**Supplementary File 2.** Detailed performance comparisons between tuned autoencoder (AE) and HMM-based imputation tools (Minimac4, Beagle5, and Impute5).

|  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Dataset** | **MESA** | | | **Wellderly** | | | **HGDP** | | |
| **array** | **Affymetrix 6.0** | **UKB Axiom** | **Omni 1.5M** | **Affymetrix 6.0** | **UKB Axiom** | **Omni 1.5M** | **Affymetrix 6.0** | **UKB Axiom** | **Omni 1.5M** |
| **AE (tuned) vs Minimac4** | 9.47e-185\*\*\* | 0.00e+00\*\*\* | 7.22e-89\*\*\* | 6.27e-209\*\*\* | 0.00e+00\*\*\* | 2.75e-198\*\*\* | 2.35e-151\*\*\* | 0.00e+00\*\*\* | 5.55e-67\*\*\* |
| **AE (tuned) vs Beagle5** | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* |
| **AE (tuned) vs Impute5** | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 5.37e-191\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* | 0.00e+00\*\*\* |
| **Minimac4 vs Beagle5** | 1.73e-259\*\*\* | 6.06e-16\*\*\* | 0.00e+00\*\*\* | 2.68e-86\*\*\* | 1.65e-11\*\*\* | 6.87e-64\*\*\* | 0.00e+00\*\*\* | 2.62e-185\*\*\* | 0.00e+00\*\*\* |
| **Minimac4 vs Impute5** | 4.87e-38\*\*\* | 3.59e-48\*\*\* | 1.17e-22\*\*\* | 3.05e-74\*\*\* | 5.94e-15\*\*\* | 1.00e-25\*\*\* | 9.43e-261\*\*\* | 0.00e+00\*\*\* | 1.73e-251\*\*\* |
| **Beagle5 vs Impute5** | 1.92e-96\*\*\* | 2.75e-09\*\*\* | 1.23e-175\*\*\* | 3.65E-01 | 1.98E-01 | 9.53e-09\*\*\* | 1.22e-25\*\*\* | 2.61e-17\*\*\* | 8.36e-57\*\*\* |
| **AE (tuned)** | 0.410±0.001 | 0.395±0.001 | 0.452±0.001 | 0.537±0.001 | 0.605±0.001 | 0.586±0.001 | 0.363±0.001 | 0.364±0.001 | 0.392±0.001 |
| **Minimac4** | 0.390±0.001 | 0.364±0.001 | 0.436±0.001 | 0.500±0.001 | 0.557±0.001 | 0.551±0.001 | 0.350±0.001 | 0.340±0.001 | 0.385±0.001 |
| **Beagle5** | 0.383±0.001 | 0.379±0.001 | 0.420±0.001 | 0.484±0.001 | 0.549±0.001 | 0.534±0.001 | 0.326±0.001 | 0.328±0.001 | 0.353±0.001 |
| **Impute5** | 0.384±0.001 | 0.356±0.001 | 0.429±0.001 | 0.485±0.001 | 0.547±0.001 | 0.539±0.001 | 0.328±0.001 | 0.314±0.001 | 0.359±0.001 |

Validation accuracies were stratified by dataset (MESA, Wellderly, HGDP) and genotype array platform (Affymetrix 6.0, UKB Axiom, Omni 1.5M). We applied Wilcoxon rank-sum tests to compare the HMM-based tools to the reference tuned autoencoder (AE). \* represents p-values ≤ 0.05, \*\* indicates p-values ≤ 0.001, and \*\*\* indicates p-values ≤ 0.0001