MHIF Research Highlights: JANUARY 2020

SHARING EXPERTISE:

Dr. Miedema was featured in *Men's Health Magazine...* in an article, "The Great Millennial Blood Pressure Problem," addressing why high blood pressure is rising for millennials.

MHIF on KSTP Channel 5 News!

Dr. Scott Sharkey and patient, Kristen Bowlds were interviewed by the local KSTP, Channel 5 news for a story about women's heart research and Kristen's experience with SCAD.

FEATURED MHIF STUDIES

Open for Enrollment and Referrals!

HITSOVA for heparin induced thrombocytopenia

CONTACTS: Carina Benson, 612-863-4393 and Jane Fox, 612-863-6289

VESALIUS for high cardiovascular risk without prior myocardial infarction or stroke

CONTACT: Ezi Ebere, 612-863-4393

REDUCE LAP-HF RCT II for heart failure

CONTACT: Jane Fox, 612-863-6289

Shout out of gratitude for Dr. Wang's support of research...

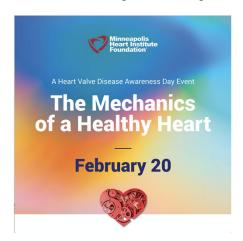


Dr. Wang is appreciated by research staff for always being so open to research and speaking with his patients about the studies! After he gives the introduction, patients are often interested in participating and we are grateful!

MARK YOUR CALENDARS

Heart Valve Awareness Event for Patients!

Thursday, February 20 Minnesota Valley Country Club



REGISTER: Mplsheart.org/valveday







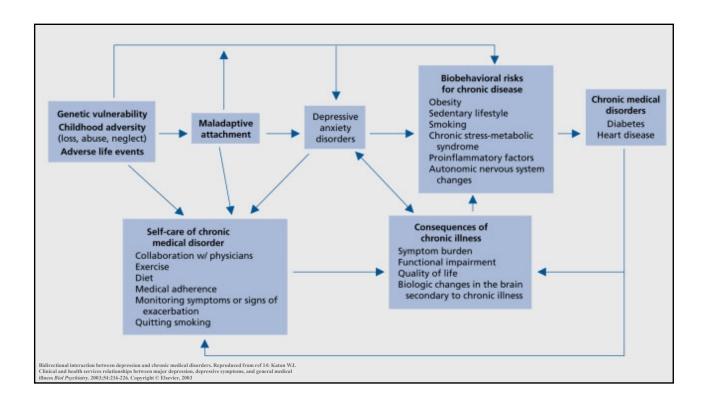
MHIF GRAND ROUNDS JANUARY 27TH 2020 Kristin Hjartardottir, DNP No affiliations to disclose



ANW PSYCHIATRY CONSULT-LIAISON SERVICE

- Two psychiatrists and a NP on a given day
- Cover the ANW med/sur, cardiology, neurology, ICU, MBC, CKRI.
- Mon-Frid and urgent coverage on the weekends
- Largest portion of consults are for depression and anxiety.
- Delirium, dementia with behavioral disturbance
- · Dual diagnosis
- Conversion disorders, somatic symptom disorders
- Capacity
- · Other mental illnesses
- Medications

Psychiatric illnesses as a risk factor for medical problems Psychiatric treatment as a risk factor Medical illnesses causing psychiatric problems Psychosomatic issues Mental health is not only the absence of diagnosable illnesses





