

## Supplementary Materials

**Supplementary Table 1. eQTL analysis for rs9831894 and rs6773363**

| GWAS SNP  | SNPs in LD  | #Chr | Position  | eQTL Cluster (GTEx)                      |
|-----------|-------------|------|-----------|--|
| rs9831894 | rs9831894   | 3    | 121800487 | CD86;testis,                             |
| rs9831894 | rs9828869   | 3    | 121748360 | CD86;testis,                             |
| rs9831894 | rs1962532   | 3    | 121751531 | 0  |
| rs9831894 | rs527769191 | 3    | 121759584 | 0  |
| rs9831894 | rs62269210  | 3    | 121760603 | 0  |
| rs9831894 | rs62269211  | 3    | 121760639 | 0  |
| rs9831894 | rs28876421  | 3    | 121764257 | 0  |
| rs9831894 | rs2255214   | 3    | 121770539 | 0  |
| rs9831894 | rs17203397  | 3    | 121781426 | 0  |
| rs9831894 | rs6786977   | 3    | 121782806 | 0  |
| rs9831894 | rs2681411   | 3    | 121785631 | 0  |
| rs9831894 | rs4308217   | 3    | 121793187 | 0  |
| rs9831894 | rs35187120  | 3    | 121796389 | 0  |
| rs9831894 | rs11717893  | 3    | 121806825 | 0  |
| rs9831894 | rs9836399   | 3    | 121811811 | CD86;testis,                             |
| rs9831894 | rs9289187   | 3    | 121815461 | CD86;testis,                             |
| rs9831894 | rs2681416   | 3    | 121817613 | CD86;testis,                             |
| rs9831894 | rs2332096   | 3    | 121821143 | CD86;testis,                             |
| rs9831894 | rs1915092   | 3    | 121823375 | 0  |
| rs9831894 | rs2262495   | 3    | 121841801 | 0  |
| rs9831894 | rs2681401   | 3    | 121843127 | 0  |
| rs9831894 | rs2715272   | 3    | 121843241 | 0  |
| rs6773363 | rs6773363   | 3    | 27793632  | 0  |
| rs6773363 | rs2371108   | 3    | 27757018  | RP11-222K16.2;Esophagus -<br>Muscularis, |
| rs6773363 | rs530939637 | 3    | 27757867  | 0  |
| rs6773363 | rs2371109   | 3    | 27758274  | 0  |
| rs6773363 | rs2887944   | 3    | 27758275  | 0  |
| rs6773363 | rs9866625   | 3    | 27759338  | 0  |
| rs6773363 | rs56306375  | 3    | 27760524  | 0  |
| rs6773363 | rs368178421 | 3    | 27763427  | 0  |
| rs6773363 | rs3806624   | 3    | 27764623  | 0  |
| rs6773363 | rs3840481   | 3    | 27765595  | 0  |
| rs6773363 | rs34269949  | 3    | 27769551  | 0  |
| rs6773363 | rs1353286   | 3    | 27772014  | 0  |
| rs6773363 | rs5847475   | 3    | 27772346  | 0  |
| rs6773363 | rs549698945 | 3    | 27774208  | 0  |
| rs6773363 | rs6801231   | 3    | 27774245  | 0  |
| rs6773363 | rs34860778  | 3    | 27776242  | 0  |
| rs6773363 | rs9880772   | 3    | 27777779  | 0  |
| rs6773363 | rs12635205  | 3    | 27779362  | 0  |
| rs6773363 | rs34024889  | 3    | 27780638  | 0  |
| rs6773363 | rs13327021  | 3    | 27783015  | 0  |
| rs6773363 | rs4680838   | 3    | 27783476  | 0  |
| rs6773363 | rs9310852   | 3    | 27784997  | 0  |
| rs6773363 | rs145548114 | 3    | 27786022  | 0  |
| rs6773363 | rs11129295  | 3    | 27788780  | 0  |
| rs6773363 | rs12330493  | 3    | 27791614  | 0  |

|           |             |   |          |   |
|-----------|-------------|---|----------|---|
| rs6773363 | rs6806966   | 3 | 27793862 | 0 |
| rs6773363 | rs13093489  | 3 | 27794907 | 0 |
| rs6773363 | rs7371953   | 3 | 27795157 | 0 |
| rs6773363 | rs12497690  | 3 | 27795397 | 0 |
| rs6773363 | rs113106250 | 3 | 27797023 | 0 |
| rs6773363 | rs199900263 | 3 | 27797030 | 0 |
| rs6773363 | rs12493245  | 3 | 27797810 | 0 |
| rs6773363 | rs4680843   | 3 | 27798071 | 0 |
| rs6773363 | rs4680844   | 3 | 27798172 | 0 |
| rs6773363 | rs9860067   | 3 | 27798275 | 0 |
| rs6773363 | rs4680845   | 3 | 27799034 | 0 |
| rs6773363 | rs17020283  | 3 | 27799105 | 0 |
| rs6773363 | rs572217447 | 3 | 27799483 | 0 |
| rs6773363 | rs572217447 | 3 | 27799483 | 0 |
| rs6773363 | rs9862284   | 3 | 27800325 | 0 |
| rs6773363 | rs6795177   | 3 | 27800734 | 0 |
| rs6773363 | rs17020287  | 3 | 27801879 | 0 |
| rs6773363 | rs60047326  | 3 | 27801932 | 0 |
| rs6773363 | rs17020288  | 3 | 27801959 | 0 |
| rs6773363 | rs9814971   | 3 | 27802163 | 0 |
| rs6773363 | rs9872851   | 3 | 27802302 | 0 |
| rs6773363 | rs9815263   | 3 | 27802364 | 0 |
| rs6773363 | rs9815419   | 3 | 27802488 | 0 |
| rs6773363 | rs56143766  | 3 | 27802504 | 0 |
| rs6773363 | rs9816235   | 3 | 27803017 | 0 |
| rs6773363 | rs4680730   | 3 | 27804731 | 0 |
| rs6773363 | rs4680851   | 3 | 27805177 | 0 |
| rs6773363 | rs4680852   | 3 | 27805204 | 0 |
| rs6773363 | rs5847478   | 3 | 27805726 | 0 |
| rs6773363 | rs2371150   | 3 | 27806147 | 0 |
| rs6773363 | rs4450763   | 3 | 27806852 | 0 |
| rs6773363 | rs4522733   | 3 | 27807006 | 0 |
| rs6773363 | rs4371473   | 3 | 27807063 | 0 |
| rs6773363 | rs12497514  | 3 | 27807665 | 0 |
| rs6773363 | rs55998928  | 3 | 27807730 | 0 |
| rs6773363 | rs11711117  | 3 | 27809342 | 0 |
| rs6773363 | rs200433332 | 3 | 27809401 | 0 |
| rs6773363 | rs1000361   | 3 | 27810319 | 0 |
| rs6773363 | rs9868208   | 3 | 27810888 | 0 |
| rs6773363 | rs5012869   | 3 | 27811220 | 0 |
| rs6773363 | rs62253101  | 3 | 27811637 | 0 |
| rs6773363 | rs9878524   | 3 | 27812457 | 0 |
| rs6773363 | rs9878720   | 3 | 27812594 | 0 |
| rs6773363 | rs9823274   | 3 | 27812666 | 0 |
| rs6773363 | rs4680861   | 3 | 27812802 | 0 |
| rs6773363 | rs4279053   | 3 | 27813319 | 0 |
| rs6773363 | rs11915329  | 3 | 27814628 | 0 |
| rs6773363 | rs62253104  | 3 | 27815285 | 0 |
| rs6773363 | rs201123277 | 3 | 27815370 | 0 |
| rs6773363 | rs6551228   | 3 | 27815535 | 0 |
| rs6773363 | rs10049011  | 3 | 27815942 | 0 |
| rs6773363 | rs13083047  | 3 | 27817959 | 0 |
| rs6773363 | rs9872426   | 3 | 27818453 | 0 |
| rs6773363 | rs7628641   | 3 | 27818887 | 0 |

