DIETS®



Product Data - DIO SERIES DIETS

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The "Original" High-Fat Diets for Diet Induced Obesity

Formulated by E. A. Ulman, Ph.D., Research Diets, Inc., 8/26/98 and 3/11/99.

Research Diets, Inc. formulated the "original" high-fat diet for diet induced obesity (DIO) studies in 1996. Today, our high-fat diets are the research standard for DIO mice and rats worldwide.



DIO Low-Fat Control Diets

Matched, Purified Ingredient Diet

We recommend that you use a matched, purified ingredient diet and not a grain-based 'chow' diet. There are many, many differences between purified diets and chow diets and these variables make it difficult to interpret your data from a study in which one group was fed a purified ingredient high-fat and the other a low-fat chow diet. Differences between your groups could be due to the level of fat, but could also be due to differences in fiber type and level, source of carbohydrate, and the presence or absence of plant chemicals (such as phytoestrogens), just to name a few.

See next page for low-fat control formulas.



(DIO) Formulas										
Product #	D1	2451	D12492							
	gm%	kcal%	gm%	kcal%						
Protein	24	20	26	20						
Carbohydrate	41	35	26	20						
Fat	24	45	35	60						
Total		100		100						
kcal/gm	4.73		5.24							
Ingredient	gm	kcal	gm	kcal						
Casein, 30 Mesh	200	800	200	800						
L-Cystine	3	12	3	12						
Corn Starch	72.8	291	0	0						
Maltodextrin 10	100	400	125	500						
Sucrose	172.8	691	68.8	275						
Cellulose, BW200	50	0	50	0						
Soybean Oil	25	225	25	225						
Lard	177.5	1598	245	2205						
Mineral Mix S10026	10	0	10	0						
DiCalcium Phosphate	13	0	13	0						
Calcium Carbonate	5.5	0	5.5	0						
Potassium Citrate, 1 H2O	16.5	0	16.5	0						
		(0)		(0)						
Vitamin Mix V10001	10	40	10	40						
Choline Bitartrate	2	0	2	0						
	0.05									
FD&C Ked Dye #40	0.05	0	0.05							
FD&C Blue Dye #1			0.05	0						
Total	858.15	4057	773.85	4057						

*Typical analysis of cholesterol in lard = 72 mg per 100 gram. **D12451** -Cholesterol (mg)/4057 kcal = 167.8 Cholesterol (mg)/kg = 195.5 **D12492** -Cholesterol (mg)/4057 kcal = 216.4 Cholesterol (mg)/kg = 279.6

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Open formula purified diets for lab animals

Low-Fat Control Formulas

There are many options for low-fat control diets. Here are a few examples:

D12450B

Contains 35% sucrose by energy, unlike D12451 and D12492. Sucrose is made up of glucose and fructose and it has been shown that diets very high in sucrose or fructose (~60-70% by energy) can induce hypertriglyceridemia, insulin resistance and fatty liver.

Match Sucrose Calories

Keep the amount of sucrose (as a percent of calories) constant across low and high-fat diets. D12450H matches the sucrose calories in D12451. D12450J matches the sucrose calories in D12492. These diets are formulated in such a way that when animals in the low and high-fat groups consume the same number of calories, they will also consume the same amount of sucrose.

Replace Sucrose with Corn Starch

D12450K contains no sucrose but only corn starch and maltodextrin as the sources of carbohydrate. Maltodextrin is a partially hydrolyzed form of corn starch that allows us to produce a quality pellet in a high corn starch diet.

There are more possible low-fat diet formulas. Please contact one of our scientists and we can help you decide if one of the above diets or perhaps a different, custom diet is better for your research.



*Typical analysis of cholesterol in lard = 72 mg per 100 gram. D12450B -Cholesterol (mg)/4057 kcal = 54.4 Cholesterol (mg)/kg = 51.6