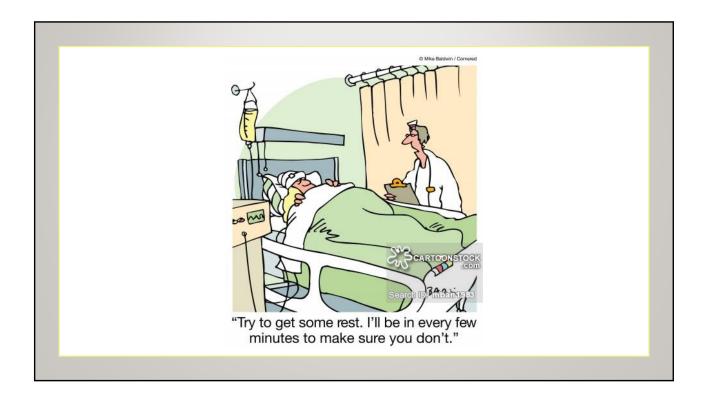
PSYCHIATRIC ISSUES IN CARDIOLOGY

PSYCHIATRIC ILLNESSES AS A RISK FACTOR

- It is estimated that patients with severe mental illness lose 25 or more years of life expectancy. Majority of those due to cardiovascular diseases.
- Chronic stress. Anxiety disorders. Panic disorders.
- Patients suffering from PTSD have increase in basal HR, and BP. PTSD can result in physiological dysregulation of the hypothalamic pituitary adrenal axis (HPA) which may contribute to cardiovascular risk factors.

OTHER PSYCHIATRIC CONSIDERATIONS

- Eating disorders
- Substance use disorders
- White Coat Syndrome/hypertension
- · Broken heart syndrome



PSYCHIATRIC TREATMENT AS RISK FACTOR

- Antipsychotic medications
- Lithium
- Tricyclic medications
- Stimulants

DEPRESSION AND CVD

- Many years of research on the bidirectional relationship with depression and CVD.
- One out of 5 patients with CVD have depression.
- 2.5 times higher the mortality rate for depressed patients
- American Heart Association advisory focuses on screening, referral, and treatment of depression for primary care providers and cardiologists.
- Outcomes have shown treatment can reduce depression, but not enough research has been on if treating depression improves cardiac outcomes.

DEPRESSION OVERVIEW

DSM 5(I diagnosis of Major Depressive disorder

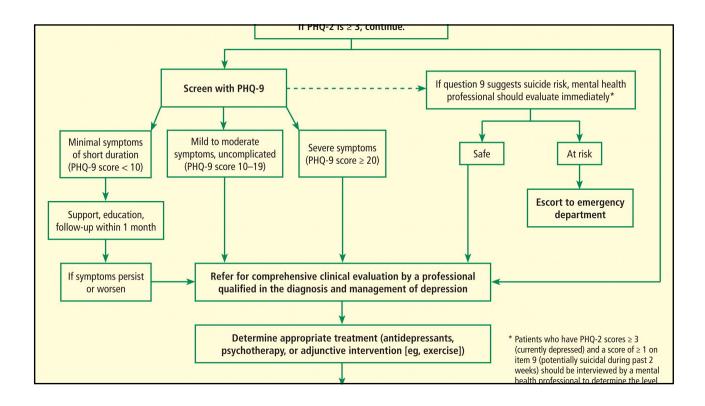
Five or more for the following symptoms during at least 2 weeks

- Depressed mood
- Diminished interest or pleasure
- Significant weight loss/gain, or decrease/increase in appetite
- Insomnia or hypersomnia nearly every day
- Psychomotor agitation/slowing nearly every day
- Fatigue or loss of energy
- Feelings of worthlessness, or excessive guild
- Diminished ability to thing or concentrate, indecisiveness.
- Recurrent thoughts of death, recurrent SI with or without a plan.

DIFFERENTIAL DIAGNOSIS

- Mood disorder due to another medical condition
- · Adjustment disorder with depressed mood
- Substance/medication induced depressive (mood disorder)
- Mixed episode of bipolar disorder
- Sadness, grief, post-partum depression, dysthymia (persistent depressive disorder)

PHQ-9 PATIENT HEALTH QUESTIONNAIRE (PHQ-9) A nine-item self-report, standardized 3 2. Feeling down, depressed, or hopeless rating scale that measures severity of 3 3. Trouble falling or staying asleep, or sleeping too much depressive symptoms and response to 5. Poor appetite or overeating treatment. Feeling bad about yourself_or that you are a failure or have let yourself or your family down Trouble concentrating on things, such as reading the newspaper or watching television 3 >20 indicates severe depression 2 <5 indicates remission A decrease ≥50% indicates a clinically significant response