|           |             |   |          |                               |
|-----------|-------------|---|----------|-------------------------------|
| rs6773363 | rs7628648   | 3 | 27818893 | 0                             |
| rs6773363 | rs7621606   | 3 | 27819783 | 0                             |
| rs6773363 | rs12633504  | 3 | 27819955 | 0                             |
| rs6773363 | rs9838021   | 3 | 27820995 | 0                             |
| rs6773363 | rs60294567  | 3 | 27821359 | 0                             |
| rs6773363 | rs7646728   | 3 | 27822171 | 0                             |
| rs6773363 | rs7647094   | 3 | 27822530 | 0                             |
| rs6773363 | rs545045422 | 3 | 27823712 | 0                             |
| rs6773363 | rs545045422 | 3 | 27823712 | 0                             |
| rs6773363 | rs7627710   | 3 | 27823887 | 0                             |
| rs6773363 | rs11710954  | 3 | 27824022 | 0                             |
| rs6773363 | rs7653004   | 3 | 27824089 | 0                             |
| rs6773363 | rs11711718  | 3 | 27824134 | 0                             |
| rs6773363 | rs57583032  | 3 | 27824709 | 0                             |
| rs6773363 | rs11129297  | 3 | 27825386 | 0                             |
| rs6773363 | rs7633952   | 3 | 27825912 | 0                             |
| rs6773363 | rs7626965   | 3 | 27826752 | 0                             |
| rs6773363 | rs4491854   | 3 | 27826932 | 0                             |
| rs6773363 | rs34089683  | 3 | 27827398 | 0                             |
| rs6773363 | rs9817112   | 3 | 27827733 | 0                             |
| rs6773363 | rs4680868   | 3 | 27829161 | RP11-222K16.1;nerve - Tibial, |
| rs6773363 | rs11925920  | 3 | 27829393 | 0                             |
| rs6773363 | rs561194513 | 3 | 27829950 | 0                             |
| rs6773363 | rs11712606  | 3 | 27830379 | 0                             |
| rs6773363 | rs543125886 | 3 | 27830535 | 0                             |
| rs6773363 | rs556526857 | 3 | 27830865 | 0                             |
| rs6773363 | rs60579266  | 3 | 27831914 | 0                             |
| rs6773363 | rs6551229   | 3 | 27831978 | 0                             |

eQTL, expression quantitative trait loci; GTEX, Genotype-Tissue Expression

**Supplementary Table 2.** Summary of the participating studies and genotyping methods in discovery.

| Study    | Study name   | No. cases | No. controls | Design  | Description of the study   | Genotyping platform  | Genotype calling algorithm | Imputation Software |
|----------|--|-----------|--------------|---|--|--|----------------------------|---------------------|
| NCI-SEER | National Cancer Institute-Surveillance, Epidemiology, and End Results Interdisciplinary Case-Control Study of Non-Hodgkin Lymphoma           | 2661      | 6221         | Pooling of case-control studies, nested case-control studies from cohorts     | 8 Population-based case-control studies. 5 clinic or hospital-based or mixed case-control studies. 9 cohort studies.   | Illumina OmniExpress   | BeadStudio (GenCall)       | IMPUTE2             |
| GELA     | Treatment program of DLBCL patients from the Groupe d'Etude des Lymphomes de l'Adulte (GELA) consisting in LNH03-1B, 2B, 3B, 39B, 6B and 7B. | 549       | 525          | Cases from randomized clinical trial with population-based controls from EPIC | Patients recruited in the study that consented to DNA analysis. Controls were from the EPIC cohort   | Illumina HumanHap 610K (cases)<br>Illumina HumanHap 610K and 660W<br>BeadStudio (controls) | BeadStudio (GenCall)       | IMPUTE2             |
| MAYO     | Mayo Clinic Case-Control Study of NHL and CLL and Iowa-Mayo SPORE Molecular Epidemiology Resource  | 393       | 172          | Clinic-based case-control study and clinic-based registry                     | Consecutive patients with newly diagnosed, histologically-confirmed non-Hodgkin lymphoma (excluding HIV-infected cases) who were residents of Minnesota, Iowa or Wisconsin. Controls were selected from patients seen in the general medicine clinics at Mayo with a pre-scheduled general medical examination, frequency on age, sex, and geographic region | Illumina HumanHap 660W   | BeadStudio (GenCall)       | IMPUTE2             |

|           |   |     |     |  |  |                                 |                         |         |
|-----------|---|-----|-----|--|--|---------------------------------|-------------------------|---------|
| UCSF<br>2 | Molecular Epidemiology of<br>non-Hodgkin lymphoma | 254 | 748 | Population-based<br>case-control study | RCA/SEER Incident NHL<br>diagnosis for patients diagnosed in<br>hospitals in 6 San Francisco Bay<br>Area Counties and who were<br>residents of the Bay Area at the<br>time of diagnosis.<br>Controls ascertained through RDD<br>were frequency matched to cases<br>on age in 5-year groups, sex and<br>county of residence; Random<br>sampling of CMS lists for person<br>residing in the same 6 Bay Area<br>counties were used to supplement<br>recruitment of controls aged 65+. | Illumina<br>HumanCNV3<br>70-Duo | BeadStudio<br>(GenCall) | IMPUTE2 |
|-----------|---|-----|-----|--|--|---------------------------------|-------------------------|---------|

