

Clinical guidelines from the Priority Epilepsy Program of the public health sector in Mexico

J.C. Reséndiz-Aparicio

Hospital Psiquiátrico Infantil Dr. Juan N. Navarro y P.P.E, Instituto Nacional de Neurología y Neurocirugía Dr. Manuel Velasco Suárez. Mexico City, Mexico



The *Programa Prioritario de Epilepsia* (PPE - Priority Epilepsy Program) was created based on the accord published in the Mexican Official Gazette of the Federation on October 24th, 1984. This program has labored in an uninterrupted manner to regulate, coordinate, methodize, and optimize the strategies in favor of patients with epilepsy, as well as their families and society. There are currently 78 centers of integral treatment for epilepsy in Mexico, located in various hospitals belonging to Mexico's health sector.

The headquarters for the national coordination is in the Instituto Nacional de Neurología y Neurocirugía (National Institute of Neurology and Neurosurgery) "Dr. Manuel Velasco Suarez" (INNN due to its acronym in Spanish) in Mexico City, from where all actions are

planned for this task. The national coordination is led by its creator and founder Francisco Rubio Donnadiu, MD and by the author.

The development of the first Clinical Guidelines (CGs) has been a laborious effort, one that has been finished due to the work of all the coordinators of the PPE, who are neurologists and pediatric neurologists certified by the Mexican Board of Neurology and who work in one of the many institutions of the health sector in Mexico. To elaborate the CG, all the coordinators of the PPE met in person in two meetings, the first in the city of Leon and the second in the city of Puebla, where we formed workgroups for each CG. These meetings were possible due to the support of the federal government and the contributions of the

Correspondence:

Juan Carlos Reséndiz-Aparicio

E-mail: jc_doc@yahoo.com

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pharmaceutical laboratories that aid in training primary health-care physicians. These CGs are designed to aid the primary health-care physicians and the specialists in making adequate decisions when approaching epileptic patients of different age groups and genders. These are the culmination of the experience of their authors, who have followed the necessary steps for proper and updated scientific research, using the criterion of the American Epilepsy Society 2016 to analyze the levels of evidence and recommendations including the benefit for the patients. To evaluate the quality of the CG, two experts coordinated each table and applied the Spanish version of the AGREE instrument of 2001.

Due to the advances in the knowledge of epilepsy, the PPE group aims to update the CG every 5 years.

These CGs constitute a series of recommendations developed by a group of medical physicians that have a particular interest in the field of epilepsy and work

throughout the various institutions of the health sector; however, it is understood that the application of said recommendations depends on many factors. It is important to state that there are no conflicts of interest in these CGs due to the fact that they are editorially independent of any external funding.

Finally, I would like to thank the INNN for their hospitality and support in the coordination of the Program, the Mexican Academy of Neurology for the publication in their magazine, the Mexican Society of Pediatric Neurology who supported the process of translating the CG to English, and the authorities of the Hospital Psiquiátrico Infantil (Children's Psychiatric Hospital) of Mexico City who have allowed me to work as the adjunct executive member of the Priority Epilepsy Program and above all, thank you to the coordinators of the Priority Epilepsy Program who worked on this project; this work is dedicated to them and their families and to which I express my most ample recognition.

Juan Carlos Reséndiz Aparicio, MD, Adjunct Executive Member

American Epilepsy Society 2016

Article Classification: Evidence

Class I: A randomized, prospective and controlled clinical trial with masked outcome assessment, in a representative population. The following are required:

- a. No more than two specified primary results.
- b. Blind allocation of subjects.
- c. Exclusion/inclusion criteria are clearly defined.
- d. Relevant baseline characteristics are presented and substantially equivalent among treatment groups or there is appropriate statistical adjustment for differences.
- e. Adequate accounting for dropouts with numbers sufficiently low to have a minimal potential for bias. (study was completed with at least 80% of the enrolled subjects).
- f. Demonstration of superior design of the studies or demonstration of non-inferiority with a 10% non-inferior design margin.

