

# Republic of the Philippines Department of Health OFFICE OF THE SECRETARY .

JAN 14 2021

ADMINISTRATIVE ORDER No. 2020 - 0012

> **SUBJECT:** Implementing Guidelines on the Medicine Access Program for

> > Mental Health (MAP-MH)

### I. **RATIONALE**

Mental health (MH) issues can adversely affect one's well-being, behavior, personal relationships, contributions to society, and overall quality of life. Mental disorders account for 10% of the global burden of disease (GBD) and 30% of the non-fatal burden of disease. In the Philippines, 3.3% of the population suffer from depressive disorders and 3.1% suffer from anxiety disorders (WHO Mental Health Atlas, 2017). The World Economic Forum (WEF) estimated in 2013 that the cumulative global impact of mental disorders alone in terms of economic output will amount to US\$ 16 trillion over the next 20 years. Yet on average, governments spend less than 2% of their health budget on mental health, with most low- and middle-income countries spending less than 1%. To address the looming mental health crisis, it is crucial for governments to invest in mental health initiatives.

Recognizing the role of mental health and well-being in national development, the Department of Health (DOH) released Administrative Order (AO) No. 8 series of 2001 entitled "National Mental Health Policy". This policy set forth the guidelines for the establishment of a sustainable mental health program in the Philippines. The program was supported by the issuance of AO 2016-0039 "Revised Operational Framework for a Comprehensive National Mental Health Program (NMHP)" which provided among others, equitable access to the rational use of a wide range of pharmacologic interventions in the treatment and management of mental, neurologic, and substance use (MNS) disorders. Moreover, mental health is included as a priority agenda of the current administration as articulated in Republic Act (RA) No. 11223 "Universal Health Care (UHC) Act", as well as in RA 11036 "Mental Health Act" — which also mandated the improvement of MH services delivery nationwide.

The Medicine Access Program for Mental Health (MAP-MH), started in 2012 by the DOH Pharmaceutical Division (PD) and operationalized by the National Center for Mental Health (NCMH), was designed to ensure availability of mental health drugs in the community. Since then, 207 access sites have been opened with around 39,000 service user beneficiaries. With the transfer of MAP-MH to the National Mental Health Program (NMHP) under the DOH Disease Prevention and Control Bureau (DPCB) and the goal of expanding coverage of beneficiaries and medicines being provided, there is a need to establish standards and guidelines to aid in the proper implementation of MAP-MH nationwide.

### Π. **OBJECTIVES**

This Order aims to set the overall guidelines on the implementation of MAP-MH in access sites such as DOH hospitals, Centers for Health Development (CHDs), Ministry of Health -Bangsamoro Autonomous Region in Muslim Mindanao (MOH-BARMM), Treatment and Rehabilitation Centers (TRCs), and other health facilities. Specifically, this Order aims to:

- 1. Increase service user's access to quality essential medicines in the treatment of various mental, neurologic, and substance use (MNS) disorders, taking into consideration rational drug use and availability up to the grassroots level.
- 2. Establish a functional electronic information management system for MAP-MH.
- 3. Improve primary health care in the poorest communities by addressing the needs of the population for essential medicines as part of primary and secondary prevention of MNS disorders.
- 4. Conduct efficient monitoring and evaluation of the utilization of essential medicines for MNS disorders.

### III. SCOPE AND COVERAGE

This Order shall apply to DOH hospitals, CHDs, TRCs, local government units (LGUs), rural health units (RHUs), Department of Social Welfare and Development (DSWD), Bureau of Jail Management and Penology (BJMP), and other institutions and agencies that will qualify as access sites, for an increased access to quality essential medicines for MNS disorders for the benefit of the Filipino people, especially the marginalized sector. This Order shall include MOH-BARMM pursuant to RA 11054 "Organic Law for the Bangsamoro Autonomous Region in Muslim Mindanao (BARMM)".

### IV. DEFINITION OF TERMS

- 1. Access site refers to a health facility where needed essential medicines for MNS disorders are being provided for enrolled service users.
- 2. Enrolled service user refers to a service user with any MNS disorder who is in the community and is seeking regular consultation in an identified access site as a beneficiary of the MAP-MH.
- 3. Essential medicines refers to a list of medicines for treating MNS disorders identified by a Therapeutic Committee that will be organized for the purpose.
- 4. Medicine Access Program for Mental Health (MAP-MH) refers to a medicine access program designed to provide needed essential medicines for all enrolled service users with MNS disorders who are in the community and are seeking regular consultation in an identified access site.
- 5. Mental Health Gap Action Programme (mhGAP) refers to a training program developed by the World Health Organization (WHO) for primary care practitioners in non-specialized settings as an intervention guide in the treatment and management of MNS disorders and adapted for use in the Philippine context.
- 6. National Drug Policy Compliance Officer (NDPCO) refers to pharmacists under CHDs or MOH-BARMM that are designated to oversee the implementation of all programs and activities of the DOH Pharmaceutical Division (PD).
- 7. National Mental Health Program (NMHP) refers to a program offering a wide range of promotive, preventive, curative, and rehabilitative services for persons who suffer from MNS disorders. It has five (5) components: Wellness of Daily Living, Extreme Life Experience, Mental Disorders, Neurologic Disorders, and Substance Abuse and other Forms of Addiction.

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- 8. Near expiry medicine refers to a medicine that is six (6) months prior to expiration date.
- 9. Starter kit refers to a set of essential medicines provided initially to a health facility whose primary care physician has been newly trained in mhGAP.

### V. GENERAL GUIDELINES

- 1. The Technical Working Group (TWG) created through a Department Personnel Order (DPO) shall be regularly convened to identify essential medicines for MNS disorders that shall be procured for the MAP-MH. The TWG may change or add medicines based on their assessment of the current needs and situations such as cost-effectiveness, treatment guidelines, most prescribed and fast-moving medicines, in accordance with the latest version of the Philippine National Formulary (PNF).
- 2. Essential medicines for MNS disorders shall be subjected to periodic review by the Health Technology Assessment Council (HTAC). Investment on any health technology or development of any benefit package by the DOH and the Philippine Health Insurance Corporation (PhilHealth) shall be based on positive recommendation of the HTAC.
- 3. Essential medicines for MNS disorders are classified as individual-based health services. The DOH shall finance the procurement until such time that essential mental health services are included in PhilHealth's primary care service package as stipulated in the Implementing Rules and Regulations (IRR) of RA 11223 "Universal Health Care Act".
- 4. Procurement of medicines shall follow the existing rules and regulations provided for in RA 9184 "Government Procurement Reform Act".
- 5. Quality analysis for samples of the MAP-MH shall be conducted by the Food and Drug Administration (FDA) before delivery to the access sites.
- 6. The MAP-MH medicines shall be packed using FDA-approved packaging materials for the purpose of promoting and advocating the program.
- 7. The MAP-MH shall be implemented in identified access sites by the DOH-NMHP.
- 8. The MAP-MH medicines shall be prescribed, dispensed, and used following existing standard treatment guidelines recognized by DOH.
- 9. The Pharmacotherapeutic Guidelines for the MAP-MH shall serve as the basis for clinical intervention in the primary and secondary health facilities (see Annex A).

### VI. **SPECIFIC GUIDELINES**

### A. Access Sites

- 1. The CHD Regional Coordinators shall recommend inclusion of access sites to the DOH-NMHP for approval based on the following criteria:
  - a. A licensed health facility where service users with MNS disorders are consulted, diagnosed, treated, managed, and regularly followed up in a specialty care setting (e.g. mental hospital, acute care psychiatric facility, chronic care psychiatric

- facility, treatment and rehabilitation center (TRC), neuroscience/neurology departments and clinics, and the like).
- b. An outpatient or primary care clinic/health facility where there is an mhGAP trained nonspecialist medical personnel, or other trained primary care physician recognized by DOH e.g. rural health units (RHU).
- 2. Identified access sites shall be provided with essential medicines according to the needs of the service users. To this end, the access site shall be responsible for implementing MAP-MH by procuring essential medicines through a Requisition Issuance Slip (RIS) with a coverage of one (1) year. This shall allow new enrolled service users to avail of the medicines. Submission of this request form shall be in the first week of November of the previous year, and first week of May of the current year. See Annex B for the template of the RIS Form.
- 3. Access sites shall regularly submit utilization reports. Reports shall be submitted quarterly on the first weeks of April, July, October, and January accordingly. See Annex C for the template of the Reporting Form.
- 4. Access sites shall ensure that medicines are stored properly. It is a must for every access site to ensure that a dedicated cabinet storage with functional lock is provided for the purpose in compliance with the Guide to Good Storage Practices (GSP) for Pharmaceuticals discussed in Annex 9 of the WHO Technical Report Series No. 908, 2003.
- 5. In case there is a shortage or stock-out of medicines in access sites, requests shall be made to the DOH-NMHP for augmentation purposes. However, requests shall be made known three (3) months prior to allow for processes to be completed.
- 6. Near expiry medicines shall be reported to the nearest CHDs or MOH-BARMM while expired medicines shall be disposed in accordance with the Joint Department of Environment and Natural Resources (DENR)-DOH AO No. 02, s. of 2005 and its amendments. A witness from the CHD or MOH-BARMM, FDA, and Commission on Audit (COA) shall be present whenever disposal of expired drugs is conducted.
- 7. Access sites shall inform and coordinate with the DOH Central Office regarding slow moving drugs and inventories, and other issues and concerns regarding the program.
- 8. Access sites shall diligently enforce the mechanism of stock transfer of medicines among access sites to prevent wastage in coordination with the CHD Regional Coordinators and NDPCO.

# **B.** Enrolled Service Users

- 1. Service user shall visit an identified access site and follow the process below:
  - a. Register in the access site
  - b. Undergo assessment by a physician to receive a clinical diagnosis
  - c. Present a valid prescription from a physician to avail the essential medicines for MNS disorders
  - d. Adhere to the treatment given and comply with the follow-up schedules as advised by the physician

- 2. Enrolled service users shall be registered in the electronic information management system developed by the DOH Knowledge Management and Information Technology Service (KMITS).
- 3. In case of change of address, the enrolled service user shall have options to re-enroll to another access site that is more accessible to him/her and shall be properly endorsed by the previous access site including all pertinent records of the service user.
- 4. In cases wherein the service user was not endorsed by a previous access site, the new access site shall determine the needs of the service user and enroll him/her to the program.

### VII. MONITORING AND EVALUATION

- 1. Access sites shall be monitored and evaluated regularly, either by announced or unannounced visits. To this end, a Monitoring and Evaluation Committee shall be organized for this purpose. This Committee shall include, among others, representatives from the NMHP, NCMH, PD, and CHD following the provisions stated in Department Order (DO) No. 2016-0269 "Guidelines on Planning, Monitoring and Evaluation of Programs, Activities, and Projects in the DOH" and its amendments.
- 2. The Committee shall identify training and mentoring needs of access sites particularly in the proper dispensing of essential medicines procured for MAP-MH.

### VIII. ROLES AND RESPONSIBILITIES

- 1. The National Mental Health Program (NMHP) shall:
  - a. Lead the implementation of MAP-MH nationwide through identified access sites for all enrolled service users.
  - b. Ensure funding support for the procurement of essential medicines until such time that PhilHealth can develop and implement a benefit package for the purpose.
- 2. The Pharmaceutical Division (PD) and National Drug Policy Compliance Officer (NDPCO) shall provide technical assistance and monitoring support to NMHP in the procurement, distribution, and utilization of procured essential medicines for MAP-MH.
- 3. The Knowledge Management and Information Technology Service (KMITS) in collaboration with the NMHP shall develop a functional electronic information management system for MAP-MH.
- 4. The **Philippine Health Insurance Corporation (PhilHealth)** shall develop and implement an outpatient benefit package for MNS disorders.
- 5. The Centers for Health Development (CHDs) shall:
  - a. Identify health facilities to be identified as access sites for MAP-MH.
  - b. Monitor and supervise the implementation of MAP-MH in the identified access sites in their regions.
  - c. Collect, consolidate, and analyze the reports from access sites.
  - d. Prepare a summary of reports collected from access sites and submit this to NMHP with recommendations on improving the implementation.

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- e. Resolve issues and other concerns needing actions on the operationalization of the program.
- f. Enforce a mechanism of moving stocks of medicines among access sites with low levels of utilization in coordination with other access sites.

# 6. The Supply Chain Management Service (SCMS) shall:

- a. Receive the goods from the local suppliers according to the quantity and specification stated in the Purchase Order.
- b. Prepare reports such as Arrival Report to End-user, Request of Inspection and Acceptance to End-User, Notice of Delivery to COA, Request of Inspection to FDA, and Request of Inspection.
- c. Oversee the inspection of medicines together with the end-user.
- d. Manage the warehousing, packaging, and distribution of medicines to the identified access sites.

### IX. REPEALING CLAUSE

Provisions of previous Orders and other related issuances inconsistent or contrary to the provision of this Administrative Order (AO) are hereby revised, modified, repealed, or rescinded accordingly. All provisions of existing issuances which are not affected by this Order shall remain valid and in effect.

### X. **EFFECTIVITY**

This Order shall take effect fifteen (15) days after following its publication in a newspaper of general circulation and upon filing with the University of the Philippines Law Center (UPLC) of three (3) certified copies of this Order.

CISCO T. DUQUE III, MD, MSc. Secretary of Health



ANNEX A

# PHARMACOTHERAPEUTIC GUIDELINES FOR THE MEDICINE ACCESS PROGRAM FOR MENTAL HEALTH (MAP-MH)

For Primary and Secondary Care Facilities

Technical Working Group
Essential Non-Communicable Disease Division
Disease Prevention and Control Bureau

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### INTRODUCTION

Mental health and well-being are intertwined with personal well-being, family relationships, and contributions to society. Problems with mental health and well-being can adversely compromise learning, creativity, productivity, contribution to family and society and overall quality of life. The value of investing in mental health is well established. The World Economic Forum estimated in 2013 that the cumulative global impact of mental disorders alone in terms of economic output will amount to US\$ 16 trillion over the next 20 years.

