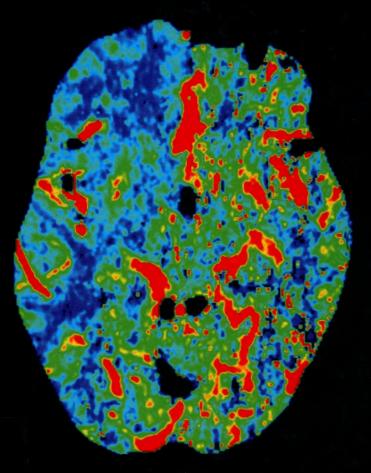
Hankey's CLINICAL Second Edition NEUROLOGY





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PREFACE

A DECADE HAS PASSED since the first edition of Clinical Neurology. Those who have embraced it have encouraged us to update it. The explosion of rigorous scientific evidence for interventions in clinical neurology, coupled with astonishing advances in the clinical neurosciences, have further inspired us to undertake a second edition. As the initial authors (GJH and JMW) are now a decade older and have gravitated toward greater subspecialization, another couple of fellow enthusiasts (PBG and FT) from Grand Rapids and Chicago, USA have joined to facilitate a re-energized, comprehensive, and more global, rather than Anglo-Australian, effort. Together we have enlisted the generosity and specialist expertise of our friends and colleagues throughout the world who are recognized leaders in their field and who have kindly agreed to enlighten us with a chapter on the subject to which they are dedicated.

The subjects and format of the first edition have been maintained and are complemented by the addition of a new chapter on sleep disorders. The chapter covering degenerative diseases of the nervous system has now been subdivided into three main sections, dementias, Parkinson's disease and parkinsonian syndromes, and hereditary ataxias. The cranial neuropathies chapter now includes an entirely new section on neuro-ophthalmology. In addition there are over 440 new illustrations.

The perspective for each chapter is also fresh, as each chapter (with the exception of the chapter on stroke) has been written by one or more of our new contributors, in contrast to the first edition which represented the perspective of GJH and JMW. The purpose of the book, nevertheless, continues to focus on the essentials for students of clinical neurology, particularly neurologists-in-training and practicing neurologists, who wish to have ready access to a comprehensive, up-to-date, and evidence-based guide to the understanding, diagnosis, and management of common and important neurologic disorders.

Many of the illustrations are images taken from our own patients, whom we would like to thank for allowing us to photograph them or the outcome of their investigations. Furthermore, we would also like to thank all the current and past contributors of figures (too many to list individually here) for providing illustrations, as indicated throughout the book. Finally, we would like to thank our families and colleagues for supporting us in this endeavor. We hope you enjoy it and we welcome any comments and criticisms.

Graeme J. Hankey Joanna M. Wardlaw Philip B. Gorelick Fernando D. Testai

DEDICATIONS

I dedicate this book in honor of Mr. Ralph Hauenstein for service to his country and his many generous commitments to the neuroscience programs at Saint Mary's Health Care and the Western Michigan area, and to Sister Myra Bergman for her dedication, devotion and spirited work as a missionary and religious leader in our region and beyond.

Philip B. Gorelick

To my wife, Flavia, for her love, patience, and endless support; to our beautiful children, Sofia and Martin, for being continuous examples of enthusiasm and dedication; to my parents, Ruben and Stella, and sisters, Alejandra and Naiara, for their motivation and support throughout the years; to our neurology residents for having chosen one of the most amazing paths in the medical sciences; and foremost, to our most brilliant mentors – our patients.

Fernando D. Testai

To the memory of my father, the late Dr. John Hankey
Graeme J. Hankey

I am grateful to my family for ongoing support and so dedicate this work to them.

