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Supplementary Table 3 (page 20): Additional details related to Table 2. Replication genotyping for these SNPs failed assay design or quality control and a suitable proxy variant was selected (rs1596117, proxy rs4859430; rs7681154, proxy rs3910105; rs13201101, proxy rs8192591; based on discovery series comparison, the minor allele for rs3910105 tags the major allele of rs7681154 therefore risk is consistent across proxy and discovery SNP). Note, only replication phase p-values are one-sided. Nearest gene or previously published proximal gene names included in table.

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Nearest gene or previously published proximal gene names included in table.

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SUPPLEMENTARY NOTE

Part 1: Discussion of novel loci.

The six novel loci we have identified (*SIPA1L2*, *INPP5F*, *MIR4697*, *GCH1*, *VPS13C*, and *DDRGK1*) in this meta-analysis include biologically plausible connections to PD etiology based on previous research. In the context of GWAS it is clear that the pathobiologically relevant gene is not always the closest or most obvious proximal candidate.¹ However, it is perhaps useful to discuss the potential pathobiological relevance of the candidate genes at our loci, with the caveat that further work is required to identify whether these are the true biological candidates. The signal-induced proliferation-associated 1 like 2 (*SIPA1L2*) locus has been implicated in inflammatory pathways by GWAS as well as in pharmacogenetic studies of smoking cessation and related behaviors.^{2,3} *INPP5F* has been suggested as an initial candidate locus for late-onset alzheimer's disease from linkage studies.⁴ *MIR4697* is a provisional microRNA identified in next generation sequencing studies of breast cancer.⁵ GTP cyclohydrolase 1 (*GCH1*) is a likely biological candidate for future therapeutics and follow-up study, as SNPs at this locus are also associated with dopamine clearance in urine.⁶ In addition, the *GCH1* locus has been offered as a risk locus in early onset and atypical PD, but results until this point have never been definitive.⁷⁻⁹ The locus containing rs2414739 and the *VPS13C* gene possesses a likely connection to neurodegenerative disease etiology as alternative splicing of *VPS13C* contributes to risk of congenital myasthenic syndrome.¹⁰ *DDRGK1* is also connected to neurodegenerative disease as it has been suggested as a candidate locus for spinocerebellar ataxias.¹¹

1. Smemo, S. *et al.* Obesity-associated variants within FTO form long-range functional connections with IRX3. *Nature* **507**, 371–375 (2014).
2. Rose, J. E., Behm, F. M., Drgon, T., Johnson, C. & Uhl, G. R. Personalized smoking cessation: interactions between nicotine dose, dependence and quit-success genotype score. *Mol. Med. Camb. Mass* **16**, 247–253 (2010).
3. Ferreira, R. C. *et al.* Association of IFIH1 and other autoimmunity risk alleles with selective IgA deficiency. *Nat. Genet.* **42**, 777–780 (2010).
4. Grupe, A. *et al.* A scan of chromosome 10 identifies a novel locus showing strong association with late-onset Alzheimer disease. *Am. J. Hum. Genet.* **78**, 78–88 (2006).
5. Persson, H. *et al.* Identification of new microRNAs in paired normal and tumor breast tissue suggests a dual role for the ERBB2/Her2 gene. *Cancer Res.* **71**, 78–86 (2011).
6. Comuzzie, A. G. *et al.* Novel genetic loci identified for the pathophysiology of childhood obesity in the Hispanic population. *PLoS One* **7**, e51954 (2012).
7. Hertz, J. M. *et al.* Low frequency of Parkin, Tyrosine Hydroxylase, and GTP Cyclohydrolase I gene mutations in a Danish population of early-onset

- Parkinson's Disease. *Eur. J. Neurol. Off. J. Eur. Fed. Neurol. Soc.* **13**, 385–390 (2006).
8. Cobb, S. A. *et al.* GCH1 in early-onset Parkinson's disease. *Mov. Disord. Off. J. Mov. Disord. Soc.* **24**, 2070–2075 (2009).
 9. Bandmann, O., Daniel, S., Marsden, C. D., Wood, N. W. & Harding, A. E. The GTP-cyclohydrolase I gene in atypical parkinsonian patients: a clinico-genetic study. *J. Neurol. Sci.* **141**, 27–32 (1996).
 10. Masuda, A. *et al.* hnRNP H enhances skipping of a nonfunctional exon P3A in CHRNA1 and a mutation disrupting its binding causes congenital myasthenic syndrome. *Hum. Mol. Genet.* **17**, 4022–4035 (2008).
 11. Kobayashi, H. *et al.* Expansion of intronic GGCCTG hexanucleotide repeat in NOP56 causes SCA36, a type of spinocerebellar ataxia accompanied by motor neuron involvement. *Am. J. Hum. Genet.* **89**, 121–130 (2011).

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111 - Barrow Neurological Institute
112 - Baylor College of Medicine
113 - Beth Israel Deaconess Medical Center
114 - Brigham & Women's Hospital
115 - Brown University (Memorial Hospital of RI)
116 - Colorado Neurological Institute
117 - Columbia University Medical Center
118 - Creighton University
119 - Evanston Northwestern Healthcare
120 - Hotel-Dieu Hospital-Chum
121 - Hunter Homes McGuire Veterans Medical Center
122 - Indiana University School of Medicine
123 - Institute For Neurodegenerative Disorders
124 - Johns Hopkins University
125 - London Health Sciences Centre
126 - Mayo Clinic Jacksonville
127 - McFarland Neurosciences
128 - Medical College of Georgia
129 - Medical College of Wisconsin
130 - Medical University of Ohio
131 - Mount Sinai School of Medicine
132 - North Shore-LIJ Health System
133 - Northwestern University
134 - Ochsner Clinic Foundation
135 - Ohio State University
136 - Ottawa Hospital Civic Site
137 - Pacific Neuroscience Medical Group
138 - Saskatoon Dist Health Board Royal Univ Hosp
139 - Scott & White Hospital/Texas A&M University

- 140 - The Parkinson's & Movement Disorder Institute
- 141 - Toronto Western Hospital, University Health
- 142 - UMDNJ-School of Osteopathic Medicine
- 143 - University of Alabama at Birmingham
- 144 - University of Alberta
- 145 - University of Calgary
- 146 - University of California Irvine
- 147 - University of California San Diego
- 148 - University of California San Francisco
- 149 - University of Chicago
- 150 - University of Cincinnati
- 151 - University of Colorado Health Sciences Center
- 152 - University of Connecticut
- 153 - University of Iowa
- 154 - University of Kansas Medical Center
- 155 - University of Maryland School of Medicine
- 156 - University of Miami
- 157 - University of Minnesota
- 158 - University of New Mexico
- 159 - University of Puerto Rico School of Medicine
- 160 - University of Rochester
- 161 - University of South Florida
- 162 - University of Tennessee Health Science Center
- 163 - University of Texas Southwestern Medical Center
- 164 - Wake Forest University School of Medicine
- 165 - Washington University
- 166 - University Southern California School of Medicine
- 167 - University of Lübeck, Germany
- 168 - UMDNJ-Robert Wood Johnson Medical School
- 169 - Massachusetts General Hospital, Harvard Medical School
- 170 - University of Virginia Health System
- 171 - University of Toronto
- 172 - Sun Health Research Institute
- 173 - Parkinson Institute, Istituti Clinici di Perfezionamento, Milano, Italy
- 174 - Cleveland Clinic Foundation
- 175 - University of Louisville School of Medicine
- 176 - University of Sydney ANZAC Research Institute, Concord Hospital, Sydney, Australia
- 177 - Struthers Parkinson's Center, Minneapolis
- 178 - Port City Neurology, Scarborough, ME
- 179 - Parkinson's Disease and Movement Disorder Center of Boca Raton
- 180 - Newcastle University, Newcastle upon Tyne, UK
- 181 - General Regional Hospital Bolzano, Bolzano, Italy
- 182 - University of Arkansas for Medical Sciences
- 183 - Aarhus University Hospital, Aarhus, Denmark
- 184 - University of Arizona

- 185 - Auckland City Hospital, Auckland, New Zealand
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Supplementary Table 3 (page 20):

Additional details related to Table 2.

Replication genotyping for these SNPs failed assay design or quality control and a suitable proxy variant was selected (rs1596117, proxy rs4859430; rs7681154, proxy rs3910105; rs13201101, proxy rs8192591; based on discovery series comparison, the minor allele for rs3910105 tags the major allele of rs7681154 therefore risk is consistent across proxy and discovery SNP). Note, only replication phase p-values are one-sided. Nearest gene or previously published proximal gene names included in table.

Significant SNPs (containing proxy SNPs)	Minor allele frequency	FC (log10) in SNP's linked SNPs	Corresponding gene information	Summary statistic for association analysis	Summary statistic for decomposition	Summary statistic from replication phase	Summary statistic to compare disease and replication genes
rs11138700	0.000	-1.0	rs11138700	Pearson (by)	Pearson (by)	Pearson (by)	Pearson (by)
rs1596117	0.000	-1.0	rs1596117	Exact test	Exact test	Exact test	Exact test
rs3910105	0.000	-1.0	rs3910105	Logistic regression	Logistic regression	Logistic regression	Logistic regression
rs13201101	0.000	-1.0	rs13201101	Rank correlation	Rank correlation	Rank correlation	Rank correlation
rs8192591	0.000	-1.0	rs8192591	Spearman	Spearman	Spearman	Spearman
rs1596117	0.000	-1.0	rs1596117	Wilcoxon signed rank	Wilcoxon signed rank	Wilcoxon signed rank	Wilcoxon signed rank
rs3910105	0.000	-1.0	rs3910105	Fisher's exact	Fisher's exact	Fisher's exact	Fisher's exact
rs13201101	0.000	-1.0	rs13201101	Z test	Z test	Z test	Z test
rs8192591	0.000	-1.0	rs8192591	T test	T test	T test	T test
rs1596117	0.000	-1.0	rs1596117	Negligible	Negligible	Negligible	Negligible
rs3910105	0.000	-1.0	rs3910105	Negligible	Negligible	Negligible	Negligible
rs13201101	0.000	-1.0	rs13201101	Negligible	Negligible	Negligible	Negligible
rs8192591	0.000	-1.0	rs8192591	Negligible	Negligible	Negligible	Negligible
rs1596117	0.000	-1.0	rs1596117	Negligible	Negligible	Negligible	Negligible
rs3910105	0.000	-1.0	rs3910105	Negligible	Negligible	Negligible	Negligible
rs13201101	0.000	-1.0	rs13201101	Negligible	Negligible	Negligible	Negligible
rs8192591	0.000	-1.0	rs8192591	Negligible	Negligible	Negligible	Negligible

Supplementary Table 4 (page
21): Summary statistics for risk
profile scoring analyses.

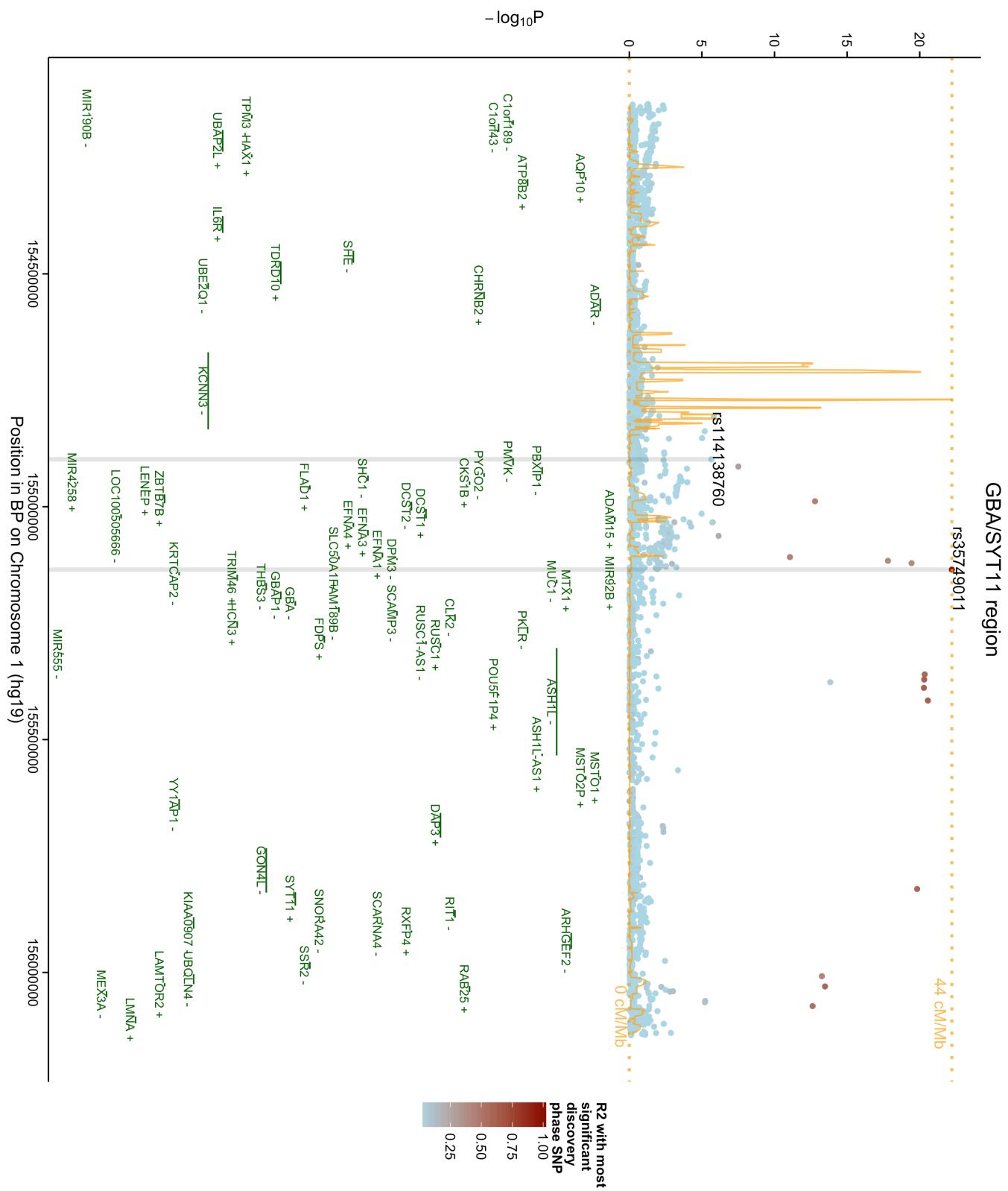
Study	Trend P-value	AUC	Trend (1 SD of change from mean)		1st quintile		2nd quintile		3rd quintile		4th quintile		5th quintile	
			OR	95% CI	OR	95% CI	OR	95% CI	OR	95% CI	OR	95% CI	OR	95% CI
USA	<2x10-16	0.69	1.5	1.41-1.59	1	-	1.47	1.24-1.76	1.66	1.39-1.99	2.14	1.79-2.56	3.18	2.65-3.79
Germany	6.4x10-12	0.59	1.4	1.14-1.44	1	-	1.34	1.14-1.53	1.17	1.09-1.27	1.66	1.36-1.99	2.06	1.67-3.58
Greece	5.12x10-11	0.58	1.38	1.25-1.52	1	-	1.38	1.03-1.85	1.47	1.09-1.97	1.61	1.2-2.17	2.48	1.83-3.36
UK	6.36x10-05	0.61	1.51	1.23-1.85	1	-	1.48	0.78-2.83	2.38	1.15-4.93	2.52	1.28-4.95	4.13	2.03-8.39
France	<2x10-16	0.67	1.92	1.65-2.22	1	-	1.43	0.93-2.18	2.24	1.48-3.39	3.01	1.96-4.62	5.97	3.79-9.38
Combined	<2x10-16	0.63	1.51	1.38-1.66	1	-	1.42	1.25-1.61	1.7	1.5-1.93	2.07	1.72-2.48	3.31	2.55-4.3
% Cases					36.14		43.92		48.44		53.62		62.34	

Supplementary Table 6 (page 23): Minor allele frequencies for all SNPs in meta-analysis stratified by case-control status for discovery and replication phases. All replication samples were included in these estimates. Discovery phase estimates include all 23andMe samples, PGPD, HIHG, NGRC and all IPDGC samples except those from the UK and Iceland.

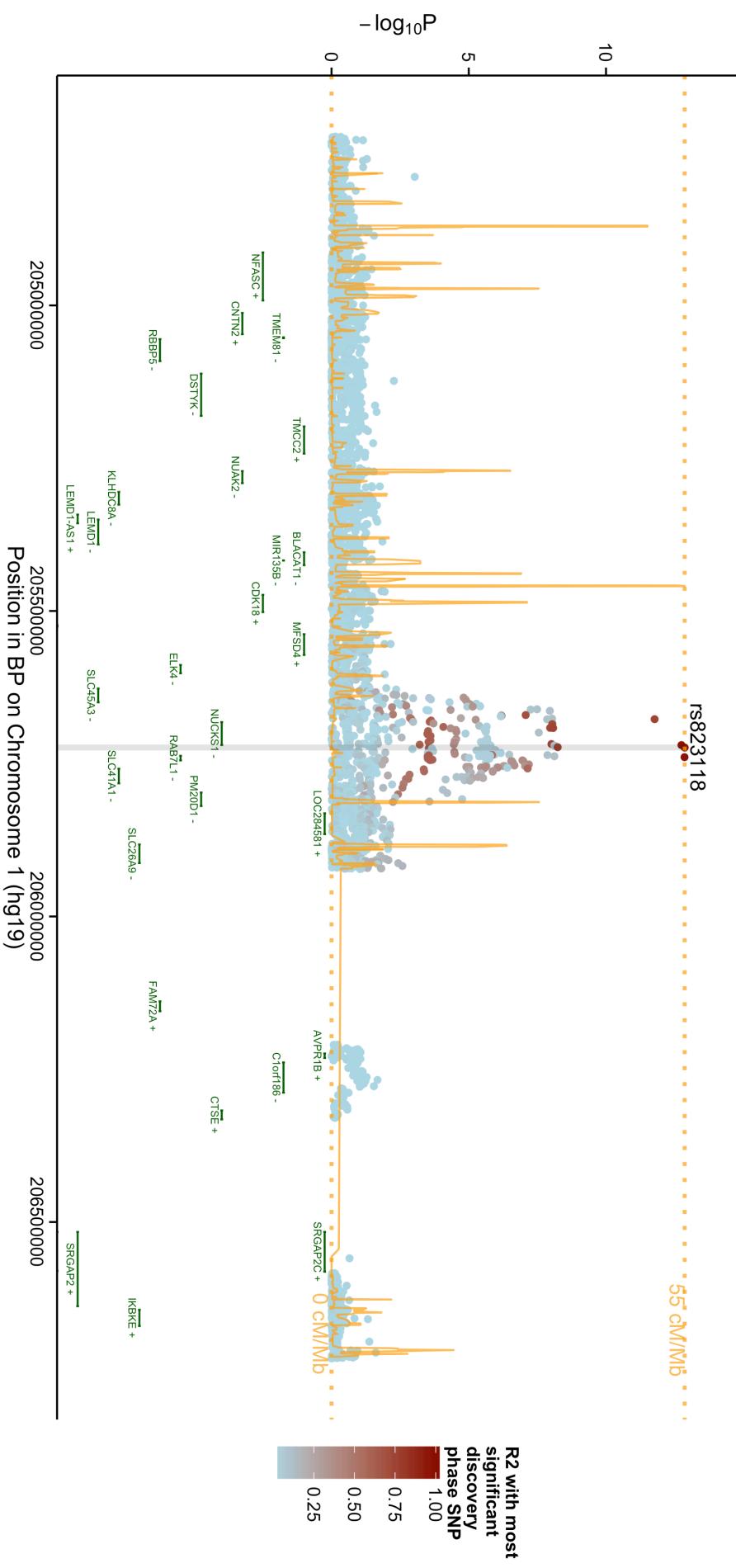
SNP	C	Position (bp)	Minor allele	Discovery and conditional phases, minor allele frequency (cases)	Discovery and conditional phases, minor allele frequency (controls)	Replication phase, minor allele frequency (cases)	Replication phase, minor allele frequency (controls)
rs114138760	1	154898185	C	0.015	0.011	0.013	0.009
rs35749011	1	155135036	A	0.026	0.014	NA	NA
rs71628662	1	155359992	C	0.028	0.018	0.015	0.007
rs823118	1	205723572	C	0.416	0.444	0.405	0.435
rs10797576	1	232664611	T	0.152	0.134	0.163	0.146
rs6430538	2	135539967	T	0.408	0.436	0.480	0.497
rs1474055	2	169110394	T	0.145	0.123	NA	NA
rs1955337	2	169129145	T	0.147	0.125	0.142	0.121
rs62267708	3	87489314	T	0.021	0.021	0.042	0.043
rs115185635	3	87520857	C	0.022	0.022	NA	NA
rs34016896	3	160992864	T	0.331	0.312	0.335	0.326
rs12637471	3	182762437	A	0.177	0.200	0.174	0.199
rs79217002	3	183011072	G	0.011	0.008	0.011	0.012
rs34884217	4	944210	C	0.062	0.073	0.096	0.105
rs34311866	4	951947	C	0.226	0.183	0.236	0.195
rs11724635	4	15737101	C	0.422	0.455	0.421	0.456
rs4859430	4	77149099	A	0.150	0.132	0.142	0.133
rs1596117	4	77151490	T	0.218	0.198	NA	NA
rs6812193	4	77198986	T	0.342	0.370	0.348	0.363
rs356182	4	90626111	G	0.421	0.355	0.397	0.337
rs3910105	4	90682571	G	0.427	0.457	0.444	0.459
rs7681154	4	90763703	C	0.502	0.500	NA	NA
rs8192591	6	32185796	T	0.039	0.036	0.033	0.028
rs13201101	6	32343604	T	0.057	0.050	NA	NA
rs9275326	6	32666660	T	0.083	0.097	0.083	0.093
rs199347	7	23293746	G	0.385	0.413	0.396	0.416
rs591323	8	16697091	A	0.261	0.279	0.252	0.272
rs60298754	8	89373041	T	0.025	0.023	NA	NA
rs7077361	10	15561543	C	0.116	0.130	0.124	0.131
rs10886515	10	121343589	C	0.270	0.291	0.281	0.284
rs117896735	10	121536327	A	0.017	0.012	NA	NA
rs118117788	10	121710488	T	0.013	0.008	0.021	0.015
rs12283611	11	83487277	A	0.398	0.415	0.418	0.426
rs3793947	11	83544472	A	0.428	0.446	NA	NA
rs329648	11	133765367	T	0.372	0.346	0.388	0.359
rs76904798	12	40614434	T	0.162	0.141	0.159	0.144
rs11060180	12	123303586	G	0.426	0.444	0.420	0.446
rs11158026	14	55348869	T	0.309	0.334	0.307	0.318
rs1077989	14	67975822	C	0.484	0.486	0.480	0.487
rs1555399	14	67984370	A	0.478	0.481	NA	NA
rs2414739	15	61994134	G	0.244	0.269	0.240	0.264
rs14235	16	31121793	A	0.400	0.374	0.427	0.395
rs11868035	17	17715101	A	0.287	0.301	0.303	0.314
rs17649553	17	43994648	T	0.189	0.231	0.187	0.228
rs12456492	18	40673380	G	0.328	0.300	0.337	0.316
rs117022814	19	2209647	T	0.027	0.023	0.022	0.022
rs10402629	19	2324458	G	0.244	0.225	0.217	0.218
rs62120679	19	2363319	T	0.321	0.300	NA	NA
rs55785911	20	3153503	A	0.357	0.374	0.368	0.386
rs8118008	20	3168166	G	0.341	0.359	NA	NA
rs2823357	21	16914905	A	0.382	0.375	0.383	0.378

Supplementary Figure 1 (pages 24 - 55): Regional association plots. 32 regional association plots for SNPs from discovery phase analyses +/- 1 Mb from most significant SNP per locus in Table 1. The r² pattern is based on most significant SNP per locus, based on the 283 European ancestry samples from the August 2010 release of the 1000 genomes project dataset.

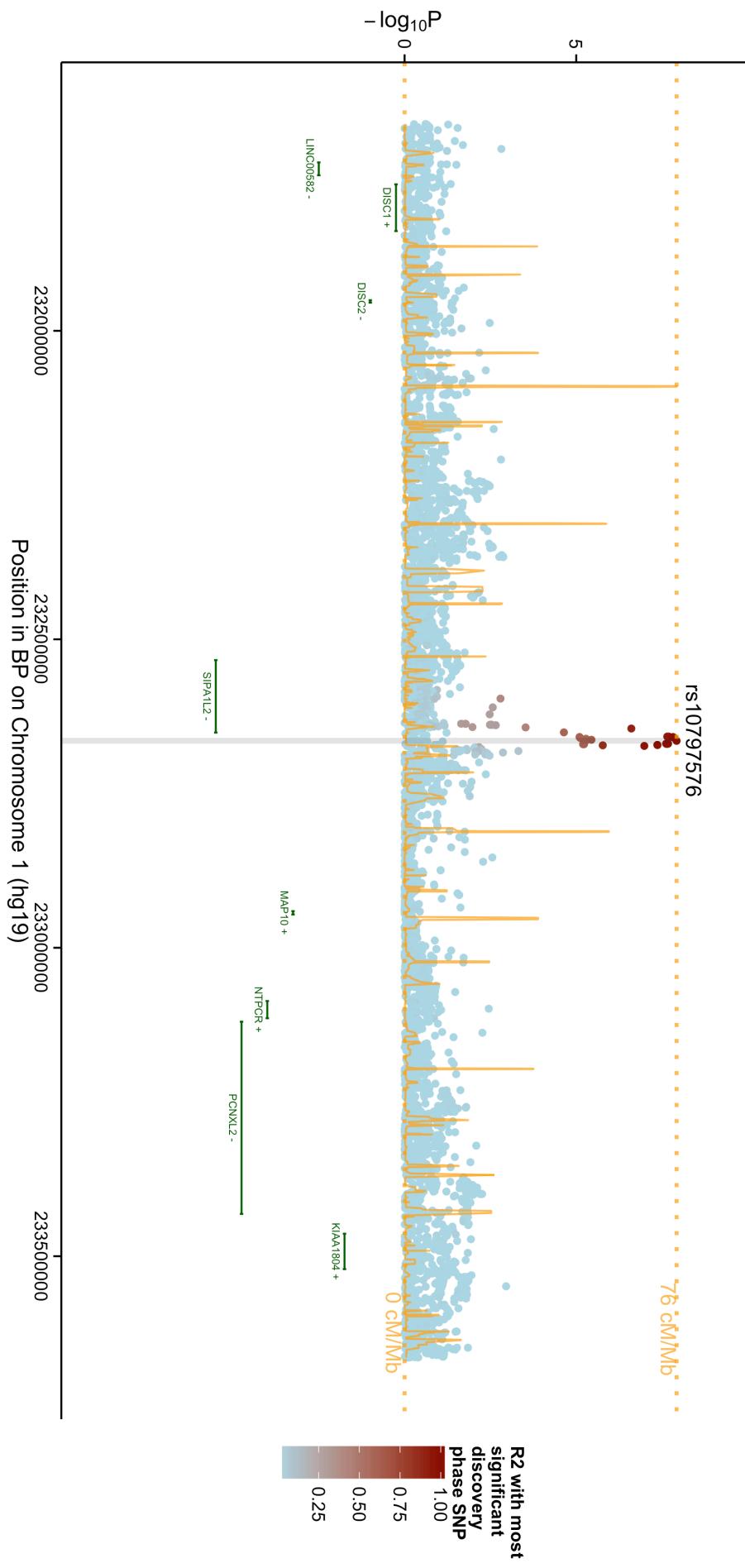
Secondary signals are annotated in text as per their description in the conditional analysis section of Table 2. Recombination rates are as per HapMap phase 2 European ancestry samples. Nearest gene or previously published proximal gene names included in table.



