

Table 1. Methods for increasing BDNF levels in mammals and their beneficial effects.

1. Exogenous injection of BDNF				
Research	Material	Methods	Effect on serum BDNF	Other effects
Ono M et al [20]	mice	Subcutaneous administration of BDNF	No data	-decrease of elevated blood glucose level,
Mitsugu et al. [23]	obese diabetic mice	subcutaneous administration of BDNF to obese diabetic mice	No data	-decrease glucose concentration, -decrease concentration of pancreatic glucagon - increase concentration of pancreatic insulin, -increase beta cell producing insulin in pancreatic islet, -decrease non-producing insulin cells area
Kuroda et al. [24]	obese rats	administration of exogenous BDNF	No data	-improve glucokinase activity, -decrease hepatic gluconeogenesis, -improve insulin sensitivity, -decrease level of fasting glucose and postprandial blood glucose -decrease hyperinsulinemia
Meek et al [25]	mice	intracerebroventricular or into the ventromedial hypothalamic nucleus injections of either BDNF or its vehicle	No data	- decrease hyperglycemia, -decrease blood glucose levels -suppress hepatic glucose production through inhibition of glucagone
Masaaki Seki et al.[38]	streptozotocin-induced diabetic rats	intraocular administration of BDNF	No data	-prevention dopaminergic amacrine cells from degeneration -potential role in the treatment of early retinal neuropathy
Lei li et al. [40]	diabetic rats	continued intrathecal administration of BDNF	No data	-alleviation of mechanical and thermal hyperalgesia -reduce hyperexcitability of dorsal root ganglion neurons -potential role in the treatment of painful diabetic neuropathy

2. Dietary restrictions				
Research	Material	Methods	Effect on BDNF	Other effects
Duan et al. [22]	Mice	dietetary restrictions	increase BDNF production in brain cells	decrease concentration of glucose, insulin, and leptin in mice whose level of these factors was increased
3. Physical activity				
Research	Material	Methods	Effect on BDNF	Other effects
Babaei et al. [11]	well-trained individuals and people living a sedentary lifestyle	Comparison of well-trained individuals and people living a sedentary lifestyle aerobic and anaerobic exercises	-basic BDNF concentration in serum is significantly lower in the well-trained group than in the control group -In both groups aerobic and anaerobic exercises increased BDNF level about its initial value	Well-trained people had better results in a picture recall memory test.
Lee et al. [29]	adolescents with type 2 diabetes mellitus	12 weeks aerobic exercise	-adolescents with type 2 diabetes mellitus and control group had similar resting BDNF levels. -after the 12 weeks aerobic exercise,, there was no significant increase in the resting BDNF levels in the patients with type 2 diabetes mellitus group	
Tonoli et al. [31]	patients with type 1 diabetes	high-intensity exercise	-serum BDNF levels were significantly higher in people with type 1 diabetes compared with the control healthy group. -in both groups BDNF increased after exercise.	
Cancela JM et al. [43]	patients with dementia	riding a bicycle for minimum 15 minutes instead of taking part in recreational activities		-improved memory -improved mobility

Rasoul Eslami et al. [44]	rats	endurance training	-diabetes reduce the expression of BDNF in the sensory and motor roots. -this phenomenon can be reversed through endurance training.	
Tang et al. [45]	rats	weight-bearing ladder and aerobic treadmill exercise	upregulate expression of BDNF and CREB in hippocampus	- learning ability increased
Floel et al. [46]	healthy patients	comparison of patients according to physical activity, measured with physical activity questionnaire, parameters of aerobic exercises, or both factors	-physically active patients had higher concentration of neurotrophin – a Granulocyte-Colony Stimulating Factor (G-CSF) -physical activity level was not significantly correlated with the BDNF level	- Physically active patients showed increased grey matter volume
Hyun-chul Cho et al. [47]	healthy patients	treadmill VO2max performance	-increase in BDNF concentration in serum, plasma and platelets immediately after exercises	
Tonoli et al. [48]	patients with type 1 diabetes	two different intensities of exercises (high-intensity and continuous medium-intensity)	-BDNF levels increased significantly after both exercise intensities. The BDNF increase had a dose-response effect for exercise intensity	
Brinkmann et al. [49]	elderly people with type 2 diabetes	examine the level of neurotrophic factors before and after exercise	-depending on the exercise mode, acute submaximal exercise can increase levels of neurotrophic factors (BDNF, VEGF)	