Ioanna M. Wardlaw

ABBREVIATIONS

5HT	5-hydroxytryptamine		ARSA	arylsulfatase A
AA	anaplastic astrocytoma		ARSACS	autosomal recessive spastic ataxia of Charlevoix-
AAD				Saguenay
AASM	American Association of Sleep Medicine		ARX	Aristaless-related homeobox
Αβ			ASA	atrial septal aneurysm
Aca	aceruloplasminemia		ASD	atrial septal defect
ACE	angiotensin-converting enzyme		AST	aspartate aminotransferase
ACE-R	Addenbrooke's Cognitive Examination Revised	1/1	AT	ataxia telangiectasia
AChR	acetylcholine receptors		ATM	acute transverse myelitis
ACTH			AVM	arteriovenous malformation
AD			AVS	acute vestibular syndrome
ADAMTS	a disintegrin and metalloprotease with thrombo		AZA	azathioprine
HSP2	spondin motif	THE	BAEP	brainstem auditory evoked potential
ADC			BAL	British anti-Lewisite
ADCA	autosomal dominant cerebellar ataxia		BBS	Bardet-Biedl syndrome
ADEM	acute disseminated encephalomyelitis		BDNGF	brain-derived nerve growth factor
ADHD			BF	blood flow
ADL			bFGF	basic fibroblast growth factor
ADLP	adrenoleukodystrophy protein		BHC	benign hereditary chorea
ADP			BMD	
AED	antiepileptic drug		BMI	Becker's muscular dystrophy
AF				body mass index
AFB			BNCT	boron neutron capture therapy
AFP			BP	blood pressure
	I I		BPAP	bilevel positive airways pressure
AHI	71 71		BPPV	benign paroxysmal positional vertigo
AICA	,		BSE	bovine spongiform encephalopathy
AIDP	acute inflammatory demyelinating polyradiculo		BSK	Barbour-Stoenner-Kelly
ATDC	Processor Processor		BWSTT	body weight supported treadmill training
AIDS			CADASIL	cerebral autosomal dominant arteriopathy with
AION	T		0111	subcortical infarcts and leukoencephalopthy
AIP	acute intermittent porphyria		CAM	computer assisted myelography
ALD	adrenoleukodystrophy		CAS	carotid artery stenting
ALDP	, , , ,		CBD	corticobasal degeneration
ALS	amyotrophic lateral sclerosis		CBF	cerebral blood flow
ALT	alanine aminotransferase		CBS	corticobasal syndrome/cystathionine β-synthase
AMAN	acute motor axonal neuropathy			deficiency
AML	angiomyolipoma		CBT	cognitive behavioral therapy
AMN			CDC	Centers for Disease Control and Prevention
(c)AMP			CEA	carotid endarterectomy/carcinoembryonic antigen
AMSAN			cEEG	continuous electroencephalography
ANCL	adult neuronal ceroid lipofuscinoses		CE-MRA	contrast-enhanced magnetic resonance
AO	anaplastic oligodendroglioma			angiography
AOA	oligoastrocytoma		CGRP	calcitonin gene-related peptide
ApoE	apolipoprotein E		CI	confidence interval/cholinesterase inhibitor
APP	amyloid precursor protein		CIDP	chronic inflammatory demyelinating poly-
APS	antiphospholipid syndrome			neuropathy
aPTT			CIM	critical illness myopathy
ARAS	ascending reticular activating system		CIMT	constraint-induced movement therapy
ARDS			CIP	critical illness polyneuropathy
ARI			CIS	clinically isolated syndrome
ARR			CISC	clean intermittent self-catheterization

