

Data Supplement for Mullins et al., GWAS of Suicide Attempt in Psychiatric Disorders and Association With Major Depression Polygenic Risk Scores. Am J Psychiatry (doi: 10.1176/appi.ajp.2019.18080957).

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Supplementary Note

Phenotype definition

Items from structured clinical interviews provided information on self-harm, suicidal ideation, plans and suicide attempt for Psychiatric Genomics Consortium (PGC) cohorts (Supplementary Table 4). Lifetime suicide attempt (SA) was defined across cohorts as a deliberate act of self-harm with at least some intent to result in death. Individuals who did not endorse suicide attempt were included in the non-attempter group and individuals missing information on suicide attempt were excluded. Phenotypic analyses were performed to assess the consistency of the suicide attempt phenotype across the 46 discovery cohorts. First, the association between the psychiatric interview used and the prevalence of suicide attempt in the cohorts was tested, using a linear regression model including psychiatric disorder as a covariate. The only psychiatric interview which showed an association with the prevalence of suicide attempt was the Schedule for Affective Disorders and Schizophrenia (SADS), which was used by eight cohorts and correlated with a higher prevalence of suicide attempt ($P = 0.038$). We note that the SADS interview has a specific item on suicide attempt and does not include self-harm or suicidal ideation (which are assessed using other items in the SADS) (Supplementary Table 4). The interview also specifies that evidence of intent to die is essential for suicide attempt and in the absence of intent “non-suicidal self-injurious behavior” is instead indicated. Therefore, we do not anticipate that the suicide attempter groups from cohorts assessed using the SADS may incorrectly include cases of suicidal ideation only. Furthermore, of the eight cohorts which used the SADS interview, five recruited psychiatric cases from inpatient hospital admissions and hence these cohorts may be more severe and have a higher prevalence of suicide attempt than other cohorts (Supplementary Tables 1-3).

Second, we assessed whether there was any association between prevalence of suicide attempt and interview items which are specific to suicide attempt only or those which assess a spectrum of suicidality (ranging from absence, to self-harm, ideation, plan and suicide attempt) in a single item

(Supplementary Table 4). In the latter, psychiatric cases were split into suicide attempters only versus other psychiatric cases according to the coding of the item. Individuals with missing data were excluded. Using a linear regression model, controlling for psychiatric disorder, there was no association between the prevalence of suicide attempt and the specificity of the item used. Since interviews assessing the spectrum of suicidality in a single item did not produce higher estimates of suicide attempt, this suggests that these do not report cases of suicidal ideation only as suicide attempters. In summary, these analyses suggest that the psychiatric interview used across cohorts does not result in heterogeneity in the definition of suicide attempt.

Polygenic risk scoring

Polygenic risk scoring was used to investigate the genetic relationship between suicide attempt and the psychiatric disorders and to test for overlap in the genetic etiology of suicide attempt between major depressive disorder (MDD), bipolar disorder (BIP) and schizophrenia (SCZ). Table S5 summarises the polygenic scoring analyses conducted, showing the discovery and test datasets used to investigate these hypotheses. PRSice software was used to generate polygenic risk scores (PRS), according to standard protocol (1). Discovery GWAS results were pruned for linkage disequilibrium (LD) using the P value informed clumping method in PLINK (--clump-p1 1 --clump-p2 1 --clump-r2 0.1 --clump-kb 250). This preferentially retains SNPs with the strongest evidence of association and removes SNPs in LD ($r^2 > 0.1$) that show weaker evidence of association within 250Kb windows, based on the LD structure in the test dataset. Subsets of SNPs were selected from the results at nine increasingly liberal P value thresholds ($P < 0.0001$, $P < 0.001$, $P < 0.01$, $P < 0.05$, $P < 0.1$, $P < 0.2$, $P < 0.3$, $P < 0.4$, $P < 0.5$). In the test datasets, the SNP probabilities were converted to best-guess data with a genotype call probability cut-off of 0.8. Sets of alleles, weighted by their log odds ratios (OR) from the discovery GWAS, were summed into PRS for each individual in the test datasets using PLINK. PRS were tested for association with suicide attempter status in the test datasets using a logistic regression model, including five genetic principal components (PCs) and a covariate for each cohort in the test dataset.

The amount of variance explained by the PRS (R^2) is presented on the liability scale, which accounts for the proportion of cases in the test dataset (2).

First, PRS for BIP, major depression and SCZ were used to investigate whether suicide attempters and non-attempters differ in genetic liability for the psychiatric disorder they are affected by. To ensure no overlap between the discovery and test datasets, PRS for psychiatric disorders were generated using PGC cohorts not included in the suicide attempt analyses. All cohorts have been described in previous publications on GWAS of psychiatric disorders conducted by the PGC (3-5). The discovery GWAS for BIP consisted of 11 PGC cohorts totaling 8,711 BIP cases and 15,283 controls, and for SCZ included 25,756 SCZ cases and 35,686 controls from 40 PGC cohorts. The discovery GWAS for major depression is a recent meta-analysis of PGC MDD cohorts and samples from deCODE, GERA, *i*PSYCH, Generation Scotland and UK Biobank (3). The phenotype analysed in this study included clinically defined MDD cases as well as self-reported MDD symptoms or treatment and thus is referred to as 'major depression' (3). Results of this meta-analysis were available in turn excluding each of the 16 PGC MDD cohorts in the suicide attempt study, which allowed us to generate independent PRS for each of the 16 cohorts while maximising the discovery GWAS sample size. These discovery GWAS had approximately 59,000 cases and 112,000 controls. The PRS for BIP, major depression and SCZ were tested for association with suicide attempter versus non-attempter status in the same disorder using logistic regression as described previously. Second, based on the results of these analyses, PRS for major depression were also tested for association with suicide attempt in BIP and SCZ.

Third, in order to investigate genetic overlap in suicide attempt across psychiatric disorders, the results of the three GWAS on suicide attempt (SA in MDD, SA in BIP and SA in SCZ) were used in turn as discovery studies and PRS for suicide attempt were tested for association with SA in the other disorders. The Bonferroni corrected significance threshold for the polygenic scoring analyses is 0.006, adjusting for eight independent tests (Supplementary Table S5).

Replication studies

UK Biobank

Genetic associations with suicide attempt were tested for replication in two independent samples of patients with mood disorders drawn from the UK Biobank and *i*PSYCH. The UK Biobank is a prospective cohort study of 501,726 individuals, recruited at 23 centres across the United Kingdom (6). Genotypic data were available for 488,380 individuals and were imputed to the HRC, UK10K and 1,000 Genomes Phase 3 reference panels using IMPUTE4 to identify \approx 93M variants for 487,409 individuals (7). Variants for analysis were limited to those with minor allele frequency \geq 0.01, imputation INFO-score \geq 0.4, and which were either genotyped or imputed to the HRC reference panel, leaving a total of 7794483 SNPs for analysis. Using the genotyped SNPs, individuals were removed if: recommended by the UK Biobank core analysis team for unusual levels of missingness or heterozygosity; SNP genotype call rate $<$ 98%; related to another individual in the dataset (KING $r <$ 0.044, equivalent to removing up to third-degree relatives inclusive); phenotypic and genotypic gender information was discordant (X-chromosome homozygosity (FX) $<$ 0.9 for phenotypic males, FX $>$ 0.5 for phenotypic females). Removal of relatives was performed using a greedy algorithm, which minimises exclusions (for example, by excluding the child in a mother-father-child trio). All analyses were limited to individuals of White Western European ancestry, as defined by 4-means clustering on the first two genetic principal components provided by the UK Biobank (7). Principal component analysis was also performed on the European-only subset of the data using the software flashpca2 (8).

Extensive phenotypic data are available for UK Biobank participants from health records and questionnaires, including an online follow-up questionnaire focussing on mental health (Mental Health Questionnaire, MHQ). Participants were classified as having a mood disorder if they either self-reported a professional diagnosis of depression or bipolar disorder as part of the MHQ [UK Biobank field 20544] or if they met criteria for depression on MHQ questions derived from the Composite International Diagnostic Interview (CIDI). To meet these latter criteria, participants must have

reported ever feeling depressed [UK Biobank field 20446] or anhedonic [UK Biobank field 20441] for two weeks in a row, for at least most of the day [UK Biobank field 20436] almost every day [UK Biobank field 20439] with more than a little interference with daily activities [UK Biobank field 20440]. In addition, they must have reported experiencing at least five of the following symptoms in this period of depression or anhedonia: depression [UK Biobank field 20446], anhedonia [UK Biobank field 20441], tiredness [UK Biobank field 20449], weight change [UK Biobank field 20536], sleep change [UK Biobank field 20532], loss of concentration [UK Biobank field 20435], worthlessness [UK Biobank field 20450] and thoughts of death [UK Biobank field 20437]. The MHQ additionally contained screening questions for bipolar disorder (9). However, for the purpose of defining potential bipolar disorder, all individuals scoring positively on these screening questions were also required to meet the CIDI depression criteria defined above, and as such participants with potential bipolar disorder were a subset of those meeting criteria for depression. Individuals who self-reported a professional diagnosis of psychosis on the MHQ [UK Biobank field 20544] were excluded. Suicide attempters with mood disorders (n=2149) were defined as those who answered yes to the question “Have you ever harmed yourself with the intention to end your life?” [UK Biobank field 20483]. Non-attempters with mood disorders were defined as those who reported no self-harm on the MHQ (n=35912). A genome-wide association study was performed comparing suicide attempters versus non-attempters with mood disorders using BGenie v.1.2 (7), covarying for 6 PCs, and factors capturing site of recruitment and genotyping batch.

