#### MOOD FLIPS and BAD TRIPS : WAYS OUT OF THE MAZE

Co-Occurring Disorder Conference October 1, 2007 Daniel E. Wolf, D.O.



#### "Face it Fred, you're lost."



#### WAYS OUT OF THE MAZE

- Me; me and my patient; me and my patient and his/her spouse or SO and family / friends
- You; you and your patient, you and your patient and his/her spouse or SO and family / friends

#### WAYS OUT OF THE MAZE

#### "There is nothing about a caterpillar which suggests that it will one day become a butterfly."

**Buckminster Fuller** 

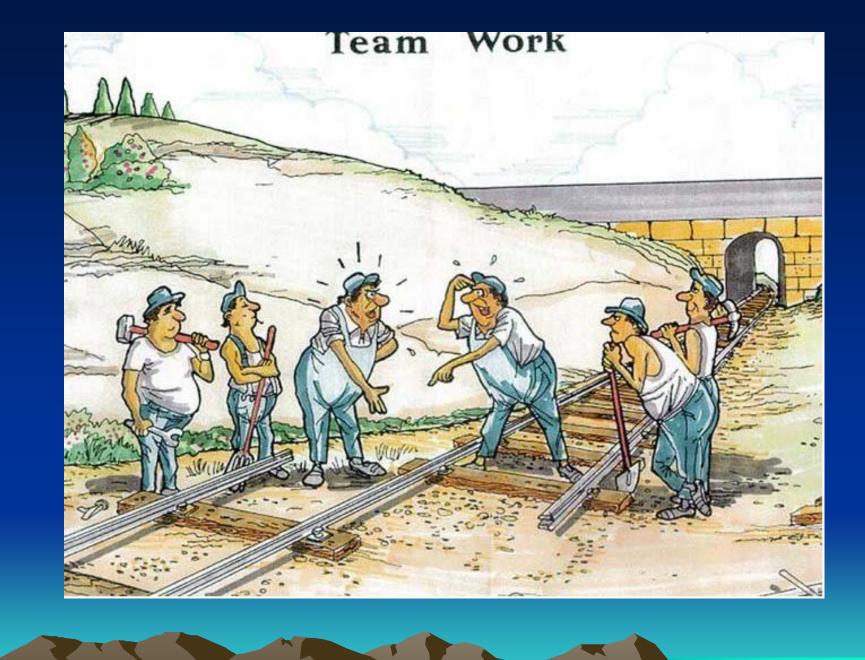
#### WAYS OUT OF THE MAZE

#### • Our patients' treatment team:

- CDC (inpatient or outpatient CD treatment)
- Therapists
- Primary care providers
- Specialists, inc. psychiatrist, PhD, ARNP, etc.
- Agencies
  - Community Mental Health Centers
  - DSHS, ADATSA, etc.
- Other



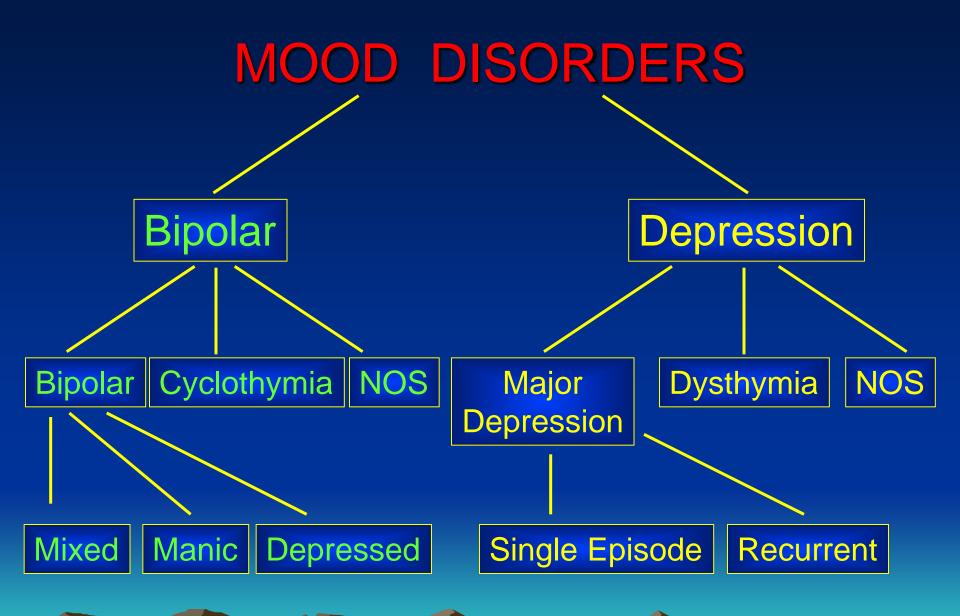
What most companies need is a disorganization chart.



