

1 Supporting information

2 Supplemental Tables

3 Table S1 The compositions of the experimental diets (g/100g)

Ingredient	CON ¹	D+CON ²	D+MR ³
Soy protein ⁴	7.59	7.59	7.59
L-Arginine	0.66	0.66	0.66
L-Histidine	0.18	0.18	0.18
L-Isoleucine	0.51	0.51	0.51
L-Leucine	0.64	0.64	0.64
L-Lysine	0.11	0.11	0.11
L-Methionine	0.69	0.69	0.00
L-Phenylalanine	0.85	0.85	0.85
L-Threonine	0.65	0.65	0.65
L-Tryptophan	0.10	0.10	0.10
L-Valine	0.57	0.57	0.57
L-Glutamic acid	1.37	1.37	2.06
L-Glycine	2.08	2.08	2.08
Corn starch	64.09	64.09	64.09
maltodextrin	5.00	5.00	5.00
Sucrose	0.10	0.10	0.10
Soybean oil	2.00	2.00	2.00
Pork Lard	2.20	2.20	2.20

Cellulose	5.00	5.00	5.00
Mineral mixture-AIN-76A	3.50	3.50	3.50
Mineral vitamin-AIN-76A	1.00	1.00	1.00
Choline chloride	0.11	0.11	0.11
CMC	1.00	1.00	1.00
Total	100.00	100.00	100.00

4 ¹CON, control diet, was mainly based upon the AIN-76A formulation.

5 ²D+CON, control diet, was mainly based upon the AIN-76A formulation.

6 ³D+MR, control diet in which methionine was restricted from 0.86% to 0.17%. The glutamic acid was
7 increased to compensate for the reduced methionine content and to create equal amounts of total amino
8 acids.

9 ⁴Amino acid composition of soy protein was as follows: 6.19% leucine, 4.11% isoleucine, 5.49%
10 valine, 1.18% methionine, 1.66% cysteine, 4.09% phenylalanine, 2.57% tyrosine, 4.83% lysine, 2.21%
11 threonine, 1.07% tryptophan, 1.99% histidine, 6.11% arginine, 3.30% serine, 3.25% alanine, 5.56%
12 proline, 3.27% glycine, 17.49% glutamic acid 9.44% aspartic acid. 1 g cysteine is equal to 0.64 g
13 methionine.¹

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15 Table S2 Sequence of primers in quantitative real-time reverse transcription polymerase chain reaction

Genes	Forward (5'–3')	Reverse (5'–3')
Nrf2	TCTTGGAGTAAGTCGAGAAGTGT	GTTGAAACTGAGCGAAAAAGGC
HO-1	GGAAATCATCCCTTGCACGC	TGTTTGAACCTTGGTGGGGCT
NQO-1	AGGATGGGAGGTACTCGAATC	AGGCGTCCTTCCTTATATGCTA

Nr1	AGAGCCCGACCCTAAAAAGAA	CCCTCCTCCCTCTCAATAGC
Nr2a	ACGTGACAGAACGCGAACTT	TCAGTGCGGTTTCATCAATAACG
Nr2b	GGCTCCGAGACTTCTACCTG	GCTTATCGCCTGTTCCGT
CAMK2A	AAACACTCAACAAAATCAAACGAC	GCCACAGAGAGACCAAAGCA
CAMK2B	GAGAACCTGAGATACTGGACGGAT	ATTTCATCAAACACTTGTATGGAC
	A	C
CAMK2D	CATCTTGACAACACTATGCTGGCTACG	TTGATGATCTCCTGTTTTCGTGCT
NF- κ B	ATGGCAGACGATGATCCCTAC	TGTTGACAGTGGTATTTCTGGTG
BDNF	TCATACTTCGGTTGCATGAAGG	TCATACTTCGGTTGCATGAAGG
TrkB	CTGGGGCTTATGCCTGCTG	GTACACCAAATCCTAGCGGAAC
RC3	TCCAAGCCAGACGACGATATT	CACACTCTCCGCTCTTTATCTTC
Gap-43	TGGTGTCAAGCCGGAAGATAA	GCTGGTGCATCACCCCTTCT
PSD-95	TCTGTGCGAGAGGTAGCAGA	AAGCACTCCGTGAACTCCTG
SYNAPO	CCTGCCCGTAACTTCCGTG	GAGCGGCGGTAGGGAAAAG
β -actin	GGCTGTATTCCTCCATCG	CCAGTTGGTAACAATGCCATG

