167	*				
2:Ctrl vs. 2:AD		0.122	49	>0.9999	-				
2:Ctrl vs. 12:Ctrl		0.915	49	0.9867	-				
2:Ctrl vs. 12:AD		0.100	49	>0.9999	-				
2:Ctrl vs. 24:Ctrl		2.526	49	0.4839	-				

	Rearing No.	Tukey's multiple comparisons tests	2:Ctrl vs. 24:AD	5.985	49	0.0013	**	2k
			2:AD vs. 12:Ctrl	1.180	49	0.9596	-	
			2:AD vs. 12:AD	0.252	49	>0.9999	-	
			2:AD vs. 24:Ctrl	2.988	49	0.2979	-	
			2:AD vs. 24:AD	6.731	49	0.0002	***	
			12:Ctrl vs. 12:AD	0.982	49	0.9817	-	
			12:Ctrl vs. 24:Ctrl	2.161	49	0.6482	-	
			12:Ctrl vs. 24:AD	6.522	49	0.0004	***	
			12:AD vs. 24:Ctrl	2.937	49	0.3161	-	
			12:AD vs. 24:AD	6.908	49	0.0002	***	
EPM: Sunrise (males)	Total open arm time	Two-way ANOVA	24:Ctrl vs. 24:AD	4.635	49	0.0224	*	2n
			Age	5.484	2, 56	0.0067	**	
EPM: Sunrise (females)	Total open arm time	Two-way ANOVA	Genotype	1.203	1, 56	0.2773	-	2o
			Age x Genotype	0.731	2, 56	0.4858	-	
OF: Sunrise (males)	Distance traveled	Two-way ANOVA	Age	1.360	2, 26	0.7128	-	2r
			Genotype	0.343	1, 26	0.7468	-	
OF: Sunrise (females)	Distance traveled	Two-way ANOVA	Age x Genotype	0.107	2, 26	0.2743	-	2s
			Age	1.046	2, 37	0.3614	-	
OF: Sunrise (males)	Rearing No.	Two-way ANOVA	Genotype	0.237	1, 37	0.6293	-	2t
			Age x Genotype	2.565	2, 37	0.0905	-	
OF: Sunrise (females)	Rearing No.	Two-way ANOVA	Age	5.245	2, 38	0.0097	**	2u
			Genotype	0.755	1, 38	0.3905	-	
EPM: Sunrise or Sundown (12-month-old females)	Total open arm time	Two-way ANOVA	Age x Genotype	2.012	2, 38	0.1477	-	3b
			Age	2.616	2, 47	0.0837	-	
EPM: Sunrise or Sundown (12-month-old females)	Total open arm time	Šidák's multiple comparisons tests (conducted based on trends observed and a priori interest)	Genotype	0.865	1, 47	0.3570	-	3c
			Age x Genotype	0.907	2, 47	0.4106	-	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Two-way ANOVA	Age	0.065	2, 47	0.9374	-	3d
			Genotype	0.366	1, 47	0.5480	-	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Age x Genotype	0.937	2, 47	0.3990	-	3e
			Period	8.961	1, 91	0.0036	**	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Two-way ANOVA	Genotype	19.020	1, 91	<0.0001	****	3f
			Period x Genotype	1.738	1, 91	0.1907	-	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	2.080	91	0.1518	-	3g
			Sundown:Ctrl vs. Sundown:AD	4.164	91	0.0003	***	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Sunrise:AD vs. Sundown:AD	3.017	91	0.0132	*	3h
			Sunrise:Ctrl vs. Sundown:Ctrl	1.197	91	0.6562	-	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Two-way ANOVA	Period	2.064	1, 39	0.1588	-	3i
			Genotype	5.204	1, 39	0.0281	*	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Period x Genotype	4.317	1, 39	0.0444	*	3j
			Sunrise:Ctrl vs. Sunrise:AD	3.014	39	0.0179	*	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Sundown:Ctrl vs. Sundown:AD	0.147	39	0.9998	-	3k
			Sunrise:AD vs. Sundown:AD	0.431	39	0.9879	-	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Šidák's multiple comparisons tests	Sunrise:Ctrl vs. Sundown:Ctrl	2.625	39	0.0483	*	3l
			Syllable	8.808	18	0.0000	****	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Three-way ANOVA	Genotype	9.092	1	0.0026	**	3m
			Period	12.174	1	0.0005	***	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Three-way ANOVA	Syllable x Genotype	4.245	18	0.0000	****	3n
			Syllable x Period	5.980	18	0.0000	****	
OF: Sunrise or Sundown (12-month-old females)	Distance traveled	Three-way ANOVA	Genotype X Period	5.592	1	0.0183	*	3o