Recognizing the role of mental health and well-being in national development, the Department of Health articulated a National Mental Health Policy in 2001, DOH AO No. 8 s. 2001 that provided for the establishment of a sustainable mental health program. Since then a viable mental health program has been established and is currently strengthening its implementation as mandated by DOH AO No. 2016-0039 which provided among others, the equitable access to the rationale use of a wide range of pharmacologic interventions in the treatment and management of mental, neurologic and substance use disorders and improvement in the quality of life. This mandate is further strengthened by the inclusion of mental health among the priority agenda of the current administration as articulated in the Philippine Health Agenda 2016-2022.

The Medicine Access Program for Mental Health (MAP-MH), started in 2012 by the Pharmaceutical Division and operationalized by the National Center for Mental Health was designed to ensure availability of mental health drugs in the community. Since then 138 access sites have been opened with around 20,000 patient beneficiaries. With the transfer of the Medicine Access Program for Mental Health to the National Mental Health Program under the Disease Prevention and Control Bureau of the Department of Health and the goal of expanding coverage of beneficiaries and medicines being provided, there is a need to establish standards and guidelines to aid in the proper implementation of the MAP-MH nationwide. Thus, the issuance of this Pharmacotherapeutic Guidelines on the Medicine Access Program for Mental Health.

The Pharmacotherapeutic Guidelines will serve as the basis for clinical intervention in the primary and secondary health facilities in the community. It can be utilized in conjunction with other community guidelines such as the Mental Health Gap Action Program (MHGAP) from the World Health Organization (WHO). The guideline is focuses on 6 mental health disorders, namely: Anxiety Disorders, Mood Disorders, Psychosis, Dementia, Epilepsy and Substance Abuse Disorders. All these diseases presented within the guideline is given a brief description of its clinical criteria and preferred regimen to be utilized for treatment.

This guideline was made from the collective efforts of various members of the Technical Working Group and various other stakeholders who aim for the improvement of health service provision in mental health and strengthening the implementation of the R.A. 11036 otherwise known as the Mental Health Act for the community.

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# CONSENSUS TREATMENT GUIDELINES ON ANXIETY DISORDERS

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# ANXIETY DISORDERS

Anxiety disorder reflect disorders that share a general feature of excessive fear (i.e. anticipation of emotional response to perceived or real threat) and/ or anxiety (anticipation of future threat) and demonstrate behavioral, physiologic and functional disturbance as a result. Panic attacks are a feature that can occur in the context of many anxiety disorders and reflect a type of fear response. There are several types of anxiety disorders.

# A. PANIC DISORDER

Panic disorder refers to recurrent unexpected panic attacks. A panic attack is an abrupt surge of intense fear or intense discomfort that reaches peak within minutes, and during which time four or more of a list of 13 physical and cognitive symptoms occur. The term *recurrent* literally means more than one unexpected panic attack. The term *unexpected* refers to a panic attack for which there is no obvious cue or trigger at the time of occurrence – that is, the attack appears to occur from out of the blue, such as when the individual is relaxing or emerging from sleep (nocturnal panic attack) (Philippine Psychiatric Association, 2017).

### CLINICAL CRITERIA:

Recurrent unexpected panic attacks – Palpitations, sweating, trembling/shaking, sensations of shortness of breath/smothering, feelings of choking, chest pain/discomfort, nausea or abdominal distress, feeling dizzy/faint, chills/heat sensations, paresthesias, de-realization or depersonalization, fear of losing control, fear of dying (PPA, 2017).

# B. GENERALIZED ANXIETY DISORDER

The essential feature of generalized anxiety disorder (GAD) is excessive anxiety and worry about several events or activities. The intensity, duration, or frequency of the anxiety and worry is disproportionate to the actual likelihood or impact of the anticipated event. An individual with GAD finds it difficult to control the worry and to keep worrisome thoughts from interfering with attention to tasks at hand (PPA, 2017).

### **CLINICAL CRITERIA:**

Should have a duration of more than 6 months that affects his/her social or occupational functioning, excessive anxiety and worry, difficulty in controlling worry, 3 or more of the following: restlessness, easily fatigued, difficulty, irritability, muscle tension, sleep disturbance (PPA, 2017).

# C. SOCIAL ANXIETY DISORDER

The essential feature of social anxiety disorder (SAD) is a marked or intense fear, or anxiety of social situations in which the individual may be scrutinized by others (PPA, 2017).

# **CLINICAL CRITERIA:**

First-Line | Selective Serotonin Reuptake Inhibitors (SSRIs)

Marked fear or anxiety about one or more social situations in which the individual is exposed to possible scrutiny by others. The fear, anxiety or avoidance is persistent, typically 6 months or more.

**Treatment Choice** 

# II. PREFERRED REGIMEN

2.1.00	- SSRIs are e and in imp types of SS effect profi disorder (P)	roving associated depressive mood. It RIs in dealing with panic disorder. Do le of SSRIs, they are recommended as	tensity of panic attacks, anticipatory anxiety, No difference in efficacy is seen in different Due to the relatively favorable safety and side is the best initial choice for patients with panic
	Medication	Caution	Side Effects
	Sertraline	Avoid use when driving, operating	Common:
		machineries, etc. (due to sedation)	Erectile dysfunction in men, decreased sexual desires in women, gastrointestinal
		Drug-lab interaction:	disturbances (decreased appetite, nausea,
		Sertraline can cause a false positive	diarrhea, and constipation), insomnia,
		urine drug screen for	sedation, agitation, tremors, headache,
		benzodiazepines	dizziness. May cause prolonged erection
			and delayed ejaculation
			Serious:
		/	Bruising and rare bleeding, rare
		•	hyponatremia, rare hypotension
!	Fluoxetine	Caution in persons with history of	Common:
		seizure (can lower the seizure	Insomnia, headache, dizziness,
		threshold)	gastrointestinal disturbances, changes in
			appetite, and sexual dysfunction.
	/	Drug-drug interactions: Avoid	
		combination with warfarin	Serious:
		(may increase bleeding risk).	Bleeding abnormalities in those who use
		May increase levels of TCAs,	aspirin or other non-steroidal anti-
		antipsychotics, and beta-blockers.	inflammatory drugs, low sodium levels.
		Caution in combination with	
		tamoxifen, codeine, and tramadol	
		(reduces the effect of these drugs).	
	Escitalopram	Use with caution among elderly.	Common:
Į.		(Elderly are more prone to	Nausea, sweating, somnolence, dizziness,
		SSRI/SNRI induced	insomnia, constipation, diarrhea, appetite
		hyponatremia)	decrease, sexual dysfunction, fatigue,
i	- 1/4		pyrexia.

		Used with caution in patient using	
		medication that will affect clotting	
		of blood.	
		Caution with other QTc-prolonging	
		medications like some	
		antipsychotics, macrolides,	
		fluoroquinolones, ondansetron, and	
		HIV protease inhibitors.	
Second-	Atypical Antips	ychotics	
Line		· · ·	ts to efficacy of quetiapine in the treatment of
	GAD and be	eing equally efficacious as antidepres	sants. However, quetiapine use is associated
			ed to placebo or antidepressants. As this might
	affect the pa	tient's adherence, quetiapine is recom	mended as a second line treatment for GAD
	for those who	o cannot tolerate the use of SSRIs (PP.	A, 2017).
	for those who	cannot tolerate the use of SSRIs (PP.  Caution	A, 2017). Side Effects
			Side Effects Common:
	Medication	Caution	Side Effects Common:
	Medication	Caution  Use with caution among elderly.	Side Effects Common:
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain),
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain,
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain),
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)  Caution with other QT-prolonging	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)  Caution with other QT-prolonging medications like fluoroquinolones,	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)  Caution with other QT-prolonging medications like fluoroquinolones, macrolides, ondansetron, and HIV	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)  Caution with other QT-prolonging medications like fluoroquinolones, macrolides, ondansetron, and HIV	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased
	Medication	Caution  Use with caution among elderly. (Elderly are more prone to SSRI/SNRI induced hyponatremia)  Caution with other QT-prolonging medications like fluoroquinolones, macrolides, ondansetron, and HIV protease inhibitors.	Side Effects  Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased

# III. TREATMENT DOSE

Disorder	Dosages				
	1 <sup>st</sup> Line			2 <sup>nd</sup> Line	
	Sertraline	Fluoxetine	Escitalopram	Quetiapine	
Panic	Initial dose:	Initial dose:	Initial dose:	Not recommended as	
Disorder	12.5 mg (Oral)	20 mg capsule (Oral)	10 mg (Oral)	second line. May be used as third line.	
	Maximum	Maximum dose:	Maximum dose:		
	dose: 150-200 mg	80 mg daily* (Oral)	20 mg/day (Oral)		
	(Oral)	*Total Daily dose >20 mg/day should be divided in two (2) separate doses, each given every 12 hours.			

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# PHARMACOTHERAPEUTIC GUIDELINES OF THE MEDICINE ACCESS PROGRAM FOR MENTAL HEALTH (MAP-MH)

Generalized	Initial dose:	Initial dose:	Initial dose:	Initial dose:
Anxiety	12.5 mg (Oral)	20 mg (Oral)	10 mg (Oral)	25 mg (Oral)
Disorder				
(GAD)	Maximum	Maximum dose:	Maximum dose:	Maximum dose:
	dose:	80 mg daily* (Oral)	20 mg/day (Oral)	150 mg (Oral)
	150-200 mg			
	(Oral)	*Total Daily dose >20		
		mg/day should be divided in		
		two (2) separate doses, each		
		given every 12 hours.		
Social	Initial dose:	Initial dose:	Initial dose:	Initial dose:
Anxiety	12.5 mg (Oral)	10 mg (Oral)	10 mg (Oral)	25 mg (Oral)
Disorder				
	Maximum	Maximum dose:	Maximum dose:	Maximum dose:
	dose:	80 mg daily* (Oral)	20 mg/day (Oral)	150 mg (Oral)
	150-200 mg			,
	(Oral)	*Total Daily dose >20		
		mg/day should be divided in		
		two (2) separate doses, each		
		given every 12 hours.		

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# CONSENSUS TREATMENT GUIDELINES ON PSYCHOSIS

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# **PSYCHOSIS**

Psychosis is characterized by distorted thoughts and perceptions, as well as disturbed emotions and behaviors. Incoherent or irrelevant speech may also be present. Symptoms such as hallucinations-fixed, false belief; severe abnormalities of behavior-disorganized behavior, agitation, excitement, inactivity or hyperactivity; disturbance in emotions-marked apathy, or disconnect between reported emotion and observed affect, such as facial expression and body language, may also be detected. The most common disorder is schizophrenia.

# A. SCHIZOPHRENIA

Schizophrenia is characterized by delusions, hallucinations, disorganized speech and behavior, and other symptoms that cause social or occupational dysfunction. For diagnosis to be made, symptoms must have been present for six months with at least one month of active symptoms.

### **CLINICAL CRITERIA:**

Two or more of the following, each present for a significant portion of time during a 1-month period: delusions, hallucinations, disorganized speech, grossly disorganized or catatonic behavior, negative symptoms (diminished emotional expression or avolition). Continuous signs of disturbance persist for at least 6 months.

The course of treatment and management of schizophrenia can be divided into three phases, each with corresponding goals: (1) acute phase, (2) stabilization phase, (3) maintenance phase.

### ACUTE PHASE

The Acute Phase is defined as the period of an acute psychotic episode. This phase begins with a new onset or acute exacerbation of symptoms and lasts until symptoms are reduced to a level considered to be the patient's expected "baseline". Before administration of medications, simple interventions such as reorientation and de-escalation techniques may be done to assist in defusing the situation (Castle et al., 2017). The patient's Advance Directive should also be taken into consideration.

For patient in the acute phase, Rapid Neuroleptization can be performed which is a common method to calm an agitated patient with Schizophrenia (Battaglia, 1997). It is defined further as a method of administering repeated doses of medication under close clinical supervision that provides rapid control of acute functional psychotic symptoms (Donion, 1979).

### 2. STABILIZATION PHASE

The stabilization period follows the acute phase and constitutes a time-limited transition to continuing treatment in the stable phase (APA Guidelines, 2010). The stabilization phase will be defined as the phase occurring from the 3<sup>rd</sup> to 6<sup>th</sup> months. Recommendation to Pharmacotherapy is to continue the same dose of antipsychotic medication which produced an adequate response for at least 6 months.

### 3 MAINTENANCE PHASE

The maintenance phase aims to sustain the patient's control of symptoms and remission.

# B. PSYCHOSIS DUE TO OTHER MENTAL DISORDERS

Psychoses could be seen in several mental disorder-like severe cases of depression. They can also be seen in substance use disorder, seizure, dementia, etc.

### **CLINICAL CRITERIA:**

Patient who present with psychosis and was diagnosed of having mental disorder aside from Schizophrenia like major depression, substance use disorder, seizure, dementia, HIV, etc.

Psychosis having marked Behavioral Changes; neglecting usual responsibilities related to work, school, domestic or social activities, agitated and aggressive behavior with decreased or increased activity, fixed false beliefs not shared by others in the person's culture, hearing voices or seeing things that are not there.