EPIDEMIOLOGY

- Life time prevalence in the general population: >20 %
- 12 month prevalence ~7-10%.
- Patients with chronic illnesses have been found to have 2-3 fold higher rates of MDD
 - Diabetes 12-18%, CHD 15-23%, COPD/Asthma 20-50%
 - MS 40-60%, Alzheimer's 30-50%, stroke 14-19%

HEALTH OUTCOMES

- · Comorbid depression is associated with
 - · Increased medical symptom burden
 - · Increased functional impairment
 - Higher medical cost
 - · Poorer adherence to treatment recommendations/self care
 - Increased risk of morbidity and mortality.

Treating depression



Setting

- Most patients are being seen primarily in primary care
- Multiple studies show depression underrecognized and therefore undertreated
- Collaborative care models improving outcomes



Treatment options

- Medications
- Psychotherapy
- Exercise

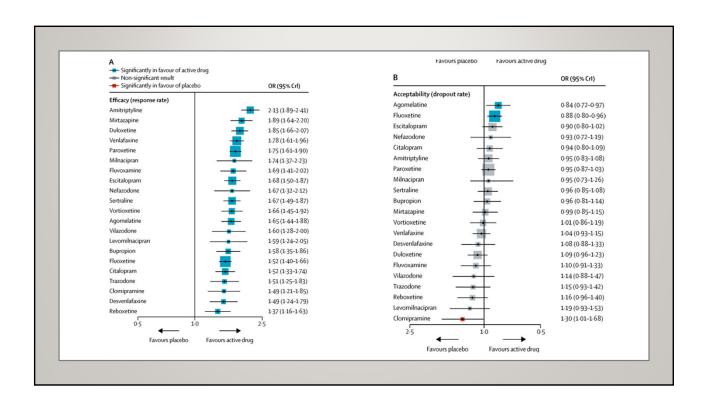


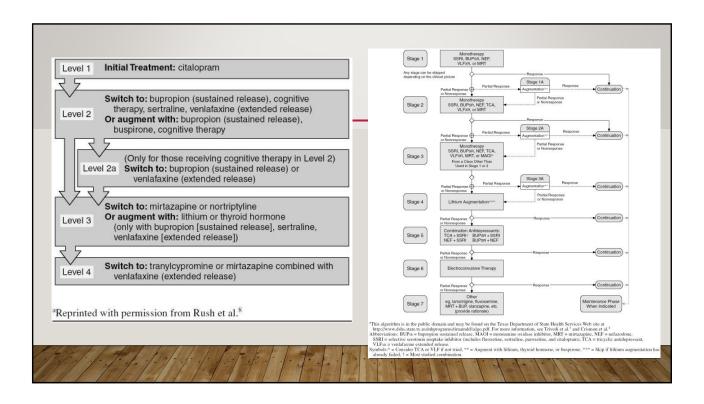
Outcomes

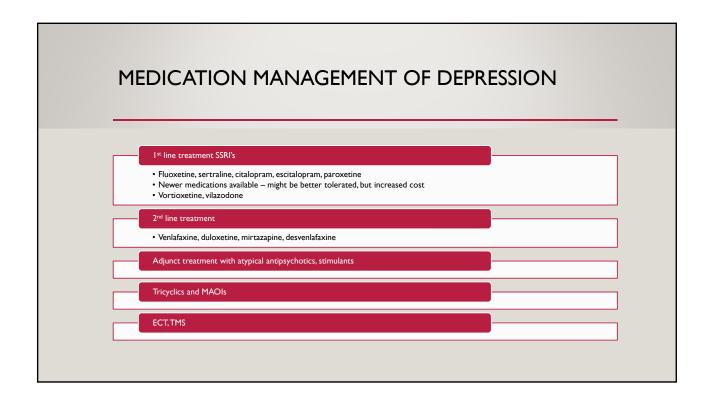
- Improved depression
- Not necessarily improved chronic illness
- Secondary gain including adherence, cost
- Ongoing studies on preventative outcomes