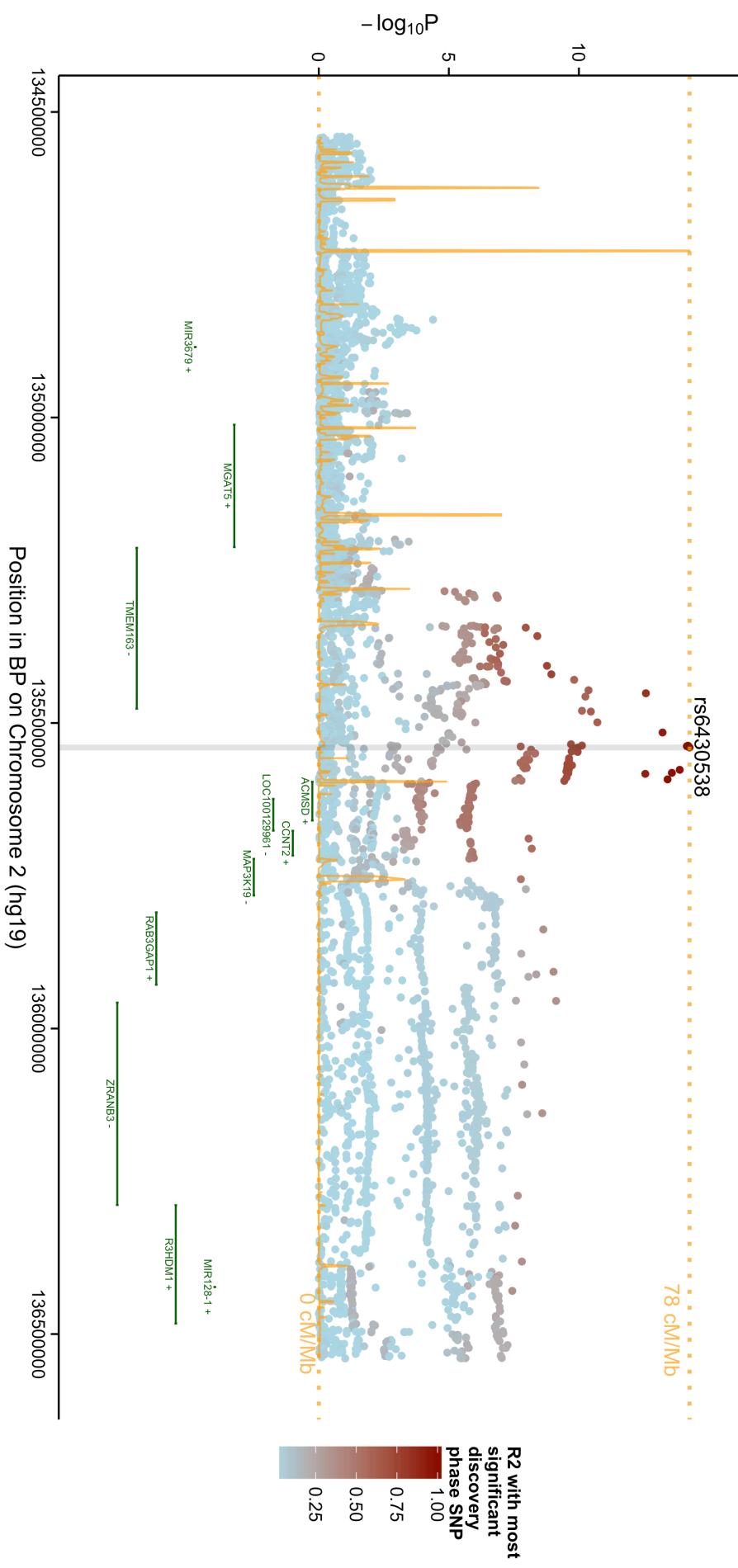
RAB7L1/NUCKS1 region



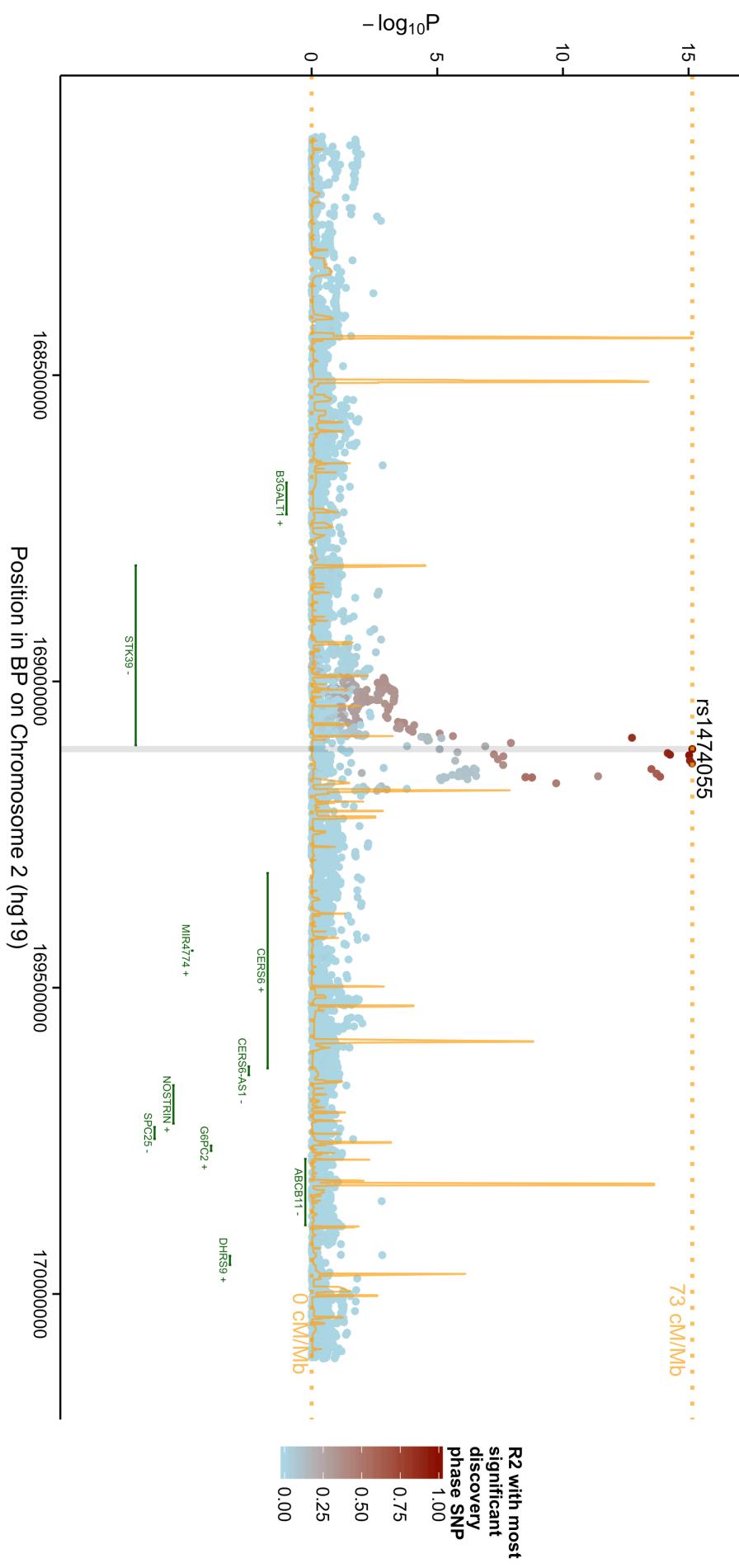
SIPA1L2 region



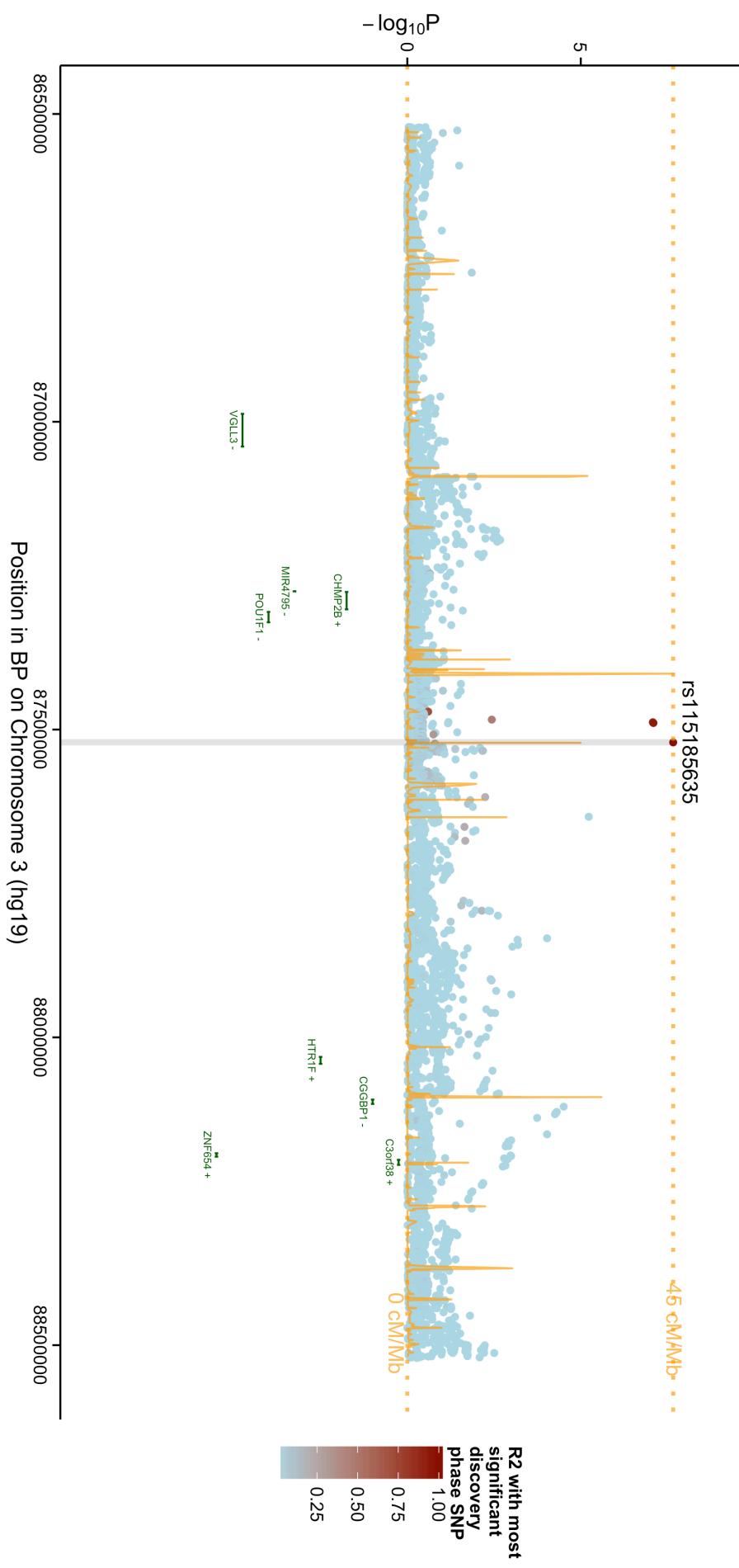
ACMSD/TMEM163 region

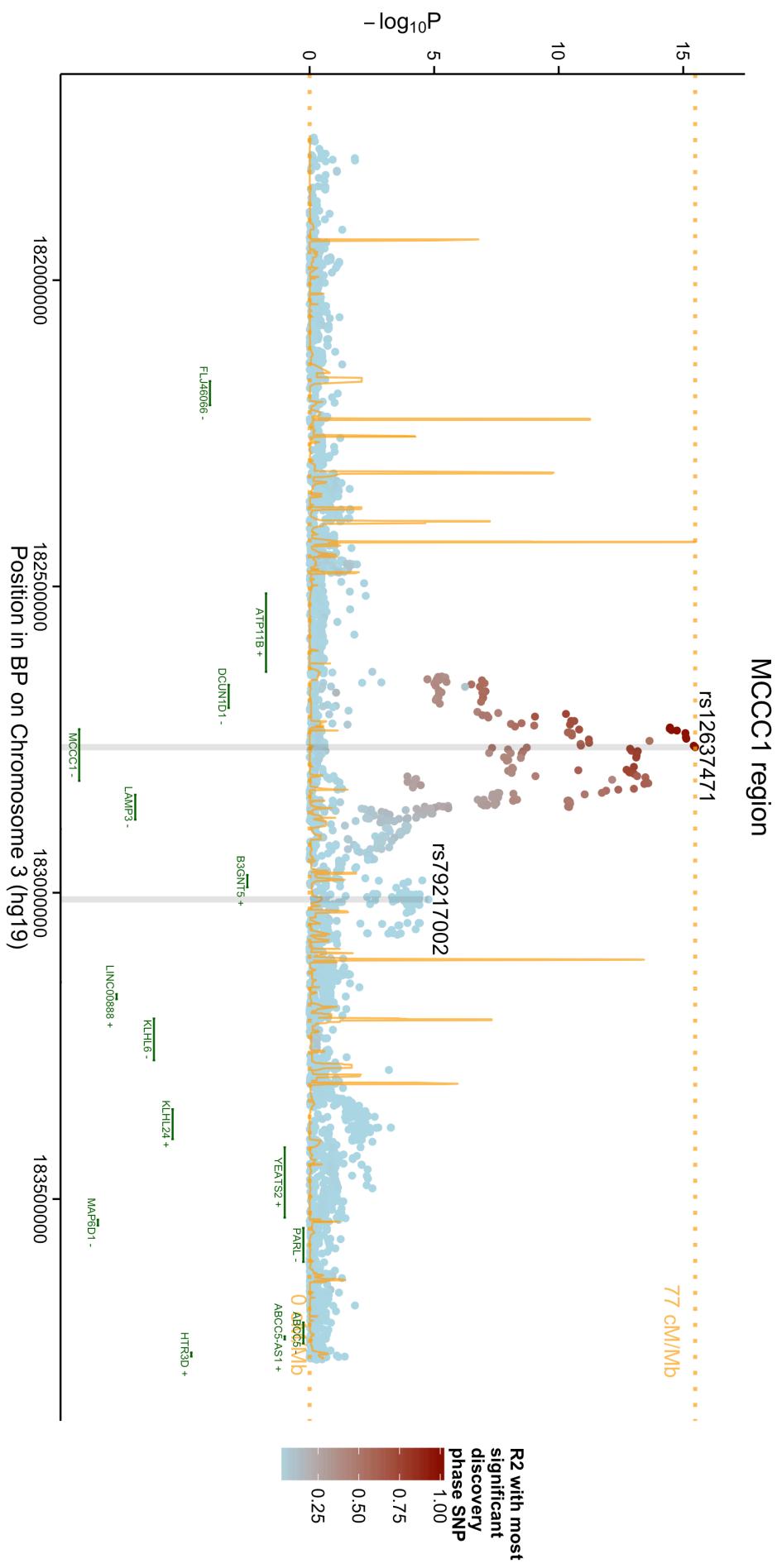


STK39 region



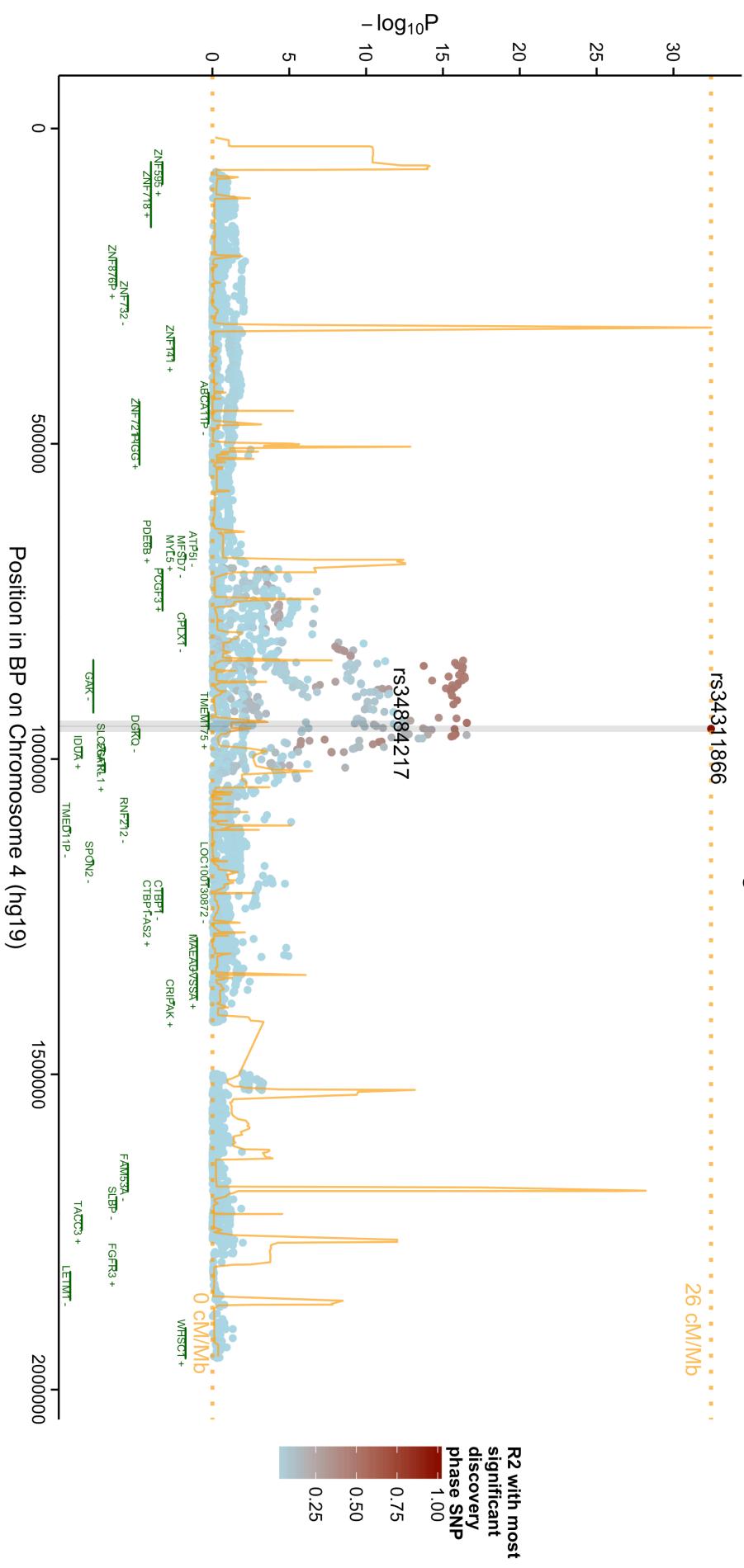
KRT8P25/APOOP2 region



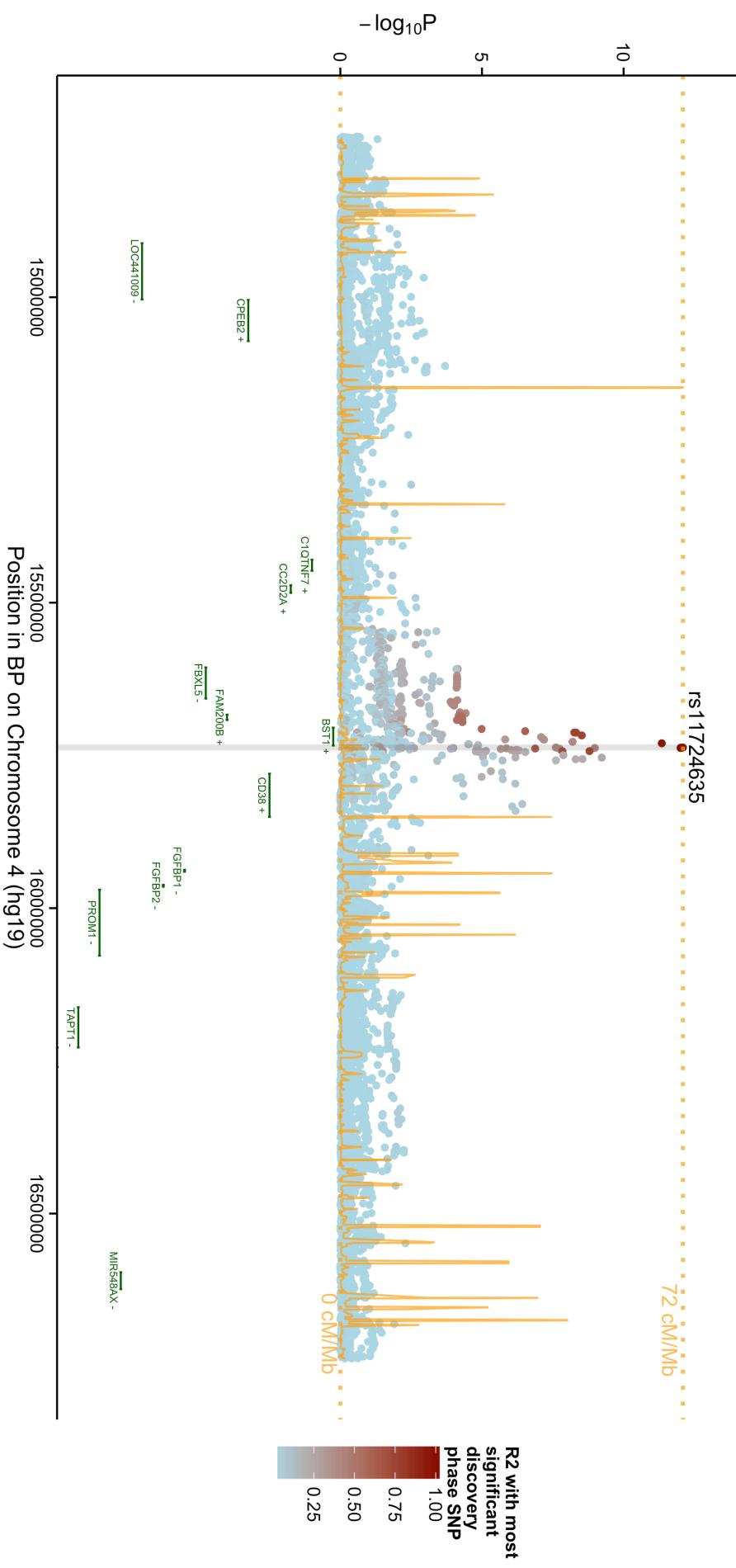


TMEM175/GAK/DGKQ region

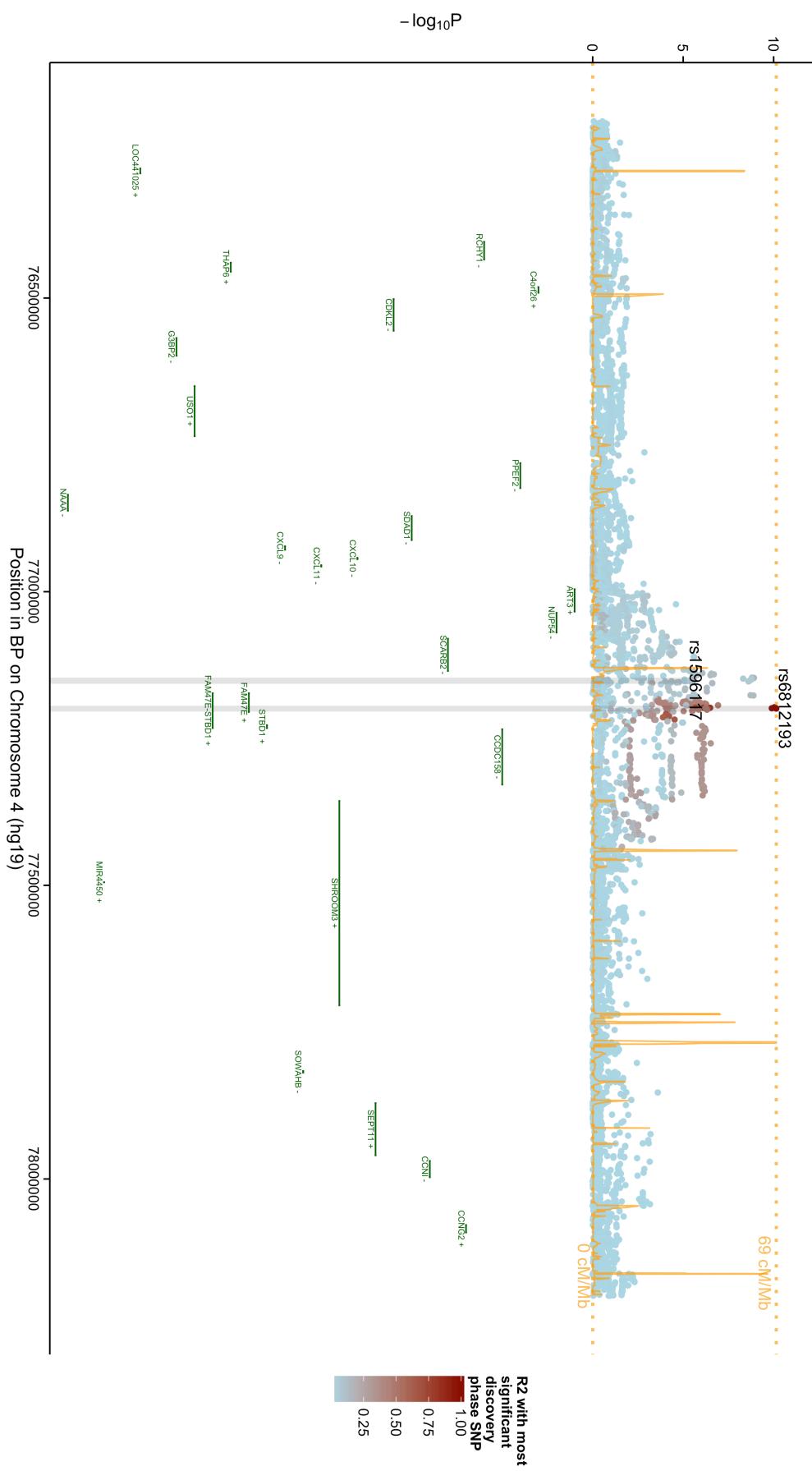
rs34311866 26 cM/Mb

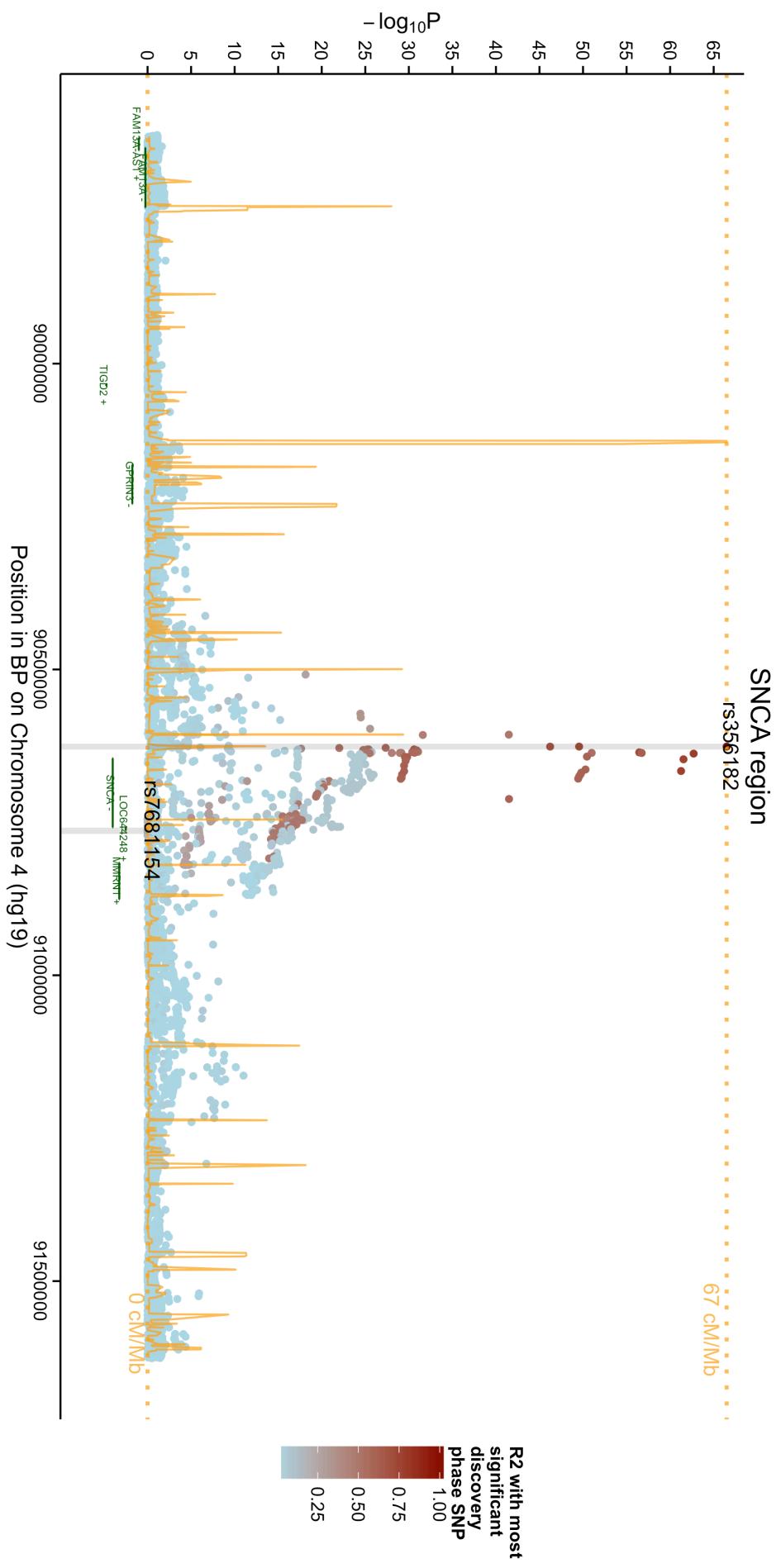


BST1 region

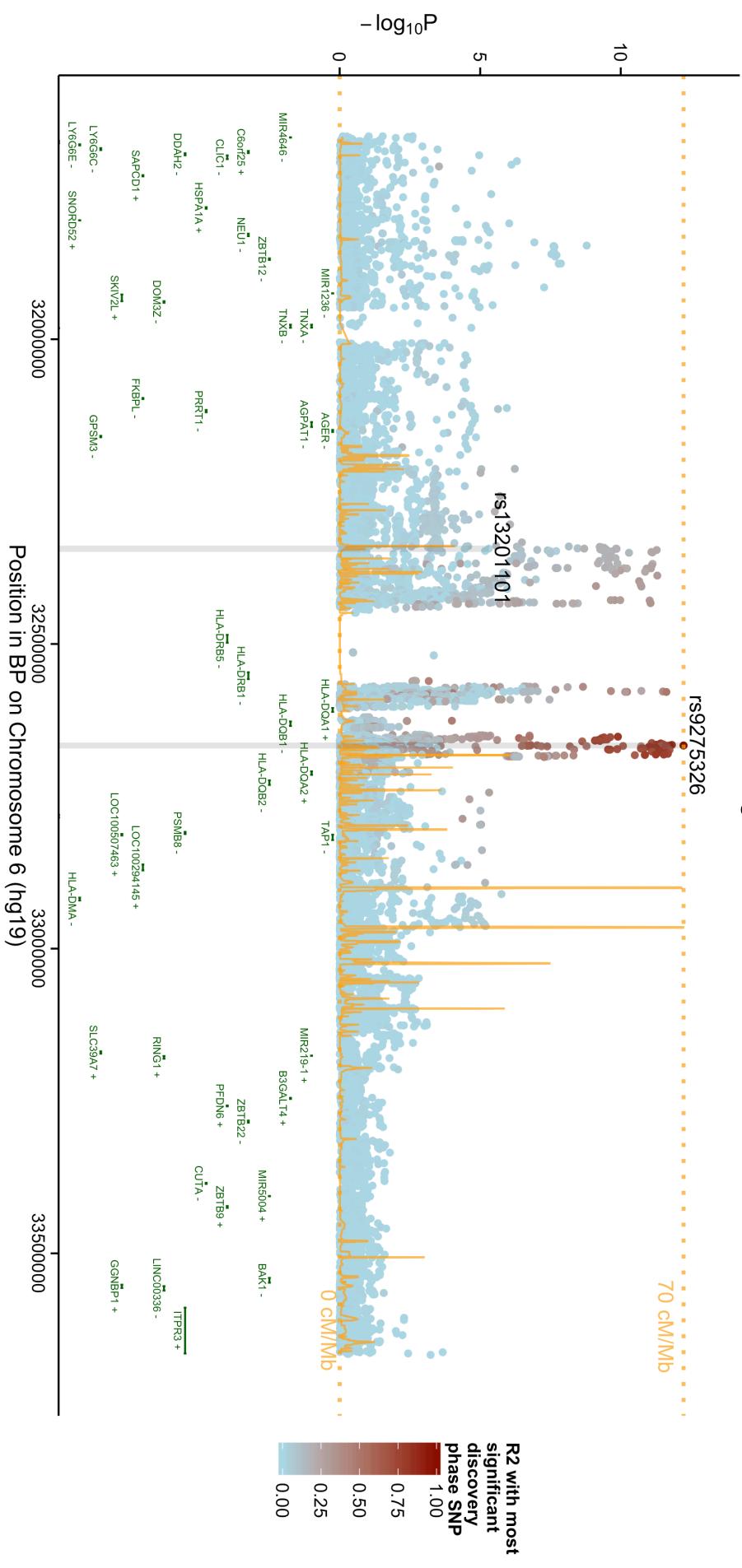


FAM47E/SCARB2 region

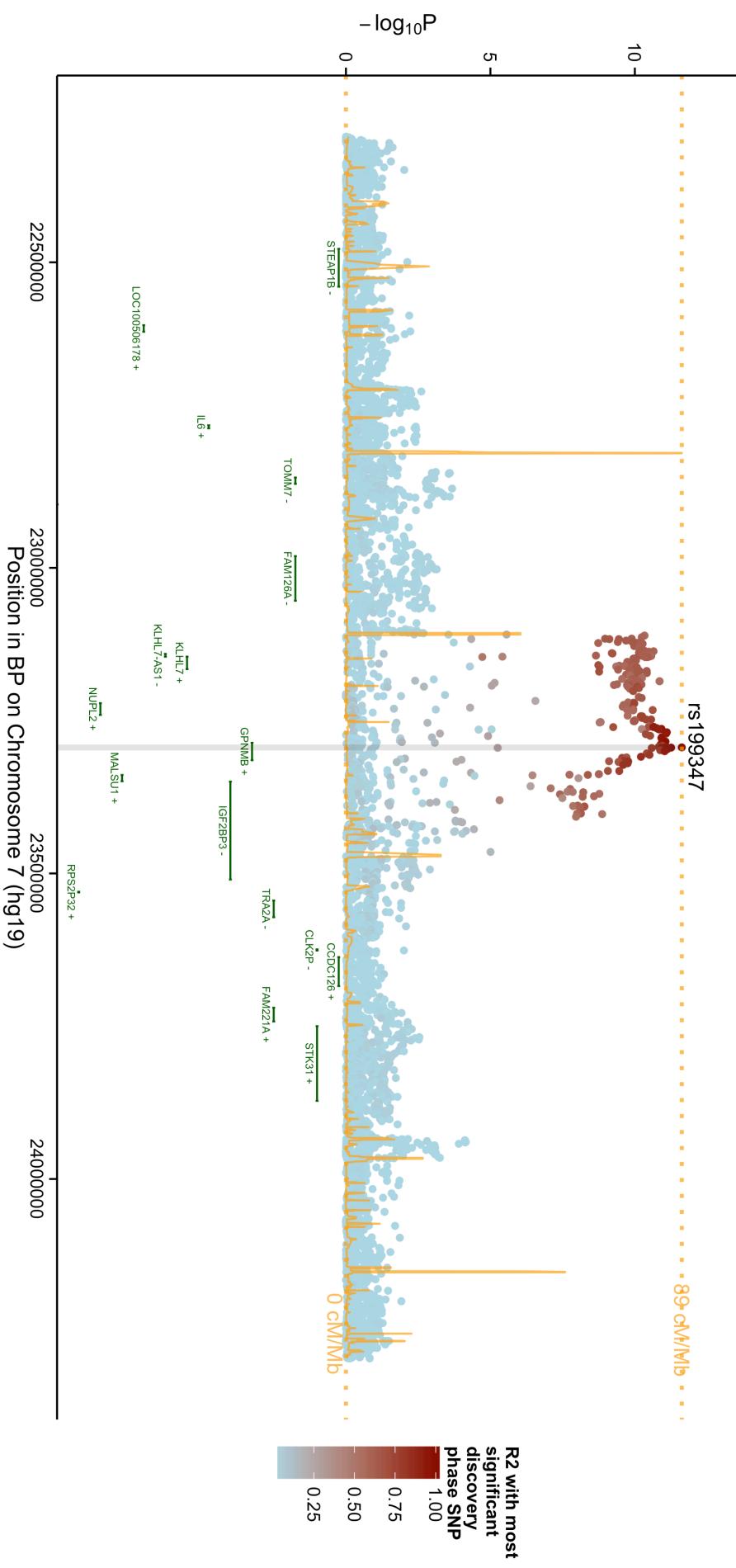




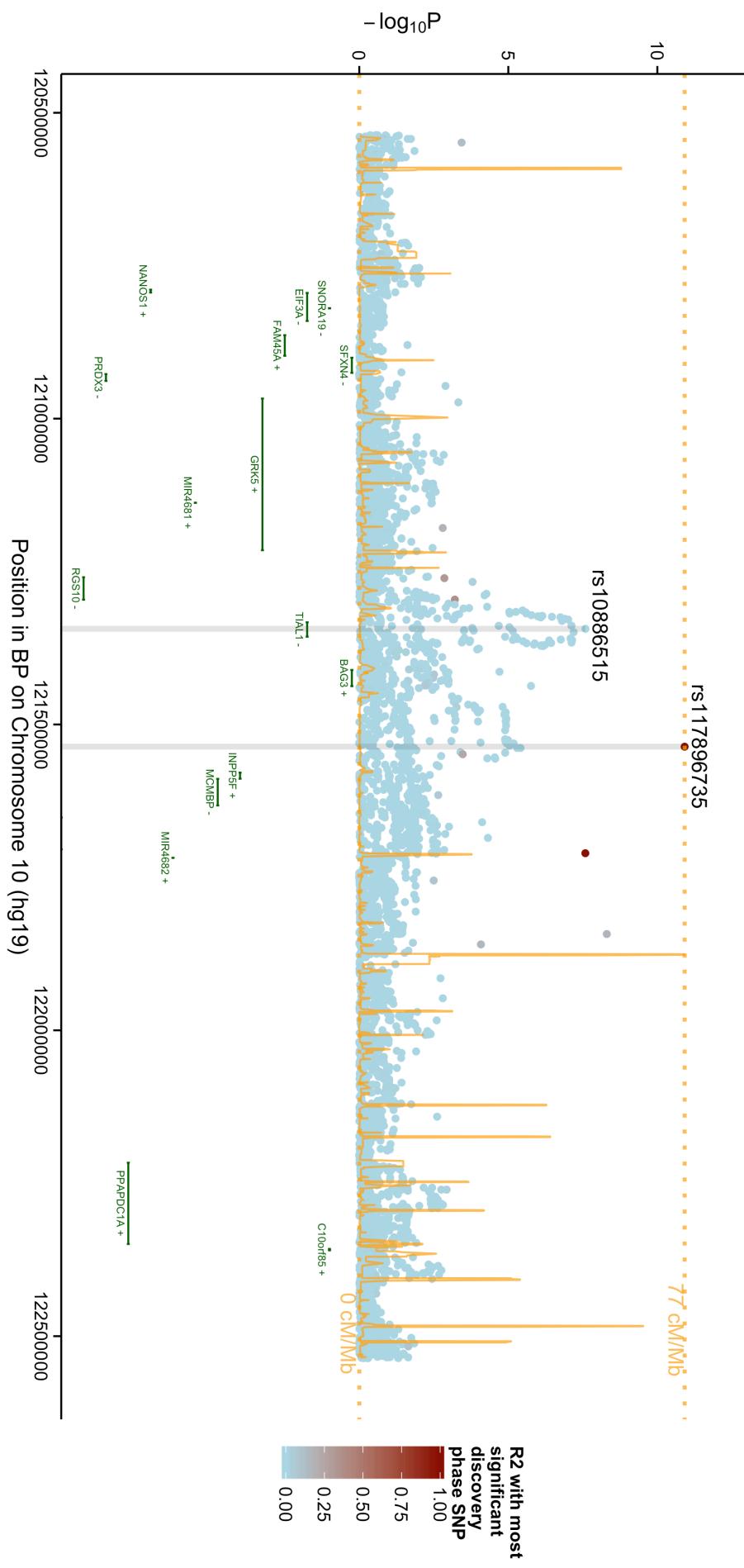
HLA-DQB1 region