# II. PREFERRED REGIMEN

		Treatment Choice	
First Line	re-hospitalizat of positive sy	ts are associated with lower risk of n ion. Both conventional and atypical a	nedication change, medication gaps and agents are associated with improvement bjects of atypical agents at follow-up symptoms. (Serra-Arain, et.al 2015)
	Medication	Cautions	Side Effects
	Risperidone (tablet or ODT*)	Use with caution in patients with cardiac disease.	Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal
	*ODT is used for extremely agitated patients	For elderly patients: start with 1 mg/day	cramps, diarrhea, constipation), other effects (dyspepsia, epistaxis, abnormal vision), cardiac dysrhythmia – QT
		Drug-drug interactions: Carbamazepine can reduce levels of risperidone, whereas fluoxetine can increase levels.	Prolongation, endocrine – diabetes mellitus (increased cholesterol and FBS)
		Monitor weight, BP, FBS, and lipid profile, if possible.	

J.	Olanzapine	Use with caution in patients with	Common:
	(tablet or ODT*)	diabetes mellitus (DM), seizures,	CNS (dizziness, drowsiness, ataxia,
		benign prostatic hyperplasia	headache), GI (dry mouth, abdominal
	*ODT is used for	(BPH), narrow angle glaucoma,	pain, diarrhea, constipation), hepatic
	extremely	and hepatic Disease	(increase in alanine aminotransferase /
	agitated patients		ALT – 3 times greater), endocrine –
		Decreased serum concentrations up	diabetes mellitus (increased
		to 50% with cigarette smoke	cholesterol and FBS)
		Monitor weight, BP, FBS, and	
		lipid profile, if possible.	
	Quetiapine	Use with caution for elderly	Common:
		patients.	CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal
		Caution with other QT-prolonging	pain, diarrhea, constipation, weight
		medications like fluoroquinolones,	gain), endocrine – diabetes mellitus
		macrolides, ondansetron, and HIV	(increased cholesterol and FBS)
		protease inhibitors.	(mercased enoiesteror and 1 BS)
		protease numbrors.	
		Monitor weight, BP, FBS, and lipid	
		profile, if possible.	
	1		
Second	Typical Antipsycl	hotic	
Second Line	Typical Antipsycl - Typical antip		ered since the risks and benefits of
	- Typical antip	osychotic medications may be conside	
	- Typical antip	osychotic medications may be consider ments for each patient change over tire	ered since the risks and benefits of me. It can improve positive symptoms
	- Typical antip various treat of patients w	psychotic medications may be consider ments for each patient change over tire with schizophrenia (PPA, 2017).	ne. It can improve positive symptoms
	- Typical antipy various treats of patients we Medication	psychotic medications may be consider ments for each patient change over tire with schizophrenia (PPA, 2017).  Cautions	ne. It can improve positive symptoms  Side Effects
	- Typical antip various treat of patients w	osychotic medications may be consider ments for each patient change over tire with schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic	Side Effects Common:
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the consideration of the considera	Side Effects Common: Sedation, anticholinergic effect (dry
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the consideration of the considera	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia,
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the consideration of the considera	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision),
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the consideration of the considera	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS),
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia,
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities,
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome,
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS
	- Typical antipy various treats of patients we Medication	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common:
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly patients.	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term therapy). Involuntary movements of
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly patients.  Contraindications:1	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term therapy). Involuntary movements of extremities may also occur. Dry
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly patients.  Contraindications:  Bone marrow suppression,	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term therapy). Involuntary movements of extremities may also occur. Dry mouth, constipation, urinary retention,
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly patients.  Contraindications:1	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term therapy). Involuntary movements of extremities may also occur. Dry mouth, constipation, urinary retention, mydriasis, agitation, insomnia,
	- Typical antipy various treats of patients we Medication Haloperidol	considerations may be considerated by the schizophrenia (PPA, 2017).  Cautions  Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended  Sedative effects are most marked during the few days of administration  Use with caution for elderly patients.  Contraindications:  Bone marrow suppression,	Side Effects  Common: Sedation, anticholinergic effect (dry mouth, urinary retention, tachycardia, constipation. blurred vision), extrapyramidal symptoms (EPS), Weight Gain, tardive dyskinesia, photosensitivity, EEG abnormalities, neuroleptic malignant syndrome, increased susceptibility to developing EPS  Common: Tardive dyskinesia (on long-term therapy). Involuntary movements of extremities may also occur. Dry mouth, constipation, urinary retention,

<sup>&</sup>lt;sup>1</sup> MIMS Philippines (n.d.): Chlorpromazine, retrieved from: https://www.mims.com/philippines/drug/info/chlorpromazine/?type=brief&mtype=generic

Drug-lab interaction:	skin reaction, amenorrhea,
Chlorpromazine can cause a false positive in urine drug screening for	,
amphetamines.	
	Fatal:
	Agranulocytosis, neuroleptic
	malignant syndrome, extrapyramidal
	dysfunction.

# III. TREATMENT DOSE

Disorder			Dosages		
	1st Line				d Line
	Risperidone	Olanzapine	Quetiapine	Haloperidol	Chlorpromazine
Schizophrenia					
a. Acute	Initial dose:	Initial dose:	Initial dose:	Initial dose:	Initial dose:
Phase*	1-2 mg/day	10 mg/day	200 mg/day	5 mg/day	75-100 mg / day
	(Oral)	(Oral)	(Oral)	(Oral)	(Oral)
	Maximum	Maximum	Maximum	Maximum	Maximum dose:
	dose:	dose:	dose:	dose:	400 mg/day
	6 mg/day	20 mg/day	800 mg/day	20 mg/day	(Oral)
	(Oral)	(Oral)	(Oral)	(Oral)	
	*For elderly start with I mg/day				
b. Stabilization	Initial dose:	Initial dose:	Initial dose:	Initial dose:	Initial dose:
Phase	1 mg/day	10 mg/day	300-400	1.5 - 3  mg	25-50 mg daily
	(Oral)	(Oral)	mg/day (Oral)	daily (Oral)	(Oral)
			Maximum	Maximum	
	Maximum	Maximum	dose:	dose:	Maximum dose:
	dosé:	dose:	800 mg/day	20 mg daily	300 mg daily (up
	4 mg/day	20 mg/day	(Oral)	(Oral)	to 1000 mg may
	(Oral)	(Oral)	()	(0101)	be necessary for
					severe cases)
					(Oral)
c. Maintenance			Same as above		
Phase			ru		
Psychosis Due to	Initial dose:	Initial dose:	Initial dose:	Initial dose:	Initial dose:
Other Mental	0.5 mg/day	2.5 mg/day	300-800	20-60 mg/day	300-1000 mg/
Disorders	(Oral)	(Oral)	mg/day (Oral)	(Oral)	day (Oral)

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## Special Considerations:

- In women with psychosis who are planning pregnancy or pregnant or breastfeeding, low-dose oral haloperidol or chlorpromazine may be considered.
- In adolescent with psychotic or bipolar disorder, risperidone can be offered as a treatment option only under supervision of a specialist, if treatment with risperidone is not feasible, haloperidol or chlorpromazine may be used only under supervision of a specialist.
- Antipsychotics carry an increased risk of cerebrovascular events and death in older adults with dementia-related psychosis.
- In those with Parkinson's disease psychosis, clozapine may be less likely to worsen Parkinson's symptoms (given proper monitoring is done) and quetiapine may be useful.
- Try the medication at a typically effective dose for at least 4-6 weeks with adherence and interactions noted before considering it ineffective.

# C. MANAGEMENT FOR TREATMENT-RESISTANCE AND/OR NON-ADHERENCE

Due to the extensive treatment and monitoring of the treatment phase, existence of treatment resistance and non-adherence. The efficient treatment course for such incidences involves the use of Long Acting Injectable (LAI) and other anti-psychotics.

# I. PREFERRED REGIMEN

	<u> </u>	Treatn	ent Choice	
First Line	1 -	onsidered for patients		nerence with oral medications. It minimizes fluctuations in blood
	Medication	Dosage and Frequency	Cautions	Side Effects
	Fluphenazine decanoate 25 mg, 1 ml Ampule (Intramuscular)*	12.5 mg – 50 mg every 2-4 weeks	Contraindications: Impaired Consciousness, Parkinsonism  Cautions in patients with: Cardiac Disease, Kidney Disease, Liver Disease. Use with caution in older adults	Common: Sedation, dizziness, blurred vision, dry mouth, urinary retention, constipation, tachycardia  Serious: Orthostatic hypotension, syncope, extrapyramidal symptoms, photosensitivity, weight gain, galactorrhea,

Flupenthixol	20mg-40mg every 2	Drug to Drug Interactions: Increases effects to blood pressure lowering medications Can lower blood pressure if used with epinephrine Contraindications:	amenorrhea, sexual dysfunction, priapism, neuroleptic malignant syndrome (NMS), agranulocytosis, jaundice  Common and serious:
decanoate 20mg/mL, amplue (Intramuscularly)	to 4 weeks  (Initial dose of 20mg for patients who have not been exposed to long-acting depot antipsychotics, 40mg for patients who have previously demonstrated tolerance to long-acting depot antipsychotics; after 4-10 days can give additional 20mg dose; maximum of 200mg every 1-4 weeks.)	<ul> <li>diminished consciousness due to any cause</li> <li>collapse due to very low blood pressure</li> <li>brain damage</li> <li>diseases of the blood with a reduced number of red or white blood cells or platelets</li> <li>phaeochromocytoma, a rare tumour of the adrenal gland which sits near the kidney.</li> <li>Do not give Fluanxol to anyone who currently has alcohol poisoning, or poisoning with medicines used to produce calmness or to help you sleep, or medicines used to treat epilepsy or strong pain.</li> <li>Do not give Fluanxol to anyone who is unconscious or in a coma.</li> <li>Do not give Fluanxol to a child or adolescent or in pregnant women.</li> <li>Cautions in patients with:         <ul> <li>Renal impairment, hepatic impairment, cardiac impairment, and elderly.</li> </ul> </li> <li>Drug to Drug Interactions:</li> </ul>	Neuroleptic-induced deficit syndrome Extrapyramidal symptoms (more common at the start of treatment), parkinsonism Insomnia, restlessness, agitation, sedation Tardive dyskinesia (risk increases with duration of treatment and with dose) Galactorrhea, amenorrhea
 			<del></del>

may decrease the
Paliperidone palmitate 78, 117, 156, 234 mg prefilled syringes  Starting dose. 25 mg (25 – 100 mg) No need for oral supplementation. Some elderly patients may tolerate lower doses better  Paliperidone palmitate 78, 117, 156, 234 mg prefilled syringes  Starting dose. 25 mg (25 – 100 mg) No need for oral supplementation. Some elderly patients may tolerate lower doses better  Precautions:  Some elderly patients may tolerate lower doses better  Procautions:  Use with caution in patients with sondificate that predispose to hypotension (delydration, overheating) Dysphagia has been associated with antipsychotics use, and paliperidone should be used cautiously in patients with antipsychotics, more than for others.  Precautions:  Some elderly patients may tolerate lower doses better  Precautions:  Use with caution in patients with captione or insperidone Procession (delydration, overheating) Dysphagia has been associated with antipsychotics use, and paliperidone should be used cautiously in patients with antipsychotics, more than for others.

- aspiration pneumonia
- Paliperidone prolongs QTc interval more than some other antipsychotics
- Priapism has been reported with other antipsychotics, including risperidone

### Contraindications:

- If patient is taking agents capable of significantly prolonging QTc interval (eg. pimozide, thioridazine, selected antiarrhythmics, moxifloxacin, sparfloxacin)
- If there is a history of QTc prolongation or cardiac arrhythmia, recent acute myocardial infarction, uncompensated heart failure
- If patient has a preexisting severe gastrointestinal narrowing
- If there is a proven allergy to paliperidone or risperidone

# **Drug Interactions:**

- May increase effects of antihypertensive agents
- May antagonize levodopa, dopamine agonists
   May enhance QTc

prolongation of other drugs capable of prolonging QTc interval

# Life-threatening or Dangerous side effects

- Hyperglycemia, in some extreme cases and associated with ketoacidosis or hyperosmolar coma or death, has been reported in patients taking atypical antipsychotics
- Increased risk of death and cerebrovascular events in elderly patients with dementia-related psychosis
- Rare neuroleptic malignant syndrome (much reduced risk compared to conventional antipsychotics)
- Rare seizure

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Clozapine 100	12.5 mg to 100 mg	Contraindications: <sup>2</sup>	Common:
mg tablet PO		Impaired bone	Decreased gastrointestinal
		marrow function,	motility, urinary retention,
		uncontrolled epilepsy,	dyslipidemia, extrapyramidal
		CNS depression,	symptoms, deep vein
		severe cardiac	thrombosis
		disorder, severe renal	
		and hepatic	Serious:
		impairment	Orthostatic Hypotension,
			bradycardia, Severe
		Drug to Drug	agranulocytosis/neutropenia,
		Interactions:	myocarditis and
		- Increased risk of	cardiomyopathy,
		neuroleptic	hepatotoxicity including
		malignant	hepatic failure, hepatic
		syndrome with	necrosis and hepatitis,
		lithium	torsade de pointes, cardiac
1		- Increased risk of	arrest, neuroleptic malignant
		seizures with	syndrome, respiratory
		valproic acid	depression or failure,
		- Increased risk of	paralytic ileus, intestinal
		myelosuppression	obstruction
		with long-acting	
		depot	
		antispychotics	

# **II. TREATMENT DOSE**

Disorder	Dosages			
	1 <sup>st</sup> Line	2 <sup>nd</sup> Line		
	Fluphenazine decanoate	Clozapine		
Treatment Resistance / Non-Adherence	Initial dose: 12.5 mg every 2-4 weeks(Intramuscular in the gluteal muscle)	,		
		Maximum dose:450 mg/day(up to 900 mg may be necessary for severe cases) (Oral) in divided doses		

<sup>&</sup>lt;sup>2</sup> MIMS Philippines (n.d.): Clozapine, retrieved from: https://www.mims.com/philippines/drug/info/clozapine?mtype=generic

# CONSENSUS TREATMENT GUIDELINES ON MOOD DISORDERS

# MOOD DISORDERS

# A. MAJOR DEPRESSIVE DISORDER

Major Depressive Disorder (MDD) is a chronic, brain disorder characterized by low mood and loss of interest that can affect a person's thoughts, behavior, feelings, and social functioning. As such, depression poses a great impact to a person's relationships and productivity. It may also complicate existing medical conditions adding to the burden of chronic diseases and may lead to possible suicide.