Class II: A randomized, prospective and controlled clinical trial with masked outcome assessment that lacks one or two criteria of Class I a-e above, or a prospective matched cohort study with masked objective outcome assessment in a representative population that meets a-e.

Class III: All other control trials in a representative population, where outcome was independently assessed by objective outcome measurement.

Class IV: Evidence from non-controlled trials, including series reports, case reports, consensus, or expert opinion.

Evidence for the Recommendation	Conclusion and Recommendation
Level A	Conclusion, Level A
One or more Class I trials or two or more Class II trials.	Established as effective, ineffective, or harmful for the given condition in the specified population. Recommendation: Must be done or must not be done.
Level B	Conclusion, Level B
One or more Class II trials or three or more class III trials.	Probably effective, ineffective, or harmful for the given condition in the specified population. Recommendation: It must be considered or must not be considered.
Level C	Conclusion, Level C
Two or more Class III trials.	Possibly effective, ineffective, or harmful for the given condition in the specified population. Recommendation: It could be considered or should not be considered.
Level U	Conclusion, Level U
Absence of trails that complement levels A, B, or C.	Data is insufficient or inadequate given current knowledge, treatment is unproven Recommendation: Should not be performed
Level R-PPE	Conclusion, Level R-PPE
	No evidence from levels A, B, or C, but it is a recommendation by consensus of the group that elaborated the Clinical Guidelines of the PPE.

In all of the CGs of the Priority Epilepsy Program (PPE), the abbreviations we published are the same as the book "Epilepsia" by authors Rubio, Reséndiz, Alonso, and Sentías, by the editorial Alfil in 2016, page numbers IX, X y XI; ISBN 978-607-741-168-0.

Glossary and Abbreviations

Channelopathies

SCN4A, SCN2A, SCN1B

KCNA1, KCNQ2, KCNQ3

CACNA1A

CHRNA4, CHRN2

GLRA1

GABRG2

Ion: NA: sodium; K: potassium; CA: calcium;

CH: acetylcholine; GL: glycine; GABA: gamma-aminobutyric acid

Channel or receptor: CN: channel; R: receptor;

N: nicotinic

Subunit: A: α ; B: β ; Q: M; G: γ

Seizure

ES: epileptic seizure

GS: generalized seizure

GTCS: generalized tonic-clonic seizure

FS: focal seizure

FIAS: focal impaired awareness seizures (or disconnection from medium)

FAS: focal aware seizure/simple partial seizure

FBTCS: focal to bilateral tonic-clonic seizure

SE: status epilepticus

FeS: febrile seizure

Electrolytes and neurotransmitters

Ca⁺⁺: calcium

Cl⁻: chlorine

K⁺: potassium

Mg⁺⁺: magnesium

Na⁺: sodium

GABA: gamma-aminobutyric acid

NMDA: N-methyl-D-aspartate

AMPA: a-amino-3-hydroxy-5-methyl-4-isoxazolepropionic acid

Neurologic structures

BBB: blood–brain barrier
CSF: cerebrospinal fluid
CNS: central nervous system

Diagnostic tests

fMRI: functional magnetic resonance imaging
MRI: magnetic resonance imaging
PET: positron emission tomography
SPECT: single-photon emission computed tomography
CT: computed tomography scan
ECoG: electrocorticography/intracranial electroencephalography
EEG: electroencephalogram
MEG: magnetoencephalogram
Video-EEG: video electroencephalogram
PSG: polysomnogram
EKG: electrocardiogram
LP: lumbar puncture/spinal tap

Genetics

AD: autosomal dominant
AR: autosomal recessive.
p: short arm of a chromosome
q: long arm of a chromosome
DNA: deoxyribonucleic acid
RNA: ribonucleic acid
NB: newborn/neonate