**Supplementary Table 3.** Summary of the replication studies conducting de novo genotyping for this study.

| Study | Study name  | Design   | Description of the study   | SNP       | Genotyping platform | Quality Control                                    | Final No. Cases | Final No. Controls |
|-------|---|--|--|-----------|---------------------|--|-----------------|--------------------|
| MAYO  | Iowa-Mayo SPORE Molecular Epidemiology Resource Mayo Clinic Case-Control Study of NHL | Clinic-based case registry and Clinic-based case-control study | See supplementary Table 1.   | rs9831894 | Sequenom            | 99% concordance on 48 duplicates                   | 782             | 1053               |
|       |   |  |  | rs6773363 | Taqman              | 100% concordance on 11 duplicates                  | 659             | 1059               |
| MDA   | MD Anderson lymphoma case-control study   | Case-control   | Cases from MD Anderson Cancer Center. Controls from Kelsey Seybold Clinics.  | rs9831894 | Taqman              | 100% concordance on 6% replicates + 2 CEPH samples | 371             | 370                |
| MSKCC | Memorial-Sloan Kettering Lymphoproliferative disorders Study                          | Hospital-based case-study and NYCP controls                    | Cases were hospital clinic based ascertainment in a tertiary referral center. NYCP controls from same geographic area. | rs9831894 | Sequenom            | 100% concordance on 24 duplicates                  | 653             | 148                |

**Supplementary Table 4.** Public epigenetic data from B cells, lymphoma cell lines and patients

| Sample                     | Mark     | No. sample | ID         | Format   | Ref. |
|----------------------------|----------|------------|------------|----------|------|
| B cells                    | CREBBP   | 1          | GSM2386721 | sra      | (1)  |
| B cells                    | DNaseI   | 1          | GSM701492  | sra      | (2)  |
| B cells                    | H3K27ac  | 1          | GSM1027287 | sra      | (2)  |
| B cells                    | H3K27ac  | 1          | GSM1519135 | sra      | (3)  |
| B cells                    | H3K27ac  | 1          | GSM1519136 | sra      | (3)  |
| B cells                    | H3K27ac  | 1          | GSM1519134 | sra      | (3)  |
| B cells                    | H3K27ac  | 1          | GSM1519137 | sra      | (3)  |
| B cells                    | H3K27ac  | 1          | GSM2386722 | sra      | (1)  |
| B cells                    | H3K27ac  | 1          | GSM2386720 | sra      | (1)  |
| B cells                    | H3K27me3 | 1          | GSM971339  | sra      | (4)  |
| B cells                    | H3K27me3 | 1          | GSM537634  | tagalign | (2)  |
| B cells                    | H3K27me3 | 1          | GSM537638  | tagalign | (2)  |
| B cells                    | H3K4me1  | 1          | GSM916053  | sra      | (2)  |
| B cells                    | H3K4me1  | 1          | GSM1648036 | sra      | (5)  |
| B cells                    | H3K4me3  | 1          | GSM537622  | tagalign | (2)  |
| B cells                    | H3K4me3  | 1          | GSM1648037 | sra      | (5)  |
| DLBCL cell line OCI-LY1    | BRD4     | 1          | GSM1296634 | sra      |      |
| DLBCL cell line OCI-Ly1    | BRD4     | 1          | GSM1133650 | sra      | (6)  |
| DLBCL cell line OCI-LY7    | DHS      | 1          | GSM2308524 | sra      | (2)  |
| DLBCL cell line OCI-LY7    | DHS      | 1          | GSM2308525 | sra      | (2)  |
| DLBCL cell line OCI-Ly1    | H3K27ac  | 1          | GSM1133646 | sra      | (6)  |
| DLBCL cell line DHL6       | H3K27ac  | 1          | GSM1254194 | sra      | (6)  |
| DLBCL cell line HBL1       | H3K27ac  | 1          | GSM1254196 | sra      | (6)  |
| DLBCL cell line OCI-LY3    | H3K27ac  | 1          | GSM1254198 | sra      | (6)  |
| DLBCL cell line OCI-LY4    | H3K27ac  | 1          | GSM1254200 | sra      | (6)  |
| DLBCL cell line TOLEDO     | H3K27ac  | 1          | GSM1254202 | sra      | (6)  |
| DLBCL cell line DOHH2      | H3K27ac  | 1          | GSM1703895 | sra      | (7)  |
| DLBCL cell line KARPAS-422 | H3K27ac  | 1          | GSM1703909 | sra      | (7)  |
| DLBCL cell line OCI-LY1    | H3K27ac  | 1          | GSM1703914 | sra      | (7)  |
| DLBCL cell line OCI-LY7    | H3K27ac  | 1          | GSM1703918 | sra      | (7)  |
| DLBCL cell line PFEIFFER   | H3K27ac  | 1          | GSM1703923 | sra      | (7)  |
| DLBCL cell line SU-DHL4    | H3K27ac  | 1          | GSM1703927 | sra      | (7)  |
| DLBCL cell line TOLEDO     | H3K27ac  | 1          | GSM1703931 | sra      | (7)  |
| DLBCL cell line OCI-LY1    | H3K27ac  | 1          | GSM2037789 | sra      | (8)  |
| DLBCL cell line Pfeiffer   | H3K27me3 | 1          | GSM1006149 | sra      | (9)  |
| DLBCL cell line KARPAS-422 | H3K27me3 | 1          | GSM1006151 | sra      | (9)  |
| DLBCL cell line WSU-DLCL2  | H3K27me3 | 1          | GSM1006153 | sra      | (9)  |
| DLBCL cell line OCI-Ly1    | H3K4me3  | 1          | GSM1572767 | sra      | (10) |
| DLBCL cell line OCI-Ly1    | H3K4me3  | 1          | GSM1133648 | sra      | (6)  |
| DLBCL cell line OCI-LY1    | MED1     | 1          | GSM1296637 | sra      |      |
| DLBCL cell line DB         | MEF2B    | 1          | GSM1703891 | sra      | (7)  |
| DLBCL cell line DB         | MEF2B    | 1          | GSM1703892 | sra      | (7)  |
| DLBCL cell line DOHH2      | MEF2B    | 1          | GSM1703898 | sra      | (7)  |
| DLBCL cell line KARPAS-422 | MEF2B    | 1          | GSM1703911 | sra      | (7)  |
| DLBCL cell line SU-DHL4    | MEF2B    | 1          | GSM1703929 | sra      | (7)  |
| DLBCL cell line SU-DHL4    | MEF2B    | 1          | GSM1703930 | sra      | (7)  |
| DLBCL cell line OCI-LY1    | RNAPII   | 1          | GSM1296635 | sra      |      |
| DLBCL cell line OCI-Ly1    | RNAPII   | 1          | GSM1133652 | sra      | (6)  |















