The Clinical Utility of Serum Brain Derived Neurotrophic Factor in Bipolar Affective Disorder

Thesis

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List of Abbreviations

AD : Atopic dermatitis

ADHD : attention deficit hyperactive disorder

AHR : Airway hyper-reactivity **BAD** : Bipolar affective disorder

Bcl2 : B cell lymphoma 2
BD : Bipolar disorder

BDNF : Brain derived neurotrophic factor

BD-NOS : Bipolar disorder not other wised specified CaMK : Calcium/calmodulin-dependent protein kinase

CBP : CREB binding protein
 CE : Capillary electrophoresis
 CNS : Cental nervous system
 CNTF : Ciliary neurotrophic factor

CPE : Carboxypeptidase E

CREB : cAMP response element-binding protein

CSF : Cerebrospinal fluid **DAG** : Diacyl glycerol

DD-NOS : Depressive disorder not other wised specified

DEP : Diesel exhausts particulatesDNMT : DNA methyl transferase

DSM IV TR: Diagnostic and Statistical Manual of Mental Disorder

text revision

EAE : Experimental autoimmune encephalomyelitis

ECT : Electroconvulsive therapy
ELISA : Enzyme linked immunoassays

ERK : Extracellular receptor-coupled kinase FEV1 : Forced expiratory volume in 1 second

FGF : Fibroblast growth factor

FGFR : Fibroblast growth factor receptor

GABA : Gamma amino butyric acid HAT : Histone acetyl transferase

HCV : Hepatitis C virusHDACs : Histone deacetylases

HMTs : Histone methyl transferases

List of Abbreviations (Cont...)

ICE : Immunoaffinity capillary electrophoresis

IFN γ : Interferon γ

IHC : Immunohistochemistry

IL : Interleukin

JCA : Juvenile chronic arthritis
JNK : Jun N terminal kinase
KD : Kawasaki disease

Kda : Kilo Dalton

LDL : Low-density lipoproteinLIF : Laser induced fluorescenceMabs : Monoclonal antibodies

MAPK : Mitogen-activated protein kinase

MDD : Major depressive disorderMeCP2 : Methyl CPG binding protein 2

MEK : Methyl ethyl ketone

Met : Methionine

MRI : magnetic resonance imaging

MS : Multiple sclerosis

MSP : Methylation specific PCR

NAc : Nucleus accumbens

NAD : Nicotinamine adenine dinucleotide

NF-KB : Nuclear factor kappa B
NGF : Nerve growth factor
NGFR : Nerve growth factor
NMDA : N-methyl-D-aspartate

NT-3 : Neurotrophin-3 NT-4/5 : Neurotrophin 4/5

PBS : Phosphate buffer saline

PC : Proconvertase
PFC : Prefrontal cortex

PI3K : Phosphatidylinositol 3- kinase

PKC : Protein kinase CPLC : Phospholipase C

PMD : Psychotic major depression

List of Abbreviations (Cont...)

Poly A : Polyadenylation

PPD: Postpartum depression

Pro NT: Pro neurotrophin

PTSD : Post traumatic stress disorder

QM-MSP: Quantitative multiplex methylation specific PCR

Q-MSP : Quantitative methylation specific PCR

RA : Rheumatoid arthritis

RBD : recurrent brief depression

RIA : Radioimmunoassays ROIs : Regions of interest

RT-PCR : Reverse transcription polymerase chain reaction

SAD : seasonal affective disorder

SLE : Systemic lupus erythematosusTGFβ : Transforming growth factor beta

Th : T helper

Trk : Tyrosine kinase

Val : Valine

VEGF : Vascular endothelial growth factor

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Introduction

Bipolar affective disorder, historically known as manic—depressive disorder, is a psychiatric diagnosis that describes a category of mood disorders defined by the presence of one or more episodes of abnormally elevated energy levels, cognition, and mood with or without one or more depressive episodes (*Blumer*, 2002). The elevated moods are clinically referred to as mania or, if milder, hypomania. Individuals who experience manic episodes also commonly experience depressive episodes, or symptoms, or a mixed state in which features of both mania and depression are present at the same time (*Basco and Ramirez*, 2006). These events are usually separated by periods of "normal" mood; but, in some individuals, depression and mania may rapidly alternate (*Yatham and Lakshmi*, 2010).

Bipolar disorder can cause suicidal ideation that leads to suicidal attempts (*Novick et al.*, *2010*). The annual average suicide rate is 0.4%, which is 10 to 20 times that of the general population (*Benjamin et al.*, *2010*).

Neurotrophins are a family of proteins that belong to a class of growth factors, which are capable of signaling particular cells to survive, differentiate, or grow (*Allen and Dawbarn, 2006*). These growth factors are known as neurotrophic factors. They are secreted by target tissue and act by preventing the associated neuron from initiating programmed cell death; thus allowing the neurons to survive (*Hempstead, 2006 and Reichardt, 2006*).