TREATMENT OF MDD - MEDICATIONS

- Medications are recommended for treatment of moderate to severe depression.
- 60% of patients with MDD fail to remit with initial pharmacotherapy and with each subsequent trial, smaller proportions remit.
- Treatment resistant or refractory depression that fails to remit after at least separate and adequate trials for antidepressants from two different classes.
 - · Risk factors; the longer the episode of depression increased atrophy in hippocampus
 - ? Bipolar depression
 - · Lack of symptomatic improvement in the first few weeks of treatment
 - Comorbid symptoms/disorders
 - Genetic variants for the P450 metabolizing system







FIRST LINETREATMENT SSRI/SNRI

FLUOXETINE, SERTRALINE, CITALOPRAM, ESCITALOPRAM PAROXETINE VENLAFAXINE DESVENLAFAXINE DULOXETINE Side effect profile guides a lot of our choices in treatment

- Headaches, activation/sedation, insomnia/drowsiness, upset stomach, dizziness – go away
- Dose related; tremor, QT prolongation (citalopram/escitalopram), sexual side effects
- Increased risk for bleeding during the perioperative period. In combination with warfarin due to increased prothrombin ration or INR response due to decreased platelet aggregation secondary to depletion of serotonin in platelets.
- Can induce SIADH.
- Osteoporosis
- SNRI anticholinergic effects, modest sustained increase in blood pressure, tachycardia.
- · Discontinuation syndrome,

BUPROPION

- NE and DA modulator
 - No weight gain, sedation, sexual dysfunction
 - Helpful for neurovegetative symptoms
- Caution with
 - Seizure
 - Brain tumor
 - Alcohol
 - Eating disorder
- Caution with anxiety

MIRTAZAPINE

- Increases NE and 5HT3
 - Blocks alpha 2 adrenergic presynaptic receptor
- Receptor antagonism
 - Serotonin (2A, 2C, 3)
 - Muscarinic
 - Histaminergic (H1)
 - Peripheral alpha I adrenergic

- Use of side effects antiemetic, sleep aid, appetite stimulation
- Caution with orthostasis
- No sexual side effects
- Earlier onset
- Does not affect p450
- · Caution with renal impairment
- "California rocket fuel" venlafaxine

TRICYCLICS AND TETRACYCLICS

- Amitriptyline
- Nortriptyline
- Doxepin
- Imipramine
- Clomipramine
- Desipramine

- Analgesic, sedative
- Anticholinergic, antihistaminic
- Caution
 - Cardiac conduction / hypotension / arrhythmia
 - Cognitive blunting / deliriogenic
 - · Lethal in overdose

CNS STIMULANTS

METHYLPHENIDATE

- DA, NE reuptake inhibitor and releaser
- · Optimal for TRD, medical patients
 - · Rapid onset
 - Analgesic properties
 - Palliative / demoralization
 - Post-stroke
 - Cognitive dysfunction
- SE: CV effects, anorexia, seizures, psychosis, abuse potential

MODAFANIL

- DA reuptake inhibitor
- Augmentation for antidepressants
 - Fatigue, sleepiness sx
- · Less abuse potential
- No appetite impact

WHAT IS ADEQUATE?

- Duration: generally treat for 6 to 12 weeks before deciding whether a response is sufficient.
- If patients show very little improvement (a reduction of baseline symptoms ≤25 percent) after four to six weeks, move to the next step.

