## **Supplementary material corresponding to the manuscript:**

## Metformin potentiates the benefits of dietary restraint: a metabolomic study

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## Supplementary material and methods S1: Food intake measurement

Food intake measurement was performed as previously reported [1]. Mice were group housed and given a known, pre-weighed amount of food in their home cage hopper. Once a week, the remaining food is weighed. Then, new amount of food is administered. Calculations were performed as follow: Food intake = Food provided – Food ingested – food wasted. Food intake measurements were divided by the number of animals per home cage, so assuming a meanintake per animals, and represented in g, although it can be easily converted to kcal, by using data on Table S1.

[1] Ellacott KLJ et al., Cell Metab, 2010.

Table S1: Nutritional composition of experimental diets..

	CD (Chow diet)	HFD (High-fat diet)
Crude Nutrients	OD (Onlow diet)	in b (mgn-lat dict)
Crude protein (%)	16.1	24.10
Crude fat (%)	3.1	34.60
Crude fibre (%)	3.9	7.2
Carbohydrate (available) (%)	71.8	24.80
Crude ash (%)	4.2	8.5
Nitrogen free extracts (%)	43.5	23.30
Energy	0.0 (40.07)	5.70 (04)
Energy density (Kcal/g (Kj/g)) Calories from protein (%)	3.3 (13.97) 19.3	5.70 (24) 19
Calories from fat (%)	8.4	60
Calories from carbohydrate (%)	72.4	21
Minerals		
Calcium (%)	1.38	0.98
Phosphorus (%)	0.99	0.65
Sodium (%)	0.47	0.20
Magnesium (%)	0.34	0.17
Potassium (%)	1.45	0.98
Chloride (%)	0.69	-
Fatty acids	ND	0.04
Caprid acid (C10:0) (%) Lauric acid (C12:0) (%)	ND ND	0.04 0.07
Myristic acid (C14:0) (%)	ND ND	0.07
Palmitic acid (C14.0) (%)	0.87	7.72
Palmitoleic acid (C16:1) (%)	0.03	0.94
Margaric acid (C17:0) (%)	ND	-
Stearic acid (C18:0) (%)	0.08	4.34
Oleic acid (C18:1) (%)	0.53	13.57
Linoleic acid (C18:2) (%)	1.34	4.75
Linolenic acid (C18:3) (%)	0.15 ND	0.51
Arachidic acid (C20:0) (%) Paullinic acid (C20:1) (%)	ND ND	0.02 0.01
Arachidonic acid (C20:4) (%)	ND ND	0.53
Cholesterol (mg/kg)	ND	265
Amino acids		
Lysine (%)	3.9	1.98
Methionine (%)	1.49	0.83
Cystine (%)	1.08	0.46
Met+Cys (%)	ND	1.28
Threonine (%)	ND 0.00	1.07
Tryptophan (%)	0.88 4.75	0.31
Arginine (%)   Histidine (%)	ND	0.88 0.76
Valine (%)	ND ND	1.64
Isoleucine (%)	ND	1.25
Leucine (%)	ND	2.36
Phenylalanine (%)	ND	1.29
Phe+Tyr (%)	ND	2.57
Glycine (%)	0.41	0.50
Glutamic acid (%) Aspartic acid (%)	ND ND	5.41 1.79
Proline (%)	ND ND	2.76
Alanine (%)	ND	0.79
Serine (%)	ND	1.43
Vitamins		
Vitamin A (IU/kg)	7500	15000
Vitamin D <sub>3</sub> (IU/kg)	1000	1500
Vitamin E (IU/kg)	120	225
Vitamin K (as menadione) (mg/Kg) Vitamin C (mg/Kg)	2.5 ND	20 30
Thiamin (B <sub>1</sub> ) (mg/Kg)	7	16
Riboflavin (B <sub>2</sub> ) (mg/Kg)	6.5	16
Pyridoxime (B <sub>6</sub> ) (mg/Kg)	2.6	18
Cyanocobalamin (B <sub>12)</sub> (µg/Kg)	20	30
Nicotinic acid (mg/Kg)	75	45
Pantothenic acid (mg/Kg)	17	55
Folic acid (mg/Kg)	0.5	19
Biotin (µg/Kg) Choline-Chloride (mg/Kg)	40 1600	310 2300
Inositol (mg/Kg)	ND	80
Trace elements	110	
Iron (mg/Kg)	280	139
Manganese (mg/Kg)	90	82
Zinc (mg/Kg)	64	56
Copper (mg/Kg)	18	12
lodine (mg/Kg)	ND ND	0.97
Selenium (mg/Kg)	ND ND	0.13
Cobalt (mg/Kg)	IND	0.13