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中枢神经系统整合外周信号调节采食量的分子机制

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Molecular Mechanisms of Feed Intake Regulation through Integration of Various Peripheral Signals by Central Nervous System

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摘要 采食量是动物生产性能充分发挥的基石,其受多种因素的共同调节。胃肠调节肽、肥胖信号分子(瘦素和胰岛素)及营养物质等外周调节因子在采食量的调节中发挥重要的作用。中枢神经系统是采食量调节的关键。近年来对中枢神经系统调节采食量的分子机制认识越来越深入,已鉴定出一些关键信号转导通路及转录因子,其中哺乳动物下丘脑雷帕霉素靶蛋白(mTOR)通路、一般性调控阻遏蛋白激酶2(GCN2)介导的一般性氨基酸应答通路及叉头转录因子1(FoxO1)和信号转导与转录激活因子3(STAT3)在中枢的采食量调节中起着重要作用。本文在简要回顾采食量调节生理机制的基础上,主要对外周信号在中枢神经系统调节采食量分子机制的研究进展进行综述。

关键词: 采食量 调节机制 中枢神经系统 信号通路 mTOR GCN2

Abstract: Feed intake is the cornerstone of assuring adequate production performance of animals and a variety of environmental stimuli influences the feed intake. Numerous peripheral stimuli involved gastrointestinal peptides, adiposity signaling molecules (insulin and leptin), and nutrients signal to central nervous system (CNS) have an important physiological role in regulating feed intake. CNS plays the central role in feed intake regulation, while the molecular pathway and mechanism has been increasingly recognized for the past few years. In mammals, it has already identified some key CNS intracellular signal transduction pathways and transcription factors as sensors/integrators in regulating feed intake, such as some highly conserved key pathways—the hypothalamic mammalian target of rapamycin (mTOR) and general control nonderepressible-2 kinase (GCN2)-mediated general amino acid control (GAAC) pathway, and transcription factors involved forkhead transcription factor 1 (FoxO1) and signal transducer and activator of transcription 3 (STAT3). This review provided an overview of the research progress of physiological and CNS molecular mechanisms in the overall control of feed intake.

Keywords: feed intake, regulation mechanisms, central nervous system, signal pathway, mTOR, GCN2

