260-2017-03-15-bipolar

Rick Gilmore 2017-03-15 12:37:04

Prelude

Today's Topics

• Bipolar disorder

Major affective (emotional) disorders

- Types
 - Bipolar disorder
 - Depression
 - Anxiety
- Heritability
 - proportion of variance in trait accounted for by genetic factors
 - Monozygotic: .69
 - Dizygotic: .13

Up/Down Documentary

Bipolar disorder

- Formerly "manic depression" or "manic depressive disorder
- Alternating mood states
 - Mania or hypomania (milder form)
 - Depression
- Cycles 3-6 mos in length, but
 - Rapid cycling (weeks or days)
- Suicide risk 20-60x normal population, (Baldessarini, Pompili, and Tondo 2006)

Symptoms

http://www.nimh.nih.gov/health/topics/bipolar-disorder/index.shtml

Prevalence, subtypes

- 1-3% prevalence, subthreshold affects another 2%
- Subtypes
 - Bipolar I: manic episodes, possible depressive ones
 - Bipolar II: no manic episodes but hypomania + depression

Related symptoms

- Psychosis (hallucinations or delusions)
- Anxiety, attention-deficit hyperactivity disorder (ADHD)
- Substance abuse

Genetics

- Overlap between bipolar disorder and schizophrenia
- Genes for voltage-gated Ca++ channels
 - Regulate NT, hormone release
 - Gene expression, cell metabolism
- (Craddock and Sklar 2013)

Brain responses to emotional faces depression

```
(Lawrence et al. 2004)
```

(Lawrence et al. 2004)

(Lawrence et al. 2004)

Amyg, Hip volume reduced,

(Hallahan et al. 2011)

(Hallahan et al. 2011)

Drug treatments

- Mood stabilizers
 - Lithium (Li)
 - Valproate (Depakote)
- Anticonvulsants
 - GABA agonists
 - Usually to treat epilepsy
 - e.g. lamotrigine (Lamictal)
- Atypical antipsychotics

Lithium "discovered" accidentally

- Injections of manic patients' urine with lithium compound (chemical stabilizer) into guinea pig test animals.
- \bullet Had calming effect
- John Cade discovered in 1948.
- Earliest effective medications for treating mental illness.

Effects of Lithium

- Reduces mania, minimal effects on depressive states
- Increases/preserves PFC, hip, amyg volume
- downregulates DA, glu; upregulates GABA
- modulates 5-HT, NE
- (Malhi et al. 2013)

Other treatment options

- Psychotherapy
- Electroconvulsive Therapy (ECT)
- Sleep medications

Prospects

- STEP-BD cohort (n=1469)
 - 58% achieved recovery
 - 49% had recurrences within 2 years
 - Residual depressive symptoms can persist
- (Geddes and Miklowitz 2013)

An Unquiet Mind

BP summed-up

- Changes in mood, but depression
- Genetic + environmental risk
- Changes in emotion processing network activity, size of hippocampus
- Heterogeneous
- No simple link to a specific NT system

Next time...

• Depression

References

Baldessarini, Ross J., Maurizio Pompili, and Leonardo Tondo. 2006. "Suicide in Bipolar Disorder: Risks and Management." CNS Spectrums 11 (06): 465–71. doi:10.1017/S1092852900014681.

Craddock, Nick, and Pamela Sklar. 2013. "Genetics of Bipolar Disorder." The Lancet 381 (9878): 1654-62. doi:10.1016/S0140-6736(13)60855-7.

Geddes, John R, and David J Miklowitz. 2013. "Treatment of Bipolar Disorder." The Lancet 381 (9878): 1672-82. doi:10.1016/S0140-6736(13)60857-0.

Hallahan, Brian, John Newell, Jair C. Soares, Paolo Brambilla, Stephen M. Strakowski, David E. Fleck, Tuula Kieseppä, et al. 2011. "Structural Magnetic Resonance Imaging in Bipolar Disorder: An International

Collaborative Mega-Analysis of Individual Adult Patient Data." *Biological Psychiatry*, Bipolar Disorder: Genes and Brain Development, 69 (4): 326–35. doi:10.1016/j.biopsych.2010.08.029.

Lawrence, Natalia S, Andrew M Williams, Simon Surguladze, Vincent Giampietro, Michael J Brammer, Christopher Andrew, Sophia Frangou, Christine Ecker, and Mary L Phillips. 2004. "Subcortical and Ventral Prefrontal Cortical Neural Responses to Facial Expressions Distinguish Patients with Bipolar Disorder and Major Depression." *Biological Psychiatry* 55 (6): 578–87. doi:10.1016/j.biopsych.2003.11.017.

Malhi, Gin S., Michelle Tanious, Pritha Das, Carissa M. Coulston, and Michael Berk. 2013. "Potential Mechanisms of Action of Lithium in Bipolar Disorder." *CNS Drugs* 27 (2): 135–53. doi:10.1007/s40263-013-0039-0.