

SUPPLEMENTARY TABLES

Table S1. Examination, demographic, and laboratory variables excluded from the study due to having zero variance.

Examination	Demographic	Laboratory
OPDDSNV	FIAPROXY	CARB E, URDMTOLC
OPDSBCO	RIDSTATR	CARB E, URDOMOLC
OPDSSNV	SDDSRVYR	CARB E, URXMTO
OPDUSNV		CARB E, URXOMO
VIQ110		FOLFMS E, LBDSFILC
VIQ130		HEPBD E, LBDHD
		PAH E, URDP01LC
		PAH E, URDP02LC
		PAH E, URDP04LC
		PAH E, URDP05LC
		PAH E, URDP06LC
		PAH E, URDP17LC
		PERNT E, URDUP8LC
		PHTHTE E, URDMEPLC
		PSA E, KIQ282
		SSHCV E, SSAHCVI
		UAM E, URDAAZLC
		UAM E, URDATZLC
		UAM E, URDDTZLC
		UAM E, URDSIMLC
		UAM E, URDSISLC
		UAM E, URXAAZ
		UAM E, URXATZ
		UAM E, URXDTZ
		UAM E, URXSIS
		UAM E, URXSISM
		UHM E, URDUCSLC
		UPP E, URDBSMLC
		UPP E, URDCHSLC
		UPP E, URDEMMLC
		UPP E, URDFRMLC
		UPP E, URDHLSLC
		UPP E, URDMMSLC
		UPP E, URDOXSLC
		UPP E, URDPIMLC
		UPP E, URDPROLC
		UPP E, URDRIMLC
		UPP E, URDSMMLC
		UPP E, URDSSFLC
		UPP E, URDTHFLC
		UPP E, URDTRALC
		UPP E, URDTRNLC
		UPP E, URXBMS
		UPP E, URXCHS
		UPP E, URXEMM
		UPP E, URXFRM
		UPP E, URXHLS
		UPP E, URXMSM
		UPP E, URXOXS
		UPP E, URXPIM
		UPP E, URXPRO

UPP E, URXRIM
UPP E, URXSMM
UPP E, URXSSF
UPP E, URXTHF
UPP E, URXTRA
UPP E, URXTRN
VID E, LBDVD3LC
VID E, LBDVIDLC
VOCWB E, LBD4CELC
VOCWB E, LBDV1ALC
VOCWB E, LBDV1ELC
VOCWB E, LBDV2CLC
VOCWB E, LBDV2ELC
VOCWB E, LBDV2PLC
VOCWB E, LBDV2TLC
VOCWB E, LBDV3BLC
VOCWB E, LBDV4ELC
VOCWB E, LBDVCBLC
VOCWB E, LBDVDEL
VOCWB E, LBDVHEL
VOCWB E, LBDVTPLC
VOCWB E, LBDVTPLC
VOCWB E, LBX4CE
VOCWB E, LBXV1A
VOCWB E, LBXV1E
VOCWB E, LBXV2C
VOCWB E, LBXV2E
VOCWB E, LBXV2P
VOCWB E, LBXV2T
VOCWB E, LBXV3B
VOCWB E, LBXV4E
VOCWB E, LBXVCB
VOCWB E, LBXVDE
VOCWB E, LBXVHE
VOCWB E, LBXVTP

Table S2. Statistically significant principal components (PCs) associated with diabetic retinopathy (DR).

PC	Beta	Standard Error	Z	p-Value	OR (95% CI)
PC1	-0.23	0.03	-6.73	<0.0001	0.79 (95% CI: 0.74–0.85)
PC9	0.19	0.05	3.92	<0.0001	1.21 (95% CI: 1.1–1.33)
PC211	-16.23	4.53	-3.58	0.0003	0 (95% CI: 0–0)
PC3	-0.14	0.04	-3.54	0.0004	0.87 (95% CI: 0.81–0.94)
PC2	-0.10	0.03	-3.20	0.001	0.9 (95% CI: 0.85–0.96)
PC21	0.20	0.07	3.02	0.003	1.22 (95% CI: 1.07–1.39)
PC27	0.21	0.08	2.80	0.005	1.24 (95% CI: 1.07–1.43)
PC110	0.48	0.17	2.76	0.006	1.62 (95% CI: 1.15–2.28)
PC8	-0.13	0.05	-2.66	0.008	0.88 (95% CI: 0.8–0.97)
PC142	-0.65	0.24	-2.65	0.008	0.52 (95% CI: 0.32–0.84)
PC38	0.22	0.09	2.49	0.01	1.24 (95% CI: 1.05–1.47)
PC34	0.19	0.08	2.26	0.02	1.21 (95% CI: 1.03–1.43)
PC25	0.16	0.07	2.24	0.03	1.18 (95% CI: 1.02–1.36)
PC105	-0.35	0.16	-2.16	0.03	0.7 (95% CI: 0.51–0.97)
PC117	-0.38	0.18	-2.12	0.03	0.68 (95% CI: 0.48–0.97)
PC30	0.17	0.08	2.10	0.04	1.18 (95% CI: 1.01–1.38)
PC152	-0.56	0.27	-2.08	0.04	0.57 (95% CI: 0.34–0.97)
PC37	-0.17	0.09	-2.04	0.04	0.84 (95% CI: 0.71–0.99)
PC13	0.10	0.05	1.88	0.06	1.11 (95% CI: 1–1.23)