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- [1] CUMMINGS D E, OVERDUIN J. Gastrointestinal regulation of food intake[J]. *Journal of Clinical Investigation*, 2007, 117: 13-23.
- [2] COTA D, PROULX K, BLAKE SMITH K A, et al. Hypothalamic mTOR signaling regulates food intake[J]. *Science*, 2006, 312: 927-930.
- [3] WOODS S C, SEELEY R J, COTA D. Regulation of food intake through hypothalamic signaling networks involving mTOR[J]. *Annual Reviews*, 2008, 28: 295-311.
- [4] HAO S Z, SHARP J W, ROSS-INTA C M, et al. Uncharged tRNA and sensing of amino acid deficiency in mammalian piriform cortex [J]. *Science*, 2005, 307: 1776-1778.
- [5] GIETZEN D W, HAO S Z, ANTHONY T G, et al. Mechanisms of food intake repression in indispensable amino acid deficiency[J]. *Annual Review of Nutrition*, 2007, 27: 63-78.
- [6] BLEVINS J E, SCHWARTZ M W, BASKIN D G. Evidence that paraventricular nucleus oxytocin neurons link hypothalamic leptin action to caudal brainstem nuclei controlling meal size[J]. *American Journal of Physiology Regulatory, Integrative and Comparative Physiology*, 2004, 287: R87-R96.
- [7] KELLEY A E, BALDO B A, PRATT W E, et al. Corticostriatal-hypothalamic circuitry and food motivation: integration of energy, action and reward [J]. *Physiology & Behavior*, 2005, 86: 773-795.
- [8] KELLEY A E, BERRIDGE K C. The neuroscience of natural rewards: relevance to addictive drugs[J]. *The Journal of Neuroscience*, 2002, 22: 3306-3311.
- [9] GERAEDTS M C P, TTOOST F J, SARIS W H M. Gastrointestinal targets to modulate satiety and food intake[J]. *Obesity Reviews*, 2010, 12: 470-477.
- [10] KARHUNEN L J, JUVONEN K R, HUOTARI A, et al. Effect of protein, fat, carbohydrate and fibre on gastrointestinal peptide release in humans [J]. *Regulatory Peptides*, 2008, 149: 70-78.
- [11] HAHN T M, BREININGER J F, BASKIN D G, et al. Coexpression of *Agrp* and *NPY* in fasting-activated hypothalamic neurons[J]. *Nature Neuroscience*, 1998, 1: 271-272.
- [12] CONE R D. Anatomy and regulation of the central melanocortin system[J]. *Nature Neuroscience*, 2005, 8: 571-578.
- [13] FAN W, BOSTON B A, KESTERSON R A, et al. Role of melanocortinergic neurons in feeding and the agouti obesity syndrome [J]. *Nature*, 1997, 385: 165-168.
- [14] PORTE D Jr., BASKIN D G, SCHWARTZ M W. Leptin and insulin action in the central nervous system[J]. *Nutrition Reviews*, 2003, 60: S20-S29.
- [15] MURPHY K G, DHILLO W S, BLOOM S R. Gut peptides in the regulation of food intake and energy homeostasis[J]. *Endocrine Reviews*, 2006, 27 (7): 719-727.
- [16] VAN DEN HEIDE L P, RAMAKERS M J, SMIDT M P. Insulin signaling in the central nervous system: learning to survive[J]. *Progress in Neurobiology*, 2006, 79: 205-221.
- [17] PARDRIDGE W M, EISENBERG J, YANG J. Human blood-brain barrier insulin receptor[J]. *Journal of Neurochemistry*, 1985, 44: 1771-1778.
- [18] HAVEL P J. Peripheral signals conveying metabolic information to the brain: short-term and long-term regulation of food intake and energy homeostasis[J]. *Experimental Biology and Medicine*, 2001, 226(11): 963-977.
- [19] AIR E L, BENOIT S C, BLAKE SMITH K A, et al. Acute third ventricular administration of insulin decreases food intake in two paradigms [J]. *Pharmacology Biochemistry and Behavior*, 2002, 72: 423-429.
- [20] OBICI S, FENG Z H, KARKANIAS G, et al. Decreasing hypothalamic insulin receptors causes hyperphagia and insulin resistance in rats[J]. *Nature Neuroscience*, 2002, 5: 566-572.
- [21] FLIER J S. Obesity wars: molecular progress confronts an expanding epidemic[J]. *Cell*, 2004, 116: 337-350.
- [22] HAVEL P J, OWNSEND R, CHAUMP L, et al. High-fat meals reduce 24-h circulating leptin concentrations in women[J]. *Diabetes*, 1999, 48: 334-341.
- [23] KORBONITS M, TRAINER P J, LITTLE J A, et al. Leptin levels do not change acutely with food administration in normal or obese subjects, but are negatively correlated with pituitary-adrenal activity[J]. *Clinical Endocrinology*, 1997, 46: 751-757.
- [24] BANKS W A, KASTIN A J, HUANG W, et al. Leptin enters the brain by a saturable system independent of insulin[J]. *Peptides*, 1996, 17: 305-311.
- [25] FEI H, OKANO H J, LI C, et al. Anatomic localization of alternatively spliced leptin receptors (Ob-R) in mouse brain and other tissues [J]. *Proceedings of the National Academy of Sciences of the United States of America*, 1997, 94: 7001-7005.
- [26] CHEN H, CHARLAT O, TARTAGLIA L A, et al. Evidence that the diabetes gene encodes the leptin receptor: identification of a mutation in the leptin receptor gene in *db/db* mice[J]. *Cell*, 1996, 84: 491-495.
- [27] SCHWARTZ M W, WOODS S C, PORTE D, et al. Central nervous system control of food intake[J]. *Nature*, 2000, 404: 661-671.
- [28] LENARD N R, BERTHOUD H. Central and peripheral regulation of food intake and physical activity: pathways and genes[J]. *Obesity*, 2008, 16: S11-S22.
- [29] OBICI S, FENG Z H, MORGAN K, et al. Central administration of oleic acid inhibits glucose production and food intake[J]. *Diabetes*, 2002, 51: 271-275.
- [30] HARDIE D G, CARLING D. The AMP-activated protein kinase-fuel gauge of the mammalian cell?[J]. *European Journal of Biochemistry*, 1997, 246: 259-273.
- [31] WULLSCHLEGER S, LOEWITH R, HALL M N. TOR signaling in growth and metabolism[J]. *Cell*, 2006, 124: 471-484.

