DSM-5 and Autism

Changes are in the air

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“New Definition of Autism Will Exclude Many, Study Suggests”

- "proposed changes in the definition of autism would sharply reduce the skyrocketing rate at which the disorder is diagnosed"

- "proposed changes would probably exclude people with a diagnosis who were higher functioning"

- "experts…strongly disagree"
  - “I don’t know how they’re getting those numbers”, Catherine Lord

“New Definition of Autism Will Exclude Many, Study Suggests”

- “45% of children and adults with higher functioning autism will qualify for proposed DSM-5 criteria”
- Access to services may be threatened
- Increased clarity

• New York Times, Jan 19, 2012 by Benedict Carey
Autism Prevalence
On The Rise
There has been a 600% increase in prevalence over the last two decades.

AUTISM SPEAKS™
It’s time to listen.
www.AutismSpeaks.org

*Recent research has indicated that changes in diagnostic practices may account for at least 25% of the increase in prevalence over time, however much of the increase is still unaccounted for and may be influenced by environmental factors.
Prevalence continues to grow

- Current CDC estimates 1 in 88 children aged 8 years have an autism spectrum disorder
  - 1 in 54 boys
  - 1 in 252 girls


- Reasons for increase
  - Awareness
  - Newer community studies
  - Better methods
Definitions

- **Sensitivity**
  - Measures a test’s ability to find those patients with a disease
  - If a person has a condition, how often will the test find it
  - Finds true positives

- **Specificity**
  - Measures a test’s ability to identify those patients without a disease
  - If a person is healthy, how often will the test find it
  - Finds true negatives

- [wikipedia.org](http://wikipedia.org)
### Contingency Table

<table>
<thead>
<tr>
<th>Expert Truth</th>
<th>AP Detection Algorithm</th>
</tr>
</thead>
<tbody>
<tr>
<td>T</td>
<td>T</td>
</tr>
<tr>
<td></td>
<td>TP</td>
</tr>
<tr>
<td></td>
<td>FN</td>
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<tr>
<td>F</td>
<td>FP</td>
</tr>
<tr>
<td></td>
<td>TN</td>
</tr>
</tbody>
</table>

**Figure 2.4.** Contingency table used in APCAT to score the fuzzy recognizer performance against the truth field defined by the human experts. The “TP” represents True Positive results, “FN” is False Negative results, “FP” is False Positive results, and “TN” is True Negative results.
Reliability vs. Validity
History of DSM and autism

♦ Autism was first recognized in DSM-III
  ♦ Monothetic (individuals had to meet all diagnostic criteria)
  ♦ Classical autism
  ♦ Emphasis on categories not etiology
History of DSM and autism

- DSM-III-R
  - Developmental orientation
  - Polythetic (individuals could meet a specific number of sub-criteria from a set of criteria)
  - Broader diagnostic group
Table 1. *Diagnostic and Statistical Manual of Mental Disorders (DSM-IV-TR)*

**Diagnostic Criteria for Autistic Disorder**

A. A total of six (or more) items from (1), (2), and (3), with at least two from (1), and one each from (2) and (3):

1. Qualitative impairment in social interaction, as manifested by at least two of the following:
   a. Marked impairment in the use of multiple nonverbal behaviors such as eye-to-eye gaze, facial expression, body postures, and gestures to regulate social interaction
   b. Failure to develop peer relationships appropriate to developmental level
   c. A lack of spontaneous seeking to share enjoyment, interests, or achievements with other people (e.g., by lack of showing, bringing, or pointing out objects of interest)
   d. Lack of social or emotional reciprocity

2. Qualitative impairments in communication as manifested by at least one of the following:
   a. Delay in, or total lack of, the development of spoken language (not accompanied by an attempt to compensate through alternative modes of communication such as gesture or mime)
   b. In individuals with adequate speech, marked impairment in the ability to initiate or sustain a conversation with others
   c. Stereotyped and repetitive use of language or idiosyncratic language
   d. Lack of varied, spontaneous make-believe play or social imitative play appropriate to developmental level

