

## SUPPLEMENTARY TABLES

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**Suppl. Table 1.** Primers developed for this work

Polymorphism	Allele-specific forward primers	Reverse primer
rs1997794	AACTCGAACTCCCTGGCCCT	CCTGGAAACGCATCAAAAACCTG
	AACTCGAACTCCCTGGTCCC	
rs2235751	AGCCCTTTCTAGTTGCCGGGA	CAGTGCTGTCTCAGTGCAGGA
	AGCCCTTTCTAGTTGCCCGGG	
rs6045819	TAGCGTTTGACAGGTCATCA	TCCCACGCAGAAGAGAGATA a
	AGCGTTTGACAGGTCATCG	

a: from Yuferov *et al.* (2009)

**Suppl. Table 2.** Allele frequencies for the populations extracted from 1000 Genomes Project

Population	rs1997794 (C)	rs2235751 (A)	rs6045819 (A)	rs10485703 (T)	rs910080 (T)	rs910079 (T)	rs2235749 (C)	rs3808627 (G)	rs6985606 (G)	(n)
JPT	0.798	0.226	1.000	1.000	0.255	0.255	0.255	0.625	0.668	104
MXL	0.359	0.719	0.914	0.914	0.695	0.688	0.688	0.703	0.633	64
CEU	0.343	0.737	0.904	0.914	0.763	0.758	0.758	0.788	0.515	99
YRI	0.875	0.269	0.611	0.806	0.537	0.523	0.523	0.991	1.000	108

References: JPT = Japanese in Tokyo; MXL = Mexican ancestry from Los Angeles, California USA; CEU = Residents of Utah with North and Western European ancestry; YRI = Yoruba in Ibadian, Nigeria.




Suppl. Table 3. Genotype frequencies of the polymorphisms for the four analysed populations

Gene	Polymorphism	Genotype	CABA	La Plata	Resistencia	MNP
<i>PDYN</i>	rs35286281	339/339	0.198	0.061	0.115	0.170
		339/407	0.321	0.394	0.438	0.528
		339/475	0	0	0.010	0
		407/407	0.434	0.455	0.406	0.283
		407/475	0.028	0.030	0.021	0
		271/339	0	0.030	0	0.019
		271/407	0.019	0.030	0.010	0
			<i>n</i> (106)	<i>n</i> (33)	<i>n</i> (95)	<i>n</i> (53)
	rs1997794	C/C	0.236	0.212	0.156	0.185
		C/T	0.368	0.394	0.458	0.444
		T/T	0.396	0.394	0.385	0.370
			<i>n</i> (106)	<i>n</i> (33)	<i>n</i> (95)	<i>n</i> (54)
	rs2235751	A/A	0.544	0.485	0.531	0.453
		A/G	0.262	0.394	0.375	0.434
		G/G	0.194	0.121	0.094	0.113
			<i>n</i> (103)	<i>n</i> (33)	<i>n</i> (96)	<i>n</i> (53)
	rs6045819	A/A	0.774	0.818	0.813	0.882
		A/G	0.198	0.121	0.187	0.118
		G/G	0.028	0.061	0	0
			<i>n</i> (106)	<i>n</i> (33)	<i>n</i> (96)	<i>n</i> (51)
	rs10485703	C/T	0.146	0.094	0.172	0.135
		T/T	0.854	0.906	0.828	0.865
			<i>n</i> (82)	<i>n</i> (32)	<i>n</i> (57)	<i>n</i> (52)
	rs910080	C/C	0.172	0.031	0.040	0.140
		C/T	0.414	0.469	0.507	0.560
		T/T	0.414	0.500	0.453	0.300
			<i>n</i> (87)	<i>n</i> (32)	<i>n</i> (70)	<i>n</i> (50)
	rs910079	C/C	0.172	0.063	0.039	0.135
C/T		0.403	0.406	0.500	0.538	
T/T		0.425	0.531	0.461	0.327	
		<i>n</i> (87)	<i>n</i> (32)	<i>n</i> (73)	<i>n</i> (52)	
rs2235749	C/C	0.414	0.516	0.429	0.333	
	C/T	0.414	0.452	0.529	0.542	
	T/T	0.172	0.032	0.042	0.125	
		<i>n</i> (87)	<i>n</i> (32)	<i>n</i> (70)	<i>n</i> (48)	
<i>OPRK1</i>	rs35566036	del/del	0.560	0.438	0.740 <sup>a</sup>	0.407
		in/del	0.380	0.438	0.240 <sup>a</sup>	0.519
		in/in	0.060	0.124	0.020 <sup>a</sup>	0.074
			<i>n</i> (100)	<i>n</i> (32)	<i>n</i> (96)	<i>n</i> (54)
	rs3808627	A/A	0.147	0.152	0.045 <sup>a</sup>	0.120
		A/G	0.306	0.212	0.334 <sup>a</sup>	0.380
		G/G	0.547	0.636	0.621 <sup>a</sup>	0.500
			<i>n</i> (95)	<i>n</i> (33)	<i>n</i> (96)	<i>n</i> (50)
	rs6985606	A/A	0.120	0.240	0.069 <sup>a</sup>	0.038
		A/G	0.420	0.209	0.387 <sup>a</sup>	0.396
		G/G	0.460	0.551	0.544 <sup>a</sup>	0.566
			<i>n</i> (100)	<i>n</i> (33)	<i>n</i> (96)	<i>n</i> (53)

CABA= Ciudad Autónoma de Buenos Aires; MNP= Misión Nueva Pompeya  
a: data from Raggio et al. (2018)

**Suppl. Table 4.** Linkage Disequilibrium for PDYN and OPRK in CABA, La Plata (LP), Resistencia (RES) and Misión Nueva Pompeya (MNP) respectively. Above the diagonal D' values, below the diagonal r values. Values with  $p < 0.01$  are highlighted in red, P values between 0.01 and 0.05 are highlighted in orange and non significant P values are highlighted in beige.