PC19	-0.12	0.06	-1.88	0.06	0.89 (95% CI: 0.79–1.01)
PC10	0.09	0.05	1.85	0.06	1.1 (95% CI: 0.99–1.21)
PC90	-0.26	0.14	-1.82	0.07	0.77 (95% CI: 0.58–1.02)
PC148	-0.46	0.25	-1.80	0.07	0.63 (95% CI: 0.38–1.04)
PC29	-0.14	0.08	-1.80	0.07	0.87 (95% CI: 0.75–1.01)
PC185	0.89	0.50	1.78	0.07	2.44 (95% CI: 0.91–6.52)
PC70	-0.21	0.12	-1.76	0.08	0.81 (95% CI: 0.64–1.02)
PC17	-0.11	0.06	-1.76	0.08	0.9 (95% CI: 0.8–1.01)
PC35	0.15	0.09	1.72	0.09	1.16 (95% CI: 0.98–1.39)
PC208	-4.82	2.84	-1.70	0.09	0.01 (95% CI: 0–2.11)
PC205	-2.43	1.45	-1.68	0.09	0.09 (95% CI: 0.01–1.5)
PC62	-0.18	0.11	-1.66	0.10	0.83 (95% CI: 0.67–1.03)
PC114	-0.29	0.18	-1.65	0.10	0.75 (95% CI: 0.53–1.06)
PC98	-0.25	0.15	-1.64	0.10	0.78 (95% CI: 0.58–1.05)
PC46	0.15	0.09	1.63	0.10	1.17 (95% CI: 0.97–1.4)
PC94	-0.23	0.15	-1.55	0.12	0.8 (95% CI: 0.6–1.06)
PC18	-0.09	0.06	-1.54	0.12	0.91 (95% CI: 0.81–1.03)
PC60	-0.17	0.11	-1.52	0.13	0.85 (95% CI: 0.68–1.05)
PC45	-0.14	0.09	-1.50	0.13	0.87 (95% CI: 0.72–1.04)
PC197	1.00	0.68	1.47	0.14	2.72 (95% CI: 0.72–10.29)
PC118	0.26	0.18	1.42	0.15	1.3 (95% CI: 0.91–1.85)
PC77	0.18	0.13	1.41	0.16	1.2 (95% CI: 0.93–1.54)
PC150	-0.36	0.26	-1.40	0.16	0.7 (95% CI: 0.42–1.16)
PC89	-0.19	0.14	-1.36	0.17	0.83 (95% CI: 0.63–1.09)
PC184	0.68	0.52	1.32	0.19	1.98 (95% CI: 0.72–5.46)
PC79	-0.17	0.13	-1.32	0.19	0.84 (95% CI: 0.65–1.09)
PC20	-0.09	0.07	-1.31	0.19	0.92 (95% CI: 0.81–1.04)
PC64	-0.15	0.11	-1.31	0.19	0.86 (95% CI: 0.69–1.08)
PC74	-0.16	0.12	-1.30	0.19	0.85 (95% CI: 0.68–1.08)
PC216	120.42	92.41	1.30	0.19	1.98482520108455E+52 (95% CI: 0–9.137746010503E+130)
PC93	0.20	0.15	1.30	0.19	1.22 (95% CI: 0.9–1.64)
PC47	-0.13	0.10	-1.30	0.19	0.88 (95% CI: 0.72–1.07)
PC86	-0.18	0.14	-1.29	0.20	0.84 (95% CI: 0.64–1.1)
PC158	0.36	0.29	1.26	0.21	1.44 (95% CI: 0.82–2.52)
PC124	-0.24	0.19	-1.24	0.21	0.79 (95% CI: 0.54–1.15)
PC33	0.10	0.08	1.24	0.21	1.11 (95% CI: 0.94–1.3)
PC11	0.06	0.05	1.24	0.22	1.06 (95% CI: 0.96–1.18)
PC200	1.04	0.85	1.23	0.22	2.84 (95% CI: 0.54–14.98)
PC16	-0.07	0.06	-1.22	0.22	0.93 (95% CI: 0.83–1.04)
PC190	-0.68	0.56	-1.22	0.22	0.51 (95% CI: 0.17–1.52)
PC138	-0.28	0.23	-1.21	0.23	0.76 (95% CI: 0.49–1.19)
PC68	0.14	0.12	1.18	0.24	1.15 (95% CI: 0.91–1.44)
PC175	0.42	0.36	1.15	0.25	1.52 (95% CI: 0.75–3.07)
PC66	0.13	0.11	1.15	0.25	1.14 (95% CI: 0.91–1.43)
PC39	-0.10	0.09	-1.13	0.26	0.9 (95% CI: 0.75–1.08)
PC202	1.06	0.94	1.13	0.26	2.87 (95% CI: 0.46–18.04)
PC87	-0.16	0.14	-1.12	0.26	0.86 (95% CI: 0.65–1.12)
PC157	0.31	0.28	1.12	0.26	1.36 (95% CI: 0.79–2.36)
PC134	0.24	0.22	1.11	0.27	1.28 (95% CI: 0.83–1.96)
PC122	0.21	0.19	1.09	0.27	1.23 (95% CI: 0.85–1.78)
PC192	-0.63	0.57	-1.09	0.27	0.53 (95% CI: 0.17–1.65)
PC176	0.39	0.37	1.06	0.29	1.48 (95% CI: 0.72–3.04)
PC144	-0.25	0.24	-1.05	0.29	0.78 (95% CI: 0.48–1.25)
PC104	-0.17	0.16	-1.04	0.30	0.84 (95% CI: 0.61–1.16)
PC214	-17.04	16.48	-1.03	0.30	0 (95% CI: 0–4213127.42)
PC57	0.11	0.11	1.03	0.30	1.11 (95% CI: 0.91–1.37)
PC149	-0.27	0.26	-1.02	0.31	0.77 (95% CI: 0.46–1.28)
PC102	0.15	0.15	1.00	0.32	1.16 (95% CI: 0.87–1.57)