|                           |         |   |  |        |      |
|---------------------------|---------|---|--|--------|------|
| DLBCL cell line DOHH2     | BACH2   | 1 | GSM1703893   | sra    | (7)  |
| DLBCL cell line OCI-LY7   | BACH2   | 1 | GSM1703916   | sra    | (7)  |
| DLBCL cell line OCI-LY3   | BATF    | 1 | GSM1370272   | sra    | (11) |
| DLBCL cell line OCI-LY10  | BATF    | 1 | GSM1370277   | sra    | (11) |
| DLBCL cell line OCI-LY1   | BCL6    | 1 | GSM2754202   | sra    | (12) |
| DLBCL cell line OCI-LY1   | BCL6    | 1 | GSM2754203   | sra    | (12) |
| DLBCL cell line OCI-Ly1   | BCL6    | 1 | GSM1133657   | sra    | (6)  |
| DLBCL cell line OCI-Ly1   | E2F1    | 1 | GSM1133656   | sra    | (6)  |
| DLBCL cell line OCI-LY3   | IRF4    | 1 | GSM1370274   | sra    | (11) |
| DLBCL cell line OCI-LY10  | IRF4    | 1 | GSM1370279   | sra    | (11) |
| DLBCL cell line DOHH2     | PAX5    | 1 | GSM1703899   | sra    | (7)  |
| DLBCL cell line OCI-LY7   | PAX5    | 1 | GSM1703921   | sra    | (7)  |
| DLBCL cell line DOHH2     | PU.1    | 1 | GSM1703900   | sra    | (7)  |
| DLBCL cell line OCI-LY7   | PU.1    | 1 | GSM1703922   | sra    | (7)  |
| DLBCL cell line OCI-LY3   | SPIB    | 1 | GSM1370276   | sra    | (11) |
| DLBCL cell line OCI-LY10  | SPIB    | 1 | GSM1370281   | sra    | (11) |
| DLBCL cell line OCI-Ly19  | STAT3   | 1 | GSM1227205   | sra    | (13) |
| DLBCL cell line OCI-Ly3   | STAT3   | 1 | GSM1227206   | sra    | (13) |
| DLBCL cell line OCI-Ly7   | STAT3   | 1 | GSM1227207   | sra    | (13) |
| DLBCL cell line SU-DHL10  | STAT3   | 1 | GSM1227208   | sra    | (13) |
| DLBCL cell line SU-DHL2   | STAT3   | 1 | GSM1227209   | sra    | (13) |
| DLBCL cell line SU-DHL4   | STAT3   | 1 | GSM1227210   | sra    | (13) |
| DLBCL patients            | H3K27ac | 6 | GSM1254206,GSM1254208,GSM1254210,GSM1254212,GSM1254214,GSM1254216                                  | sra    | (6)  |
| HGB patients              | H3K27ac | 7 | GSM1703933,GSM1703935,GSM1703937,GSM1703939,GSM1703941,GSM1703943,GSM1703945                       | tdf    | (7)  |
| MCL cell lines GRANTA-519 | H3K27ac | 1 | GSM1703901   | tdf    | (7)  |
| MCL cell lines JEKO-1     | H3K27ac | 1 | GSM1703903   | tdf    | (7)  |
| MCL cell lines MINO       | H3K27ac | 1 | GSM1703912   | tdf    | (7)  |
| MCL cell lines REC-1      | H3K27ac | 1 | GSM1703925   | tdf    | (7)  |
| MCL cell lines HBL2       | H3K27ac | 1 | GSM2571789   | tdf    | (14) |
| MCL cell lines JVM2       | H3K27ac | 1 | GSM2571791   | tdf    | (14) |
| MCL cell lines MAVER-1    | H3K27ac | 1 | GSM2571793   | tdf    | (14) |
| MCL cell lines SP-49      | H3K27ac | 1 | GSM2571828   | tdf    | (14) |
| MCL cell lines Z138       | H3K27ac | 1 | GSM2571832   | tdf    | (14) |
| MCL patients              | H3K27ac | 4 | GSM1703947,GSM1703949,GSM1703951,GSM1703953  | tdf    | (7)  |
| MCL patients              | H3K27ac | 3 | GSM2760324,GSM2760326,GSM2760328   | tdf    | (14) |
| SLL patients              | H3K27ac | 3 | GSM1703955,GSM1703957,GSM1703959   | tdf    | (7)  |
| FL patients               | H3K27ac | 9 | GSM1523532,GSM1523535,GSM1523544,GSM1523550,GSM1523556,GSM1523559,GSM1523565,GSM1523571,GSM1523573 | bigwig | (15) |