CABA-PDYN	rs35286281	rs1997794	rs2235751	rs6045819	rs10485703	rs910080	rs910079	rs2235749
rs35286281	*	0.6881	0.6433	0.7678	0.3661	0.4785	0.5291	0.5291
rs1997794	0.6171	*	0.7059	0.5465	0.7905	0.7219	0.7562	0.7562
rs2235751	0.5854	0.5761	*	0.9987	0.9981	0.4959	0.5578	0.5578
rs6045819	-0.2238	0.2454	-0.2649	*	0.915	0.543	0.5178	0.5178
rs10485703	-0.0785	0.2611	-0.1947	0.673	*	0.999	0.999	0.999
rs910080	0.4728	0.6553	0.4459	0.2686	0.3635	*	0.975	0.975
rs910079	0.5164	0.695	0.4954	0.253	0.3591	0.9631	*	0.9997
rs2235749	0.5164	0.695	0.4954	0.253	0.3591	0.9631	0.9997	*

 P<0.01  
 0.01<P<0.05  
 p>0.05

LP-PDYN	rs35286281	rs1997794	rs2235751	rs6045819	rs10485703	rs910080	rs910079	rs2235749
rs35286281	*	0.9048	0.5881	0.3295	0.2725	0.5611	0.5407	0.5989
rs1997794	0.717	*	0.8176	0.7529	0.9982	0.8994	0.8954	0.8741
rs2235751	0.5677	0.6712	*	0.1205	0.2086	0.5557	0.5338	0.4486
rs6045819	0.1856	0.3361	0.0655	*	0.9988	0.3952	0.3952	0.41
rs10485703	0.0917	0.2661	0.0677	0.5964	*	0.9986	0.9986	0.9986
rs910080	0.5118	0.6501	0.4892	0.244	0.3682	*	0.833	0.7988
rs910079	0.4931	0.6472	0.4699	0.244	0.3682	0.833	*	0.997
rs2235749	0.5356	0.6196	0.3873	0.2582	0.3755	0.7833	0.9777	*

RES-PDYN	rs35286281	rs1997794	rs2235751	rs6045819	rs10485703	rs910080	rs910079	rs2235749
rs35286281	*	0.7104	0.8159	0.5638	0.9979	0.3959	0.4096	0.4143
rs1997794	0.6363	*	0.7055	0.7589	0.7846	0.6257	0.6306	0.5939
rs2235751	0.7134	0.5526	*	0.5506	0.9975	0.4837	0.4784	0.508
rs6045819	-0.1297	0.3056	-0.1108	*	0.8095	0.8657	0.8545	0.8473
rs10485703	-0.2193	0.3017	-0.1916	0.773	*	0.999	0.999	0.999
rs910080	0.3533	0.5	0.474	0.4362	0.4807	*	0.9996	0.9996
rs910079	0.3639	0.5018	0.4709	0.4325	0.4828	0.9952	*	0.9996
rs2235749	0.3856	0.4951	0.4773	0.4093	0.4608	0.9583	0.9541	*

MNP-PDYN	rs35286281	rs1997794	rs2235751	rs6045819	rs10485703	rs910080	rs910079	rs2235749
rs35286281	*	0.7097	0.737	0.4238	0.4069	0.697	0.6594	0.7153
rs1997794	0.6468	*	0.7245	0.998	0.998	0.8187	0.8211	0.8141
rs2235751	0.5688	0.6135	*	0.0002	0.9964	0.8231	0.8277	0.8048
rs6045819	0.1165	0.3009	0	*	0.8608	0.998	0.998	0.998
rs10485703	-0.0994	0.3233	-0.1879	0.8011	*	0.998	0.998	0.998
rs910080	0.6306	0.8127	0.7022	0.3031	0.3257	*	0.9997	0.9997
rs910079	0.6168	0.8	0.6829	0.2932	0.315	0.9669	*	0.9997
rs2235749	0.6365	0.7948	0.6981	0.3083	0.3312	0.9832	0.9509	*

<b>CABA-OPRK</b>	<b>rs35566036</b>	<b>rs3808627</b>	<b>rs6985606</b>
rs35566036	*	0.8278	0.9992
rs3808627	-0.3129	*	0.7887
rs6985606	-0.4048	-0.3623	*

<b>LP-OPRK</b>	<b>rs35566036</b>	<b>rs3808627</b>	<b>rs6985606</b>
rs35566036	*	0.3645	0.9995
rs3808627	-0.1554	*	0.9992
rs6985606	-0.5291	-0.4304	*

<b>RES-OPRK</b>	<b>rs35566036</b>	<b>rs3808627</b>	<b>rs6985606</b>
rs35566036	*	0.9981	0.5604
rs3808627	-0.1859	*	0.2503
rs6985606	-0.1343	-0.0821	*

<b>MNP-OPRK</b>	<b>rs35566036</b>	<b>rs3808627</b>	<b>rs6985606</b>
rs35566036	*	0.8536	0.9992
rs3808627	-0.4046	*	0.7483
rs6985606	-0.3925	-0.2786	*