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- 16 Nrf2: nuclear factor erythroid-derived 2-like 2, HO-1: heme oxygenase 1, NQO-1: NADPH quinone
- 17 oxidoreductase-1, Nr1: N-methyl-D-aspartate (NMDA) receptor 1, Nr2a: N-methyl-D-aspartate
- 18 (NMDA) receptor 2a, Nr2b: N-methyl-D-aspartate (NMDA) receptor 2b, CAMK2A:
- 19 Ca²⁺/calmodulin-dependent protein kinase II alpha chain, CAMK2B: Ca²⁺/calmodulin-dependent
- 20 protein kinase II beta chain, CAMK2D: Ca²⁺/calmodulin-dependent protein kinase II delta chain, NF-
- 21 κ B: nuclear factor kappa B, BDNF: brain-derived neurotrophic factor, TrkB: tyrosine kinase receptor B,
- 22 RC3: neurogranin, Gap-43: neuromodulin, PSD-95: postsynaptic density protein 95, SYNAPO:

23 synaptopodin.

24 Table S3 ¹H chemical shift assignment of the metabolites of short-chain fatty acid production

25 pathways in feces

Keys	Metabolites	Moieties	$\delta^1\text{H}$ (ppm) and multiplicity	Annotated chemical shift (ppm)	Integral region (ppm)
1	butyrate	CH ₃ , β CH ₂ , α CH ₂	0.90(t), 1.56(m), 2.15(t)	0.90(CH ₃)	0.88-0.92
2	α -ketoisocaproate	2*CH ₃ , CH, CH ₂	0.92(d), 2.06(m), 2.61(d)	2.61(CH ₂)	2.60-2.62
3	propionate	CH ₃ , CH ₂	1.07(t), 2.19(q)	1.07(CH ₃)	1.06-1.08
4	α -keto- β -methyl-valerate	δ CH ₃ , γ' CH ₃ , γ CH, γ' CH, β CH	0.88(t), 1.10(d), 1.47(m), 1.69(m), 2.93(m)	1.10(CH ₃)	1.10-1.11
5	α -ketoisovalerate	CH ₃ , CH	1.13(d), 3.02(m)	1.13(CH ₃)	1.12-1.13
6	lactate	α CH, β CH ₃	1.33(d), 4.11(q)	1.33(CH ₃)	1.31-1.35
7	acetate	CH ₃	1.92(s)	1.92(CH ₃)	1.91-1.93
8	5-aminovalerate	2CH ₂ , 3CH ₂ , 1CH ₂ , 4CH ₂	1.62(m), 1.65(m), 2.26(t), 3.02(t)	2.26(CH ₂)	2.25-2.27
9	pyruvate	CH ₃	2.37(s)	2.37(CH ₃)	2.37-2.38
10	succinate	CH ₂	2.41(s)	2.41(CH ₂)	2.40-2.42
11	malate	β CH ₂	2.38(dd), 2.70(dd)	2.70(CH ₂)	2.69-2.71

		β' CH ₂ , α CH	4.31(dd)		
12	malonate	CH ₂	3.11(s)	3.11(CH ₂)	3.10-3.12
13	fumarate	CH	6.52(s)	6.52(CH)	6.52-6.53
14	formate	CH	8.46(s)	8.46(CH)	8.46-8.47

26 s, singlet, d, doublet, t, triplet, q, quartet, dd, doublet of doublets, m, multiplet.

27 References

28 M. Di Buono, L. J. Wykes, R. O. Ball and P. B. Pencharz, Dietary cysteine reduces the methionine
 29 requirement in men, *Am. J. Clin. Nutr.*, 2001, **74**, 761-766.