NURSE PRACTITIONER, PRIMARY CARE AND THE BIPOLAR PATIENT

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#### NURSE PRACTITIONERS FIT

 Primary Care Nurse Practitioners (NP) identify mental health issues, provide initial medication and specialist referrals.

 NP's also monitor general health, medication levels, adjust and add adjunct therapies as well as treat comorbid conditions such as hypertension and diabetes.

**DEALING WITH MENTAL HEALTH CONCERNS** • Research found approximately 33% of the primary care (PC) population qualify as a mentally disordered. • A PC based study reported a 9.8% prevalence of positive screening results for Bipolar Disorder.

## **BIPOLAR DISORDER**

• Bipolar disorder (BPD) is a genetic, neurochemically complex, recurrent, and potentially progressive neuropsychiatric disorder, effecting 1-3% of US population. BPD involves dysregulation of mood, sleep, cognition, endocrine and motor systems.

## **DEFINING TERMS**

- Bipolar 1 is one or more episodes of mania, typically both depressive and manic episodes alternating,
- Bipolar 2 is alternating major depression and hypomania.

## **MAJOR DEPRESSION**

 DSM-IV defines Major Depression as a clinical syndrome including anhedonia (depressed mood) every day, all day for at least 2 weeks with significant functional impairment.

## ANHEDONIA

• Anhedonia is accompanied by at least four other symptoms, sleep or appetite changes, fatigue, psychomotor retardation or agitation, lack of concentration, difficulty in decision making, helplessness, hopelessness, guilt, or frequent suicidal thoughts.

## DYSTHYMIA

• Dysthymia is a mild, chronic depressive disorder characterized by fatigue, pessimism, low self-esteem, low motivation and decreased capacity for joy, not meeting major depression criteria.

#### MANIA

 Mania is observable heightened mood, faster speech, quicker thought, brisker physical & mental activity levels, greater energy, irritability, perceptual acuity, paranoia, heightened sexuality & impulsivity for at least seven days.

## HYPOMANIA & MIXED STATES

- Hypomania is a moderate reflection of the above changes and may or may not result in serious problems for the individual experiencing them.
- Mixed states episodes when major depression & mania criteria are met simultaneously.

## CYCLOTHYMIA & CYCLING

- Cyclothymia represents dysthymia with a component of hypomania. Cyclothymia as related to bipolar I & II, is less severe and associated with less impairment.
- Cycling the movement over time between episodes (mood swings).
- Cycling's 3 dimensions –frequency, time between episodes, duration

#### **IDENTIFYING BIPOLAR SPECTRUM DISORDER** Look for:

- A family hx Bipolar in first degree relative
- Antidepressant induced mania or hypomania
- Hyperthymic personality
- Recurrent major depressive episode (>3 months)
- Brief major depressive episode (<3 months)

#### BIPOLAR SPECTRUM CONT. • Atypical features (increased sleep or

- appetite)
- Psychotic major depressive episode
- Early age onset depressive episode (before age 25)
- Anti-depressant tolerance
- Lack of response to >3 antidepressant treatment trials

## BIPOLAR SPECTRUM CONT.

- Co-morbid conditions frequently occur such as:
- Anxiety Disorder
- Attention Deficit Hyperactivity Disorder
- Obsessive Compulsive Disorder
- Substance Abuse
- Suicidal Ideations

## **Mnemonic DIGFAST**

- Distractibility
- Insomnia
- Grandiosity

Flight of ideas
Activity
Speech
Thoughtlessness

**NEUROBIOLOGY & NEUROPHYSIOLOGY** • Research suggests Bipolar & spectrum conditions arise from abnormalities in cellular plasticity leading to aberrant processing in synapses & circuits altering affective, cognitive, motoric & neurovegetative functions not simply deficits excesses in individual neurotransmitters.

