

**Figure S1. Resting membrane potential of 12 PD neurons does not change after 8 hr. of impalement with two microelectrodes, Related to Figure 2.** Each line shows the membrane potential measurement of a single neuron at the start of the experiment in TTX (t=0) and then after 8 hr. of two-electrode voltage clamp (t=8). *P* value generated using a two-tailed paired t test.

Activity-Dependent Correlations	<u>p values</u>			<u>Pearson's r</u>		
Same slope (Figure 3A)	Control	Silent	TEVC	Control	Silent	TEVC
IH vs. BKKCa	0.003	0.280	0.001	0.805	0.358	0.836
IRK vs. CaV2	0.022	0.717	0.005	0.678	-0.132	0.746
NALCN vs. CaV2	0.006	0.804	0.002	0.737	0.090	0.801
IH vs. SKKCa	0.033	0.142	0.013	0.643	0.473	0.691
Shaw1 vs. Shaker	<0.0001	0.301	<0.0001	0.879	0.344	0.933
Shaw2 vs. CaV2	0.001	0.610	0.041	0.849	-0.198	0.596
IRK vs. NALCN	0.015	0.153	0.045	0.705	-0.462	0.586
Shaw2 vs. IRK	0.001	0.630	0.041	0.864	-0.174	0.596
Shaw2 vs. Shaw1	0.019	0.478	0.015	0.691	0.254	0.680
SKKCa vs. Shaker	0.003	0.068	0.001	0.772	0.569	0.822
Shal vs. IH	0.007	0.070	0.001	0.790	0.566	0.846
Shab vs. NALCN	0.036	0.255	0.012	0.607	0.375	0.696
Shaw2 vs. Shaker	0.041	0.315	0.003	0.622	-0.354	0.780
BKKCa vs. SKKCa	0.033	0.127	0.003	0.615	0.490	0.801
Different slope (Figure 3B)						
NALCN vs. Shaker	<0.0001	0.059	0.009	0.871	0.585	0.712
NALCN vs. SKKCa	0.001	0.182	0.021	0.838	0.434	0.654
Shaker vs. IRK	0.019	0.942	0.003	0.691	-0.025	0.769
Shaw1 vs. IRK	0.007	0.456	0.001	0.758	0.251	0.821
Shaw1 vs. CaV2	0.015	0.417	<0.0001	0.855	0.289	0.927
NALCN vs. Shaw1	<0.0001	0.831	0.003	0.843	-0.073	0.772
Shaker vs. CaV2	0.018	0.102	<0.0001	0.666	0.547	0.918

Table S1. Correlated patterns of gene expression that arise due to voltage activity of the membrane, Related to Figure 2 and Figure 3.

Feedback-Independent	p			Dooroor	o'o r		
Correlations (Figure 202)	values	011	Pearson's r		157		TEVO
	Control	Silent	TEVC	Control	Sile	nt	IEVC
BKKCa vs. NaV	0.028	8 0.033	0.04	1 0.6	86 0	.643	0.652
Shab vs. CaV2	0.01	1 0.026	s <0.000	1 0.7	01 0	.695	0.932
NaV vs. Shal	0.014	4 0.010	0.02	3 0.7	75 0	.735	0.673
Shab vs. SKKCa	0.02	1 0.010	0.002	2 0.6	53 0	.737	0.786
Shaw1 vs. SKKCa	0.004	4 0.027	0.002	2 0.7	56 0	.660	0.792
Shaw1 vs. Shab	0.00	1 0.006	s <0.000	1 0.8	20 0	.764	0.923
Shal vs. BKKCa	0.024	4 0.024	0.00	7 0.6	72 0	.671	0.757
Shab vs. Shaker	0.00	1 0.030	< 0.000	1 0.8	14 0	.650	0.922
Activity-Independent Correlations	p values			Pearson's	r		
	, Control	Silent	TEVC	Control	Silent	TEV	2
Same Trend as Figure 2C <sub>3</sub>							
CaV2 vs. CaV1	<0.0001	0.376	0.234	0.857	0.315	0.3	391
IRK vs. CaV1	0.004	0.793	0.310	0.783	-0.090	0.3	338
Shaw1 vs CaV1	0.015	0.071	0.549	0.680	0.564	0.2	203
Shaw2 vs. CaV1	<0.0001	0.840	0.995	0.945	0.074	0.0	003
Activity-Restricted Correlations	p values		Pearson's r				
	Control	Silent	TEVC	Control	Silent	TEVC	;
Same Trend as Figure 2C <sub>4</sub>							
SKKCa vs. CaV1	0.418	<0.0001	0.816	0.258	0.894	-0.0	80
NaV vs. CaV1	0.186	0.013	0.696	0.455	0.716	-0.1	42
Shab vs CaV1	0.177	0.036	0.555	0.417	0.635	0.2	200
NaV vs. IH	0.378	0.014	0.302	0.335	0.710	0.3	42
Shal vs. CaV1	0.765	0.041	0.286	-0.102	0.622	-0.3	54