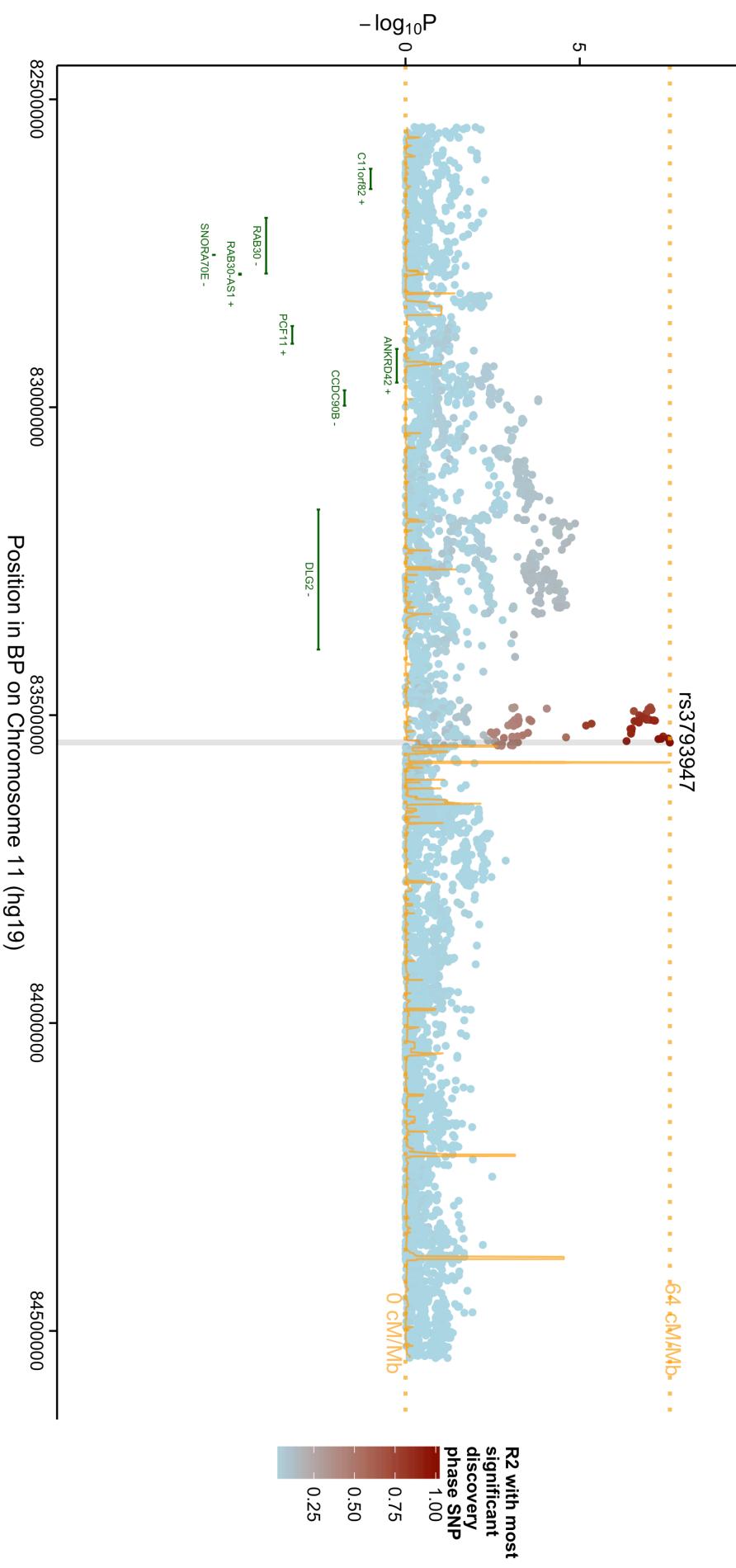
GPNMB region



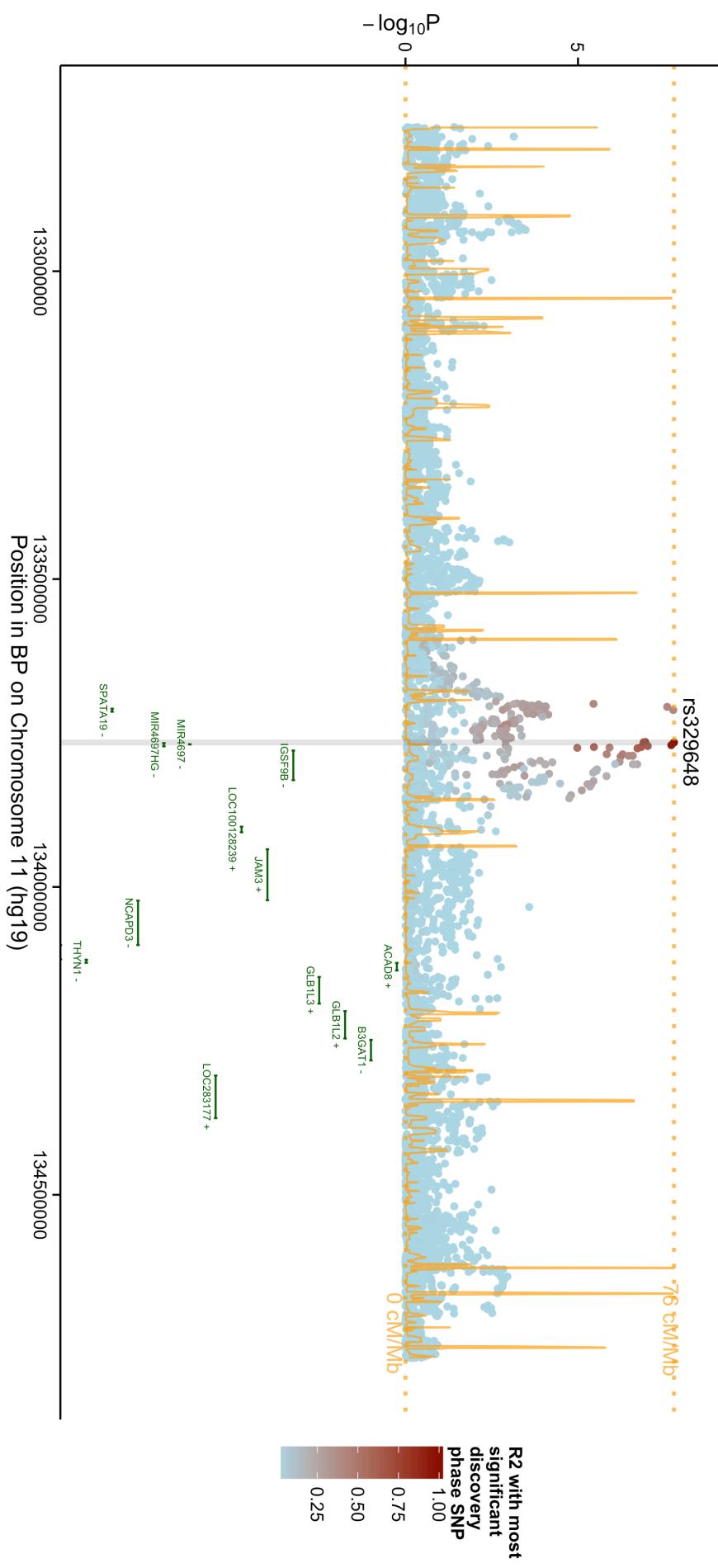
INPP5F region

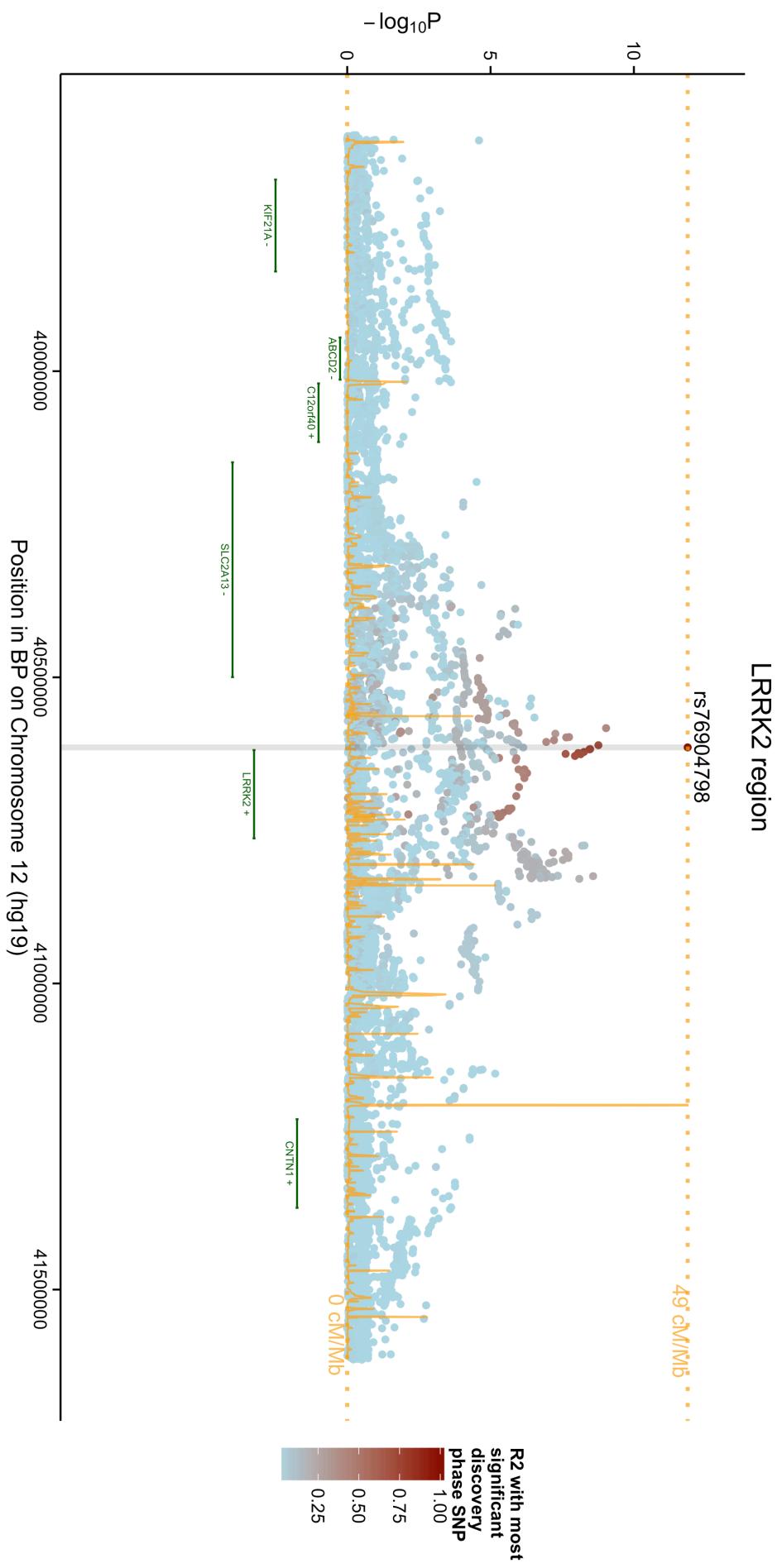


DLG2 region

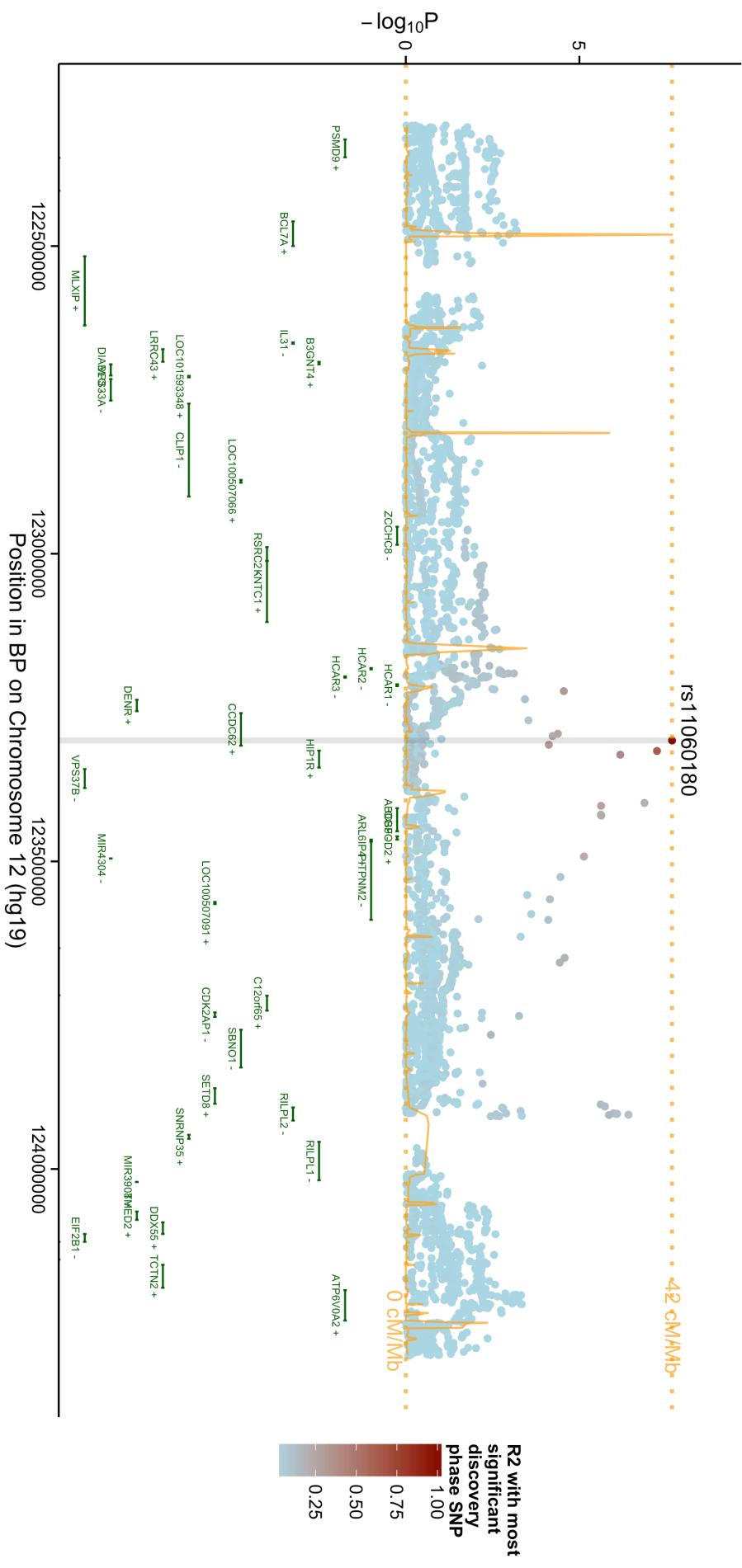


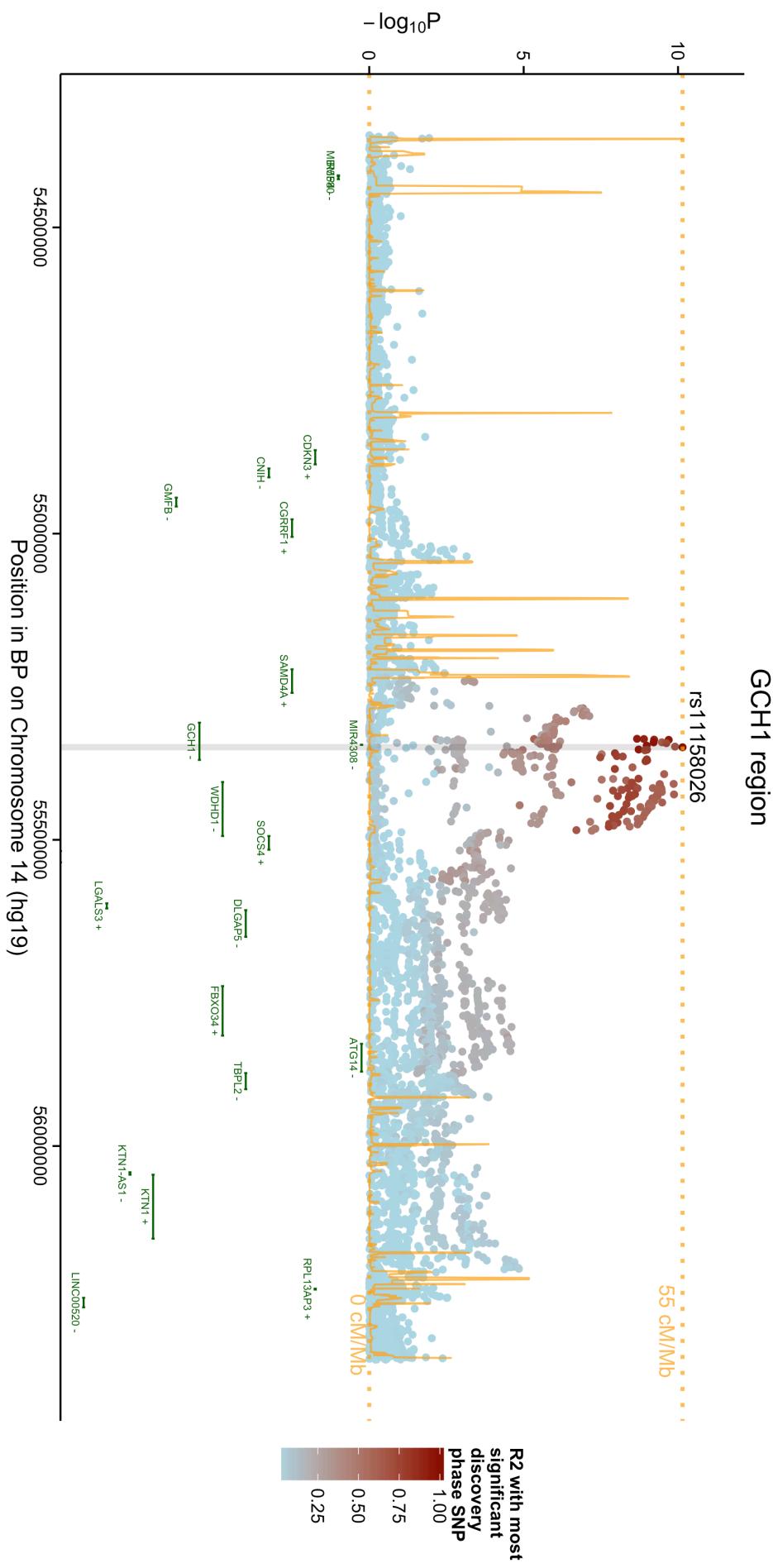
LOC283174 region



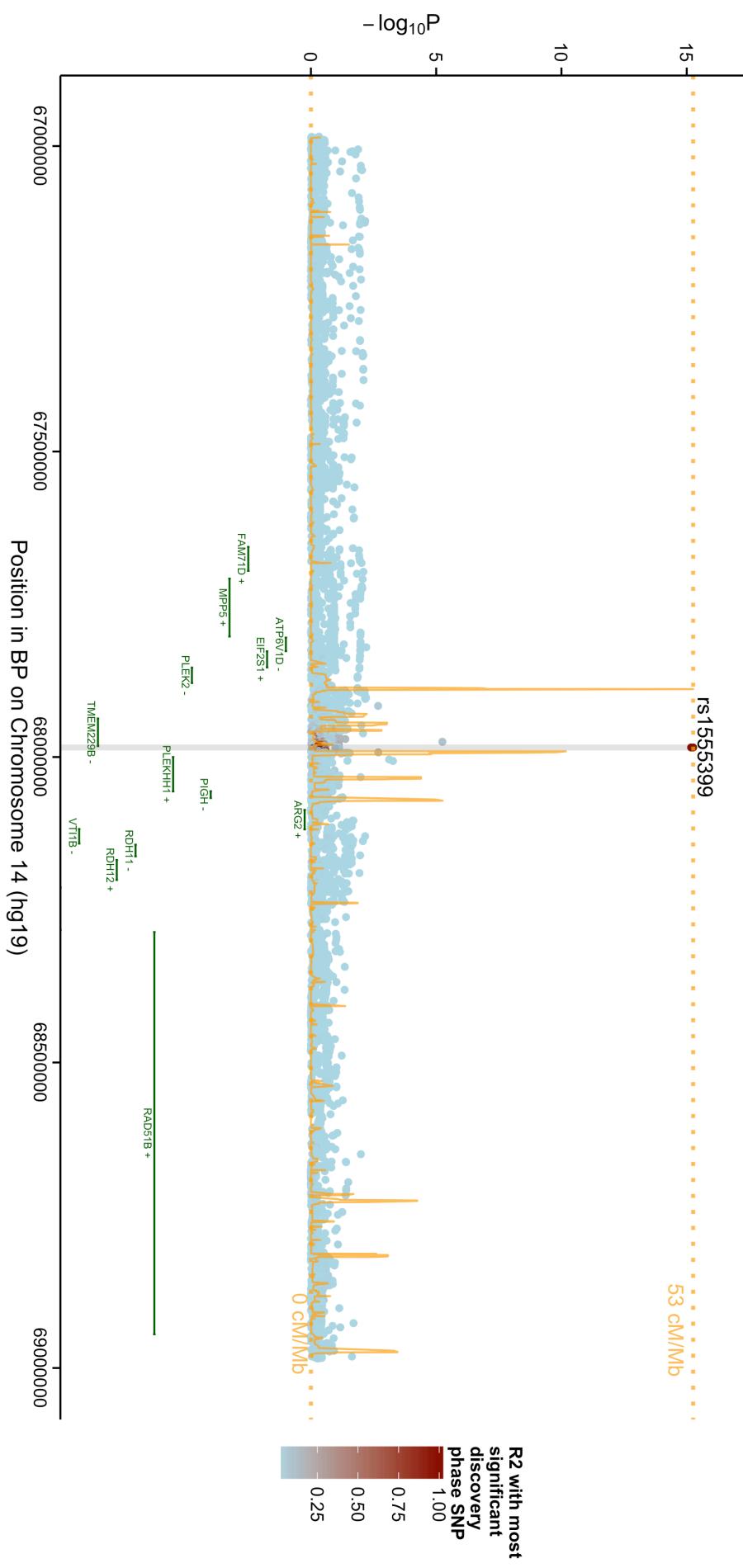


CCDC62 region

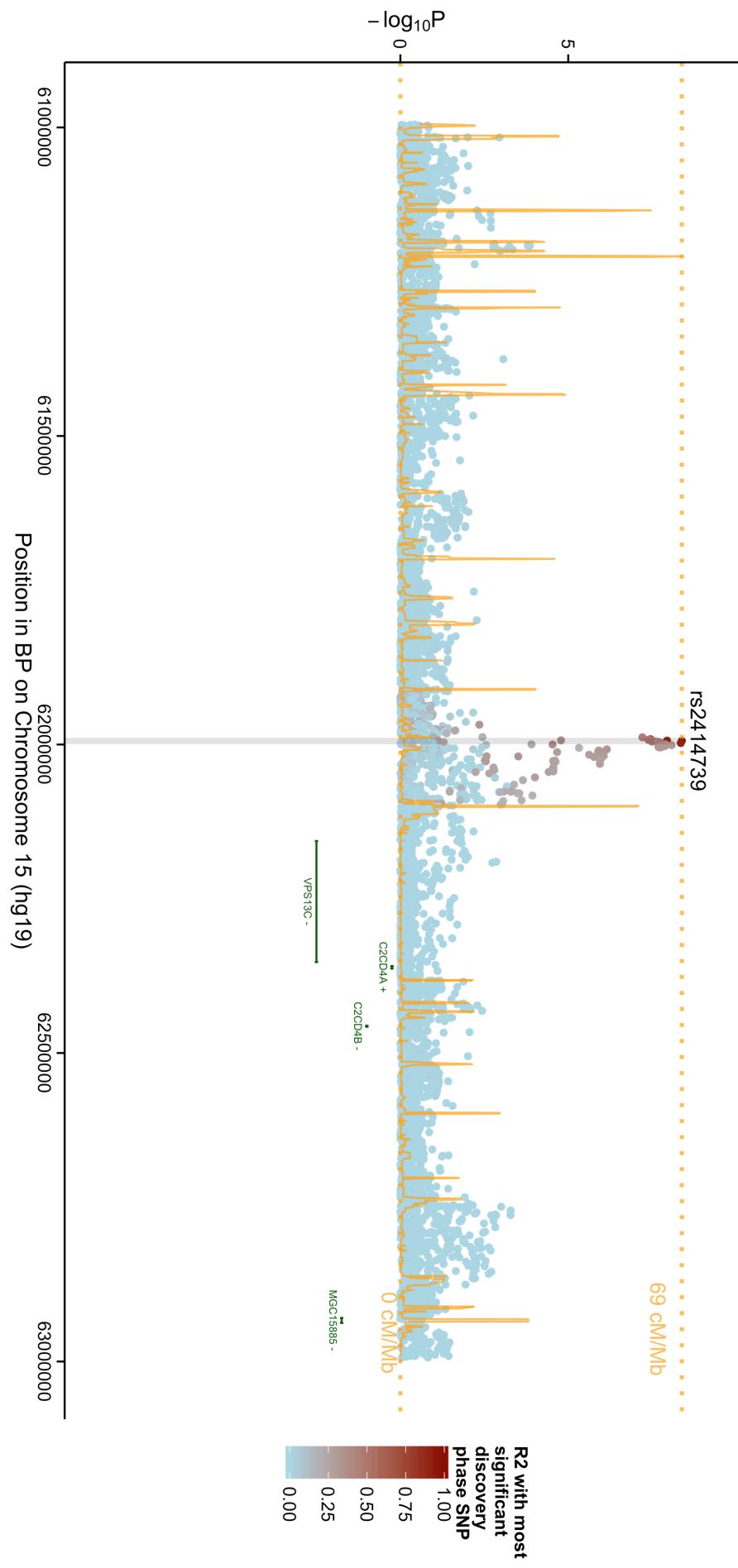




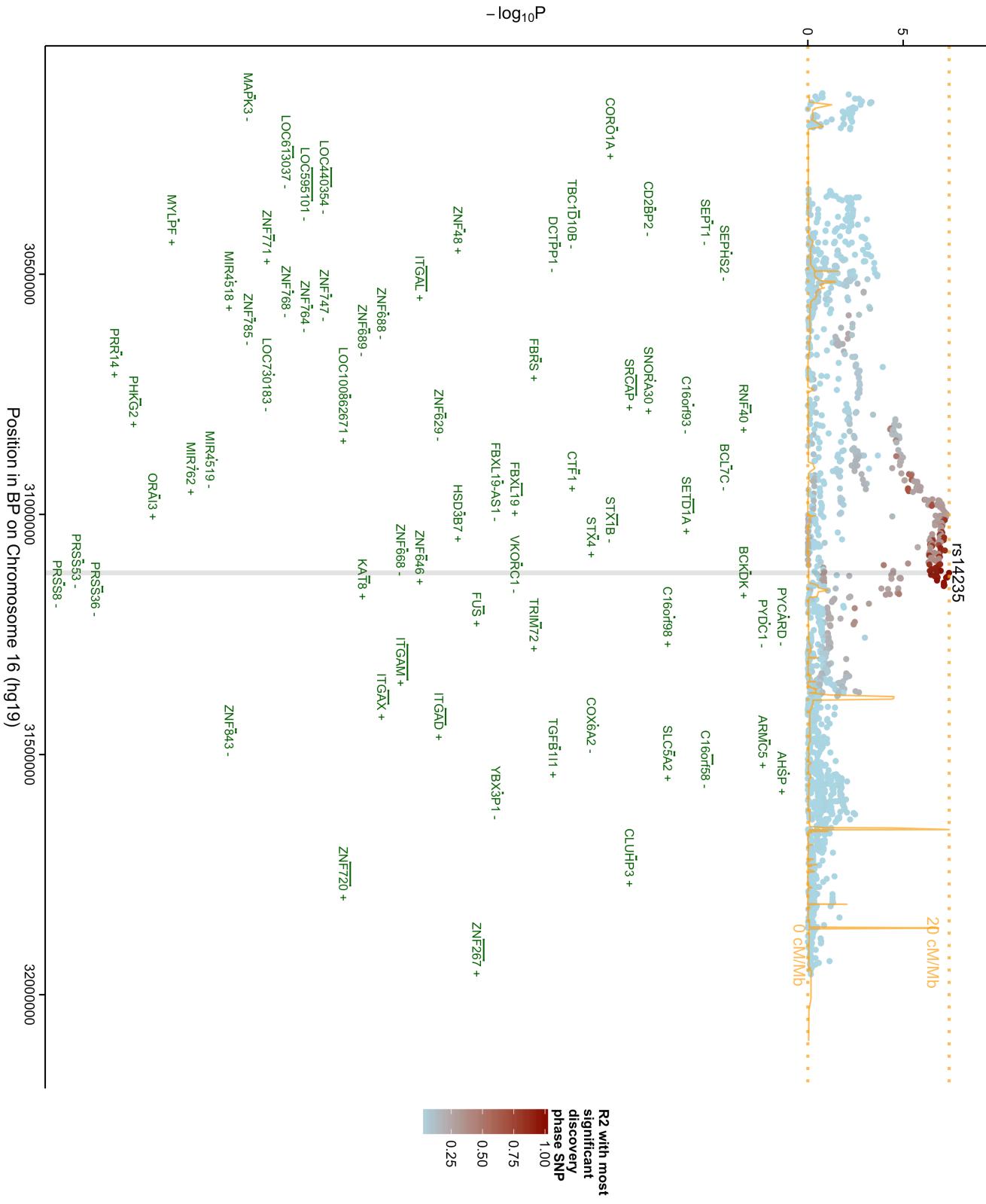
TMEM229B region

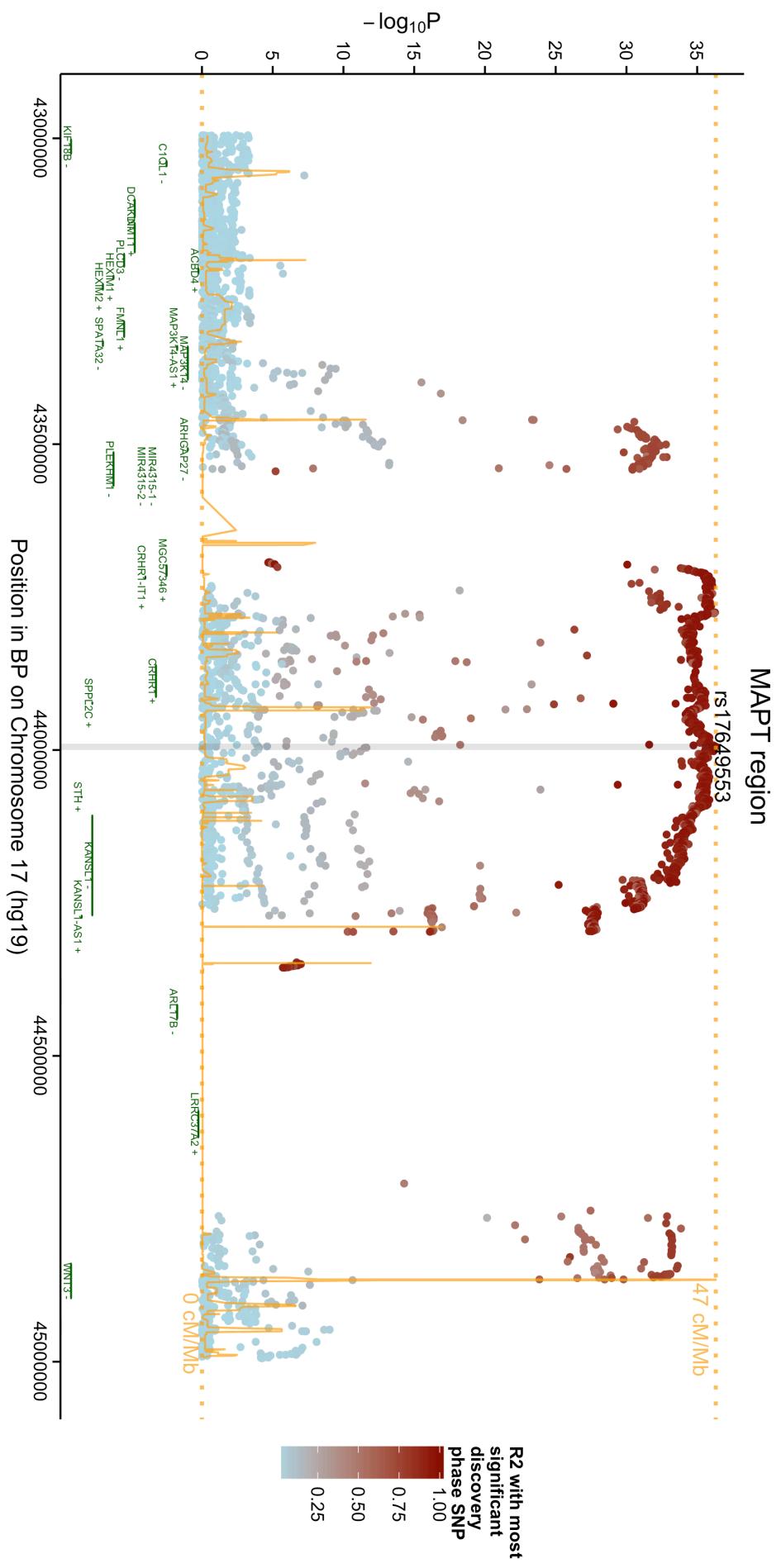


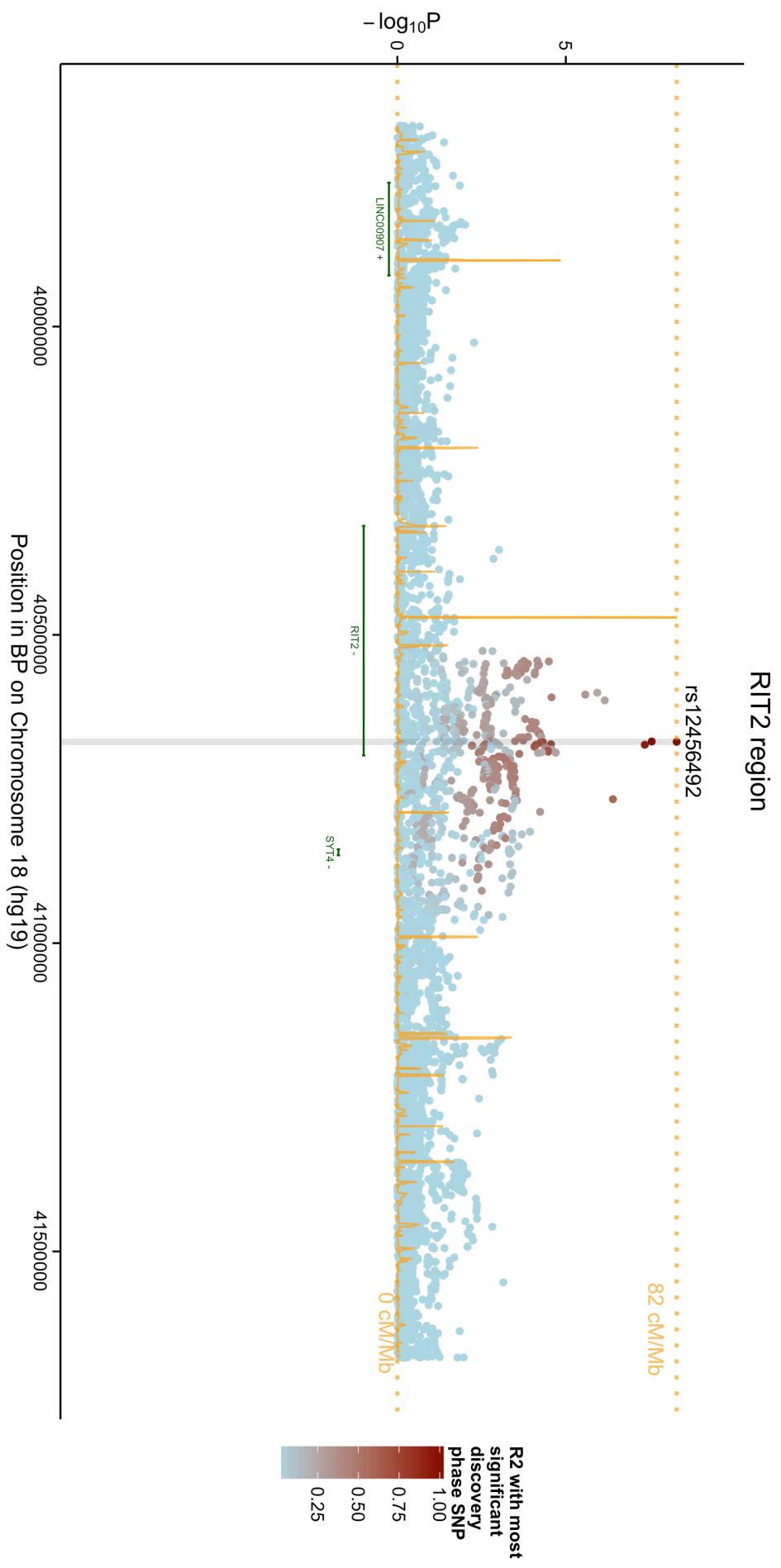
VPS13C region



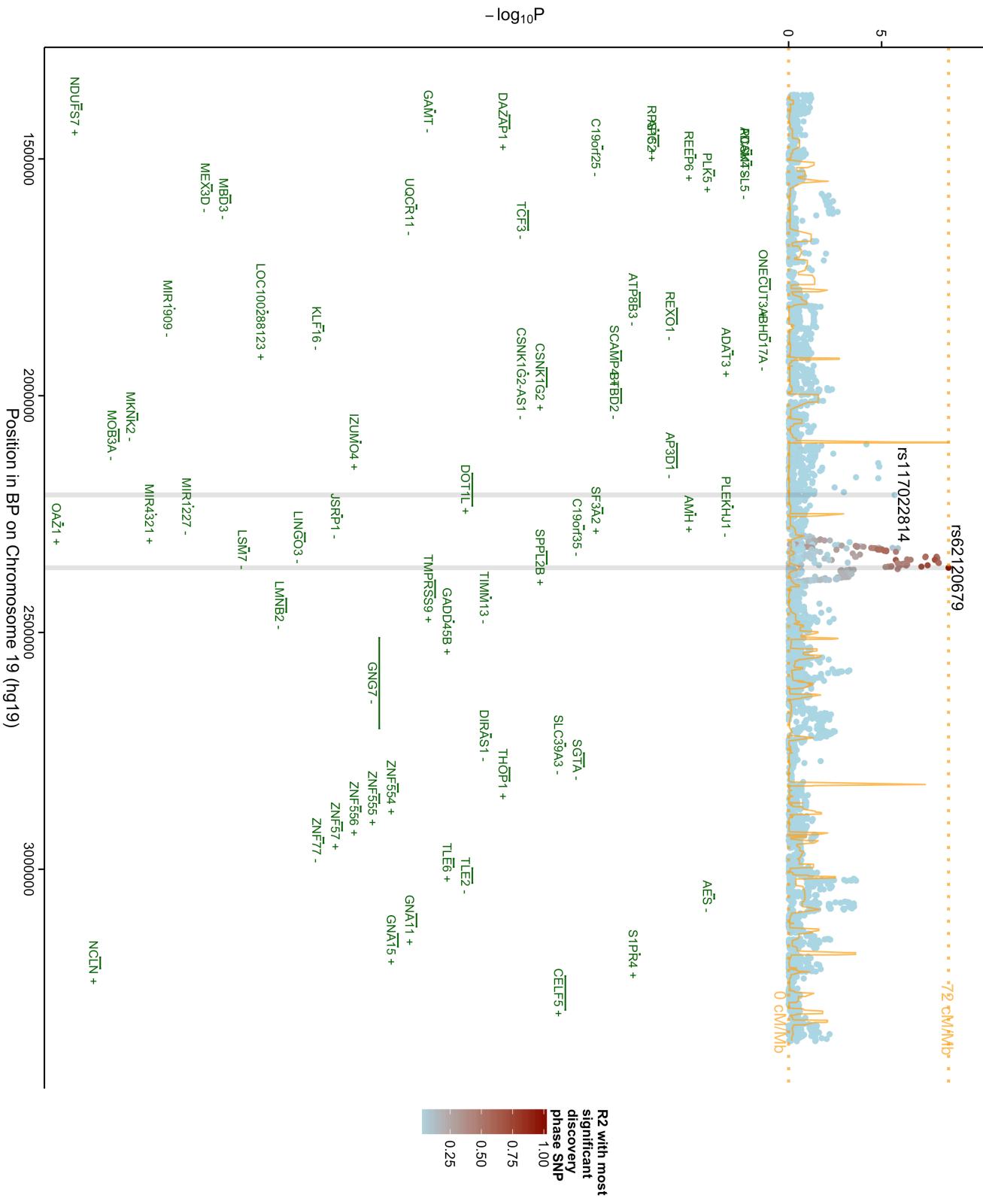
BCKDK/ STX1B region

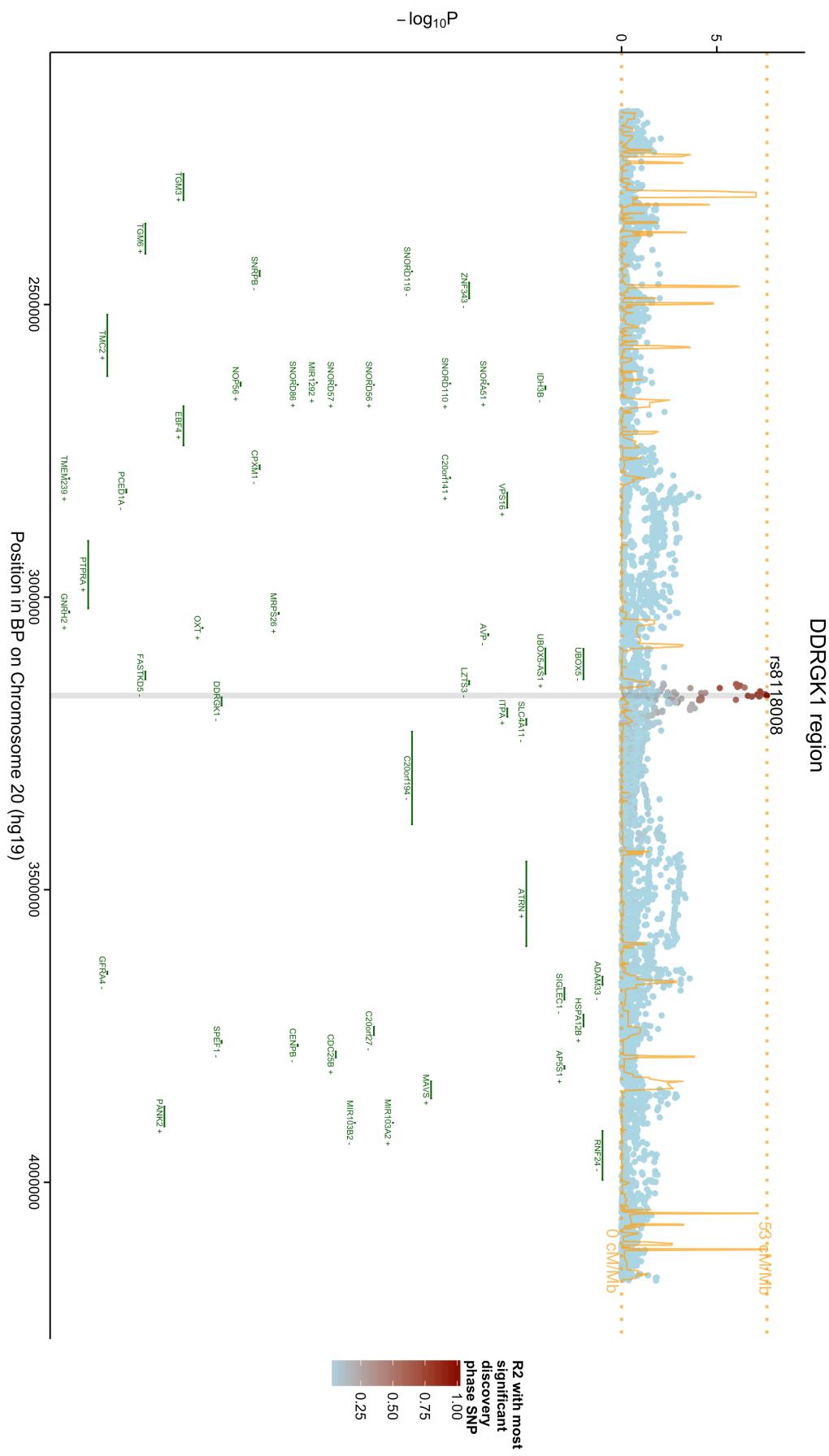