#### CLOSE TO HOME

#### BY JOHN McPHERSON





American Psychiatric Association. DSM-IV. Washington, DC: APP;1994.

#### **BIPOLAR TYPES**

 Bipolar type I At least one mania - Bipolar type II At least one hypomania -Bipolar NOS At least one mania or hypomania but insufficient duration for I or II

#### MANIC EPISODE

- Elevated, expansive, or irritable mood for ≥1 wk, plus ≥3 of the following:
  - Inflated self-esteem or grandiosity
  - Decreased need for sleep
  - Pressured speech
  - Racing thoughts/flight of ideas
  - Distractibility
  - Psychomotor agitation/increase goal-directed activity
  - Excessive involvement in high-risk activities

American Psychiatric Association. DSM-IV. Washington, DC: APP;1994.

#### MANIA: DIGFAST

- Distractibility -- poorly focused
- Insomnia -- decreased need for sleep
- Grandiosity -- inflated self-esteem
- Flight of ideas -- racing thoughts
- Activities -- inc'd goal-directed activities
- Speech -- pressured or more talkative
- Thoughtlessness -- risk-taking behaviors
   www.medscape.com
   (Mood change plus 3+ for 1 wk + )

#### **HYPOMANIA**

- Elevated, expansive, or irritable mood, lasting ≥ 4 days, *plus*
- $\geq$  3 symptoms of mania that
  - Are not severe enough to impair functioning markedly
  - Do *not* necessitate hospitalization
  - Are not accompanied by psychotic features
- Produces enough change to be noticed by others

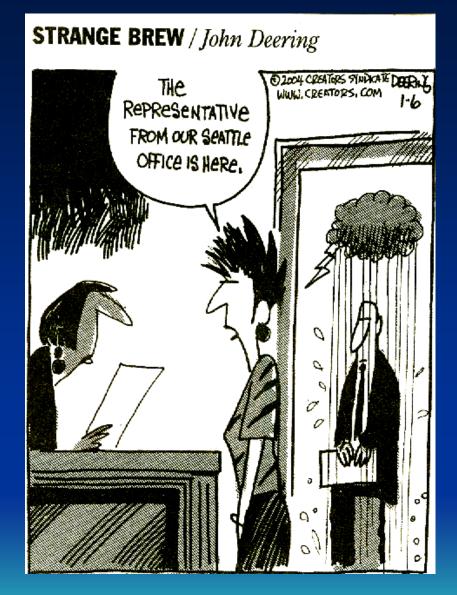
American Psychiatric Association. DSM-IV. Washington, DC: APP;1994.