PC153	-0.26	0.26	-0.99	0.32	0.77 (95% CI: 0.46–1.29)
PC12	0.05	0.05	0.98	0.33	1.05 (95% CI: 0.95–1.17)
PC212	5.01	5.28	0.95	0.34	149.65 (95% CI: 0–4697021.47)
PC28	-0.07	0.08	-0.95	0.34	0.93 (95% CI: 0.8–1.08)
PC26	-0.07	0.07	-0.93	0.35	0.93 (95% CI: 0.81–1.08)
PC130	-0.19	0.21	-0.93	0.35	0.83 (95% CI: 0.55–1.24)
PC54	-0.09	0.10	-0.91	0.36	0.91 (95% CI: 0.74–1.11)
PC42	-0.08	0.09	-0.90	0.37	0.92 (95% CI: 0.77–1.1)
PC187	-0.47	0.53	-0.88	0.38	0.63 (95% CI: 0.22–1.77)
PC126	-0.17	0.19	-0.88	0.38	0.84 (95% CI: 0.58–1.23)
PC168	-0.28	0.32	-0.88	0.38	0.76 (95% CI: 0.41–1.41)
PC188	0.48	0.55	0.87	0.38	1.61 (95% CI: 0.55–4.71)
PC145	0.21	0.24	0.87	0.38	1.23 (95% CI: 0.77–1.98)
PC215	10.14	11.66	0.87	0.38	25387.49 (95% CI: 0–212202584172954)
PC196	0.57	0.66	0.86	0.39	1.76 (95% CI: 0.49–6.37)
PC113	-0.15	0.18	-0.86	0.39	0.86 (95% CI: 0.61–1.21)
PC95	0.13	0.15	0.86	0.39	1.14 (95% CI: 0.85–1.52)
PC217	#####	#####	0.85	0.39	NA
PC108	0.14	0.17	0.85	0.40	1.15 (95% CI: 0.83–1.59)
PC112	-0.14	0.17	-0.83	0.41	0.87 (95% CI: 0.62–1.21)
PC65	-0.09	0.11	-0.83	0.41	0.91 (95% CI: 0.73–1.14)
PC195	0.53	0.64	0.83	0.41	1.71 (95% CI: 0.48–6.03)
PC167	-0.26	0.32	-0.83	0.41	0.77 (95% CI: 0.41–1.43)
PC88	0.12	0.14	0.83	0.41	1.12 (95% CI: 0.85–1.48)
PC67	0.10	0.12	0.82	0.41	1.1 (95% CI: 0.88–1.38)
PC171	-0.27	0.34	-0.79	0.43	0.76 (95% CI: 0.39–1.49)
PC136	-0.18	0.22	-0.79	0.43	0.84 (95% CI: 0.54–1.3)
PC15	-0.04	0.06	-0.77	0.44	0.96 (95% CI: 0.85–1.07)
PC191	0.43	0.57	0.76	0.45	1.54 (95% CI: 0.5–4.73)
PC166	0.24	0.31	0.75	0.45	1.27 (95% CI: 0.68–2.34)
PC209	2.14	2.84	0.75	0.45	8.48 (95% CI: 0.03–2222.54)
PC225	#####	#####	-0.72	0.47	NA
PC99	-0.11	0.15	-0.72	0.47	0.9 (95% CI: 0.66–1.21)
PC23	-0.05	0.07	-0.71	0.47	0.95 (95% CI: 0.83–1.09)
PC143	0.17	0.24	0.71	0.48	1.19 (95% CI: 0.74–1.89)
PC40	-0.06	0.09	-0.70	0.48	0.94 (95% CI: 0.79–1.12)
PC170	-0.23	0.33	-0.70	0.48	0.79 (95% CI: 0.42–1.51)
PC181	0.29	0.42	0.68	0.50	1.33 (95% CI: 0.59–3.02)
PC219	#####	#####	-0.68	0.50	NA
PC44	-0.06	0.09	-0.66	0.51	0.94 (95% CI: 0.78–1.13)
PC120	0.12	0.19	0.66	0.51	1.13 (95% CI: 0.78–1.63)
PC75	-0.08	0.12	-0.65	0.51	0.92 (95% CI: 0.72–1.18)
PC147	-0.16	0.25	-0.64	0.52	0.85 (95% CI: 0.52–1.39)
PC220	#####	#####	0.63	0.53	NA
PC51	0.06	0.10	0.63	0.53	1.06 (95% CI: 0.88–1.29)
PC84	0.08	0.14	0.63	0.53	1.09 (95% CI: 0.83–1.42)
PC164	-0.18	0.30	-0.60	0.55	0.83 (95% CI: 0.46–1.51)
PC137	0.14	0.22	0.60	0.55	1.14 (95% CI: 0.74–1.77)
PC32	-0.05	0.08	-0.60	0.55	0.95 (95% CI: 0.81–1.12)
PC180	0.24	0.40	0.59	0.55	1.27 (95% CI: 0.58–2.8)
PC125	0.11	0.19	0.57	0.57	1.12 (95% CI: 0.76–1.63)
PC146	-0.14	0.25	-0.57	0.57	0.87 (95% CI: 0.53–1.41)
PC154	0.15	0.27	0.56	0.58	1.16 (95% CI: 0.69–1.97)
PC119	-0.10	0.19	-0.54	0.59	0.9 (95% CI: 0.63–1.3)
PC73	0.07	0.12	0.54	0.59	1.07 (95% CI: 0.84–1.35)
PC201	-0.46	0.87	-0.53	0.59	0.63 (95% CI: 0.11–3.47)
PC36	0.04	0.08	0.53	0.60	1.05 (95% CI: 0.89–1.23)
PC123	0.10	0.19	0.53	0.60	1.11 (95% CI: 0.76–1.61)
PC100	-0.08	0.15	-0.52	0.60	0.92 (95% CI: 0.68–1.25)