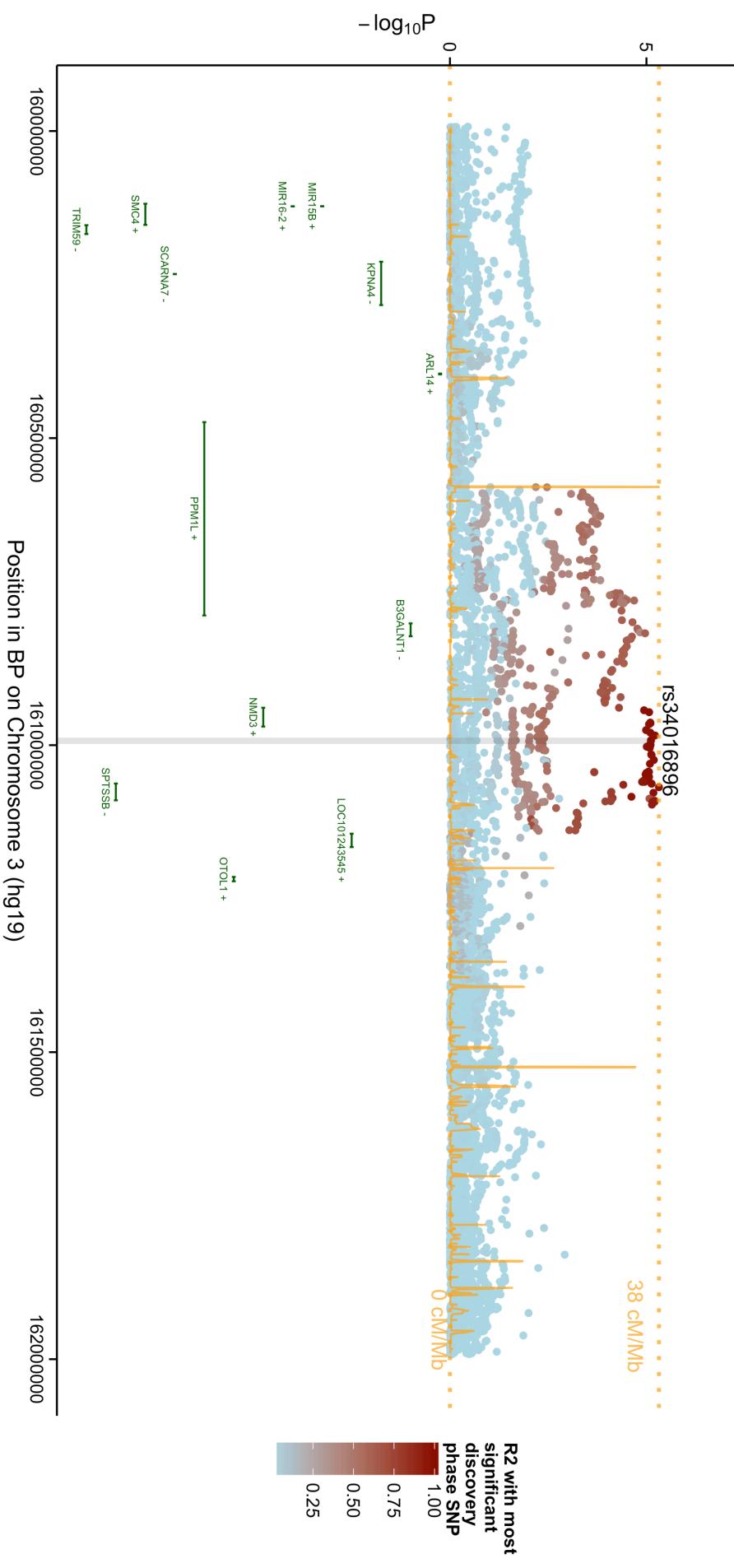


SPPL2B region

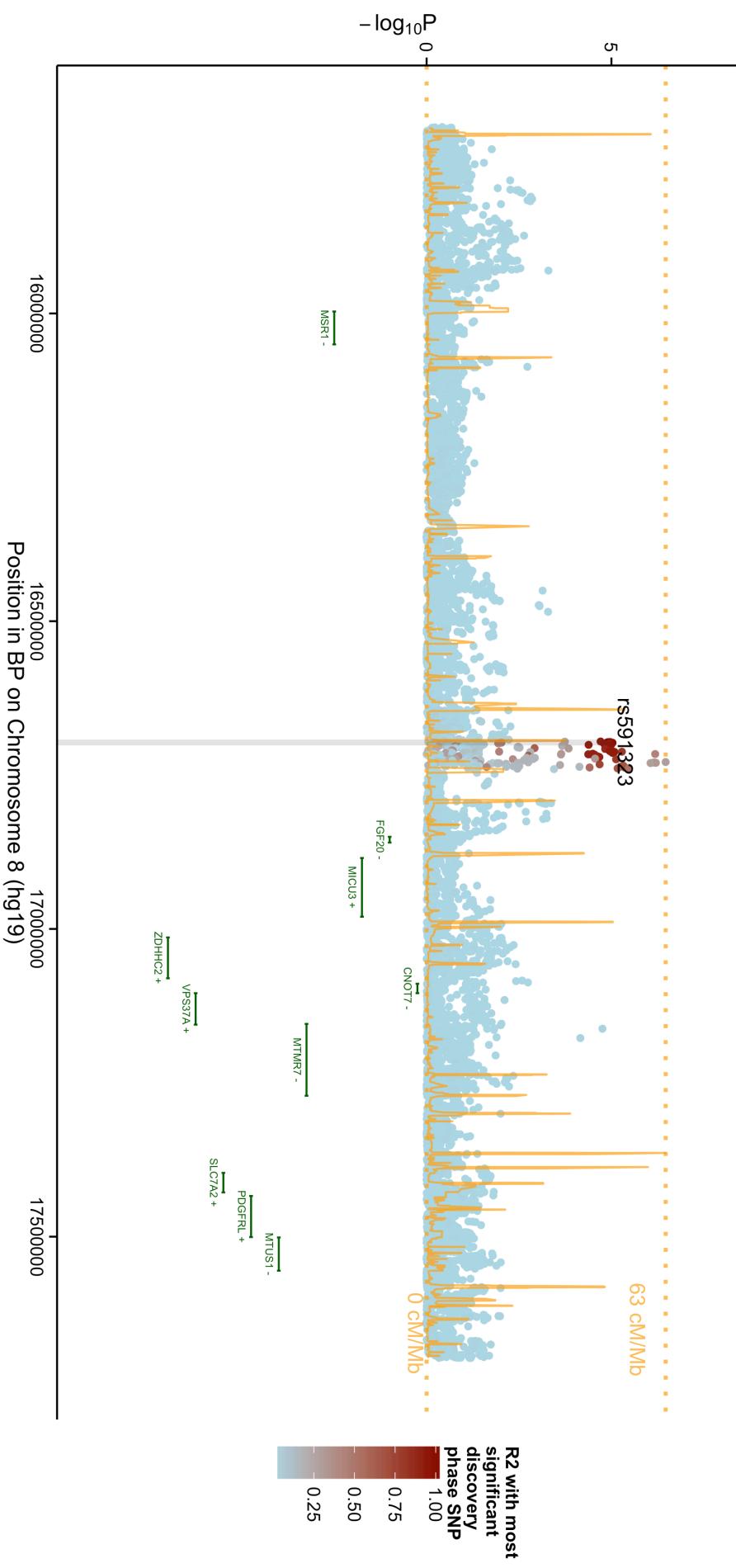




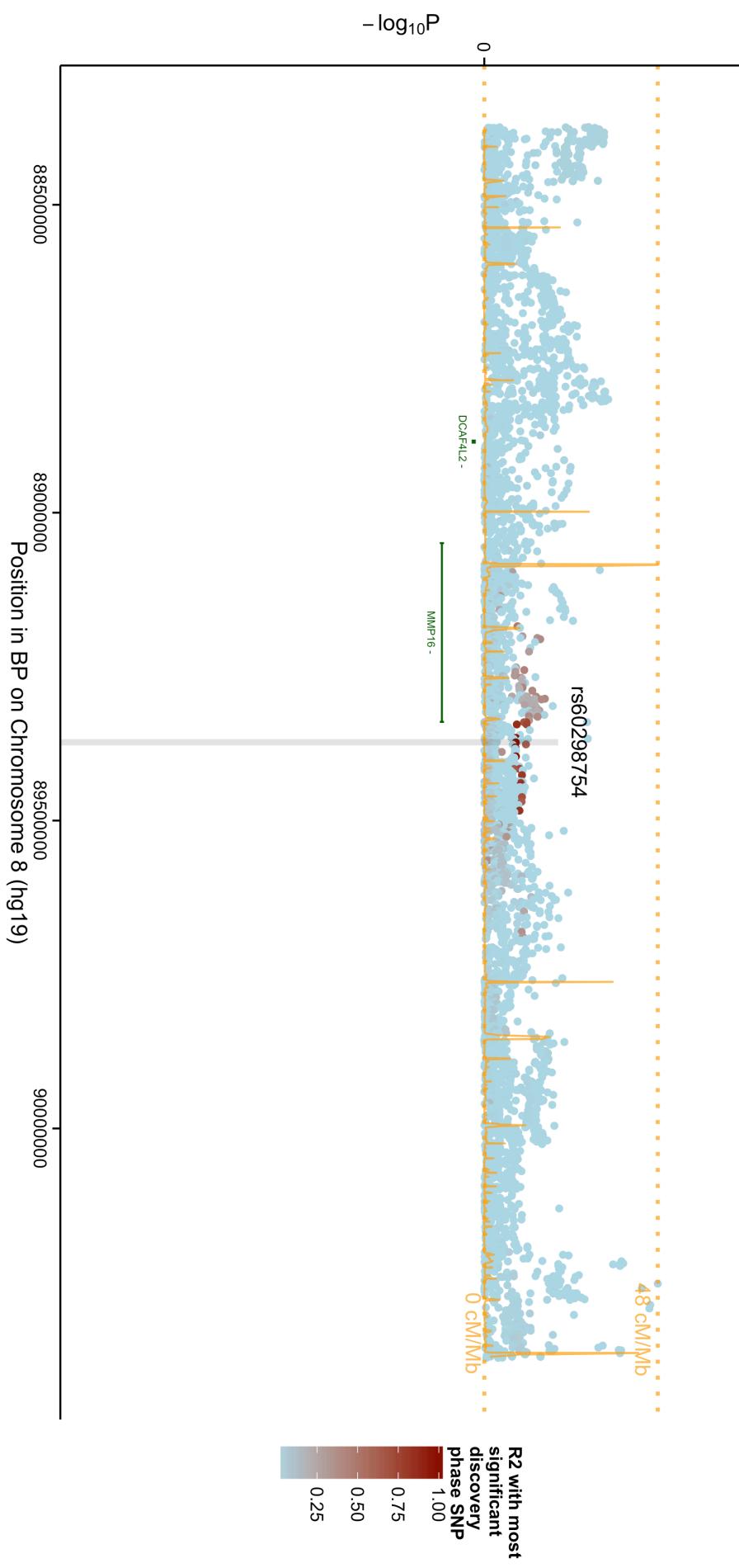
NMD3 region



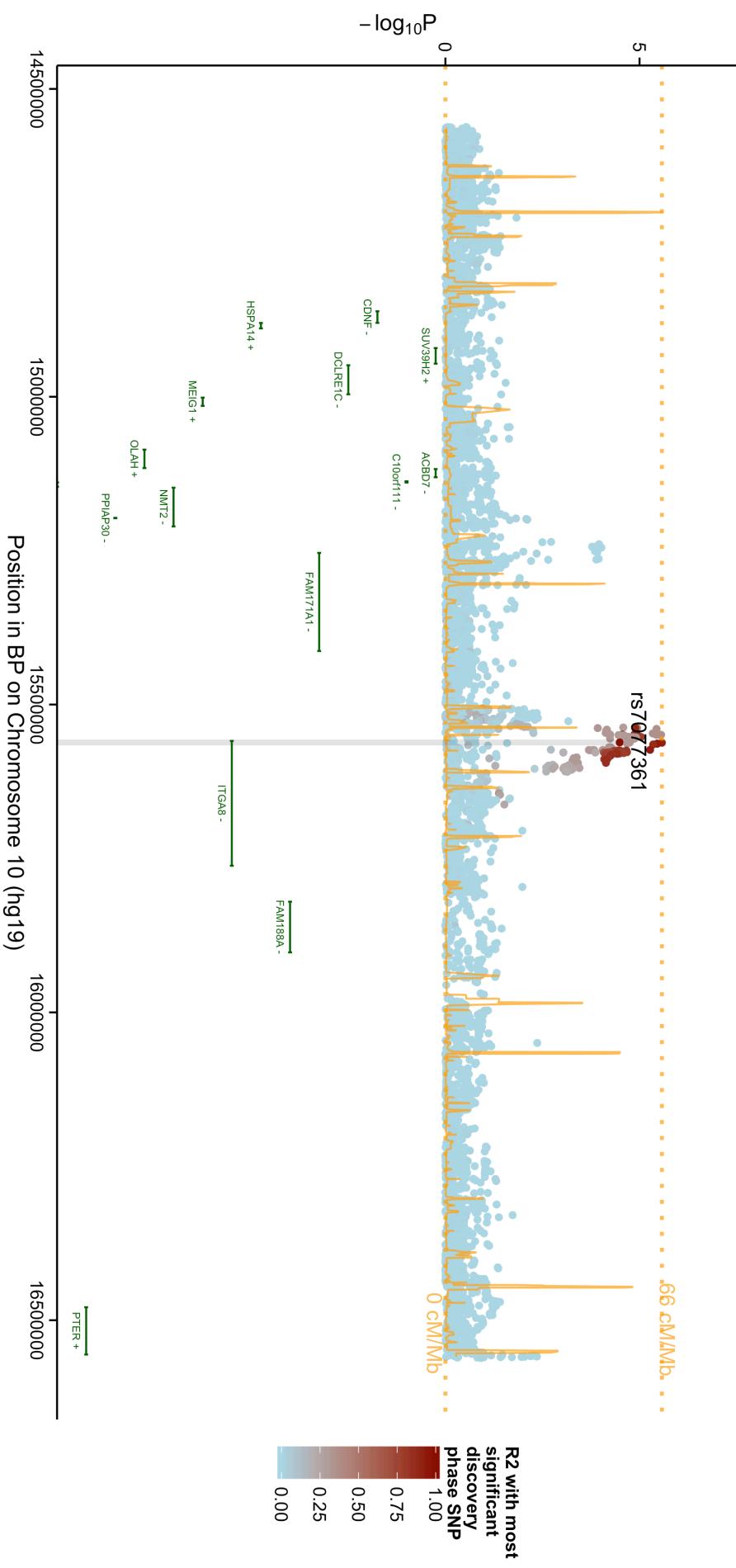
FGF20 region



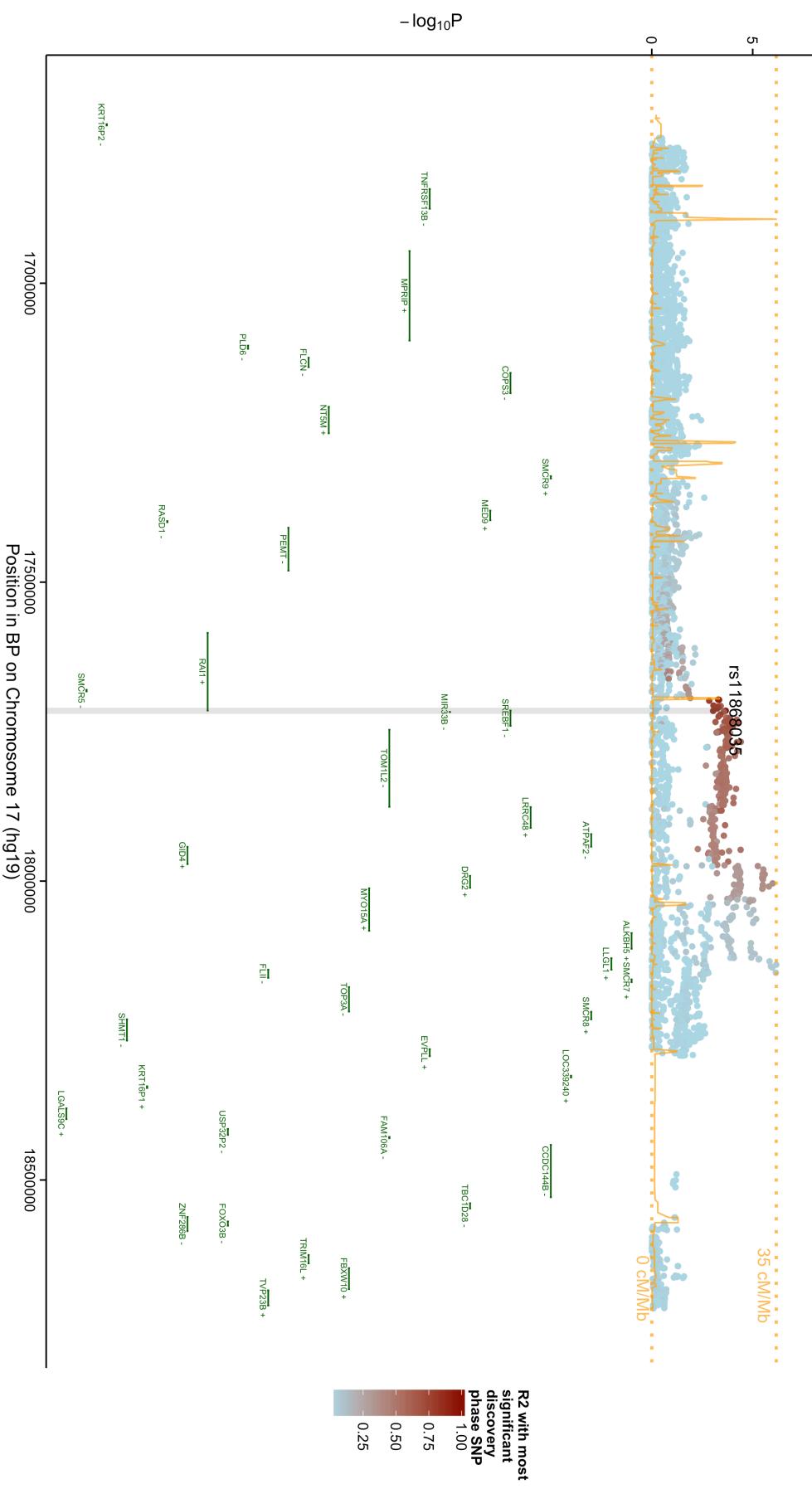
MMP16 region



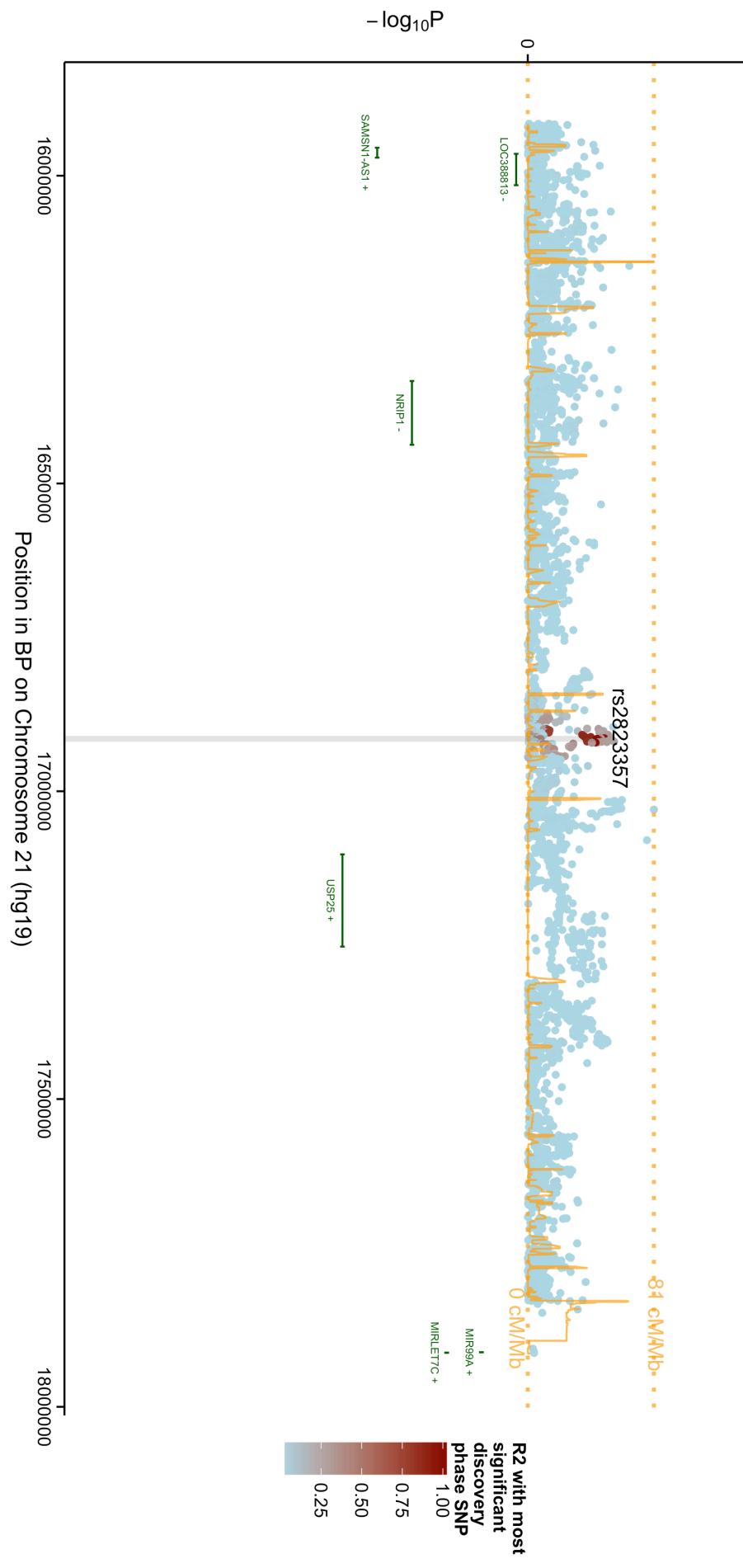
|TGA8 region



SREBF/RAI1 region

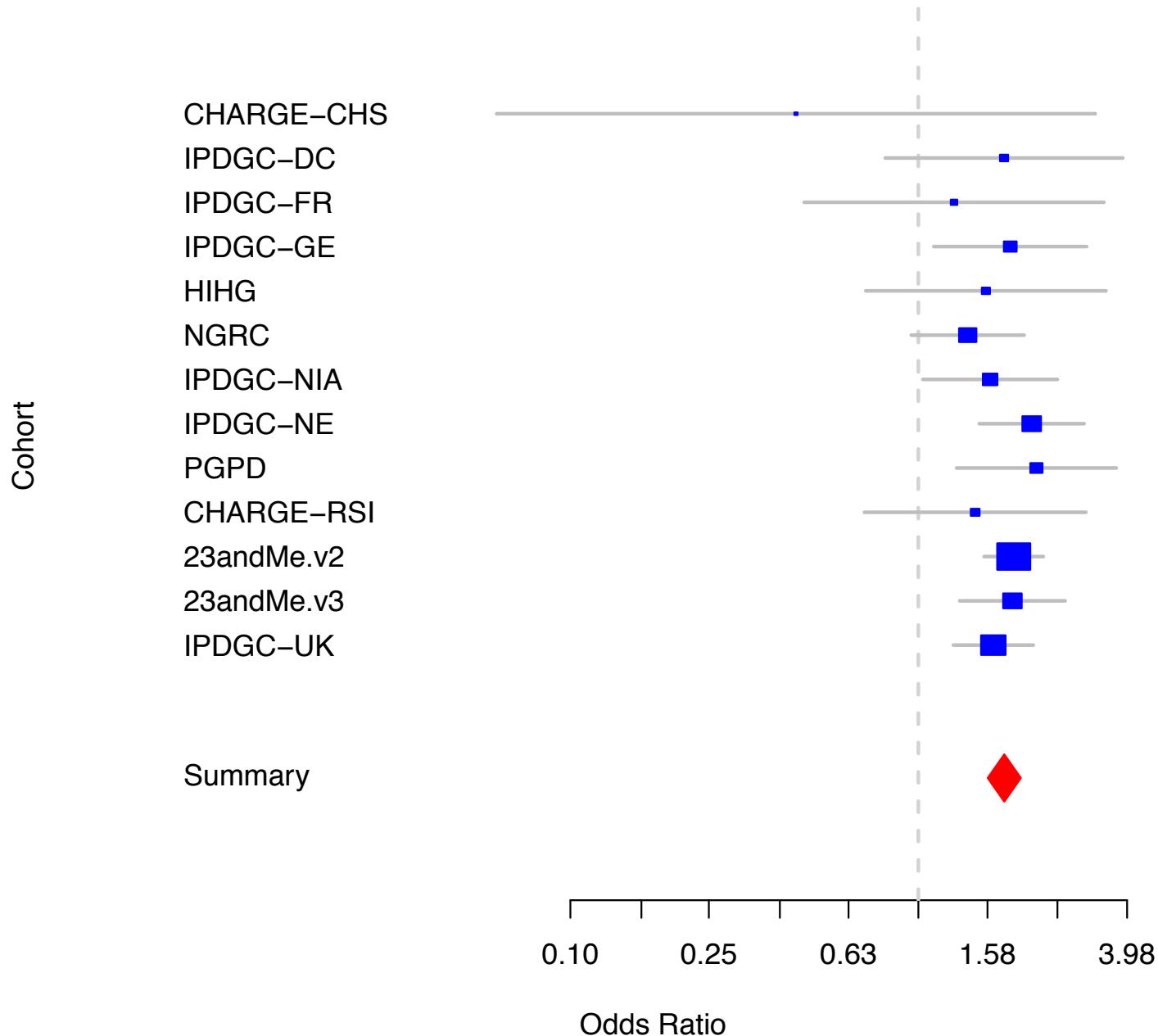


USP25 region

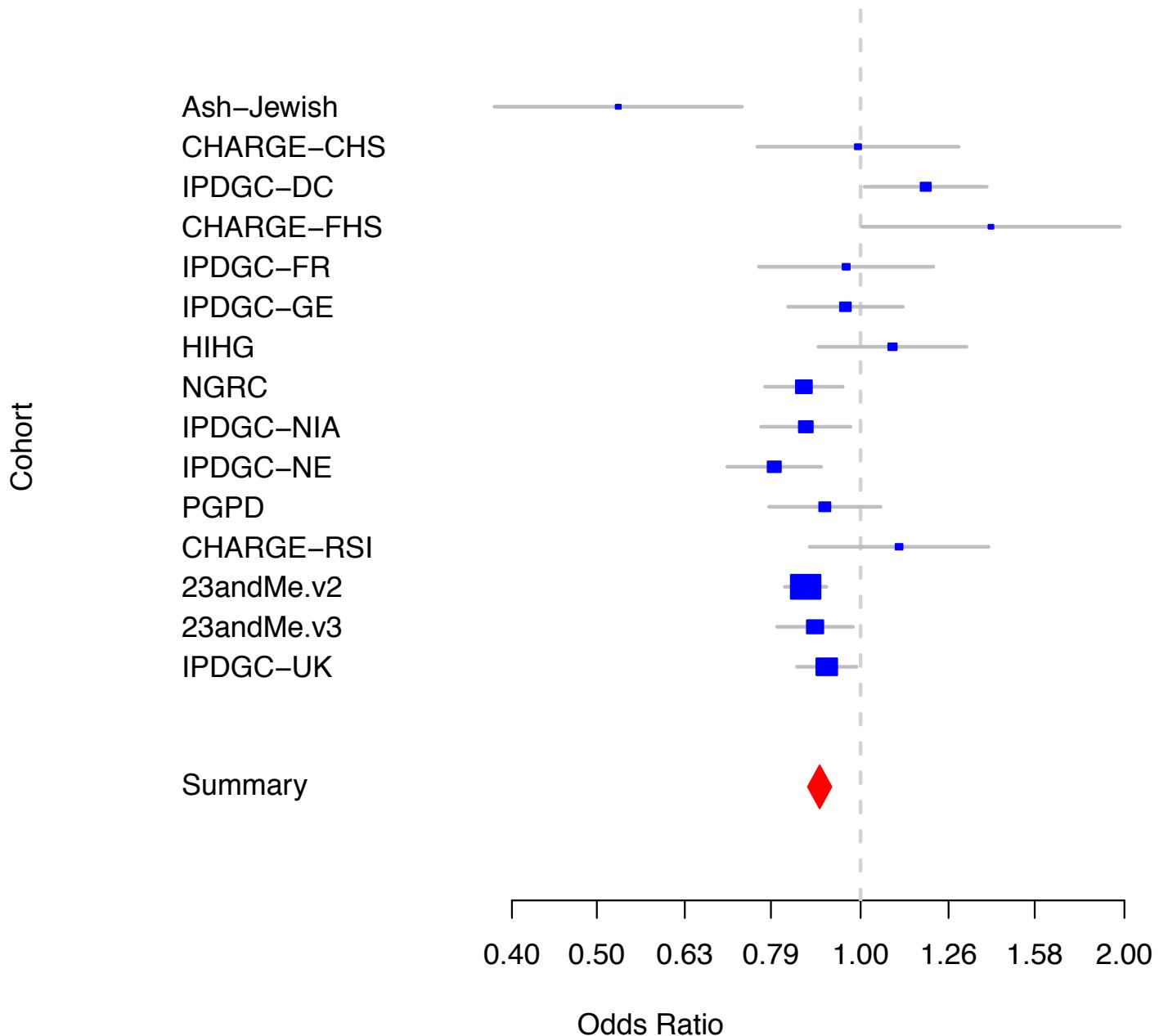


Supplementary Figure 2 (page 56 - 95): Forest plots. 40 Forest plots of SNPs from discovery and conditional phases described in Tables 1 and 2. Nearest gene or previously published proximal gene names included in table.