			Syllable x Genotype x Period	2.846	18	0.0001	****	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	77.000	20	0.2789	-	3p
			AD Sunrise vs. AD Sundown	49.000	20	0.5865	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	84.000	20	0.1397	-	3q
			AD Sunrise vs. AD Sundown	83.000	20	0.1495	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	84.000	20	0.1397	-	3r
			AD Sunrise vs. AD Sundown	78.000	20	0.2651	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	57.000	20	0.9479	-	3s
			AD Sunrise vs. AD Sundown	15.000	20	0.0103	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	13.000	20	0.0091	**	3t
			AD Sunrise vs. AD Sundown	19.000	19	0.0275	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	86.000	18	0.0103	*	3u
			AD Sunrise vs. AD Sundown	15.000	20	0.0103	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	17.000	20	0.0103	*	3v
			AD Sunrise vs. AD Sundown	85.000	18	0.0119	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	87.000	18	0.0103	*	3w
			AD Sunrise vs. AD Sundown	33.000	18	0.3920	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	9.000	20	0.0037	**	3x
			AD Sunrise vs. AD Sundown	82.000	18	0.0275	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	9.000	20	0.0037	**	3y
			AD Sunrise vs. AD Sundown	88.000	22	0.3484	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	58.000	22	0.6689	-	3z
			AD Sunrise vs. AD Sundown	85.000	22	0.4022	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	64.000	22	0.9070	-	4a
			AD Sunrise vs. AD Sundown	91.000	22	0.3001	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	65.000	22	0.9070	-	4b
			AD Sunrise vs. AD Sundown	58.000	22	0.6689	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	47.000	22	0.3484	-	4c
			AD Sunrise vs. AD Sundown	28.000	22	0.0575	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	36.000	21	0.2498	-	4d
			AD Sunrise vs. AD Sundown	93.000	19	0.0276	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	36.000	22	0.1512	-	4e
			AD Sunrise vs. AD Sundown	29.000	22	0.0575	-	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	80.000	18	0.0575	-	4f
			AD Sunrise vs. AD Sundown	108.000	21	0.0276	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	32.000	18	0.4022	-	4g
			AD Sunrise vs. AD Sundown	25.000	22	0.0479	*	
KPMS EPM analysis	Normalized syllable frequency	Mann-Whitney U test with FDR correction	AD Sunrise vs. AD Sundown	76.000	19	0.2498	-	4h
			AD Sunrise vs. AD Sundown	21.000	22	0.0276	*	
KPMS EPM analysis	Syllable 10 frequency	Two-way ANOVA	Period	7.875	1, 45	0.0409	*	4i
			Genotype	7.925	1, 45	0.0074	**	
KPMS EPM analysis	Syllable 10 frequency	Two-way ANOVA	Period x Genotype	4.429	1, 45	0.0072	**	4j
			Sunrise:Ctrl vs. Sunrise:AD	0.730	45	0.9548	-	
KPMS EPM analysis	Syllable 10 frequency	Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sundown:Ctrl	0.745	45	0.9522	-	4k
			Sunrise:Ctrl vs. Sundown:AD	5.241	45	0.0031	**	
KPMS EPM analysis	Syllable 10 frequency	Tukey's multiple comparisons tests	Sunrise:AD vs. Sundown:Ctrl	0.010	45	>0.9999	-	4l
			Sunrise:AD vs. Sundown:AD	4.656	45	0.0101	*	
KPMS EPM analysis	Syllable 10 frequency	Tukey's multiple comparisons tests	Sundown:Ctrl vs. Sundown:AD	4.797	45	0.0077	**	4m
			Period	8.528	1, 45	0.0054	**	
KPMS EPM analysis	Syllable 13 frequency	Two-way ANOVA	Genotype	8.018	1, 45	0.0069	**	4n
			Period x Genotype	3.869	1, 45	0.0554	-	