#### HYPOMANIA < 4 DAYS

• Duration criteria is  $\geq$  4 days

 Many people have hypomanic symptoms of sufficient severity but insufficient duration....such as 2 to 3 days

Bipolar Disorder, Not Otherwise Specified



#### **BIPOLAR DISORDER, DEPRESSED**

- Distinct mood change OR loss of interest in things that used to give interest or pleasure PLUS
- Daily or near daily for at least 2 weeks PLUS
- A change in at least 4 out of these 7 baseline features:
  - Sleep; Appetite; Concentration; Energy; Guilt; (feelings of H/H/W); Suicidal thoughts; Psychomotor

#### **BIPOLAR DISORDER, DEPRESSED**

 AND causes significant distress or impairment in function at work or home or school

 AND is not due to another medical disorder (drug/alcohol, thyroid, meds, etc)

AND episodes of hypomania or mania

 Bipolar type I: mania; type II: hypomania
 Major Depression: Neither mania nor hypomania

#### BIPOLAR DEPRESSION WHY IS IT HARD TO DX ?

- Hypomanic and manic episode may go unnoticed by the patient
  - > 50 % have depression as 1<sup>st</sup> episode
  - Most enjoy the hypomania, or there is great productivity so hard to see it as a "problem"
- Studies show ~ 75% of mood states in people with Bipolar Disorder are depressed

#### **BIPOLAR DISORDER, MIXED**

- Criteria met for both manic episode + major depressive episode for ≥1 week
- Symptoms
  - Are sufficient to impair functioning OR
  - Necessitate hospitalization OR
  - Are accompanied by psychotic features

American Psychiatric Association. DSM-IV. Washington, DC: APP;1994.

#### **RAPID CYCLING**

- 4 or more episodes of depression, mania or hypomania in previous 12 months
- Episodes are demarcated by a switch to the opposite polarity or by a period of remission

 Often triggered by antidepressants (esp. TCA though any antidepressant can do it)

American <u>Psy</u>chiatric Association. *DSM-IV*, Washington, DC: APP;1994

#### **RAPID CYCLING**

- 13 20% of all bipolar patients
- Initial onset 20%, later onset 80%
- Predominantly in females
- Associated with increased thyroid dysfunction

## WHAT TO ASK

- Precipitant ?

   (loss / threat of loss)

   Duration of episode

   Longest ever
   Shortest ever
   Average

   Frequency/intensity
  - Tends to increase



#### Mood Disorder Questionnaire Has there ever been a period of time when you were not your usual self and...

- ... you felt so good or so hyper that other people thought you were not your normal self or you were so hyper that you got into trouble?
- ... you were so irritable that you shouted at people or started fights or arguments?
- ... you felt much more self-confident than usual?
- ... you got much less sleep than usual and found you didn't really miss it?
- ... you were much more talkative or spoke much faster than usual?
- ... thoughts raced through your head or you couldn't slow your mind down?

#### Mood Disorder Questionnaire

- ... you were so easily distracted by things around you that you had trouble concentrating or staying on track?
- ... you had much more energy than usual?
- ... you were much more active or did many more things than usual?
- ... you were much more social or outgoing than usual; for example, you telephoned friends in the middle of the night?
- ... you were much more interested in sex than usual?
- ... you did things that were unusual for you or that other people might have thought were excessive, foolish, or risky?
- ... spending money got you or your family into trouble?

MDQ NEGATIVE (not Bipolar Disorder, type I)

- May represent :
  - Bipolar Disorder, type II or NOS
  - Cyclothymic Disorder
  - Other causes of mood swings
    - Metabolic; Structural lesions in brain; Infectious; medications (esp. Prednisone > 40 to 60 mg/d)
  - Substance-Induced Mood Disorder, cyclic

## Manic Episode: Differential Diagnoses

Differential diagnosis	Consider if
Mood disorder due to a general medical	Major medical condition present
condition	• First episode at >50 years of age
Substance-induced	<ul> <li>Symptoms in context of intoxication or withdrawal</li> </ul>
mood disorder	• H/O treatment for depression
Hypomanic episode	<ul> <li>Mood disturbance not severe enough to require hospitalization or impair functioning</li> </ul>
Mixed episode	Manic episode and major depressive episode in 1 wk

American Psychiatric Association. DSM-IV. Washington, DG: APP;1994.