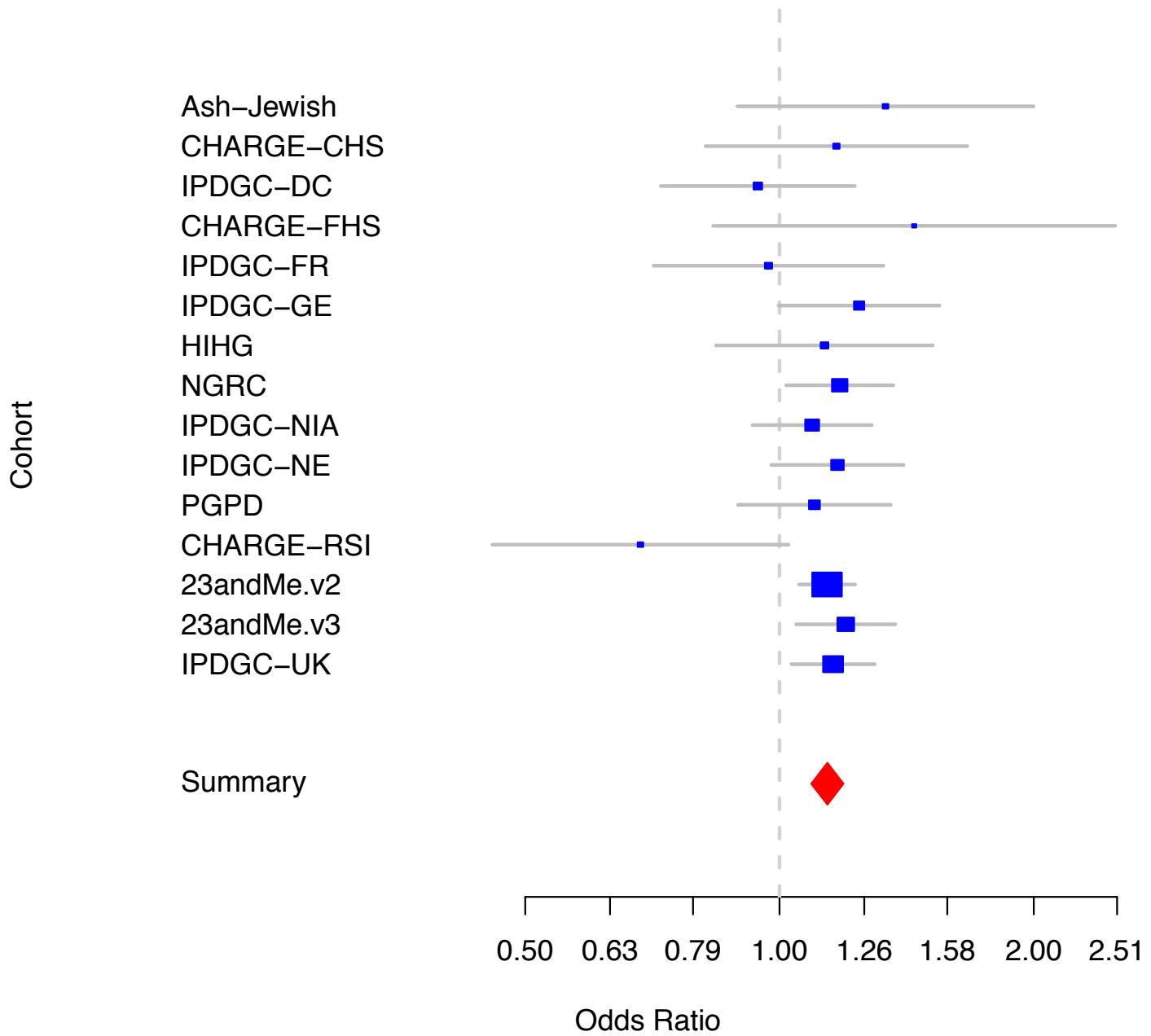
rs35749011 GBA/SYT11 Discovery SNP



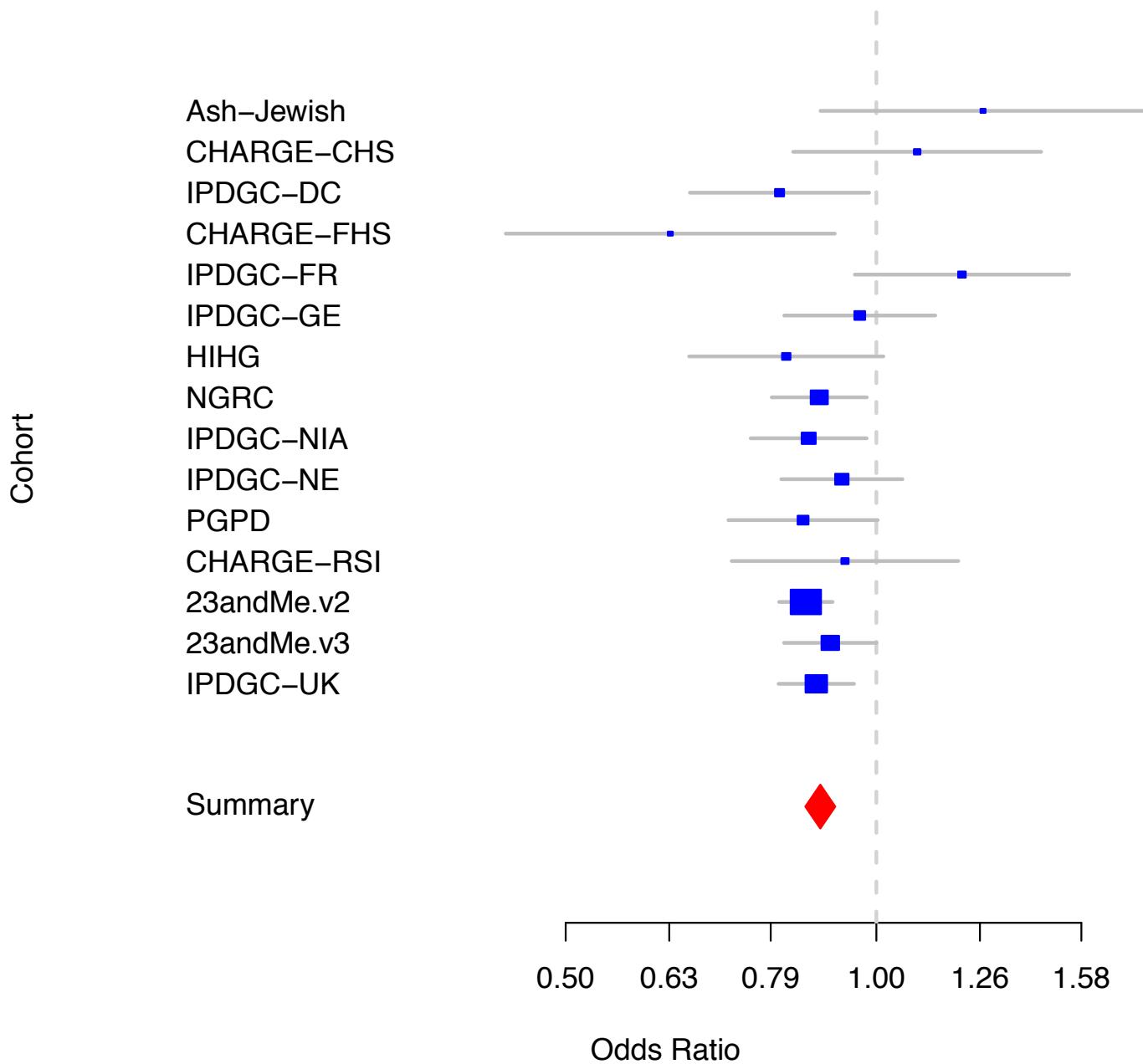
rs823118 RAB7L1/NUCKS1 Discovery SNP



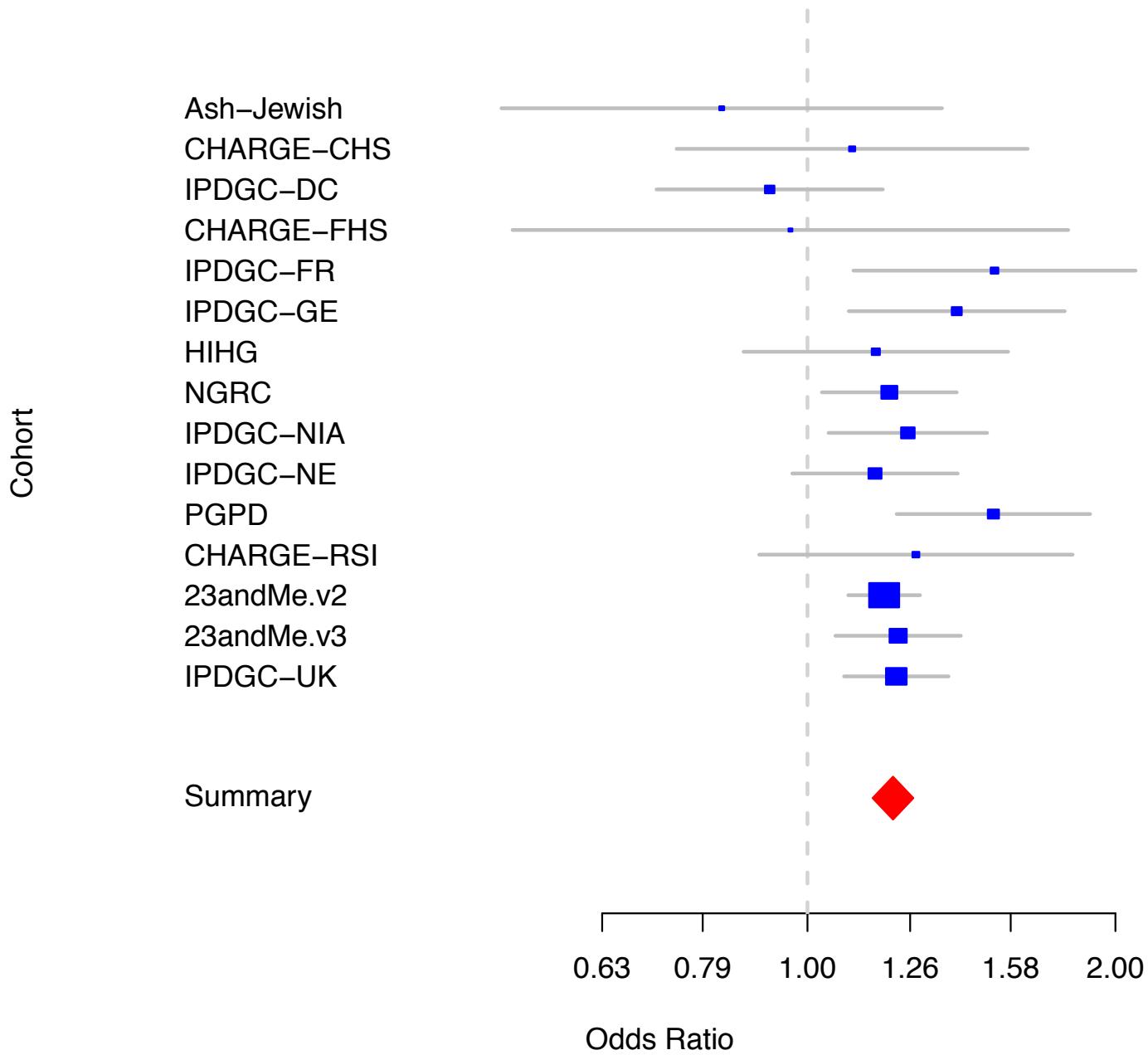
rs10797576 SIPA1L2 Discovery SNP



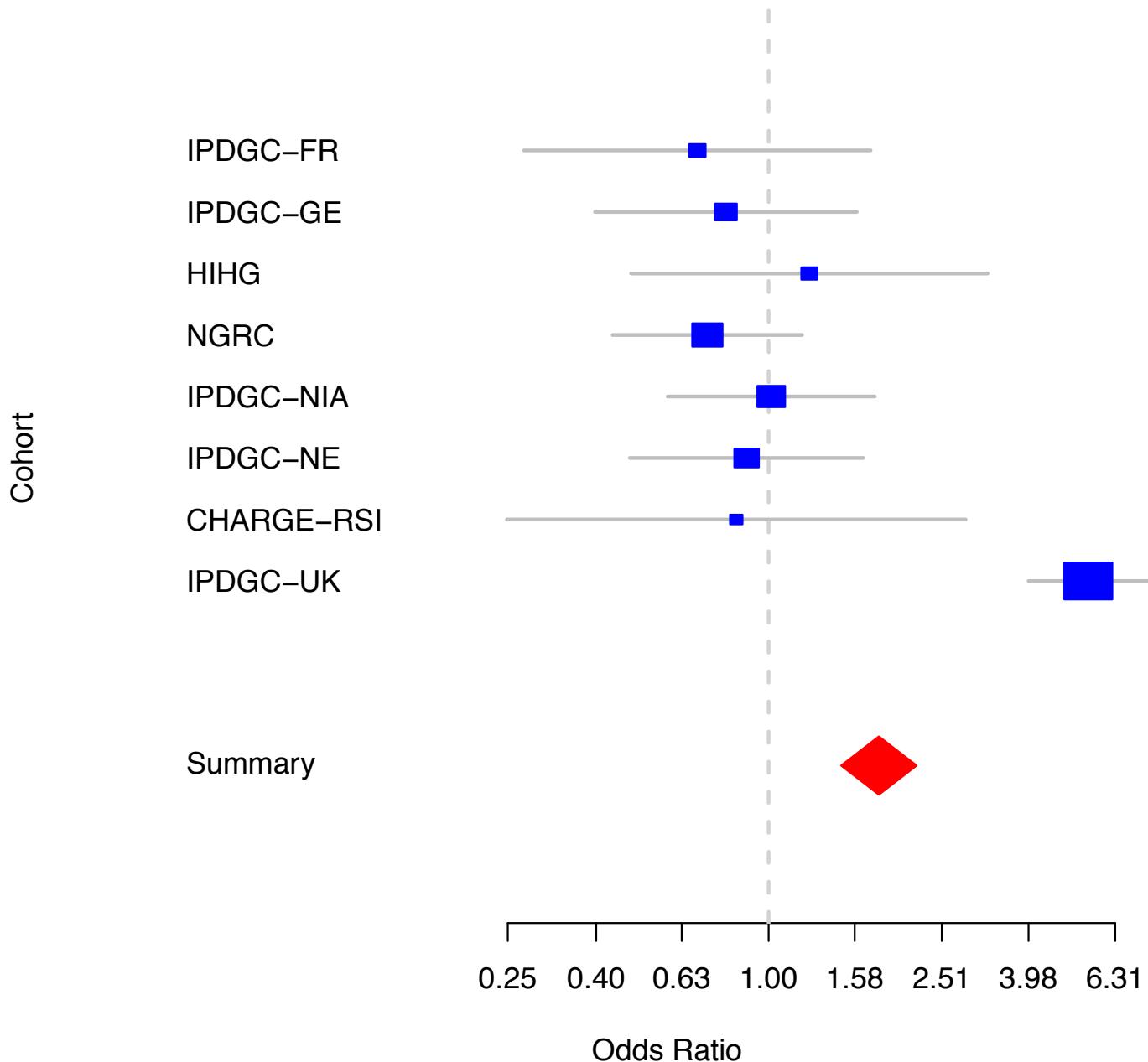
rs6430538 ACMSD/TMEM163 Discovery SNP



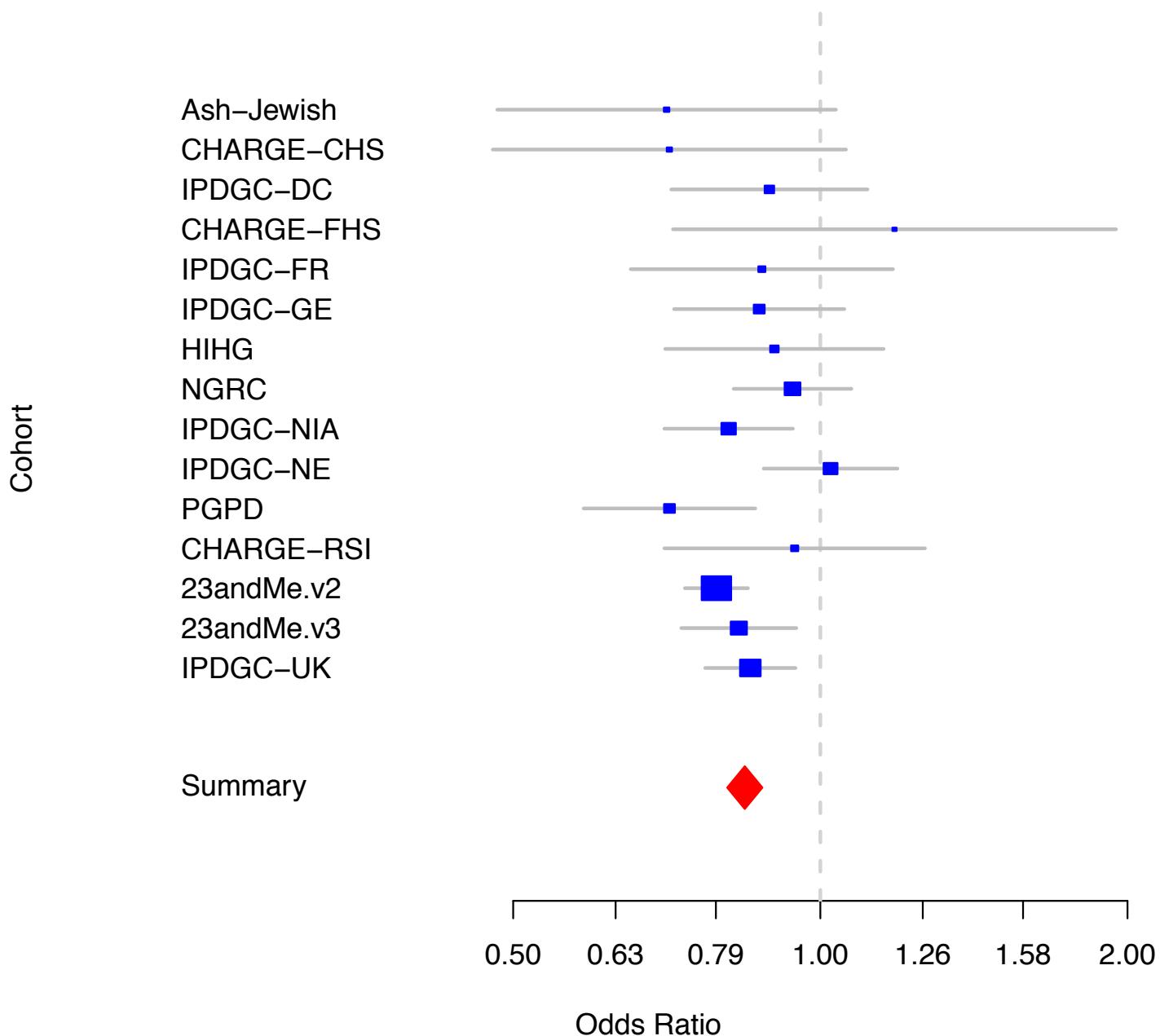
rs1474055 STK39 Discovery SNP



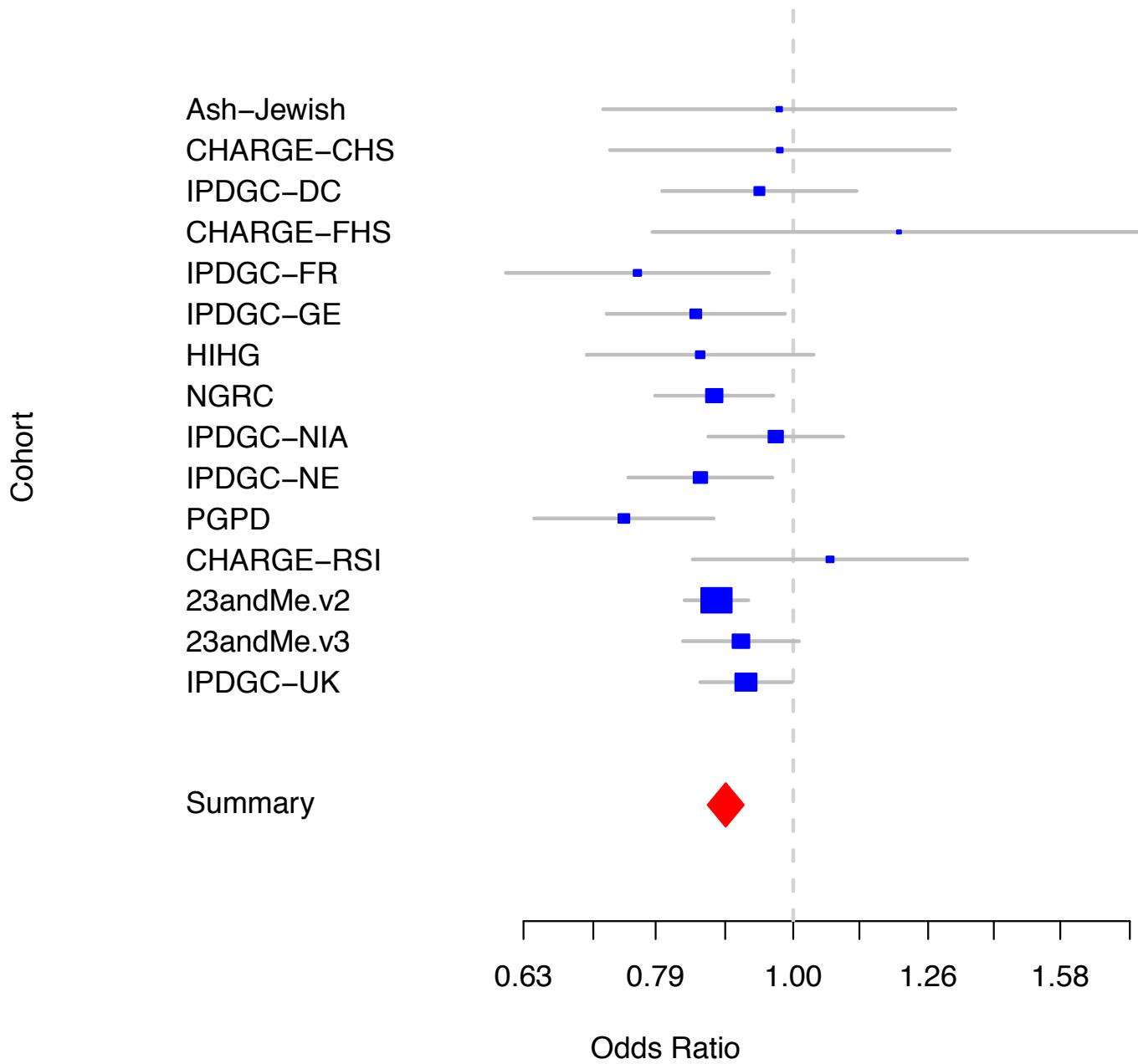
rs115185635 KRT8P25/APOOP2 Discovery SNP



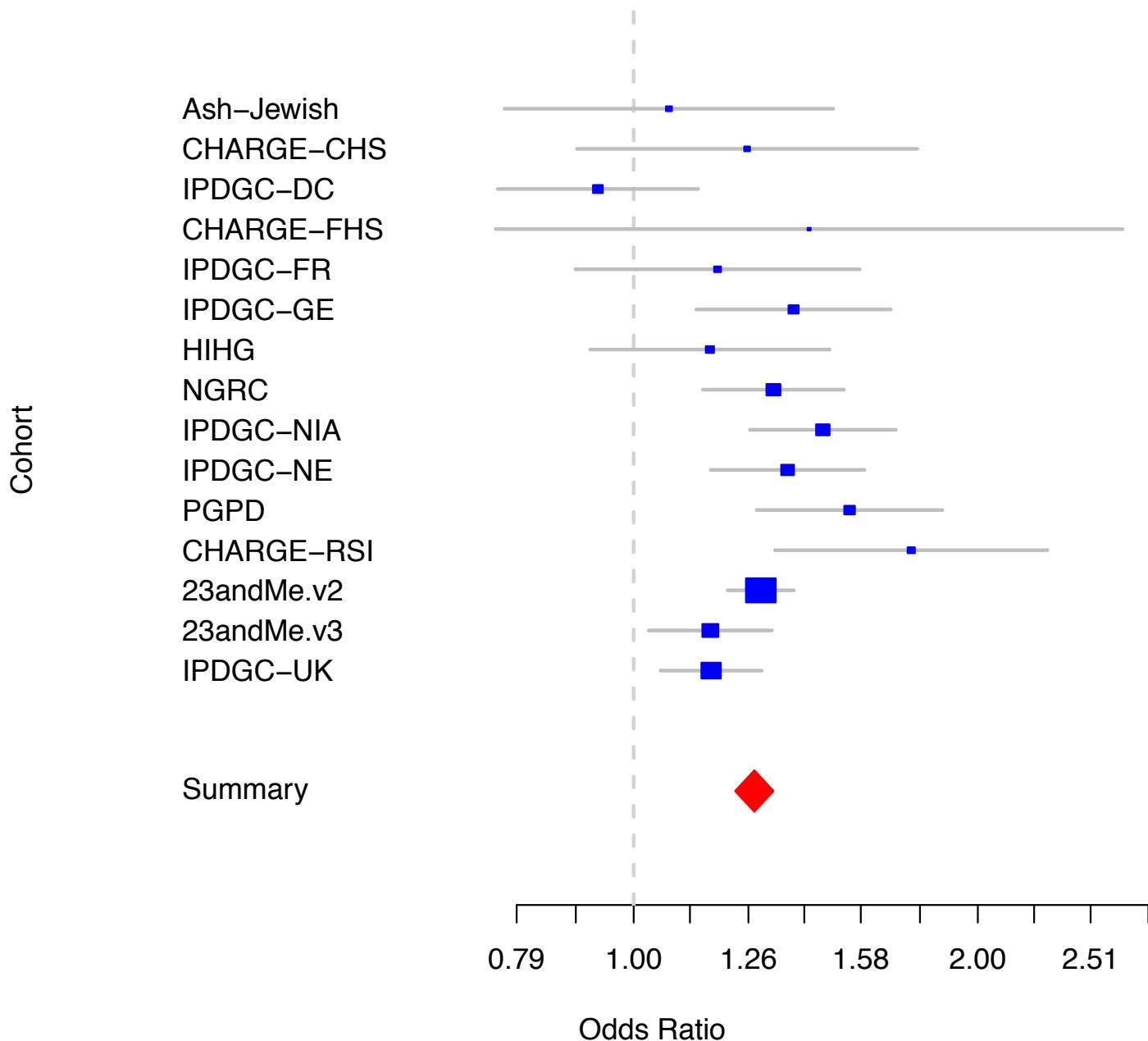
rs12637471 MCCC1 Discovery SNP



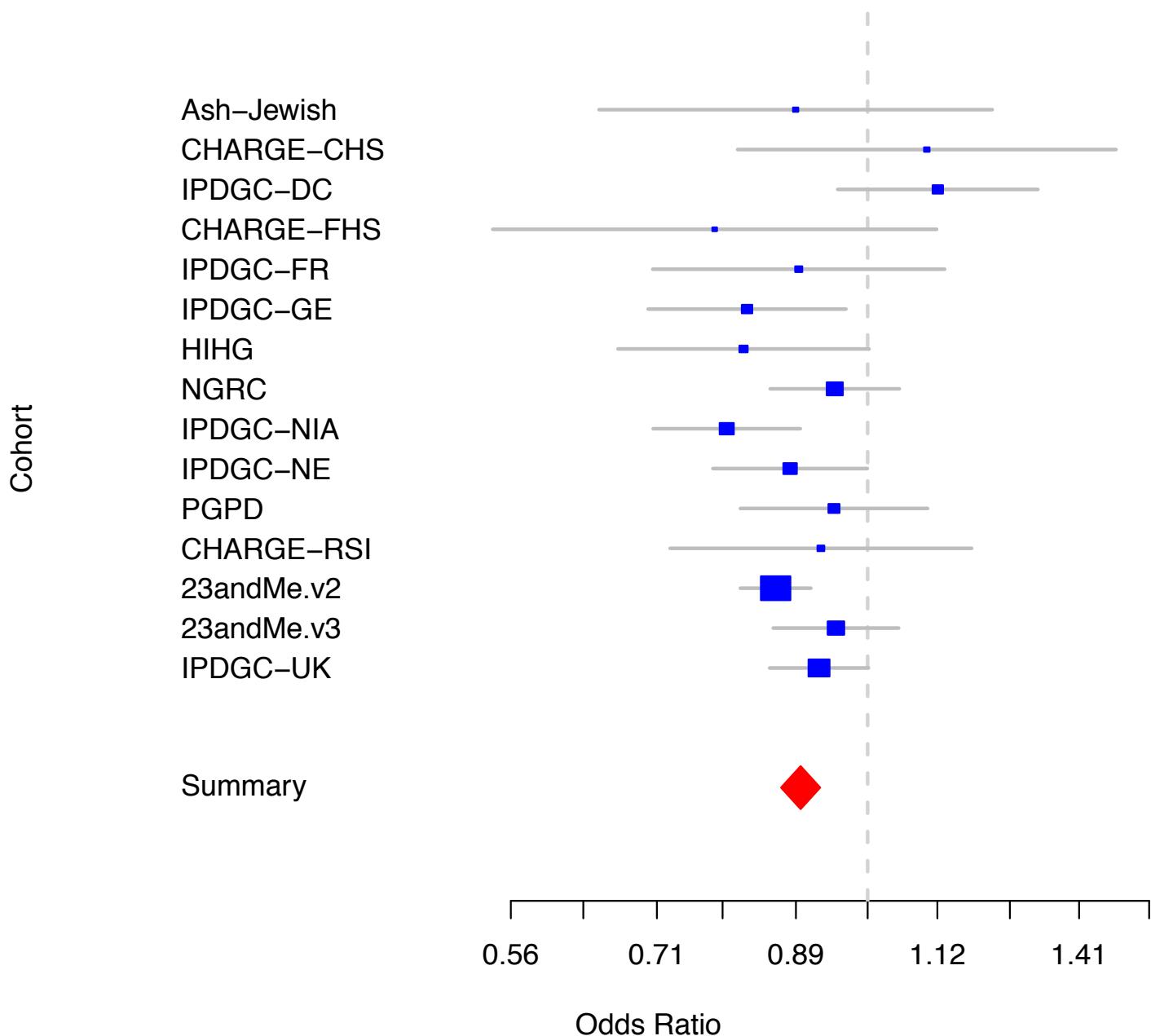
rs11724635 BST1 Discovery SNP



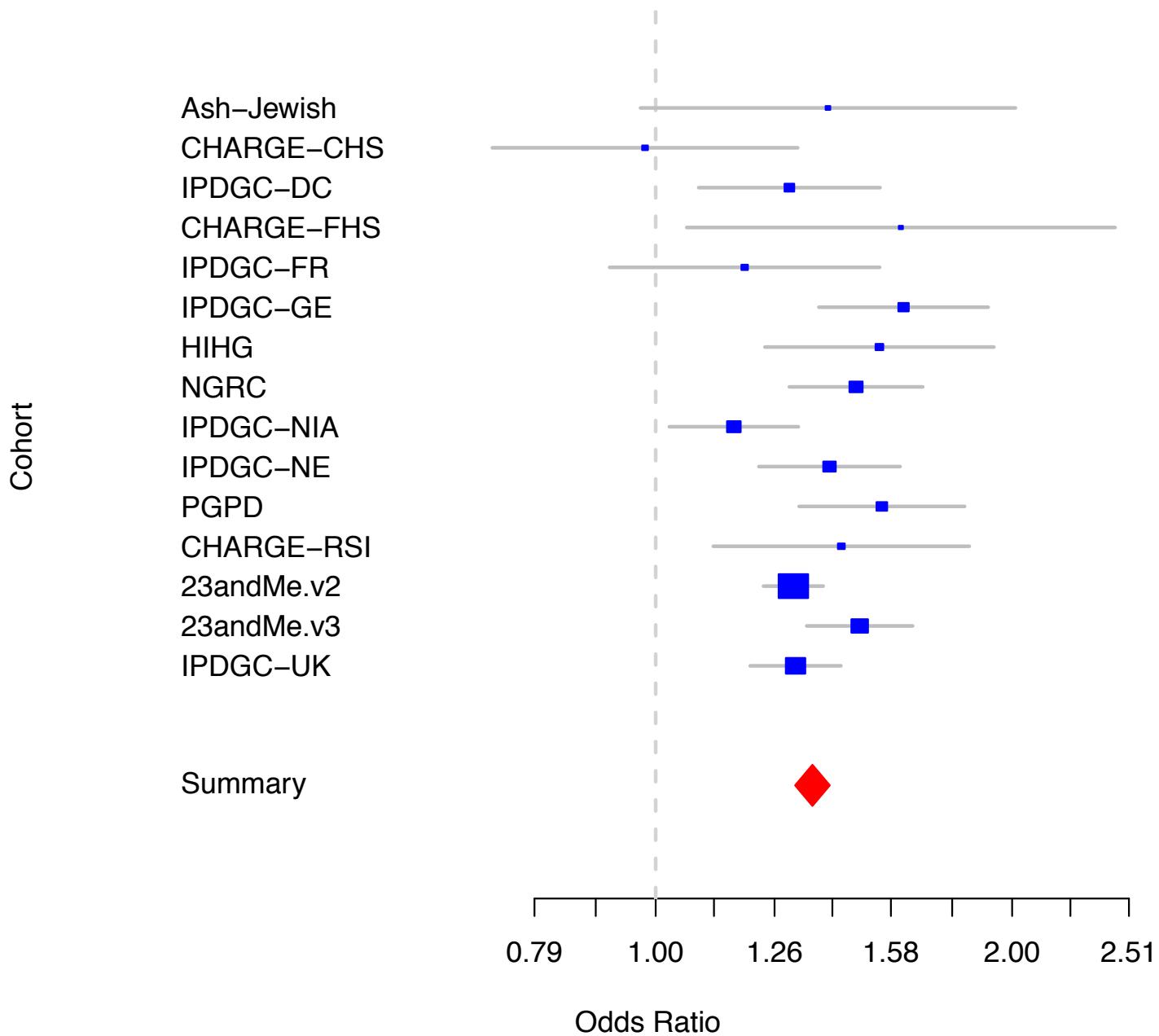
rs34311866 TMEM175/GAK/DGKQ Discovery SNP