PC174	-0.18	0.35	-0.51	0.61	0.84 (95% CI: 0.42–1.67)
PC50	-0.05	0.10	-0.50	0.62	0.95 (95% CI: 0.78–1.16)
PC151	0.13	0.26	0.49	0.62	1.14 (95% CI: 0.68–1.9)
PC226	#####	#####	0.49	0.63	NA
PC199	0.36	0.77	0.47	0.64	1.44 (95% CI: 0.32–6.51)
PC182	-0.20	0.43	-0.46	0.64	0.82 (95% CI: 0.36–1.89)
PC69	0.05	0.12	0.46	0.65	1.06 (95% CI: 0.84–1.33)
PC169	-0.15	0.32	-0.45	0.65	0.86 (95% CI: 0.46–1.63)
PC179	0.18	0.40	0.45	0.65	1.2 (95% CI: 0.55–2.6)
PC97	-0.07	0.15	-0.44	0.66	0.94 (95% CI: 0.7–1.26)
PC173	-0.15	0.35	-0.43	0.67	0.86 (95% CI: 0.44–1.7)
PC189	0.23	0.55	0.42	0.67	1.26 (95% CI: 0.43–3.73)
PC133	-0.09	0.21	-0.42	0.68	0.92 (95% CI: 0.61–1.39)
PC71	0.05	0.12	0.41	0.68	1.05 (95% CI: 0.83–1.33)
PC224	#####	#####	0.41	0.68	NA
PC160	0.12	0.29	0.41	0.68	1.13 (95% CI: 0.63–2.01)
PC78	-0.05	0.13	-0.40	0.69	0.95 (95% CI: 0.74–1.22)
PC76	0.05	0.13	0.40	0.69	1.05 (95% CI: 0.82–1.34)
PC156	-0.11	0.28	-0.40	0.69	0.9 (95% CI: 0.52–1.54)
PC163	-0.12	0.30	-0.39	0.69	0.89 (95% CI: 0.49–1.61)
PC109	0.07	0.17	0.39	0.69	1.07 (95% CI: 0.77–1.49)
PC162	0.12	0.30	0.39	0.70	1.12 (95% CI: 0.63–2.01)
PC59	0.04	0.11	0.39	0.70	1.04 (95% CI: 0.84–1.29)
PC58	-0.04	0.10	-0.38	0.70	0.96 (95% CI: 0.78–1.18)
PC193	0.24	0.63	0.38	0.70	1.27 (95% CI: 0.37–4.4)
PC115	-0.07	0.18	-0.37	0.71	0.94 (95% CI: 0.66–1.32)
PC198	-0.26	0.71	-0.37	0.71	0.77 (95% CI: 0.19–3.09)
PC177	-0.14	0.38	-0.36	0.72	0.87 (95% CI: 0.42–1.82)
PC72	-0.04	0.12	-0.36	0.72	0.96 (95% CI: 0.75–1.21)
PC91	-0.05	0.14	-0.36	0.72	0.95 (95% CI: 0.72–1.26)
PC22	0.02	0.07	0.35	0.73	1.02 (95% CI: 0.9–1.17)
PC41	0.03	0.09	0.35	0.73	1.03 (95% CI: 0.86–1.23)
PC101	0.05	0.16	0.35	0.73	1.06 (95% CI: 0.78–1.43)
PC165	-0.11	0.31	-0.34	0.73	0.9 (95% CI: 0.49–1.65)
PC139	-0.08	0.23	-0.33	0.74	0.93 (95% CI: 0.59–1.45)
PC210	-1.07	3.21	-0.33	0.74	0.34 (95% CI: 0–187.7)
PC128	-0.07	0.20	-0.33	0.74	0.94 (95% CI: 0.63–1.38)
PC129	0.07	0.20	0.32	0.75	1.07 (95% CI: 0.72–1.59)
PC83	-0.04	0.13	-0.31	0.76	0.96 (95% CI: 0.74–1.25)
PC132	-0.06	0.21	-0.30	0.76	0.94 (95% CI: 0.62–1.42)
PC5	-0.01	0.04	-0.29	0.77	0.99 (95% CI: 0.92–1.07)
PC4	-0.01	0.04	-0.28	0.78	0.99 (95% CI: 0.92–1.06)
PC43	-0.03	0.09	-0.28	0.78	0.97 (95% CI: 0.81–1.17)
PC159	0.08	0.29	0.27	0.79	1.08 (95% CI: 0.61–1.9)
PC82	-0.03	0.13	-0.27	0.79	0.97 (95% CI: 0.75–1.25)
PC6	-0.01	0.04	-0.26	0.79	0.99 (95% CI: 0.91–1.08)
PC31	-0.02	0.08	-0.26	0.80	0.98 (95% CI: 0.84–1.14)
PC49	-0.03	0.10	-0.26	0.80	0.98 (95% CI: 0.81–1.18)
PC127	0.05	0.20	0.26	0.80	1.05 (95% CI: 0.71–1.55)
PC161	0.07	0.29	0.25	0.80	1.08 (95% CI: 0.61–1.91)
PC85	-0.03	0.14	-0.24	0.81	0.97 (95% CI: 0.74–1.26)
PC111	0.04	0.17	0.23	0.82	1.04 (95% CI: 0.74–1.45)
PC203	0.22	1.02	0.22	0.83	1.25 (95% CI: 0.17–9.23)
PC55	0.02	0.10	0.21	0.83	1.02 (95% CI: 0.83–1.26)
PC207	0.30	1.45	0.20	0.84	1.34 (95% CI: 0.08–23.25)
PC141	-0.05	0.23	-0.20	0.84	0.95 (95% CI: 0.6–1.51)
PC223	#####	#####	-0.18	0.85	NA
PC14	-0.01	0.06	-0.18	0.86	0.99 (95% CI: 0.89–1.1)
PC7	-0.01	0.04	-0.18	0.86	0.99 (95% CI: 0.91–1.08)