3. Restricted repetitive and stereotyped patterns of behavior, interests, and activities, as manifested by at least one of the following:
   a. Encompassing preoccupation with one or more stereotyped and restricted patterns of interest that is abnormal either in intensity or focus
   b. Apparently inflexible adherence to specific, nonfunctional routines or rituals
   c. Stereotyped and repetitive motor manners (e.g., hand or finger flapping or twisting, or complex whole-body movements)
   d. Persistent preoccupation with parts of objects

B. Delays or abnormal functioning in at least one of the following areas, with onset prior to age 3 years:

1. Social interaction
2. Language as used in social communication
3. Symbolic imaginative play

C. The disturbance is not better accounted for by Rett's Disorder or Childhood Disintegrative Disorder

From autism to Asperger’s Disorder
DSM-IV criteria for Asperger’s Disorder

• Impaired social interaction, with at least two of four criteria

• Restricted, repetitive, and stereotypical behaviors and interests with at least one of four criteria

• Significant impairment in important areas of function

• No overall delay in language

• No significant delay in cognitive development, (mental retardation), or in adaptive skills

• Criteria are not met for autism or schizophrenia
Problems with DSM-IV and autism

- Reliability
  - Best-estimate clinical diagnoses compared to those using ADI-R and ADOS
  - Experts differ on how they interpreted DSM-IV-TR data
    - Regional differences
    - Influenced by verbal IQ and language
  - DSM-IV is difficult to implement
Problems with DSM-IV and autism

- Validity
  

- Asperger’s disorder may not be distinct from high-functioning autism
  
  
  Difficult to retrospectively measure language delays by age 3 years.

- Limited neuropsychological distinctions from autism

- Precedence rule

- PDD-NOS
  
  Residual category
  
  Poorly defined, poor reliability of diagnosis
DSM-5 criteria for Autism Spectrum Disorder

♦ A. Persistent deficits in social communication and social interaction across multiple contexts, as manifest by the following, currently or by history:
  ♦ 1. Deficits in social-emotional reciprocity
  ♦ 2. Deficits in nonverbal communicative behaviors used for social interaction
  ♦ 3. Deficits in developing and maintaining, and understanding relationships
DSM-5 criteria for Autism Spectrum Disorder

B. Restricted, repetitive patterns of behavior, interests, or activities, as manifest by at least two of the following, currently or by history.

1. Stereotyped or repetitive motor movements, use of objects, or speech
2. Insistence on sameness, inflexible adherence to routines, or ritualized patterns of verbal or nonverbal behavior
3. Highly restricted, fixated interests that are abnormal in intensity or focus
4. Hyper-or hypo-reactivity to sensory input or unusual interest in sensory aspects of environment
DSM-5 criteria for Autism Spectrum Disorder revised

♦ C. Symptoms must be present in early developmental period childhood (but may not become fully manifest until social demands exceed limited capacities)

♦ D. Symptoms cause clinical significant impairment in function
<table>
<thead>
<tr>
<th>Severity level for ASD</th>
<th>Social communication</th>
<th>Restricted interests and repetitive behavior</th>
</tr>
</thead>
<tbody>
<tr>
<td>Level 3 (requiring very substantial support)</td>
<td>Severe deficits with very limited initiation of social interactions and minimal response to social overtures from others.</td>
<td>markedly interfere with functioning in all spheres. Marked distress, very difficult to redirect.</td>
</tr>
<tr>
<td>Level 2 (requiring substantial support)</td>
<td>Marked deficits with limited initiation of social interactions and reduced or abnormal response to social overtures from others.</td>
<td>obvious interference with functioning. Distress or frustration is apparent, difficult to redirect</td>
</tr>
<tr>
<td>Level 1 (requiring support)</td>
<td>Without supports in place, deficits in social communication cause noticeable impairments.</td>
<td>significant interference with functioning in one or more contexts. Resists interruption or redirection</td>
</tr>
</tbody>
</table>
Summary of DSM-5 changes for Autism

- Autism, Asperger’s disorder, and PDD-NOS will be combined into a single category

- Monothetic for social-communication symptoms
  - Combines DSM-IV criteria for social interaction and impairments of communication

- Polythetic for restricted, repetitive behaviors
  - 2 of 4 symptoms must be present
  - Sensory symptoms are included