35% Sucrose

D12451 Match 17% Sucrose

D12492 Match No Sucrose

	35% Sucrose		17% Sucrose		7% Sucrose		No Sucrose		
Product #	D12450B		D124	D12450H		D12450J		D12450K	
	gm%	kcal%	gm%	kcal%	gm%	kcal%	gm%	kcal%	
Protein	19.2	20.0	19.2	20.0	19.2	20.0	19.2	20.0	
Carbohydrate	67.3	70.0	67.3	70.0	67.3	70.0	67.3	70.0	
Fat	4.3	10.0	4.3	10.0	4.3	10.0	4.3	10.0	
Total		100.0		100.0		100.0		100.0	
kcal/gm	3.85		3.85		3.85		3.85		
Ingredient	gm	kcal	gm	kcal	gm	kcal	gm	kcal	
Casein, 80 Mesh	200	800	200	800	200	800	200	800	
L-Cystine	3	12	3	12	3	12	3	12	
Corn Starch	315	1260	452.2	1808.8	506.2	2024.8	550	2200	
Maltodextrin 10	35	140	75	300	125	500	150	600	
Sucrose	350	1400	172.8	691.2	68.8	275.2	0	0	
Cellulose, BW200	50	0	50	0	50	0	50	0	
Soybean Oil	25	225	25	225	25	225	25	225	
Lard	20	180	20	180	20	180	20	180	
Mineral Mix S10026	10	0	10	0	10	0	10	0	
DiCalcium Phosphate	13	0	13	0	13	0	13	0	
Calcium Carbonate	5.5	0	5.5	0	5.5	0	5.5	0	
Potassium Citrate, 1 H2O	16.5	0	16.5	0	16.5	0	16.5	0	
Vitamin Mix V10001	10	40	10	40	10	40	10	40	
Choline Bitartrate	2	0	2	0	2	0	2	0	
FD&C Yellow Dye #5	0.05	0	0.04	0	0.04	0	0	0	
FD&C Red Dye #40	0	0	0.01	0	0	0	0.025	0	
FD&C Blue Dye #1	0	0	0	0	0.01	0	0.025	0	
Total	1055.05	4057	1055.05	4057	1055.05	4057	1055.05	4057	

D12451, D12492 The "Original High-Fat Diets"

These are just a sample of the hundreds of citations comprising the body of scientific work built around these OpenSource diet formulas

References

DIO Rat Models

- Anderson EJ, Lustig ME, Boyle KE, et al. Mitochondrial H 2 O 2 emission and cellular redox state link excess fat intake to insulin resistance in both rodents and humans. J Clin Invest. 2009;119(3):573–581. 1.
- Burgmaier M, Sen S, Philip F, et al. Metabolic adaptation follows contractile dysfunction in the heart of obese zucker rats fed a high-fat "Western" diet. Obesity (Silver Spring, Md.). 2010;18(10):1895-901. 2.
- Chan C, Leo DD, Joseph J, McQuaid T. Increased uncoupling protein-2 levels in β -cells are associated with impaired glucose-stimulated insulin secretion. Diabetes. 2001;50(June):1302-1310.
- 2001;50(June):1502-1510. Chelikani PK, Haver AC, Reidelberger RD. Intermittent intraperitoneal infusion of peptide YY(3-36) reduces daily food intake and adiposity in obese rats. American Journal of Physiology. Regulatory, Integrative and Comparative Physiology. 2007;293(1):R39-46. 4.
- 5.
- 6.
- 7.
- 2007;293(1):R39-46.
 Dorfman SE, Laurent D, Gounarides JS, et al. Metabolic implications of dietary trans-fatty acids. Obesity (Silver Spring, Md.). 2009;17(6):1200-7.
 Erickson SD, Banner B, Berthel S, et al. Potent, selective MCH-1 receptor antagonists. Bioorganic & Medicinal Chemistry Letters. 2008;18(4):1402-6.
 Esler WP, Rudolph J, Claus TH, et al. Small-Molecule Ghrelin Receptor Antagonists Improve Glucose Tolerance, Suppress Appetite, and Promote. Receptor. 2007;148(11):5175-5185.
 Fong TM, Guan X-M, Marsh DJ, et al. Antiobesity efficacy of a novel cannabinoid-1 receptor inverse agonist, N-[(15,2S)-3-(4-chlorophenyl)-2-(3-cyanophenyl)-1-methylpropyl]-2-methyl-2-[[5-(trifluoromethyl)pyrinin-2-yi]oxylpropanamide (MK-0364), in rodents. The Journal of Pharmacology and Experimental Therapeutics. 2007;321(3):1013-22.
 Furnes MW, Zhao C-M, Chen D. Development of obesity is associated with 8.
- Furnes MW, Zhao C-M, Chen D. Development of obesity is associated with increased calories per meal rather than per day. A study of high-fat diet-induced obesity in young rats. Obesity Surgery. 2009;19(10):1430-8.
 Gollisch KSC, Brandauer J, Jessen N, et al. Effects of exercise training on subcutaneous and visceral adipose tissue in normal- and high-fat diet-fed rats. American Journal of Physiology. Endocrinology and Metabolism. 2009;297(2):E495-504.
 Hägerkvist R, Jansson L, Welsh N. Imatinib mesylate improves insulin sensitivity and glucose disposal rates in rats fed a high-fat diet. Clinical Science (London, England : 1979). 2008;114(1):65-71.
 Johnson J a, Trasino SE, Ferrante AW, Vasselli JR. Prolonged decrease of adipocyte size after rosiglitazone treatment in high- and low-fat-fed rats. Obesity (Silver Spring, Md.). 2007;15(11):2653-63.
 Kishino E, Ito T, Fujita K, Kiuchi Y. A Mixture of the Salacia reticulata (Kotala