PC56	0.02	0.10	0.17	0.86	1.02 (95% CI: 0.83–1.25)
PC121	-0.03	0.19	-0.17	0.86	0.97 (95% CI: 0.67–1.4)
PC221	#####	#####	-0.17	0.87	NA
PC186	0.08	0.51	0.17	0.87	1.09 (95% CI: 0.4–2.93)
PC80	0.02	0.13	0.15	0.88	1.02 (95% CI: 0.79–1.32)
PC131	-0.03	0.21	-0.15	0.88	0.97 (95% CI: 0.64–1.46)
PC222	#####	#####	-0.15	0.88	NA
PC52	-0.01	0.10	-0.13	0.89	0.99 (95% CI: 0.81–1.2)
PC140	0.03	0.23	0.13	0.90	1.03 (95% CI: 0.65–1.63)
PC48	0.01	0.10	0.13	0.90	1.01 (95% CI: 0.84–1.22)
PC155	-0.03	0.27	-0.12	0.90	0.97 (95% CI: 0.56–1.65)
PC172	-0.04	0.34	-0.12	0.90	0.96 (95% CI: 0.49–1.88)
PC106	0.02	0.16	0.12	0.91	1.02 (95% CI: 0.74–1.4)
PC116	-0.02	0.18	-0.11	0.91	0.98 (95% CI: 0.69–1.39)
PC63	-0.01	0.11	-0.10	0.92	0.99 (95% CI: 0.79–1.23)
PC81	-0.01	0.13	-0.09	0.93	0.99 (95% CI: 0.76–1.28)
PC194	0.05	0.61	0.08	0.94	1.05 (95% CI: 0.32–3.48)
PC135	-0.01	0.22	-0.07	0.95	0.99 (95% CI: 0.64–1.52)
PC92	-0.01	0.15	-0.06	0.95	0.99 (95% CI: 0.75–1.32)
PC183	-0.02	0.44	-0.05	0.96	0.98 (95% CI: 0.41–2.31)
PC107	0.01	0.17	0.05	0.96	1.01 (95% CI: 0.73–1.39)
PC61	0.00	0.11	0.04	0.97	1 (95% CI: 0.81–1.24)
PC53	0.00	0.10	-0.03	0.98	1 (95% CI: 0.82–1.21)
PC213	0.19	6.20	0.03	0.98	1.21 (95% CI: 0–228449.56)
PC178	-0.01	0.38	-0.03	0.98	0.99 (95% CI: 0.47–2.1)
PC96	0.00	0.15	0.03	0.98	1 (95% CI: 0.75–1.35)
PC206	-0.03	1.31	-0.02	0.98	0.97 (95% CI: 0.07–12.6)
PC103	0.00	0.16	0.02	0.99	1 (95% CI: 0.74–1.37)
PC218	#####	#####	-0.01	0.99	NA
PC24	0.00	0.07	0.00	1.00	1 (95% CI: 0.87–1.14)
PC204	0.00	1.23	0.00	1.00	1 (95% CI: 0.09–11.13)

Notes: Principal component analysis was performed on laboratory variables in an unsupervised fashion, with resulting PCs regressed against DR outcome.

Table S3. RandomForest™-selected features for replication cohort NHANES 2005-2006.