(f/i/s/v)CJI	O (familial/iatrogenic/sporadic/variant) Creutzfeldt– Jakob disease	EITB ELISA	enzyme-linked immunoelectrotransfer blot assay enzyme-linked immunosorbent assay
CK	creatine kinase	EM	erythema migrans
CLAM	cholesterol-lowering agent	EMD	Emery–Dreifuss muscular dystrophy
CM	congenital myopathy	EMG	electromyography
CMAP	compound muscle action potential	EOG	electro-oculogram
CMD	congenital muscular dystrophy	EPP	endplate potential
CMT	Charcot–Marie–Tooth disease	EPT	enhanced physiologic tremor
CMV	cytomegalovirus	ER	extended-release
CNS	central nervous system	ERG	electroretinography
COACH	cerebellar vermis hypo/aplasia, oligophrenia, ataxia	ESR	erythrocyte sedimentation rate
CONCII	congenital, coloboma, and hepatic fibrosis	ESRD	end-stage renal disease
COMT	catechol-O-methyltransferase	ET	essential tremor
CORS	cerebello-oculo-renal syndrome	EV	Eustachian valve
COX	cyclo-oxygenase	EVD	extraventricular drain
CPA	cerebello-pontine angle	FA	Friedreich's ataxia
CPAP	continuous positive airway pressure	FAST	
CPK		FDA	Functional Assessment Staging Test
CPM	creatine phosphokinase		Food and Drug Administration
	central pontine myelinolysis	FES	functional electrical stimulation
CPP	cerebral perfusion pressure	FFI	fatal familial insomnia
Cr CRAO	creatinine	FHM	familial hemiplegic migraine
	central retinal artery occlusion	FIESTA	fast imaging employing steady state acquisition
CRP	C-reactive protein	ET A ID	sequence Management and Management a
CRVO	central retinal vein occlusion	FLAIR	fluid attenuated inversion recovery
CS	Cowden's syndrome	FMD	fibromuscular dysplasia
CSA	central sleep apnea	FSHD	facioscapulohumeral muscular dystrophy
CSF	cerebrospinal fluid	FTA	fluorescent treponemal antibody
CT	computed tomography	FTD	frontotemporal dementia
CTA	computed tomography angiography	FTLD	frontotemporal lobar degeneration
CTV	computed tomography venography	FVC	forced vital capacity
CV	color vision HAMA	FXTAS	fragile X-associated tremor/ataxia syndrome
CVA	cerebrovascular accident	GABA	gamma-aminobutyric acid
CVT	cerebral venous thrombosis	GAD	glutamic acid decarboxylase
DAI	diffuse axonal injury	GALC	galactocerebrosidase
DALY	disability-adjusted life year	GBM	glioblastoma multiforme
DBS	deep-brain stimulation	GBS	group B streptococci/Guillain-Barré syndrome
DFA	direct immunofluorescent antibody	GCI	glial cytoplasmic inclusion
DGC	dystrophin glycoprotein complex	GCS	Glasgow Coma Scale
DIC	disseminated intravascular coagulation	GCSE	generalized convulsive status epilepticus
DLB	dementia with Lewy bodies	GCT	undifferentiated germinoma
DMD	Duchenne's muscular dystrophy	GFR	glomeruler filtration rate
DNA	deoxyribonucleic acid	Glut 1	glucose transporter type 1 (deficiency)
DNET	dysembryoplastic neuroepithelial tumor	GMP	guanosine monophosphate
DRPLA	dentato-rubro-pallido-luysian atrophy	GPi	globus pallidus internus
DSA	digital subtraction cerebral angiography	GSS	Gerstmann–Straüssler–Scheinker syndrome
DSPN	distal symmetric polyneuropathy	GTN	glyceryl trinitrate
DUB	deubiquitinating enzyme	GTP	guanosine triphosphate
DVT	deep vein thrombosis	H&E	hematoxylin and eosin
DWI	diffusion-weighted imaging	HAART	highly-active antiretroviral therapy
EACA	epsilon-aminocaproic acid	HAM/ISP	HTLV-associated myelopathy/tropical spastic
EBRT	external beam radiation therapy	******	paraparesis Introducido de dor
EBV	Epstein–Barr virus	HANAC	hereditary angiopathy, nephropathy, aneurysm,
ECG	electrocardiogram/electrocardiography	TTIPE	and muscle cramps
ECT	electroconvulsive therapy	HARP	hypoprebetalipoproteinemia, acanthocytes,
EDH	extradural hematoma	LICE	retinitis pigmentosa, pallidal degeneration
EEG	electroencephalography	HCD	hepatocerebral degeneration
EGF(R)	epidermal growth factor (receptor)	HCG	human chorionic gonadotropin
EIAC	enzyme-inducing anticonvulsant	HCP	hereditary coproporphyria