KPMS EPM analysis	Syllable 13 frequency	Two-way ANOVA	Sunrise:Ctrl vs. Sunrise:AD	0.888	45	0.9226	-	4o
			Sunrise:Ctrl vs. Sundown:Ctrl	1.012	45	0.8904	-	
KPMS EPM analysis	Syllable 13 frequency	Tukey's multiple comparisons tests (conducted based on trends observed and a priori interest)	Sunrise:Ctrl vs. Sundown:AD	5.363	45	0.0024	**	4p
			Sunrise:AD vs. Sundown:Ctrl	0.096	45	0.9999	-	
KPMS EPM analysis	Syllable 13 frequency	Tukey's multiple comparisons tests	Sunrise:AD vs. Sundown:AD	4.634	45	0.0106	*	4q
			Sundown:Ctrl vs. Sundown:AD	4.679	45	0.0097	**	
KPMS EPM analysis	Syllable 13 frequency	Two-way ANOVA	Period	8.506	1, 45	0.0055	**	4r
			Genotype	7.405	1, 45	0.0092	**	
KPMS EPM analysis	Syllable 13 frequency	Two-way ANOVA	Period x Genotype	4.181	1, 45	0.0467	*	4s

	Syllable 14 frequency	Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	0.695	45	0.9607	-	3i	
			Sunrise:Ctrl vs. Sundown:Ctrl	0.925	45	0.9135	-		
			Sunrise:Ctrl vs. Sundown:AD	5.256	45	0.0030	**		
			Sunrise:AD vs. Sundown:Ctrl	0.212	45	0.9988	-		
			Sunrise:AD vs. Sundown:AD	4.704	45	0.0092	**		
			Sundown:Ctrl vs. Sundown:AD	4.647	45	0.0103	*		
SCN subpopulation analysis	Core c-Fos+ cells / mm3 (x 1000)	Two-way ANOVA	Period	15.390	1, 21	0.0008	***	4o	
			Genotype	0.051	1, 21	0.8238	-		
			Period x Genotype	2.353	1, 21	0.1399	-		
	Shell c-Fos+ cells / mm3 (x 1000)	Two-way ANOVA	Period	8.719	1, 21	0.0076	**	4p	
			Genotype	2.274	1, 21	0.1465	-		
			Period x Genotype	0.132	1, 21	0.7205	-		
	Core VIP+ cells / mm3 (x 1000)	Two-way ANOVA	Period	0.758	1, 21	0.3937	-	4q	
			Genotype	1.610	1, 21	0.2184	-		
			Period x Genotype	0.823	1, 21	0.3746	-		
	Shell AVP+ cells / mm3 (x 1000)	Two-way ANOVA	Period	0.511	1, 21	0.4825	-	4r	
			Genotype	0.016	1, 21	0.9011	-		
			Period x Genotype	1.281	1, 21	0.2705	-		
	(VIP+c-Fos+)/VIP+	Two-way ANOVA	Period	2.152	1, 21	0.0197	*	4s	
			Genotype	6.370	1, 21	0.2355	-		
			Period x Genotype	1.491	1, 21	0.1572	-		
	(AVP+c-Fos+)/AVP+	Two-way ANOVA	Period	16.340	1, 21	0.0006	***	4t	
			Genotype	8.236	1, 21	0.0092	**		
			Period x Genotype	5.240	1, 21	0.0325	*		
		Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	5.296	21	0.0061	**		
			Sunrise:Ctrl vs. Sundown:Ctrl	5.971	21	0.0020	**		
			Sunrise:Ctrl vs. Sundown:AD	7.095	21	0.0003	***		
			Sunrise:AD vs. Sundown:Ctrl	1.143	21	0.8498	-		
			Sunrise:AD vs. Sundown:AD	1.873	21	0.5584	-		
			Sundown:Ctrl vs. Sundown:AD	0.567	21	0.9777	-		
Period			4.670	1, 12	0.0516	-			
Brain-wide activity mapping at Sundown and Sunrise	Total open arm time	Mixed (Two-way repeated measures) ANOVA	Genotype	5.544	1, 12	0.0364	*	5b	
			Period x Genotype	0.890	1, 12	0.3842	-		
			Subject	0.998	12, 12	0.5014	-		
		Fischer's LSD (conducted based on trends observed and a priori interest)	Sunrise:Ctrl vs. Sunrise:AD	0.997	24	0.3288	-		
			Sundown:Ctrl vs. Sundown:AD	2.331	24	0.0285	*		
			Sunrise:Ctrl vs. Sundown:Ctrl	0.806	12	0.4362	-		
	Sunrise:AD vs. Sundown:AD	2.371	12	0.0353	*				
	All regions (eYFP): absolute correlation coefficient (Pearson's r)	Unpaired t-test, two-tailed	Ctrl vs. AD	36.040	18791	<0.0001	****	5h	
			Cluster 1 regions (eYFP): absolute correlation coefficient (Pearson's r)	6.027	1637	<0.0001	****	5k	
	Piezo-sleep recordings	Amplitude	Two-way ANOVA	Age	8.188	3, 163	<0.0001	****	Extended data figure 1f
Genotype				0.149	1, 163	0.8999	-		
Age x Genotype				4.856	3, 163	0.0029	**		