***i*PSYCH**

The *i*PSYCH study was approved by the regional Danish ethics committee and the Danish Data Protection Agency (10). DNA preparation, genotyping on the Illumina PsychChip array and quality control were performed as described previously (11, 12). Individuals with mood disorders were identified based on ICD-10 codes (F30-F39) from the Danish Psychiatric Central Research Register and the National Registry of Patients, both complete until December 31, 2016 (10). Suicide attempters with mood disorders (n=4943) were defined as those with diagnoses of suicide attempt (ICD-10: X60-

X84, equivalent to intentional self-harm), those with suicide attempt indicated as 'reason for contact', and with a main diagnosis of poisoning (ICD-10: T39, T42, T43, and T58) or those with a diagnosis in the ICD-10: F chapter as main diagnosis and report of poisoning by drugs or other substances (ICD-10: T36–T50, T52–T60) or injuries to hand, wrist, and forearm (ICD-10: S51, S55, S59, S61, S65, S69). Only contacts starting at age 10 years old or older were considered. Individuals who died by suicide according to the Cause of Death Register were also included in the suicide attempter group. Non-attempters were defined as mood disorder cases not fulfilling any of these criteria (n=15849).

Analysis of depressive symptoms and suicide attempt in schizophrenia

Data on symptoms of illness in schizophrenia were available for eight of the PGC SCZ cohorts included in the GWAS on suicide attempt. Clinical symptoms were assessed using the OPCRIT (Operational Criteria for Psychotic Illness), PANSS (Positive and Negative Syndrome Scale), Lifetime Dimensions of Psychosis Scale (LDPS) or the Comprehensive Assessment of Symptoms and History (CASH) (13-16). As previously described, factor analyses were performed on these data to identify a quantitative depressive symptom dimension, harmonized across instruments and cohorts (17). Data were available for 1426 suicide attempters and 2428 non-attempters with schizophrenia. The association between standardized depressive symptom-based factor scores and suicide attempt was investigated using a logistic regression model, covarying for sex and cohort. Higher depressive symptom factor scores were significantly associated with suicide attempt (OR = 1.67, C.I. 1.56 -1.79, $P = 2.7 \times 10^{-47}$).

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Supplementary Tables and Figures

Table S1: Description of 16 major depressive disorder cohorts

| Cohort (References) | PGC label | Country | Ascertainment and Evaluation of Cases | Inclusion criteria (lifetime) of Cases | Exclusion criteria (lifetime) of Cases |
|------------------------|----------------|-------------|---|--|---|
| BOMA 1-3 | boma | Germany | Consecutive inpatients; SCID or SADS interview; medical records | DSM-IV MDD; German ancestry; age \geq 18 | BIP, hypomania, NAP, MDD related to SUD |
| CoFaMS 4 | cof3 | Australia | Opportunistic; inpatient & outpatient; SCID or MINI | DSM-IV MDD | BIP, NAP, MDD related to SUD |
| PsyCoLaus 5 | col3 | Switzerland | Random population sample; DIGS | DSM-IV MDD, age 35-66 | BIP, hypomania, NAP, MDD related to SUD |
| GenRED1 6,7 | gens | USA | Opportunistic; DIGS3; medical records or informant (subset) | DSM-IV MDD (recurrent or episode $>$ 3 yrs) & onset $<$ 31 yrs; FHx MDD in sibling or parent | BIP, NAP, mod-severe ID; FHx BIP; if SUD, MDD onsets without $<$ 2y of sobriety |
| GenRED2 6 | grnd | USA | Opportunistic; DIGS; medical records or informant (subset) | DSM-IV MDD (recurrent or episode $>$ 3 yrs) & onset $<$ 31 yrs; FHx MDD in sibling or parent | BIP, NAP, mod-severe ID; FHx BIP; if SUD, MDD onsets without $<$ 2y of sobriety |
| GSK/MPIP 8 | gsk2 | Germany | Inpatients; SCAN | DSM-IV MDD (recurrent, mod-severe) | BIP, NAP, SUD, mood-incongruent psychosis, OCD, PTSD, secondary MD |
| MARS 9-11 | mml2 mmo4 | Germany | Inpatients; CIDI | DSM-IV MDD | BIP, SUD, secondary MD, severe medical conditions |
| NESDA/NTR: NESDA 12,13 | nes1 | Netherlands | Psychiatric outpatients, primary care, & population; CIDI | DSM-IV MDD | BIP, NAP, SUD |
| NESDA/NTR: NTR 12,13 | nes1 | Netherlands | Twin registry; longitudinal MDD sx; CIDI (subset) | DSM-IV MDD | Mania (if interviewed) |
| QIMR 14,15 | qi3c qi6c qi02 | Australia | Australian Twin Registry (proband most severe, sx, or earlier onset); SSAGA | DSM-IV MDD | MDD related to SUD |
| RADIANT-UK 16 | rad3 | UK | UK outpatients from DeNT, DeCC, GENDEP studies; SCAN | DSM-IV MDD (recurrent in DeCC & DeNT; MDD FHx in DeNT) | BIP, NAP, MDD related to SUD; BIP FHx |
| RADIANT-GER 16 | rage | Germany | German outpatients from DeNT, DeCC, GENDEP studies; SCAN | DSM-IV MDD (recurrent in DeCC & DeNT; MDD FHx in DeNT) | BIP, NAP, MDD related to SUD; BIP FHx |
| SHIP 0 17 | shp0 | Germany | Study of Health in Pomerania; CIDI | DSM-IV MDD | BIP, MDD related to SUD |
| STAR*D 18 | stm2 | USA | Outpatients in clinical trial; clinical interviews | DSM-IV MDD | BIP, NAP |

Abbreviations: SCID=Structured Clinical Interview for DSM-IV, SADS=Schedule for Affective Disorders and Schizophrenia, MDD = major depressive disorder, BIP = bipolar disorder, NAP=non-affective psychosis, SUD = substance use disorder, MINI = MINI International Neuropsychiatric Interview, DIGS=Diagnostic Interview for Genetic Studies, FHx = family history, ID = intellectual disability, SCAN = Schedules for Clinical Assessment in Neuropsychiatry, OCD = obsessive compulsive disorder, PTSD = post-traumatic stress disorder, CIDI = Composite International Diagnostic Interview, sx = symptoms

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Table S2: Description of 21 bipolar disorder cohorts

| Cohort (References) | PGC label | Country | Ascertainment and Evaluation of Cases | Inclusion criteria (lifetime) of Cases | Exclusion criteria (lifetime) of Cases |
|--|-----------|------------|---|--|---|
| BOMA-Germany I1-4 | bonn | Germany | Consecutive admissions to in-patient units, SCID-I, SADS-L, medical records, FHx, OPCRIT | DSM-IV BIP I or BIP II | |
| Trinity College Dublin 5 | dub1 | Ireland | Hospitals and Community psychiatric facilities, SCID, case note review | DSM-IV BIP I | |
| FaST, TGEN1, TGEN2 6 | fat2 | USA | Hospitals, ADE, MINI | DSM-IV BIP I or BIP II | |
| French PGC2 6 | fran | France | DIGS, FIGS, medical case notes, mood scales, self-rating questionnaires assessing dimensions | DSM-IV BIP I or BIP II | |
| BACCs 7 | gsk1 | UK, Canada | Advertisements in hospitals, clinics, primary care physician offices, patient support groups, SCAN CATEGO algorithm | DSM-IV or ICD-10 BIP I or BIP II | Dx of intravenous drug dependency/use, mood incongruent psychotic sx, manic episodes only with alcohol/substance abuse/dependence/medical illnesses/medications |
| Mayo Clinic 8 | may1 | USA | Mayo Clinic Bipolar Biobank, patients ascertained through routine clinical appointments, in-patients in mood disorder units and recruitment advertising, SCID | DSM-IV-TR BIP I/ BIP II/ schizoaffective | |
| Pritzker Neuropsychiatric Disorders Research Consortium 7, 9 | mich | USA | NIMH Genetics Initiative Repository, DIGS, FIGS, medical record review | DSM-III or IV BIP I | Suspected major depression |
| STEP1 5,7 | stp1 | USA | ADE, MINI | DSM-IV BIP I | |
| STEP2 | stp2 | USA | Hospitals, ADE, MINI | DSM-IV BIP I or BIP II | |
| TOP7 10 | top7 | Norway | Out-patient and in-patient psychiatric units, SCID-I, case note review, follow up interview | DSM-IV BIP I, BIP II, SAB, BIP-NOS | IQ score < 70 |
| TOP8 10 | top8 | Norway | Out-patient and in-patient psychiatric units, SCID-I, case note review, follow up interview | DSM-IV BIP I, BIP II, SAB, BIP-NOS | IQ score < 70 |
| UCL 5, 11 | ucl | UK | Clinical diagnosis according to UK National Health Service (NHS) psychiatrists at interview, SADS-L, OPCRIT | DSM-IV BIP I | |
| UMEA | ume4 | Sweden | MINI, DIGS, FIGS, SCAN | DSM-IV-TR BIP | |
| WTCCC 5, 7, 12 | wfcc | UK | Individuals in contact with mental health services, SCAN | RDC BIP I, BIP II, SAB, BIP-NOS | |
| GAIN 7, 13 | gain | USA | Multiplex families, sibling pair families or individuals, DIGS, FIGS, medical records | DSM-III-R & IV BIP I or SAB | |
| BOMA-Germany II 14 | bmg2 | Germany | Consecutive admissions to in-patient units, AMDP, medical records, family history, OPCRIT | DSM-IV lifetime BIP | |
| BOMA-Germany III 14 | bmg3 | Germany | Recruited from psychiatric hospitals, AMDP, CID-5, SADS-L, SCID, medical records, family history, OPCRIT | DSM-IV lifetime BIP | |
| BOMA-Poland 14 | bmpo | Poland | Recruited from Department of Psychiatry, SCID | DSM-IV lifetime BIP | |
| BOMA-Spain 14 | bmsp | Spain | Recruited from hospital mental health departments, SADS-L, OPCRIT, medical records, FISC | DSM-IV and RDC BIP | |
| Nova Scotia | hal2 | Canada | Recruited from specialty mood disorder clinics, SADS-L | DSM-IV and RDC BIP | |
| BOMA-Romania 15 | rom3 | Romania | Consecutive admissions to psychiatric hospital, DIGS, FIGS, medical records, family reports | DSM-IV BIP I | |

