Integrated Health Topic: Anxiety in Pediatrics

Jason Schweitzer, MD Transforming Mental Health Leading the way for children to live their best lives

Introduction

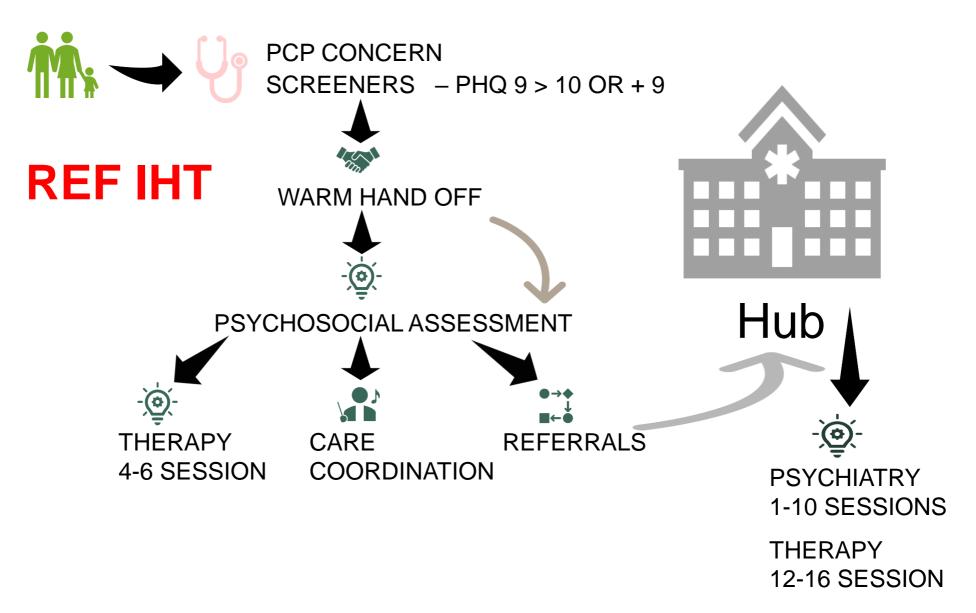
No disclosures







MHI Flow



- Infants
 - Loud Noises
 - Dropped
 - Startle Reflex



- 9 Months
 - Stranger Anxiety
- Toddlers
 - Monsters, Darkness



- Early School age
 - Physical wellbeing, injuries
 - Natural Disasters, storms
 - Spiders, Darkness



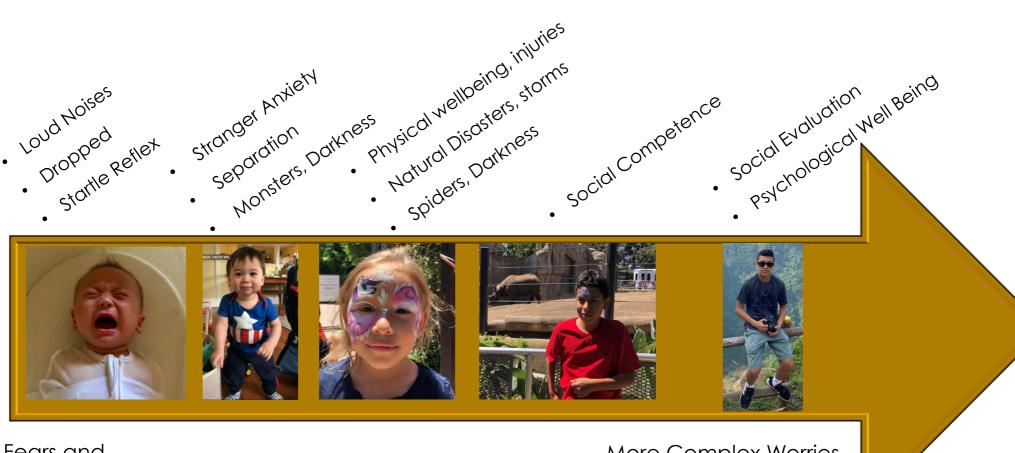
- Early Adolescence
 - Social Competence





- Social Evaluation
- Psychological Well Being





More Complex Worries Context Specific Fears

Fears and Worries

What is Anxiety?