- Universal age onset by early childhood
DSM-5 diagnostic simplicity

DSM-IV
2,027 different combinations

DSM-5 only has 11 different ways to meet diagnosis

McPartland JAACAP, 2012;51(4)368-383.
Goals for DSM-5 and autism

- Recognize “essential shared features” of ASD
  - Clearer, simpler diagnosis
  - Failure of social communication development

- Combine categorical and dimensional measures
  - Individualized diagnosis

- Reintegrate Asperger’s disorder and PDD-NOS into ASD
  
DSM-5 Neurodevelopmental Disorders Workgroup

- 13 members chaired by Susan Swedo, M.D.
  - Chief, Pediatrics & Developmental Neuroscience Branch, NIMH

- Over 6,000 hours of meetings and teleconferences
  - Formulate draft criteria
  - Field trials and data collection
  - Revision of draft criteria due Spring 2012
  - Public feedback and APA processing
  - Final draft data due December 31, 2012 with release date set for May 18-22, 2013
DSM-IV-TR vs. DSM-5

- 5,484 8-year olds in Finland rated for autism
  - 82 subjects with autism found by DSM-IV criteria
  - DSM-5 less sensitive for Asperger’s disorder and high-functioning autism
    - 46% identification rate in those with an IQ > 50.
    - None of the 11 patients with Asperger’s were seen
  - Early more stringent version of DSM-5
DSM-IV-TR vs. DSM-5

- DSM-5 had improved “construct validity” over DSM-IV-TR
  - 708 subjects from autism specialty clinic, 3Di
  - Removes
  - “Stereotyped and repetitive use of language”
  - “Lack of varied, spontaneous make-believe play or social imitative play appropriate to developmental level”
- Adds “Hyper- or hypo-reactivity to sensory input or unusual interest in sensory aspect of environment”
DSM-5 validated


- 14,744 siblings from the Interactive Autism Network
- Hybrid model of categorical and dimensional criteria supported
- "superior specificity"
- At risk for false negative
  - Females
  - Asperger’s disorder

<table>
<thead>
<tr>
<th></th>
<th>DSM-IV-TR</th>
<th>DSM-5</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sensitivity</td>
<td>0.95</td>
<td>0.81</td>
</tr>
<tr>
<td>Specificity</td>
<td>0.86</td>
<td>0.97</td>
</tr>
</tbody>
</table>
Sensitivity and Specificity of DSM-5 criteria

McPartland et al. JAACAP, 2012;51(4)368-383.

- 933 patients evaluated from DSM-IV field trial exposed to DSM-5 criteria
- Overall 60% of cases with ASD met revised DSM-5 criteria
- Overall specificity was 94.9%

<table>
<thead>
<tr>
<th>Diagnosis</th>
<th>Sensitivity</th>
</tr>
</thead>
<tbody>
<tr>
<td>Autism</td>
<td>0.76</td>
</tr>
<tr>
<td>Asperger’s disorder</td>
<td>0.25</td>
</tr>
<tr>
<td>PDD-NOS</td>
<td>0.28</td>
</tr>
<tr>
<td>IQ &lt; 70</td>
<td>0.70</td>
</tr>
<tr>
<td>IQ &gt; 70</td>
<td>0.46</td>
</tr>
</tbody>
</table>
News and Views

♦ DSM-5 results in 30-45% fewer children, adolescents, and adults diagnosed with ASD than DSM-IV-TR

♦ 10% of those with autism, 16.6% of those with Asperger’s disorder, and 50% of those with PDD-NOS will not meet criteria for ASD with DSM-5
DSM-5 vs. DSM-IV


♦ Data set of 4,453 children assessed with ADOS and ADI-R

♦ Overall sensitivity of 0.91 comparable to DSM-IV
  ♦ Lowest sensitivity for Asperger’s disorder was 0.69

♦ Specificity of 0.53 better than DSM-IV
  ♦ Specificity improved when both parent report and clinical observation was required to identify impaired social reciprocity and nonverbal behavior