## Manic Episode: Differential Diagnoses

#### Differential diagnosis Consider if . . .

**ADHD** 

- Early childhood mood disturbance onset
- Chronic rather than episodic course
- No clear onsets and offsets
- No abnormally elevated mood
- No psychotic features

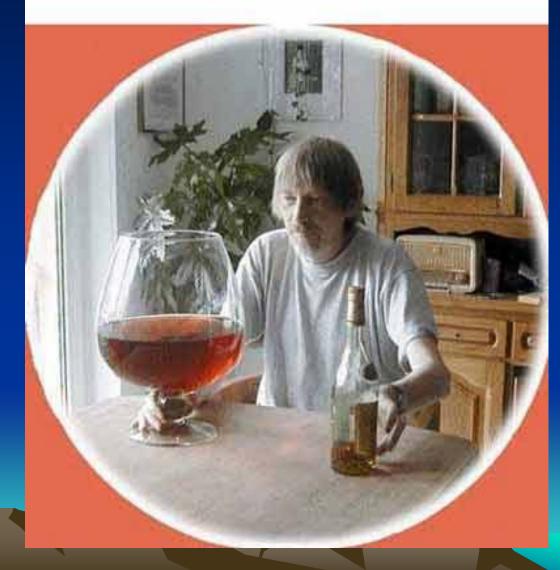
American Psychiatric Association. DSM-IV. Washington, DC: APP;1994

#### COMMONLY ABUSED DRUGS

- Alcohol
- Marijuana
- Opiates
- Cocaine, meth and other stimulants, including caffeine

- Cigarettes / Nicotine
- Benzodiazepines and barbiturates
- Hallucinogens
- Inhalants
- Rave drugs

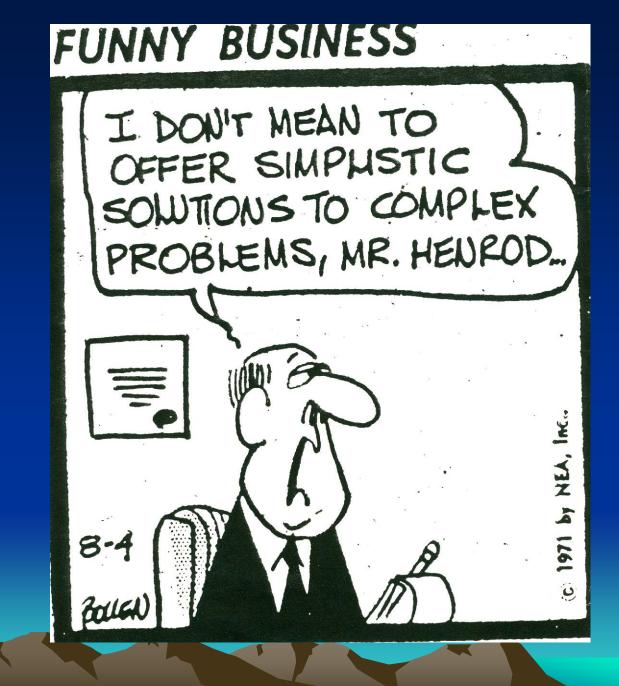
# My Doctor said "Only 1 glass of alcohol a day". I can live with that.



#### WHAT IS ONE DRINK ?

- 12 grams of pure alcohol
  - = one 12 oz beer
    = one wine cooler
    = one 5 oz glass wine
    = 1.5 ounces of distilled spirits (a jigger, vs. shot = 1 oz)