HDL Huntington's disease HDL Huntington's disease-like HE hepatic encephalopathy HELLP hemolysis, elevated liver enzymes, low-platelet count syndrome HHT hereditary hemorrhagic telangiectasia (Osler- Rendu-Weber syndrome) HHT hereditary inclusion body myopathy HHF hypoxia-inducible factor HIF horizontal head impulse test HIF horizontal head impulse test HIV human immunoedheiciency virus HAA human leukocyte antigen HNP hereditary neuropathy with liability to pressure palsies HPE holoprosencephaly HR hazard rationheart rate HIRG human rabies immune globulin HSP hereditary spastic paraparasis HSV heres simplex virus MAPT HTIG human Tetanus immune globulin HTIG human retanus immune globulin HTIG human retanus immune globulin HTILV human Flymphotropic virus MAPT MyperPP hypoPP hypokalemic periodic paralysis MCP HTILV human retanus immune globulin HSP inclusion body myositis BPN immune-mediated brachial plexus neuropathy HTICA interacrial alressure ICCI intracranial pressure ICRA intracranial hypertension III interleukin II.AE International League against Epilepsy III.AE International League against Epilepsy III.AE International League against Epilepsy III.AE International normalized ratio INN intracribatic and market Examination INNE intervenican on MNE III.AE International normalized ratio INNE intervenican on MNE III.AE International	
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IPC intermittent pneumatic compression MND motor neuron disease	
IRIS immune reconstitution inflammatory syndrome MOTSA multiple overlapping thin-slab acqu	
IVIG intravenous immune globulin MPR multi-planar reformat	
JCV John Cunningham virus MRA magnetic resonance angiography	
JME juvenile myoclonic epilepsy MRI magnetic resonance imaging	
KBS Klüver–Bucy syndrome mRS Modified Rankin Score	
KD Krabbe disease MRSA methicillin-resistant Staphylococcus	
KSS Kearns–Sayre syndrome MRV magnetic resonance venography	
LAA left atrial appendage MS multiple sclerosis	
LCMV lymphocytic choriomeningitis virus MSA multiple system atrophy	
LD Lhermitte–Duclos disease MSLT Multiple Sleep Latency Test	

MSM	men who have sex with men	PCD)	paraneoplastic cerebellar degeneration
MSPNST	malignant peripheral nerve sheath tumor	PCN	ISL	primary CNS lymphoma
MTHFR	methylenetetrahydrofolate reductase	PCo	m	posterior communicating artery
MTR	methionine synthase	PCR	Dielet	polymerase chain reaction
MUP	motor unit action potential	PCV		vincristine
MuSK	muscle-specific receptor tyrosine kinase	PD		Parkinson's disease
MUT	methylmalonyl-CoA mutase	PDD)	Parkinson's disease dementia
MWT	Maintenance of Wakefulness Test	PDC	GF .	platelet-derived growth factor
MZ	monozygotic/marginal zone	PDV	V	proton density-weighted
NAAT	nucleic acid amplication testing	PE		plasma exchange
NAc	neuroacanthocytosis	PEG		percutaneous endoscopic gastrostomy
NAD	nicotinamide adenine dinucleotide	PEN	1	paraneoplastic encephalomyelitis
NADP	nicotinamide adenine dinucleotide phosphate	PEC)	progressive external ophthalmoplegia
NAION	nonarteritic anterior ischemic optic neuropathy	PET		positron emission tomography
NARP	neurogenic weakness with ataxia and retinitis	PFK		phosphofructokinase
	pigmentosa	PFO		patent foramen ovale
NBIA	neurodegeneration with brain iron accumulation	n PIO	N	posterior ischemic optic neuropathy
NCL	neuronal ceroid lipofuscinosis	Pi-T	ON	posterior indirect traumatic optic neuropathy
NCS	nerve conduction studies	PKA	N	pantothenate kinase-associated neurodegeneration
NCSE	nonconvulsive status epilepticus	PLE	D	periodic lateralized epileptiform discharge
NDT	B. C.	PLE	X	plasmapheresis
NF		PLN	1	periodic leg movement
NF-1	neurofibromatosis 1	PLN	ID	periodic leg movement disorder
NFT	neurofibrillary tangle	PMA	A	progressive myoclonic ataxia
NFG		PMI	Ξ	progressive myoclonic epilepsy
NFLE		PMI		progressive multi-focal leukoencephalopathy
NGGCT	nongerminoma			polymorphonuclear
NHL	non-Hodgkin's lymphoma	PMz	D	Pelizeaus–Merzbacher disease
NIF	negative inspiratory pressure	PNE		primitive neuroectodermal tumor
NIH-SS	National Institutes of Health Stroke Scale	PNF		progressive nonfluent aphasia
NIID	neuronal intranuclear inclusion disease	POC	CI	posterior circulation infarct
NMDA	N-methyl-D-aspartate	POC	CS	posterior circulation syndrome
NMJ	neuromuscular junction	POE	MS	polyneuropathy, organomegaly, endocrinopathies,
NMO				M-protein, skin changes including thickening and
NMS	neuroleptic malignant syndrome			hyperpigmentation, clubbing of the fingers
NO	nitrous oxide	POS	T	positive occipital sharp transients of sleep
NPC	Niemann–Pick type C	POV	/L	postoperative visual loss
NPH	normal pressure hydrocephalus			preplate zone/perfusion pressure
NPHP		PPA		primary progressive aphasia
NSE	neuron-specific enolase	PPR		paremedian pontine reticular formation
NTD		PRE		posterior reversible encephalopathy syndrome
O-AA		PRO		pontine respiratory group
OAA		PrP		prion protein
OCD		PSN		paraneoplastic sensory neuropathy
ONH		PSP		progressive supranuclear palsy
OP	*	PSV		peak systolic velocity
OPCA		PSW		periodic sharp wave complex
OPV		PTH		parathyroid hormone
OR	odds ratio			post-traumatic stress disorder
OSA		PWI		perfusion-weighted imaging
OTR		PXA		pleomorphic xanthoastrocytoma
PA		PXE		pseudoxanthoma elasticum
PACI		RBC		red blood cell
PACS		RCT		randomized controlled trial
PAF		RCV		reversible cerebral vasoconstriction syndrome
PAM		RDI		respiratory disturbance index
PAS		REN		rapid eye movement
PC		RER		respiratory event-related arousal