Abbreviations: SCID = Structured Clinical Interview for DSM-IV, SADS-L = Schedule for Affective Disorders and Schizophrenia Lifetime Version, FHx = family history, OPCRIT = Operational Criteria Checklist, BIP = bipolar disorder, ADE = Affective Disorders Evaluation, MINI = MINI International Neuropsychiatric Interview, DIGS = Diagnostic Interview for Genetic Studies, FIGS = Family Interview for Genetic Studies, SCAN = Schedules for Clinical Assessment in Neuropsychiatry, Dx = diagnosis, sx = symptoms, SAB = Seasonal affective disorder, BIP-NOS = bipolar disorder not otherwise specified, AMDP = Association Methodology and Documentation in Psychiatry, CID-5 = Composite International Diagnostic Screener, FISC = Family Informant Schedule and Criteria, RDC = Research Diagnostic Criteria

| References | Number | Citation |
|------------|--------|--|
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Table S3: Description of 9 schizophrenia cohorts

| Cohort (References) | PGC label | Country | Ascertainment and Evaluation of Cases | Inclusion criteria (lifetime) of Cases | Exclusion criteria (lifetime) of Cases |
|---------------------------------------|-----------|----------------|---|---|---|
| Bonn/ Mannheim 1 | boco | Germany | Consecutive hospital admissions, SCID, SADS-L, OPCRIT, medical records, FHx | DSM-IV SCZ | |
| Bulgaria 2 | butr | Bulgaria | Family trios where proband had SCZ/ SA, SCAN | DSM-IV SCZ / SA | Mental retardation |
| Denmark 1 | denm | Denmark | Psychiatric departments and twin pair studies, OPCRIT | ICD-10 SCZ | Mania/bipolar illness |
| Molecular Genetics of Schizophrenia 3 | mgs2 | USA, Australia | Clinical settings and community residences, DIGS 2.0, FIGS 2.0, Medical records | DSM-IV SCZ / SA | |
| Munich 1 | munc | Germany | Cases diagnosed with SCZ from the Munich area, SCID interview | DSM-IV SCZ | Head injury/ neurological diseases |
| Portugal 4 | port | Portugal | Probands from families segregating SCZ, DIGS, SIS, SANS, SAPS, OPCRIT | DSM-IV SCZ | Bipolar disorder |
| Thematic Organized Psychosis Research | top8 | Norway | Out-patient and in-patient psychiatric units, SCID-I interview | DSM-IV SCZ/ SA/ schizophreniform disorder | IQ score < 70 |
| UCLA 1 | ucla | Netherlands | Inpatients and outpatients recruited through psychiatric hospitals and institutions, CASH | DSM-IV SCZ | Short-term drug-induced psychoses, psychoses with learning disability/ head injury, other symptomatic psychoses |
| University College London 4 | uclo | UK | SCZ diagnosis recorded in medical case-history, SADS-L, RDC | ICD-10 SCZ | SA, bipolar disorder, schizomania |

Abbreviations: SCID = Structured Clinical Interview for DSM-IV, SADS = Schedule for Affective Disorders and Schizophrenia, OPCRIT = Operational Criteria Checklist, FHx = family history, SCZ = schizophrenia, SA = schizoaffective disorder, SCAN = Schedules for Clinical Assessment in Neuropsychiatry, DIGS = Diagnostic Interview for Genetic Studies, FIGS = Family Interview for Genetic Studies, SIS = Kendler's Structured Interview for Schizotypy, SANS = Schedule for the Assessment of Negative Symptoms, SAPS = Schedule for the Assessment of Positive Symptoms, CASH = Comprehensive Assessment of Symptoms and History, RDC = Research Diagnostic Criteria

| References | Number | Citation |
|------------|--------|---|
| | 1 | Stefansson, H. <i>et al.</i> Common variants conferring risk of schizophrenia. <i>Nature</i> 460, 744-7 (2009). |
| | 2 | Kirov G. <i>et al.</i> De novo CNV analysis implicates specific abnormalities of postsynaptic signalling complexes in the pathogenesis of schizophrenia. <i>Molecular Psychiatry</i> 17, 142-153 (2012). |
| | 3 | Shi, J. <i>et al.</i> Common variants on chromosome 6p22.1 are associated with schizophrenia. <i>Nature</i> 460, 753-7 (2009). |
| | 4 | International Schizophrenia Consortium. Rare chromosomal deletions and duplications increase risk of schizophrenia. <i>Nature</i> 455, 237-41 (2008). |
| | 5 | Athanasou, L. <i>et al.</i> Gene variants associated with schizophrenia in a Norwegian genome-wide study are replicated in a large European cohort. <i>Journal of Psychiatric Research</i> 44, 748-53 (2010). |

Table S4. Items on suicide from psychiatric interviews

| Psychiatric Interview | Section/ Question | Information Collected |
|---|--|--|
| SCAN (Schedules for Clinical Assessment in Neuropsychiatry) | 6.011 Suicide attempt and self-harm during episode of depression | 0=absent, 1=deliberately considered suicide or self-injury but made no attempt, 2= injured self or made an attempt but no serious harm results, 3 = as 2 but with serious self-harm, 4 = made an attempt at suicide designed to result in death |
| SCID (Structured Clinical Interview for DSM-IV) | Sections on Depression, Mania, Mixed states | Recurrent thoughts of death (not just fear of dying), recurrent suicidal ideation without a specific plan, or a suicide attempt or a specific plan for committing suicide |
| SADS (Schedule for Affective Disorders and Schizophrenia) | Section O Suicidal Behavior | Ever made a suicide attempt, describe the most serious attempt, most serious attempt is rated by the interviewer in terms of intent and lethality |
| DIGS (Diagnostic Interview for Genetic Studies) | Section O Suicidal Behavior | Ever made a suicide attempt, describe the most serious attempt, medical treatment or hospitalisation required, whether the patient wanted to die or thought they would die, most serious attempt is rated by the interviewer in terms of intent and lethality |
| OPCRIT (Operational Criteria Checklist) | Past psychiatric history | Ever made suicide attempt |
| MINI (MINI International Neuropsychiatric Interview) | Section C Suicidality | Lifetime suicide attempt, in the past month thoughts about suicide, suicide plan, suicide attempt, hoped to survive or expected to die |
| CIDI (Composite International Diagnostic Interview) | Section on Major Depression | During worst two weeks in the last 12 months - thought about committing suicide, suicide plan, suicide attempt |
| SSAGA (Semi-Structured Assessment for the Genetics of Alcoholism) | Section I Depression, Section N Suicidal Behavior | Thoughts of death or suicide, suicide plan, suicide attempt, describe the most serious attempt, method, medical treatment, hospitalisation, whether the patient wanted to die or thought they would die, interviewer rates both the lethality and intent from unclear to extreme |
| FIGS (Family Interview for Genetic Studies) | During depression | Did the family member talk about death or suicide, try suicide |
| CASH (Comprehensive Assessment of Symptoms and History) | Major Depressive Syndrome | Thoughts about death and suicide, plus possible wishes to be dead, suicide plans, suicide attempts, rated from mild to severe |

Table S5: Summary of polygenic risk scoring analyses with arrows showing direction from discovery to test dataset

| Test for genetic overlap between psychiatric disorders and suicide attempt in the same disorder | | |
|--|---|---------------------------------------|
| PGC BIP 1 | → | Suicide attempt vs non-attempt in BIP |
| PGC Major Depression 2 | → | Suicide attempt vs non-attempt in MDD |
| PGC SCZ 3 | → | Suicide attempt vs non-attempt in SCZ |
| Test for genetic overlap between major depression and suicide attempt in other disorders | | |
| PGC Major Depression 2 | → | Suicide attempt vs non-attempt in BIP |
| PGC Major Depression 2 | → | Suicide attempt vs non-attempt in SCZ |
| Test for genetic overlap in suicide attempt across psychiatric disorders | | |
| Suicide attempt vs non-attempt in BIP | ↔ | Suicide attempt vs non-attempt in MDD |
| Suicide attempt vs non-attempt in BIP | ↔ | Suicide attempt vs non-attempt in SCZ |
| Suicide attempt vs non-attempt in SCZ | ↔ | Suicide attempt vs non-attempt in MDD |

PGC, Psychiatric Genomics Consortium; BIP, bipolar disorder; MDD, major depressive disorder, SCZ, schizophrenia. 1. Stahl et al. Genomewide association study identifies 30 loci associated with bipolar disorder. *bioRxiv*, 2017. 2. Wray et al. Genome-wide association analyses identify 44 risk variants and refine the genetic architecture of major depression. *Nat Genet.* 2018;50(5):668-81. 3. Schizophrenia Working Group of the Psychiatric Genomics Consortium. Biological insights from 108 schizophrenia-associated genetic loci. *Nature.* 2014;511(7510):421-7.

Table S6: Summary of suicide attempt in major depressive disorder cohorts *

| Cohort | N Suicide attempters | N Non-attempters |
|--------------|----------------------|------------------|
| CoFaMS | 27 | 74 |
| PsyCoLaus | 65 | 442 |
| GenRED2 | 168 | 653 |
| GSK MPIP | 115 | 763 |
| MARS 650 | 137 | 407 |
| MARS OMNIex | 38 | 187 |
| NTR/NESDA | 229 | 1146 |
| QIMR I317 | 61 | 521 |
| QIMR I610 | 32 | 263 |
| QIMR COEX | 48 | 299 |
| RADIANT-UK | 150 | 1424 |
| RADIANT-Ger | 35 | 276 |
| STAR*D | 126 | 807 |
| BOMA | 170 | 361 |
| SHIP 0 | 18 | 348 |
| GenRED1 | 203 | 815 |
| Total | 1622 | 8786 |

*Individuals missing information on suicide attempt were excluded.