- [32] MINOKOSHI Y,ALQUIER T,FURUKAWA N,et al.AMP-kinase regulates food intake by responding to hormonal and nutrient signals in the hypothalamus[J].Nature,2004,428: 569-574.
- [33] MORRISON C D,XI X C,WHITE C L,et al.Amino acids inhibit *Agrp* gene expression via an mTOR-dependent mechanism[J].American Journal of Physiology Endocrinology and Metabolism,2007,293: E165-E171.
- [34] ZONCU R,EFEYAN A,SABATINI D M.mTOR:from growth signal integration to cancer,diabetes and ageing[J].Nature Reviews Molecular Cell Biology,2011,12: 21-35.
- [35] LAPLANTE M,SABATINI D W.mTOR signaling in growth control and disease[J].Cell,2012,149: 274-293.
- [36] WU Q,ZHANG Y,XU J,et al.Regulation of hunger-driven behaviors by neural ribosomal S6 kinase in *Drosophila*[J].Proceedings of the National Academy of Sciences of the United States of America,2005,102: 13289-13294.
- [37] DAGON Y,HUR E,ZHENG B,et al.p70S6 kinase phosphorylates AMPK on serine 491 to mediate leptin's effect on food intake[J].Cell Metabolism,2012,16: 104-112.
- [38] SHARP J W,ROSS-INTA C M,HAO S Z,et al.Co-localization of phosphorylated extracellular signal-regulated protein kinases 1/2 (ERK1/2) and phosphorylated eukaryotic initiation factor 2 α (eIF2 α) in response to a threonine-devoid diet[J].Journal of Comparative Neurology,2006,494: 485-494.
- [39] SOOD R,PORTER A C,OLSEN D,et al.A mammalian homologue of GCN2 protein kinase important for translational control by phosphorylation of eukaryotic initiation factor-2 α [J].Genetics,2000,154: 787-801.
- [40] ZHANG P,MCGRATH BC,REINERT J,et al.The GCN2 eIF2 α kinase is required for adaptation to amino acid deprivation in mice[J].Molecular and Cellular Biology,2002,22: 6681-6688.
- [41] DONG J S,QIU H F,GARCIA-BARRIO M,et al.Uncharged tRNA activates GCN2 by displacing the protein kinase moiety from a bipartite tRNA-binding domain[J].Molecular Cell,2000,6: 269-279.
- [42] BLAIS A,HUNEAU J F,MAGRUM L J,et al.Threonine deprivation rapidly activates the system A amino acid transporter in primary cultures of rat neurons from the essential amino acid sensor in the anterior piriform cortex[J].The Journal of Nutrition,2003,133: 2156-2164.