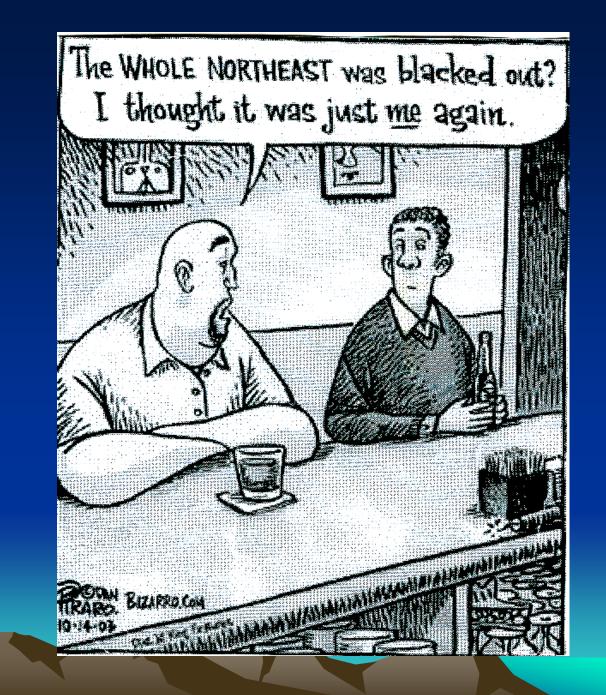
#### By Roger Bollen



#### ALCOHOL / DRUG HISTORY

- Age 1<sup>st</sup> drink / use
  - Amount consumed / used
- Last intoxication w alcohol; last drug use
  - Amount able to "hold" w/o gross impairment
  - ? Increased tolerance
- Episodes of loss of control and what was done to contain or reduce future episodes

– ? Success of control efforts ?



# DRUGS AND ALCOHOL Anything that causes a problem, is a problem

 If your alcohol or drug use causes you problems, then you have an alcohol and/or drug problem
 PLUS whatever problems the alcohol and/or drugs caused you

# • 3 C's:

#### C ompulsive Use

– Loss of C ontrol

# C ontinued use despite adverse consequences

# SUBSTANCE-INDUCED MOOD DISORDER

- Mood changes due to direct or indirect effects of drugs and/or alcohol in the brain
- Different from mood variability seen in the first several weeks to months of recovery from SUD
- Often requires medication and concurrent mental health tx and usual SUD tx

#### ABSTINENCE

- Staying "clean and sober" is no guarantee that problematic mood disruptions will go away
- Untreated problematic mood disruptions will undermine effective treatment of the SUD
- Cannot get full benefit of treatment of mood disorder if person continues to drink or use

# (IDEAL) OBJECTIVE

• Differentiate:

- Bipolar Disorder

#### from

- Substance-induced mood disorder, cyclic from
- Mood swings of other etiologies

#### HAS BD AND SUD: NOW WHAT ?

- Is patient Safe ?
  - Assess risks:

- Suicide; homicide; self-care; shelter; support

Is patient Sane ?

Any impairing psychotic symptoms

Is patient "Sober and clean ?"

- Encourage / motivate ideal of abstinence

• Is patient Stable ?