2:Ctrl vs. 2:AD				4.062	163	0.0046	**		
6:Ctrl vs. 6:AD				1.352	163	0.3403	-		
12:Ctrl vs. 12:AD				3.054	163	0.0323	*		
24:Ctrl vs. 24:AD				1.301	163	0.3588	-		
2:Ctrl vs. 6:Ctrl				0.051	163	>0.9999	-		
2:Ctrl vs. 12:Ctrl				0.320	163	0.9959	-		
2:Ctrl vs. 24:Ctrl				1.781	163	0.5900	-		
6:Ctrl vs. 12:Ctrl				0.283	163	0.9971	-		
6:Ctrl vs. 24:Ctrl				1.822	163	0.5716	-		
12:Ctrl vs. 24:Ctrl			1.599	163	0.6713	-			
2:AD vs. 6:AD			2.659	163	0.2406	*			
2:AD vs. 12:AD			7.283	163	<0.0001	****			
2:AD vs. 24:AD			6.285	163	<0.0001	****			
6:AD vs. 12:AD			4.271	163	0.0154	*			
6:AD vs. 24:AD			3.811	163	0.0386	*			
12:AD vs. 24:AD			0.214	163	0.9988	-			
Acrophase			Two-way ANOVA	2 vs. 6	1.998	163	0.4933	-	
				2 vs. 12	5.502	163	0.0008	***	
				2 vs. 24	5.938	163	0.0003	***	
				6 vs. 12	3.426	163	0.0769	-	
				6 vs. 24	4.121	163	0.0210	*	
		12 vs. 24		1.169	163	0.8418	-		
		Two-way ANOVA	Age	2.284	3, 162	0.0810	-		
			Genotype	0.001	1, 162	0.9698	-		
			Age x Genotype	2.157	3, 162	0.0951	-		
			Age	10.780	3, 164	<0.0001	****		
			Genotype	5.320	1, 164	0.0223	*		
			Age x Genotype	3.775	3, 164	0.0118	*		
Light Cycle Sleep (%)		Two-way ANOVA	2:Ctrl vs. 2:AD	2.437	164	0.0868	-		
			6:Ctrl vs. 6:AD	2.374	164	0.0951	-		
			12:Ctrl vs. 12:AD	3.393	164	0.0176	*		
			24:Ctrl vs. 24:AD	3.147	164	0.0274	*		
			2:Ctrl vs. 6:Ctrl	5.103	164	0.0023	**		
			2:Ctrl vs. 12:Ctrl	0.173	164	0.9993	-		
		Tukey's multiple comparisons tests	2:Ctrl vs. 24:Ctrl	1.021	164	0.8884	-		
			6:Ctrl vs. 12:Ctrl	5.360	164	0.0012	**		
			6:Ctrl vs. 24:Ctrl	4.093	164	0.0222	*		
			12:Ctrl vs. 24:Ctrl	0.926	164	0.9137	-		
			2:AD vs. 6:AD	0.053	164	>0.9999	-		
			2:AD vs. 12:AD	5.466	164	0.0009	***		
			2:AD vs. 24:AD	4.338	164	0.0133	*		
			6:AD vs. 12:AD	5.297	164	0.0014	**		
			6:AD vs. 24:AD	4.258	164	0.0158	*		
			12:AD vs. 24:AD	0.224	164	0.9986	-		
			Tukey's multiple comparisons tests (age only)	2 vs. 6	3.481	164	0.0700	-	
	2 vs. 12			3.845	164	0.0361	*		
2 vs. 24	2.659	164		0.2405	-				
6 vs. 12	7.510	164		<0.0001	****				
6 vs. 24	5.874	164		0.0003	***				
12 vs. 24	0.752	164		0.9513	-				
Dark Cycle Sleep (%)	Two-way ANOVA	Age	4.991	3, 164	0.0024	**	Extended data figure 1i		
		Genotype	5.364	1, 164	0.0218	*			
		Age x Genotype	1.357	3, 164	0.2580	-			
	Tukey's multiple comparisons tests (age only)	2 vs. 6	4.384	164	0.0121	*			
		2 vs. 12	4.654	164	0.0066	**			
		2 vs. 24	4.288	164	0.0148	*			
Two-way ANOVA	6 vs. 12	0.072	164	>0.9999	-				
	6 vs. 24	0.260	164	0.9978	-				
	12 vs. 24	0.204	164	0.9989	-				
	Age	16.070	3, 164	<0.0001	****				
	Genotype	15.340	1, 164	0.0001	***				
	Age x Genotype	4.266	3, 164	0.0062	**				

	Light Cycle Sleep Bout (s)	Tukey's multiple comparisons tests	2:Ctrl vs. 2:AD	0.876	164	0.5367	-	Extended data figure 1j			
			6:Ctrl vs. 6:AD	6.277	164	<0.0001	****				
			12:Ctrl vs. 12:AD	3.022	164	0.0341	*				
			24:Ctrl vs. 24:AD	2.755	164	0.0531	-				
			2:Ctrl vs. 6:Ctrl	3.867	164	0.0347	*				
			2:Ctrl vs. 12:Ctrl	2.090	164	0.4532	-				
			2:Ctrl vs. 24:Ctrl	4.045	164	0.0244	*				
			6:Ctrl vs. 12:Ctrl	6.418	164	<0.0001	****				
			6:Ctrl vs. 24:Ctrl	8.177	164	<0.0001	****				
			12:Ctrl vs. 24:Ctrl	2.247	164	0.3877	-				
	2:AD vs. 6:AD	3.301	164	0.0946	-						
	2:AD vs. 12:AD	5.675	164	0.0005	***						
	2:AD vs. 24:AD	6.889	164	<0.0001	****						
	6:AD vs. 12:AD	2.074	164	0.4600	-						