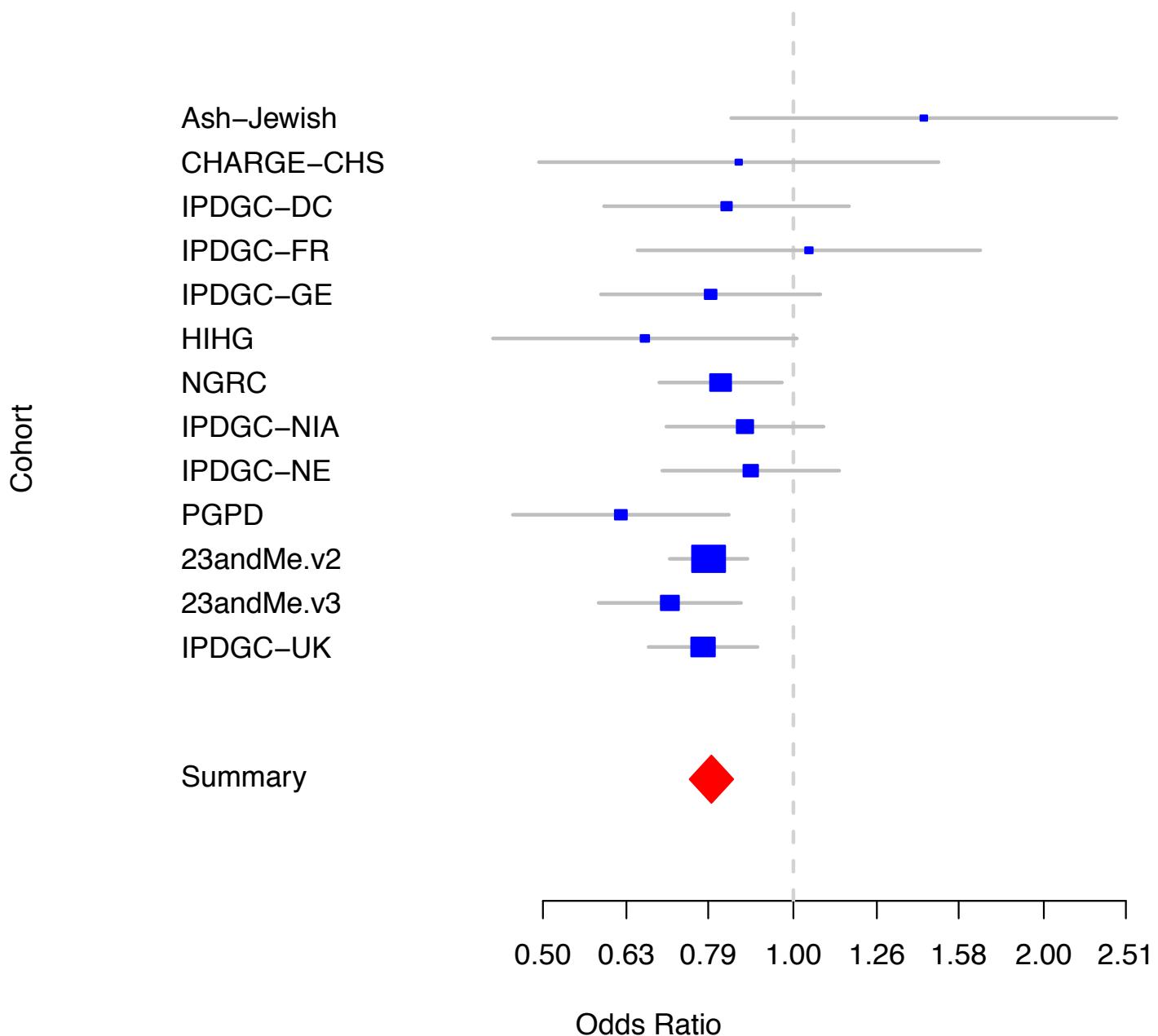
rs6812193 FAM47E/SCARB2 Discovery SNP



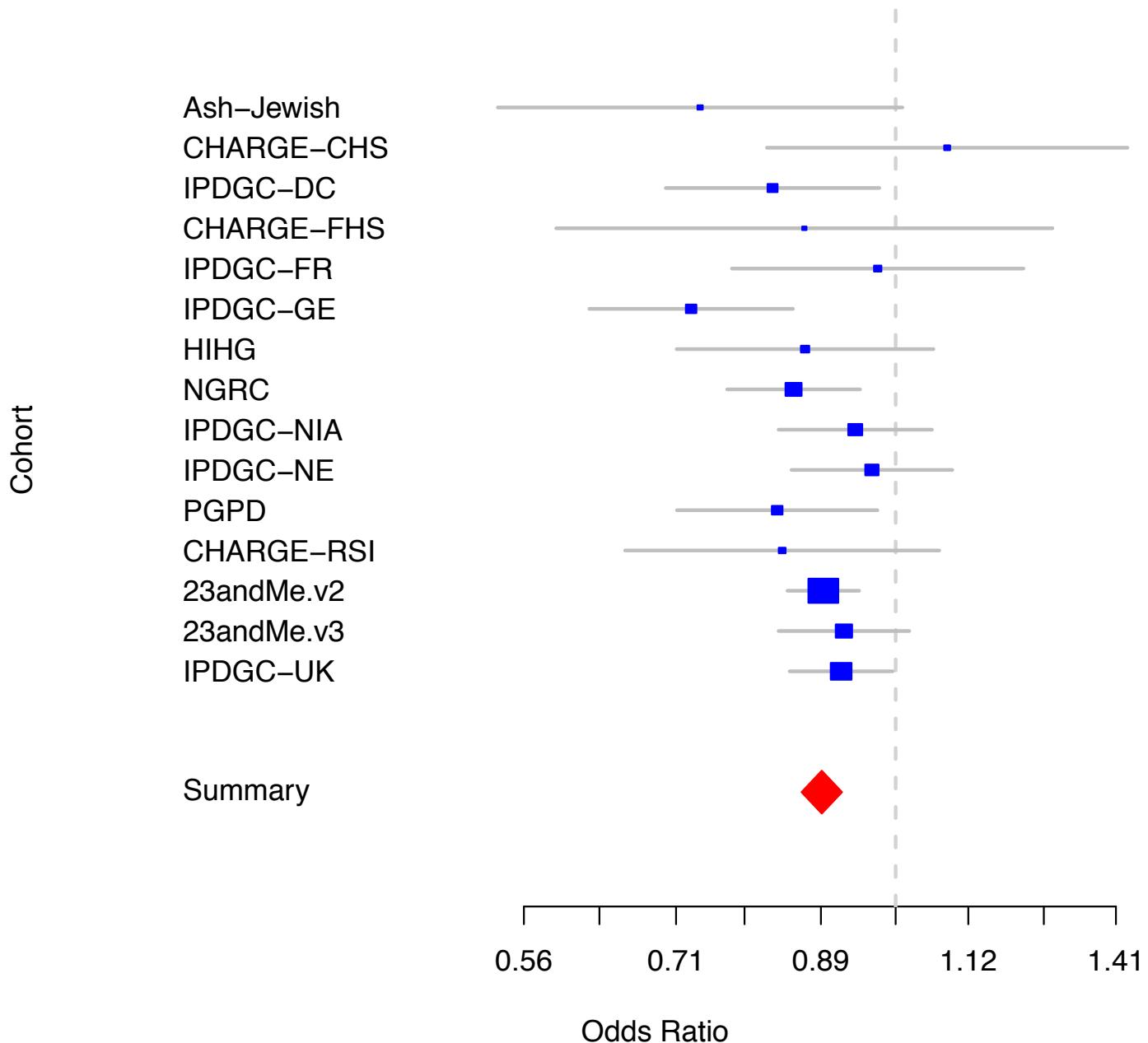
rs356182 SNCA Discovery SNP



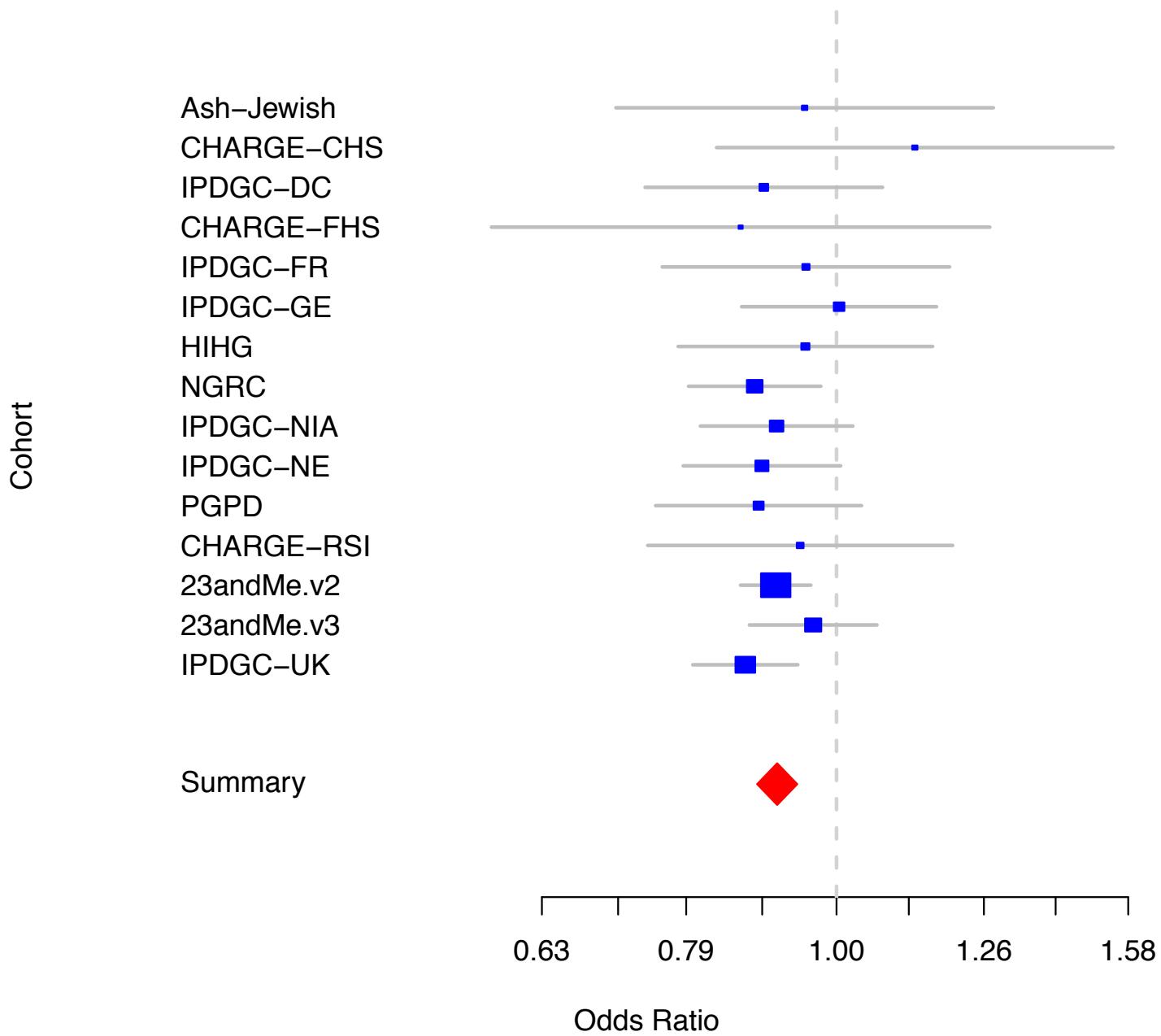
rs9275326 HLA-DQB1 Discovery SNP



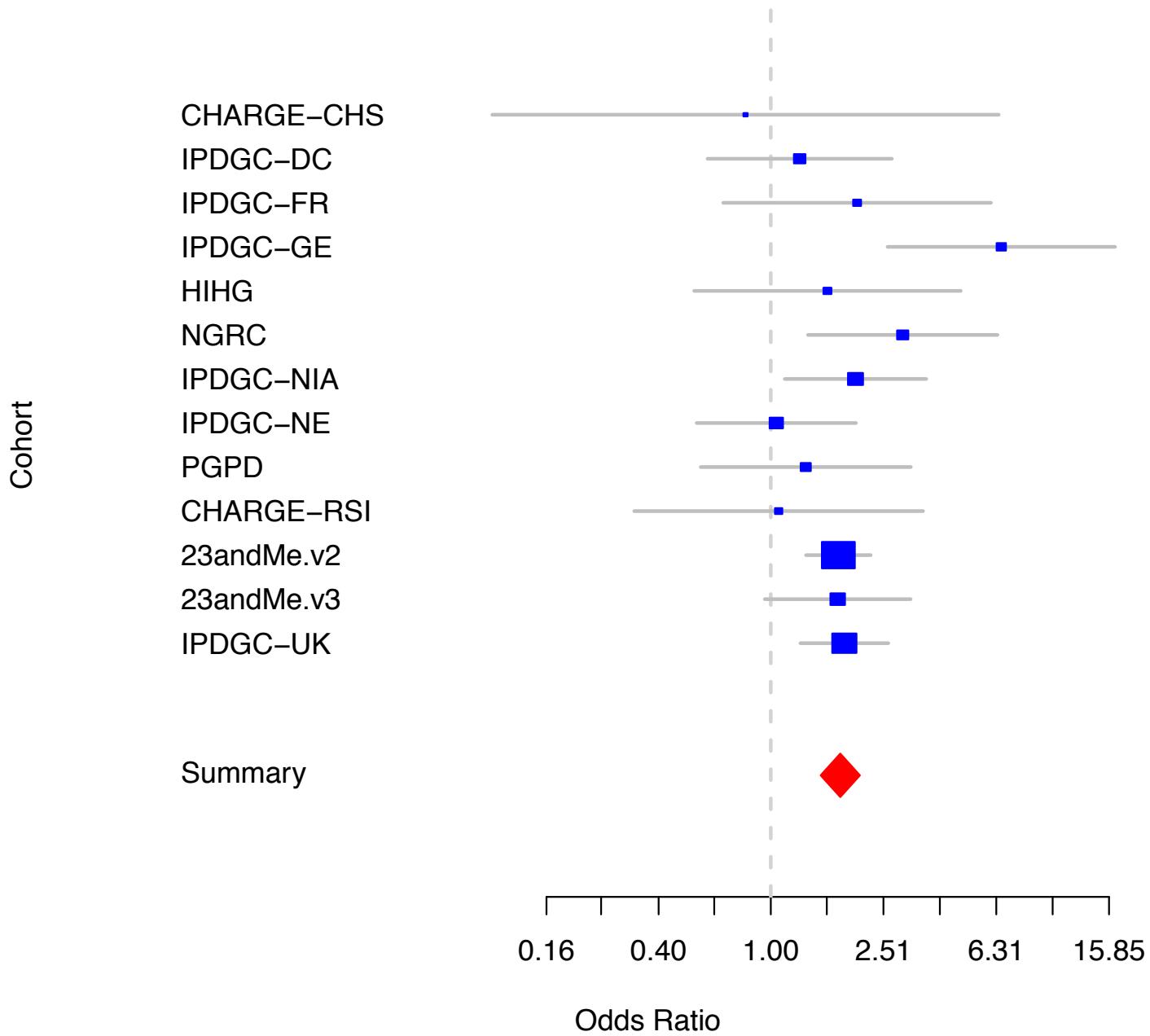
rs199347 GPNMB Discovery SNP



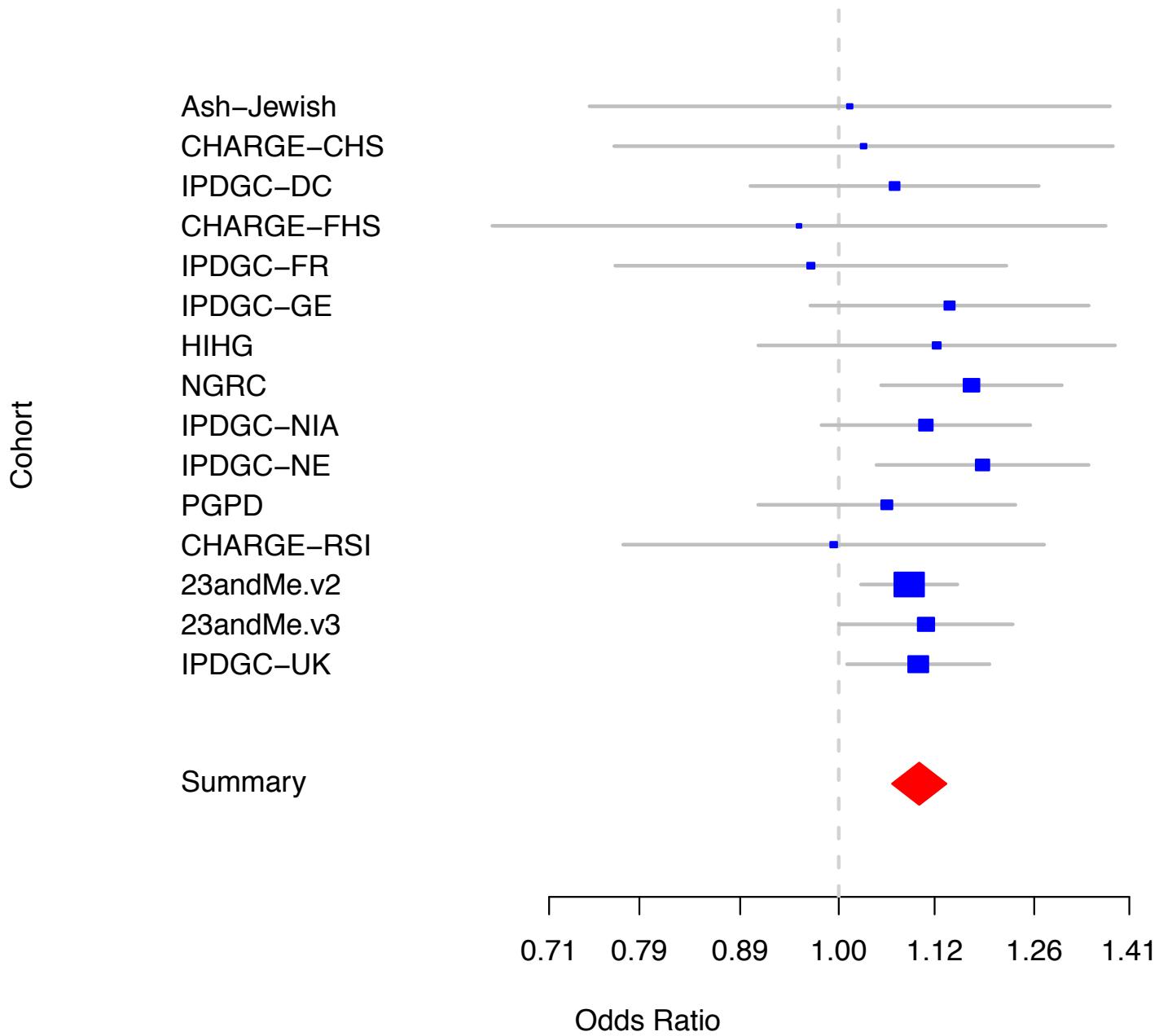
rs3793947 DLG2 Discovery SNP



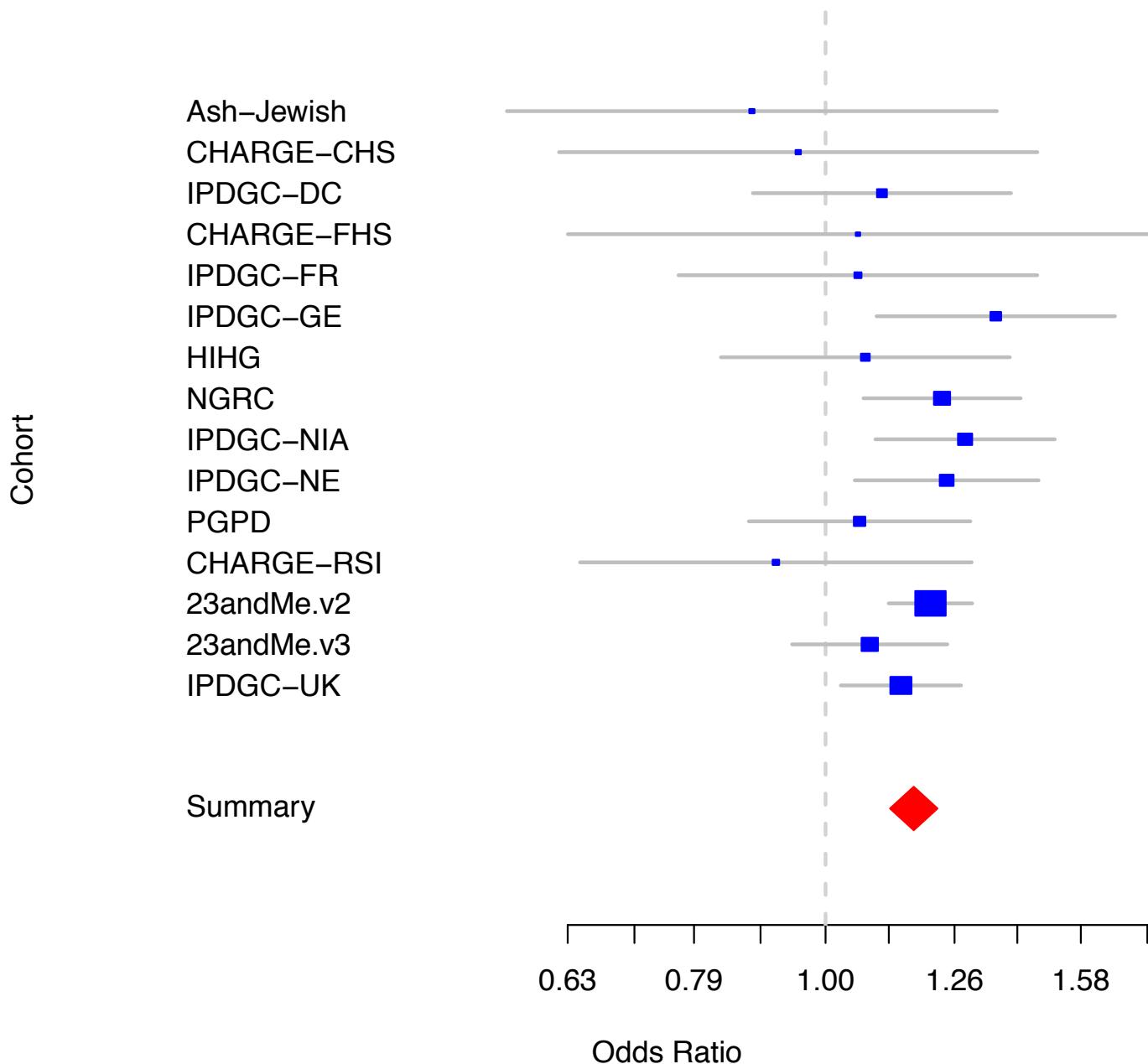
rs117896735 INPP5F Discovery SNP



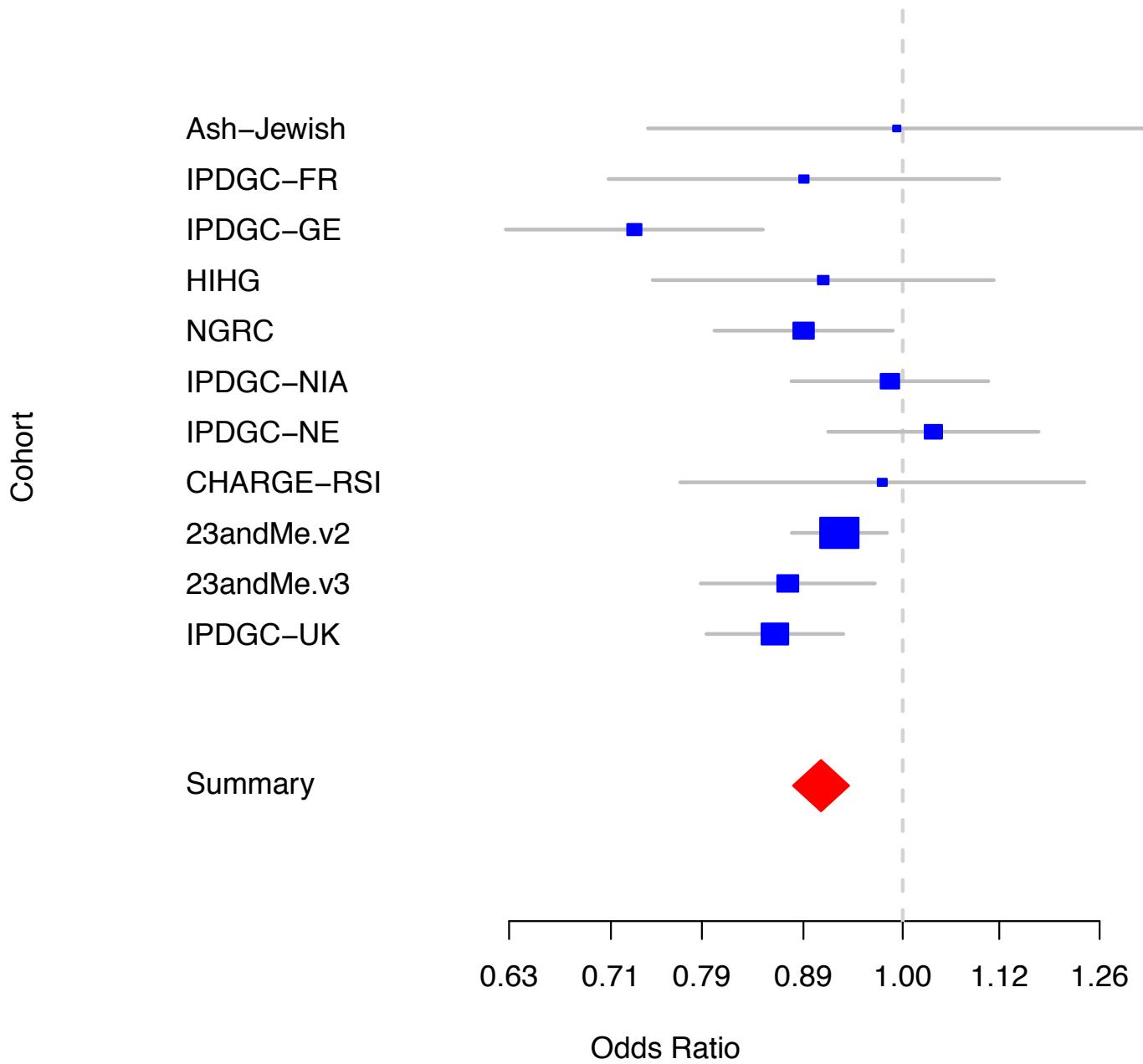
rs329648 LOC283174 Discovery SNP



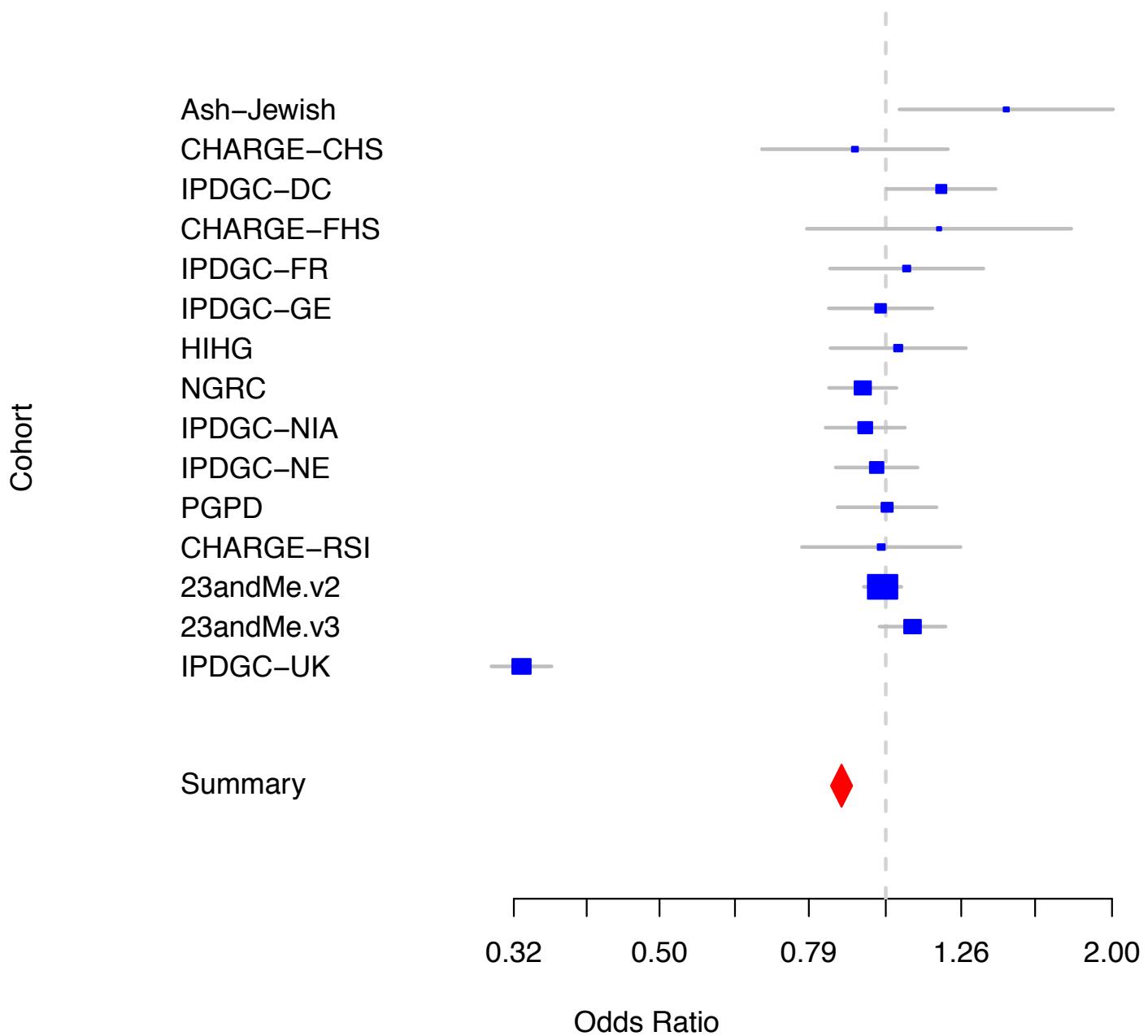
rs76904798 LRRK2 Discovery SNP



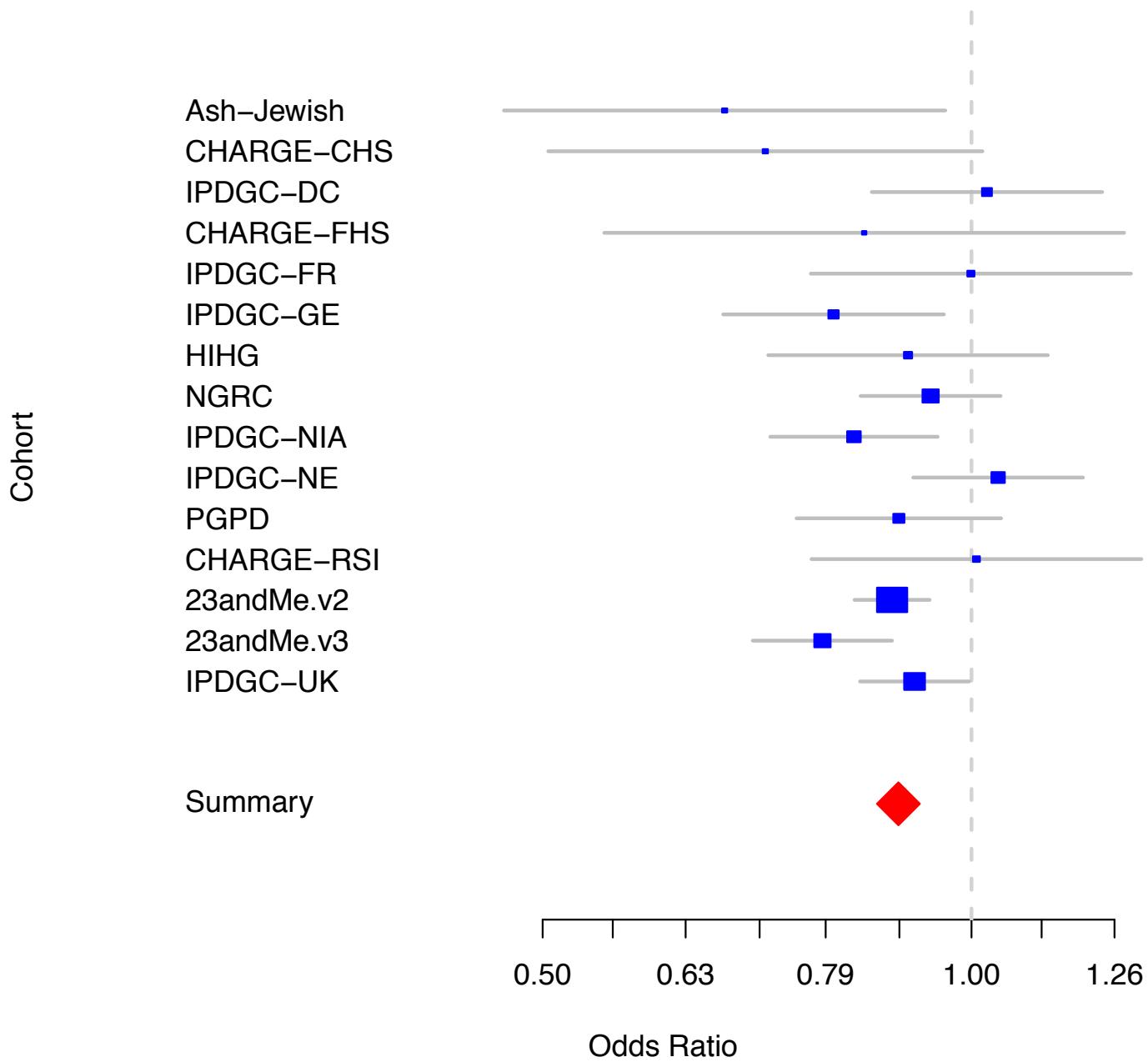
rs11060180 CCDC62 Discovery SNP



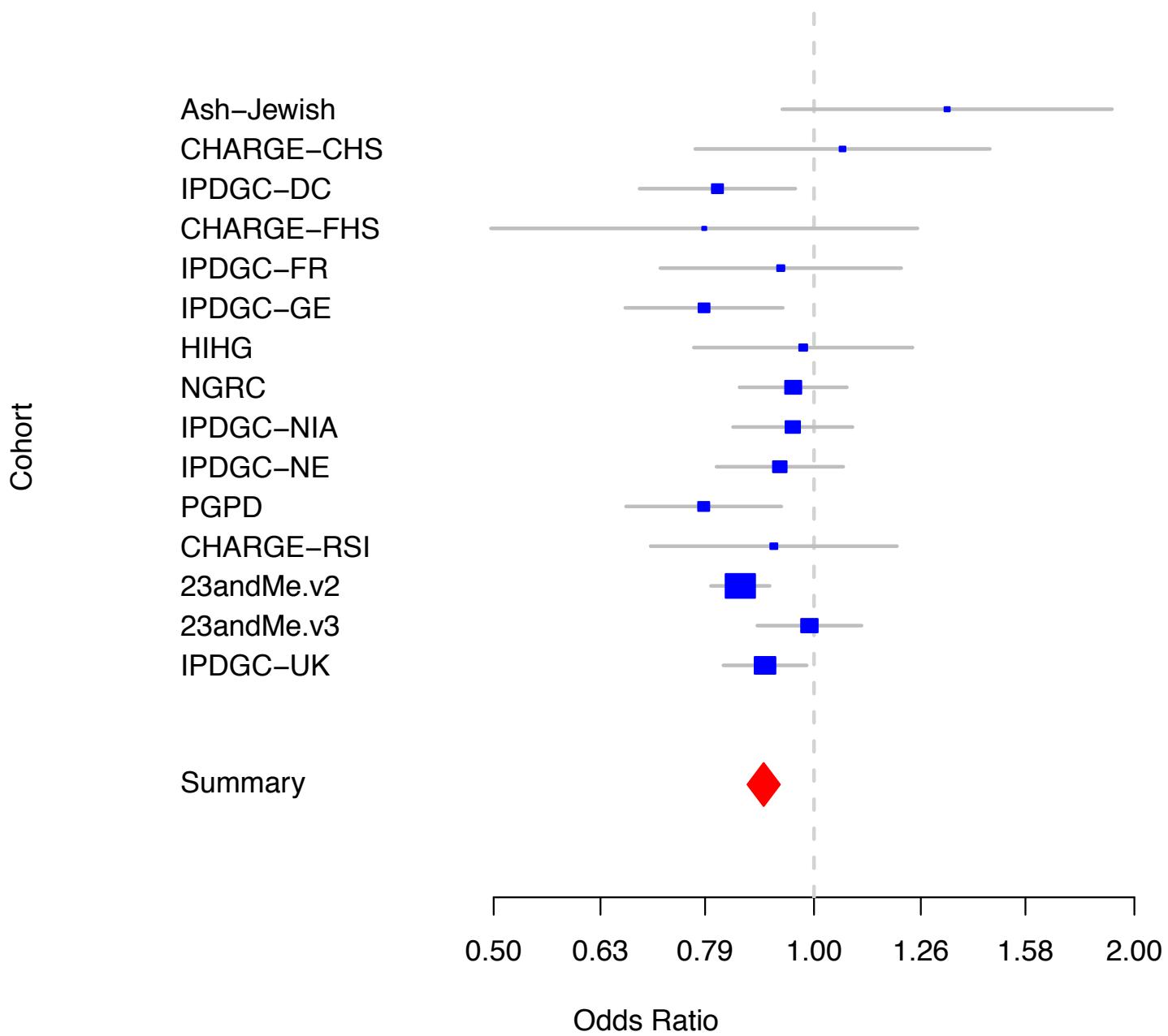
rs1555399 TMEM229B Discovery SNP



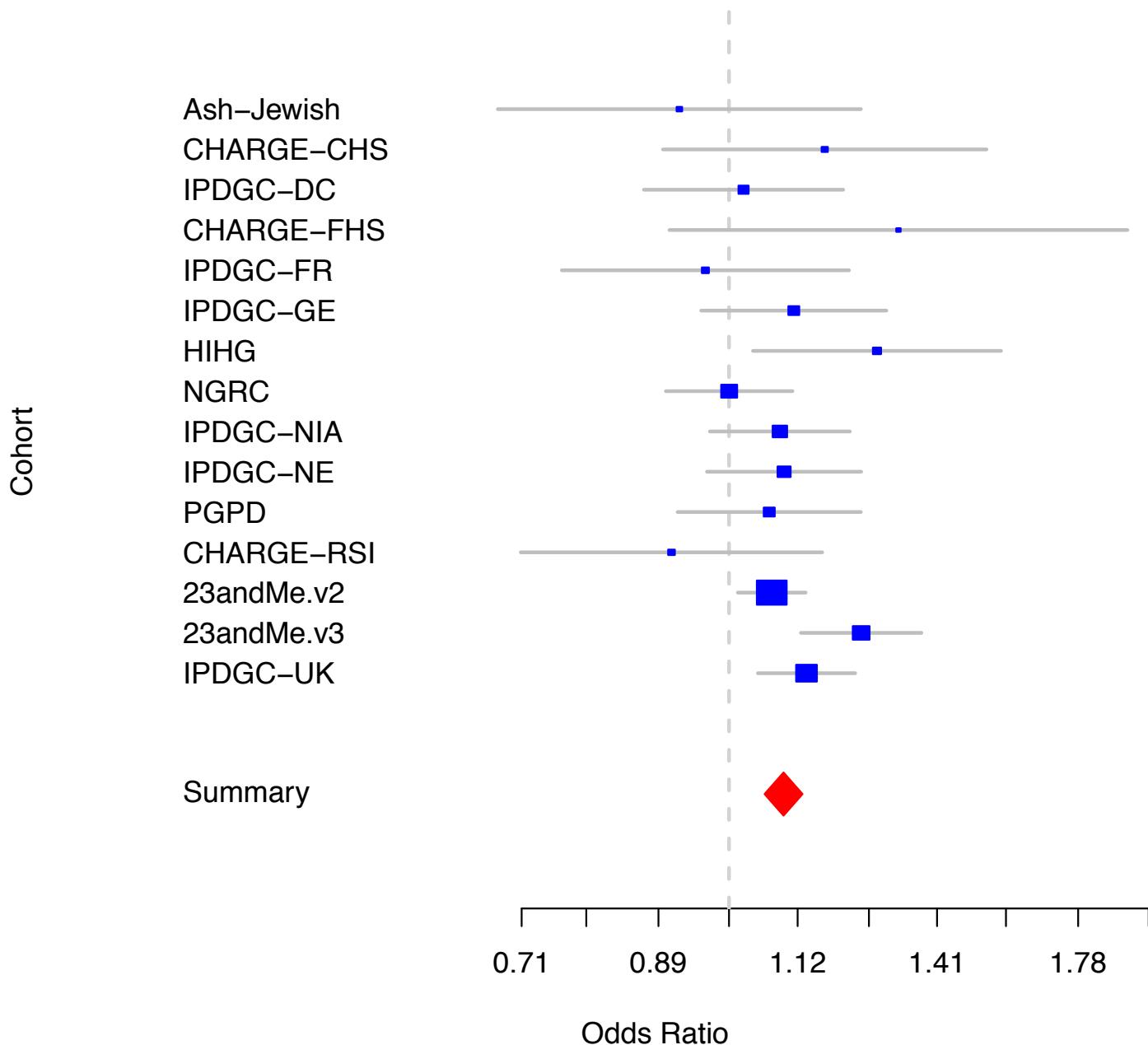
rs11158026 GCH1 Discovery SNP



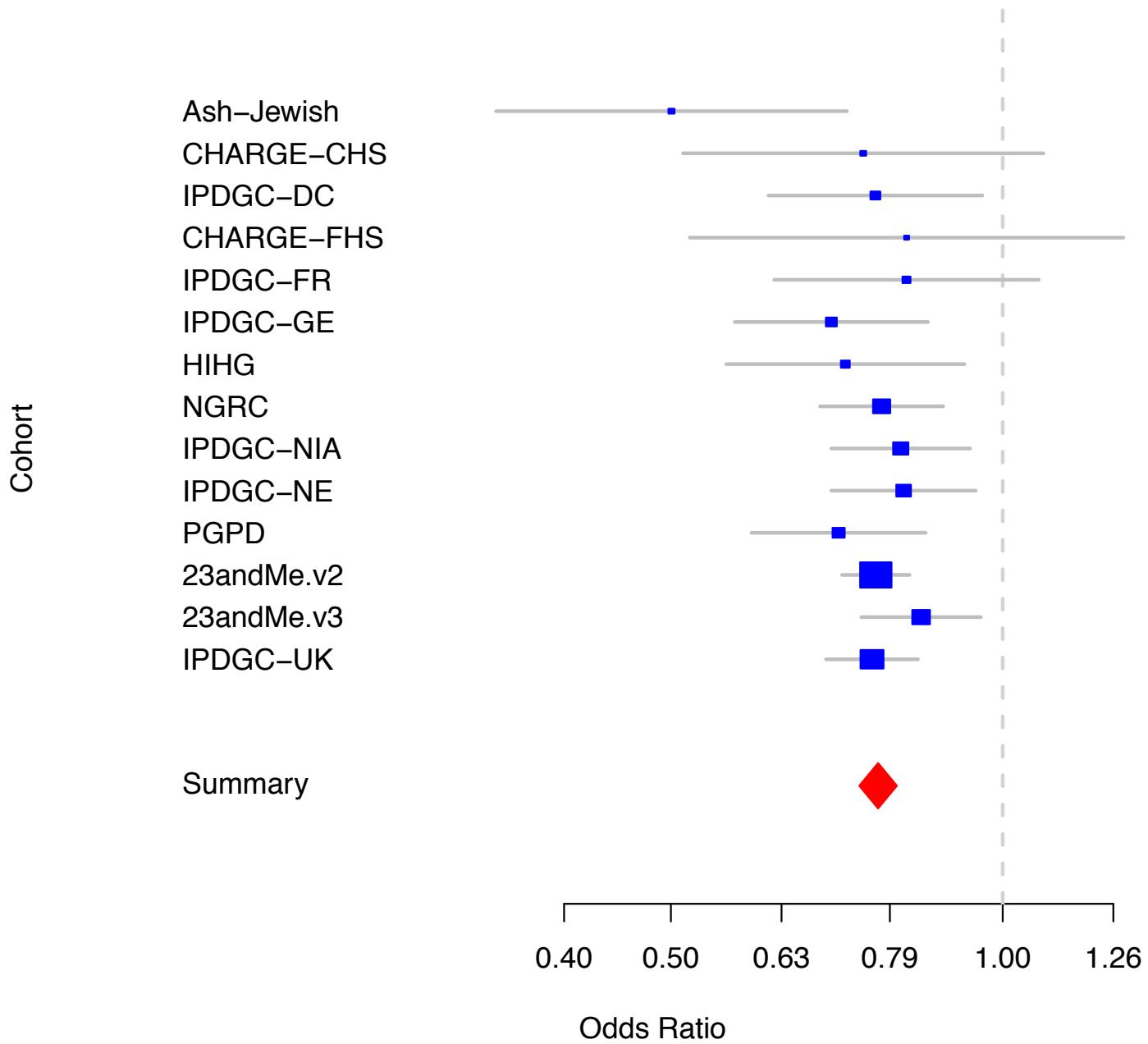
rs2414739 VPS13C Discovery SNP



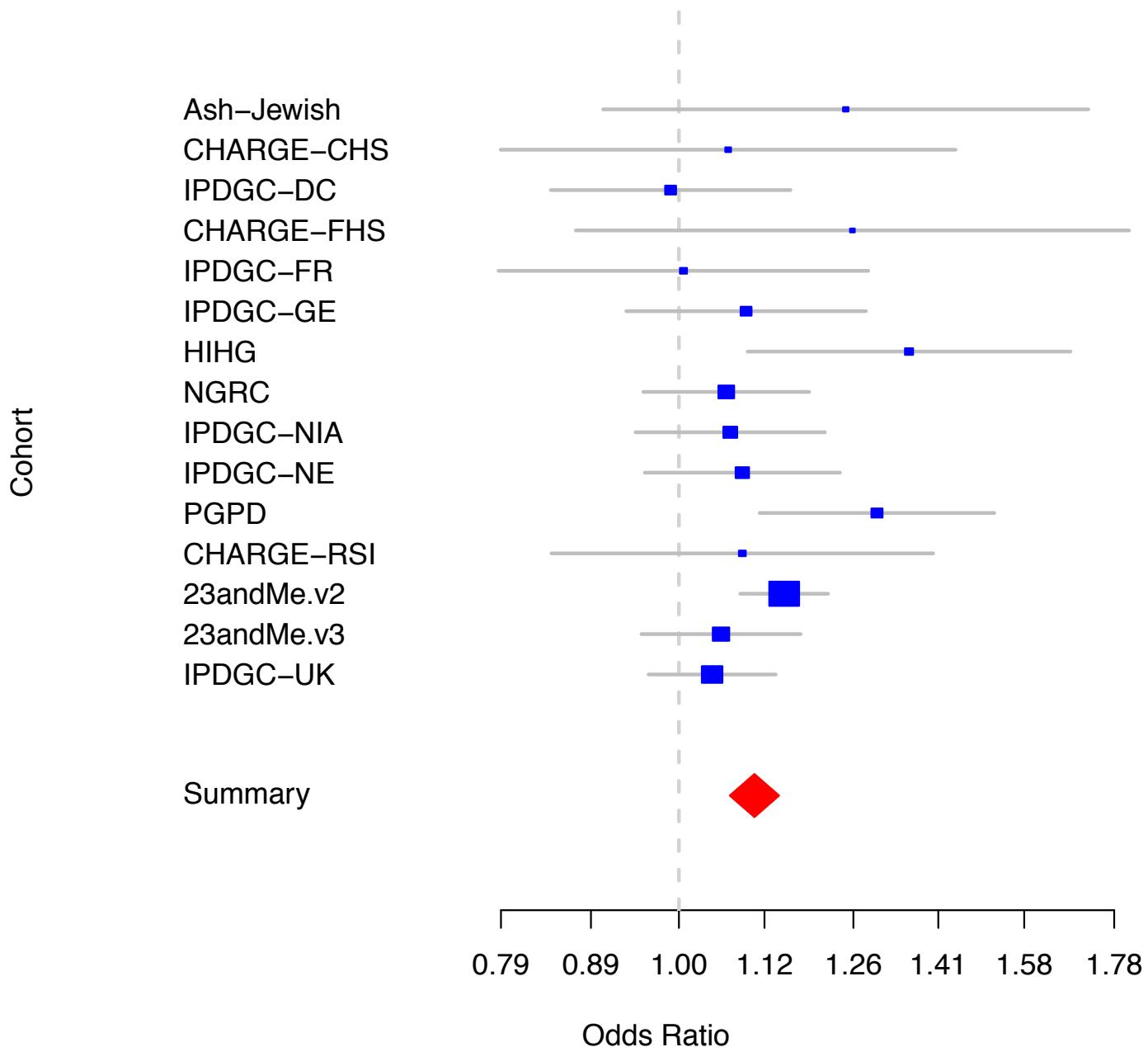