Table S7: Summary of suicide attempt in bipolar disorder cohorts*

| Cohort | N Suicide attempters | N Non-attempters |
|---|----------------------|------------------|
| BOMA-Germany I | 241 | 365 |
| Trinity College Dublin | 26 | 26 |
| FaST, TGEN1, TGEN2 | 120 | 124 |
| French PGC2 | 185 | 254 |
| BACCs | 77 | 505 |
| Mayo Clinic | 307 | 610 |
| Pritzker Neuropsychiatric Disorders Research Consortium | 161 | 310 |
| STEP2 | 170 | 363 |
| STEP1 | 392 | 453 |
| TOP7 | 116 | 207 |
| TOP8 | 48 | 94 |
| UCL | 182 | 74 |
| UMEA | 74 | 124 |
| WTCCC | 423 | 673 |
| GAIN | 233 | 277 |
| BOMA-Germany II | 62 | 119 |
| BOMA-Germany III | 132 | 234 |
| Boma Poland | 150 | 251 |
| Boma Spain | 22 | 66 |
| Nova Scotia Canada | 78 | 223 |
| BOMA Romania | 65 | 148 |
| Total | 3264 | 5500 |

*Individuals missing information on suicide attempt were excluded.

Table S8: Summary of suicide attempt in schizophrenia cohorts*

| Cohorts | N Suicide attempters | N Non-attempters |
|---|----------------------|------------------|
| Bonn/ Mannheim | 287 | 310 |
| Bulgaria | 103 | 208 |
| Denmark | 28 | 98 |
| Molecular Genetics of Schizophrenia | 754 | 1202 |
| Munich | 156 | 263 |
| Portugal | 80 | 243 |
| Thematic Organized Psychosis Research Study | 100 | 214 |
| UCLA | 86 | 304 |
| University College London | 89 | 104 |
| Total | 1683 | 2946 |

*Individuals missing information on suicide attempt were excluded.

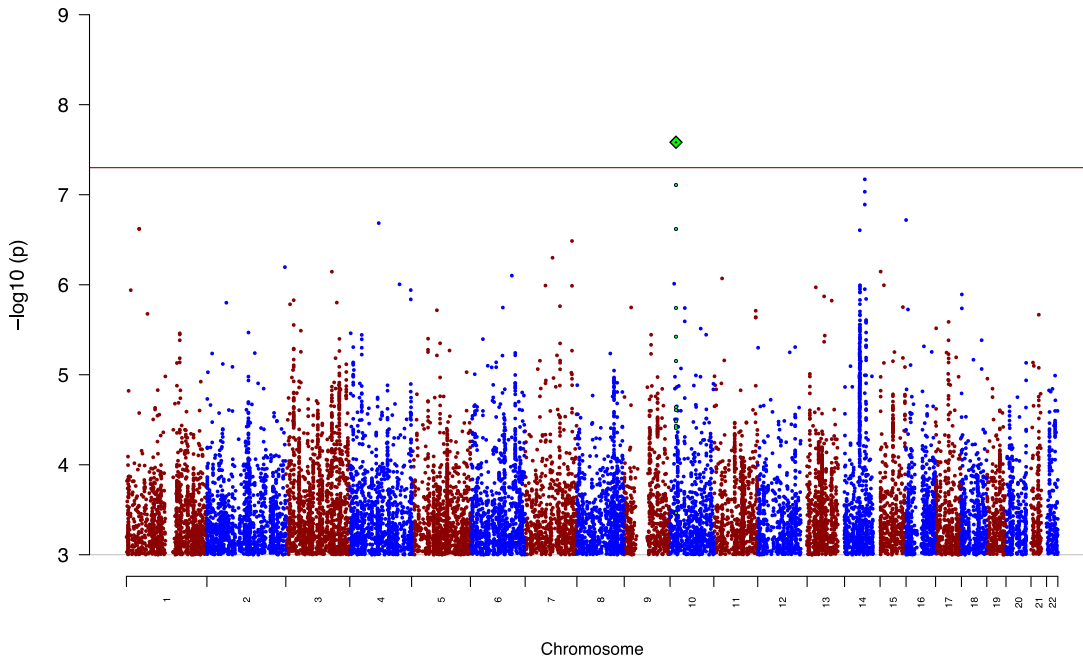


Figure S1: Manhattan plot from GWAS of suicide attempt in major depressive disorder

Table S9: Top 20 results from GWAS of suicide attempt in major depressive disorder showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non-attempters | P value | OR (C.I.) | Direction in each cohort |
|------------------|-----|-----------|-------|--------------------|------------------------|-----------------|-----------------|--------------------------|
| rs45593736 | 10 | 18954937 | A/G | 0.02 | 0.01 | 2.61E-08 | 2.38(1.75-3.23) | +++?+++++?+-?+ |
| rs111625585 | 14 | 82804332 | T/C | 0.08 | 0.06 | 6.75E-08 | 1.57(1.33-1.84) | +++++-----+----- |
| rs116428372 | 16 | 589359 | A/G | 0.07 | 0.06 | 1.91E-07 | 1.74(1.41-2.14) | +++++-----+----- |
| rs77033326 | 4 | 89777618 | A/G | 0.03 | 0.02 | 2.07E-07 | 2.24(1.65-3.03) | +-----+-----+----- |
| rs183414028 | 1 | 40442026 | T/C | 0.98 | 0.99 | 2.39E-07 | 0.39(0.27-0.56) | -----+----- |
| rs113330417 | 14 | 67249421 | A/G | 0.96 | 0.98 | 2.48E-07 | 0.57(0.46-0.71) | -----+----- |
| rs111367251 | 7 | 144968289 | C/G | 0.98 | 0.99 | 3.26E-07 | 0.45(0.33-0.61) | -?+-----+----- |
| rs62460873 | 7 | 84977966 | T/C | 0.98 | 0.99 | 5.01E-07 | 0.37(0.25-0.54) | ?-???-?-----+----- |
| rs186736781 | 2 | 240473090 | T/C | 0.03 | 0.02 | 6.37E-07 | 2.27(1.64-3.13) | +?+?+-----+----- |
| chr15_24344805_D | 15 | 24344805 | D/I3 | 0.37 | 0.41 | 7.14E-07 | 0.79(0.73-0.87) | -----+-----+----- |
| rs111326206 | 3 | 142438431 | T/C | 0.95 | 0.97 | 7.16E-07 | 0.62(0.51-0.75) | +-----+?+----- |
| chr6_128178230_I | 6 | 128178230 | I2/D | 0.08 | 0.06 | 7.92E-07 | 1.58(1.32-1.90) | -----+----- |
| rs184924771 | 11 | 25885205 | A/C | 0.98 | 0.99 | 8.51E-07 | 0.39(0.27-0.57) | -?+?+-----+----- |
| rs113386487 | 10 | 13358583 | A/T | 0.97 | 0.98 | 9.73E-07 | 0.47(0.35-0.64) | -?+?+-----+----- |
| rs13137453 | 4 | 153907879 | A/G | 0.97 | 0.98 | 9.88E-07 | 0.46(0.34-0.63) | -?+?+-----+----- |
| rs9972552 | 15 | 34396913 | A/C | 0.03 | 0.02 | 1.01E-06 | 2.18(1.59-2.98) | +-----+-----+----- |
| rs191852465 | 7 | 63164142 | T/C | 0.96 | 0.97 | 1.02E-06 | 0.46(0.34-0.63) | +-----+-----+----- |
| chr13_46256859_D | 13 | 46256859 | D/I6 | 0.02 | 0.01 | 1.07E-06 | 2.27(1.63-3.16) | -----+?+-----+----- |
| rs145440507 | 4 | 188400537 | A/T | 0.98 | 0.99 | 1.15E-06 | 0.37(0.25-0.55) | -?+?+-----+----- |
| rs76347430 | 1 | 14395819 | A/G | 0.97 | 0.98 | 1.15E-06 | 0.50(0.38-0.66) | ?-?+-----+----- |

CHR, chromosome; BP, basepair position; freq, frequency; OR, odds ratio; CI, confidence interval