- Worry Cognitions Apprehension, thoughts about how things could or will go wrong
 - Gets more complex over time
- Fear Perceived imminent or likely threat, more immediate
 - Diminishes over time
- Somatic Symptoms

Cognitions of Anxiety

- High frequency and salience of negative predictions
- Catastrophizing thought style
- All or Nothing Thinking
- Mental Filter
- Often co-morbid with depression

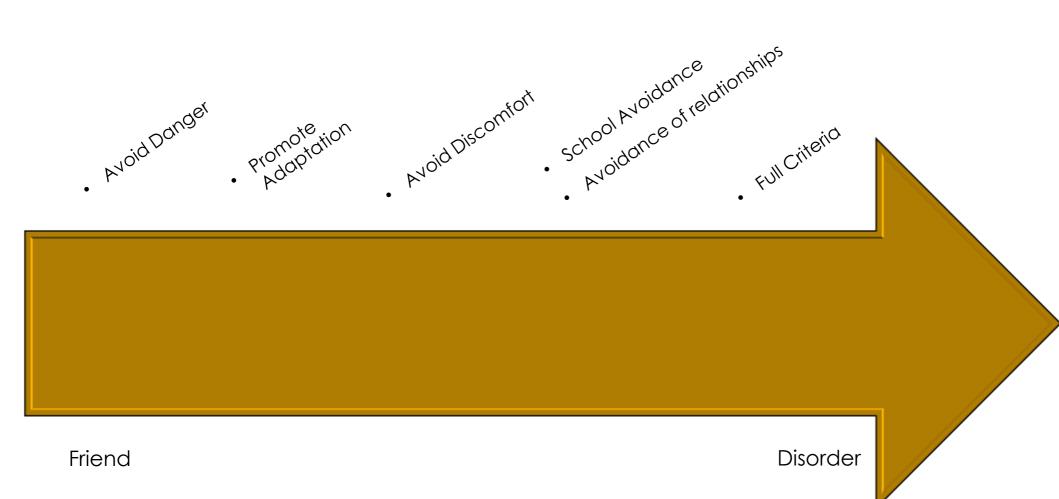
Symptoms of Anxiety

- Hand wringing, stomach/GI upset, headaches
- Panic Attacks
 - Abrupt occurrence of
 - Palpitations, trembling, sweating, cp, n/v, dizziness, chills, heat sensations, parasthesias, derealization / depersonalization, fear of dying or "going crazy"
- Sympathetic symptoms
- Somatic symptoms

Child and Adolescent Understanding Anxiety

- Problematic or Disorder
 - Symptoms
 - Distress
 - Dysfunction
 - Inflexibility
 - Avoidance
 - Eventually Anxiety in the absence of stimulus

Anxiety Friend or Foe?



Developmental Anxiety



More Complex Worries Context Specific Fears

Fears and Worries

2016 National Survey of Children's Health

- Anxiety Becomes more prevalent in Teen years
- 7.1% from 3-17 Years of Age
 - 3 5 yo : 1.3 %
 - 5 11 yo: 6.6 %
 - 12 17 yo: 10.5 %

Table I. Prevalence of currently diagnosed depression, anxiety, and behavioral/conduct problems among children aged 3-17 years, by sociodemographic and health characteristics, NSCH 2016

Approach to Anxiety

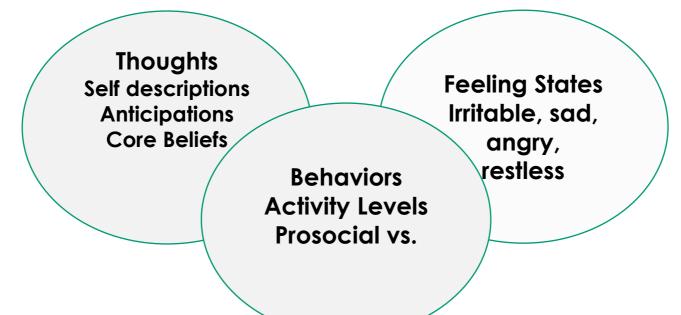
- Normalize
 - "Anxiety is a good thing, we all have it, and need it"
 - Fight or fight examples (snake, car accident)
 - Sometimes this reaction gets too strong, and people forget to turn it off. Sometimes they can't turn it off
- Does the pattern match age expected anxiety syndromes?
- Give a SCARED: 8 17 Years old
- http://www.midss.org/content/screen-child-anxiety-related-disorders-scared