rs14235 BCKDK/ STX1B Discovery SNP



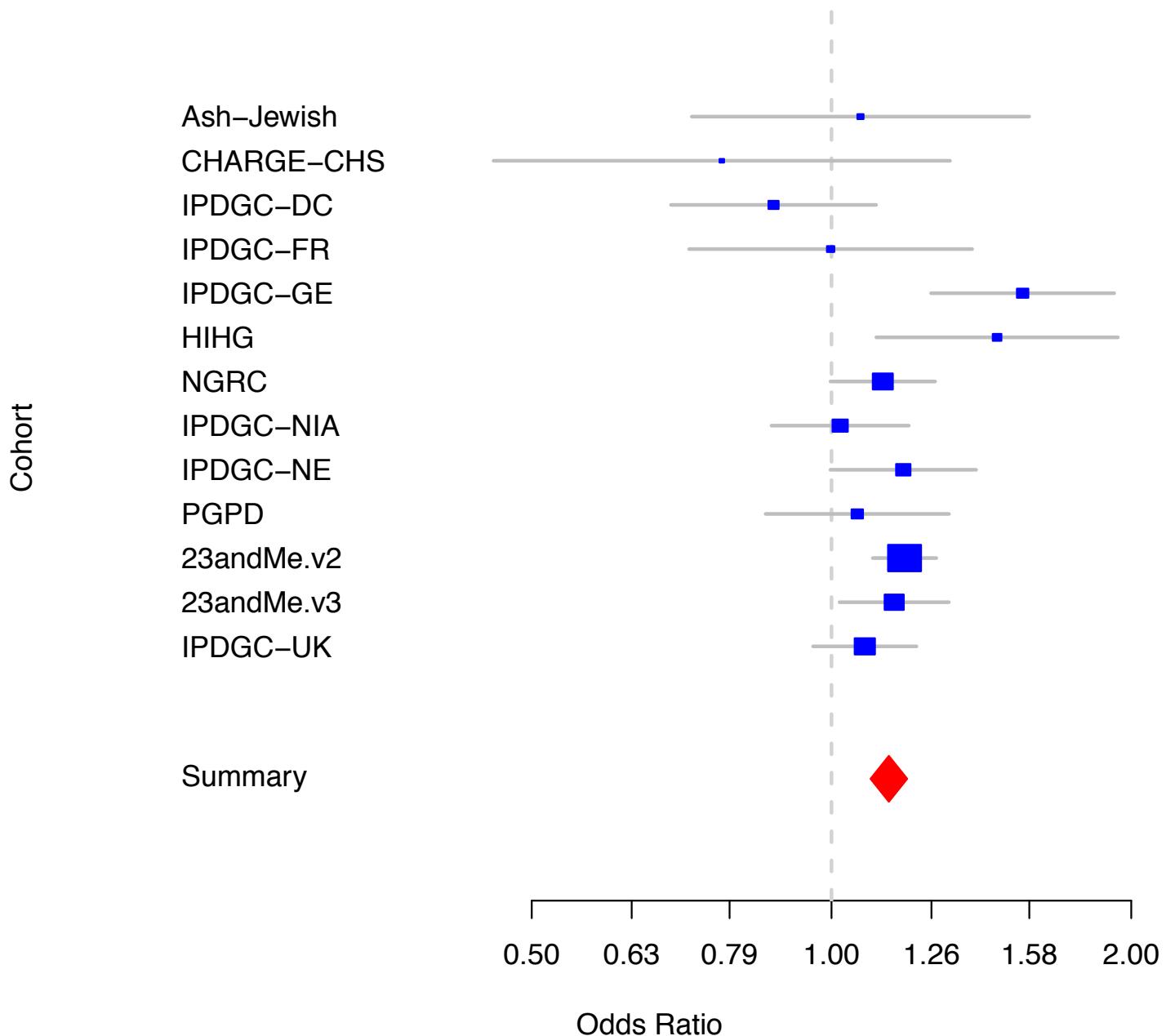
rs17649553 MAPT Discovery SNP



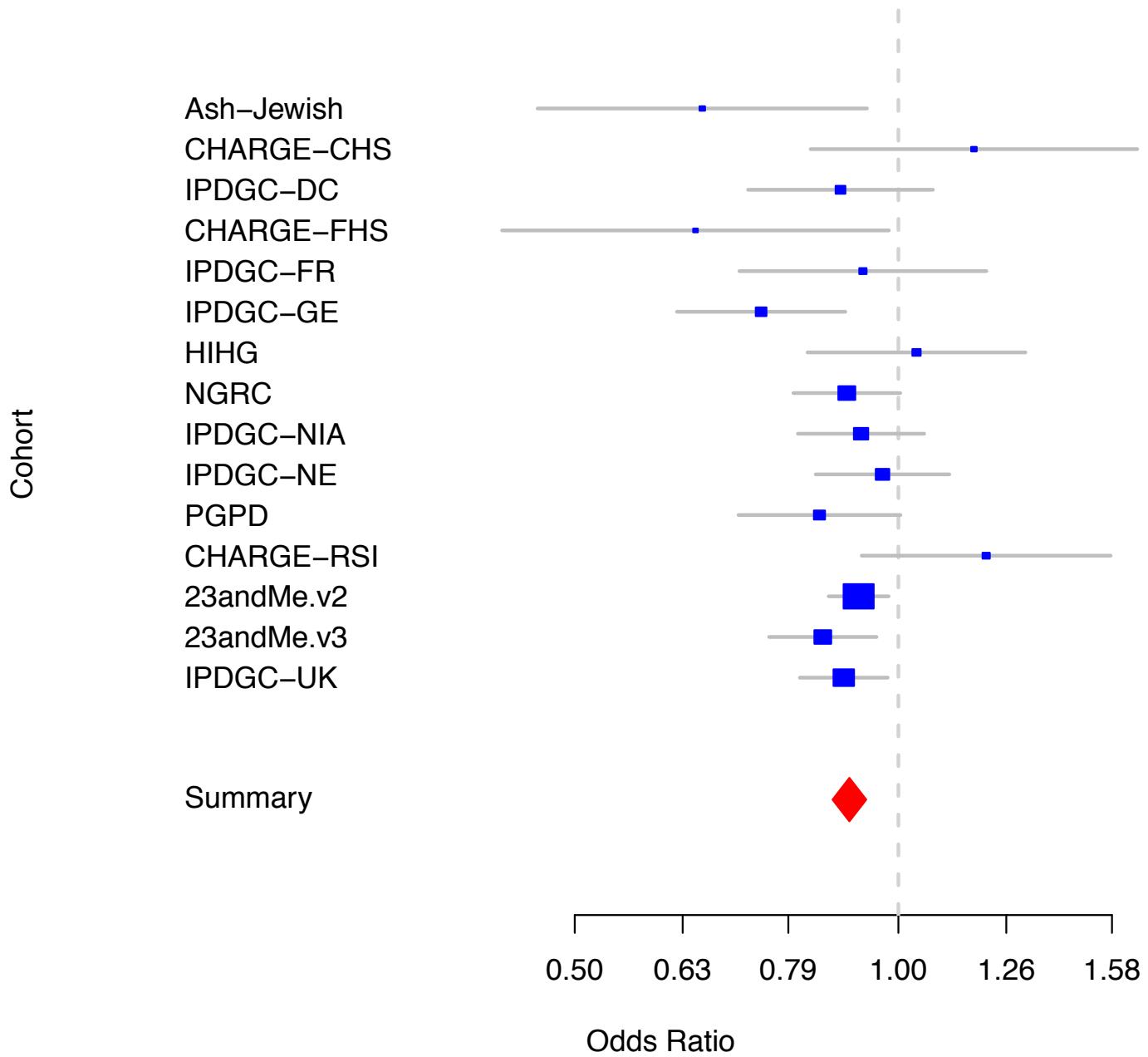
rs12456492 RIT2 Discovery SNP



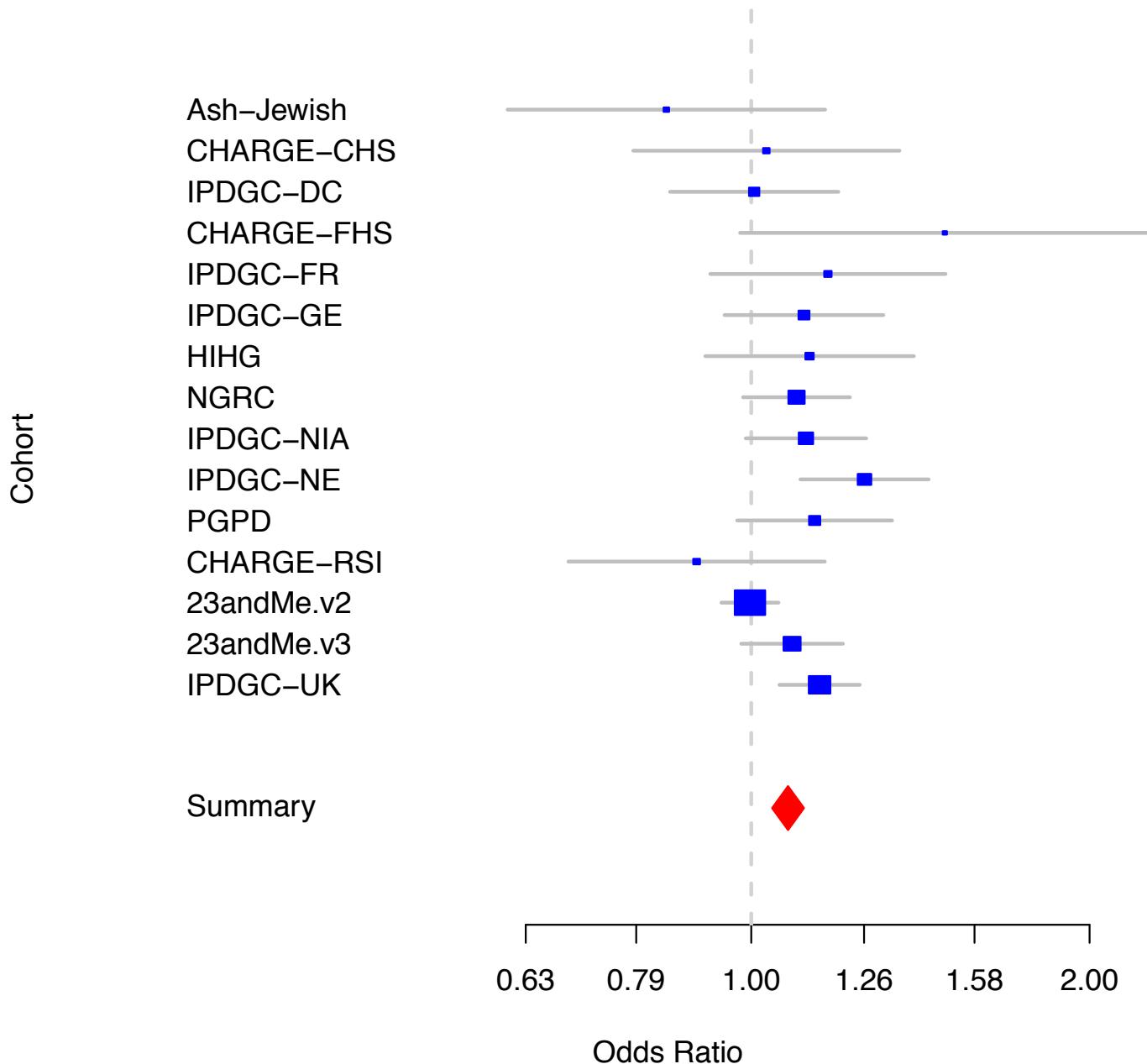
rs62120679 SPPL2B Discovery SNP



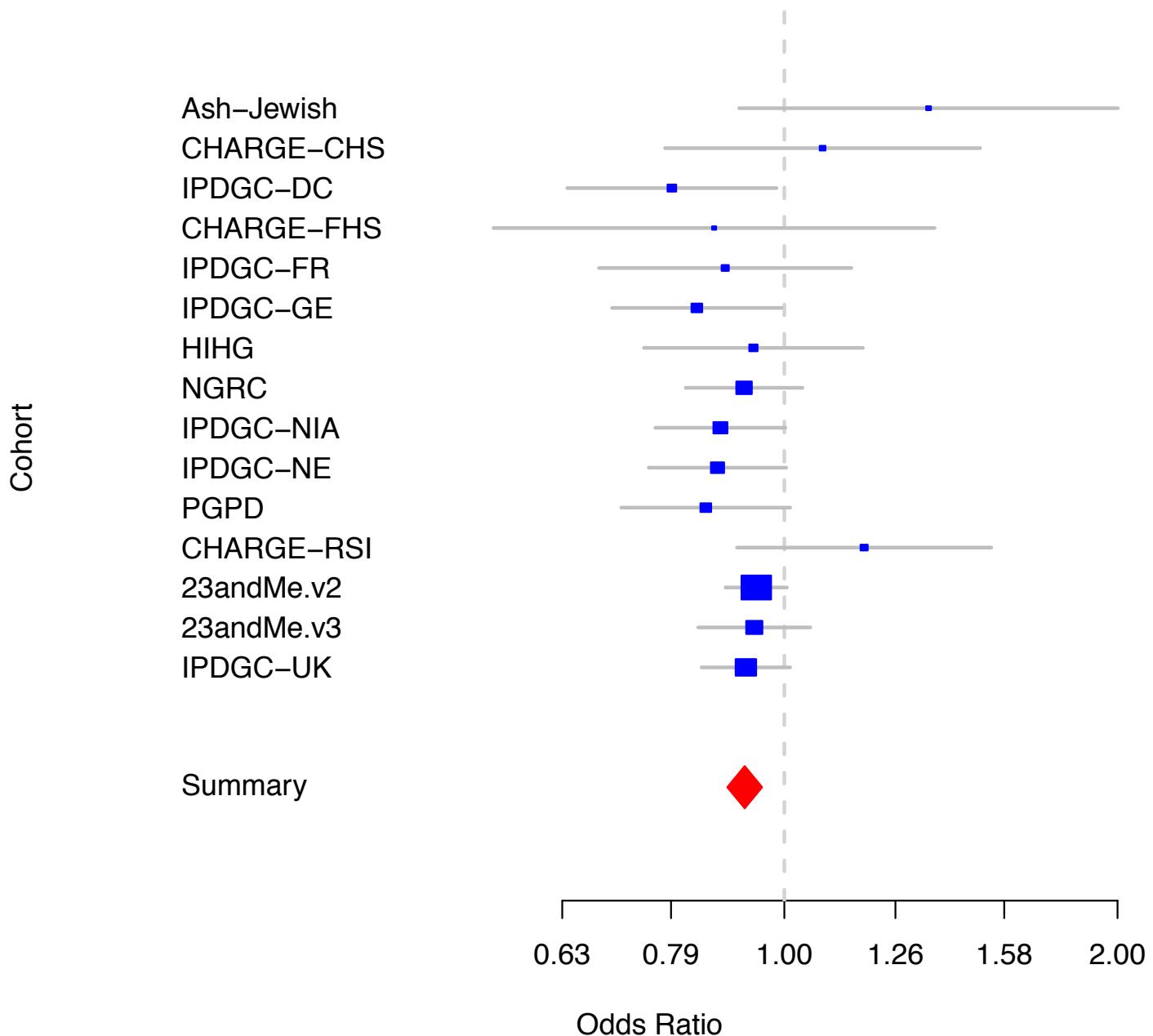
rs8118008 DDRGK1 Discovery SNP



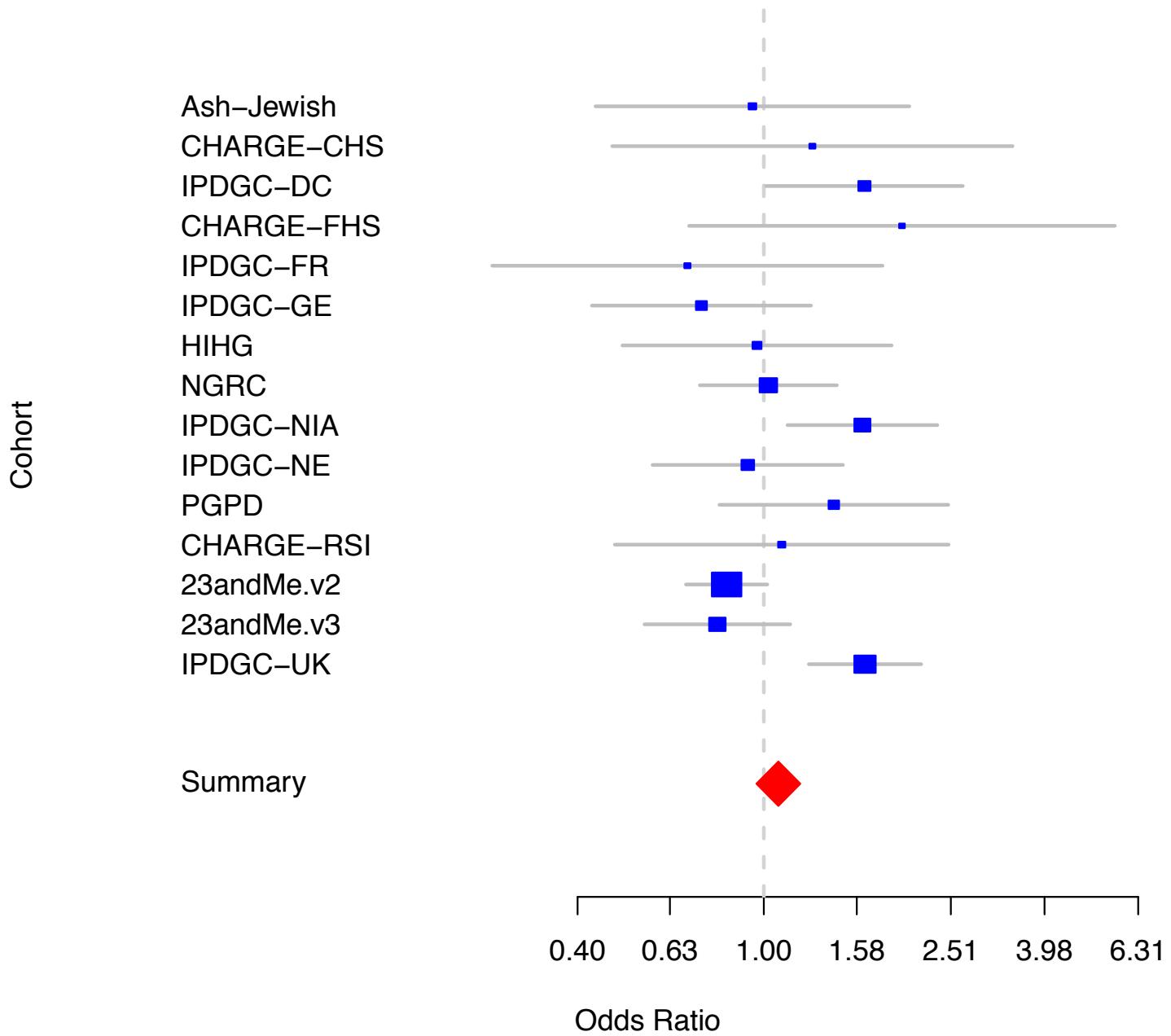
rs34016896 NMD3 Known SNP



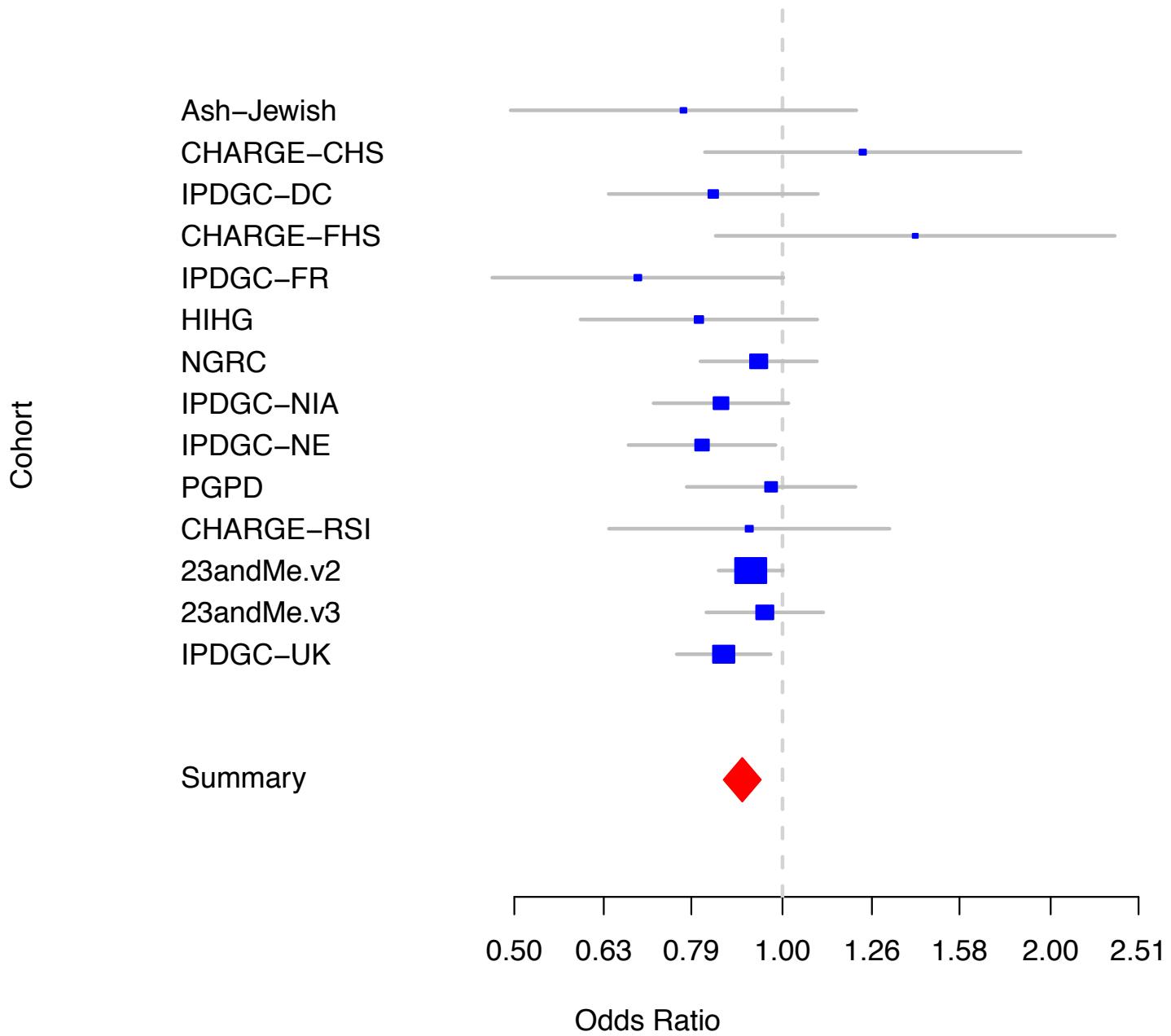
rs591323 FGF20 Known SNP



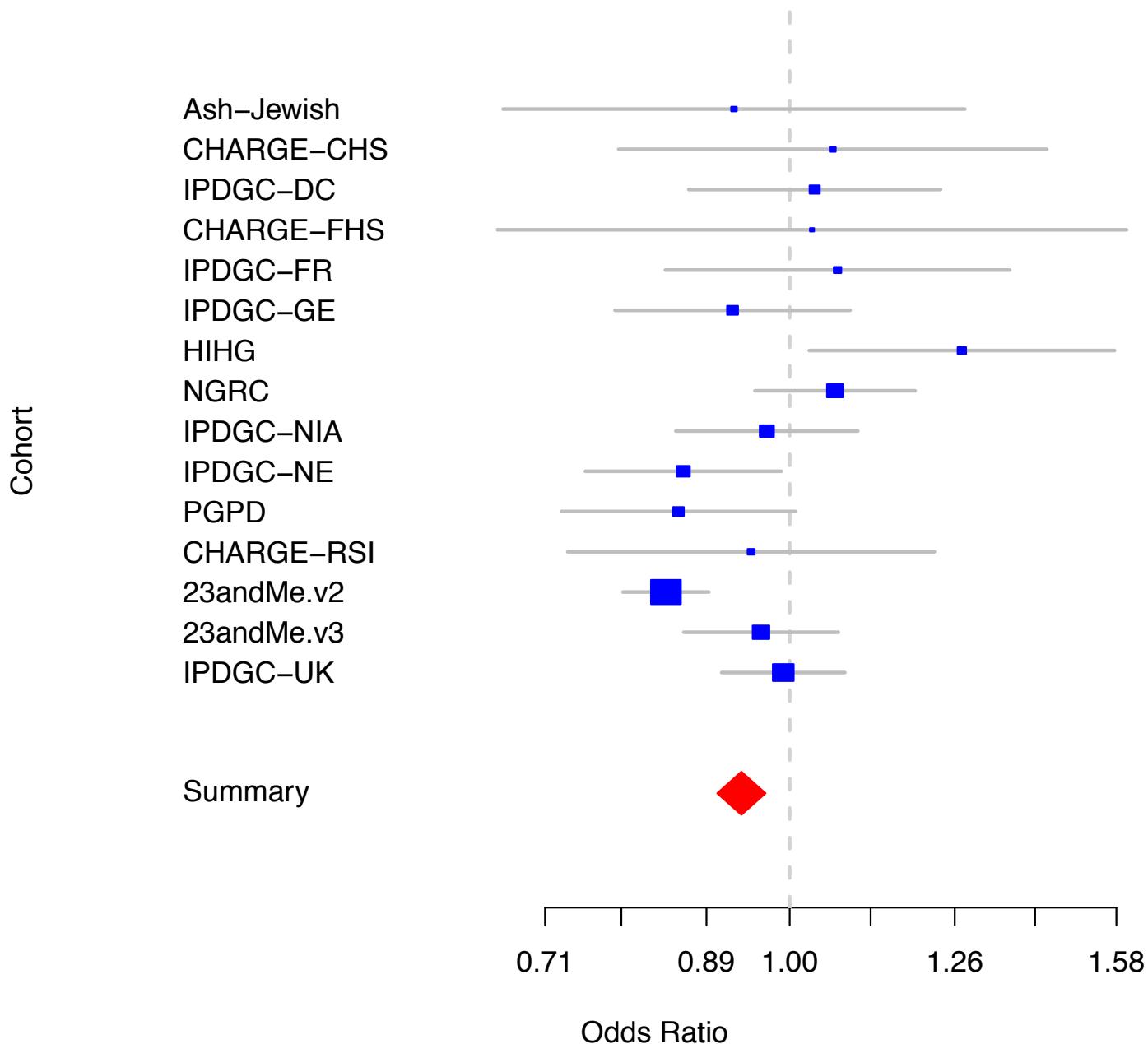
rs60298754 MMP16 Known SNP



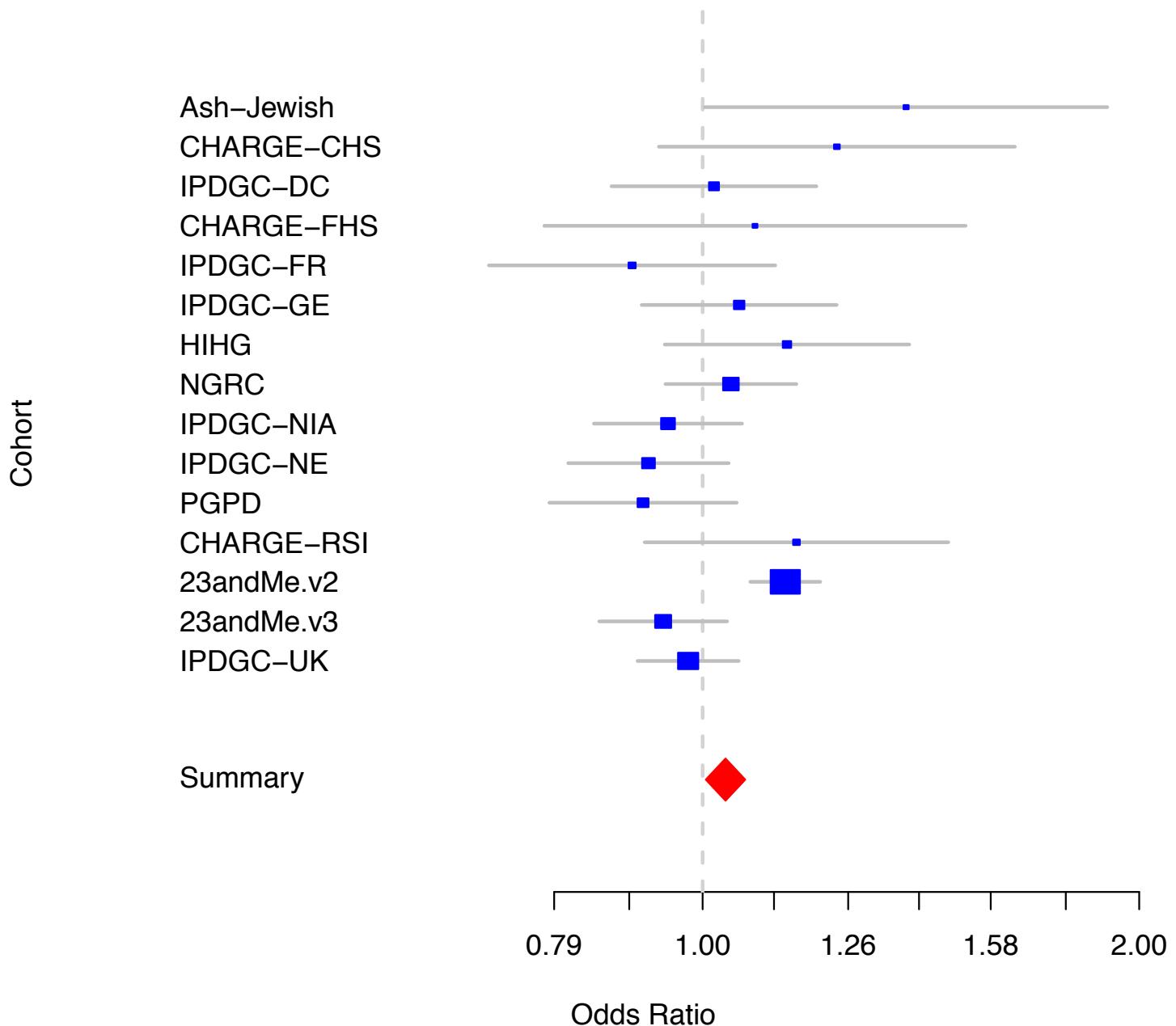
rs7077361 ITGA8 Known SNP



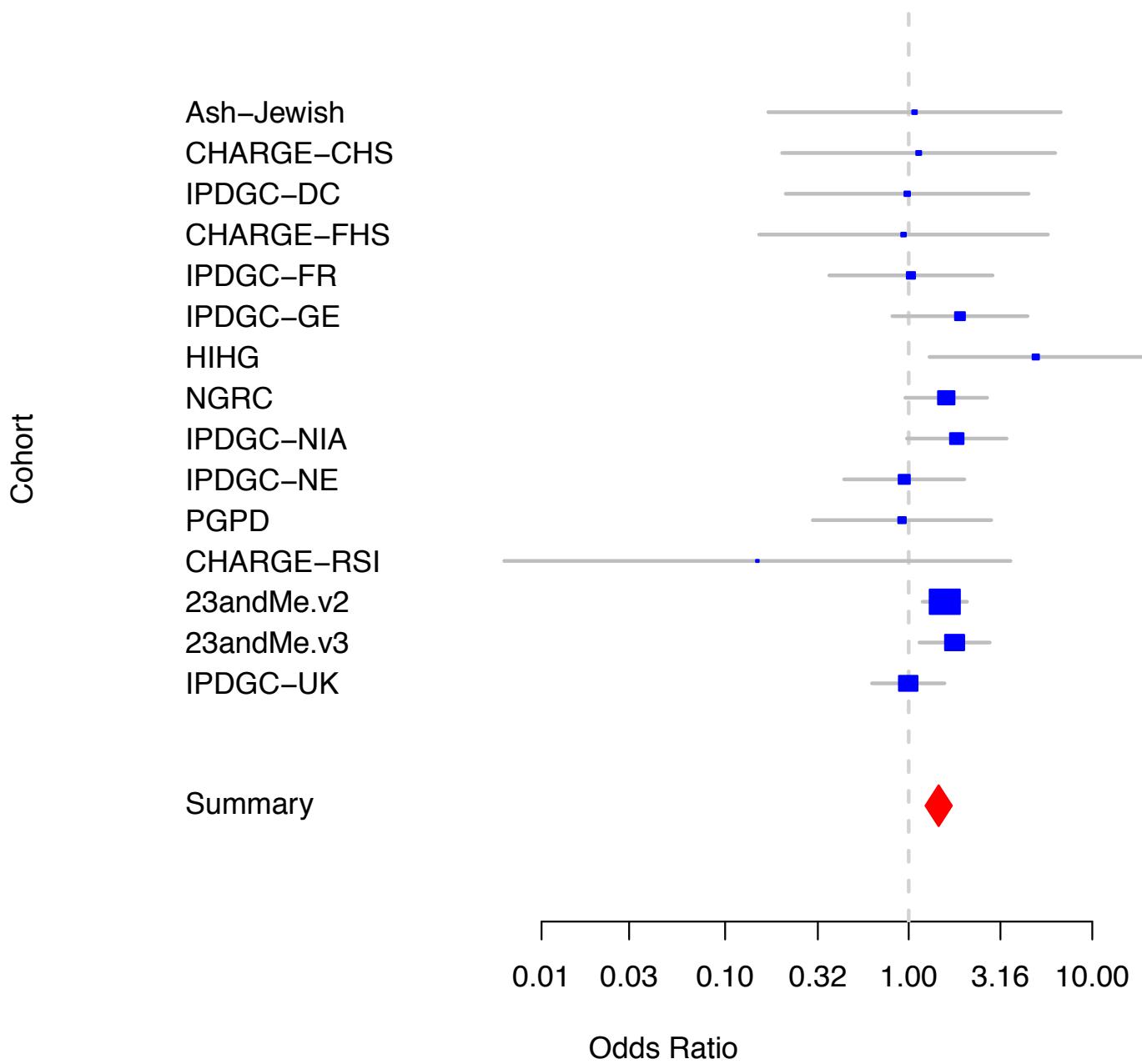
rs11868035 SREBF/RAI1 Known SNP



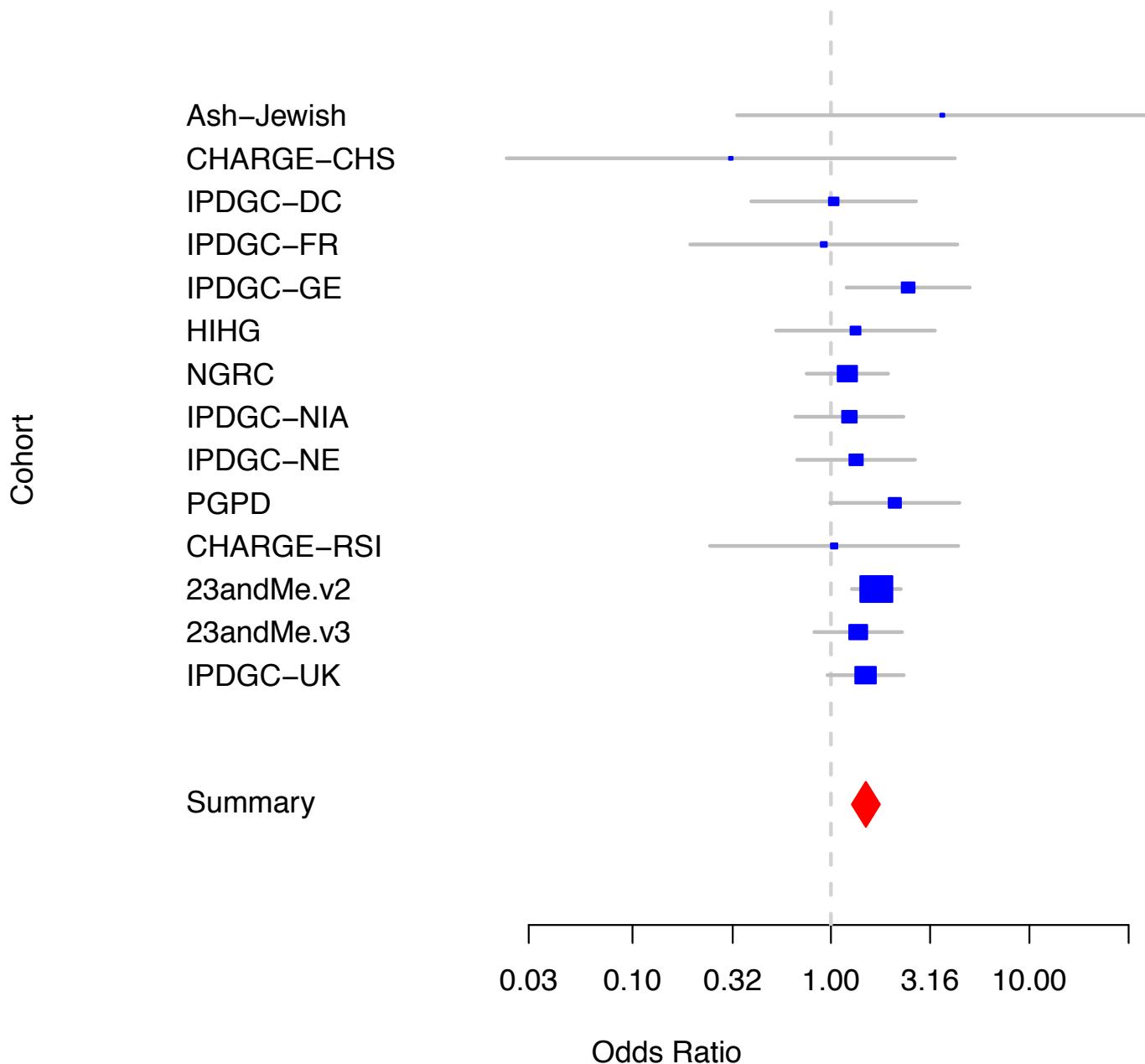
rs2823357 USP25 Known SNP



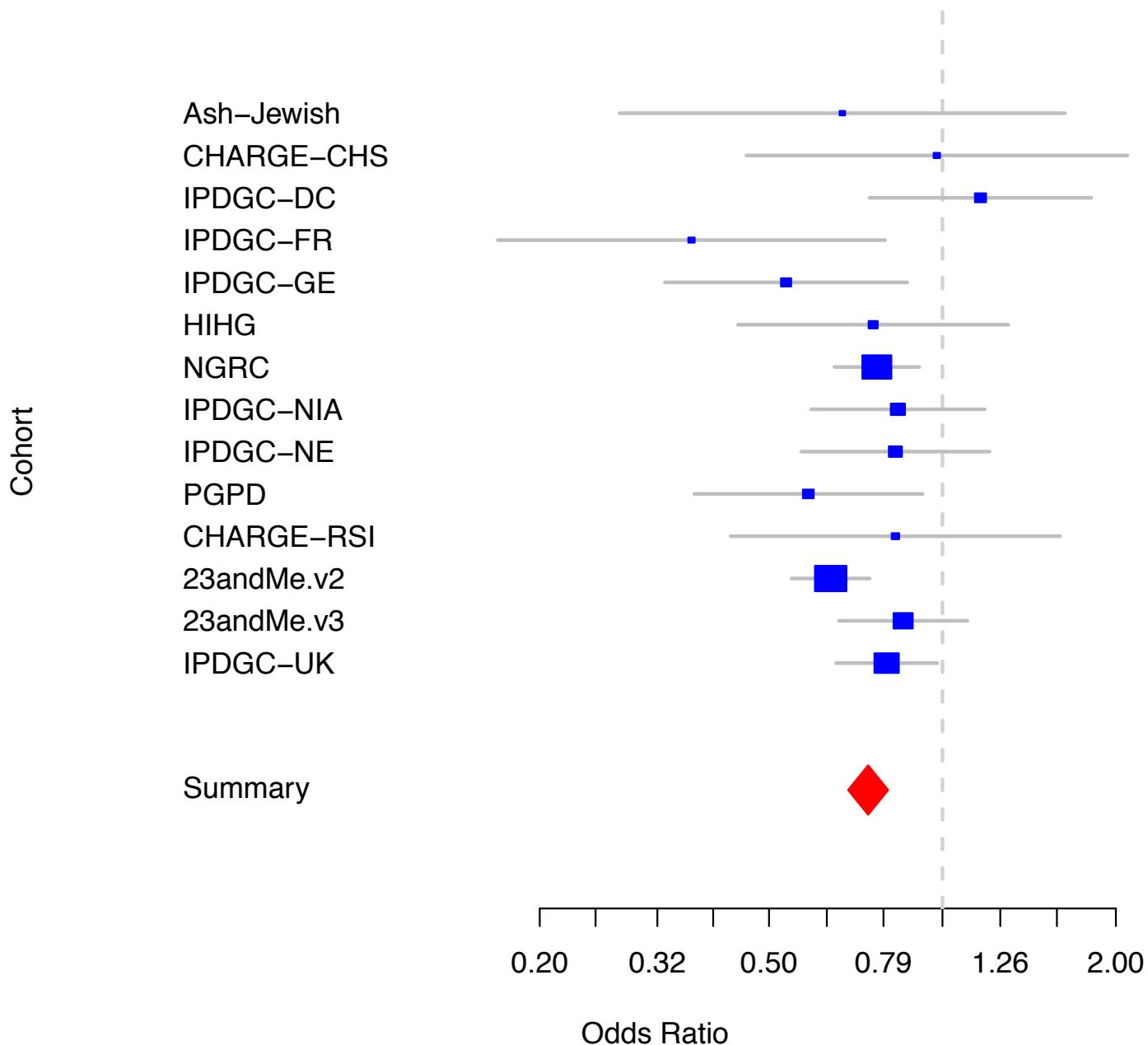
rs79217002 MCCC1 Conditional SNP



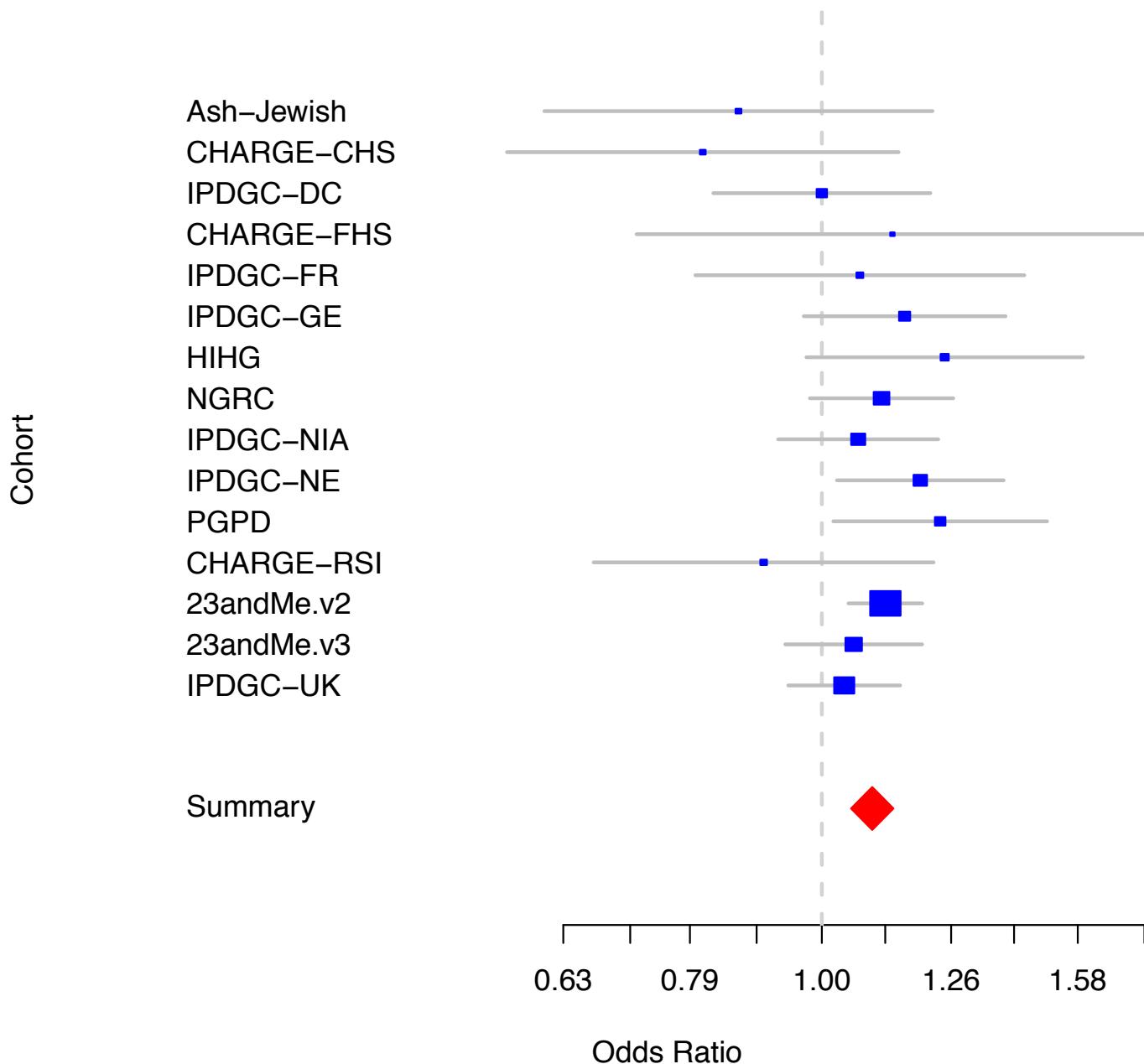
rs114138760 GBA/SYT11 Conditional SNP