♦ Most children will remain eligible for ASD diagnosis
<table>
<thead>
<tr>
<th>Study / year</th>
<th>Number of Subjects</th>
<th>Type of Sample</th>
<th>Instruments Used</th>
<th>Results</th>
<th>Limitations</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mattila et al\textsuperscript{19} / 2011</td>
<td>82</td>
<td>Screened epidemiological sample diagnosed with DSM-IV criteria</td>
<td>ADI-R, ADOS, early DSM-IV criteria</td>
<td>DSM-5 was less sensitive than DSM-IV for ASD, (0.46)</td>
<td>Early DSM-5 criteria, prevalence for PDD-NOS was not examined</td>
</tr>
<tr>
<td>Mandy et al\textsuperscript{20} / 2012</td>
<td>708</td>
<td>Consecutive referrals to an autism specialty clinic</td>
<td>3Di</td>
<td>DSM-5 model was superior to DSM-IV</td>
<td>Higher functioning sample, 3Di is a DSM-IV derived tool</td>
</tr>
<tr>
<td>Frazier et al\textsuperscript{21} / 2012</td>
<td>14,744 siblings (8,911 with autism)</td>
<td>Family-selected internet registry</td>
<td>Mapped caregiver rated SRS and SCQ to DSM-5 criteria</td>
<td>DSM-5 had lower sensitivity, (0.81 vs. 0.95) but greater specificity, (0.97 vs. 0.86) than DSM-IV</td>
<td>Early DSM-5 criteria, self-selected sample, reliance on caregiver reports only</td>
</tr>
<tr>
<td>McPartland et al\textsuperscript{4} / 2012</td>
<td>933 (657 diagnosed with ASD)</td>
<td>Multicenter DSM-IV field trial database, with clear reliability data</td>
<td>Algorithm of items from DSM-IV mapped to match DSM-5 criteria</td>
<td>60.6% of cases with ASD met DSM-5 criteria with a specificity of 94.9%</td>
<td>Included only 48 DSM-IV subjects with Asperger's disorder, modified a historical data set to new criteria</td>
</tr>
<tr>
<td>Matson et al\textsuperscript{22,23} / 2012</td>
<td>2,721 toddlers aged 17-36 months</td>
<td>EarlySteps participants</td>
<td>Clinical judgment using diagnostic algorithms</td>
<td>52.2% of toddlers were diagnosed with ASD by DSM-5</td>
<td>Single author review of evaluations based on DSM-IV</td>
</tr>
<tr>
<td>Gibbs et al\textsuperscript{13} / 2012</td>
<td>132 youth</td>
<td>Referred to tertiary autism clinic for initial evaluation</td>
<td>ADOS, ADI-R</td>
<td>76.5% of participants were diagnosed with ASD by DSM-5</td>
<td>ADOS and ADI-R are DSM-IV based tools</td>
</tr>
<tr>
<td>Taheri and Perry\textsuperscript{33} / 2012</td>
<td>131 children aged 2-12 years</td>
<td>Retrospective file review</td>
<td>CARS, DSM-IV checklist</td>
<td>62.6% of total sample met diagnosis of ASD by DSM-5</td>
<td>No Asperger's disorder patients, DSM-5 criteria were evaluated by checklist</td>
</tr>
<tr>
<td>Huerta et al\textsuperscript{24} / 2012</td>
<td>5,143 subjects, 4453 had PDD</td>
<td>Data sets from family genetics study, university and autism center databases</td>
<td>ADI-R and ADOS matched to DSM-IV and DSM-5 criteria. Included parent report and/or direct observation</td>
<td>DSM-5 identified 91% of children with PDD diagnoses. Overall specificity was low, (0.53) but improved over DSM-IV</td>
<td>More severe clinical sample, retrospective data analysis</td>
</tr>
</tbody>
</table>

*3Di = Developmental, Dimensional, and Diagnostic Interview; ADI-R = Autism Diagnostic Interview; ADOS = Autism Diagnostic Observation Schedule; ASD = autism spectrum disorder; CARS = Childhood Autism Rating Scale; NOS = not otherwise specified; PDD = pervasive developmental disorder; Revised; SCQ = Social Communication Questionnaire; SRS = Social Responsiveness Scale.*
The view

- All of the studies have been retrospective in nature
- Larger samples sizes support DSM-5
- Only Heurta, et al., matched a sample to both DSM-IV-TR and DSM-5
- Prospective field trials are the only answer
DSM-5 criteria for Social communication disorder