Table S2. Correlated patterns of gene expression that arise as a result of factors other than the naturalistic waveform activity of PD, Related to Figure 2. *Feedback-Independent Correlations:* Correlated patterns of gene expression that arise independent of voltage activity of the membrane and chemical feedback. *Activity-Independent Correlations:* Correlated relationships that are present in the control group, but eliminated in both TTX (silent) and TTX+Activity (TEVC) groups. *Activity-Restricted Correlations:* Correlated relationships that are present only when activity was suppressed.

Never Correlated <u>TTX + TEVC</u>	
BKKCa vs. CaV1Shab vs. BKKCaCaVBKKCa vs. Shaw 2NaV vs. CaV2IRKIH vs. CaV1Shal vs. NALCNNALNACLN vs. CaV1SKKCa vs. NALCNShaShaker vs. CaV1SKKCa vs. ShalShaIH vs. CaV2Shaker vs. ShalShaIH vs. CaV2NaV vs. SKKCaShaIH vs. CaV2Shaker vs. NaVNaVShal vs. CaV2Shaker vs. NaVShaIH vs. IRKShab vs. NaVShaIH vs. ShabShaker vs. NaVShaIH vs. ShakerShaker vs. NaVShaIH vs. ShakerShaker vs. NaVShaIH vs. ShakerShaker vs. NaVShaIH vs. ShakerShaker vs. NaVShaIH vs. ShakerShaShaIH vs. ShakerShaShaIH vs. ShakerShaShaIH vs. ShakerShaShaIH vs. ShakerShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaShaIH vs. Shaw1ShaIH vs. Shaw1ShaIH vs. Shaw1ShaIH vs. ShaShaIH v	V2 vs. BKKCa K vs. BKKCa LCN vs. BKKCa aker vs. BKKCa aw1 vs. BKKCa KCa vs. CaV2 ab vs. IRK V vs. NALCN aw1 vs. Shal aw2 vs. Shal aw1 vs. NaV aw2 vs. NaV
IH vs. Shaker Sha	aw1 vs. NaV
IH vs. Shaw1ShaIH vs. Shaw2Sha	aw2 vs. NaV aw2 vs. Shab
Shal vs. IRK SKKCa vs. IRK NaV vs. IRK	
Shaw2 vs. BKKCa	
Shab vs. Shal Shaw2 vs. SKKCa	

Table S3. Channel pairs that were never correlated, became with either TTX or TTX+TEVC, or were only correlated with TTX+TEVC, Related to Figure 2.

Became correlated with TTX and <u>TTX + TEVC</u>	Significant difference between slopes?	ANCOVA p value		
Shah ya DKKOa	Na	0.000		
Shad vs. BKKCa	INO	0.299		
NaV vs. CaV2	Yes	0.044		
Shal vs. NALCN	No	0.560		
SKKCa vs. Shal	Yes	0.021		
Shaker vs. Shal	No	0.109		
NaV vs. SKKCa	No	0.225		
Shab vs. NaV	No	0.227		
Shaker vs. NaV	No	0.146		

Table S4. ANCOVA results for correlations that emerged after loss of inputs with or without activity, Related to Figure 2.