	6:AD vs. 24:AD	3.845	164	0.0362	*						
	12:AD vs. 24:AD	2.191	164	0.4108	-						
	2 vs. 6	0.171	164	0.9994	-						
	2 vs. 12	5.555	164	0.0007	***						
	2 vs. 24	7.875	164	<0.0001	****						
	6 vs. 12	5.754	164	0.0004	***						
6 vs. 24	8.054	164	<0.0001	****							
12 vs. 24	3.113	164	0.1272	-							
Age	1.608	3, 165	0.1895	-							
Genotype	17.550	1, 165	<0.0001	****							
Age x Genotype	0.834	3, 165	0.4769	-							
Dark Cycle Sleep Bout (s)	Two-way ANOVA						Extended data figure 1k				
EEG/EMG recordings		Unpaired t -test, two-tailed	2:Ctrl vs. 2:AD	Mean no. bouts per day in Wake	2,314	15	0.0353	*	Extended data figure 2c		
				Mean no. bouts per day in NREM	2,428	15	0.0282	*			
				Mean no. bouts per day in REM	2,334	15	0.0339	*			
				Wake-NREM transitions per day	2,451	15	0.0270	*			
				NREM-REM transitions per day	2,426	15	0.0283	*			
				NREM-Wake transitions per day	0,353	15	0.7288	-			
				REM-NREM transitions per day	1,540	15	0.1444	-			
				REM-Wake transitions per day	2,292	15	0.0368	*			
				Time per light cycle in Wake	2,013	15	0.0625	-			
				Time per light cycle in NREM	2,008	15	0.0630	-			
				Time per light cycle in REM	0,900	15	0.3824	-			
				Time per dark cycle in Wake	2,109	15	0.0522	-			
				Time per dark cycle in NREM	2,175	15	0.0461	*			
				Time per dark cycle in REM	0,423	15	0.6780	-			
				Bouts per light cycle in Wake	3,417	15	0.0038	**			
				Bouts per light cycle in NREM	3,347	15	0.0044	**			
				Bouts per light cycle in REM	2,639	15	0.0186	*			
				Bouts per dark cycle in Wake	0,457	15	0.6544	-			
				Bouts per dark cycle in NREM	0,540	15	0.5971	-			
				Bouts per dark cycle in REM	1,242	15	0.2332	-			
				Mean bout duration per light cycle in Wake	0,989	15	0.3383	-			
				Mean bout duration per light cycle in NREM	3,414	15	0.0038	**			
				Mean bout duration per light cycle in REM	2,352	15	0.0327	*			
				Mean bout duration per dark cycle in Wake	1,303	15	0.2122	-			
				Mean bout duration per dark cycle in NREM	2,446	15	0.0273	*			
				Mean bout duration per dark cycle in REM	2,121	15	0.0510	-			
				Mean no. bouts per day in Wake	0,481	14	0.6379	-			
				Mean no. bouts per day in NREM	0,790	14	0.4425	-			
				Mean no. bouts per day in REM	1,004	14	0.3326	-			
				Wake-NREM transitions per day	1,409	14	0.1807	-			
				NREM-REM transitions per day	2,256	14	0.0406	-			
				NREM-Wake transitions per day	0,029	14	0.9789	-			
				REM-NREM transitions per day	0,296	14	0.7719	-			
				REM-Wake transitions per day	2,367	14	0.0329	-			
				Time per light cycle in Wake	0,568	14	0.5794	-			
				Time per light cycle in NREM	2,839	14	0.0131	*			
				Time per light cycle in REM	1,931	14	0.0740	-			
				Time per dark cycle in Wake	2,258	14	0.0405	*			
				Time per dark cycle in NREM	3,593	14	0.0029	**			
				Time per dark cycle in REM	0,106	14	0.9171	-			
				Bouts per light cycle in Wake	0,612	14	0.5501	-			
				Bouts per light cycle in NREM	0,329	14	0.7468	-			
				Bouts per light cycle in REM	1,620	14	0.1276	-			
				Bouts per dark cycle in Wake	1,402	14	0.1826	-			
				Bouts per dark cycle in NREM	1,764	14	0.0995	-			
				Bouts per dark cycle in REM	0,105	14	0.9176	-			
				Mean bout duration per light cycle in Wake	0,006	14	0.9956	-			
				Mean bout duration per light cycle in NREM	1,910	14	0.0788	-			
				Mean bout duration per light cycle in REM	0,161	14	0.8746	-			