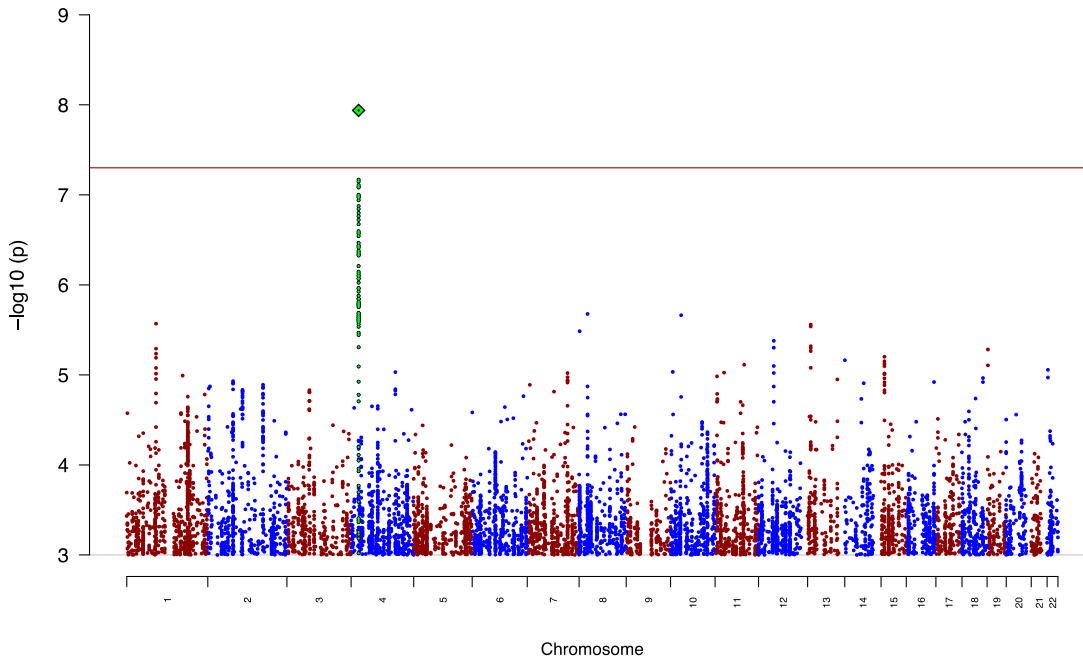


Figure S2: Manhattan plot from GWAS of suicide attempt in bipolar disorder

Table S10: Top 20 results from GWAS of suicide attempt in bipolar disorder showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non-attempters | P value | OR (C.I.) | Direction in each cohort |
|------------------|-----|-----------|-------|--------------------|------------------------|-----------------|-----------------|--------------------------|
| chr4_23273116_D | 4 | 23273116 | D/I10 | 0.20 | 0.17 | 1.15E-08 | 1.29(1.18-1.40) | +++++++----- |
| rs1052873 | 8 | 27667793 | T/C | 0.19 | 0.22 | 2.10E-06 | 0.83(0.76-0.89) | ----- |
| rs118167891 | 10 | 32946009 | T/C | 0.02 | 0.02 | 2.17E-06 | 1.93(1.47-2.53) | +++++?+--- |
| rs6428588 | 1 | 90825206 | T/C | 0.30 | 0.34 | 2.69E-06 | 0.85(0.79-0.91) | ----- |
| rs7982251 | 13 | 28909835 | T/C | 0.85 | 0.87 | 2.76E-06 | 0.79(0.72-0.87) | -----+--- |
| rs67658161 | 8 | 3286733 | A/C | 0.48 | 0.44 | 3.27E-06 | 1.16(1.09-1.24) | +++++----- |
| rs7979008 | 12 | 47479528 | A/C | 0.31 | 0.34 | 4.18E-06 | 0.85(0.79-0.91) | -----+--- |
| rs55893662 | 19 | 2955759 | T/C | 0.11 | 0.10 | 5.22E-06 | 1.46(1.24-1.73) | +++++----- |
| rs5016373 | 15 | 34314041 | T/C | 0.61 | 0.65 | 6.28E-06 | 0.86(0.80-0.92) | -----+--- |
| rs190572487 | 14 | 19675351 | T/C | 0.75 | 0.73 | 6.86E-06 | 1.35(1.18-1.54) | +++++----- |
| rs3847511 | 11 | 91454034 | T/G | 0.08 | 0.06 | 7.70E-06 | 1.36(1.19-1.55) | +++++----- |
| rs165774 | 22 | 19952561 | A/G | 0.33 | 0.30 | 8.78E-06 | 1.18(1.10-1.27) | +++++----- |
| chr10_7228436_I | 10 | 7228436 | I2/D | 0.14 | 0.12 | 9.25E-06 | 1.31(1.16-1.47) | ----- |
| rs12639760 | 4 | 135958271 | A/T | 0.22 | 0.19 | 9.31E-06 | 1.21(1.11-1.31) | +++++----- |
| rs141199126 | 11 | 29761274 | T/C | 0.02 | 0.01 | 9.43E-06 | 2.05(1.49-2.83) | +++++?+?+--- |
| chr7_123745464_I | 7 | 123745464 | I2/D | 0.08 | 0.07 | 9.54E-06 | 1.32(1.17-1.49) | ----- |
| chr1_172543621_I | 1 | 172543621 | I2/D | 0.03 | 0.04 | 1.02E-05 | 0.60(0.47-0.75) | +++++----- |
| rs12799429 | 11 | 8097070 | T/C | 0.54 | 0.56 | 1.04E-05 | 0.86(0.80-0.92) | -----+--- |
| rs17077064 | 18 | 64976306 | T/C | 0.83 | 0.80 | 1.08E-05 | 1.21(1.11-1.31) | +++++----- |
| rs117018753 | 13 | 110581806 | T/C | 0.07 | 0.06 | 1.12E-05 | 1.37(1.19-1.58) | +++++----- |

CHR, chromosome; BP, basepair position; freq, frequency; OR, odds ratio; CI, confidence interval

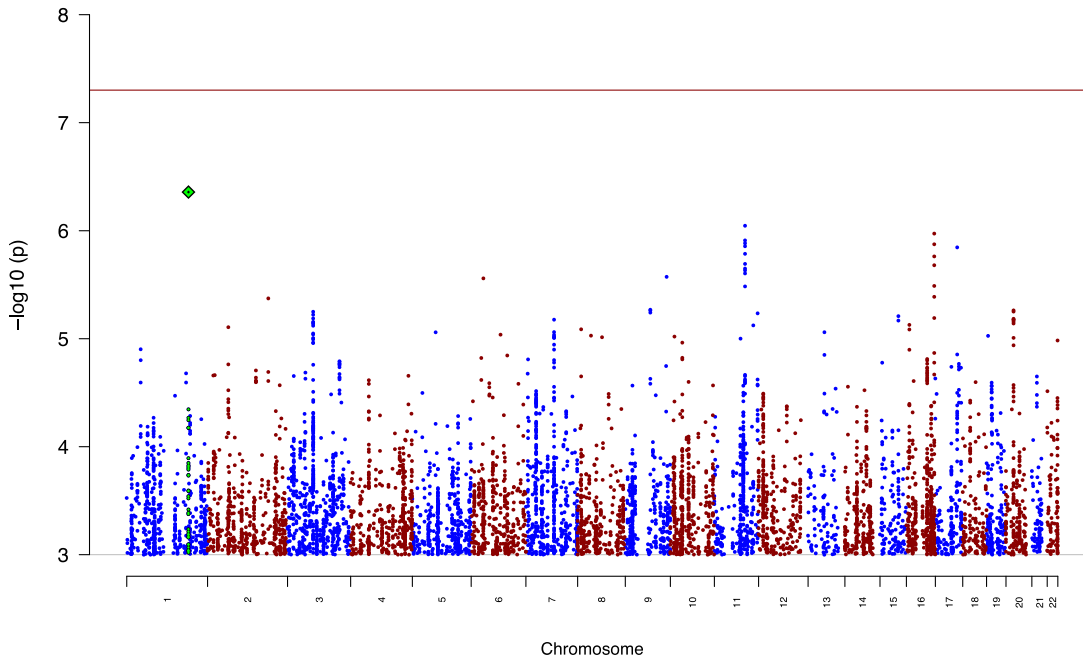


Figure S3: Manhattan plot from GWAS of suicide attempt in schizophrenia

Table S11: Top 20 results from GWAS of suicide attempt in schizophrenia showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non- attempters | P value | OR (C.I.) | Direction in each cohort |
|-------------------|-----|-----------|-------|-----------------------|----------------------------|----------|-----------------|--------------------------|
| rs482039 | 1 | 190777567 | T/C | 0.03 | 0.02 | 4.39E-07 | 2.37(1.70-3.31) | +++++ |
| rs3858375 | 11 | 95077167 | T/C | 0.07 | 0.05 | 8.99E-07 | 1.60(1.33-1.94) | +++++ |
| rs4843180 | 16 | 86753070 | T/C | 0.44 | 0.39 | 1.06E-06 | 1.25(1.14-1.36) | +++++ |
| rs180697792 | 17 | 67294430 | A/G | 0.23 | 0.20 | 1.43E-06 | 1.34(1.19-1.50) | +++++ |
| rs72756712 | 9 | 128902906 | A/G | 0.93 | 0.95 | 2.67E-06 | 0.60(0.49-0.74) | ----- |
| rs191312301 | 6 | 38975727 | A/C | 0.86 | 0.89 | 2.76E-06 | 0.71(0.62-0.82) | ----- |
| chr2_188481671_D | 2 | 188481671 | D/I3 | 0.34 | 0.38 | 4.23E-06 | 0.79(0.71-0.87) | ----- |
| rs73650494 | 9 | 78831443 | T/C | 0.96 | 0.98 | 5.39E-06 | 0.56(0.43-0.72) | ----- |
| rs6114731 | 20 | 24427180 | A/G | 0.04 | 0.03 | 5.46E-06 | 1.78(1.39-2.27) | +++++ |
| rs75305337 | 3 | 84333133 | A/C | 0.12 | 0.09 | 5.63E-06 | 1.39(1.20-1.60) | +++++ |
| chr11_133764404_I | 11 | 133764404 | I5/D | 0.79 | 0.75 | 5.82E-06 | 1.29(1.15-1.43) | ----- |
| rs57729539 | 15 | 78524199 | A/G | 0.80 | 0.83 | 6.18E-06 | 0.76(0.68-0.86) | ----- |
| rs875777 | 7 | 86745380 | T/C | 0.16 | 0.20 | 6.66E-06 | 0.76(0.67-0.86) | ----- |
| rs6497871 | 16 | 10364163 | A/G | 0.64 | 0.60 | 7.44E-06 | 1.24(1.13-1.37) | +++++ |
| rs151336980 | 11 | 120470737 | T/C | 0.04 | 0.03 | 7.52E-06 | 1.78(1.38-2.29) | +++++ |
| rs4494728 | 2 | 65589513 | T/C | 0.50 | 0.55 | 7.82E-06 | 0.82(0.75-0.89) | ----- |
| rs2739958 | 8 | 12232534 | T/C | 0.58 | 0.60 | 8.18E-06 | 0.68(0.57-0.80) | ----- |
| rs73215273 | 13 | 72403250 | A/C | 0.88 | 0.90 | 8.71E-06 | 0.66(0.54-0.79) | ----- |
| rs11739808 | 5 | 72745041 | A/G | 0.03 | 0.02 | 8.72E-06 | 2.30(1.59-3.31) | +++++ |
| rs13198361 | 6 | 91673413 | T/C | 0.11 | 0.14 | 9.17E-06 | 0.65(0.54-0.79) | ----- |

CHR, chromosome; BP, basepair position; freq, frequency; OR, odds ratio; CI, confidence interval