Marker	Mean decrease accuracy	Mean decrease gini
Glycohemoglobin (%)	16.96	18.06
Pyridoxal 5'-phosphate (nmol/L)	8.15	5.24
Total Cholesterol (mmol/L)	8.14	6.33
Cholesterol (mmol/L)	7.53	6.10
Blood urea nitrogen (mmol/L)	5.62	5.14
Diastolic blood pressure	4.76	5.94
Albumin, urine (mg/L)	4.25	6.00
Hematocrit (%)	4.18	3.35
Lactate dehydrogenase LDH (U/L)	3.97	4.44
Mono-(2-ethyl-5-hydroxyhexyl) phthalate	3.72	1.75
Uranium, urinary (µg/L)	3.57	2.55
2-amnothiazolne-4-carboxylic acid(ng/mL)	3.51	3.25
Mono-n-methyl phthalate	3.22	1.85
Urinary nitrate (ng/mL)	3.20	7.25
Blood Carbon Tetrachloride (ng/mL)	3.06	0.70
Perchlorate, water (ng/mL)	2.61	3.50
Iron, refrigerated (µmol/L)	2.59	4.87
Potassium (mmol/L)	2.47	4.06
Nitrate, water (ng/mL)	2.35	2.61
Red blood cell count (million cells/µL)	1.89	4.61
Mono-2-ethyl-5-carboxypentyl phthalate	1.67	2.02
Hemoglobin (g/dL)	1.47	2.47

2-hydroxyphenanthrene (ng/L)	1.24	2.50
Dimethyldithiophosphate ($\mu\text{g/L}$)	1.05	1.24
Alkaline phosphatase (U/L)	0.80	4.56
Propyl paraben (ng/ml)	0.80	3.02

Notes: The model was initially trained on all laboratory variables in an unsupervised fashion, with Kappa-based model tuning to select the optimum values for "mtry" (the ideal number of variables to randomly sample) and "ntrees" (the ideal number of trees). Only features contributing >1% mean decrease in accuracy from initial model were retained.

Supplementary figures

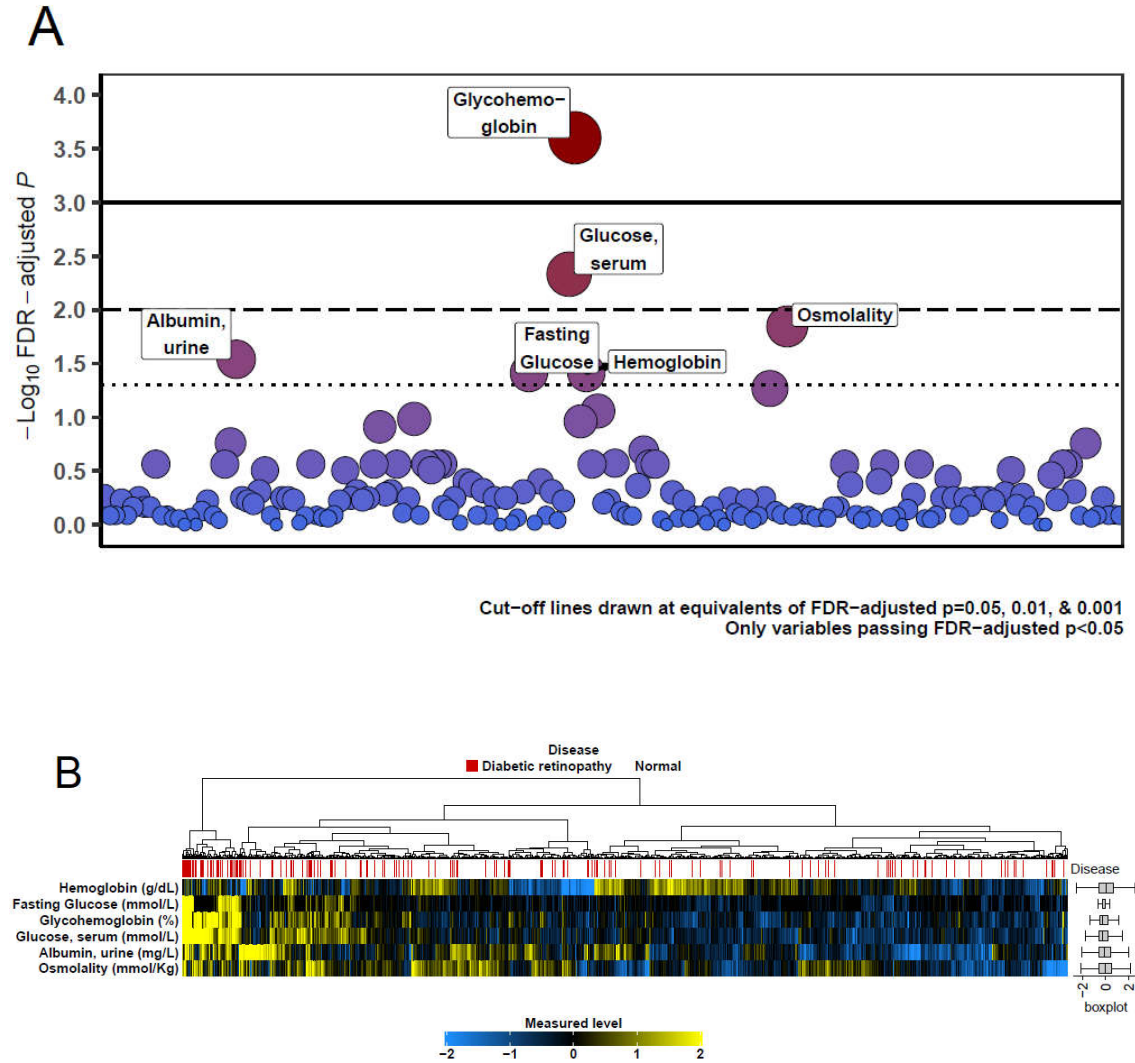


Figure S1. Laboratory variables statistically significantly associated with diabetic retinopathy.