Approach to Anxiety

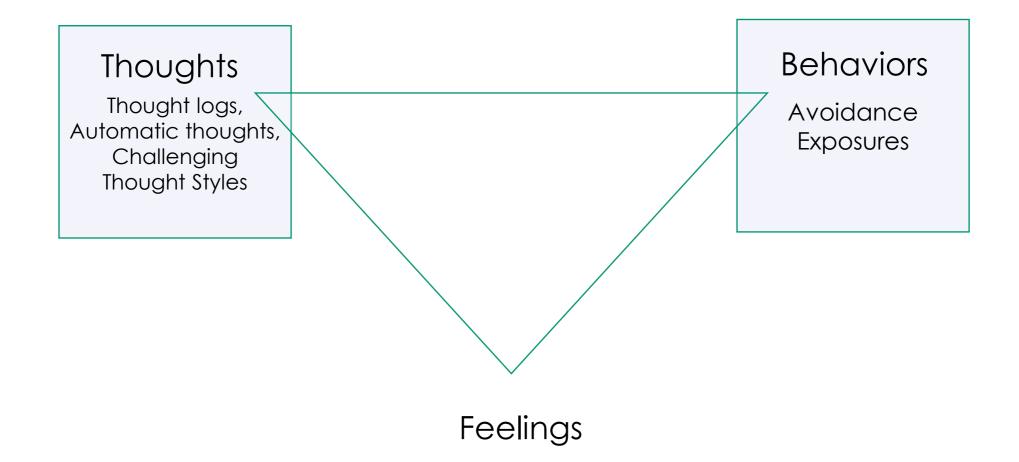
- Connect family with resources
- Therapy is first line
 - CBT
 - Anxiety scales followed by exposures targeting avoidance
- SSRIs Can be helpful
 - Medication without therapy is not first-line

Treatment

- Therapy
- Medication
- Combination therapy has best data
 - CBT for Anxiety



Treatment



- Social Anxiety Disorder
 - Fear about social settings with PEERS (not just adults), out of proportion to context
 - Concern with scrutiny of others
 - CBT
 - SSRI > SNRI
 - Multiple RCTs



- First Line: Big Three
 - Fluoxetine
 - Escitalopram
 - Sertraline
- Return in 1-4 weeks to see if tolerating, ask about SI.
 - Follow-up meetings can be brief
 - The pay off comes in the form of improved compliance, placebo response and better titration
- Up-titrate
 - Week 8 good response or maximal dose



- Second Line: A different of the big Three
 - Fluoxetine
 - Escitalopram
 - Sertraline

SSRIs for Anxiety

General

Headache, Gi Upset, SI Rare activation or mania/Hypo Mania

Black Box Warning

Cross placenta and breast milk but Generally Safe

Fluoxetine

Start at 5 -10 mg 40 - 60 mg MAX Can be more activating Withdrawal is usually well tolerated