#### NEUROLOGY & BIOLOGY CONT.

• Multiple systems are implicated in BPD including Serotonergic system, Noradrenergic system, Dopaminergic system, Cholinergic system, Glutamatergic system, GABergic system, Cortisol releasing factor (CRF)& Hypothalmicpituitary (HPA) axis, & Peptids

## MULTIPLE FACTORS (MAY INFLUENCE)

- Genetic Factors
- Chemical Factors
- Electrical Factors
- Emotional Factors
- Physical Factors

- Medication Factors
- Environmental Factors
- Reaction Factors
- Factors that = Balance

- If yGENET fest degree relative (mother, father, or sibling) you are (7) times more like FACTOR op a bipolar disorder. Understanding that this disorder may not manifest the same in your self or relatives.
- Male vs. Female Genetic/Brain Structure/Social Roles will make women more prone.
- Each disorder is unique and will need to be treated the same once the underlying causes are understood.

### ENVIRONMENTAL FACTORS

• What are we exposed to today?

- -Stress and Stress Management
- -Sleep or Lack of Sleep
- -Work
- Managing Family and Self

 Biorhythms – Circadian Rhythms, sun / moon /seasons affect secretion of catecholamines (norepinephrine)

#### NEUROLOGY & BIOLOGY CONT.

- Investigators using high resolution 3-D MRI studies recorded diminished gray matter volumes in the orbital, medial prefrontal cortex, ventral striatum, amygdala & hippocampus.
- Imaging studies and post mortem neuropathology investigations reveal a reduction in glial cell numbers and cell body size.

#### LooBALTANGE "feel good" feeling or response? – FAGTGRONS

- Natural Remedies
- Exercise / Diet
- Group therapy learning new brain responses
- Alleviating the Anxiety
- Life Style
- Health Maintenance

#### • CerMEPIGATIONes:

- **FACTOR** Medications for bipolar / schizophrenia diagnosis consider the effect to the illness itself and the side effects to the person
- Other medications (concomitant) influences with or without treatment
- Self-Medicating

#### NEUROLOGY & BIOLOGY CONT.

• Lithium and the mood stabilizer valproate demonstrate neuroprotective effects and in the case of lithium with chronic use enhances neuroplasticity & increasing brain cellular mass and numbers.

## HERITABILITY

- Evidence for genetic links between family members is documented in family, twin & adoptions studies.
- Family studies of BPD show spectrum mood disorders are found among first degree relatives including MDD, BPD1, BPD2 & schizoaffective spectrum disorders.
- Although linkage studies suggest several regions in the genome, no established universal genetic risk factor or causative gene is identified.

## ASSESSMENT

- Physical Assessment
- Interview Patient symptoms
- History Injuries, illness, genetics
- Social History Work, habits, addictions
- Verifying information
- Previous Medical Records

## ASSESSMENT TOOLS

PHQ9
Adult Wellbeing
Bipolar
Hypomania
Mania

#### ASSOCIATED SYMPTOMS INVENTORIES • GAD-7

- CAGE (Co-morbid Substance Abuse)
- ADHD (Adult ADHD Self-Report Scale, ASRS-v1.1)
- Suicidal Ideation Questions
- Pain Scale

## EVALUATION

- Differential Diagnosis
  ID Co-morbid Conditions
- R/O Organic Disease
- Drug or Alcohol use/abuse
- Develop Plan

## COLLABORATIVE TREATMENT

• Neuro-psychiatric Evaluation Neurology Consultation Psychology Consultation Diagnostic Imaging • Laboratory Testing

#### PHARMACOTHERAPEUTICS

For Bipolar Patients

## ACUTE MANIA OR HYPOMANIA

In a frankly manic or hypomanic state urgent psychiatric hospitalization may be required. Anti-depressants may need to be discontinued until the manic episode is resolved. Intramuscular injection of medication may be required, hence the need for hospitalization. Current first line choices are aripiprazole or olanzapine, haloperidol or lorazepam may be beneficial

## HYPOMANIA OR MILD MANIA

Monotherapy Lithium Carbamazepine Divalproex Risperidone Olanzapine Aripiprazole

# MANIA

Monotherapy may be attempted Combination therapy is most often recommended

Dosage maximization is imperative

If change agents in combined therapy change only one agent at a time, otherwise effectiveness or SE's will be confused

## **ACUTE DEPRESSION**

Available studies support a concern over use of antidepressants exacerbating mania. When used, anti depressants are used in combination with mood stabilizers such as: Olanzapine and fluoxetine **Bupropion and topiramate** Escitalipram and olanzapine or Citalopram and aripiprazole

### **RAPID CYCLING**

4 or more mood cycles in one year May occur in a monthly cycle of switching between depression and mania Choice of treatment difficult with the presence of severe depression prompting increased use of anti depressants that lead to extreme manic responses.