- Kishino E, Ito T, Fujita K, Kiuchi Y. A Mixture of the Salacia reticulata (Kotala himbutu) Aqueous Extract and Cyclodextrin Reduces the Accumulation of Visceral Fat Mass in Mice and Rats with High-Fat Diet–Induced Obesity. The Journal of Nutrition. 2006;136(2):433.
- Laurent D, Didier L, Yerby B, et al. Diet-induced modulation of mitochondrial activity in rat muscle. American Journal of Physiology. Endocrinology and Metabolism. 2007;293(5):E1169-77.
- Metabolism. 2007;295(5):E1109-77.
 15. Li G, Zhang Y, Cheng KY, Scarpace PJ. Lean rats with hypothalamic pro-opiomelanocortin overexpression exhibit greater diet-induced obesity and impaired central melanocortin responsiveness. Diabetologia. 2007;50(7):1490-9.
 16. Lomba A, Milagro FI, García-Díaz DF, et al. Obesity induced by a pair-fed High-Fat sucrose diet: methylation and expression pattern of genes related to energy a present of the in Houhle and Director 2010;9(60).
- homeostasis. Lipids in Health and Disease. 2010;9(60).
 17. Morgan EE, Rennison JH, Young ME, et al. Effects of chronic activation of peroxisome proliferator-activated receptor-alpha or high-fat feeding in a rat infarct model of heart failure. American Journal of Physiology-Heart and Circulatory Physiology-Heart and Circulatory
- Physiology. 2006;290(5):H1899.
 Nagae A, Fujita M, Kawarazaki H, et al. Effect of High-Fat loading in Dahl salt-sensitive rats. Clinical and Experimental Hypertension. 2009;31(5):451–461.
 Nagae A, Fujita M, Kawarazaki H, et al. Sympathoexcitation by oxidative stress in

- Nagae A, Fujita M, Kawarazaki H, et al. Sympathoexcitation by oxidative stress in the brain mediates arterial pressure elevation in obesity-induced hypertension. Circulation. 2009;119(7):978-86.
 Okere IC, Young ME, McElfresh TA, et al. Low carbohydrate/high-fat diet attenuates cardiac hypertrophy, remodeling, and altered gene expression in hypertension. Hypertension. 2006;48(6):1116.
 Omagari K, Kato S, Tsuneyama K, et al. Effects of a long-term high-fat diet and switching from a high-fat to low-fat, standard diet on hepatic fat accumulation in Sprague-Dawley rats. Digestive Diseases and Sciences. 2008;53(12):3206-12.
 Picó C, Oliver P, Sánchez J, et al. The intake of physiological doses of leptin during lactation in rats prevents obesity in later life. International Journal of Obesity (2005). 2007;31(8):1199-209.
 Rizi S, Tiwari S, Sharma N, Rash A, Ecclbarger CM, Abundance of the Na-K-2CI
- Riazi S, Tiwari S, Sharma N, Rash A, Ecelbarger CM. Abundance of the Na-K-2Cl cotransporter NKCC2 is increased by high-fat feeding in Fischer 344 X Brown Norway (F1) rats. American Journal of Physiology. Renal physiology. 2009;296(4):F762-70.
- Stylopoulos N, Hoppin AG, Kaplan LM. Roux-en-Y gastric bypass enhances energy expenditure and extends lifespan in diet-induced obese rats. Obesity (Silver Spring, Md.). 2009;17(10):1839-47.
- 25. Tamashiro KLK, Terrillion CE, Hyun J, Koenig JI, Moran TH. Prenatal stress or high-fat diet increases susceptibility to diet-induced obesity in rat offspring. Diabetes. 2009;58(5):1116.
- 26. Tanaka S, Hayashi T, Toyoda T, et al. High-fat diet impairs the effects of a single bout of endurance exercise on glucose transport and insulin sensitivity in rat skeletal muscle. Metabolism: Clinical and Experimental. 2007;56(12):1719-28.
- Wickers MH, Gluckman PD, Coveny AH, et al. The effect of neonatal leptin treatment on postnatal weight gain in male rats is dependent on maternal nutritional status during pregnancy. Endocrinology. 2008;149(4):1906-13.
 Wang Y, Liu J. Plasma ghrelin modulation in gastric band operation and sleeve gastrectomy. Obesity Surgery. 2009;19(3):357-62.
 Yang L, Scott K a, Hyun J, et al. Role of dorsomedial hypothalamic neuropeptide Y in modulating food intake and energy balance. The Journal of Neuroscience : the