### **CLINICAL FEATURES:**

MDD is characterized by either a depressed mood or loss of interest or pleasure as the core symptoms, persisting for at least 2 weeks and associated with a change from previous functioning.

There must have a significant effect on the individual's functioning and must have at least five other symptoms of depression: Depressed mood, markedly diminished interest or pleasure in almost all activities, significant weight loss, insomnia, psychomotor agitation or retardation, fatigue or loss of energy, feelings of worthlessness or excessive or inappropriate guilt, diminished ability to think or concentrate or indecisiveness.

An adequate trial of an anti-depressant should be a minimum of three weeks at the recommended therapeutic dose. If there is no improvement at 2 to 4 weeks of adequate antidepressant dose, it is recommended to increase dose or to switch to another antidepressant if the patient experiences intolerable side effects (Kennedy et al, 2016). Once remission sets in, maintenance antidepressant treatment for at least six months up to one year is recommended.

# II. PREFERRÉD REGIMEN

	Treatment Choice					
First Line	<ul> <li>Selective Serotonin Reuptake Inhibitors (SSRIs)</li> <li>These are the first-line treatment recommendations for adult MDD (e.g. escitalopram, sertraline, fluoxetine).</li> </ul>					
	Medication	Dosage and Frequency	Cautions	Side Effects		
	Escitalopram	20 - 60 mg/day	Elderly are more prone to SSRI induced	*Adverse effects are usually transient:		
		Elderly/Medically Ill: Preferred	Hyponatremia	Nausea, Sweating, Somnolence, Dizziness,		
		Choice, Start 10 mg daily, then increase to 20 mg	Used with caution in patient medication that will affect clotting of blood.	Insomnia, Constipation, Diarrhea, Appetite Increase,		

# PHARMACOTHERAPEUTIC GUIDELINES OF THE MEDICINE ACCESS PROGRAM FOR MENTAL HEALTH (MAP-MH)

			Do not take at night unless sedation occurs.	
	Fluoxetine	Adult: Start 10 mg daily for one week then 20 mg daily. If no response in 6 weeks, increase	Elderly are more prone to SSRI induced Hyponatremia	Headache, Nausea, Incomnia, Anorexia, Anxiety, Asthenia, Diarrhea, Nervousness, Somnolence
		to 40 mg (maximum of 80 mg)	Combination with Tamoxifen, codeine and tramadol can lead to treatment failure	Less Common: Dizziness, Dry mouth, dyspepsia, sweating, tremor, decreased libido, abnormal taste, agitation, chest pain, sleep disorder
		Adolescent: Start 10 mg daily. Increase 20 mg daily if no response in 6 weeks (maximum 40 mg)	Do not take at night unless sedation occurs  Cautions in persons with history of seizure.	Serious: Bleeding abnormalities in those who use aspirin or other non-steroidal anti-inflammatory drugs, low sodium levels
			Drug-Drug Interactions: Avoid combination	
		Elderly/Medically Ill: Preferred Choice, Start 10 mg daily, then increase to 20 mg (maximum of 40 mg)	with warfarin (may increase levels of TCAs, antipsychotics and betablockers.	
	Sertraline	Initial Dose: 12.5 mg  Maximum Dose:	Elderly are more prone to SSRI induced Hyponatremia	Headache, Somnolence, drowsiness, fatigue, dizziness, insomnia, tremor, anxiety, paresthesia, agitation, sexual
		150-200 mg  Elderly: maximum dose = 50 mg/day	Do not take at night unless sedation occurs	dysfunction, nausea, dry mouth, diarrhea, constipation, abnormal vision
Second -	Atypical antipsy	chotic	<u> </u>	<u></u>
Line	The conventional and atypical antipsychotics show modest efficacy in the treatment of psychosis, agitation, and aggression in patient with depression.			
	Quetiapine	ion, and aggression in p	Special Instructions:	Common:
	Anenahine	150-5001118	Metabolic Effects Monitoring (need specificity)	CNS (dizziness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine (increased cholesterol and FBS)

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# III. TREATMENT DOSE

Disorder	Dosages				
		2 <sup>nd</sup> Line			
	Escitalopram	Fluoxetine	Sertraline	Quetiapine	
Major	Initial dose:	Adult:	Initial dose:	150-300 mg/day	
Depressive	10-20 mg/day	Start 10 mg once daily for one	12.5 mg once		
Disorder	(Oral)	week then increase to 20 mg once	daily (Oral)		
		daily. If no response in 6 weeks,			
	Elderly/Medically	increase to 40 mg once daily	Maximum dose:		
	Ill: Start 10 mg	(Maximum dose: 80 mg/day)	150 – 200 mg		
	once daily, then		once daily		
	increase to 20 mg	Adolescent:			
		Start 10 mg once daily for one	*Elderly		
		week then increase to 20 mg once	maximum dose:		
		daily if no response in 6 weeks	50 mg/day		
		(Maximum dose: 40 mg/day)			
		Elderly:			
		Start 10 mg once daily then			
		increase to 20 mg once daily			
		(Maximum dose: 40 mg/day)			

## **Special Considerations:**

- Antidepressant medications usually need to be continued for at least 9 12 months after resolution of symptoms
- If the person develops a manic episode stop the antidepressant immediately, it may trigger a manic episode in untreated bipolar disorder
- Do not combine with other depressants and serotonergic agents (e.g. tramadol, ginseng, St. John's wort, etc.), this may cause serotonin syndrome (these conditions consist of a combination of mental status changes, neuromuscular hyperactivity and autonomic hyperactivity)<sup>3</sup>
- Antidepressants may increase suicidal ideation, especially in adolescents and young adults of aged 24 and below, however the effect decreases amongst adults ages 25 and above.
- If symptoms persist or worsen despite interventions, consider FLUOXETINE. Ask adolescent to return weekly for the first 4 weeks to monitor thoughts or plans of suicide.
- If the woman is breastfeeding, avoid long acting anti-depressant medication such as Fluoxetine.
- For adults with thoughts or plans of suicide SSRIs are the first choice.

<sup>&</sup>lt;sup>3</sup> Volpi-Abadie, J., et al (2013): Serotonin Syndrome, National Institute of Health, The Oshner Journal Vol 13(4) pp 553-540, retrieved from: https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3865832/

# B. BIPOLAR DISORDER4

Bipolar Mood Disorder is a chronic mental health condition characterized by manic and/or hypomanic episodes, either with or without depressive episodes, which cause significant impairment in social, occupational and/or academic functioning. It is often co-morbid with other psychiatric conditions such as anxiety disorders, substance use disorders, personality disorders, and attention deficit hyperactivity disorder.

### 1. BIPOLAR I DISORDER

### **CLINICAL CRITERIA:**

The DSM-5 criteria for Bipolar I disorder require at least one episode of mania. In most cases, the manic episode may be preceded or followed by a hypomanic or depressive episode.

### 2. BIPOLAR MOOD DISORDER WITH MIXED FEATURES

### **CLINICAL CRITERIA:**

Manic or Hypomanic Episodes with mixed features are characterized by episodes that meet all criteria for mania or hypomania, with at least three of the following symptoms during most days of the episode: Depressed mood, Diminished interest or pleasure in most activities, Psychomotor Retardation, Low Energy, Excessive Guilt or Thoughts of Worthlessness, Recurrent thoughts about death or suicide, or suicide attempt.

\*The purpose of the treatment is similar and thus treated with same medications

# 3. HYPOMANIA

### **CLINICAL CRITERIA:**

This is associated with unequivocal change in functioning that is uncharacteristic of the individual when not symptomatic. It is not severe enough to cause marked impairment in social or occupational functioning, or to necessitate hospitalization.

\*Clinical practice suggests that Mood Stabilizers used in acute mania are also effective in hypomanic episodes.

# 4. RAPID CYCLING BIPOLAR MOOD DISORDER

### **CLINICAL CRITERIA:**

At least four mood episodes in a 12-month period, the episodes meet both the symptom and duration criteria for mania, hypomania, or major depression; the episodes that occur as part of a rapid cycling pattern are no different from episodes that occur as part of a non-rapid cycling pattern; the episodes are demarcated by a period of partial or full remission for at least two months, or by a switch to an episode of opposite polarity.

<sup>&</sup>lt;sup>4</sup> PPA (February 2017): Consensus Treatment Guidelines on Bipolar Mood Disorder.

The treatment of manic and hypomanic episodes in patients with rapid cycling bipolar disorder is comparable to the treatment of mania or hypomania in general. The choice of treatment for rapid cycling is governed by current mood state and the need to prevent depressive state. The treatment options include valproate, lithium, olanzapine, and lamotrigine. Quetiapine has also been shown to be effective.

# I. PREFERRED REGIMEN

Treatment Choice					
First Line	Mood Stabilizers + Anti-Psychotics				
			ood and psychotic symptom		
		polar Mood Disorder is	s preferably a combination	of Mood Stabilizers and	
	Anti-Psychotics.				
			n to be more efficacious that	n Lithium in the	
		and psychotic manic e		\$ -MICROSCO	
	Medication	Dosage and	Cautions	Side Effects	
		Frequency			
	Divalproex Sodium	600 - 750  mg/day	Contraindicated in	Common:	
	ER Tablet	in divided disease.	patients with liver	Sedation, Headache,	
		Increase dose by	disease or severe liver	tremor, ataxia, nausea,	
		200 – 500 mg/day	dysfunction; take with	vomiting, diarrhea,	
		at 3 day intervals	large amount of water or	weight gain, transient	
		until desired	food to avoid GI upset;	hair loss.	
		response.	caution on Pregnant		
		/	women, Monitor liver	*Caution on pregnant	
		Or ′	function during the 1st	women / breastfeeding	
			six months of therapy	women	
		Administer loading	and monitor platelet		
		dose of 20	function before major	Serious:	
		/mg/kg/day in	surgery.	Impaired hepatic	
	,	divided doses		function,	
	/		Drug to Drug	thrombocytopenia,	
			Interactions: Valproate	leucopenia,	
			levels decreased by	drowsiness/confusion,	
	,		Carbamazepine	liver failure,	
				hemorrhagic	
				pancreatitis	
	+ Risperidone	0.5 mg	Special Instructions:	Common:	
			Fasting Blood Sugar	CNS (dizziness,	
			(GBS) and lipid profile	drowsiness, ataxia,	
			monitoring	headache), GI (dry	
				mouth, abdominal	
			Avoid abrupt	cramps, diarrhea,	
			withdrawal	constipation), Other	
				effects (Dyspepsia,	
			Cautions in patients with	epistaxis, abnormal	
			cardiac disease.	vision)	
L			<u> </u>		

# PHARMACOTHERAPEUTIC GUIDELINES OF THE MEDICINE ACCESS PROGRAM FOR MENTAL HEALTH (MAP-MH)

	+ Olanzapine	5 mg	Drug-Drug Interactions: Carbamazepine can reduce levels of risperidone, whereas fluoxetine can increase levels. Use with caution in patients with Diabetes Mellitus (DM), seizures, Benign Prostatic Hyperplasia (BPH), Narrow Angle Glaucoma	Common: CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation), Hepatic (increase in Alanine Aminotransferase –
Second- Line	Lithium Carbonate	In Acute Mania: 450 – 2000 mg in divided doses in 5-7 days	Severe cardiac or kidney disease. Dehydration can increase lithium levels.  Drug-Drug Interactions: Nonsteroidal Antiinflammatory Drugs (NSAIDs), Angiotensin-Converting Enzyme Inhibitor (ACE Inhibitor), thiazide diuretics, Metronidazole, and Tetracycline can increase lithium levels.  Lithium toxicity can cause seizures, delirium, coma and death.	Common: Sedation, cognitive problems, tremor, impaired coordination, hypotension, leukocytosis, polyuria, polydipsia, nausea, diarrhea, weight gain, hair loss, rash.  Serious: Diabetes insipidus, hypothyroidism, ECG changes (arrhythmia, sick sinus syndrome, twave changes)
	+ Haloperidol	0.5 – 2 mg	Watch out for orthostatic hypotension, increase water intake, regular eye exams recommended, Sedative effects are most marked during the few days of administration	Sedation, anticholinergic effect (dry mouth, urinary retention. Tachycardia, constipation. Blurred vision), EPS, Weight Gain, Tardive dyskinesia, photosensitivity, EEG Abnormalities