				Mean bout duration per dark cycle in Wake	1,899	14	0.0784	-			
				Mean bout duration per dark cycle in NREM	1,055	14	0.3091	-			
				Mean bout duration per dark cycle in REM	0,273	14	0.7890	-			
				Theta power (AUC) during NREM	2:Ctrl vs. 2:AD	1,072	15	0.3009		-	Extended data figure 3c
					12:Ctrl vs. 12:AD	0,921	14	0.3726		-	Extended data figure 3d
				Alpha power (AUC) during NREM	2:Ctrl vs. 2:AD	1,521	15	0.1490		-	Extended data figure 3e
					12:Ctrl vs. 12:AD	5,115	14	0.0002		***	Extended data figure 3f
				Beta power (AUC) during NREM	2:Ctrl vs. 2:AD	1,646	15	0.1206		-	Extended data figure 3g
					12:Ctrl vs. 12:AD	4,624	14	0.0004		***	Extended data figure 3h
				Gamma power (AUC) during NREM	2:Ctrl vs. 2:AD	2,745	15	0.0150		*	Extended data figure 3i
					12:Ctrl vs. 12:AD	1,854	14	0.0849		-	Extended data figure 3j
					2:Ctrl vs. 2:AD	0,072	15	0.9435		-	Extended data figure 3k

	Delta power (AUC) during REM		12:Ctrl vs. 12:AD	1.001	14	0.3337	-	Extended data figure 3l		
	Theta power (AUC) during REM		2:Ctrl vs. 2:AD	0.054	15	0.9577	-	Extended data figure 3m		
	Alpha power (AUC) during REM		12:Ctrl vs. 12:AD	2.252	14	0.0409	*	Extended data figure 3n		
			2:Ctrl vs. 2:AD	2.483	15	0.0254	*	Extended data figure 3o		
	Beta power (AUC) during REM		12:Ctrl vs. 12:AD	5.238	14	0.0001	***	Extended data figure 3p		
			2:Ctrl vs. 2:AD	2.374	15	0.0314	*	Extended data figure 3q		
	Gamma power (AUC) during REM		12:Ctrl vs. 12:AD	5.866	14	<0.0001	****	Extended data figure 3r		
			2:Ctrl vs. 2:AD	1.157	15	0.2654	-	Extended data figure 3s		
EPM: Sundown (males)	Distance Traveled	Two-way ANOVA	Age	1.925	2.89	0.0002	***	Extended data figure 4a		
			Genotype	9.330	1.89	0.5507	-			
		Age x Genotype	0.359	2.89	0.1519	-				
		Tukey's multiple comparisons tests	2 vs. 12	3.068	89	0.0821	-			
			2 vs. 24	3.157	89	0.0713	-			
			12 vs. 24	6.091	89	0.0001	***			
			Age	2.930	2.70	0.0600	-			
		EPM: Sundown (females)	Two-way ANOVA	Genotype	0.008	1.70	0.9273		-	Extended data figure 4b
Age x Genotype	0.711			2.70	0.4945	-				
OF: Sundown (males)	Center Time	Two-way ANOVA	Age	2.813	2.49	0.0697	-	Extended data figure 4d		
			Genotype	0.395	1.49	0.5328	-			
			Age x Genotype	0.606	2.49	0.5494	-			
OF: Sundown (females)	Center Time	Two-way ANOVA	Age	0.803	2.59	0.4527	-	Extended data figure 4f		
			Genotype	5.283	1.59	0.0251	*			
			Age x Genotype	2.288	2.59	0.1104	-			
OF: Sunrise (males)	Center Time	Two-way ANOVA	Age	1.046	2.37	0.3614	-	Extended data figure 4j		
			Genotype	0.237	1.37	0.6293	-			
			Age x Genotype	2.565	2.37	0.0905	-			
OF: Sunrise (females)	Center Time	Two-way ANOVA	Age	2.616	2.47	0.0837	-	Extended data figure 4i		
			Genotype	0.865	1.47	0.3570	-			
			Age x Genotype	0.907	2.47	0.4106	-			
EPM: Sunrise (males)	Distance Traveled	Two-way ANOVA	Age	36.350	2.57	<0.0001	****	Extended data figure 4g		
			Genotype	2.712	1.57	0.1051	-			
		Age x Genotype	3.242	2.57	0.0464	*				
		Tukey's multiple comparisons tests	2:Ctrl vs. 2:AD	1.642	57	0.8532	-			
			2:Ctrl vs. 12:Ctrl	0.800	57	0.9928	-			
			2:Ctrl vs. 12:AD	0.658	57	0.9971	-			
			2:Ctrl vs. 24:Ctrl	6.750	57	0.0002	***			
			2:Ctrl vs. 24:AD	9.473	57	<0.0001	****			
			2:AD vs. 12:Ctrl	0.888	57	0.9885	-			
			2:AD vs. 12:AD	2.183	57	0.6379	-			
			2:AD vs. 24:Ctrl	4.588	57	0.0239	*			
			2:AD vs. 24:AD	7.399	57	<0.0001	****			
			12:Ctrl vs. 12:AD	1.413	57	0.1664	-			