# RAPID CYCLING CONTINUED

You must look for psychostimulants, prescription or recreational drug use, alcohol use, excessive caffeine intake and even over the counter diet pills.

The focus in rapid cycling is to approach mood stabilization as the primary goal.

The role of lithium should not be ignored in this setting as an effective treatment.

## **OLDER PATIENTS**

The prevalence of bipolar disorder is believed to decrease in late life.

The onset of mania in later life, after the age of 60, may be more often associated with medical conditions such as stroke and other brain lesions.

The same medications used for younger patients are effective at lower doses. As always, start low and go slow.

# MAINTENANCE THERAPY

Begin after the first manic episode Discussion of the maintenance plan with the patient needs to occur immediately after mood stabilization

Medications used to successfully stabilize the patient from the initial episode should be continued

### MEDICATIONS

The medications recommended overlap in frequency and are often used as combined therapy. For this reason medications will be covered regarding their dosing and effects under each section. In cases where medications repeat they will be mentioned and not reviewed. Many recommendations concern the use of atypical antipsychotics which have significant severe adverse side effects including; extrapyramidal symptoms, tardive dyskinesia, hyperglycemia and diabetes. Common side effects are; increased appetite, somnolence, fatigue and nausea and vomiting.

Acute mania or hypomania with agitation will require acute admission due to the need for intramuscular medication until stabilization. Standard recommended therapy is:

Aripiprazole: 9.75-15 mg IM q 2 hours, max 30 mg/day Olanzapine: 2.5 to 10 mg IM q 2-4 hours, max 30 mg/day Secondary option Haloperidol: 5 mg IM q 30-60 minutes max 8mg/day Lorazepam: 1-2 mg IM q 30-60 min max 8 mg/day

Acute mania or hypomania without agitation can be treated on an outpatient basis with oral monotherapy using a mood stabilizer or atypical antipsychotic such as; Lithium: 900 mg/day divided into 3 to 4 doses daily. Maximum dose is 1800 mg/day. Therapeutic serum level is 0.6-1.2 mEq/L. >1.5 mEq/L is considered toxic. Divalproex Sodium ER: 25-60 mg/kg/day in a single daily dose. Start at 25 mg/kg and dose rapidly to the lowest effective dose. Therapeutic drug level in mania 50-125 mcg/ml, >175 mcg/ml is considered toxic.

Acute mania or hypomania without agitation cont.

Carbamazepine: 200-800 mg/day. Start at 100-200 mg po twice daily and increase by 200 mg/day every 3-4 days. Maximum dose is 1600 mg/day. Therapeutic drug levels 4-12 mcg/ml, toxic level > 12 mcg/ml.

Risperidone: Start at 2 mg daily and adjust by doses of 1 mg over more than 24 hours. Maximum dose 6 mg/day.

Acute mania or hypomania without agitation cont.

Olanzapine: Start at 5 mg daily and adjust dose by 5 mg daily as needed. Maximum dose is 20 mg daily.

Quetiapine: Start at 50 mg at bedtime. Increase to 100 mg qhs, then 200 mg qhs, then 300 mg qhs. Maximum dose is 600 mg/day, though doses greater than 300 mg/day are rarely more effective. Aripiprazole: Start at 15 mg/day. Maximum daily dose 30 mg/day. Recommended combination therapy with Risperidone, Olanzapine, Quetiapine, and Aripiprazole are lithium and valproate.