Variables were tested in a univariate fashion, with all p -values then adjusted for false discovery rate (FDR). Six variables reached statistical significance at FDR-adjusted $p \leq 0.05$. **A)** Manhattan plot of laboratory variables associated with diabetic retinopathy from univariate testing. Variables were tested via a compute-parallelized procedure, with p -values then adjusted for false discovery rate (FDR). Six variables reached statistical significance at FDR-adjusted $p \leq 0.05$. Y-axis is plotting $-\log_{10}$ (FDR-adjusted p -value). **B)** clustering and heatmap.

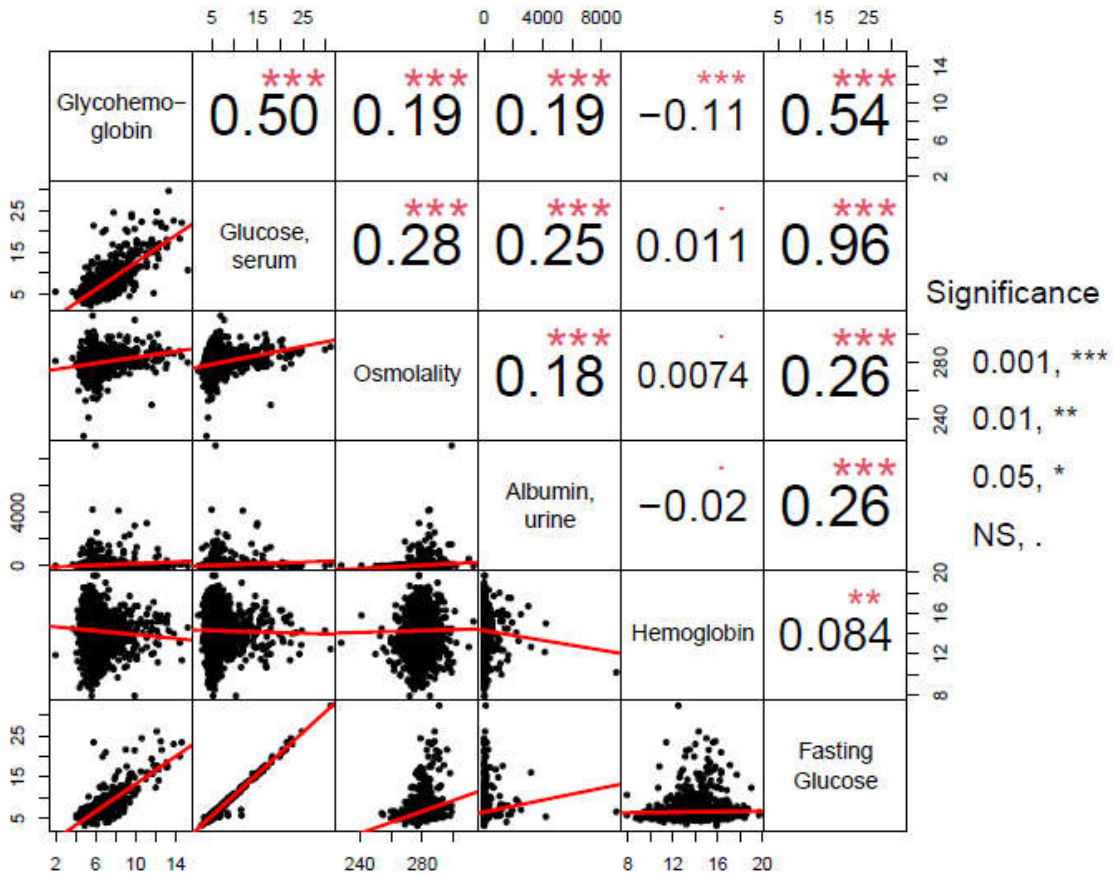


Figure S2. Inter-correlation among laboratory variables statistically significantly associated with diabetic retinopathy.

Bottom triangle, scatter plots with linear regression fit line. Top triangle, Spearman ρ coefficients, with p -value from correlation test indicated by asterisks.