- Yang L, Scott K a, Hyun J, et al. Role of dorsomedial hypothalamic neuropeptide M in modulating food intake and energy balance. The Journal of Neuroscience : the official journal of the Society for Neuroscience. 2009;29(1):179-90.
 Zhang J, Matheny MK, Tümer N, Mitchell MK, Scarpace PJ. Leptin antagonist reveals that the normalization of caloric intake and the thermic effect of food after high-fat feeding are leptin dependent. American Journal of Physiology. Regulatory, Integrative and Comparative Physiology. 2007;292(2):R868-74.

D12451, D12492 The "Original High-Fat Diets"

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References

DIO Mouse Models

- Ahmadie R, Santiago J-jon, Walker J, et al. A High-Lipid Diet Potentiates Left Ventricular Dysfunction in Nitric Oxide Synthase 3-Deficient Mice after Chronic Pressure Overload. Journal of Nutrition. 2010;140(June):1438-1444.
 Atsui YM, Irasawa YH, Ugiura TS, et al. Metformin Reduces Body Weight Gain
- and Improves Glucose Intolerance in High-Fat Diet-Fed C57BL / 6J Mice. Biol.
- and Improves Glucose Intolerance in High-Fat Diet-Fed C5/BL/6J Mice. Biol. Pharm. Bull. 2010;33(June):963-970. Bradford EM, Miller ML, Prasad V, et al. CLIC5 mutant mice are resistant to diet-induced obesity and exhibit gastric hemorrhaging and increased susceptibility to torpor. American Journal of Physiology-Regulatory, Integrative and Comparative Physiology. 2010;298(6):R1531. Chung S, Wong T, Nagasaki H, Civelli O. Acute Homeostatic Responses to Increased Ext. Concurption in MCMLIR. Koochout Mice. Journal of Malegular.
- Increased Fat Consumption in MCH1R Knockout Mice. Journal of Molecular Neuroscience. 2010;Online(April 22).
- Church C, Moir L, Mcmurray F, et al. Overexpression of Fto leads to increased food intake and results in obesity. Nature Genetics. 2010;42(November):1086–1092. Cooksey RC, Jones D, Gabrielsen S, et al. Dietary iron restriction or iron chelation protects from diabetes and loss of \$\beta\$-cell function in the obese (ob/ob lep-/-) 6. mouse. American Journal of Physiology-Endocrinology And Metabolism. 2010;298(6):E1236.
- Coppey L, Davidson E, Lu B, Gerard C, Yorek M. Neuropharmacology Vasopeptidase inhibitor ilepatril (AVE7688) prevents obesity- and diabetes-induced neuropathy in C57Bl / 6J mice. Neuropharmacology. 2011;60(February-March):259-266. Deiuliis J, Shah Z, Shah N, et al. Visceral Adipose Inflammation in Obesity Is
- 8 Associated with Critical Alterations in Tregulatory Cell Numbers. Plos ONE. 2011;6(1):e16376.
- Drake AJ, Raubenheimer PJ, Kerrigan D, et al. Prenatal Dexamethasone Programs Expression of Genes in Liver and Adipose Tissue and Increased High-Fat Diet. 9. Endocrinology. 2010;151(April):1581-1587
- Enriori, PJ, Evans, A.E., Sinnayah, P, et al. Diet-induced obesity causes severe but reversible leptin resistance in arcuate melanocortin neurons. Cell Metabolism. 2007;5(3):181–194.
- Eric D, Lawrence C, Bao L, Victor A, Nigel A, others. The Roles of Streptozotocin Neurotoxicity and Neutral Endopeptidase in Murine Experimental Diabetic Neuropathy. Experimental Diabetes Research. 2010;2009(431980).
- 12. Faleck D, Áli K, Roat R, et al. Adipose differentiation-related protein regulates lipids and insulin in pancreatic islets. American Journal of Physiology-Endocrinology And
- Metabolism. 2010;299(2):E249.
 13. Gallou-Kabani C, Gabory A, Al. E. Sex- and Diet-Specific Changes of Imprinted Gene Expression and DNA Methylation in Mouse Placenta under a High-Fat Diet. Plos ONE. 2010;5(12):e14398.
