

Table S2. Transcriptional Regulators, Kinases and Phosphatases that Affect Amino Acid Metabolism. Related to Figure 3 B-C.

| amino acid | transcriptional regulator (number) | kinase (number) | phosphatase (number) | total No |
|---------------|--|--|-----------------------------|-------------|
| alanine | <i>ADA2 GAT1 GCR2 HAP5 HMO1 HOT1 LEU3 MOT2 RGT1 SPT10 USV1 YRR1 (12)</i> | <i>ADK1 BUB1 CLB2 DBF2 ELM1 FAB1 GIN4 MCK1 PCL7 PHO80 PHO85 SKY1 TCO89 VIP1 (14)</i> | <i>OCA1 PTC6 SAP155 (3)</i> | 29 |
| aspartate | <i>HAP5 HIR2 IXR1 MOT2 SIR3 SPT10 UME6 XBP1 (8)</i> | <i>MCK1 PHO80 SCH9 SLT2 SPS1 VIP1 (6)</i> | <i>OCA1 PTC1 SIT4 (3)</i> | 17 |
| glutamate | <i>GCR2 GLN3 NRG1 RGT1 SPT10 YRR1 (6)</i> | <i>ADK1 DBF2 ELM1 FAB1 KIN4 MRK1 NPR1 SKM1 (8)</i> | <i>OCA1 SAP155 SIT4 (3)</i> | 17 |
| phenylalanine | <i>SPT10 YRR1 (2)</i> | <i>CTK3 PHO85 VIP1 (3)</i> | <i>SIT4 (1)</i> | 6 |
| glycine | <i>SKO1 SNF2 SOK2 TYE7 YRR1 (5)</i> | <i>FRK1 PHO80 SSN3 (3)</i> | <i>OCA1 (1)</i> | 9 |
| histidine | <i>PHO2 YRR1 (2)</i> | <i>CGII21 FAB1 SSN8 TCO89 (4)</i> | <i>SIT4 (1)</i> | 7 |
| isoleucine | <i>AFT1 SFL1 SPT10 SWI6 YRR1 (5)</i> | <i>YGK3 (1)</i> | <i>OCA1 PTC6 (2)</i> | 8 |
| lysine | <i>ADA2 CIN5 HIR2 LYS14 PHO2 SFL1 SFP1 SIR2 SNF2 ZAP1 (10)</i> | <i>ADK1 MRK1 PHO85 SSN8 (4)</i> | <i>OCA1 SIT4 (2)</i> | 16 |
| leucine | <i>YRR1 (1)</i> | | | 1 |
| methionine | <i>AFT1 MIG1 MOT2 SFP1 SPT10 YRR1 (6)</i> | <i>ADK1 (1)</i> | <i>SIT4 (1)</i> | 8 |
| asparagine | <i>CIN5 GLN3 HAP2 HAP5 LYS14 SFP1 UME6 YRR1 (8)</i> | <i>ADK1 CKA2 FAB1 SCH9 TCO89 YCK3 (6)</i> | | 14 |
| proline | <i>AFT1 GCR2 GLN3 HMO1 LEU3 LYS14 SPT10 SWI6 (8)</i> | <i>ADK1 CKA2 CTK3 FAB1 NPR1 PHO85 SCH9 (7)</i> | <i>OCA1 SAP155 (2)</i> | 17 |
| glutamine | <i>ADA2 GLN3 HAP2 HAP5 LEU3 SWI6 YRR1 (7)</i> | <i>CKA2 PHO85 SCH9 (3)</i> | <i>OCA1 PTC1 (2)</i> | 12 |
| arginine | <i>AFT1 ARG81 DOT6 LEU3 LYS14 PHO2 SWI6 (7)</i> | <i>CGII21 MCK1 NPR1 PBS2 SCH9 (5)</i> | <i>SIT4 (1)</i> | 13 |
| serine | <i>CHA4 HMO1 IXR1 LEU3 LYS14 RPH1 SKO1 SNF2 SPT2 USV1 YRR1 (11)</i> | <i>ADK1 ELM1 FAB1 PCL1 PCL5 PHO80 PRO1 SKY1 SSN3 TCO89 (10)</i> | <i>OCA1 PTC1 (2)</i> | 23 |
| threonine | <i>ADR1 DAL81 GZF3 HOT1 SIR2 SNF2 SPT10 UME6 YRR1 (9)</i> | <i>ADK1 CKA2 CTK3 FAB1 PHO80 YGK3 (6)</i> | <i>PTC1 SAP4 (2)</i> | 17 |
| valine | <i>AFT1 AFT2 DAL81 LEU3 SNF2 SPT10 (6)</i> | <i>FAB1 FRK1 PKH2 PRO1 TCO89 (5)</i> | <i>OCA1 PTC1 PTC6 (3)</i> | 14 |
| tryptophan | <i>GCN4 LYS14 SUM1 YRR1 (4)</i> | <i>VIP1 (1)</i> | | 5 |
| tyrosine | <i>AFT1 ASH1 LEU3 LYS14 MAC1 MOT2 PHO2 SNF2 SPT10 SUM1 ZAP1 (11)</i> | <i>ADK1 PBS2 PHO85 PRO1 SSN8 TCO89 VIP1 (7)</i> | <i>OCA1 PTC1 SIT4 (3)</i> | 21 |