#### Practicing Safe Sex at the Olympics

#### TREAT IMPAIRING SYMPTOMS

Mood stabilizing medication

Antipsychotic medication
 Atypical vs. Conventional

#### Combination

### **TX CONSIDERATIONS**

#### Psychiatric features

- psychotic; suicide, homicide, and violence

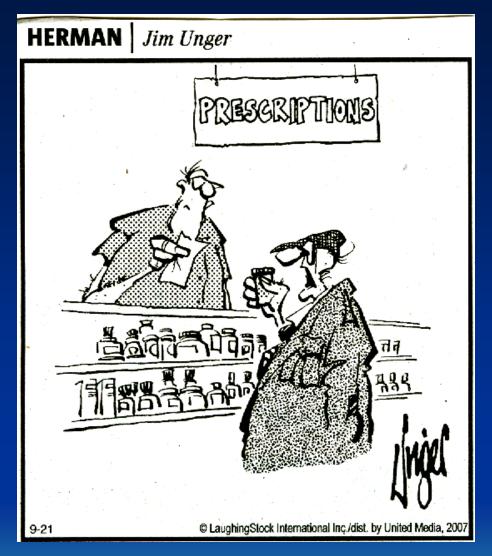
#### Demographic & psychosocial factors

- gender, cross-cultural issues, geriatric patients

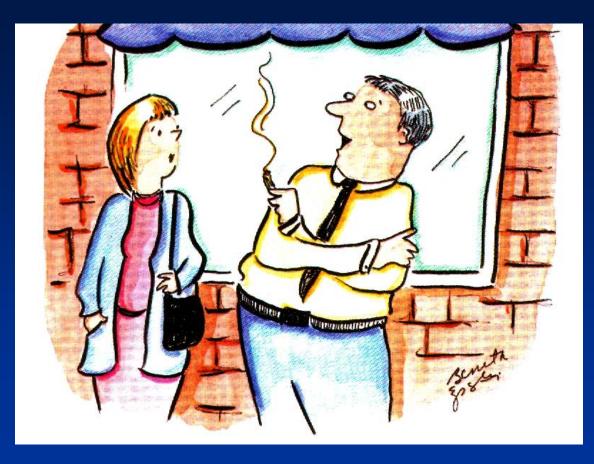
#### Concurrent medical conditions

- HIV infection, other medical

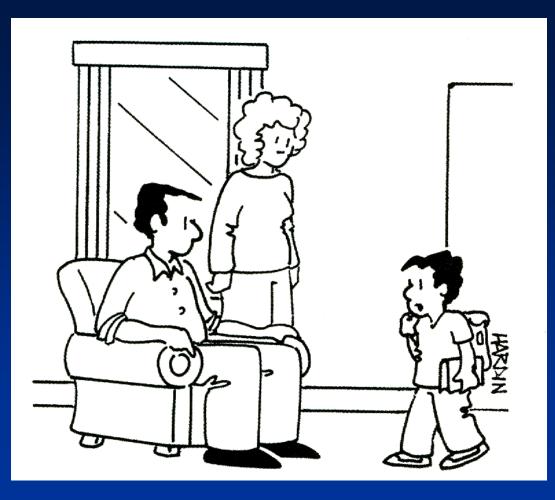
Hirschfeld RMA et al. Am Journal of Psychiatry 2002;159(4):1-50



"Are there any side-effects to these besides bankruptcy?"



#### "I only smoke now so that I can sue someone for it later."



"Boy, what a depressing day. We studied hereditary."

- Lithium \*
   Lamotrigine \*
- Divalproex \*

 Equetro \* (Carbamazepine ER)

\* FDA-approved for use in Mania for Bipolar Disorder

(why this is important: Liability risk and "off-label" term used more and more by HIPs to deny coverage)

- Lithium carbonate
  - Eskalith, Lithobid, Lithium
- Divalproex
  - Depakote ER, Valproic acid
- Carbamazepine
  - Tegretol, Carbatrol ER
  - Equetra
- Oxcarbazepine
  - Trileptal

- Neurontin
  - Gabapentin
- Lamotrigine
  - Lamictal
- Topiramate
  - Topamax
- Tiagibine
  - Gabatril

Keppra

 Levetiracetam

 Zonegran

 Zonisamide

 Newer ones

## **ATYPICAL ANTIPSYCHOTICS**

- Clozaril (Clozapine)
- Risperidone (Risperdal)
- Quetiapine (Seroquel)
- Olanzapine (Zyprexa)
- Ziprasidone (Geodon)
- Aripiprazole (Abilify)
- Invega (Paliperidone)

#### **CONVENTIONAL ANTIPSYCHOTICS**

- Haldol (Haloperidol)
- Stelazine (Trifluoperazine)
- Thorazine (Chlorpromazine)
- Mellaril (Thioridazine)
- Trilafon (Perphenazine)
- Navane (Thiothixene)
- Prolixin (

### ACUTE MANIA

• With or without psychosis: Meds of choice:

Lithium OR Divalproex + Risperidone OR Olanzapine

ARE BETTER THAN ANY ONE ALONE

#### **MEDICATION COMBINATIONS**

 Combinations often needed to achieve symptom control, if not remission, and to prevent future episodes