Most interactions

Escitalopram

Start 5 – 10 mg Max is 20 mg Can be more sedating, /

orthostasis

Fewest Interactions

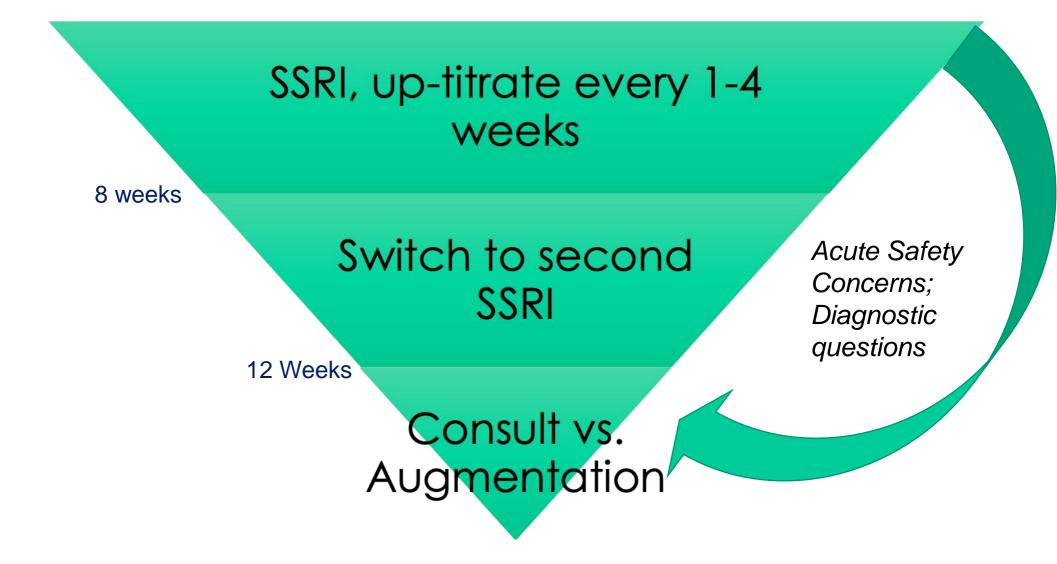
When stopping taper over 1-2 weeks, or slower

Sertraline

Start 12.5 – 25 mg

Target 125 -150 mg or remission When stopping, taper

Stepped Treatment



SSRI : Adverse Effects

- Headache, Glupset
- Black Box Warning: Suicide Events
- Antidepressants increased the risk compared to placebo of suicidal thinking and behavior (suicidality) in children, adolescents, and young adults in short-term studies of major depressive disorder (MDD) and other psychiatric disorders

Medications: SSRIS

- Rates of suicide events vary .7% 4%
- FDA and TADS study show no completed suicides
- <u>https://www.uptodate.com/contents/effect-of-antidepressants-on-suicide-risk-in-children-and-adolescents</u>
- For Families:
 - Communication about safety
 - Monitoring

Fluoxetine

- Fluoxetine 10 to 40 mg po daily (TADS Study; Cochrane Review)
- FDA Approved for Depression
- Adolescents: Start at 10 mg, within 1-4 weeks increase to 20 mg, true target is remission
- School Age: Start lower 5; Use Liquid; go Slow

Fluoxetine

- Long Half-life! (2-3 days parent, 2 week metabolite)
 - Good: no withdrawal
 - Bad: 5 week washout for MAOi
- Can be activating
 - Less often can be sedating
- CYP inhibition (2D6 \rightarrow Codeine, B blockers; 3A4 \rightarrow some benzos, Statins)

SSRI : General Approach

- Lowest effective dose, target remission
- Symptomatic and tolerating? Increase
- Effect can take 3-5 Weeks per dose change
- Monitor weekly or bimonthly
 - Suicidal ideation
 - Mania or hypomania (SLEEP, personality change, etc.)



- At baseline: IF clinical exam is concerning for biological underpinnings or any association with eating disorder
 - CBC, TSH, CMP, B12, Vit D.
- Follow-up Labs if symptoms of electrolyte abnormality

SSRI: Other Cautions

- Use with other serotonergic medications can cause
 Serotonin Syndrome
- Risk Category C, present in breast milk
- Rare adverse effects can be serious: Suicidal Ideation (Black Box), Bleeding, Electrolyte abnormalities
- Likely safe with OCPS, more studies needed (Berry-Bibee et al, 2016)

SSRI: Other Considerations

- Higher doses tend to be helpful for more anxiety
- Co-treatment with stimulants is usually safe
 - There are studies that show increase hypomania risk with this combination

Resource List

- SmartCare for Families : 858-956-5900
- SmartCare for Providers : 858-880-6405
- Psychologytoday.com
- County Sevices
 - <u>https://www.optumsandiego.com/content/dam/sandiego/documents/socdirectory/SBC_DBH-SUDRS_Provider_Directory_English.pdf</u>

References

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Ipser JC., Dj, S., Hawkridge, S., & Hoppe, L. (2010). Pharmacotherapy for anxiety disorders in children and adolescents (Review), (3). doi:10.1002/14651858.CD005170.pub2.www.cochranelibrary.com

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Strawn, J. R., Welge, J. A., Ph, D., Wehry, A. M., Keeshin, B., & Rynn, M. A. (2015). EFFICACY AND TOLERABILITY OF ANTIDEPRESSANTS IN PEDIATRIC ANXIETY DISORDERS : A SYSTEMATIC REVIEW AND META-ANALYSIS, *157* (November 2014), 149–157. doi:10.1002/da.22329