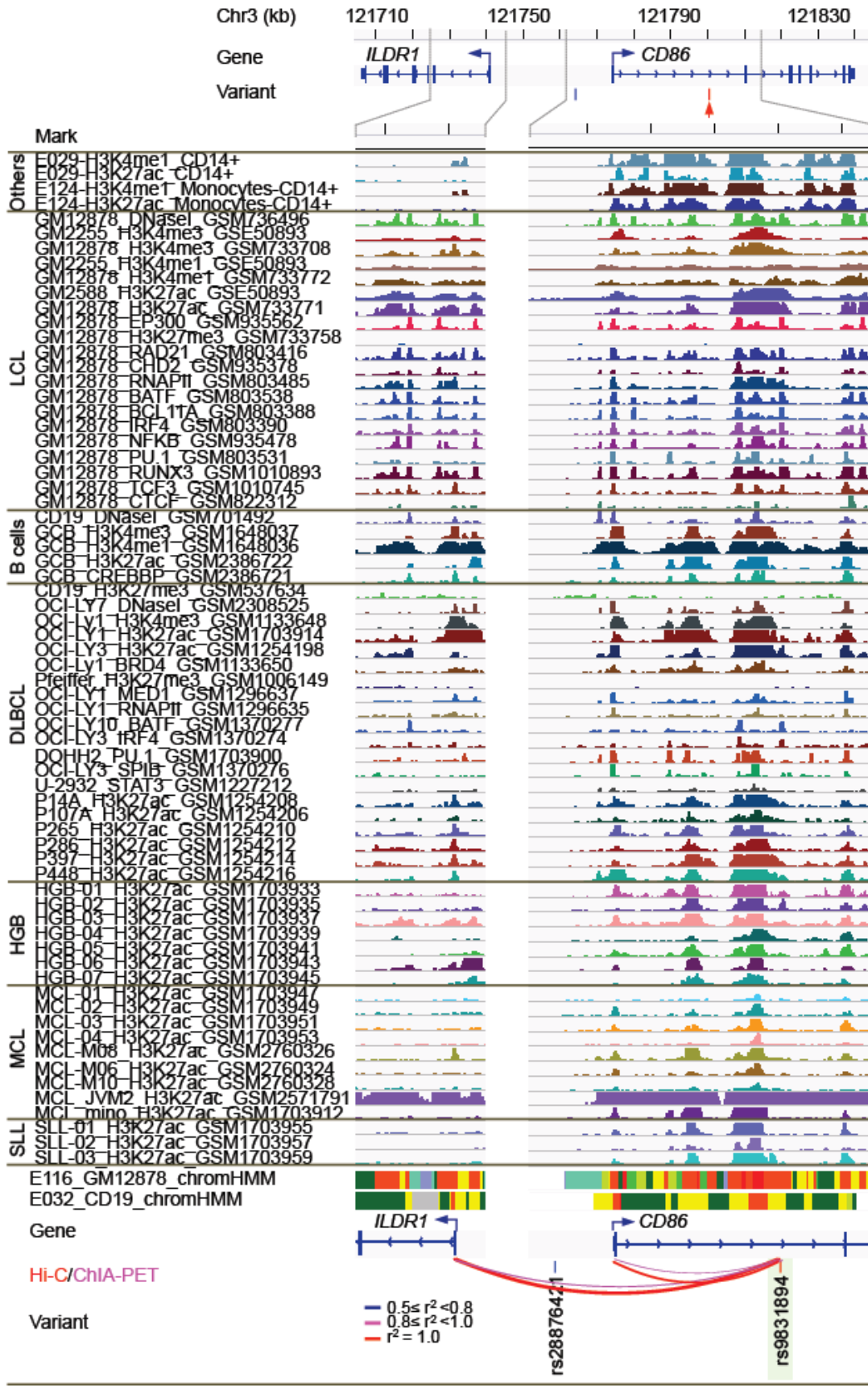
Except DNase I hypersensitive site (DHS), other marks were assayed by ChIP-seq. FL, follicular lymphoma; HGB, high-grade B-cell lymphoma; MCL, mantle cell lymphoma; SLL, small lymphocytic lymphoma.



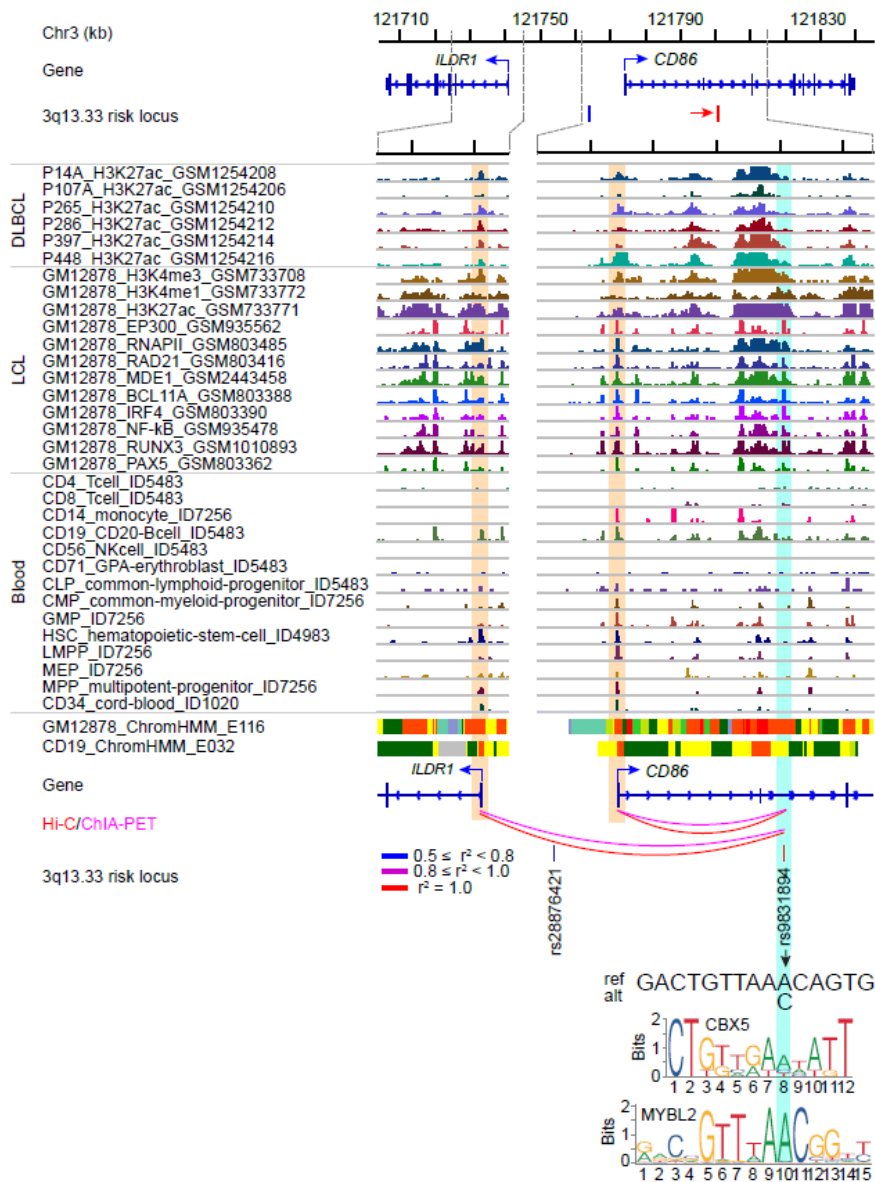
**Supplementary Figure 1. Epigenetic features of rs9831894 at independent loci associated with the risk of DLBCL.** Epigenetic features at the 3q13.33 risk locus. Tag SNP rs9831894 has a single linked variant ( $r^2 \geq 0.5$ ). Only rs9831894 was shown for its overlap with epigenetic marks, while the other one was in a region without the enrichment of the assayed epigenetic marks. In particular, The GWAS SNP rs9831894 is in a super-enhancer. The region spanning rs9831894 showed looping interactions with the promoters of *CD86* and *ILDR1*. E029, primary monocytes from peripheral blood (CD14+); E124, monocytes CD14+ RO01746 primary cells.