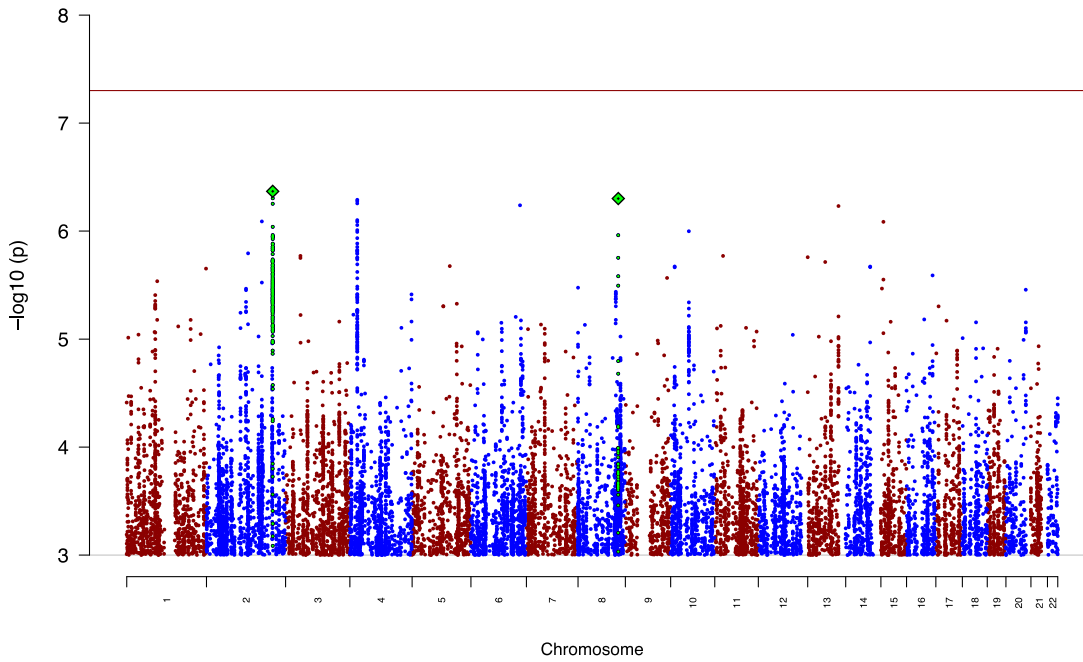


Figure S4: Manhattan plot from meta-analysis of suicide attempt in major depressive disorder, bipolar disorder and schizophrenia

Table S12: Top 20 results from meta-analysis of suicide attempt in MDD, BIP and SCZ showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non-attempters | P value | OR (C.I.) | Direction in each cohort |
|------------------|-----|-----------|-------|--------------------|------------------------|----------|-----------------|--------------------------|
| rs149268645 | 2 | 203833018 | A/G | 0.14 | 0.15 | 4.28E-07 | 0.85(0.80-0.90) | --- |
| rs4870888 | 8 | 125108977 | T/C | 0.52 | 0.54 | 5.00E-07 | 0.89(0.86-0.93) | --- |
| chr4_23273116_D | 4 | 23273116 | D/I | 0.19 | 0.18 | 5.13E-07 | 1.16(1.09-1.23) | +++ |
| rs141252918 | 6 | 151828058 | A/G | 0.02 | 0.01 | 5.76E-07 | 1.73(1.39-2.14) | +++ |
| rs9577511 | 13 | 113991823 | A/G | 0.86 | 0.87 | 5.86E-07 | 0.83(0.77-0.89) | --- |
| rs62173322 | 2 | 170611029 | A/G | 0.86 | 0.87 | 8.13E-07 | 0.84(0.78-0.90) | --- |
| rs76371172 | 15 | 31814455 | T/G | 0.98 | 0.98 | 8.20E-07 | 0.61(0.50-0.74) | --- |
| rs11004733 | 10 | 56849344 | T/C | 0.04 | 0.04 | 1.00E-06 | 1.34(1.19-1.50) | +++ |
| rs138689899 | 2 | 128288162 | T/C | 0.02 | 0.02 | 1.61E-06 | 1.53(1.29-1.82) | ++- |
| rs142055939 | 3 | 45995554 | T/C | 0.02 | 0.02 | 1.69E-06 | 1.48(1.26-1.75) | +++ |
| rs35107435 | 11 | 27249330 | A/T | 0.16 | 0.15 | 1.70E-06 | 1.17(1.10-1.24) | +++ |
| rs113988902 | 13 | 19525105 | T/C | 0.05 | 0.04 | 1.74E-06 | 1.52(1.28-1.80) | ?++ |
| chr13_73243177_I | 13 | 73243177 | I/D | 0.04 | 0.03 | 1.93E-06 | 1.59(1.32-1.93) | ?-- |
| rs186672572 | 5 | 116878032 | T/C | 0.02 | 0.01 | 2.11E-06 | 1.64(1.33-2.00) | +++ |
| rs113386487 | 10 | 13358583 | A/T | 0.98 | 0.98 | 2.12E-06 | 0.67(0.56-0.79) | --- |
| rs73348245 | 14 | 96158072 | A/G | 0.06 | 0.06 | 2.13E-06 | 1.27(1.15-1.40) | +++ |
| rs6426297 | 1 | 246538381 | T/C | 0.02 | 0.01 | 2.22E-06 | 1.65(1.34-2.04) | +++ |
| rs73577700 | 16 | 80280761 | A/T | 0.83 | 0.85 | 2.57E-06 | 0.87(0.82-0.92) | --- |
| rs72756712 | 9 | 128902906 | A/G | 0.94 | 0.95 | 2.71E-06 | 0.78(0.70-0.86) | --- |
| rs75633108 | 1 | 96735185 | T/C | 0.02 | 0.02 | 2.91E-06 | 1.59(1.31-1.93) | +++ |

MDD, major depressive disorder; BIP, bipolar disorder; SCZ, schizophrenia; CHR, chromosome; BP, basepair position; OR, odds ratio; CI, confidence interval

Table S13: Top 20 results from meta-analysis of suicide attempt in mood disorders showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non-attempters | P value | OR (C.I.) | Direction in each cohort |
|------------------|-----|-----------|-------|-----------------------|---------------------------|-----------------|-----------------|--------------------------|
| rs138689899 | 2 | 128288162 | T/C | 0.02 | 0.01 | 2.50E-08 | 1.75(1.44-2.14) | ++ |
| rs28591567 | 4 | 23253912 | A/G | 0.78 | 0.80 | 3.11E-08 | 0.84(0.79-0.89) | -- |
| chr6_151835609_D | 6 | 151835609 | I/D | 0.94 | 0.95 | 3.66E-07 | 0.74(0.66-0.83) | ++ |
| chr13_61834504_D | 13 | 61834504 | I/D | 0.91 | 0.92 | 5.77E-07 | 0.78(0.71-0.86) | ++ |
| rs186672572 | 5 | 116878032 | T/C | 0.02 | 0.01 | 7.19E-07 | 1.78(1.42-2.24) | ++ |
| rs112944737 | 8 | 134677667 | T/C | 0.91 | 0.92 | 1.11E-06 | 0.80(0.73-0.87) | -- |
| rs9577511 | 13 | 113991823 | A/G | 0.86 | 0.87 | 1.35E-06 | 0.81(0.75-0.88) | -- |
| rs150795632 | 6 | 37439376 | A/G | 0.98 | 0.98 | 1.69E-06 | 0.59(0.48-0.73) | -- |
| rs113051785 | 11 | 20167807 | C/G | 0.02 | 0.01 | 1.70E-06 | 1.75(1.39-2.20) | ++ |
| rs1355048 | 1 | 90830490 | T/C | 0.37 | 0.40 | 1.91E-06 | 0.88(0.84-0.93) | -- |
| rs115833694 | 4 | 99918226 | C/G | 0.98 | 0.98 | 1.99E-06 | 0.63(0.52-0.76) | -- |
| rs17764923 | 6 | 159820779 | A/G | 0.16 | 0.14 | 2.04E-06 | 1.19(1.11-1.27) | ++ |
| rs117020391 | 12 | 107099751 | T/C | 0.98 | 0.98 | 2.09E-06 | 0.63(0.52-0.76) | -- |
| rs72832403 | 2 | 115551269 | A/G | 0.92 | 0.93 | 2.16E-06 | 0.79(0.72-0.87) | -- |
| rs150320200 | 4 | 23450330 | A/G | 0.97 | 0.98 | 2.19E-06 | 0.66(0.55-0.78) | -- |
| rs114598476 | 1 | 224438315 | A/G | 0.96 | 0.97 | 2.30E-06 | 0.67(0.57-0.79) | -- |
| rs76400344 | 22 | 26697120 | A/C | 0.03 | 0.02 | 3.01E-06 | 1.52(1.28-1.82) | ++ |
| chr2_124851208_D | 2 | 124851208 | I/D | 0.96 | 0.96 | 3.48E-06 | 0.72(0.62-0.82) | ++ |
| rs143457262 | 7 | 82072544 | A/G | 0.03 | 0.03 | 4.16E-06 | 1.51(1.26-1.79) | ++ |
| rs3829881 | 1 | 117531813 | T/C | 0.49 | 0.47 | 4.45E-06 | 1.12(1.07-1.18) | ++ |

CHR, chromosome; BP, basepair position; freq, frequency; OR, odds ratio; CI, confidence interval

