

Supplemental Table 1. Number of SNPs with evidence of association with methylation of sites associated with lupus nephritis.

| CpG | Gene | # SNPs tested | # SNPs | # SNPs FDR q<0.05 |
|-------------------|-----------------------------------|---------------|-------------------|----------------------|
| | | | regression p<0.05 | |
| cg15262954 | <i>HELZ2</i> | 37 | 4 | 1 |
| cg19168338 | <i>CORO7</i> | 18 | 1 | 0 |
| cg00356916 | <i>CASQ2</i> | 26 | 0 | 0 |
| cg01079652 | <i>IFI44</i> | 57 | 2 | 0 |
| cg05883128 | <i>DDX60</i> | 51 | 0 | 0 |
| cg25114611 | <i>FKBP5;</i> <i>LOC285847</i> | 26 | 6 | 0 |
| cg22891070 | <i>HIF3A</i> | 49 | 13 | 3 |
| cg05552874 | <i>IFIT1</i> | 49 | 4 | 0 |
| cg18468088 | No gene w/in 5KB | 49 | 5 | 0 |
| cg00791854 | <i>IL6R</i> | 23 | 2 | 0 |
| cg23510764 | No gene w/in 5kb | 28 | 0 | 0 |
| cg02812482 | No gene w/in 5kb | 16 | 1 | 0 |
| cg04845871 | <i>MIR-29B-2, LOC148696</i> | 49 | 12 | 1 |
| cg23256579 | <i>PRR4</i> | 27 | 14 | 13 |
| cg16672562 | <i>HIF3A</i> | 49 | 14 | 4 |
| cg16032408 | <i>SLC6A6</i> | 92 | 1 | 0 |
| cg13437525 | <i>METTL7A</i> | 30 | 3 | 0 |
| cg14519664 | <i>MLLT3</i> | 39 | 1 | 0 |
| cg10959651 | <i>RSAD2</i> | 65 | 0 | 0 |

Supplemental Table 2. Association between methylation status and lupus nephritis adjusting for SNPs. “No SNP” results reflect the association observed between methylation of the site and lupus nephritis as presented in Table 1 and are included for reference.

| CpG | Gene | SNP | SNP association results | | | Lupus nephritis association results | |
|------------|-------------------------------|------------|---|--------------------|-------------|-------------------------------------|--------------------|
| | | | Regression coefficient for SNP (95% CI) | Regression p-value | FDR q-value | β (95% CI) | Regression p-value |
| cg15262954 | <i>HELZ2</i> | rs1151621 | 0.011 (0.005 to 0.016) | 2.66E-04 | 0.01 | 0.014 (0.009 to 0.019) | 1.65E-07 |
| | | No SNP | | | | 0.014 (0.009 to 0.019) | 2.84E-08 |
| cg22891070 | <i>HIF3A</i> | rs2072491 | -0.06 (-0.97 to -0.023) | 1.79E-03 | 0.029 | -0.094 (-0.132 to -0.055) | 2.56E-06 |
| | | rs10412620 | -0.046 (-0.068 to -0.023) | 9.79E-05 | 0.002 | -0.108 (-0.146 to -0.071) | 3.70E-08 |
| | | rs11880808 | 0.051 (0.027 to 0.074) | 3.33E-05 | 0.002 | -0.109 (-0.146 to -0.072) | 2.62E-08 |
| | | No SNP | | | | -0.094 (-0.130 to -0.058) | 5.69E-07 |
| cg16672562 | <i>HIF3A</i> | rs2072491 | -0.07 (-0.108 to -0.033) | 2.91E-04 | 0.007 | -0.091 (-0.130 to -0.053) | 5.49E-06 |
| | | rs10412620 | -0.042 (-0.066 to -0.019) | 4.63E-04 | 0.007 | -0.106 (-0.145 to -0.068) | 1.20E-07 |
| | | rs11880808 | 0.05 (0.026 to 0.074) | 6.02E-05 | 0.003 | -0.107 (-0.146 to -0.070) | 7.02E-08 |
| | | rs16980543 | -0.059 (-0.098 to -0.021) | 2.99E-03 | 0.035 | -0.102 (-0.141 to -0.064) | 3.67E-07 |
| | | No SNP | | | | -0.092 (-0.129 to -0.055) | 1.55E-06 |
| cg04845871 | <i>MIR-29B-2b, LOC148696b</i> | rs2076221 | 0.010 (0.005 to 0.016) | 4.78E-04 | 0.023 | 0.020 (0.012 to 0.028) | 8.98E-07 |
| | | No SNP | | | | 0.019 (0.012 to 0.027) | 1.22E-06 |
| cg23256579 | <i>PRR4</i> | rs10772380 | 0.026 (0.012 to 0.041) | 5.00E-04 | 0.007 | -0.050 (-0.070 to -0.030) | 1.66E-06 |
| | | rs1015443 | 0.025 (0.013 to 0.037) | 5.16E-05 | 0.003 | -0.054 (-0.074 to -0.035) | 1.71E-07 |
| | | rs10845242 | 0.026 (0.014 to 0.038) | 2.72E-05 | 0.003 | -0.055 (-0.074 to -0.035) | 1.36E-07 |
| | | rs10492102 | 0.028 (0.014 to 0.043) | 1.86E-04 | 0.005 | -0.050 (-0.070 to -0.030) | 2.19E-06 |
| | | rs3851583 | 0.028 (0.014 to 0.043) | 2.05E-04 | 0.005 | -0.049 (-0.069 to -0.029) | 2.31E-06 |
| | | rs10845252 | 0.026 (0.012 to 0.041) | 5.00E-04 | 0.007 | -0.050 (-0.070 to -0.030) | 1.65E-06 |

| | | | | | |
|------------|--------------------------|----------|-------|---------------------------|----------|
| rs7307031 | -0.02 (-0.033 to -0.007) | 3.50E-03 | 0.028 | -0.057 (-0.077 to -0.037) | 8.57E-08 |
| rs10845279 | -0.02 (-0.033 to -0.007) | 3.50E-03 | 0.028 | -0.057 (-0.077 to -0.036) | 1.01E-07 |
| rs12226919 | -0.02 (-0.033 to -0.007) | 3.50E-03 | 0.028 | -0.057 (-0.077 to -0.037) | 8.57E-08 |
| rs12226920 | -0.02 (-0.033 to -0.007) | 3.67E-03 | 0.028 | -0.057 (-0.077 to -0.036) | 1.05E-07 |
| rs7135018 | 0.028 (0.013 to 0.044) | 3.89E-04 | 0.007 | -0.049 (-0.069 to -0.028) | 4.42E-06 |
| rs12581501 | 0.025 (0.009 to 0.040) | 2.00E-03 | 0.025 | -0.053 (-0.073 to -0.033) | 4.23E-07 |
| rs4298989 | -0.02 (-0.033 to -0.007) | 3.50E-03 | 0.028 | -0.057 (-0.077 to -0.037) | 8.57E-08 |
| No SNP | | | | -0.050 (-0.070 to -0.030) | 1.41E-06 |

* β represents the difference in methylation between SLE cases with nephritis and SLE cases without nephritis adjusted for disease duration, population substructure, mixed cell populations, and listed SNP if indicated.