Organizations

AAN: American Academy of Neurology
AES: American Epilepsy Society
AAP: American Academy of Pediatrics
CAIE: Centros de Atención Integral para la Epilepsia or Comprehensive Care Centers for Epilepsy
FDA: Food and Drug Administration
IBE: International Bureau for Epilepsy
ILAE: International League Against Epilepsy
INNN: Instituto Nacional de Neurología y Neurocirugía or National Institute for Neurology and Neurosurgery.
WHO: World Health Organization
PAHO: Pan American Health Organization
PPE: Programa Prioritario de Epilepsia or Priority Epilepsy Program
SAdE: Sociedad Andaluza de Epilepsia or Andalusian Epilepsy Society

GPC-PPE: Guía de Práctica Clínica del Programa Prioritario de Epilepsia or Clinical Guidelines of the Priority Epilepsy Program

Additional neurological disorders

CVD: cerebrovascular disease
TBI: traumatic brain injury

Drug administration routes

IM: intramuscular
IV: intravenous
PO: oral
S/C: subcutaneous
S/L: sublingual

Syndromes and types of epilepsy

BECTS: benign epilepsy with centrotemporal spikes (Rolandic Epilepsy)
IGE: idiopathic generalized epilepsy
PME: progressive myoclonus epilepsy
JME: juvenile myoclonus epilepsy
MTS: mesial temporal sclerosis
LGS: Lennox-Gastaut syndrome
DRE: difficult to treat seizures/drug-resistant epilepsy

Miscellaneous

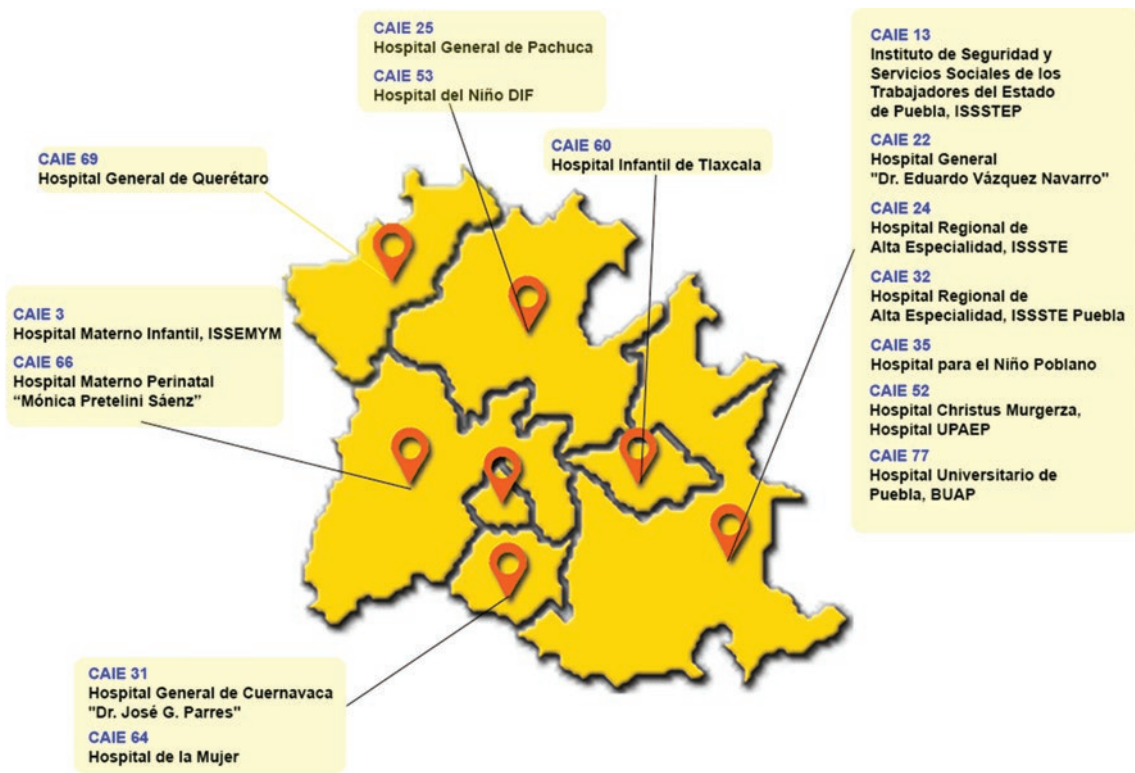
AED: antiepileptic drugs/anti-seizure medications
BZD: benzodiazepines