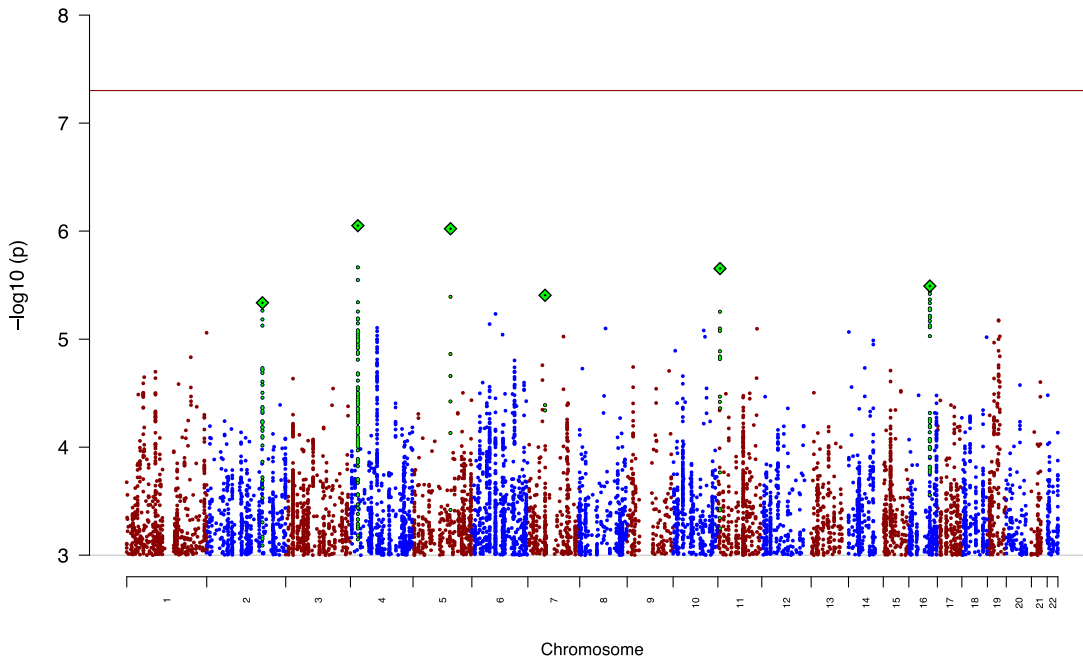


Figure S5: Manhattan plot from meta-analysis of suicide attempt in bipolar disorder and schizophrenia

Table S14: Top 20 results from meta-analysis of suicide attempt in BIP and SCZ showing the most significant SNP from each genomic region

| Variant | CHR | BP | A1/A2 | A1 freq attempters | A1 freq non-attempters | P value | OR (C.I.) | Direction in each cohort |
|-----------------|-----|-----------|-------|-----------------------|---------------------------|----------|-----------------|--------------------------|
| chr4_23273116_D | 4 | 23273116 | D/I | 0.19 | 0.17 | 8.90E-07 | 1.19(1.11-1.27) | ++ |
| rs26318 | 5 | 115687905 | T/C | 0.98 | 0.99 | 9.50E-07 | 0.52(0.40-0.67) | -- |
| rs118102650 | 11 | 9099847 | A/G | 0.02 | 0.01 | 2.22E-06 | 1.79(1.41-2.28) | ++ |
| rs12925656 | 16 | 65107920 | T/C | 0.88 | 0.90 | 3.22E-06 | 0.82(0.76-0.89) | -- |
| rs73122740 | 7 | 53695512 | T/G | 0.92 | 0.91 | 3.93E-06 | 1.26(1.14-1.39) | ++ |
| rs2353181 | 2 | 170305256 | T/C | 0.95 | 0.94 | 4.61E-06 | 1.33(1.17-1.49) | ++ |
| rs75237141 | 6 | 72966596 | A/G | 0.90 | 0.92 | 5.84E-06 | 0.80(0.73-0.88) | -- |
| rs56342621 | 19 | 35071781 | C/G | 0.98 | 0.99 | 6.67E-06 | 0.61(0.49-0.75) | -- |
| rs9475195 | 6 | 55061018 | T/C | 0.58 | 0.61 | 7.25E-06 | 0.89(0.84-0.93) | -- |
| chr4_82106147_I | 4 | 82106147 | I/D | 0.07 | 0.06 | 7.85E-06 | 1.29(1.15-1.44) | -- |
| chr8_80591790_D | 8 | 80591790 | I/D | 0.61 | 0.63 | 7.95E-06 | 0.88(0.83-0.93) | ++ |
| rs10892827 | 11 | 122210990 | T/G | 0.90 | 0.92 | 8.01E-06 | 0.80(0.73-0.88) | -- |
| rs189924441 | 10 | 94182243 | A/G | 0.97 | 0.97 | 8.30E-06 | 0.67(0.56-0.80) | -- |
| rs8022689 | 14 | 21608986 | A/G | 0.38 | 0.35 | 8.58E-06 | 1.14(1.07-1.20) | ++ |
| rs3007305 | 1 | 246862572 | C/G | 0.66 | 0.68 | 8.72E-06 | 0.88(0.83-0.93) | -- |
| rs72924216 | 6 | 94570858 | A/G | 0.95 | 0.96 | 9.08E-06 | 0.72(0.63-0.83) | -- |
| rs66666015 | 19 | 38717143 | T/C | 0.76 | 0.74 | 9.42E-06 | 1.16(1.09-1.24) | ++ |
| rs38758 | 7 | 109943767 | A/C | 0.47 | 0.44 | 9.46E-06 | 1.12(1.07-1.18) | ++ |
| rs117559494 | 10 | 98008526 | A/G | 0.03 | 0.03 | 9.50E-06 | 1.53(1.27-1.85) | ++ |
| rs117637007 | 18 | 75827605 | T/C | 0.11 | 0.13 | 9.58E-06 | 0.80(0.72-0.88) | -- |

CHR, chromosome; BP, basepair position; freq, frequency; OR, odds ratio; CI, confidence interval

Table S15: Genome-wide significant loci for suicide attempt tested in independent replication cohorts

| Cohort | Variant | CHR | A1/A2 | P value | OR (C.I.) |
|------------|--------------|-----|-------|---------|------------------|
| UK Biobank | rs45593736 | 10 | A/G | 0.985 | 1.00 (0.83-1.21) |
| UK Biobank | rs138689899 | 2 | T/C | 0.677 | 0.96 (0.79-1.16) |
| UK Biobank | rs4626184^ | 4 | C/A | 0.663 | 1.04 (0.86-1.26) |
| iPSYCH | rs117607218* | 10 | C/T | 0.776 | 0.97 (0.79-1.18) |
| iPSYCH | rs138689899 | 2 | T/C | 0.228 | 0.87 (0.69-1.09) |
| iPSYCH | rs28591567 | 4 | G/A | 0.521 | 1.02 (0.96-1.07) |

^rs28591567 was not present in the UK Biobank dataset, so the SNP in highest LD ($R^2 = 0.88$) was chosen.

*rs45593736 was not present in the iPSYCH dataset, so the SNP in highest LD ($R^2 = 0.60$) was chosen.

CHR, chromosome; OR, odds ratio; CI, confidence interval

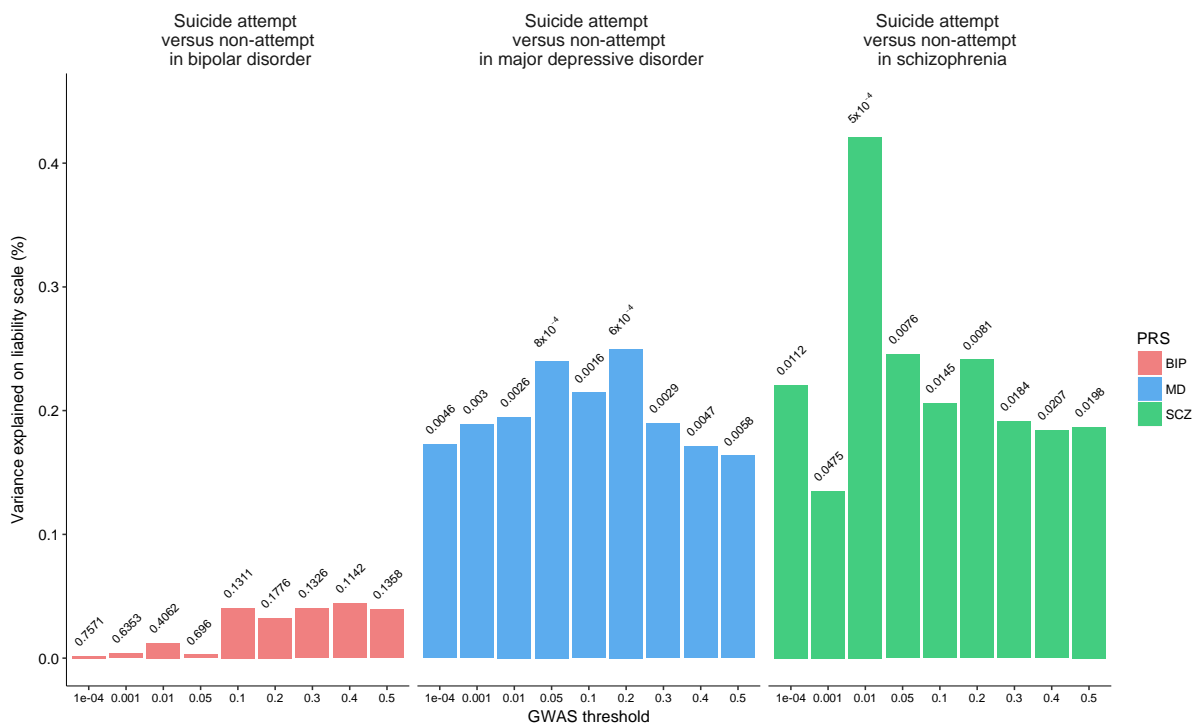
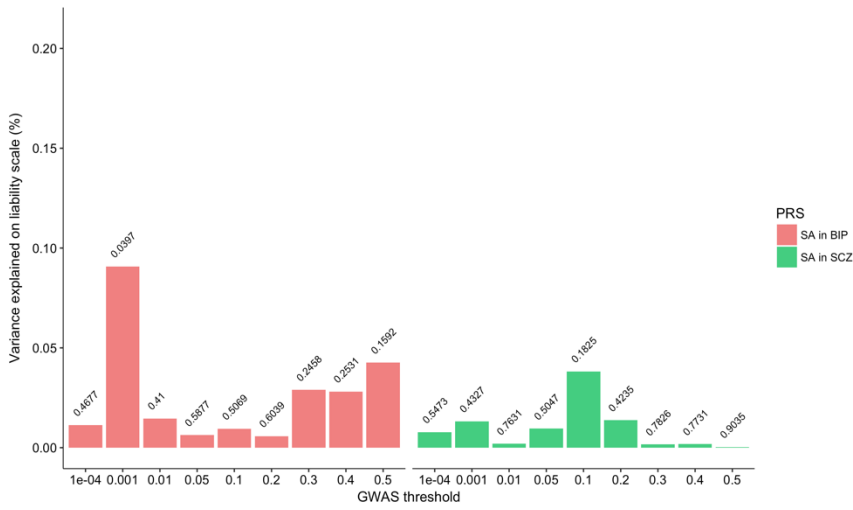