Chromatin state

- |   |                                   |   |                              |
|---|-----------------------------------|---|------------------------------|
|  | 1 Active TSS                      |  | 9 Heterochromatin            |
|  | 2 Flanking active TSS             |  | 10 Bivalent/poised TSS       |
|  | 3 Transcription at gene 5' and 3' |  | 11 Flanking bivalent TSS/Enh |
|  | 4 Strong transcription            |  | 12 Bivalent enhancer         |
|  | 5 Weak transcription              |  | 13 Repressed Polycomb        |
|  | 6 Genic enhancers                 |  | 14 Weak repressed Polycomb   |
|  | 7 Enhancers                       |   | 15 Quiescent/low             |
|  | 8 ZNF genes + repeats             |   |                              |

















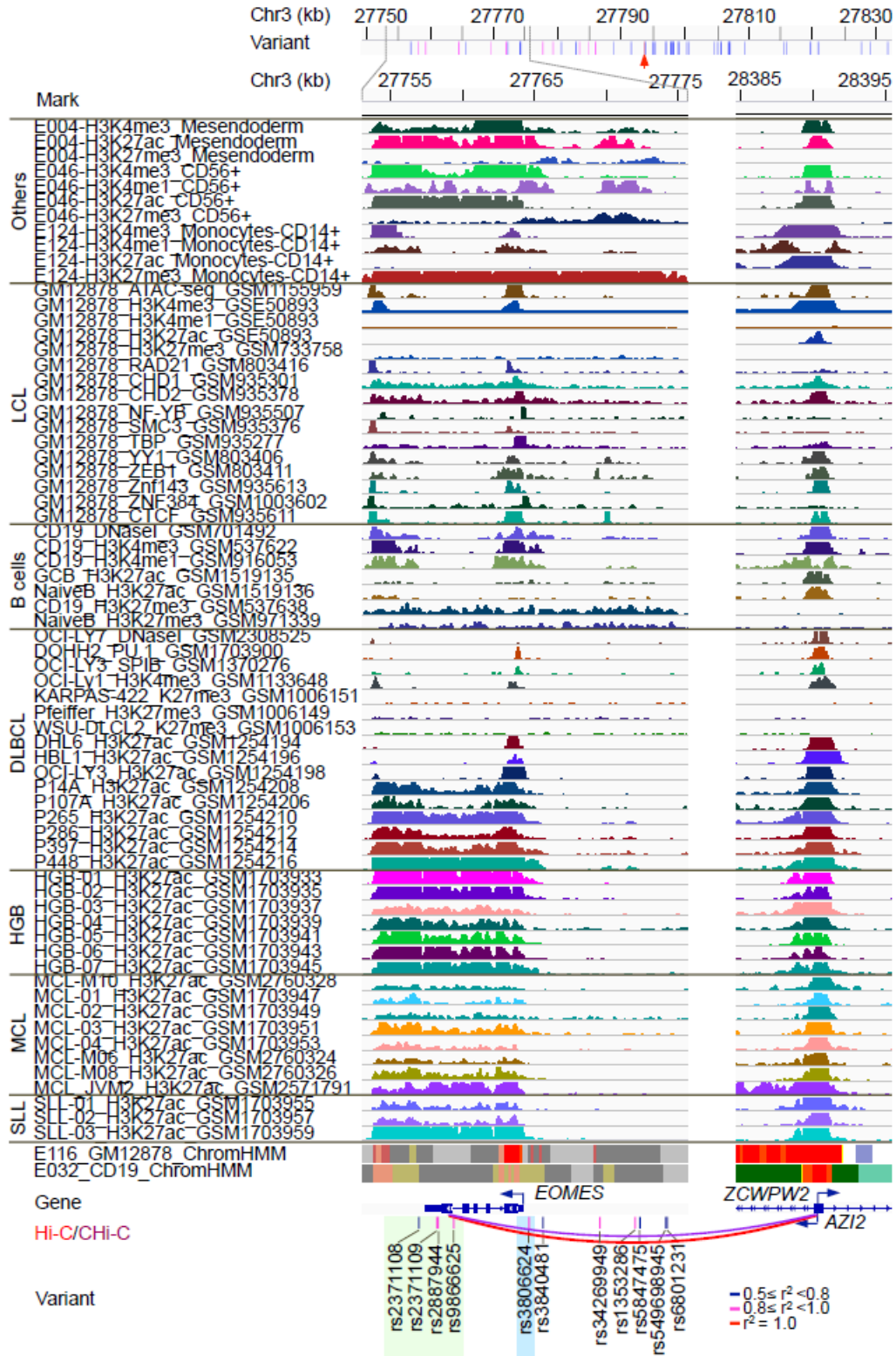
**Supplementary Figure 2. Regulatory variant at rs9831894-tagged DLBCL risk locus on 3q13.33.** The ATAC-seq data for the 14 blood cell types was downloaded from NCBI GEO under the accession GSE74912. The enhancer carrying tag SNP rs9831894 was found to be transcribed into eRNA in GM12878 (highlighted). In GM12878, this enhancer, like the *ILDR1* and *CD86* promoter it interacted with, showed the binding of RNAPII, P300, RAD21, MED1, and multiple TFs, as typically seen for transcribed enhancer. GMP, granulocyte macrophage progenitor; LMPP, lymphoid primed multipotent progenitor; MEP, megakaryocyte erythroid progenitor. See Supplementary Figure 1 legend for additional information.