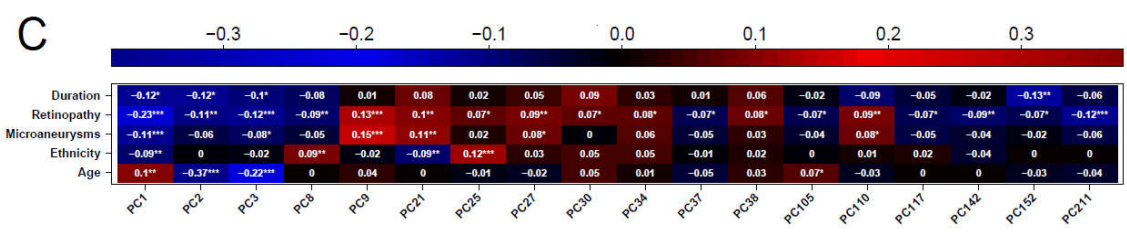
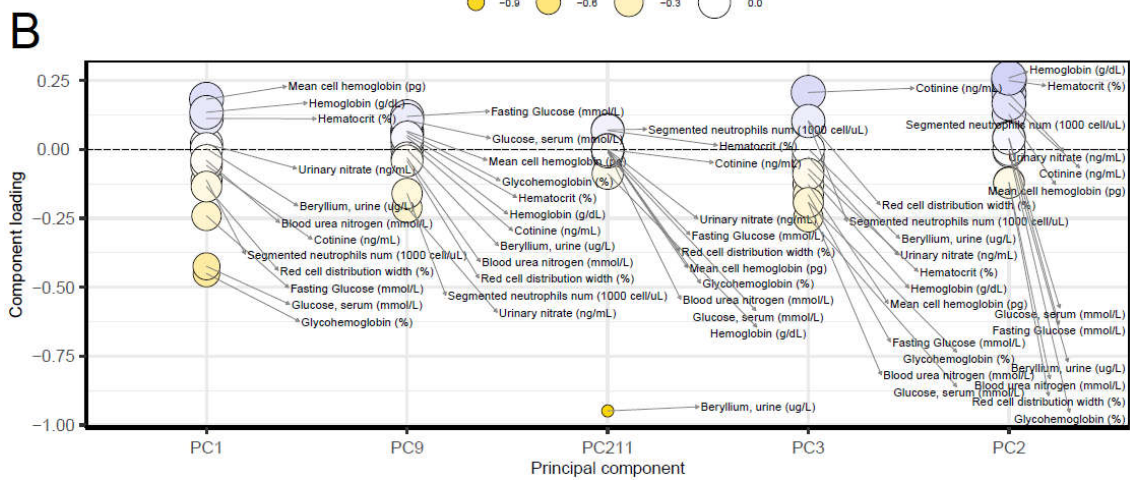
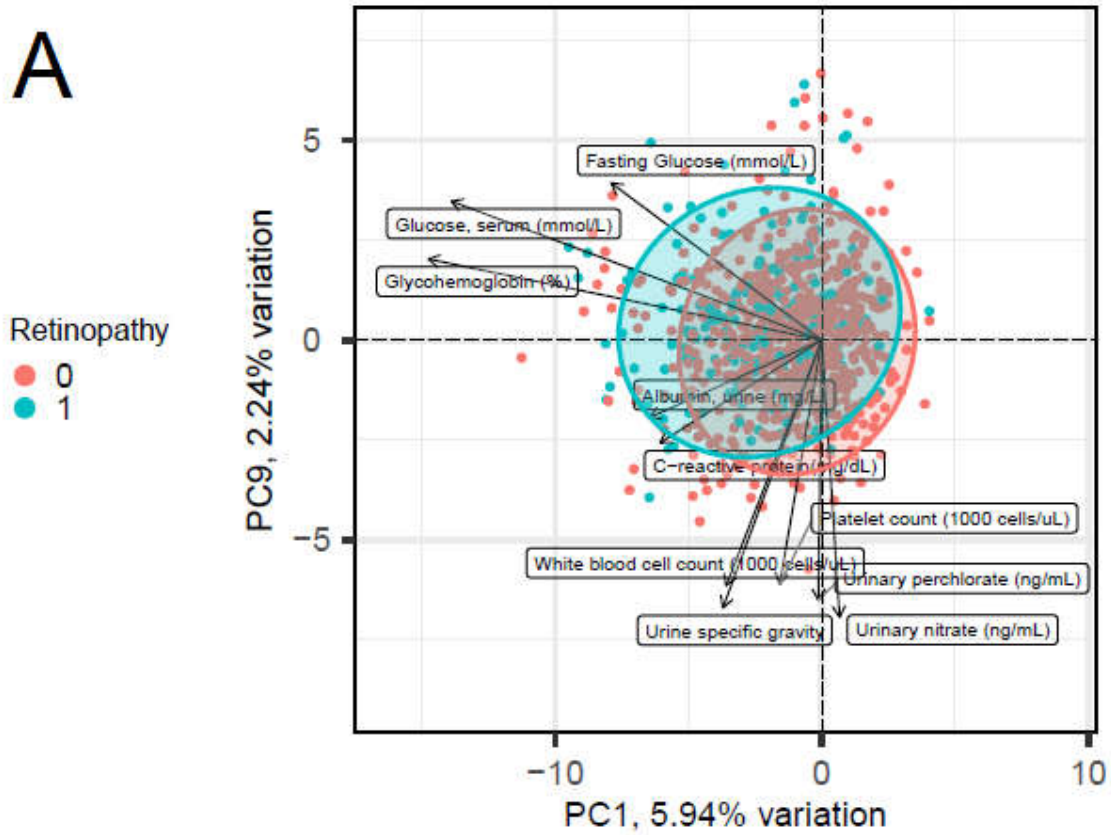


Figure S3. Principal component analysis (PCA) highlights key sources of variation in the cohort.

PCA was performed in an unsupervised fashion on laboratory variables. A, bi-plot of PC1 versus PC9, the top two PCs statistically significantly associated with diabetic retinopathy (DR) (see **Supplementary Table 2**), reveals HbA1c, CRP, and others as key drivers of variation along these PCs. 0, no retinopathy; 1, retinopathy. B, top variables contributing to variation along top five PCs statistically significantly associated with DR. C, "eigenplot" comparing covariates (age, ethnicity, and diabetes duration), retinopathy (as DR), and the top retinal lesion (microaneurysms) against all PCs statistically significantly associated with DR. The statistically significant correlations highlight relevant usage of these covariates during univariate testing. Pearson correlation values shown, with significancies: ***, $p \leq 0.001$; **, $p \leq 0.01$; *, $p \leq 0.05$.