- 14. Gallou-Kabani C, Vigé A, Gross MS, et al. C57BL/6J and A/J Mice Fed a High-Fat Diet Delineate Components of Metabolic Syndrome&ast. Obesity. 2007;15(8):1996-2005.
- 15. Ito, M., Suzuki, J., Tsujioka, S., et al. Longitudinal analysis of murine steatohepatitis model induced by chronic exposure to high-fat diet. Hepatology Research. 2007;37(1):50-57.
- Jelinek D, Heidenreich RA, Erickson RP, Garver WS. Decreased Npc1 gene dosage in mice is associated with weight gain. Obesity. 2009;18(7):1457–1459.
 Jiang T, Wang Z, Proctor G, et al. Diet-induced obesity in C57BL/6J mice causes
- increased renal lipid accumulation and glomerulosclerosis via a sterol regulatory element-binding protein-1c-dependent pathway. Journal of Biological Čhemistry. 2005;280(37):32317.
- Kalupahana NS, Voy BH, Saxton AM, Moustaid-Moussa N. Energy-Restricted High-Fat Diets Only Partially Improve Markers of Systemic and Adipose Tissue Inflammation. Obesity. 2010;19(July 2010):245-254.
- 19. King VL, Hatch NW, Chan HW, et al. A murine model of obesity with accelerated atherosclerosis. Obesity. 2009;18(1):35-41.
- 20. Kusakabe, T, Tanioka, H, Ebihara, K., et al. Beneficial effects of leptin on glycaemic
- Kusakabe, 1, Tanioka, H, Ebinara, K., et al. Beneficial effects of reptin on glycaemic and lipid control in a mouse model of type 2 diabetes with increased adiposity induced by streptozotocin and a high-fat diet. Diabetologia. 2009;52(4):675–683.
 Mu J, Woods J, Zhou Y-ping, et al. Chronic Inhibition of Dipeptidyl Peptidase-4 With a Sitagliptin Analog Preserves Pancreatic beta -Cell Mass and Function in a Rodent Model of Type 2 Diabetes. Diabetes. 2006;55(June):1695-1704.
 Rabot S, Membrez M, Bruneau A, et al. Germ-free C57BL/6J mice are resistant to
- high-fat-diet-induced insulin resistance and have altered cholesterol metabolism. The FASEB Journal. 2010;24(12):4948-4959.
- 23. Raher M, Thibault H, Buys E. A short duration of high-fat diet induces insulin Yaner W, Hubart H, By's L'Arshift editation of negrate det index insulin resistance and predisposes to adverse left ventricular remodeling after pressure overload. American Journal Of Physiology. 2008;295(6):2495-2502.
 Rajala MW, Qi Y, Patel HR, et al. Regulation of resistin expression and circulating levels in obesity, diabetes, and fasting. Diabetes. 2004;53(7):1671.
 Raubenheimer PJ, Nyirenda MJ, Walker BR. A choline-deficient diet exacerbates
- fatty liver but attenuates insulin resistance and glucose intolerance in mice fed a
- high-fat diet. Diabetes. 2006;55(7):2015.
 26. Sutton GM, Trevaskis JL, Hulver MW, et al. Diet-Genotype Interactions in the Development of the Obese, Insulin-Resistant Phenotype of C57BL / 6J Mice Lacking Melanocortin-3 or -4 Receptors. Endocrinology. 2006;147(5):2183-2196.
 27. Townsend KL, Lorenzi MM, Widmaier EP. High-fat diet-induced changes in body
- mass and hypothalamic gene expression in wild-type and leptin-deficient mice. Endocrine. 2008;33(2):176-88
- Trevaskis JL, Meyer EA, Galgani JE, Butler AA. Counterintuitive Effects of Double-Heterozygous Null Melanocortin-4 Receptor and Leptin Genes on Diet-Induced Obesity and Insulin Resistance in C57BL / 6J Mice. Endocrinology. 2008;149(1):174 -184.
- Valker C, Li X, Whiting L, Glyn-Jones S. Mice Lacking the Neuropeptide {alpha}-Calcitonin Gene-Related Peptide Are Protected Against Diet-Induced Obesity, Endocrinology, 2010;151(9):4257-4269.
- 30. Wei P, Lane PH, Lane JT, Padanilam BJ, Sansom SC. Glomerular structural and functional changes in a high-fat diet mouse model of early-stage Type 2 diabetes. Diabetologia. 2004;47(9):1541-9.