	RF	resistance to flow	TGF	transforming growth factor
	rFVIIa	recombinant activated factor VII	THB	tetrahydrobiopterin
	RLS	right-to-left shunt/restless legs syndrome	TIA	transient ischemic attack
	RMSF	Rocky Mountain spotted fever	TMJ	temporomandibular joint
	RNA	ribonucleic acid	TMP-SM>	K trimethoprim–sulfamethoxazole
	RNS	repetitive nerve stimulation	TN	trigeminal neuralgia
	ROM	range-of-motion	TNF	tumor necrosis factor
	RPR	rapid plasma reagin	TOAST	Trial of Org 10172 in Acute Stroke Treatment
	RR	risk ratio/relative risk	TOE	trans-esophageal echocardiography
	RRR	relative risk reduction	TOF	time-of-flight
	(r)-tPA	(recombinant) tissue plasminogen activator	TPHA	Treponema pallidum particle agglutination assay
	SAM	S-adenosyl-methionine	TS	Tourette's syndrome
1	SC	Sydenham's chorea	TSE	turbo spin echo/transmissible spongiform
	SCA	spinocerebellar ataxia		encephalopathy
	SCC	semicircular canal	TSH	thyroid stimulating hormone
	SCD	subacute combined degeneration	TST	thermoregulatory sweat test
	SCI	spinal cord injury	TTE	transthoracic echocardiography
	SCLC	small-cell lung carcinoma	TTR	time in therapeutic range/transthyretin
	SCN	suprachiasmatic nucleus	UBO	unidentified bright object
	ScvO ₂	central venous oxygen saturation	UFH	unfractionated heparin
	SD	semantic dementia	ULN	upper limit of normal
	SDB	sleep-related breathing disorder	UMN	upper motor neuron
	SDS	Shy-Drager syndrome	VA	visual acuity
	SE	status epilepticus	VaD	vascular dementia
	SEGA	subependymal giant cell astrocytoma	VAPP	vaccine-associated paralytic poliomyelitis
	SFEMG	single fiber electromyography	VLDL	very low-density lipoprotein
	SGCT	subependymal giant cell tumor	VDRL	Venereal Disease Research Laboratory
	SIADH	syndrome of inappropriate antidiuretic hormone	V-EEG	video electroencephalography
	SIBM	sporadic inclusion body myositis	VEGF	vascular endothelial growth factor
	SIS	second impact syndrome	VEMP	vestibular evoked potential
	SISCOM	subtraction ictal SPECT coregistered to MRI	VEP	visual evoked potential
	SLE	systemic lupus erythematosus	VF	visual field
	SMA	spinal muscular atrophy	VGCC	voltage-gated calcium channel
	SMN	survival of motor neuron	VHL	von Hippel-Lindau disease
	SNAP	sensory nerve action potential	VKA	vitamin K antagonist
	SND	striatonigral degeneration	VLCFA	very long-chain fatty acid
	SOD	septo-optic dysplasia	VLM	ventrolateral medulla
	SOREMP	sleep onset REM period	VNG	video-nystagmography
	SPECT	single photon emission tomography	VNS	vagal nerve stimulation
	SSCP	single-stranded conformational polymorphism	VOR	vestibulo-ocular reflex
	SSEP	somatosensory evoked potential	VP	vascular parkinsonism/variegate porphyria
	SSPE	subacute sclerosing panencephalitis	VPM	ventral posteromedial
	SSRI	selective serotonin-reuptake inhibitor	VR	volume rendered
	SSS	superior sagittal sinus/Scandinavian Stroke Scale	VSGP	vertical supranuclear gaze palsy
	SUDEP	sudden unexplained death in epilepsy	VSR	vestibulospinal reflex
	SVV	subjective visual vertical	VTE	venous thromboembolism
	SVZ	subventricular zone	vWF	von Willebrand factor
	SW	Sturge–Weber syndrome	VWFCP	von Willebrand factor-cleaving protease
	SWI	susceptibility-weighted imaging	VZ	ventricular zone
	TA	temporal arteritis	VZV	varicella-zoster virus
	TAB	temporal artery biopsy	WBC	white blood cell
	TACI	total anterior circulation infarct	WBRT	whole brain radiation therapy
	TACS	total anterior circulation syndrome	WD	Wilson's disease
	TAO	thyroid-associated ophthalmopathy	WHO	World Health Organization
	TB	tuberculosis	WNV	West Nile virus
	TBI	traumatic brain injury	XP	xeroderma pigmentosum
	TCD	transcranial Doppler ultrasonography		
	TCS	tuberous sclerosis complex		