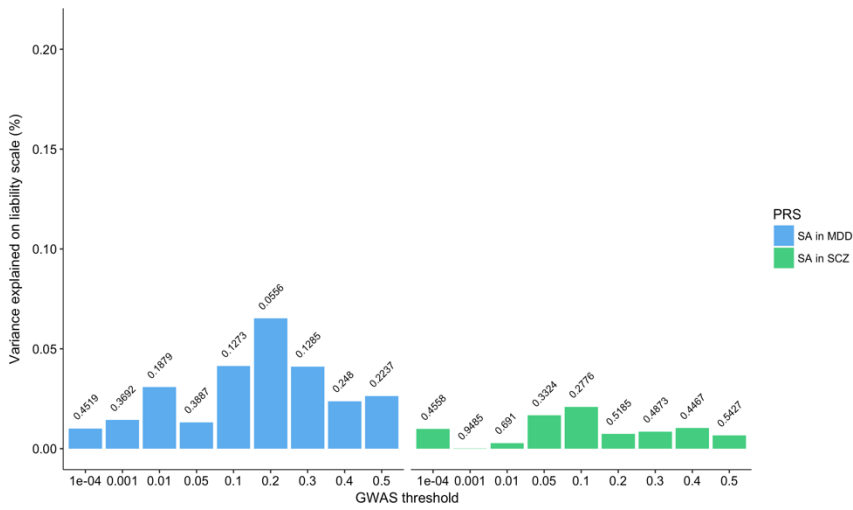
Figure S6: Polygenic risk scores for bipolar disorder, major depression and schizophrenia tested for association with suicide attempt in the same disorder.

PRS-polygenic risk score, BIP-bipolar disorder, MD-major depression, SCZ-schizophrenia. P values of association between polygenic scores and suicide attempt are shown above each bar.

A. Suicide attempt versus non-attempt in major depressive disorder



B. Suicide attempt versus non-attempt in bipolar disorder



C. Suicide attempt versus non-attempt in schizophrenia

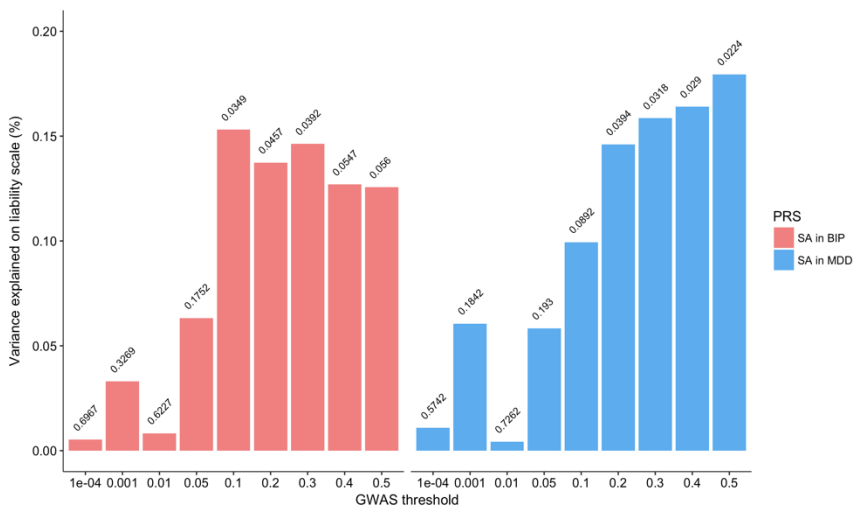


Figure S7: Polygenic risk scores for suicide attempt tested for association with suicide attempt in A - major depressive disorder, B - bipolar disorder and C - schizophrenia. PRS-polygenic risk score, SA-suicide attempt, MDD-major depressive disorder, BIP-bipolar disorder, SCZ-schizophrenia. P values of association between polygenic scores and suicide attempt are shown above each bar.

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Psychiatric Genomics Consortium Major Depressive Disorder Cohorts

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The following table lists the funding that supported the primary studies analyzed.

| Study | Lead investigator | Award number | Funder | Country |
|-------------------|----------------------------------|--|--|-------------|
| PGC | PF Sullivan | U01 MH109528 | NIMH | USA |
| PGC | A Agrawal | U01 MH109532 | NIDA | USA |
| PGC | D Posthuma | 480-05-003 | Netherlands Scientific Organization | Netherlands |
| PGC | D Posthuma | – | Dutch Brain Foundation and the VU University Amsterdam | Netherlands |
| PsyColaus | M Preisig | 3200B0–105993, 3200B0-118308, 33CSO-122661, 33CS30-139468, 33CS30-148401 | Swiss National Science Foundation | Switzerland |
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| QIMR | AC Heath | AA07535, AA07728, and AA10249 | NIAAA | USA |
| RADIANT | C Lewis, G Breen | G0701420 | MRC | UK |
| RADIANT | G Breen | G0901245 | MRC | UK |
| RADIANT | G Breen | U01 MH109528 | NIMH | UK |
| BoMa | M Rietschel | RI 908/11-1 | Deutsche Forschungsgemeinschaft | Germany |
| BoMa | MM Nöthen | NO246/10-1 | Deutsche Forschungsgemeinschaft | Germany |
| BoMa | MM Nöthen | Excellence Cluster ImmunoSensation | Deutsche Forschungsgemeinschaft | Germany |
| BoMa | MM Nöthen, M Rietschel, S Cichon | 01ZX1314A/01ZX1614A, 01ZX1314G/01ZX1614G, | BMBF Integument | Germany |
| BoMa | MM Nöthen, M Rietschel, S Cichon | 01GS08144, 01GS08147 | BMBF NGFNplus MoodS | Germany |
| CoFaMS - Adelaide | BT Baune | APP1060524 | NHMRC | Australia |
| NESDA | BWJH Penninx | ZonMW Geestkracht grant | N.W.O. | Netherlands |
| NTR | DI Boomsma | 480-15-001/674 | N.W.O. | Netherlands |
| SHIP-LEGEND/TREND | HJ Grabe | DFG: GR 1912/5-1 | German Research Foundation | Germany |
| STAR*D | SP Hamilton | R01 MH-072802 | NIMH | USA |

Psychiatric Genomics Consortium Bipolar Disorder Cohorts

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The following table lists the funding that supported the primary studies analyzed.

| Study | Lead investigator | Country, Funder, Award number |
|-------------------------------------|--------------------------|--|
| PGC | P Sullivan | USA, NIMH MH109528 |
| PGC | D Posthuma | Netherlands, Scientific Organization Netherlands, 480-05-003 |
| PGC | D Posthuma | Dutch Brain Foundation and the VU University Amsterdam Netherlands |
| BiGS, Uchicago | ES Gershon | R01 MH103368 |
| BiGS, GAIN | FJ McMahon | US, NIMH, R01 MH061613, ZIA MH002843 |
| BiGS, UCSD | J Kelsoe | US, NIMH, MH078151, MH081804, MH59567 |
| BiGS, University of Pittsburgh | V Nimgaonkar | US, NIMH MH63480 |
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| BOMA-Romania | M Grigoriou-Serbanescu | Romania, UEFISCDI, Grant no. 89/2012 |
| BOMA-Germany I, II, III | S Cichon | Germany, BMBF Integument, 01ZX1314A/01ZX1614A |
| BOMA-Germany I, II, III | S Cichon | Germany, BMBF NGFNplus MoodS, 01GS08144 |
| BOMA-Germany I, II, III | S Cichon | Switzerland, SNSF, 156791 |
| BOMA-Germany I, II, III | MM Nöthen | Germany, BMBF Integument, 01ZX1314A/01ZX1614A |
| BOMA-Germany I, II, III | MM Nöthen | Germany, BMBF NGFNplus MoodS, 01GS08144 |
| BOMA-Germany I, II, III | MM Nöthen | Germany, Deutsche Forschungsgemeinschaft, Excellence Cluster ImmunoSensation |
| BOMA-Germany I, II, III | MM Nöthen | Germany, Deutsche Forschungsgemeinschaft, NO246/10-1 |
| BOMA-Germany I, II, III | SH Witt | Germany, Deutsche Forschungsgemeinschaft, WI 3429/3-1 |
| BOMA-Germany I, II, III, BOMA-Spain | M Rietschel | Germany, BMBF Integument, 01ZX1314G/01ZX1614G |
| BOMA-Germany I, II, III, BOMA-Spain | M Rietschel | Germany, BMBF NGFNplus MoodS, 01GS08147 |

| | | |
|--|----------------------|--|
| BOMA-Germany I, II, III, BOMA-Spain | M Rietschel | Germany, Deutsche Forschungsgemeinschaft, RI 908/11-1 |
| BOMA-Germany I, II, III, PsyCourse, BiGS | TG Schulze | Germany, BMBF Integument, 01ZX1314K |
| BOMA-Germany I, II, III, PsyCourse, BiGS | TG Schulze | Germany, DFG, SCHU 1603/4-1, SCHU 1603/5-1, SCHU 1603/7-1 |
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Psychiatric Genomics Consortium Schizophrenia cohorts

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Replication cohorts

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