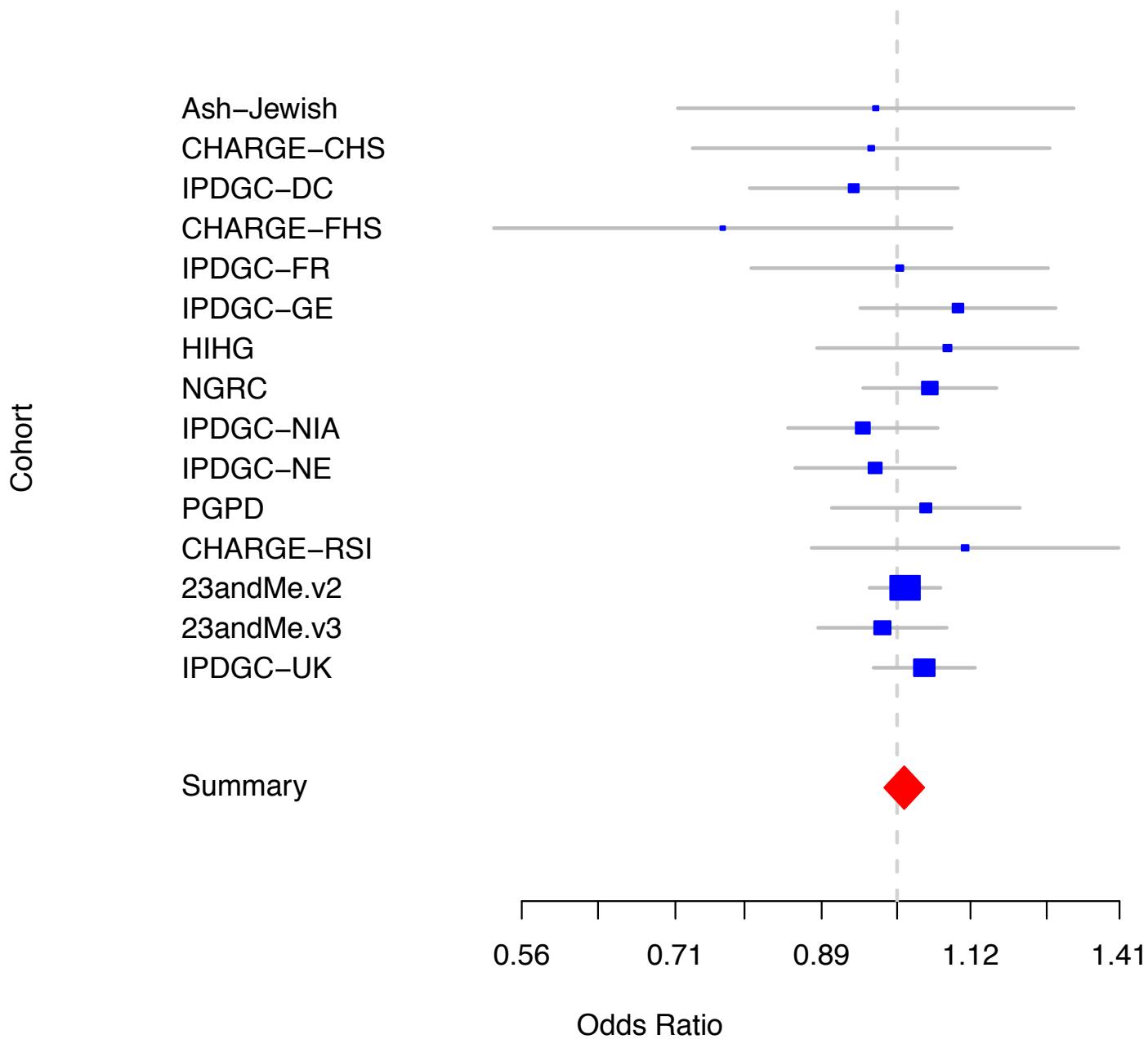
rs34884217 TMEM175/GAK/DGKQ Conditional SNP



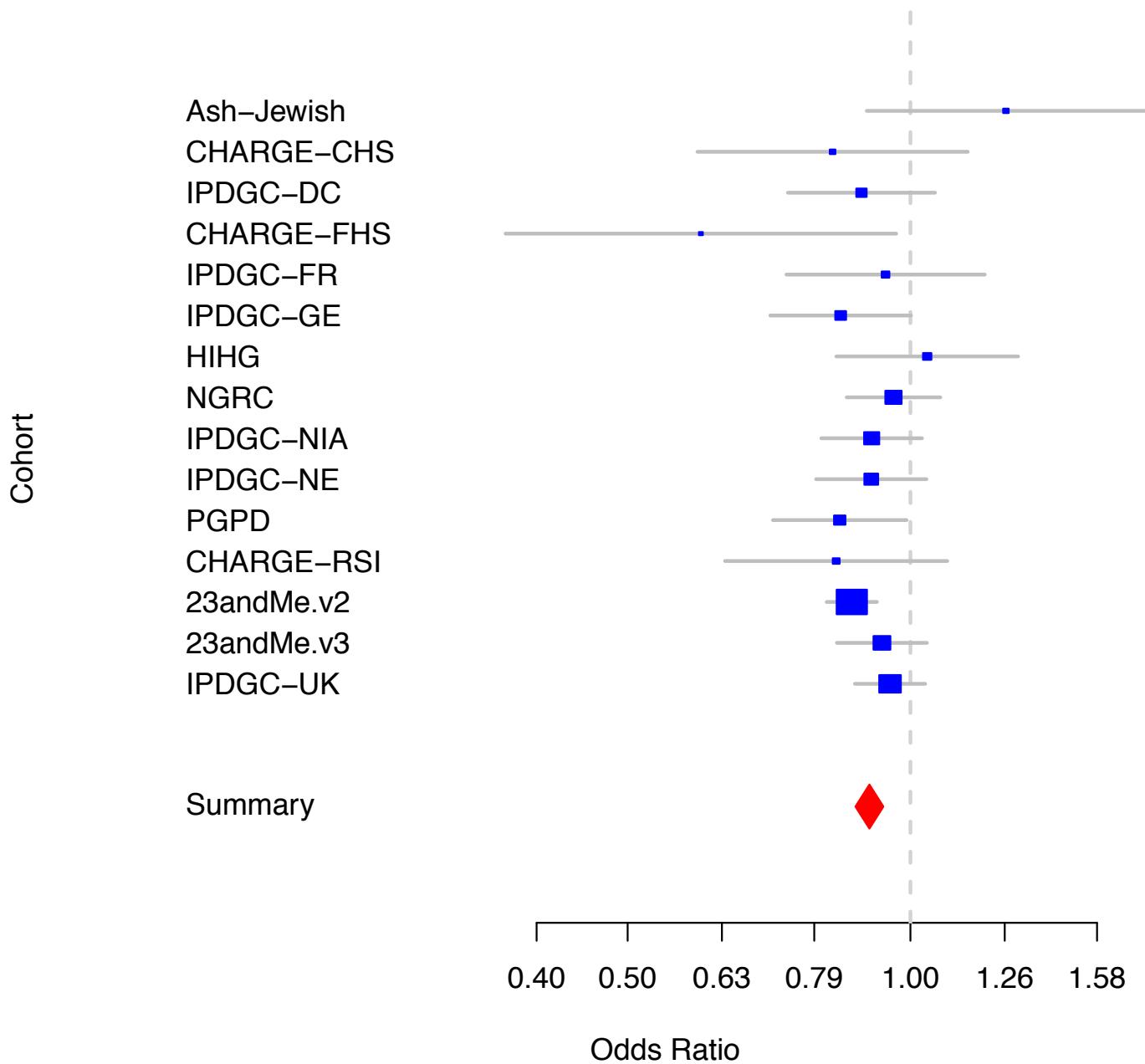
rs1596117 FAM47E/SCARB2 Conditional SNP



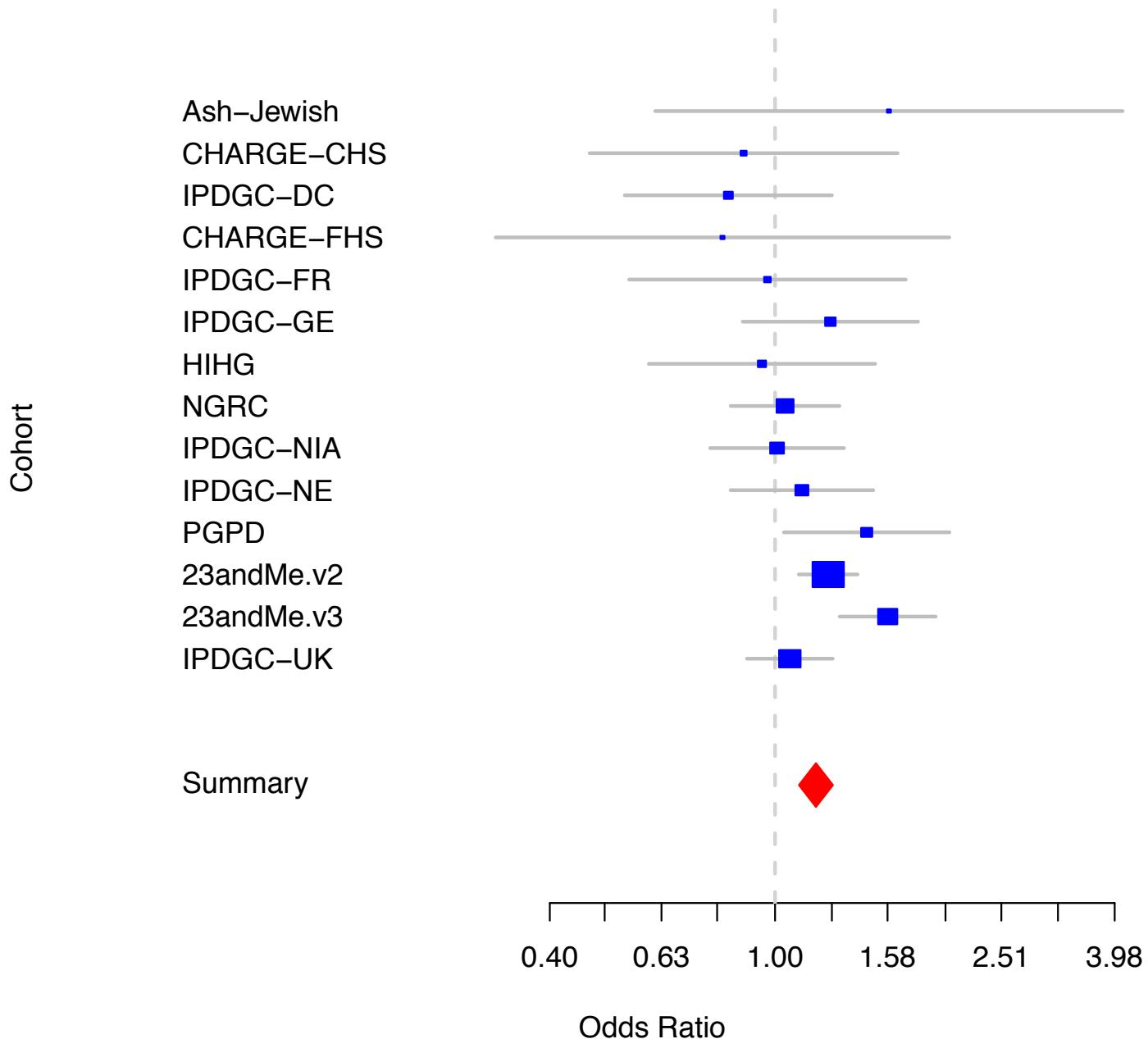
rs7681154 SNCA Conditional SNP



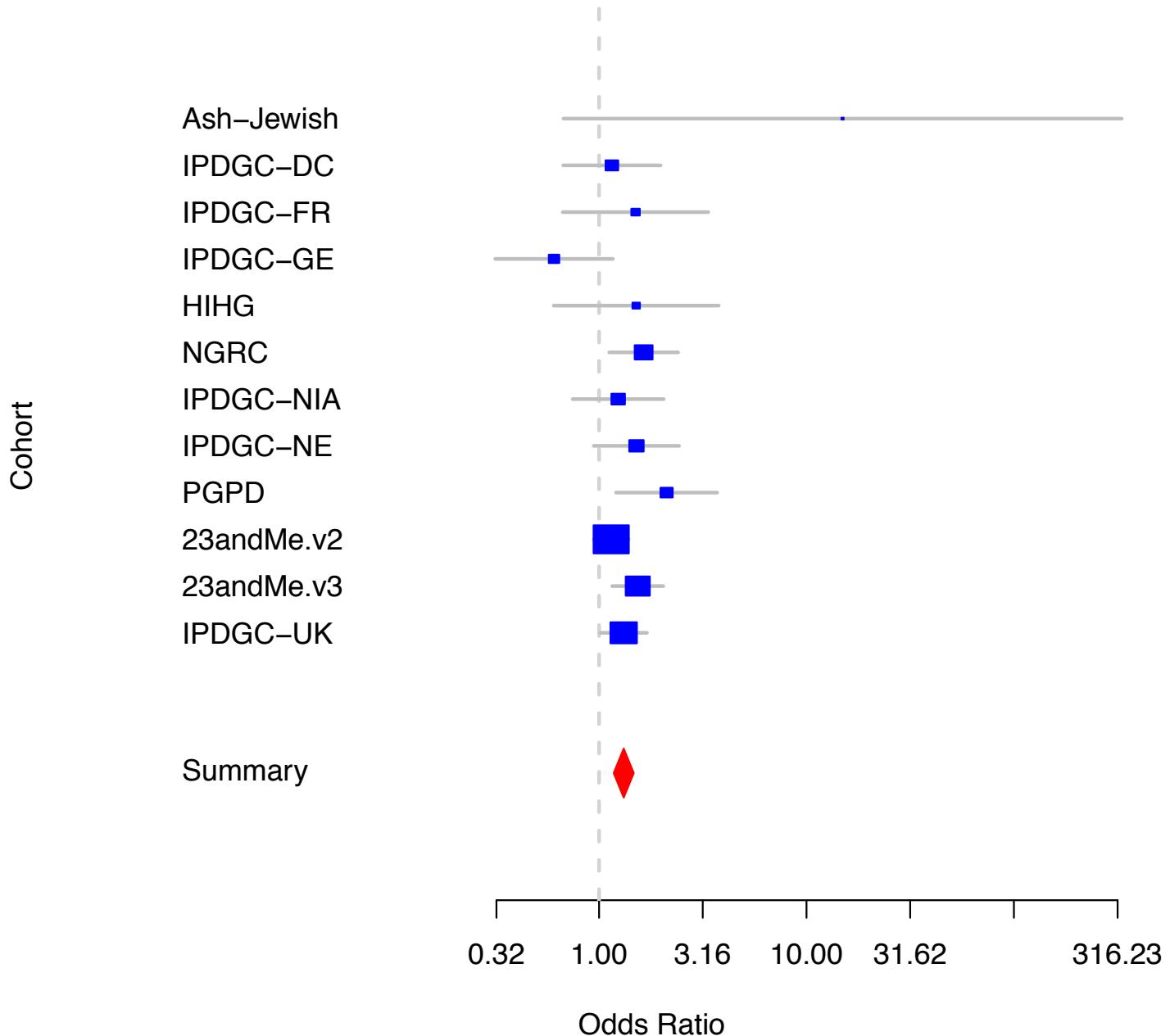
rs10886515 INPP5F Conditional SNP



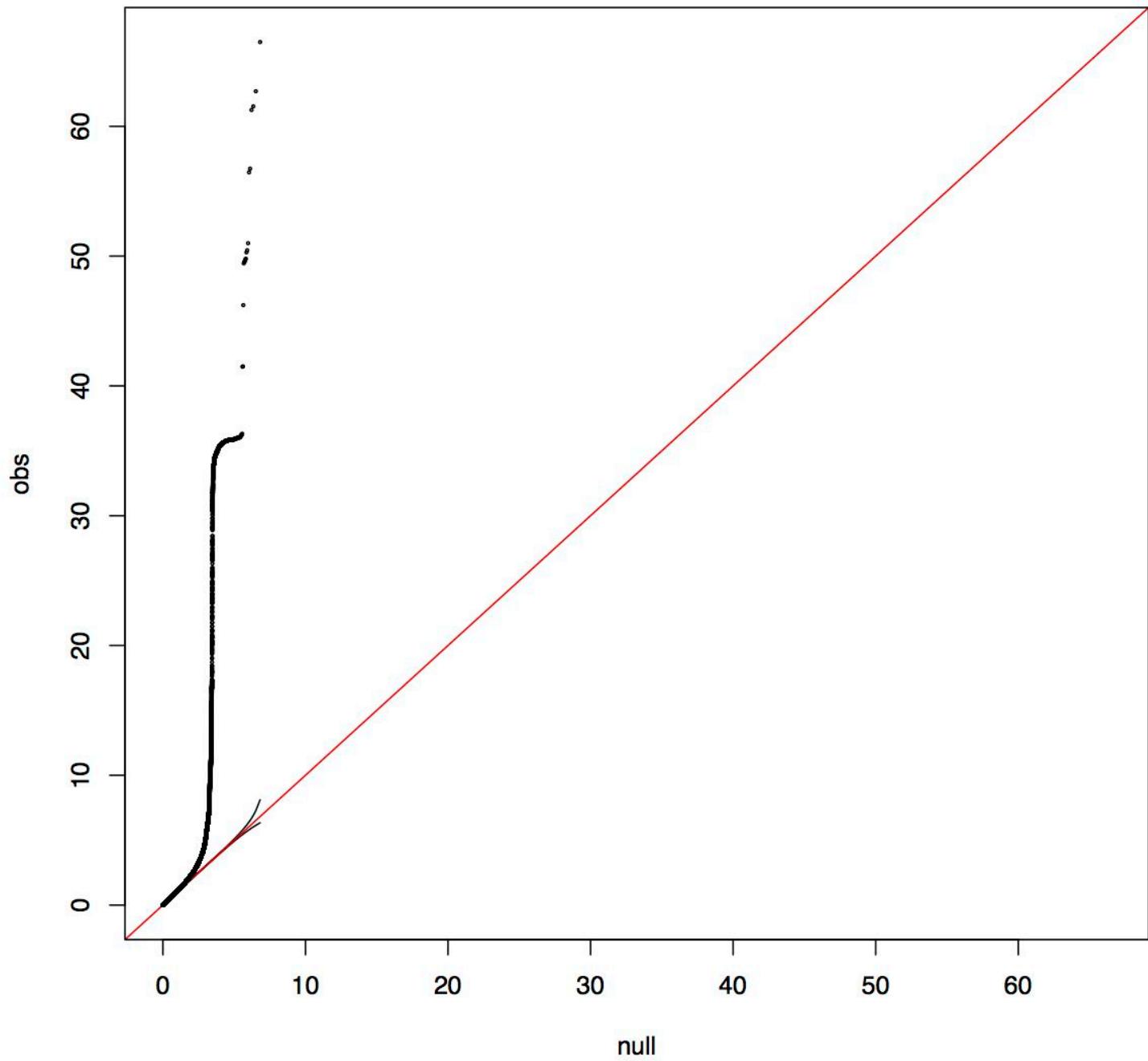
rs13201101 HLA-DQB1 Conditional SNP



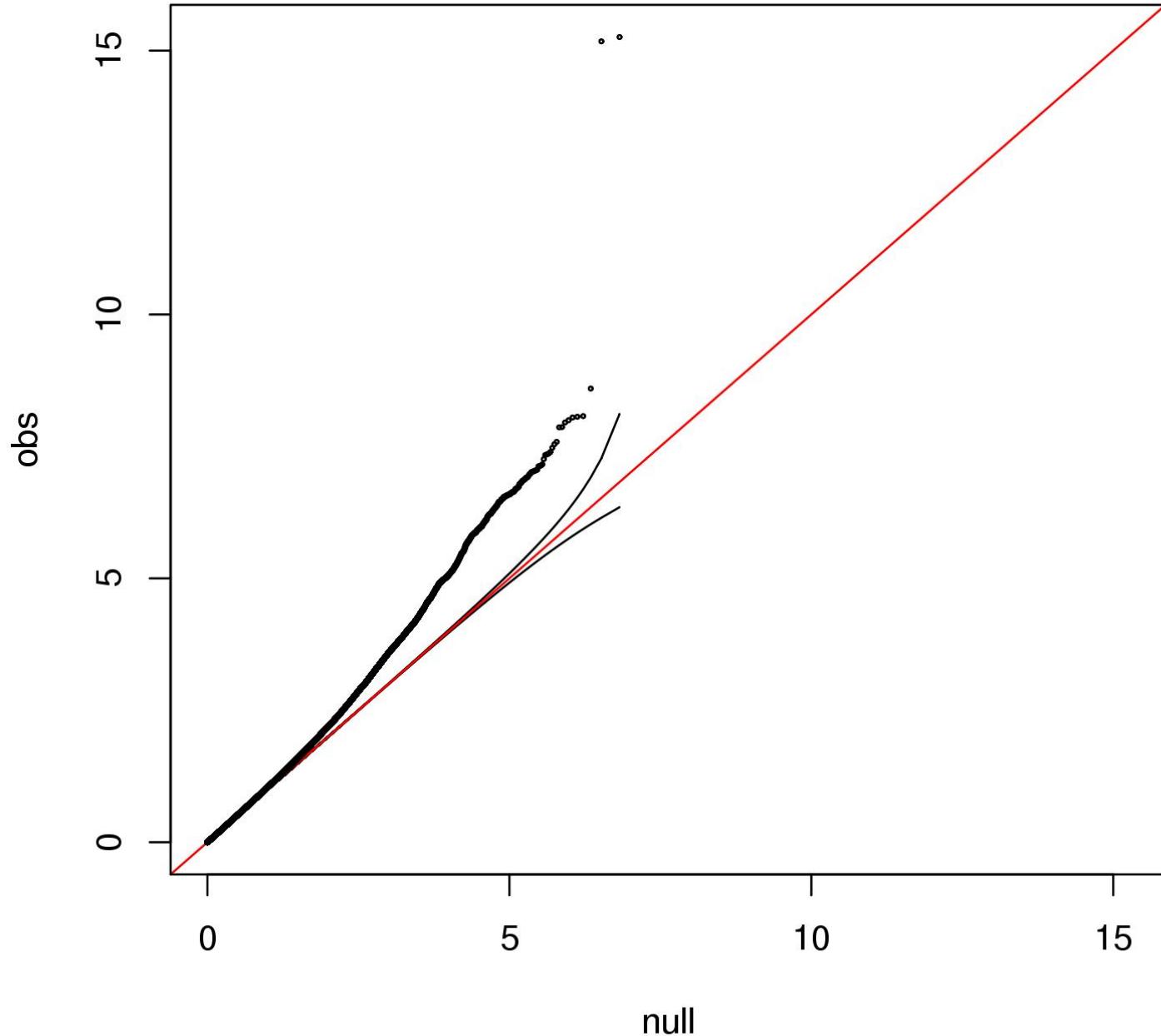
rs117022814 SPPL2B Conditional SNP



Supplementary Figure 3 (page 96):
QQ plot of p-values from discovery
meta-analysis.



Supplementary Figure 4 (page 97):
QQ plot of p-values from discovery
meta-analysis excluding significant
and replicated loci. All SNPs within
+/- 1 megabase of a replicated
genome-wide significant SNP were
excluded (unadjusted lambda =
1.045).



Supplementary Figure 5 (page 98): ROC curve for genetic risk profiles across cohorts adjusting for cohort membership, age and gender.