Target	Forward Primer	Reverse Primer	Probe	Labeling	Efficiency
NALCN	TCGCTTCCACGGTGTACATTC	GCGGTGCCTTTGTTCTCAG	TCTTCGTCTTCCTTGGCTGCATGA	FAM-BHQ1	97.1
IRK	TACAGTGGCGTTGGACTCTAC	TCCACCACACCAAGGCAAATAG	TCGTGTTCGCTATGTCATTCATCAGC	CAL Fluor Red 610-BHQ2	93.9
IH	TCGGTGCCACTAGACTACATC	GACCCGCGTGGAGAATCTG	TCCTCATCTTCAACCAGGACTTCAGC	Quasar 670-BHQ2	95.9
SKKCa	GCATCGGAGCATTGAACAGAA	GCCCGGACAGATAGTCATCAG	CAACTTCAACACTCGGTTTGTCCTCAA	Quasar 705-BHQ2	96.1
Shaw1	CGCGTCACTCCTCAGGACTT	CCCAGCACCAGGAAGAACAC	TGATACAGACTTTCCGTGCATCCGC	Quasar 705-BHQ2	87.4
Shaw2	GAACGCCATCAAGCACTATCATC	ATGGCGCCCGACAGCTTAG	TGGCTTGAAGGACGGTCTCACA	Quasar 670-BHQ2	91.5
CbNaV	TCAACGGGAGGTACCATAAGTG	TCGCTGTTCACCCAAGAGTAG	CGGAGGGATTGAAGCTCAACGCA	610-BHQ2	81
Shaker	GAGGCTCAGAAGACCAGTCAAC	TGGCGATATCACCGAGCTCAT	CACTCGATGTCTTCGCGGAGGAGAT	540-BHQ1	90.7
Shab	GAGCCGGACAGACAGGAAC	TGCGCCTCCTTCTGTAGTC	AAGAACCACGAACACCACATGGGTC	FAM-BHQ1	93.5
CaV1	CCAGGCCTTCTACTGGCTCATT	GCTGGCGATAGTGCTCACTG	TGTGCTCGTCTTCCTCAACACGG	FAM-BHQ1	96
CaV2	ATCCGGCGGACAGTAAAGC	GTTCGGCAGCAACACAAAC	TGGTTCTACTGGTTCGTCATCATACTTGT	540-BHQ1	97.9
Shal	GACACCACCTTCACCTCCATTC	GAACCATGTCGCCGTATCCTA	CGGCGTTTTGGTACACCATTGTCAC	Quasar 670-BHQ2	110.7
BKKCa	GCTCAAACTCGGCTTCATTG	CTGCGTGTCTGGAGAAGTTT	AGAATCCCGGCGCTAAACATGACT	Quasar 705-BHQ2	96
vGluT	GCGTTCGTGGACCTTCTAC	TCAGCCACCCTGTAATGGAA	ATCACAGCCAACCTACTTCAGCGAG	CAL Fluor Gold 540-BHQ1	95.3
ChAT	GGACCGCCTGGCTAAGTAC	TCGCGGAGTCCCATAAGG	AGGCGGCGCTCAAGCTTCAGAC	610-BHQ2	93.1
vAChT	GCGTCAGCTGCTTCTTCCT	CAGCAGTGCCGTGTCTATGAG	TTCGCCAGCAACTACTGGGTGTT	Quasar 670-BHQ2	92.8
ACHE	GGGCAACATGGGCATGTAC	GGTCACCACCGAAGAATTCAATG		Quasar 705-BHQ2	94.2

Table S5. Primers and probes used for multiplex qRT-PCR, Related to STAR Methods.Each box in the table represents a group of genes that were multiplexed in a single PCR run.