 Each additional medication increases risk of side - effects and/or med interactions

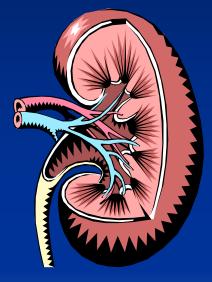
Hirschfeld RMA et al. Am Journal of Psychiatry 2002;159(4):1-50

# LITHIUM

- Lithium still effective after 50+ years
- Ability to tolerate Lithium is greater during acute mania and then decreases
- Start dose: 300 mg TID or once daily
- Serum lithium level:
  - 10 to 14 hours post-last dose

• Q 4 - 7 days until level and clinical condition stable or until Fieve R, side-effects, preclude higher doseizer: 50+ years later. Poster presented at: Annual Meeting of the American Psychiatric Association; May 18-23; Philadelphia, Pa.

## LITHIUM SIDE-EFFECTS



- Relative Contraindications:
  - Heart or kidney dis.
  - Diuretic use
  - NSAID use
    - Inc. Lithium levels 40 + %
  - Chronic diarrhea
    - Li caps vs. tabs
  - Psoriasis / acne

# LITHIUM SIDE-EFFECTS



- Hand tremors
- Urinary frequency
- GI (nausea, diarrhea)
- Metallic taste
- Weight gain
  - 20 to 25 % gainweight, often > 15 lbs

## ANTICONVULSANTS

- Except Lithium, all other mood stabilizing medications are anticonvulsants with shared side-effect risks:
  - Sedation, fatigue
  - Blurred / double vision
  - Balance problems, clumsy, falls
  - Cognition fuzziness, forgetfulness
- These can occur in therapeutic doses, too

## **DIVAPROEX** (Valproic acid)

- Loading dose for middle aged folks:
  - 30 mg/kg body wt for 2 days, then
  - 20 mg/kg body wt
- Effective serum Depakene levels achieved in 3 days and well-tolerated

Keck, Allen et al. Safety and Efficacy of Rapid-Loading Divalproex Sodium in Acutely Manic Bipolar Patients, 1999, Poster Session 152<sup>nd</sup>, Meeting APA

# **DIVAPROEX** (Valproic acid)

• Alternate strategy:

Take patient's weight in lbs, add "zero," and it converts to oral dose mg/day



### DIVALPROEX

#### Divalproex

- Peak onset 2 hours, half-life 10 hours
- TID, sometimes BID dosing

#### Depakote ER

- Peak onset 5 hours; HL 9 to 16 hours but slow release is over 18 to 24 hours, so once daily dosing, AM or HS; steady state 3 days
- More consistent efficacy through the day
- Less side-effects (less peak and trough)

#### EQUETRO (CARBAMAZEPINE ER)

- Acute mania: 200 mg BID for 3 to 7 days, then 400 mg BID
  - Serum Tegretol levels not recommended
- Average effective dose 700 mg daily
- 100, 200, and 300 mg caps
  - Can be pulled apart and sprinkled on food
- Pre-treatment CBC w plts, AST
- Many med interactions

#### Dosing to minimize risk of skin rash

- Weeks 1 2 25 mg QD
- Weeks 3 4 50 mg QD
- Week 5 100 mg QD
- After week 5: increase by 100 mg/wk, up to 300 mg/day if clinically indicated
- Not for acute mania

Calabrese JR, et al. J Clin Psychiatry 2000; 61: 841-50



• Metabolized mostly by glucuronic acid conjugaton, so: medication interactions:

Doubles Lamotrigine levels by ↓ clearance:
Divalproex (Depakote ER; Valproic acid)
Halves Lamotrigine levels by ↑ clearance:
Carbamazepine (Tegretol); Dilantin (Phenytoin); Phenobarbital (Phenobarb); Primidone (Mysoline)