CONTENTS

Contributors	7	Chapter 5: Vertigo	155
Preface	9	Jorge Kattah	
Dedications	10	Introduction	155
Abbreviations	- 11 m	Specific syndromes	169
District States and multipanistillader	ins lariV	VES-and a same two yelgometry has land operation hellerile	
Chapter 1: Neurologic diagnosis	17	Chapter 6: Hyperkinetic movement disorders	175
Robert Edis		Annu Aggarwal, Victor Fung,	
Introduction	17	Philip Thompson	
The diagnostic process in neurology	18	Introduction	175
Neurologic examination	21	Dystonia	176
Lumbar puncture and CSF examination		Chorea	187
John Dunne		Tremor	190
Neurophysiologic examination		Myoclonus	194
William McAuliffe		Tics and the arebnosib whod nois	
Imaging the brain and spine		Stereotypies	201
magnig the bruin and spine		Dyskinesias	202
Chapter 2: Disorders of consciousness	81	Drug-induced movement disorders	
Vibhav Bansal, Sean Ruland,		Psychogenic movement disorders	
Venkatesh Aiyagari		1 sychogeme movement disorders	200
Impaired consciousness	81	Chapter 7: Developmental diseases of the	
Brain death		nervous system	207
Persistent/permanent vegetative state	93	James H. Tonsgard	
Minimally conscious state		Introduction	207
Locked-in syndrome		Embryonic development of the nervous system	207
Syncope	96	Developmental malformations of the	207
Hypoxic–ischemic encephalopathy		nervous system	209
Central pontine myelinolysis	107	Defects in neural tube formation	210
Central pontine myemorysis	107	Defects in hindbrain development	215
Chapter 3: Epilepsy	111	Defects in forebrain and cerebral development	220
Paul V. Motika, Andres M. Kann		Neurocutaneous disorders	227
Definition Definition		Neurofibromatosis Type 1	228
Epidemiology		Neurofibromatosis Type 2	234
Pathophysiology		Tuberous sclerosis	237
Classification		Sturge–Weber syndrome	241
Clinical features		Hereditary hemorrhagic telangiectasia	
Differential diagnosis		(Osler–Rendu–Weber syndrome)	243
Investigation and diagnosis	116	Linear nevus sebaceous syndrome	246
Treatment			
		Cowden's syndrome and Lhermitte–Duclos disease	
Chapter A: Hondache	135		
Chapter 4: Headache	133		2 40
Peter J. Goadsby	125	Incontinentia pigmenti	
Headache	135	Hypomelanosis of Ito	250
Migraine	139		
Tension-type headache	149		
Cluster headache	150		