References

Table I. Prevalence of currently diagnosed depression, anxiety, and behavioral/conduct problems among children aged 3-17 years, by sociodemographic and health characteristics, NSCH 2016

	Currently diagnosed with depression				Currently diagnosed with anxiety				Currently diagnosed with behavioral or conduct problems			
Characteristics	Unweighted, n	Weighted, N	Weighted, %	95% CI	Unweighted, n	Weighted, N	Weighted, %	95% CI	Unweighted, n	Weighted, N	Weighted, %	95% CI
All children (3-17 y)		1 934 000	3.2	2.9-3.5		4 355 000	7.1	6.6-7.6		4 509 000	7.4	6.9-7.9
Severity of diagnosed condition												
Mild	806	885 000	46.3	41.4-51.2	1796	1 949 000	45.2	41.5-48.9	1342	1 748 000	39.5	36.1-42.9
Moderate	716	841 000	44.0	39.1-49.0	1634	2 037 000	47.2	43.4-51.1	1445	2 110 000	47.6	44.1-51.2
Severe	135	185 000	9.7	6.6-14.0	308	326 000	7.6	6.2-9.2	335	572 000	12.9	10.6-15.7
Current depression		N/A			1280	1 402 000	32.3	29.1-35.8	673	908 000	20.3	17.7-23.2
Current anxiety	1280	1 402 000	73.8	69.4-77.8		N/A			1308	1 630 000	36.6	33.2-40.1
Current behavioral or conduct problems	673	908 000	47.2	42.3-52.2	1308	1 630 000	37.9	34.3-41.6		N/A		
Sociodemographic characteristics	0.0	000 000		12.0 02.2			01.0	01.0 11.0				
Sex												
Male	715	932 000	3.0	0.2-2.6	1783	2 164 000	6.9	6.2-7.7	2205	3 155 000	10.1	9.3-10.9
Female	957	1 002 000	3.3	0.2-2.9	1980	2 191 000	7.3	6.6-8.1	968	1 354 000	4.5	4.0-5.1
Age, y	301	1 002 000	0.0	0.2 2.0	1000	2 101 000	1.5	0.0 0.1	500	1 004 000	4.0	4.0 0.1
3-5	7	9000	*0.08	0.0-0.2	113	153 000	1.3	0.9-1.7	288	410 000	3.4	2.8-4.2
6-11	271	421 000	1.7	1.3-2.2	1115	1 624 000	6.6	5.7-7.6	1390	2 259 000	9.1	8.3-10.1
12-17	1394	1 504 000	6.1	5.5-6.8	2535	2 578 000	10.5	9.7-11.3	1495	1 840 000	7.5	6.7-8.3
	1394	1 304 000	0.1	5.5-0.0	2000	2 5/6 000	10.5	9.7-11.5	1490	1 040 000	7.5	0.7-0.3
Race/ethnicity	184	330 000	2.2	1.6-2.9	375	915 000	6.0	4.8-7.5	364	837 000	5.5	4.5-6.7
Hispanic												
Non-Hispanic white	1198	1 088 000	3.4	3.1-3.8	2908	2 713 000	8.6	8.0-9.2	2158	2 394 000	7.6	7.0-8.2
Non-Hispanic black	110	331 000	4.2	3.1-5.6	136	358 000	4.5	3.4-5.9	287	848 000	10.7	9.1-12.7
Non-Hispanic multiracial/other	180	184 000	2.9	2.3-3.6	344	368 000	5.7	4.4-7.5	364	430 000	6.7	5.3-8.5
Family structure												
Two parents, married	899	904 000	2.3	2.0-2.7	2385	2 504 000	6.4	5.8-7.0	1753	1 986 000	5.1	4.6-5.6
Two parents, unmarried	130	172 000	3.5	2.3-5.2	249	404 000	8.1	5.9-11.1	253	508 000	10.2	7.7-13.4
Single mother	404	585 000	5.9	5.0-7.0	736	946 000	9.6	8.4-11.0	674	1 177 000	12.0	10.6-13.6
Other	218	254 000	4.5	3.6-5.6	340	430 000	7.6	6.3-9.3	422	715 000	12.7	10.7-15.0
Household educational attainment												
Less than high school	53	262 000	4.6	3.0-7.0	83	419 000	7.3	5.3-10.1	111	499 000	8.7	6.5-11.6
High school, GED, or vocational training	266	422 000	3.6	2.9-4.4	467	810 000	6.9	5.7-8.3	524	988 000	8.4	7.2-9.7
More than high school	1317	1 191 000	2.9	2.6-3.2	3139	3 013 000	7.2	6.7-7.8	2438	2 840 000	6.8	6.3-7.4
Household poverty												
<100% FPL	273	625 000	4.8	3.8-6.0	463	984 000	7.6	6.3-9.0	557	1 405 000	10.8	9.4-12.4
100%-199% FPL	334	420 000	3.1	2.4-3.9	659	1 010 000	7.4	6.1-9.0	650	1 024 000	7.5	6.4-8.8
200%-399% FPL	500	453 000	2.8	2.2-3.4	1136	1 124 000	6.8	5.9-7.9	923	1 066 000	6.5	5.6-7.5
≥400% FPL	565	436 000	2.4	2.0-2.8	1505	1 237 000	6.8	6.1-7.6	1043	1 014 000	5.6	5.0-6.3
												(continued)
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Cochrane Review