• Refer for • Multiple failed trials (2-4) • Ongoing mood or behavior dysregulation • Suicidal, homicidal, psychotic, or catatonic



ADDITIONAL OPTIONS/PROCEDURES

- Light therapy seasonal component
- Electroconvulsive therapy (ECT).
 - · Requires anesthesia
 - Elicits a brief seizure
 - Side effects of anterograde amnesia
 - Brief, unilateral ECT with lower cognitive impact
- $\bullet \quad \text{Repetitive transcranial magnetic stimulation (rTMS)}.$
 - Usually only after trial of ECT has failed
 - Uses an electromagnetic coil placed against the scalp to stimulate nerve cells
 - · Fewer cognitive symptoms
 - More tolerable than ECT
- Vagus nerve stimulation (VNS).
 - Usually only after other brain stimulation therapies have not been successful
 - . Stimulation of the vagus nerve with electrical impulses through an implant in the chest to the neck to the mood centers in the brain
 - 12 month response of 30% or more

SUICIDALITY

- Part of screening with instruments like the PHQ9
- Depression as a risk factor is estimated to count for 60% of suicides
- · Chronic illness increases risk for suicide
- Other risk factors include sex, age, psychosocial status

REFERENCES

- Newcomer JW, Hennekens CH. Severe Mental Illness and Risk of Cardiovascular Disease. JAMA. 2007;298(15):1794–1796. doi:10.1001/jama.298.15.1794
- Coughlin SS. Post-traumatic Stress Disorder and Cardiovascular Disease. Open Cardiovasc Med J. 2011;5:164–170. doi:10.2174/1874192401105010164
- 3) Huang Y, Huang W, Mai W, et al. White-coat hypertension is a risk factor for cardiovascular diseases and total mortality. *J Hypertens*. 2017;35(4):677–688. doi:10.1097/HJH.00000000001226
- Redfors B, Shao Y, Omerovic E. Stress-induced cardiomyopathy (Takotsubo)--broken heart and mind?. Vasc Health Risk Manag. 2013;9:149–154. doi:10.2147/VHRM.S40163
- Yekehtaz H, Farokhnia M, Akhondzadeh S. Cardiovascular considerations in antidepressant therapy: an evidence-based review. J Tehran Heart Cent. 2013;8(4):169–176.
- Steven M. Bradley, John S. Rumsfeld, Depression and cardiovascular disease, Trends in Cardiovascular Medicine, Volume 25, Issue 7, 2015, Pages 614-622, https://doi.org/10.1016/j.tcm.2015.02.002.
- 7) Celano CM,Villegas AC, Albanese AM, Gaggin HK, Huffman JC. Depression and Anxiety in Heart Failure: A Review. Harv Rev Psychiatry. 2018;26(4):175–184. doi:10.1097/HRP.000000000000162

- 8) Comparative efficacy and acceptability of 21 antidepressant drugs for the acute treatment of adults with major depressive disorder: as systematic review and network meta-analysis. A. Cipriani, T. A. Furukawa, G. Salanti, A. Chaimani, L.Z. Atkinson, Y. Ogawa, et.al. 2018, February 21st. The Lancet.
- Trivedi, Madhukar MD. et. al. Use of Treatment Algorithms for Depression. The Primary Care Companion to the Journal of Clinical Psychiatry. 2006; 8(5): ;291-298
- Textbook of Psychosomatic Medicine: Psychiatric Care of the Medically III. Second Edition. Edt by Levenson, James MD. American Psychiatric Publising Inc. 2011. Table 8-1. p. 175-197
- 11. The Mayo Clinic. http://www.mayoclinic.org/diseases-conditions/depression/in-depth/treatment-resistant-depression
- Rush, A. John, MD. et. al. Acute and Longer-Term Outcomes in Depressed Outpatients Requiring One or Several Treatment Steps: A STAR*D Report. American Journal of Psychiatry. 2006; 163: 1905-1917

KETAMINE

- What provides the antidepressant activity?
- Opioid receptor agonist
 - Analgesic
- Activate anterior cingulate cortex
 - Emotion, impulse-control, cognitive function
- Blockade of glutamatergic NMDA receptors and facilitation of AMPA receptors

- IV, IM, intranasal, subcutaneous, oral, sublingual
- Infusion
- Requires hospital setting
- Psychiatric side effects
- Quick onset, relapse