			12:Ctrl vs. 24:Ctrl	5.857	57	0.0015	**			
			12:Ctrl vs. 24:AD	8.864	57	<0.0001	****			
			12:AD vs. 24:Ctrl	7.117	57	<0.0001	****			
			12:AD vs. 24:AD	9.742	57	<0.0001	****			
		24:Ctrl vs. 24:AD	3.466	57	0.1566	-				
EPM: Sunrise (females)	Two-way ANOVA	Age	2.682	2.27	0.0866	-	Extended data figure 4h			
		Genotype	3.490	1.27	0.0726	-				
		Age x Genotype	0.691	2.27	0.5098	-				
OF: Sunrise or Sundown (12-month-old females)	Center Time	Two-way ANOVA	Period	0.344	1.39	0.5611	-	Extended data figure 5b		
			Genotype	0.274	1.39	0.6038	-			
			Period x Genotype	0.656	1.39	0.4227	-			
EPM: Sunrise or Sundown (12-month-old females)	Distance Traveled	Two-way ANOVA	Period	0.692	1.92	0.4075	-	Extended data figure 5d		
			Genotype	7.746	1.92	0.0065	**			
		Period x Genotype	4.615	1.92	0.0343	*				
		Sidák's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	3.348	92	0.0047	**			
			Sundown:Ctrl vs. Sundown:AD	0.469	92	0.9832	-			
			Sunrise:AD vs. Sundown:AD	0.905	92	0.8402	-			
	Sunrise:Ctrl vs. Sundown:Ctrl		2.170	92	0.1240	-				
	Headdips (no.)	Two-way ANOVA	Period	2.606	1.45	0.1134	-			
			Genotype	16.100	1.45	0.0002	***			
			Period x Genotype	1.894	1.45	0.1755	-			
			Syllable 10 frequency vs. Headdips	115.000	1.39	<0.0001	****			
			Syllable 13 frequency vs. Headdips	96.920	1.36	<0.0001	****			
Syllable 14 frequency vs. Headdips			114.900	1.41	<0.0001	****				
KPMS EPM analysis	-	Simple linear regression	Syllable 10 frequency vs. Headdips (high performers removed)	17.880	1.37	0.0001	***	Extended data figure 5i		
			Syllable 13 frequency vs. Headdips (high performers removed)	15.800	1.34	0.0003	***	Extended data figure 5j		
			Syllable 14 frequency vs. Headdips (high performers removed)	15.970	1.39	0.0003	***	Extended data figure 5k		
			Syllable 8 frequency	Two-way ANOVA	Period	9.408	1.45	0.0037	**	Extended data figure 5l
					Genotype	2.840	1.45	0.0989	-	
				Period x Genotype	9.403	1.45	0.0037	**		
	Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD		1.419	45	0.7484	-			
		Sunrise:Ctrl vs. Sundown:Ctrl		0.001	45	>0.9999	-			
		Sunrise:Ctrl vs. Sundown:AD		4.431	45	0.0155	*			
		Sunrise:AD vs. Sundown:Ctrl	1.499	45	0.7151	-				
	Sunrise:AD vs. Sundown:AD	5.815	45	0.0009	***					
	Sundown:Ctrl vs. Sundown:AD	4.633	45	0.0106	*					
	Syllable 12 frequency	Two-way ANOVA	Period	2.639	1.45	0.1112	-	Extended data figure 5m		
			Genotype	2.100	1.45	0.1543	-			
		Period x Genotype	4.557	1.45	0.0383	*				
		Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	0.704	45	0.9591	-			
			Sunrise:Ctrl vs. Sundown:Ctrl	0.542	45	0.9807	-			
			Sunrise:Ctrl vs. Sundown:AD	2.866	45	0.1937	-			
Sunrise:AD vs. Sundown:Ctrl	0.191		45	0.9991	-					
Sunrise:AD vs. Sundown:AD	3.564	45	0.0702	-						
Sundown:Ctrl vs. Sundown:AD	3.494	45	0.0784	-						
Syllable 16 frequency	Two-way ANOVA	Period	17.460	1.45	0.0001	***	Extended data figure 5m			
		Genotype	1.894	1.45	0.1756	-				
	Period x Genotype	6.476	1.45	0.0144	*					
	Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	1.200	45	0.8309	-				
		Sunrise:Ctrl vs. Sundown:Ctrl	1.734	45	0.6136	-				
		Sunrise:Ctrl vs. Sundown:AD	5.179	45	0.0035	**				
Sunrise:AD vs. Sundown:Ctrl		3.041	45	0.1532	-					

	Syllable 18 frequency	Two-way ANOVA	Sunrise:AD vs. Sundown:AD	6.375	45	0.0003	***	Extended data figure 5o	
			Sundown:Ctrl vs. Sundown:AD	3.823	45	0.0459	*		
			Period	9.726	1, 45	0.0032	**		