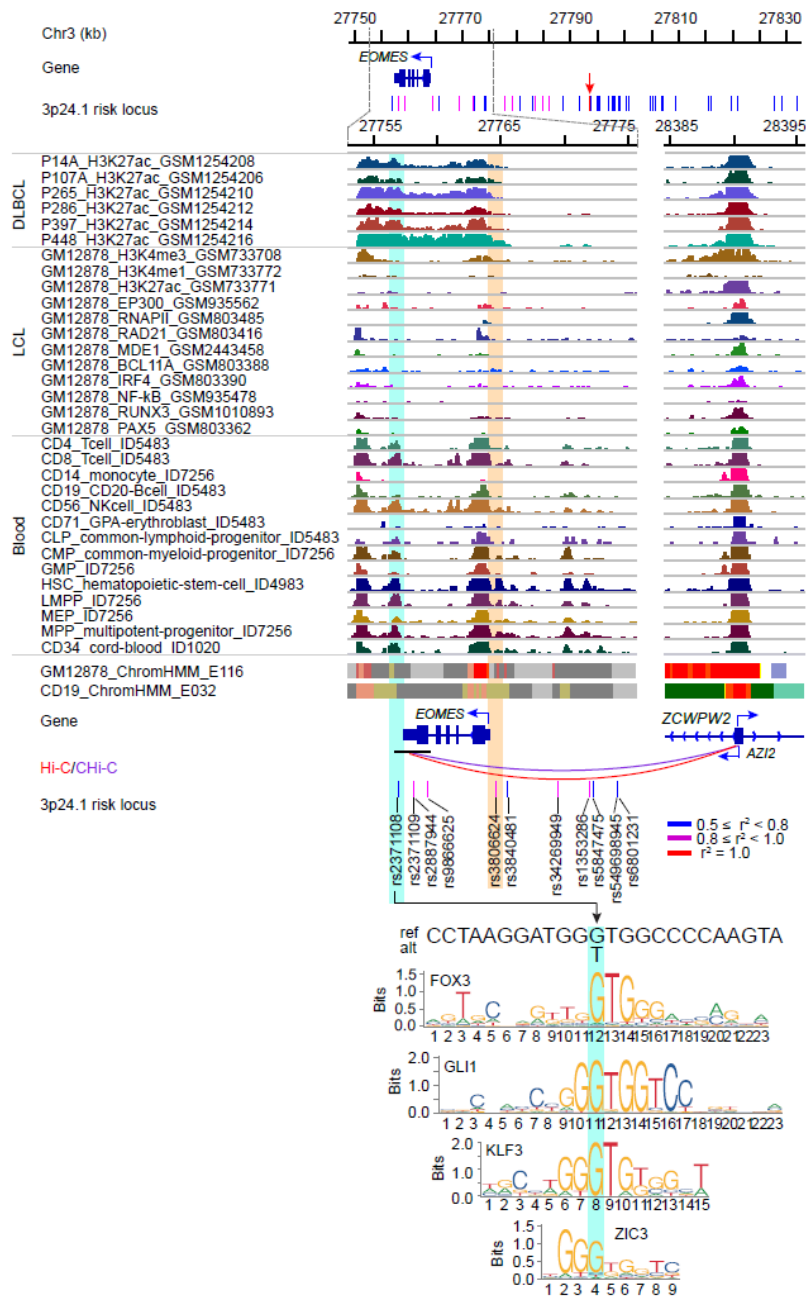
**Supplementary Figure 3. Epigenetic features of rs6773363 at independent loci associated with the risk of DLBCL.** Epigenetic features at the 3p24.1 risk locus. This 74,961-bp risk locus contains 51 variants ( $r^2 \geq 0.5$ ), with red arrow indicating the tag SNP rs6773363. For simplicity, only 11 of the variants were displayed (bottom panel) for the association with epigenetic profiles. Forty-three of the variants starting from rs34269949, including the tag SNP, did not overlap any of the active marks assayed in B-cells and DLBCL. Four variants between rs2371108 to rs9866625 ( $r^2 > 0.7$ ) overlapped an 11.6-kb super-enhancer. Based on the Hi-C data in GM12878 and capture Hi-C data from 17 blood cell types, this region showed strong long-range interactions with the promoters of *AZ12* and *ZCWPW2*. Rs3806624 (highlighted in light blue), located 417 bp upstream of the transcription start site of *EOMES*, is in the edge of the super-enhancer. E004, H1 BMP4 derived mesendoderm cultured cells; E046, primary natural killer cells from peripheral blood (CD56+); E124, monocytes-CD14+ RO01746 primary cells.

#### Chromatin state

|   |                                   |   |                              |
|---|-----------------------------------|---|------------------------------|
|  | 1 Active TSS                      |  | 9 Heterochromatin            |
|  | 2 Flanking active TSS             |  | 10 Bivalent/poised TSS       |
|  | 3 Transcription at gene 5' and 3' |  | 11 Flanking bivalent TSS/Enh |
|  | 4 Strong transcription            |  | 12 Bivalent enhancer         |
|  | 5 Weak transcription              |  | 13 Repressed Polycomb        |
|  | 6 Genic enhancers                 |  | 14 Weak repressed Polycomb   |
|  | 7 Enhancers                       |   | 15 Quiescent/low             |
|  | 8 ZNF genes + repeats             |   |                              |