# PHARMACOTHERAPEUTIC GUIDELINES OF THE MEDICINE ACCESS PROGRAM FOR MENTAL HEALTH (MAP-MH)

		Caution in patients with	
		kidney disease, liver	
		disease, cardiac disease,	
		long QT syndrome or	
		taking QT-prolonging	
		medications. Monitor	
		ECG if possible.	
+ Chlropromazine	10 – 25 mg	Watch out for orthostatic	Sedation,
Chitopromazine	10 25 mg	hypotension, increase	Anticholinergic effect
	May increase to 25-	water intake	(dry mouth, urinary
	•	water intake	
	100 mg to	Decrete and average	retention, tachycardia,
	maintenance dose	Regular eye exams	constipation, blurred
	of 100-300 mg	recommended	vision), EPS, Weight
		G 1 GG	Gain, Tardive
		Sedative effects are most	Dyskinesia,
		marked during the few	Photosensitivity, EEG
		days of administration	abnormalities
		Contraindications:	
		impaired consciousness,	
		bone marrow	
		depression,	
		pheochromocytoma	
		Caution in Patients with	
		respiratory disease,	
		kidney disease, liver	
		disease, glaucoma,	
		urinary retention,	
		cardiac disease, long QT	
		syndrome or taking QT-	1
		prolonging medications.	]
		Monitor ECG if	
		possible.	
		possible.	
		Dana Dana	
		Drug-Drug Interactions:	
		- Increases effects of	
		blood pressure	
		<u> </u>	
		lowering	
		medications	
		- Lowers blood	
		pressure if	
		combined with	
		epinephrine	
		Levels may be increased	
		by antimalarial	
		including quinine	

# II. TREATMENT DOSE

Disorder	Dosages				
	1 <sup>st</sup>	Line	2 <sup>nd</sup> Line		
	Divalpro	ex Sodium	Lithiun	Carbonate	
Bipolar I	600 – 750 mg/day in divi	ded dose. Increase dose by	In Acute M	ania: 450 – 2000	
	200 - 500 mg/day at 3-6	mg/day in divid	led doses in 5-7 days		
Bipolar Mood	respons	se. (Oral)		•	
Disorder with	(	OR			
Mixed	Administer loading dose	of 20 mg/kg/day in divided			
Features	do	oses			
	+ Risperidone	+ Olanzapine	+ Haloperidol	+ Chlorpromazine	
Hypomania	0.5 –2mg/day	5 – 10 mg/day	0.5-2 mg/day	10-25 mg/day	
				Non-in-	
				May increase to	
				25-100 mg/day to	
				maintenance dose	
				of 100-300 mg/day	

# A.BIPOLAR II

Bipolar II is characterized by the predominance of depressive episodes, its treatment mirrors the treatment of bipolar I depression.

In Bipolar II Disorder, both hypomanic episode and major depressive episode must be present to meet the criteria.

# I. PREFERRED REGIMEN

	Treatment Choice					
First Line	Treatment for acute depressive episode of Bipolar II disorder is almost the same as that of Bipolar I. <i>Quetiapine</i> and <i>Lamotrigine</i> are the most favored agents in clinical practice (PPA, 2017)					
	Medication	Cautions	Side Effects			
	Quetiapine	Use with caution for elderly patients.  Caution with other QT-prolonging medications like macrolides, fluoroquinolones, ondansetron, and HIV protease inhibitors.  Monitor weight, BP, FBS, and Lipid Profile, if possible.	CNS (dizziness, drowsiness, ataxia, headache), GI (dry mouth, abdominal pain, diarrhea, constipation, weight gain), Endocrine – diabetes mellitus (increased			

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Second Line	Lamotrigine	Caution in patients with renal, hepatic, and cardiac impairment and elderly.	
		Life-threatening rashes have developed in association with lamotrigine use; it should generally be discontinued at the first signs of serious rash	, , , , , , , , , , , , , , , , , , , ,
		Drug-Lab interaction: Lamotrigine can cause a false positive urine drug screen for phencyclidine and synthetic cannabinoids	associated with Stevens-Johnson syndrome,

# II. TREATMENT DOSE

Disorder		Dosages		
		2 <sup>nd</sup> Line		
	Quetiapine	Lamotrigine	Lamotrigine	
Acute	Initial dose:	Not recommended as first line	As monotherapy (without phenytoin,	
Depression	50mg/day at	medication	phenobarbital, carbamazepine,	
	bedtime.		rifampicin, or lopinavir/ritonavir):	
			- For the first 2 weeks: 25mg/day	
	Increase to 100		- At week 3: increase to 50mg/day	
	mg/day on day 2		- At week 5: increase to 100mg/day	
	and increase dose		- At week 6: increase to 200mg/day,	
	by 50-100mg/day		maximum dose generally	
	until desired		200mg/day	
	response,			
	maximum dose		As monotherapy (with phenytoin,	
	generally 300 mg		phenobarbital, carbamazepine,	
	once daily		rifampicin, lopinavir/ritonavir)	
Maintenance	300 mg once	As monotherapy (without	- For the first 2 weeks: 50mg/day	
	daily	phenytoin, phenobarbital,	- At week 3: increase to 100mg/day	
		carbamazepine, rifampicin, or	- At week 5: increase to 200mg/day in	
		lopinavir/ritonavir):	divided doses	
		- For the first 2 weeks: 25mg/day	- At week 6: increase to 300mg/day in	
		- At week 3: increase to 50mg/day	divided doses, maximum dose	
		- At week 5: increase to 100mg/day	generally 400mg/day	
			Adjunct to Valproate:	

- At week 6: increase to 200mg/day, maximum dose	- For the first 2 weeks: 25mg/day every other day
generally 200mg/day	- At week 3: increase to 25mg/day
	- At week 5: increase to 50mg/day
As monotherapy (with phenytoin,	- At week 6: increase to 100mg/day;
phenobarbital, carbamazepine,	maximum dose generally
rifampicin, lopinavir/ritonavir)	100mg/day
- For the first 2 weeks: 50mg/day	
- At week 3: increase to 100mg/day	
- At week 5: increase to 200mg/day	
in divided doses  - At week 6: increase to 300mg/day	
in divided doses, maximum dose	
generally 400mg/day	
governey to take any	
Adjunct to Valproate:	
- For the first 2 weeks: 25mg/day	
every other day	
- At week 3: increase to 25mg/day	
- At week 5: increase to 50mg/day	
- At week 6: increase to	
100mg/day; maximum dose	

generally 100mg/day

# CONSENSUS TREATMENT GUIDELINES ON DEMENTIA

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#### **DEMENTIA**<sup>6</sup>

Dementia is a term used to describe a large group of conditions affecting the brain which cause a progressive decline in a person's ability to function. It is NOT considered as a normal part of aging.

#### **CLINICAL CRITERIA:**

Decline or problems with memory (severe forgetfulness) and orientation (awareness of time, place and person), Mood or Behavioral Problems such as apathy (appearing uninterested) or irritability, loss of emotional control (easily upset, irritable or tearful), difficulties in carrying out usual work, domestic or social activities.

EMERGENCY PRESENTATION: Agitated and/or aggressive disorder

#### I. STAGES OF DEMENTIA:

\*These are general descriptions. Behaviors may vary.

#### A. EARLY STAGE

Forgetfulness; word-finding difficulty; lost and confused in familiar places; misplaces objects; losing track of time; difficulty handling finances; possible mood changes such as anxiety or depression.

#### B. MIDDLE STAGE

Very forgetful especially of recent events and names; more confused with time and place; needs help with personal care (ex. toileting, dressing); unable to live alone safely; disturbed sleep; hallucinations; inappropriate behavior.

#### C. LATE STAGE

Unaware of time, place and events; unable to recognize family and friends; unable to eat without assistance; may be confined to wheelchair or bed and unable to walk; may have aggressive behavior.

### II. PRINCIPLE GOALS FOR DEMENTIA CARE:

- Early diagnosis in order to promote early and optimal management
- Optimizing physical health, cognition, activity, and well-being
- Identifying and treating accompanying physical illness

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<sup>&</sup>lt;sup>6</sup> WHO (2016): MhGAP Intervention Guide for Mental, Neurological and Substance Abuse Disorders in non-specialized Health Settings, Version 2.0.

- Detecting and treating challenging behavioral and psychological symptoms
- Providing information and long-term support to carers.
- Psychosocial intervention is always first line. Provide support to carers.

#### III. TYPES OF DEMENTIA

#### A. ALZHEIMER'S DISEASE

It is the most common cause of dementia. It is a progressive disease marked by neuropathologic changes which include neuritic plaques and neurofibrillary tangles.

#### **B. VASCULAR DEMENTIA**

Vascular dementia is the second most common cause of dementia after Alzheimer's disease. It can occur when blood flow to the brain becomes reduced. Vascular dementia can also be called vascular cognitive impairment and is sometimes split into more specific types. (Alzheimer's Research UK, 2018). Examples include:

- Stroke-related dementia. This includes multi-infarct dementia (MID), which happens after a series of small strokes. It also includes dementia which happens after a stroke (called post-stroke dementia).
- Subcortical vascular dementia (also called Binswanger's disease, small vessel disease-related dementia or lacunar state). This is caused by changes to very small blood vessels in the brain.

#### IV. PREFERRED REGIMEN

		Treatment Choice							
Alzheimer's	Acetylcholine	Acetylcholinesterase (ACE) Inhibitors							
Disease		e recommended first-line therapy, although co							
and	have diff	Inhibitors and Memantine is likewise recommended since the two classes of drugs have different mechanisms of action. Monotherapy may also be a treatment option. (ADAP, 2014)							
Vascular	Medication	Medication Cautions Side Effects							
Dementia									
- Early Stage	Donepezil	Monitor for cardiac dysrhythmias, especially in those with preexisting cardiovascular disease.	GI upset, headache, fatigue, pain, common cold, anorexia, psychiatric disturbances, syncope, dizziness, insomnia,						
	·	Watch out for respiratory exacerbation in those with asthma or COPD. This medication is contraindicated during pregnancy.	rash, pruritus, muscle cramps, urinary incontinence, accident						

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			Reports of seizures,
			bradycardia, raised creatinine
	1		kinase, EPS, NMS. <sup>7</sup>
	Donepezil	See above	See above
- Middle	Memantine	Use with caution in those with impaired	Headache, dizziness,
Stage		kidney function.	constipation, HTN,
			somnolence, anxiety,
		Renal Adjustment for Memantine:	confusion, hallucinations,
		CrCl 5-29 mL/min: max 10 mg per day	fatigue, abnormal gait,
		(CrCl <5 mL/min not defined)	hypertonia, vomiting, fungal
			infections, cystitis,
			thromboembolism, increased
			libido, psychotic reactions,
			pancreatitis; agranulocytosis,
			leucopenia (including
			néutropenia),
			thrombocytopenia,
			pancytopenia, thrombotic
		/	thrombocytopenic purpura,
			CHF, hepatitis, suicidal
			ideation, acute renal failure
		,	(including increased creatinine
	1		and renal impairment),
			Stevens-Johnson syndrome.8
- Late	}	discontinuation if patient is no longer comm	unicative and completely
Stage	dependent.		

### V. TREATMENT DOSE

	Dosages							
Stage of Dementia	Alzheimér's Disease	Vascular Dementia	Notes					
Early Stage	<b>Donepezil</b> 5-10 mg/day (Oral)	Memantine 10 mg/day (gradually increase to 20 mg / day)	Memantine titration - (10mg/tablet) titrate accordingly:					
		OR	Week 1: 5 mg (1/2 tablet) once daily before bedtime					
		Donepezil 5-10 mg/day						

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 $<sup>^7</sup>$  MIMS UK (n.d.): Donepezil, retrieved from: https://www.mims.co.uk/drugs/central-nervous-system/alzheimer-s-dementia/donepezil

<sup>&</sup>lt;sup>8</sup> MIMS Philippines (2018): Memantine, retrieved from:

https://www.mims.com/philippines/drug/info/memantine/?type=brief&mtype=generic#AdverseReactions

Middle	Donepezil 5 mg/day	Continue Memantine. May add	Week 2: 5 mg (1/2 tablet)
Stage	(Oral)	Donepezil 5-10 mg per tab	twice daily
			Week 3: 5 mg (1/2 tablet)
	OR		twice daily
			Week 4 onwards: 10 mg (1
	(Donepezil 5-10 mg/day)		tablet) twice daily
	+ (Memantine* 10		
	mg/day) (Oral)		
	Gradually increase to 20		
	mg/day		
Late Stage	May consider discontinuat	ion if patient is no longer commun	icative and completely
	dependent.		

#### **Special Considerations:**

- Antipsychotic should only be considered IF
  - a. Psychotic symptoms persist (i.e. violent behavior, hallucinations, persecutory delusions)
  - b. You assess that there is imminent risk for the person and/or carer
  - c. "Start slow, go slow" titrate and review the need regularly
  - d. Use the lowest effective dose
  - e. Monitor the person for side effects such as extrapyramidal symptoms and oversedation
- Avoid anticholinergics/antihistamines for sedations (hydroxyzine, diphenhydramine)
- Avoid haloperidol (I.V.)
- Avoid diazepam
- Medications should not be routinely considered in all cases.
- Adequate supervision and monitoring of side effects are important.
- Follow up should be every 3 months at a minimum. Discontinue drugs if confusion and behavioral changes worsen.

# CONSENSUS TREATMENT GUIDELINES ON EPILEPSY

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#### I. EPILEPSY

Epilepsy is a disease of the brain defined by any of the following conditions:<sup>9</sup>

- 1. At least two unprovoked (or reflex) seizures occurring >24 hours apart.
- 2. One unprovoked (or reflex) seizure and a probability of further seizures similar to the general recurrence risk (at least 60%) after two unprovoked seizures, occurring over the next 10 years.
- 3. Diagnosis of an epilepsy syndrome

Epilepsy is considered to be resolved for individuals who had an age-dependent epilepsy syndrome but are now past the applicable age or those who have remained seizure-free for the last 10 years, with no seizure medicines for the last 5 years.

#### A. EPILEPTIC SEIZURE<sup>10</sup>

Manifestation (s) of epileptic (excessive and/or hypersynchronous) usually self-limited activity of neurons in the brain.

#### **CLINICAL CRITERIA:**

Sudden and transitory abnormal phenomena which may include alterations of consciousness, motor, sensory, automatic, or psychic events perceived by the patient or an observer.