Acute mania or hypomania without agitation cont. Adjunct Therapy:

Clonazepam: For patients failing monotherapy a benzodiazepine may be added. Start at 0.25-0.5 mg 2-3 times a day.

- Moderate to severe disease without agitation uses the same medications as the above scenario. The addition is oral typical antipsychotics such as;
- **Primary options**
- Oral combination therapy: Mood stabilizer and atypical antipsychotic
- Lithium: 900 mg/day divided into 3 to 4 doses daily.
- Divalproex Sodium: 20 mg/kg to maximum of 60 mg/kg. Target serum level 80-100 mcg/ml.
- Carbamazepine: 200-800 mg/day. Start at 100-200 mg po twice daily and increase by 200 mg/day every 3-4 days. Maximum dose is 1600 mg/day.

- Moderate to severe disease without agitation cont.
- And

Risperidone: 1 mg po daily, max 6 mg/day or Olanzapine: 5 mg/day max 20 mg/day or Quetiapine: 5 mg/day max 20 mg/day Aripiprazole: 15 mg/day max 30 mg/day Ziprasidone: 40 mg twice daily max 160 mg/day

- Moderate to severe disease without agitation cont.
- 2nd oral typical antipsychotics
- Haloperidol: 2 mg/day orally initially given in 2-3 divided doses, increase according to response, maximum 12 mg/day. D/C if ANC <1000.
- Chlorpromazine: 30 mg/day orally initially given in 1-4 divided doses, increase according to response, maximum 1000 mg/day

- Non-rapid cycling, non-pregnant, acute depression uses a mix of selective serotonin reuptake inhibitors (SSRIs) and atypical antipsychotics primarily such as;
- Quetiapine: 50 mg/day, increase according to response, max 800 mg/day in 2-3 divided doses. 300 mg ER/day max 600 mg ER/day
- Olanzapine/fluoxetine: 6 mg/25 mg orally once daily initially, increase according to response, maximum 12 mg/50 mg once daily

- Non-rapid cycling, non-pregnant, acute depression cont; And
- Lithium: 900 mg/day divided into 3 to 4 doses daily. Maximum dose is 1800 mg/day.
- Bupropion: 150 mg/day max 450 mg/day given BID
- Aripiprazole: 15 mg/day max 30 mg/day
- Divalproex sodium: 20 mg/kg ER max 60 mg/kg serum level target 80-100 mcg/ml
- Quetiapine: 50 mg/day max 800 mg/day bid or tid
- And
- Citalopram: 20 mg/day max 40 mg/day Escitalopram: 10 mg/day max 20 mg/day

Non-rapid cycling, non-pregnant, acute depression cont;

Fluoxetine: 20 mg/day max 80 mg/day Paroxetine: 20 mg/day max 50 mg/day Sertraline: 50 mg/day max 200 mg/day

- Rapid cycling and non-pregnant uses the same medications as the acute mania or hypomania setting noted above.
- Lithium: 900 mg/day divided into 3 to 4 doses daily. Maximum dose is 1800 mg/day.
- Divalproex sodium: 20 mg/kg ER max 60 mg/kg serum level target 80-100 mcg/ml
- Carbamazepine: 200-800 mg/day initially bid
- Risperidone: 1 mg po daily, max 6 mg/day
- Olanzapine: 5 mg/day max 20 mg/day
- Quetiapine: 50 mg/day max 800 mg/day bid or tid
- Aripiprazole: 15 mg/day max 30 mg/day

Rapid cycling and non-pregnant uses the same medications as the acute mania or hypomania setting noted above.

And

Clonazepam: 1 mg/day bid or tid max 6 mg/day or Risperidone: 1 mg po daily, max 6 mg/day or Olanzapine: 5 mg/day max 20 mg/day or Quetiapine: 50 mg/day max 800 mg/day bid or tid or Aripiprazole: 15 mg/day max 30 mg/day

After stabilization in the non-pregnant patient with mania maintain therapy with the medication that managed the acute mania initially. Additional medications for mood stabilization can include: Topiramate: Titration is required starting at 25 mg bid times 3 days, then 25 mg qam and 25 mg 2 qpm, then 25 mg 2 bid, the 25 mg 2 gam and 3 gpm, then 25 mg 3 gam and 3 gpm with each of these transitions occurring over one week (as in 25 mg gam and 25 mg 2 qpm times 1 week). After the 3 qam and 3 qpm the final dosing is 50 mg 2 qam and 2 qpm as a final dose, which becomes the chronic dosing.