			Genotype	3.675	1, 45	0.0616	-		
			Period x Genotype	4.245	1, 45	0.0452	*		
		Tukey's multiple comparisons tests	Sunrise:Ctrl vs. Sunrise:AD	0.147	45	0.9996	-		
			Sunrise:Ctrl vs. Sundown:Ctrl	1.123	45	0.8566	-		
			Sunrise:Ctrl vs. Sundown:AD	4.695	45	0.0094	**		
			Sunrise:AD vs. Sundown:Ctrl	1.304	45	0.7933	-		
			Sunrise:AD vs. Sundown:AD	4.910	45	0.0061	**		
			Sundown:Ctrl vs. Sundown:AD	3.878	45	0.0418	*		
PLSC: sundown-active (eYFP+) ensembles	Fixed Inertia	Permutation analysis	Overall relationship between behavioral and brain variables	112.765	-	0.0002	***	Extended data figure 6b, c	
			Relationship between behavioral and brain variables for latent dimension	80.457	-	0.0001	***		
			1st eigenvalue	17.418	-	0.8888	-		
			2nd eigenvalue	5.245	-	1.0000	-		
			3rd eigenvalue	5.199	-	1.0000	-		
			4th eigenvalue	2.663	-	1.0000	-		
			5th eigenvalue	0.860	-	1.0000	-		
			6th eigenvalue	0.528	-	1.0000	-		
			7th eigenvalue	0.212	-	1.0000	-		
			8th eigenvalue	0.105	-	1.0000	-		
			9th eigenvalue	0.075	-	0.9996	-		
			10th eigenvalue	0.004	-	0.9982	-		
PLSC: sunrise-active (c-Fos+) ensembles	Fixed Inertia	Permutation analysis	Overall relationship between behavioral and brain variables	145.655	-	0.0007	***	Extended data figure 6g, h	
			Relationship between behavioral and brain variables for latent dimension	90.173	-	0.0001	***		
			1st eigenvalue	21.787	-	0.9562	-		
			2nd eigenvalue	12.749	-	1.0000	-		
			3rd eigenvalue	10.366	-	0.9992	-		
			4th eigenvalue	5.586	-	1.0000	-		
			5th eigenvalue	2.631	-	1.0000	-		
			6th eigenvalue	1.290	-	1.0000	-		
			7th eigenvalue	0.460	-	1.0000	-		
			8th eigenvalue	0.310	-	1.0000	-		
			9th eigenvalue	0.290	-	0.9834	-		
			10th eigenvalue	0.011	-	0.9946	-		
Brain-wide activity mapping at Sundown and Sunrise	Sensorimotor regions (eYFP ⁺): absolute correlation coefficient (Pearson's r)	Unpaired t-test, two-tailed	Ctrl vs. AD	7.514	208	<0.0001	****	Extended data figure 7c	
	All regions (c-Fos ⁺): absolute correlation coefficient (Pearson's r)			9.138	26350	<0.0001	****	Extended data figure 7h	
	Sensorimotor regions (c-Fos ⁺): absolute correlation coefficient (Pearson's r)			0.308	208	0.7585	-	Extended data figure 7k	
EEG/EMG recordings	-	Simple linear regression	Hippocampal plaque burden vs. NREM delta power	0.004	1, 6	0.9519	-	Supplementary figure 1d	
	-		Cortical plaque burden vs. NREM delta power	0.037	1, 6	0.8543	-	Supplementary figure 1e	
Brexpirazole behavioral testing	OF distance traveled	Two-way ANOVA	Genotype	2.834	1, 19	0.1086	-	Supplementary figure 2b	
			Dose	9.092	1, 19	0.0071	**		
			Genotype x Dose	4.685	1, 19	0.0434	*		
		Tukey's multiple comparisons tests	Veh Ctrl vs. Veh AD	5.095	19	0.0094	**		
			Veh Ctrl vs. Brex Ctrl	0.453	19	0.9883	-		
			Veh Ctrl vs. Brex AD	1.355	19	0.7744	-		
			Veh AD vs. Brex Ctrl	4.622	19	0.0194	*		
	Veh AD vs. Brex AD	4.119	19	0.0407	*				
	Brex Ctrl vs Brex AD	0.866	19	0.9270	-				
	EPM open arms time	Two-way ANOVA	Genotype	2.834	1, 19	0.1086	-		Supplementary figure 2c
			Dose	9.092	1, 19	0.0071	**		
			Genotype x Dose	4.685	1, 19	0.0434	*		
		Tukey's multiple comparisons tests	Veh Ctrl vs. Veh AD	5.095	19	0.0094	**		
			Veh Ctrl vs. Brex Ctrl	0.453	19	0.9883	-		
Veh Ctrl vs. Brex AD			1.355	19	0.7744	-			
Veh AD vs. Brex Ctrl			4.622	19	0.0194	*			
Veh AD vs. Brex AD	4.119	19	0.0407	*					
Brex Ctrl vs Brex AD	0.866	19	0.9270	-					