Table S4. Assignment of strains to batches. Related to STAR Methods.

| batch 1 | batch 2 | batch 3 | batch 4 | batch 5 | batch 6 | batch 7 | batch 8 | batch 9 | batch 10 | batch 11 |
|---------|---------|---------|---------|---------|---------|---------|---------|---------|----------|----------|
| 43 | 23 | 16 | 38 | 25 | 13 | 21 | 27 | 17 | 32 | 26 |
| 28 | 7 | 31 | 20 | 5 | 9 | 36 | 8 | 14 | 42 | 4 |
| 41 | 1 | 37 | 11 | 12 | 29 | 19 | 10 | 18 | 34 | 30 |
| 15 | 44 | 24 | 22 | 6 | 40 | 3 | 35 | 2 | 33 | 39 |
| | | B | C | 47 | 45 | G | E | D | A | 46 |
| | | 49 | 74 | 71&F | 70 | 72 | 48 | 50 | 51&73 | 75 |

Table S5. Details for the analytical method to determine amino acids levels by LC-MS/MS; SRM transitions. Related to STAR Methods.

| amino acid | compound abbreviation | SRM transition | fragmentor (V) | collision energy (V) | retention time (min) | external calibration |
|---------------|-----------------------|----------------|----------------|----------------------|----------------------|----------------------|
| alanine | A | 90 > 44.1 | 50 | 8 | 1.87 | linear |
| arginine | R | 175.1 > 70 | 100 | 15 | 2.37 | power |
| asparagine | N | 133.1 > 74 | 80 | 9 | 2.12 | linear |
| aspartate | D | 134.1 > 74 | 80 | 10 | 2.24 | linear |
| glutamate | E | 148.1 > 84.1 | 75 | 10 | 2.09 | power |
| glutamine | Q | 147.1 > 84 | 50 | 16 | 2.07 | power |
| glycine | G | 76 > 30.1 | 50 | 5 | 1.98 | linear |
| histidine | H | 156.1 > 110.2 | 80 | 12 | 2.36 | power |
| isoleucine | I | 132.1 > 86 | 80 | 8 | 1.33 | power |
| leucine | L | 132.1 > 86 | 80 | 8 | 1.24 | power |
| lysine | K | 147.1 > 84 | 50 | 16 | 2.41 | power |
| methionine | M | 150.1 > 104 | 40 | 8 | 1.47 | linear |
| phenylalanine | F | 166.1 > 120 | 100 | 9 | 1.23 | power |
| proline | P | 116.1 > 70.1 | 100 | 13 | 1.60 | power |
| serine | S | 106 > 60 | 40 | 9 | 2.10 | power |
| threonine | T | 120.1 > 74 | 80 | 9 | 1.96 | power |
| tryptophane | W | 205.1 > 188 | 85 | 5 | 1.29 | linear |
| tyrosine | Y | 182 > 165 | 90 | 5 | 1.64 | power |
| valine | V | 118.1 > 71.9 | 100 | 10 | 1.57 | power |