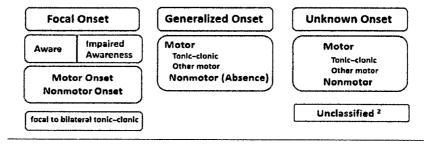
Source: ILAE 1993,1997

#### A. Classification of Seizure Types:

International League Against Epilepsy (ILAE) Classification of Seizure Types is based on 3 key features: where seizures begin in the brain, the level of awareness during a seizure and other features of seizures. The seizure types are based on the seizure onset and how the seizures spread.<sup>11</sup>

The type of seizure onset (seizure type) is important because it affects choice of antiepileptic drugs, may relate to possible etiologies and prognosis in different age groups and other treatment possibilities.

#### ILAE 2017 CLASSIFICATION OF SEIZURE TYPES BASIC VERSION<sup>12</sup>



Philippine League Against Epilepsy (PLAE) References:

<sup>9</sup> Fisher et al. (n.d.): ILAE Official Report: A Practical Clinical Definition of Epilepsy. Epilepsia 55 (4), 475-482.

<sup>&</sup>lt;sup>10</sup> ILAE Classification on Task Force and Terminology, 2001

<sup>&</sup>lt;sup>11</sup> Fisher et al. (2017): operational Classification of Seizure Types by International League Against Epilepsy – Position Paper of the ILAE Commission for Classification and Terminology. Epilepsia, 58 (4): 522-530.

- 1. Focal Seizures start in an area or network of cells on one side of the brain. Focal to Bilateral Seizure - a seizure that starts in one side or part of the brain and spreads to both sides has been called a secondary generalized seizure.
- 2. Generalized Seizures engage or involve networks on both sides of the brain at the onset.
- 3. Unknown Onset if the onset of seizure is unknown.

#### II. ANTIEPILEPTIC DRUG (AED) SELECTION:

- 1. 1ST Line Antiepileptic Drug AED of first choice for specific seizure types and group selected epilepsy syndromes.
- 2. 2<sup>ND</sup> Line Antiepileptic Drug AED of next choice when seizures are not controlled with an appropriately chosen and adequately dosed 1st line AED or development of adverse drug reactions with the use of the 1st/line AED. May also be used as an alternative AED when there is a contraindication to the use of the 1st line AED or the inability to screen for the HLAB1502 allele in cases where carbamazepine is indicated for first time users.

Patients currently on phenobarbital with good seizure control should continue to be maintained on phenobarbital, since switching to another antiepileptic drug will not ensure seizure control. Switching from one antiepileptic drug to another should be done gradually, with transient overlap of the two drugs based on their respective half- lives to prevent withdrawal seizures.

#### PEFERRED REGIMEN<sup>12</sup>:

Antiepileptic Drugs for Specific Seizure Types and Age Group								
Seizure Type		Children (<18 years old)	Girls and Women of Child Bearing Age	Adults	Elderly (>60 years old)			
Focal Onset Seizure with or	1st Line	Oxcarbazepine	Lamotrigine 17*** Levetiracetam	Carbamazepine <sup>16</sup>	Lamotrigine*** Levetiracetam <sup>14</sup>			
without Evolution to Bilateral Tonic, Clonic, Tonic-	2 <sup>nd</sup> Line	Levetiracetam Valproic Acid	Oxcarbazepine <sup>17</sup> *	Levetiracetam Valproic Acid*	Carbamazepine* Valproic Acid **			

<sup>&</sup>lt;sup>12</sup> Philippine League Against Epilepsy (PLAE) mhGAP Task Force on Epilepsy

https://www.mims.com/philippines/drug/info/tegretol/special-precautions?selectedTab=precautions

https://www.pharma.us.novartis.com/sites/www.pharma.us.novartis.com/files/tegretol.pdf

<sup>&</sup>lt;sup>16</sup> MIMS (n.d.): Tegetrol Drug Information, retrieved from:

<sup>&</sup>lt;sup>17</sup> Pharma US (n.d.): Tegetrol, retrieved from:

<sup>&</sup>lt;sup>18</sup> Ferrell and Mcleod (2018): Carbamazepine, HLA-B\*1502 and Risk of Stevens – Johnson Syndrome and Toxic Epidermal Necrolvsis – USFDA Recommendations, Pharmagerian Control of the Contr Epidermal Necrolysis - USFDA Recommendations, Pharmacogenomics 9 (10): 1543-1546

Clonic Seizures <sup>13</sup>					
Generalized	1st Line	Valproic Acid **	Levetiracetam <sup>17</sup>	Valproic Acid**	Lamotrigine
Onset Seizures <sup>13</sup>		_			Levetiracetam
15	2 <sup>nd</sup> Line	Levetiracetam	Lamotrigine <sup>17</sup>	Levetiracetam	Valproic Acid**
Unknown Onset	1st Line	Valproic Acid <sup>19 20</sup>	Levetiracetam	Valproic Acid**	Lamotrigine***
Seizure		21**			Levetiracetam
	2 <sup>nd</sup> Line	Levetiracetam	Lamotrigine***	Levetiracetam	Valproic Acid**

<sup>\*</sup>Screening for HLAB1502 required for first time users

\*\* Avoid in girls and women of childbearing potential, unless there is no other treatment option. When used in girls and women of child bearing potential, should be prescribed at lowest effective dose (DO NOT exceed 500 – 600 mg/day). There may be patients who will require higher doses to attain seizure control.

\*\*\*Lamotrigine may worsen myoclonic seizures

Antiepileptic Drugs for Selecte	d Epilepsy Syndrom	es in Primary Care
Seizure Type	,	
Child Absence Epilepsy (CAE) <sup>13</sup> 15	1st Line	Valproic Acid <sup>19 20 21</sup> **
	2 <sup>nd</sup> Line	Lamotrigine***
Rolandic Epilepsy <sup>13 15</sup>	1st Line	Oxcarbazepine
	2 <sup>nd</sup> Line	Levetiracetam
Juvenile Myoclonic Epilepsy (JME) <sup>13 15</sup>	1st Line	Valproic Acid**
	2nd Line	Levetiracetam

\*\*All girls and women with epilepsy on antiepileptic drugs of child bearing potential should receive folic acid supplementation at 400 micrograms to 5 milligrams per day.

PLAE References:

<sup>&</sup>lt;sup>13</sup> Glauser et al. (2013): Updated ILAE Evidence Review of Antiepileptic Drug Efficacy and Effectiveness as Initial Monotherapy for Epileptic Seizures and Syndromes. Epilepsia, 54 (3): 551-563.

<sup>&</sup>lt;sup>14</sup> Werhahn et al (2015): A Randomized, Double Bind Comparison of Antiepileptic Drug Treatment in the Elderly with New-onset Focal Epilepsy, Epilepsia, 56 (3): 450-459.

<sup>&</sup>lt;sup>15</sup> National Institute of Clinical Experience (NICE) Guidelines 2012, retrieved from:

http://www.nice.org.uk/guidance/cg137/chaoter/1-Guidance#pharmacological-treatment

<sup>&</sup>lt;sup>19</sup> Tomson et al (2015): Valproate in the Treatment of Epilepsy in Girls and Women of Childbearing Potential, Epilepsia, 56 (7): 1006-1019

<sup>&</sup>lt;sup>20</sup> Tomson et al (2018): Comparative Risk of Major Congenital Malformations with Eight Different Anti-Epilepic

Drugs: A Prospective Cohort Study of the EURAP Registry, Lancet Neurology, 17 (6): 530-538

<sup>&</sup>lt;sup>21</sup> European Medicines Agency (EMA), 2018

#### IV. TREATMENT DOSE

Medication	Patient	Dosage and	Mainten-	Maximum	Cautions	Side Effects
	Group	Frequency	ance	Dose .		
Carbamazepine	Adult	Initial:	10 – 20	1400	Caution in patients	Common:
		100-200 mg/day	mg/kg/day	mg/day <sup>22</sup>	with history of blood	Sedation,
		in 2-3 divided			disorders, kidney,	confusion,
		doses. <sup>22</sup>			liver or cardiac	dizziness, ataxia,
					disease <sup>22</sup>	double vision,
		Increase by 100				nausea, diarrhea,
		- 200 mg each			Dose may need to be	benign
		week, until a			adjusted after 2	leukopenia <sup>22</sup>
		dose of 400			weeks due to	
		mg/day is			induction of its own	Serious:
		reached			metabolism <sup>22</sup>	Hepatotoxicity,
						cardiac conduction
	Child	Initial:	5 - 30	< 6 y/o: 35	Screening for	delay, low sodium
		5 mg/kg/day	mg/kg/day	mg/kg/dáy	HLAB1502 required	levels <sup>22</sup>
		divided 2-4		,	for first time users to	
Į.		times per day.	*Infants:	6-15 y/o:	prevent serious	
			10-40	1000	allergic reaction	
		Titration:	mg/kg/day	mg/day	(Stevens-Johnson	
		10-20			syndrome/Toxic	
		mg/kg/day over		>15 y/o:	Epidermal	
		2-4 weeks.		1200	Necrolysis) <sup>23 24</sup>	
				mg/day		
Sodium	Adult	Initial:		3000	Use with caution	Common:
Valproate/		500 mg/day in 2		mg/day	with underlying or	Sedation,
Valproic Acid		divided doses.			suspected hepatic	headache, tremor,
					disease <sup>25</sup>	ataxia, nausea,
		Increase by 500				vomiting,
		mg/day each			*If ER tablet is used	diarrhea, weight
		week			DO NOT SPLIT	gain, transient hair
					TABLET	loss
		L			L	

Doi: 10.2217/14622416.9.10.1543

PLAE References:

<sup>&</sup>lt;sup>22</sup> MhGAP Base Course – Version 2.0 (2016)

PLAE References:

<sup>&</sup>lt;sup>23</sup> US FDA (2008): Safety Alerts for Drugs, Biologics, Medical Devices and Dietary Supplements – 2007

<sup>&</sup>lt;sup>24</sup> Ferrel and Mcleod (October 2008): Carbamazepine, HLAB-1502 and risk of Steven-Johnson Syndrome and Toxic Epidermal Necolysis, US FDA Recommendations Pharmacogenomics, 2008 Octoberr: 9(10): 1543-1546.

<sup>&</sup>lt;sup>25</sup> MhGAP Base Course – Version 2.0 (2016)

	Child	Initial:	Children	60	Drug-drug	
	Cimu	10-15			interactions:	Serious:
			<20 kg: 20-40	mg/kg/day	Valproic levels	Impaired hepatic
		mg/kg/day divided twice a	mg/kg/day		decreased by	function,
			nig/kg/day		1	thrombocytopenia,
		day or thrice a	Children		carbamazepine,	
		day	Children		increase by aspirin <sup>12</sup>	leukopenia,
		*Extended	>20 kg:	į		drowsiness,
		Release – once a	20-30		AVOID in girls and	confusion
		day	mg/kg/day		women of child	(valproate-induced
				;	bearing potential,	hyperammonemic
					unless there is no	encephalopathy, a
					other treatment	sign of toxicity),
					option <sup>26 27</sup>	liver failure,
					f. When used in	hemorrhagic
					girls and women	pancreatitis.
					of Childbearing	
					potential, should	
					be prescribed at	
				-	lowest effective	
					dose, not exceed	
				]	500-600 mg/day.	
					There may be	
					patients who will	
					require hgher	
					doses to attain	
					seizure control. <sup>26</sup>	
					Seizure Control.	
Lamotrigine	Adult <sup>28</sup>	a. Monotherapy		Monotherapy	Very slow titration to	Common:
<b>.</b>		:		: 200 mg/day	prevent serious	Dizziness,
		First 2 weeks:			idiosyncratic rash.	diplopia,
		25mg/day		With	Strictly adhere to	insomnia, ataxia <sup>22</sup>
				valproic	very slow titration	mooning attrict
		Week 3:		acid: 100	recommendation.	Serious:
		50mg/day		mg/day	(Very slow titration	Idiosyncratic
				Ing/day	precludes in patients	rash <sup>22</sup>
		Week 5:		With	<u> </u>	14511
		100mg/day				
				enzyme	seizures recurring	
		Week 6:		inducer: 400	several times daily,	
		200mg/day,		mg/day in	weekly, or monthly)	
		maximum dose		divided	<b>D</b>	
		generally		doses	Drug-lab	
		200mg/day		-	interaction:	
				:	Lamotrigine can	
		h TCi41-			cause a false	
		b. If with			positive urine drug	
		valproic			screen for	
		acid:	<u> </u>		3313311101	

<sup>&</sup>lt;sup>26</sup> Tomson, et. Al (2015): Valproate in the treatment of Epilepsy in Girls and Women of Childbearing Potential, Epilepsia, 56(7): 1006-1019, doi: 10.1111/epi.13201

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<sup>&</sup>lt;sup>27</sup> European Medicines Agency (EMA), 2018

<sup>&</sup>lt;sup>28</sup> Continuum, American Academy of Neurology – February 2016