Chapter 8: Hereditary and metabolic		Chapter 11: Infections of the central	
diseases of the central nervous		nervous system	421
system in adults	253	Jeremy D. Young, James L. Cook	
Alma R. Bicknese		Acute bacterial meningitis	421
Introduction	253	Brain abscess	426
Metabolic and degenerative diseases in adults	255	Neurosyphilis	429
Huntington's disease	255	Lyme disease	433
Cerebral autosomal dominant arteriopathy		Tuberculosis (TB)	435
with subcortical infarcts and		Botulism	439
leukoencephalopathy	257	Tetanus	441
Fragile X-associated tremor/ataxia syndrome	258	Aseptic meningitis	444
X-linked adrenoleukodystrophy	259	Viral encephalitis	447
Pantothenate kinase-associated	237	Rabies	452
	261	Poliomyelitis	454
neurodegeneration	261		434
Inherited diseases with prominent visceral	202	Progressive multifocal leukoencephalopathy	157
and neurologic findings	262	(PML)	457
Wilson's disease	262	Cryptococcosis	460
Acute intermittent porphyria	264	Coccidioidomycosis	463
Disorders of amino acid metabolism	265	Toxoplasmosis	465
Homocystinuria	265	Neurocysticercosis	468
Inclusion body disorders	269	Eosinophilic meningitis	470
Lafora body disease	269	Creutzfeldt–Jakob disease (CJD)	472
Adult neuronal ceroid lipofuscinoses	270		
Late-onset lysosomal storage diseases	271	Chapter 12: Inflammatory disorders of the	
Late-onset G _{M2} gangliosidosis	271	nervous system	481
Late-onset metachromatic leukodystrophy	272	Neil Scolding	
Fabry's disease	273	Acute disseminated encephalomyelitis	
Late-onset Krabbe disease	274	(post-infectious encephalomyelitis)	481
Niemann-Pick Type C	274	Multiple sclerosis (MS)	484
Mitochondrial disorders	275	Neurosarcoidosis	497
Diseases of mitochondrial DNA	276	Cavernous sinus syndrome	502
Chapter 9: Trauma of the brain and		Chapter 13: Tumors of the nervous system	507
spinal cord	279	Julie E. Hammack,	
James L. Stone,		Robert P. Dinapoli	
Prasad S.S.V. Vannemreddy		Neoplasia	507
Introduction	279	Gliomas	515
Head injury	279	Meningioma	522
Concussion	283	Craniopharyngioma	526
Spinal cord injury	284	Pituitary tumors	527
		Vestibular schwannoma (acoustic neuroma)	529
Chapter 10: Stroke and transient ischemic		Primary central nervous system lymphoma	530
attacks of the brain and eye	291	Germ cell tumors of the CNS	532
Graeme J. Hankey	231	von Hippel–Lindau disease	533
Introduction	291	Metastases to the CNS	536
Epidemiology	294	Paraneoplastic neurologic disease	539
		r araneopiastic neurologic disease	337
Pathology and etiology Risk factors	296		
	321		
Clinical assessment	323	SECTION AND ADDRESS OF THE SECTION ADDRESS OF THE SECTI	
Investigations	343		
Diagnosis	369		
Prognosis	376		
Treatment	381		