Maintenance therapy should involve monitoring for relapse or recurrence of the initial presenting symptoms. Also look for adverse effects of the medications, both neurological (extrapyramidal side effects and tardive dyskinesia) and metabolic (obesity, diabetes, dyslipidemia) or toxicity complications (renal, hematologic, hepatic). When using antidepressants be aware that these may trigger or exacerbate mania, cycling or even suicide.

# INITIAL TREATMENT

- Treat the worst presenting symptoms first.
- Suicidal Ideations Major Depressive Episode Manic Symptoms Psychosis

MONITORING **AND FOLLOW-UP Relationship Building Medication** Management Laboratory Monitoring **Primary Health Maintenance and** Wellness Primary Source for Referrals Frequent Follow up

## **INDIVIDUAL ISSUES**

- #1 problem Lack of Insight
- #2 problem Lack of Trust
- #3 problem Lack of Motivation
- #4 problem Medications Intolerance
- #5 problem Discontinuation of Treatment

# **REFERRAL ISSUES**

#### Access

 Availability – Growing gap between mental health services demand and available mental health (MH) professionals. • Travel – Geographic distribution of MH providers.

# AFFORDABILITY

- Insurance Until recently mental health services often not covered.
- State Aid Programs Qualifying for State programs requires ability to complete application forms.

#### NURSE PRACTITIONE R EXPANDED

- Primary Care NPs are the first to identify the need for mental health services.
- Primary Care NPs are frequently the provider to which the individual returns for further care.

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# RESOURCES / MEBSUPHysiEanStp://www.aafp.org

- American Psychological Association <a href="http://www.apa.org">http://www.apa.org</a>
- Cleveland Clinic Journal of Medicine <a href="http://www.ccjm.org">http://www.ccjm.org</a>
- http://www.freemedicaljournals.com
- Johns Hopkins University http://www.johnshopkinshealthalert.com
- National Alliance on Mental Illness (NAMI) http://www.nami.org
- National Institute of Mental Health (NIMH) http://www.nimh.nih.gov
- U. S. Preventative Services Task Force

#### **ASSESSMENT TOOL WEBSITES**

- The Mood Disorder Questionnaire is obtained online one reliable source is [http://www.dbsalliance.org/pdfs/MDQ.pdf].
- HCL-32 can be retrieved at [http://www.psycheducation.org/depression/HCL-32Listonly.pdf.]
- PROS is found at several reliable sites, the following is easily accessed, [http://lmasettlements.com/documents/psych\_review.pdf].
- PHQ-9 available at [http://www.integration.samhsa.gov/images/res/PHQ%20-%20Questions.pdf]. (World Health Organization – approved for unrestricted use with copyright acknowledgment.)
- Adult ADHD Self-Report Scale (ASRS-v1.1) Symptom Checklist available at [http://www.med.harvard.edu/ncs/ftpdir/adhd/18%20Question%20ADHD-ASRS-v1.1.pdf].
- CAGE is found
   [http://www.hopkinsmedicine.org/johns\_hopkins\_healthcare/download s/CAGE%20Substance%20Screening%20Tool.pdf].
- DAST can be obtained at [http://www.leademcounseling.com/wpcontent/uploads/DAST.pdf].
- MAST can be downloaded from [http://www.leademcounseling.com/wpcontent/uploads/MAST.pdf].
- GAD-7 is retrievable at [http://www.psychiatrictimes.com/all/editorial/psychiatrictimes/pdfs/sca le-GAD7.pdf].
- The pocket guide can be downloaded from [http://www.mentalhealth.va.gov/docs/Suicide-Risk-Assessment-Guide.pdf].