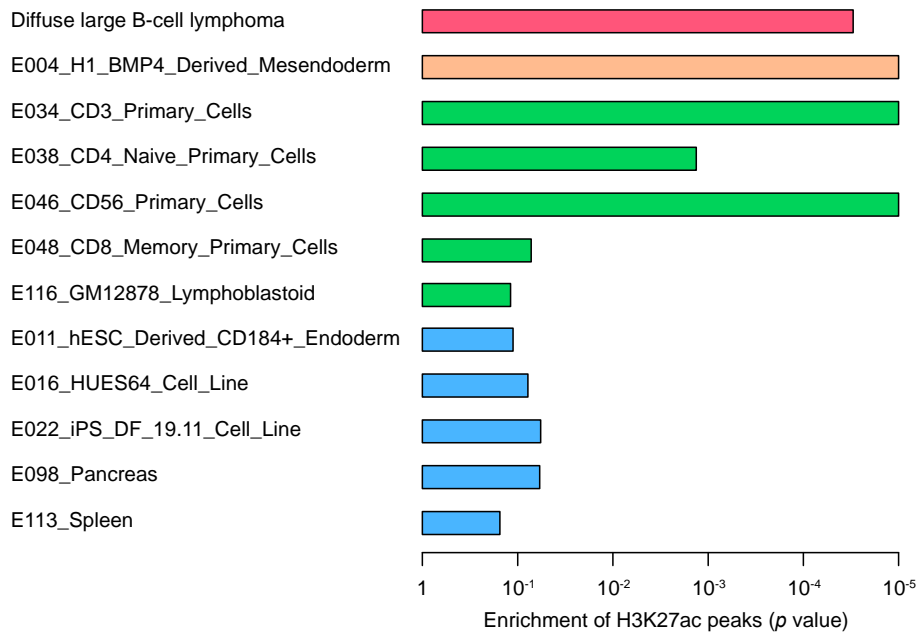


**Supplementary Figure 4. Regulatory variants at rs6773363-tagged DLBCL risk locus on 3p24.1.** The ATAC-seq data for the 14 blood cell types was downloaded from NCBI GEO under the accession GSE74912. SNP rs2371108 (highlighted), which overlapped a DLBCL super-enhancer, is located within an open chromatin region in nine blood cell types. The alternative T allele at this SNP was predicted to disrupt FOX3, GLI1, KLF3, and ZIC3 binding motif. In addition, SNP rs3806624 (highlighted) is located in an open chromatin region in hematopoietic stem cell. GMP, granulocyte macrophage progenitor; LMPP, lymphoid primed multipotent progenitor; MEP, megakaryocyte erythroid progenitor. See Supplementary Figure 3 legend for additional information.

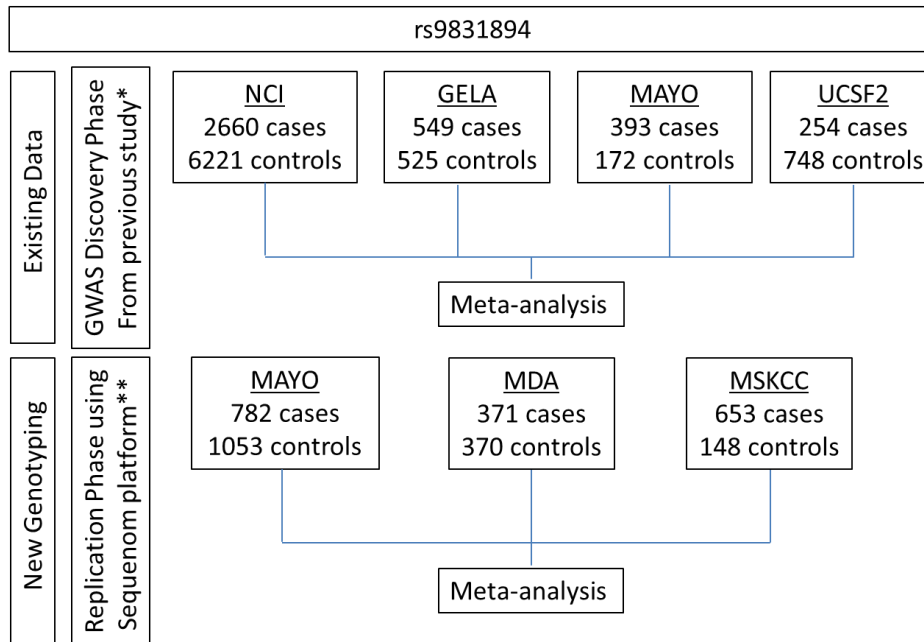




**Supplementary Figure 5. H3K27ac enrichment in DLBCL risk loci across 12 cell types.** Except DLBCL patient biopsy, the other 11 cell types are from the 127 reference epigenome, available at <https://egg2.wustl.edu/roadmap/data/byFileType/peaks/consolidated/narrowPeak/>. Only those whose H3K27ac peaks overlapped at least one of the risk loci were shown. The empirical  $p$  value was calculated by comparing to a null distribution of randomly sampling minor-allele-frequency-matched variants from the 1000 Genomes Project phase 3 release (Methods). DLBCL risk variants are enriched in DLBCL enhancer as expected, and in enhancers from three immune related cell types as well. E004-E116: reference epigenome identifier.

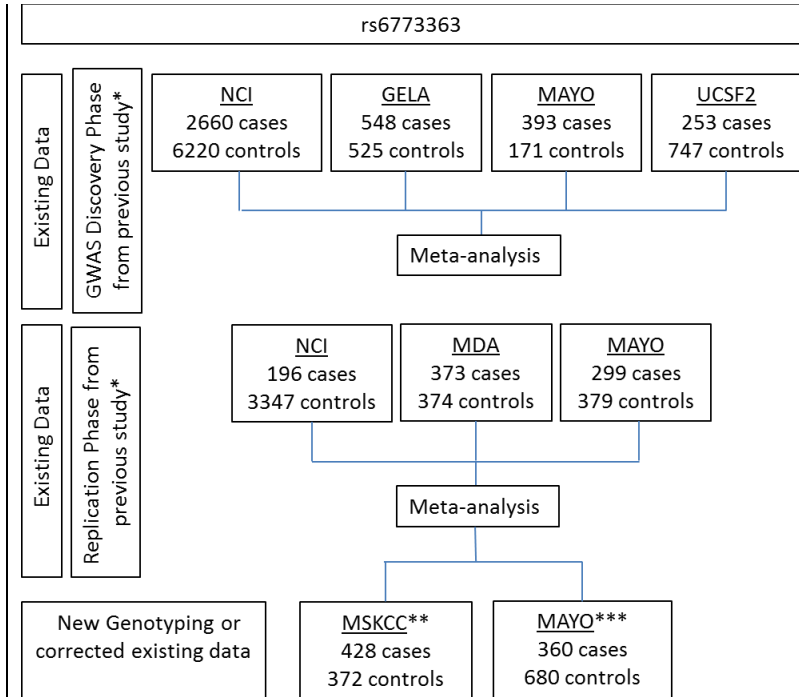


### Supplementary Figure 6. Schematic of the study design



\*Cerhan, J.R. *et al.* (2014) Genome-wide association study identifies multiple susceptibility loci for diffuse large B cell lymphoma. *Nature genetics*, **46**, 1233–8.

\*\*SNP failed design with Taqman and was unable to be analyzed in the replication stage in our previous GWAS.



\*Cerhan, J.R. *et al.* (2014) Genome-wide association study identifies multiple susceptibility loci for diffuse large B cell lymphoma. *Nature genetics*, **46**, 1233–8.

\*\*SNP miscoded in the original replication stage in our previous GWAS and now corrected

\*\*\*Additional cases and controls genotyped from Mayo in addition to our previous replication phase



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