Chapter 14: Degenerative diseases of the		Chapter 17: Cranial neuropathies I, V,	
nervous system	545	and VII–XII	679
James A. Mastrianni		Wilson Cueva, Helene Rubeiz	
Dementia	545	Olfactory nerve neuropathy (cranial nerve I)	679
Alzheimer's disease (AD)	545	Trigeminal nerve neuropathy (cranial nerve V)	682
Frontotemporal dementia (FTD)	555	Selected conditions affecting the trigeminal nerve	687
Dementia with Lewy bodies (DLB)	560	Facial nerve neuropathy (cranial nerve VII)	690
Vascular dementia	564	Selected conditions affecting the facial nerve	694
Prion diseases	568	Vestibulo-cochlear nerve neuropathy	
Normal pressure hydrocephalus	575	(cranial nerve VIII)	696
Brandon R. Barton, Kathleen M. Shannon		Glossopharyngeal nerve neuropathy	
Parkinson's disease and parkinsonian		(cranial nerve IX)	700
disorders	577	Selected conditions affecting the	
Introduction	577	glossopharyngeal nerve	702
Primary (idiopathic) parkinsonism:		Vagus nerve neuropathy (cranial nerve X)	703
Parkinson's disease	578	Spinal accessory nerve neuropathy	
Multisystem degenerations ('Parkinsonism pl		(cranial nerve XI)	705
or atypical parkinsonism)	596	Hypoglossal nerve neuropathy (cranial nerve XII)	706
Heredodegenerative and secondary		Charlette Annie Tolki diseason	
parkinsonism	608	Chapter 18: Cranial neuropathies II, III, IV,	
Sid Gilman, Vikram Shakkottai, Peter Todd		and VI	709
Hereditary ataxias	612	Molly E. Gilbert	
Introduction	612	Optic nerve neuropathy (cranial nerve II)	709
Congenital ataxias	612	Cranial nerve III, IV, and VI palsies	723
Autosomal dominant cerebellar ataxias	616	er 22 - Nauromuscular junction	
Autosomal recessive cerebellar ataxias	625	Chapter 19: Spinal cord disease	739
X-linked causes of ataxia	633	Octavia Kincaid	
		Introduction to myelopathy	739
Chapter 15: Acquired metabolic diseases of	f	Extramedullary spinal cord lesions	
the nervous system	641	(extrinsic myelopathy)	741
Fernando D. Testai		Intramedullary spinal cord lesions	
Thiamine deficiency: Wernicke-Korsakoff		(intrinsic myelopathy)	744
syndrome	641	Inflammatory myelopathy (e.g. myelitis)	744
Vitamin B12 deficiency	645	Infectious myelopathy	746
Vitamin E deficiency	650	Metabolic and toxic myelopathies	750
Niacin (nicotinic acid) deficiency	653	Vascular myelopathies	751
Marchiafava-Bignami disease	654	Structural myelopathies	755
Hepatic encephalopathy	655	Hereditary myelopathies	756
Uremic encephalopathy	661	Spinal cord tumors	761
Posterior reversible encephalopathy syndrome	662	Myelopathy associated with	
Conversion of All Book at Chicago College Sid Gilm		motor neuron disease	764
Chapter 16: Disorders of circulation of			
the cerebrospinal fluid	667	Chapter 20: Autonomic nervous system	
Daniel B. Hier, Edward A. Micha		disorders	777
Introduction	667	Robert Henderson, Judy Spies	
Aqueductal stenosis with hydrocephalus	668	Autonomic neuropathy	777
Communicating hydrocepahalus	670	Conversity of Misure, Miller School	
Normal pressure hydrocephalus	671		
Chiari malformation	672		
Syringomyelia (syringohydromyelia)	674		
Idiopathic intracranial hypertension			
(pseudotumor cerebri)	675		
Intracranial hypotension syndrome	677		

Chapter 21: Diseases of the peripheral nerve	Chapter 23: Muscle disorders	873	
and mononeuropathies	787	Kourosh Rezania, Peter Pytel,	
Andrea Swenson		Betty Soliven	
Peripheral neuropathy	787	Introduction	873
Diabetic neuropathy	793	Myopathy	875
Toxic neuropathy	796	Inflammatory myopathy	877
Nutrition-related neuropathy	801	Critical illness myopathy	884
Neuropathy in systemic disease	803	Endocrine and metabolic myopathies	886
Guillain-Barré syndrome (GBS)/		Muscular dystrophies	895
Acute inflammatory demyelinating		Muscle channelopathies (disorders of	
polyradiculoneuropathy (AIDP)	804	membrane excitability)	911
Chronic acquired demyelinating			
polyneuropathy	807	Chapter 24: Sleep disorders	917
Vasculitic neuropathy	811	John H. Jacobsen,	
Immune-mediated brachial plexus neuropathy		Michael Kohrman	
(IBPN)	815	Introduction	917
Infectious neuropathy	816	Normal sleep	917
Hereditary motor and sensory neuropathy		Commonly performed sleep tests	923
(Charcot-Marie-Tooth disease)	820	Insomnias	931
Syndromic hereditary peripheral neuropathies	824	Hypersomnias of central origin	935
Mononeuropathies	826	Sleep-related movement disorders	941
Upper extremity neuropathy	826	Sleep-related breathing disorders	945
Lower extremity neuropathy	842	Circadian rhythm sleep disorders	951
Entr-onser lysosomalistor thumis 2415 HI syrin hi		Parasomnias	956
Chapter 22: Neuromuscular junction		Conclusion	961
disorders	855		
Qin Li Jiang, Julie Rowin		Index	965
Myasthenia gravis	855		
Congenital myasthenic syndromes	865		
Lambert–Eaton myasthenic syndrome	867		
Botulism	869		

HANKEY'S SECOND EDITION CLINICAL NEUROLOGY

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