Antiepileptic drugs

ACZ: acetazolamide
ACTH : adrenocorticotrophic hormone
NE: barbitone
NE: beclamide
CBZ: carbamazepine
CLB: clobazam
CZP: clonazepam
CLP: lorazepam
DZP: diazepam
ESM: ethosuximide
FBM: felbamate
GBP: gabapentin
LTG: lamotrigine
LEV: levetiracetam
LZP: lorazepam
MDL: midazolam

MPH: methylphenidate	CBT: carabersat (SB-204269)
MPB: methylphenobarbital	TBT: tonabersat (SB-220453)
MSM: mesuximide/methsuximide	SFM: safinamide (PNU-151774E)
NTZ: nitrazepam	RUF: rufinamide (SGP33101)
OXC: oxcarbazepine	STL: soretolide (D-2916)
PAC: phenacemide	TLP: talampanel (GYKI 53773)
PTR: pheneturide	HUP: huperzine A
PB: phenobarbital	ATM: atipamezole
PSM: phensuximide	VLR: valroceamide (TV1901)
PHT: phenytoin	IVR: isovaleramide
NE: fosphenytoin	VPG: valproyl glycinamide
PGB: pregabalin	VLT: valnoctamide
PRM: primidone	VPD: valpromide
PRO: progabide	VCD: valroceamide
STM: sultiame/sulthiame	PID: propylisopropylacetamide
TGB: tiagabine	LiCBZ: licarbazepine
TPM: topiramate	EsliCBZ: eslicarbazepine (BIA 2-093)
NE: trimethadione	FI-FBM: fluorofelbamate
VPA: valproic acid	NA: ganaxolone
VGB: vigabatrin	carisbamate (RWJ-333369)
ZNS: zonisamide	perampanel
NE: 4-amino-3-hydroxybutiric Acid	ELB-139
FLN: flunarizine	JZP-4
LSG: losigamone	NS-1209
RLT: raltitoline (CI-946)	CGX-1007
REM: remacemide	SPD-421
STP: stiripentol	ICA27243
HRK: harkoseride	T2000
LCM: lacosamide	XP-13512*
RET: retigabine (D-23129)	YKP3089
BRV: brivaracetam	NE: not established
STM: seletracetam (ucb 44212)	NA: not applicable

Centers of integral care for epilepsy “CAIE” in Mexico



- CAIE 1.- Instituto Nacional de Neurología y Neurocirugía.
- CAIE 2.- Instituto Nacional de Pediatría.
- CAIE 5.- Hospital Pediátrico de Legaria.
- CAIE 11.- Instituto Nacional de Perinatología.
- CAIE 14.- Hospital General Centro Médico La Raza.
Hosp. Gral. "Dr. Gaudencio González Garza", IMSS.
- CAIE 15.- Centro Médico Nacional "20 de Noviembre".
- CAIE 28.- Hospital Psiquiátrico Infantil "Dr. Juan N. Navarro".
- CAIE 29.- Hospital Central de Alta Especialidad Sur PEMEX.
- CAIE 30.- Hospital Infantil de México "Federico Gómez".
- CAIE 33.- Centro Médico Nacional "20 de Noviembre",
ISSSTE.
- CAIE 37.- Hospital General de México "Dr. Eduardo
Liceaga", Unidad de Pediatría.
- CAIE 45.- Instituto Nacional de Ciencias Médicas y
Nutrición "Salvador Zubirán".
- CAIE 61.- Hospital General "Dr. Manuel Gea González".
- CAIE 72.- Hospital Regional "Lic. Adolfo López Mateos",
ISSSTE.
- CAIE 73.- Unidad Médica de Alta Especialidad, Hospital de
Pediatría "Dr. Silvestre Frenk Freund"
CMN Siglo XXI, IMSS.
- CAIE 76.- Hospital Central Norte, PEMEX.

CAIE CDMX



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