- Ipser JC, Stein DJ, Hawkridge S, Hoppe L (2009). Pharmacotherapy for anxiety disorders in children and adolescents (Review). Cochrane Review
- 22 short-term (<= 16 weeks) RCTs (2519 participants)
- 15/22 studies assessed the efficacy of the SSRIs
 - Medication and placebo response occurred in 58.1% and 31.5% of patients,
 - Number needed to treat = 4.
 - Medication was more effective than placebo in reducing overall symptom severity in OCD in a post-hoc comparison (N = 7, Weighted Mean Difference (WMD) = -4.45, 95%Cl = -5.94, -2.97, n = 765).
- Medication treatments can reduce core symptoms
- Best data is for OCD
- No clear evidence to show that any particular class of medication is more effective or better tolerated than any other.
- Routine use of benzodiazepines cannot be recommended, especially given concerns of dependency and AEs

- Separation anxiety
 - > 4 weeks and >= 3 of
 - Developmentally inappropriate fear concerning separation for attachment figures
 - Persistent worry about bad things happening to attachment figures
 - Alterations in sleep, absenteeism, nightmares, somatic symptoms
 - CBT and SSRIs

- Selective Mutism
 - > 1 month of consistent failure to speak in certain settings, that interferes with schooling, social relationships
 - Exposures, CBT
 - SSRI

- Panic Disorder
 - Recurrent panic attacks
 - Fear of inducing attacks (> 1 Month)
 - Resulting change in behavior
 - Treatment: CBT and SSRIs (at least 3 RCTS)

- Generalized Anxiety Disorder
 - >= 6 mo of worry that is difficult to control and about many things
 - >=]
 - Restlessness, fatigue, poor concentration, irritability, muscle tension, sleep probs
 - CBT
 - SSRI, SNRI

- Specific Phobia
 - > 6 Mo → Marked fear or anxiety of a thing that is out of proportion to associated danger
 - Can be expressed by tantrums, freezing or clinging
 - Impairing
- EXPOSURES
- Data for SSRI is less convincing, but exposures work so well, that we use medication less

Escitalopram

- Escitalopram 10 to 20 mg po daily (TADS Study; Cochrane Review)
- FDA Approved for depression
- Adolescents: Start at 5 mg, increase to 10 mg, true target is remission, increase accordingly
- School age kids go slower

Escitalopram

- Half life is 24 32 hrs
- Discontinuation withdrawal symptoms rare, but can happen \rightarrow taper
- Few interactions

Sertraline

- Sertraline 25 150 mg
- 130 mg is the average studied dose
- FDA Approved for OCD
- Adolescents: Start at 25 mg, increase to 50 mg, true target is remission, increase accordingly
- School age kids go slower

Sertraline

- Half-life is 24-36 hours, can be longer
- Taper on discontinuation 50% over 3 days to week
- Some CYP inhibition