First 2 weeks: 25mg/day every other day  Week 3: 25mg/day  Week 5: 50mg/day  Week 6: 100mg/day; maximum dose generally 100mg/day c. If with enzyme- inducer (as noted in above dosing): First 2 weeks: 50mg/day  Week 3: 100mg/day  Week 5: 200mg/day  Week 6: 300mg/day in divided doses  Week 6: 300mg/day in divided doses, maximum dose generally 400mg/day  Child <sup>28</sup> Initial: Mono-			F:	_		1' 1' 1	
other day  Week 3: 25mg/day  Week 5: 50mg/day  Week 6: 100mg/day; maximum dose generally 100mg/day c. If with enzyme- inducer (as noted in above dosing): First 2 weeks: 50mg/day  Week 3: 100mg/day  Week 3: 100mg/day  Week 6: 300mg/day in divided doses  Week 6: 300mg/day in divided doses, maximum dose generally 400mg/day			First 2 weeks:			phencyclidine and	
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Week 5: 200mg/day in divided doses  Week 6: 300mg/day in divided doses, maximum dose generally 400mg/day							
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200mg/day in divided doses  Week 6: 300mg/day in divided doses, maximum dose generally 400mg/day							
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300mg/day in divided doses, maximum dose generally 400mg/day							
300mg/day in divided doses, maximum dose generally 400mg/day			Week 6				
divided doses, maximum dose generally 400mg/day							
maximum dose generally 400mg/day					•		
generally 400mg/day							
400mg/day							
							Į
Child <sup>28</sup> Initial: Mono-			400mg/day				
Cimu   miriai.   MOHO-	<del> </del>	Ch;1428	Initial:	Mona			
		CIIIIO-,		ſ			
monotherapy – therapy:							
(for weeks 1 and   4.5-7.5							
2) 0.3 mg/kg/day				mg/kg/day			
mg/kg/day			mg/kg/day				
With				With			
With valproate:   valproate:		ı	With valproate:	valproate:		1	1
0.15 mg/kg/day 1-5							
mg/kg/day				1			
If with enzyme-			If with enzyme-				
inducer: With				With			
0.6 mg/kg/day enzyme			о.о шукудау				
inducer:				1			
Titration dose: 5-15			Titration dose:	i			
mg/kg/day				mg/kg/day			

		a. Monotherap				
		y – 1				
		Week 3-4: 0.6				
		mg/kg/day				
		Ing/kg/day				
}		337 1 5	ļ			
		Week 5				
		onwards:				
		increase by 0.6				
		mg/kg/day	ĺ			
		b. With				
		valproate:				
		Weeks 3-4:				
		1				
		0.3 mg/kg/day Week 5				
		1				
		onwards:				
		increase 0.3				
		mg/kg/day			.'	
		every 1-2 weeks				
		c. With		; 		
		enzyme		!		
		inducer:				
		Weeks 3-4: 1.2				
		mg/kg/day				
		Week 5				
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		e dose by not				
		more than 1.2				
		mg/kg/day				
		every 1-2 weeks				
Levetiracetam	Adult <sup>29</sup>	Initial:		3000	AVOID in patients	Common:
Lic voiri acciaili	. 14411	250 mg a day	1	mg/day	with psychosis,	Irritability,
				ing/uay		· -
		and increase by			irritability,	aggressiveness,
		250 mg every 2			aggressive behavior,	behavioral
		weeks to 500			unless there is no	changes,
		mg twice a day			other treatment	hallucinations,
	Child	10-20		60mg/kg/da	option.	dizziness <sup>30</sup>
Ì		mg/kg/day and		y or 3000		
		adjusted,		mg/day		Serious:
		according to				None reported <sup>30</sup>
		response by				2.ono reportou
ľ		increments of		İ		
						:
		10-20				
		mg/kg/day				
		every 2 weeks				

<sup>&</sup>lt;sup>29</sup> Product Insert, Levetiracetam. Version Number NCDS v.07, June 15 016
<sup>30</sup> Perucca and Engel (2017): Treatment of Epilepsy, 3<sup>rd</sup> Edition, Medscape Drug Reference, Blackwell Publishing Ltd.

Oxcarbazepine	Adult	Initial: 300mg	600 –	2400	Cross-reactivity with	CNS: Headache,
•		twice daily <sup>31 32</sup>	1200	mg/day	carbamazepine has	somnolence
	<u> </u>	_	mg/day	_	been reported, with	dizziness
		Titration:			25.5% of patients	GI: nausea
		300mg per week			who had a history of	Skin: rash
		based on			skin rashes on	Endo: reduces
		response 27			carbamazepine also	serum vitamin D
					developing a rash	GU: hyponatremia
					when they were	usually
					converted to	asymptomatic, but
					oxcarbazepine. A	could cause an
					few cases of	increase in
					Stevens-Johnson	seizures and other
					syndrome have been	adverse effects
					reported.	when serum levels fall below 125
					Avoid in patients	mEq/L.
					with history of skin	tarq/r.
					rash with	
					antiepileptic	
					medications.	
	Child	Initial: 8-10	20-40	<60 mg/kg/d		
}		mg/kg/d <sup>27</sup>	mg/kg/d <sup>27</sup>	or		
				2100mg/day		
		Titration:	Dose for	27		
		5-10mg/k/d	child 2-16			
		every 3-7 days	years old			
		based on	and 4-16			
		response <sup>27</sup>	years old,			
			please see			
			PNF			
			chapter on			
			Nervous			
			system			

#### V. SPECIALTY REFERRAL

#### When to refer to a specialist:

- In cases where the physician is in doubt of the diagnosis
- Uncontrolled seizures despite being on adequate doses of appropriately chosen AED: One monotherapy failure in children, Two sequential monotherapy failure in adults
- Children with epilepsy below age 2
- Children with epilepsy with developmental delay or regression in development

<sup>&</sup>lt;sup>31</sup> Abou-Khalil, B. W. (2016). Antiepileptic Drugs. CONTINUUM: Lifelong Learning in Neurology, 22(1, Epilepsy), 132–156. doi:10.1212/con.000000000000289

<sup>&</sup>lt;sup>32</sup> Shorvon, S., Perucca, E., Engel, J. The Treatment of Epilepsy 4th ed. November 2015 p.575

- Patients presenting with recurrent seizures with focal neurologic deficits or suggestion of a progressive neurologic condition
- Patients with psychiatric comorbidity or other neurologic comorbidities.
- Women who are planning pregnancy or pregnant women with epilepsy
- Patients who develop adverse drug reactions
- Patients who are considered for tapering of AEDs.
- When in doubt, for pre-employment clearance

# CONSENSUS TREATMENT GUIDELINES ON SUBSTANCE USE DISORDERS

## SUBSTANCE USE DISORDERS A. ALCOHOL USE AND ALCOHOL USE DISORDERS

Conditions resulting from different patterns of alcohol consumption include acute alcohol intoxication, harmful alcohol use, the alcohol dependence syndrome, and the alcohol withdrawal state. Acute Intoxication is a transient condition following intake of alcohol resulting in disturbances of consciousness, cognition, perception, affect or behavior. Harmful use of alcohol is a pattern of alcohol consumption that is causing damage to health. The damage may be physical (e.g. liver disease) or mental (e.g. episodes of depressive disorder). It is often associated with social consequences (e.g. family problems, or problems at work).

Alcohol dependence is a cluster of physiological, behavioral and cognitive phenomena in which the use of alcohol takes on a much higher priority for a given individual than other behaviors that once had greater value. The alcohol withdrawal state refers to a group of symptoms that may occur upon cessation of alcohol after its prolonged daily use.<sup>33</sup>

The course of treatment and management of a person with Alcohol Use Disorders undergoes 3 phases: (1) Detoxification and Withdrawal Management, (2) Rehabilitation Program, (3) After-Care and Follow-Up.

#### A.1 ACUTE ALCOHOL INTOXICATION

Acute Alcohol Intoxication is a transient condition following the administration of alcohol or resulting in disturbances in level of consciousness, cognition, perception, affect or behavior, or other psychophysiological functions and responses. Alcohol intoxication is usually closely related to dose levels. Alcohol intoxication is a transient phenomenon. Intensity of intoxication lessens with time, and effects eventually disappear in the absence of further use of the substance.<sup>34</sup>

Look for:	Assess:
Smell of alcohol on the breath	Level of Consciousness
> Slurred speech	Cognition and Perception
Uninhibited behavior	

#### If Intoxication is likely:

1.

- > Assess airway and breathing
- > Put person on the side to prevent aspiration in case they vomit

Refer to hospital if necessary or observe until effects of alcohol have worn off If methanol poisoning is suspected, refer to hospital for emergency management

<sup>33</sup> mhGAP Intervention Guide Ver. 2.0, World Health Organization, 2016

<sup>&</sup>lt;sup>34</sup> International Classification of Diseases 10, World Health Organization, 1996

#### A.2 ALCOHOL WITHDRAWAL

Alcohol withdrawal state is a group of symptoms of variable clustering and severity occurring on absolute or relative withdrawal of a substance after repeated, and usually prolonged and/or high dose, use of that substance. Onset and course of withdrawal state are time-limited and are related to the type of substance and the dose being used immediately before abstinence. The withdrawal state may be complicated by convulsions.<sup>35</sup>

#### Alcohol Withdrawal is best treated under a medical setting.

Alcohol withdrawal occurs following cessation of heavy alcohol consumption, typically between 6 hours and 6 days after the last drink

Look for:

Tremor in the hands

Sweating

Nousea

Nousea

Ask about:

Headache

Nausea

Anxiety

Increased pulse and blood pressure

Note: Seizure and confusion may

#### Refer to a Specialist/Hospital:

> Agitation

- 1. If withdrawal likely to be SEVERE:
  - a. Look for:
    - i. Past episodes of severe alcohol withdrawal including delirium and seizures

occur in severe cases

- ii. Other medical or psychiatric problems or Benzodiazepine dependence
- iii. Severe withdrawal symptoms already present only a few hours after stopping drinking
- 2. If withdrawal is complicated by Delirium
- 3. If withdrawal is complicated by a Seizure

#### B. DRUG USE AND DRUG USE DISORDERS

Conditions resulting from different patterns of drug use include acute sedative overdose, acute stimulant intoxication or overdose, harmful or hazardous use, cannabis dependence, opioid dependence, stimulant dependence, benzodiazepine dependence, and their corresponding withdrawal states. Harmful use of drugs is a pattern of drug consumption that is causing damage to health. The damage may be physical (as in cases of infection related to drug use) or mental (e.g. episodes of depressive disorders) and is often associated with damage to social functioning (e.g. family problems, legal problems, or work-related problems)Drug Dependence is a cluster of psychological, behavioral and cognitive phenomena in which drug use takes higher priority for a given individual than

<sup>35</sup> mhGAP Intervention Guide Ver. 2.0, World Health Organization, 2016

other behaviors that once had greater value. The drug withdrawal state refers to a group of symptoms occurring upon the cessation of a drug after its prolonged daily use.<sup>36</sup>

#### Opioid Overdose or other sedative overdose or mixed drug with or without alcohol overdose

#### Look for:

- > Unresponsiveness or minimally responsive
- > Slow respiratory rate
- > Pinpoint pupils (Opioid Overdose)

#### **Acute Opioid Withdrawal**

#### Look for:

- ➤ History of Opioid Dependence, recent heavy use ceasing in the last days
- Muscle aches and pains, abdominal cramps, headaches
- > Nausea, vomiting, diarrhea
- > Dilated pupils
- > Raised pulse and blood pressure
- > Yawning, runny eyes and nose, pilo-erection ("gooseflesh")
- > Anxiety, restlessness

#### Acute stimulant intoxication or overdose

#### Look for:

- > Dilated pupils
- > Excited, racing thoughts, disordered thinking, paranoia
- Recent use of stimulants (Methamphetamine, Cocaine, Ecstacy, etc.)
- > Raised blood pressure
- > Aggressive, erratic or violent behavior

#### Refer to a Specialist/Hospital

#### C. MENTALLY ILL CHEMICAL ABUSE (MICA)

Adults with severe mental illness have high rates of co-occurring substance use disorders which adversely affect their current adjustment, course, and outcome. Separate and parallel mental health and substance abuse treatment systems do not offer interventions that are accessible, integrated, and tailored for the presence of co-occurrence. Recent integrated interventions for this population have the specific goal of ameliorating substance use disorder and the general goal of improving adjustment and quality of life.<sup>37</sup>

Pharmacological management of both psychiatric and the substance use disorder is an important foundation of the treatment of clients with co-occurring severe mental illness and substance use disorders. <sup>28</sup>

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<sup>&</sup>lt;sup>36</sup> mhGAP Intervention Guide Ver. 2.0, World Health Organization, 2016

<sup>&</sup>lt;sup>37</sup> International Classification of Diseases 10, World Health Organization, 1996

#### Mentally Ill Chemical Abuser (MICA)

#### **Pharmacological Interventions**

- Medications shown to be effective for the treatment of substance use disorders are probably effective also in patients with serious mental illness
- Antidepressants reduce not only symptoms of depression but also substance abuse in clients with both mental disorder and substance use disorder
- Mood stabilizers are active not only in common mania but also on substance use in clients with bipolar disorders and co-morbid substance dependence
- > Typical Antipsychotics improve the symptoms of schizophrenia but have little effect on co-morbid substance dependence
- Atypical Antipsychotics are equally effective as the typical antipsychotics in improving schizophrenia symptoms and may offer some benefit in reducing craving or substance use, but research is preliminary

#### Recommendations:

- > Use protocols for management of Mental Disorders
- > Integrated management/Multidisciplinary approach is recommended

Refer to Specialist/Hospital for initial management and complicated cases

# CONSENSUS TREATMENT GUIDELINES ON EXTRAPYRAMIDAL SYMPTOMS (EPS)

#### EXTRAPYRAMIDAL SYMPTOMS (EPS)

Antipsychotics induced extra pyramidal symptoms include a variety of movement disorders Acute extrapyramidal symptoms are like acute dystonia, akathisia and parkinsonism develop within hours or weeks after initiating or increasing doses of antipsychotics. Tardive dyskinesia and tardive dystonia are delayed onset syndromes and usually develop after a prolonged use of antipsychotics.