- [43] HUTSON S M,LIETH E,LANOUE K F.Function of leucine in excitatory neurotransmitter metabolism in the central nervous system[J].The Journal of Nutrition,2001,131: 846S-850S.
- [44] SHARP J W,ROSS C M,KOEHNLE T J,et al.Phosphorylation of Ca²⁺/calmodulin dependent protein kinase type II and the α -amino-3-hydroxy-5-methyl-4-isoxazole propionate (AMPA) receptor in response to a threonine-devoid diet[J].Neuroscience,2004,126: 1053-62.
- [45] WEK R C,JIANG H Y,ANTHONY T G.Coping with stress:eIF2 kinases and translational control[J].Biochemical Society Transactions,2006,34: 7-11.
- [46] KITAMURA T,FENG Y,KITAMURA I Y I,et al.Forkhead protein FoxO1 mediates *Agrp*-dependent effects of leptin on food intake[J].Nature Medicine,2006,12: 534-540.
- [47] PLUM L,BELGARDT B F,BRUNING J C.Central insulin action in energy and glucose homeostasis[J].Journal of Clinical Investigation,2006,116: 1761-66.
- [48] BATES S H,STEARNS W H,DUNDON T A,et al.STAT3 signalling is required for leptin regulation of energy balance but not reproduction [J].Nature,2003,421: 856-859.
- [1] 柯丹霞,印遇龙.小G蛋白对mTOR信号通路的调控[J].动物营养学报,2013,25(8): 1663-1670
- [2] 刘南南,姚军虎.营养素和激素对乳蛋白合成过程中哺乳动物雷帕霉素靶蛋白信号通路调节作用的研究进展[J].动物营养学报,2013,25(6): 1158-1163
- [3] 朱宇旌,李艳,张勇,于姣姣,邵彩梅.转录因子E2相关因子2-抗氧化反应元件信号通路与机体抗氧化的关系[J].动物营养学报,2013,25(3): 458-463
- [4] 丁耿芝,孟庆翔.反刍动物干物质采食量预测模型研究进展[J].动物营养学报,2013,25(2): 248-255
- [5] 王钰飞,丁丽敏,付京杰,张文珍,王建梅.不同蛋白质来源饲料对不同生长阶段藏獒营养物质表观消化率及粪便质量的影响[J].动物营养学报,2013,25(10): 2345-2354
- [6] 栾兆双,宋娟,胡彩虹.丝裂原活化蛋白激酶信号通路抑制剂对断奶仔猪小肠形态和肠通透性的影响[J].动物营养学报,2013,25(1): 44-49
- [7] 张万金,李胜利,史海涛,骆雅萍,杨军香,尹义江,贾春涛.饲料添加不同种类的糖蜜对夏季热应激奶牛采食量和产奶性能的影响[J].动物营养学报,2013,25(1): 163-170
- [8] 马涛,刁其玉,邓凯东,姜成钢,屠焰,王永超,刘洁,赵一广.饲料不同采食水平下肉羊氮沉积和尿中嘌呤衍生物排出规律的研究 [J].动物营养学报,2012,24(7): 1229-1235
- [9] 杨静,谢明,侯水生,黄苇,喻俊英,汪超.精氨酸调控畜禽采食量的机制及其影响因素[J].动物营养学报,2012,24(4): 612-616
- [10] 李玉莲,田启宇,范志勇,肖勇,陈永辉,杨泰,董莲花.免疫调控技术在畜禽采食量调控中的应用[J].动物营养学报,2012,(11): 2092-2096
- [11] 栾兆双,姚丽丽,傅振宇,宋娟,胡彩虹.断奶应激对仔猪肠形态、肠黏膜屏障和p38丝裂原活化蛋白激酶信号通路的影响[J].动物营养学报,2012,(11): 2237-2242
- [12] 罗佳捷,陈宇光,张彬,李丽立,杨永生.脂联素受体表达的调节机制[J].动物营养学报,2011,23(10): 1651-1657
- [13] 王坤坤,曹崇海,孙建义,刘建新,钱利纯.锌和降钙素基因相关肽对仔猪摄食的影响及其机制研究[J].动物营养学报,2011,23(09): 1545-1552
- [14] 王改英,吴在富,杨维仁,胥保华.饲料蛋白质水平对意大利蜜蜂咽下腺发育及产浆量的影响[J].动物营养学报,2011,23(07): 1147-1152
- [15] 吕继蓉,曾凡坤,张克英.猪乳香味剂对断奶仔猪采食行为和采食量的影响[J].动物营养学报,2011,23(05): 848-853