- Increases Lamotrigine blood levels: – BCPs ; no effect on contraception
  No impact on Lamotrigine levels: – Lithium (and no impact on Lithium level)
  - Trileptal (Oxcarbazepine)
  - Keppra (Levetiracetam)

- Risk of rash: same as any med when dosing protocol is followed
- See Doctor ASAP if following occur:
  - Skin rash w/wo hives
  - Fever
  - Swollen glands in neck
  - Painful sores in mouth or around eyes
  - Swelling of lips or tongue

## NEURONTIN

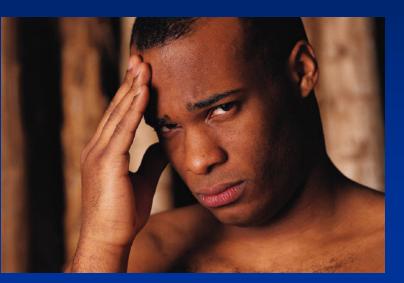


- Not effective in Bipolar Disorder when used as sole mood-stabilizer
- Effective as add-on
- No medication interactions
- Kidney metabolism

## NEURONTIN

#### Indications:

- Inadequate mood stability with 1<sup>st</sup> mood stabilizer
- Significant anxiety component
- No medication interactions



## NEURONTIN

- Caps: 100 mg, 300 mg, 400 mg
- Tabs (generic): 400 mg, 600 mg, 800 mg
- Peaks in 2 hours; ½ life 10 hours
  - Can be used prn acute anxiety or breakthrough mood reactivity
- Starting dose: 300 mg TID or more aggressive depending on acuity/severity

## NON-MED TX OPTIONS

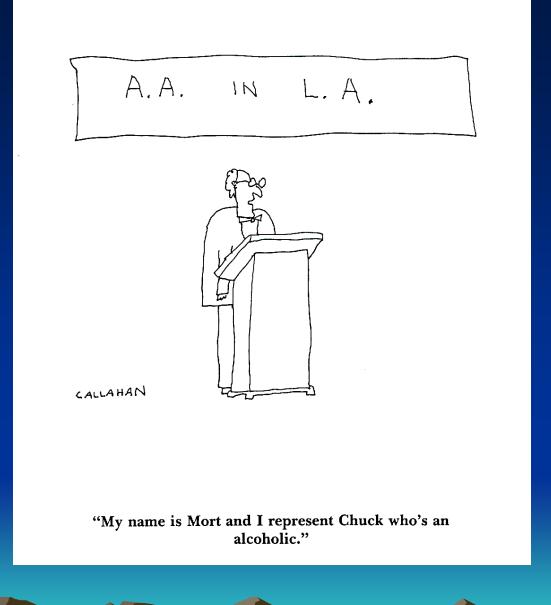




- Omega-3-fatty acid
- Psychotherapy "here and now" initially
- Exercise / Sleep
- Phototherapy
- ECT

#### **RECOVERY PSYCHOTHERAPY**

- Individual therapy
- Couples counseling
- Family therapy
- Group therapy
- Combination med / therapy group



#### NON-AA MUTUAL SELF-HELP GROUPS

- Dual Recovery Anonymous (DRA)
- Sobriety Knowledge Is Power (SKIP)
- Women For Sobriety (WFS)
- Self-Management Alcohol Recovery Training (SMART)
- Rational Sobriety (RR)

Lifering Secular Recovery

### SUMMARY

- Careful initial evaluation
  - Screen all depressed pts for Bipolar Disorder
- History from others
  - Family, friends, etc.
  - Other members of treatment team
    - CDC, therapist, primary care provider, specialists
- Encourage "clean and sober"

### SUMMARY

- Treat impairing symptoms first
- Serial, frequent assessments
- Adjust treatment as indicated
- Motivate "clean and sober"
- Collaborate and consult
- Repeat

#### CLOSE TO HOME

#### BY JOHN McPHERSON



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