- 1. Dystonia are characterized by intermittent or sustained muscle action.
- 2. Akathisia manifests as the feeling of restlessness and irresistible urge to move.
- 3. Drug--induced Parkinsonism is characterized by a triad of bradykinesia, muscle rigidity and tremor
- 4. Tardive dyskinesia is manifested by involuntary choreoathetoid movements of the orofacial region, extremities, trunk and respiratory muscles.
- 5. Neuroleptic malignant syndrome (NMS) is marked by rigidity, fever, changes in the mental status and the autonomic dysfunction. (Kirgaval et al., 2017)

#### II. PREFERRED REGIMEN

Treatment	Preferred Regimen								
Choice									
First Line	Anticholinergic								
	- Short-term use of anticholinergics may be considered only in individuals with significant								
	extrapyramidal side-effects when dose reduction and switching strategies have proven								
			le-effects are acute or se	<del></del>					
	Medication	Dosage	Cautions	Side Effects					
1		and	•						
		Frequency							
	Biperiden	1 mg twice	Caution in patients	Common:					
		daily	with cardiac, liver,	Sedation, Confusion and memory					
		(Increase to		imbalance (especially in older					
		3-12 mg		adults), tachycardia, dry mouth,					
		daily)	Drug-Drug	urinary retention and constipation					
			Interactions:						
			Caution when						
			other anticholinergic						
	medications								
	Diphenhydramine <sup>41</sup>	0.5 to 1 ml	Caution in: Patient	Chest tightness, extrasystoles,					
		(IM/IV)	w/ glaucoma,	hypotension, palpitations,					

<sup>&</sup>lt;sup>40</sup> World Health Organization (n.d.): Mental Health - Role of anticholinergic medications in patients requiring long-term antipsychotic treatment for psychotic disorders, retrieved from:

https://www.mims.com/philippines/drug/info/diphenhydramine?mtype=generic

https://www.who.int/mental\_health/mhgap/evidence/psychosis/q6/en/ MIMS Philippines (2018): Diphenhydramine, retrieved from:

50 mg/ml, 1 ml amp	urinary retention,	tachycardia; ataxia, chills,
	myasthenia gravis,	confusion, dizziness, drowsiness,
	epilepsy or seizure	euphoria, excitement, fatigue,
	disorders, bronchitis	headache, insomnia, irritability,
	or COPD, CV	nervousness, neuritis,
	disease, thyroid	paraesthesia, paradoxical
ļ	dysfunction. Hepatic	excitation, restlessness, sedation,
	and moderate to	seizure, vertigo; diaphoresis;
	severe renal	menstrual disease; GI
	impairment.	disturbances (e.g. anorexia,
	Pregnancy.	constipation, diarrhoea); difficulty
		in micturition, urinary frequency,
		urinary retention; agranulocytosis,
		haemolytic anaemia,
		thrombocytopenia; anaphylactic
		shock; tremor; blurred vision,
		diplopia; acute labyrinthitis,
		tinnitus; constriction of pharynx,
		nasal congestion, thickening of
		bronchial secretions, wheezing;
		photosensitivity, rash, urticaria
		(topical).

### APPENDICES

#### APPENDIX A

#### GLOSSARY (mhGAP Intervention Guide Version 2.0)

**DEFINITION TERMS** 

Agitation Marked restlessness and excessive motor activity, accompanied by

anxiety

A blood disorder in which there is an absence of granulocytes (a type of Agranulocytosis

> white blood cell). It is an acute condition involving a severe and dangerous leukopenia, also known as drug-induced secondary

agranulocytosis.

Akathisia A subjective sense of restlessness, often accompanied by observed

excessive movements (e.g. fidgety movements of the legs, rocking from

foot to foot, pacing, inability to sit or stand still).

The absence or lack of voluntary movement. A state of difficulty in Akinesia

initiating movements or changing from one motor pattern to another that

is associated with Parkinson's disease.

A changed level of awareness or mental state that falls short of Altered Mental Status

> unconsciousness which is often induced by substance intake or other mental or neurological conditions. Examples include confusion and

disorientation. See delirium and confusional state.

Alzheimer's Disease A primary degenerative cerebral disease of unknown etiology in the

> majority of cases with characteristic neuropathological neurochemical features. The disorder is usually insidious in onset and

develops slowly but steadily over a period of several years.

Anticholinergic medicines block the effects of acetylcholine at Anticholinergic Side-effects

> muscarinic receptors. Anticholinergic effects include dryness of the mouth, urinary frequency or retention, palpitations and sinus tachycardia. Failure of muscular coordination. People with ataxia have problems with

> coordination because parts of the nervous system that control movement and balance are affected. Ataxia may affect the fingers, hands, arms, legs,

body, speech, and eye movements.

Cerebrovascular Accident A sudden disturbance of cerebral function attributable to vascular

disease, principally thrombosis, hemorrhage, or embolism. See stroke

Mental processes associated with thinking. These include reasoning, Cognitive

remembering, judgement, problem-solving and planning.

Comorbid, Comorbidity Describing diseases or disorders that exist simultaneously

Confusion, Confusional A state of impaired consciousness associated with acute or chronic

> cerebral organic disease. Clinically it is characterized by disorientation, slowness of mental processes with scanty association of ideas, apathy, lack of initiative, fatigue, and poor attention. In mild confusional states,

rational responses and behavior may be provoked by examination but more severe degrees of the disorder render the individual unable to retain

contact with the environment.

Convulsion, Convulsive

Movement

Ataxia

State

Clinical or subclinical disturbances of cortical function due to a sudden. abnormal, excessive, and disorganized discharge of brain cells (see seizure). Clinical manifestations include abnormal motor, sensory and

psychic phenomena.

Delirium

Transient fluctuating mental state characterized by disturbed attention (i.e., reduced ability to direct, focus, sustain, and shift attention) and awareness (i.e., reduced orientation to the environment) that develops over a short period of time and tends to fluctuate during the course of a day. It is accompanied by (other) disturbances of perception, memory, thinking, emotions or psychomotor functions. It may result from acute organic causes such as infections, medication, metabolic abnormalities, substance intoxication or substance withdrawal.

Delusion

Fixed belief that is contrary to available evidence. It cannot be changed by rational argument and is not accepted by other members of the person's culture or subculture (i.e., it is not an aspect of religious faith). The process by which an individual is withdrawn from the effects of a

Detoxification

The process by which an individual is withdrawn from the effects of a psychoactive substance. Also referring to a clinical procedure, the withdrawal process is carried out in a safe and effective manner, such that withdrawal symptoms are minimized.

Disability

Any restriction or lack (resulting from an impairment) of ability to perform an activity in the manner, or within the range, considered to be normal for a human being. The term disability reflects the consequences of impairment in terms of functional performance and activity by the individual.

Disorganized / Disordered Thinking

A disturbance in the associative thought process typically manifested in speech in which the person shifts suddenly from one topic to another that is unrelated or minimally related to the first. The individual gives no indication of being aware of the disconnectedness or illogicality of his or her thinking.

Disorganized Behavior

Behavior including posture, gait, and other activity that is unpredictable or not goal-directed (e.g., shouting at strangers on the street).

Dystonia

Sustained muscle contraction or involuntary movements that can lead to fixed abnormal postures. See tardive dyskinesia.

Extrapyramidal Sideeffects/Symptoms (EPS) Abnormalities in muscle movement, mostly caused by antipsychotic medication. These include muscle tremors, stiffness, spasms and/or akathisia

Hallucination

False perception of reality: seeing, hearing, feeling, smelling or tasting things that are not real.

Hypersensitivity Reaction

Hypersensitivity reactions are the adverse effects of pharmaceutical formulations (including active drugs and excipients) that clinically resemble allergy. It belongs to type B adverse drug reactions, which are defined by the WHO as the dose-independent, unpredictable, noxious, and unintended response to a medicine taken at a dose normally used in humans. It covers many different clinical phenotypes with variable onset and severity.

Irritability, Irritable Mood

A mood state characterized by being easily annoyed and provoked to anger, out of proportion to the circumstances

Motor Twitching Neuroleptic Malignant Syndrome See convulsion

A rare but life-threatening condition caused by antipsychotic medications, which is characterized by fever, delirium, muscular rigidity and high blood pressure

Orthostatic Hypotension Sudden drop of blood pressure that can occur when one changes position

from lying to sitting or standing up, usually leading to feelings of light-

headedness or dizziness. It is not life-threatening

Pruritus Itching; an intense sensation that produces the urge to rub or scratch the

skin to obtain relief.

QT Prolongation A potential medication induced side-effect of ventricular myocardial

repolarization characterized by a prolonged QT interval on the electrocardiogram (ECG) that can lead to symptomatic ventricular

arrhythmias and an increased risk of sudden cardiac death.

Relapse A return to drinking or other drug use after a period, of abstinence, often

accompanied by reinstatement of dependence symptoms. The term is also used to indicate return of symptoms of MNS disorder after a period of

recovery.

Rigidity Resistance to the passive movement of a limb that persists throughout its

range. It is a symptom of Parkinsonism.

Seizure Episode of brain malfunction due to disturbances of cortical function

resulting in sudden, abnormal, excessive, and disorganized discharge of brain cells. Clinical manifestations include abnormal motor, sensory and

psychic phenomena.

Self-Harm Intentional self-inflicted poisoning or injury to oneself, which may or

may not have a fatal intent or outcome.

Slurred Speech Speech with indistinctive pronunciation

Status Epilepticus Defined as 5 min or more of continuous clinical and/or electrographic

seizure activity or recurrent seizure activity without recovery (returning to baseline) between seizures; it can be convulsive or non-convulsive.

Stevens-Johnson Syndrome Life-threatening skin condition characterized by painful skin peeling,

ulcers, blisters and crusting of mucocutaneous tissues such as mouth, lips, throat, tongue, eyes and genitals, sometimes associated with fever. It is most often caused by severe reaction to medications, especially

antiepileptic medicines.

Stigma A distinguishing mark establishing a demarcation between the

stigmatized person and others attributing negative characteristics to this person. The stigma attached to mental illness often leads to social exclusion and discrimination and creates an additional burden for the

affected individual.

Stroke See cerebrovascular accident (CVA).

Suicidal Thoughts/Ideation Thoughts, ideas, or ruminations about the possibility of ending one's life,

ranging from thinking that one would be better off dead to formulation of

elaborate plans.

Tardive Dyskinesia This is dystonia characterized by twisting and sustained muscle spasms

that affect regions of the head, neck, and occasionally, the back. It may

not improve after stopping the antipsychotic medicine.

Thrombocytopenia Abnormally low number of platelets in the blood. This disease may

present with increased bruising or hemorrhaging. Confirmation is by

identification of decreased platelet count in a blood sample.

Tremor Trembling or shaking movements, usually of the fingers, that is an

involuntary oscillation of a body part

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#### Republic of the Philippines Department of Health

#### FORM 1

#### **REQUISITION AND ISSUANCE SLIP (RIS)**

### MENTAL HEALTH MEDICINES ACCESS PROGRAM Name of Health Facility: \_\_\_\_\_ Contact No: \_\_\_\_\_ Address: \_\_\_\_\_

	Psychotropic Medicines	Quantity for 1 year consumption + 3-month buffer stock	Total *Number of Patients using the medicine	Target Number of Patients for the year
1.	Carbamazepine 200 mg Tablet			
2.	Lithium Carbonate 450 mg MR Tablet			
3.	Valproate Disodium + Valproic Acid 250 mg tablet			
4.	Sodium Valproate 250 mg/5 ml Syrup			
5.	Valproic Acid + Soidum Valproate 500 mg (MR)			
6.	Biperiden Hydrochloride 2 mg Tablet		.,	
7.	Chlorpromazine 200 mg Tablet			
8.	Clozapine 100 mg Tablet			
9.	Fluphenazine Decanoate 25 mg/mL, 1 mL Ampoule			
10.	Diphenhydramine 50 mg/mL, 1 mL Ampoule			
11.	Haloperidol 5 mg Tablet			
12.	Haloperidol 5 mg/mL, 1 mL Ampoule			
13.	Olanzapine 10 mg Tablet			
14.	Olanzapine 10 mg Oro-Dispersible Tablet (ODT)			
15.	Quetiapine 200 mg Tablet			
16.	Risperidone 2 mg Tablet			
17.	Risperidone 2 mg Oro-dispersible Tablet (ODT)			
18.	Escitalopram 10 mg Tablet			
19.	Fluoxetine 20 mg Capsule			
20.	Sertraline 50 mg Tablet			
21.	Lamotrigine 100 mg Tablet			
22.	Donepezil 10 mg Tablet			
23.	Fluphentixol 20 mg/ml			
24.	Paliperidone Palmitate 150 mg prefilled syringes			
25.	Paliperidone Palmitate 100 mg prefilled syringes			

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Previously number of patients	served with the following mental hed	alth conditions (2018):	
			Number of
	Mental Health Conditions		patients
Psychosis (Schizophrenia)			
Anxiety Disorders			
Mood Disorders			
Dementia			
Epilepsy			
Substance Abuse Disorders			
Other diagnosis (please speci	fy):		1
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	Prepared by:	Approved by:	
Signature:			
Name:			
Designation:			
Contact No.:			
Email Address:			

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#### ANNEX C REPORTING FORM

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ID No.	DATE OF ASSESMENT	NAME OF PATIENT	ADDRESS	AGE				CONTACT NO.	DEPRESSION
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PSYCHOSES	EPILEPSY/	CHILD AND ADOLESCENT MENTAL AND BEHAVIOURAL DISORDERS	DEMENTIA	DISORDERS DUE TO SUBSTANCE USE	SELF-HARM/ SUICIDE	OTHERS	MEDICATION (DOSAGE & FREQUENCY)	REMARKS	DATE OF NEX FOLLOW-UP
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