- Difficulties in verbal and nonverbal communication
  - Deficits in use of communication for social purposes
  - Inability to match context or needs of the listener
  - Poor ability to follow rules of conversation and storytelling
  - Poor understanding of nuances, humor, sarcasm

- Causes dysfunction in social, academic, or occupational settings

- Early developmental onset

- Not explained by autism or ID
## Proposed Modifications to DSM-5

<table>
<thead>
<tr>
<th>DSM-5 draft criteria</th>
<th>Suggested changes</th>
</tr>
</thead>
<tbody>
<tr>
<td>Must meet all 3 criteria for social communication and interaction</td>
<td>May meet 2 of 3 criteria for social communication and interaction</td>
</tr>
<tr>
<td>“deficits in nonverbal communication”</td>
<td>“deficits in verbal and/or nonverbal communication”</td>
</tr>
<tr>
<td>“Excessive adherence to routines and rituals”</td>
<td>“excessive adherence to routines and/or rituals”</td>
</tr>
<tr>
<td>“symptoms must be present in early childhood”</td>
<td>“symptoms present in childhood”</td>
</tr>
</tbody>
</table>

Increased sensitivity with modification

<table>
<thead>
<tr>
<th>Diagnosis</th>
<th>DSM-5 criteria</th>
<th>Modified criteria</th>
</tr>
</thead>
<tbody>
<tr>
<td>Autism Spectrum Disorders, FSIQ &gt; 50</td>
<td>46%, (n=26)</td>
<td>96%</td>
</tr>
<tr>
<td>High functioning autism (FSIQ &gt; 70)</td>
<td>73%, (N=11)</td>
<td>100%</td>
</tr>
<tr>
<td>Asperger’s disorder</td>
<td>0%, (n=11)</td>
<td>91%</td>
</tr>
</tbody>
</table>

Proposed Modifications to DSM-5

- Relaxing DSM-5 criteria with one less SCI or RRB criterion may increase sensitivity by 11% to 12%
  - Useful impact on those with Asperger’s disorder
  - Patients with limited early childhood history
  - Supported by Huerta, et al. 2012

- “symptoms must be present in early childhood” should be modified to allow children or adults without accurate early history can be diagnosed
Proposed Modifications to DSM-5

- Relax SCI criteria to allow 2 of 3 criteria to be used
- AND Lower RRB threshold from 2 to 1
  - McPartland JAACAP, 2012;51(4)368-383.

<table>
<thead>
<tr>
<th></th>
<th>DSM-5</th>
<th>Relaxed SCI criteria</th>
<th>AND Lowered RRB threshold</th>
</tr>
</thead>
<tbody>
<tr>
<td>sensitivity</td>
<td>0.61</td>
<td>0.75</td>
<td>0.91</td>
</tr>
<tr>
<td>specificity</td>
<td>0.95</td>
<td>0.85</td>
<td>0.75</td>
</tr>
</tbody>
</table>
DSM-5 Workgroup response

- Expanded examples of criteria
- Symptoms may be counted if observed or historical
Finding the right balance
Pros of DSM-5 and autism

- Improved specificity
  - Stability
  - Clarity

- Dimensional approach
  - Core issue of autism = social communication deficits
    - Genetic influence
    - Normal distribution in population

- Measures of severity
Pros of DSM-5 and autism

- Increased homogeneity
  - Longitudinal research
  - Endophenotypes
  - Genotypes

- Reflects evolution of research

- DSM-IV symptoms map to DSM-5
Cons of DSM-5 and autism

- Decreased sensitivity
  - Those with high functioning autism and Asperger’s disorder may not meet criteria for DSM-5
  - Social communication disorder

- Effect on service eligibility

- Compatibility with prior research subtypes

- Fit with ICD-11
Concerns with social communication disorder

- Is it valid?
  - What does it mean?
  - How to operationalize the diagnosis?

- Will it cover services?

- What about those with more RRB than SCI?
NIMH Research Domain Criteria, RDoC

- DSM-5 vs. NIMH
  - DSM-5 is a consensus document
    - Diagnosis is based on clusters of clinical symptoms

- RDoC looks for biomarkers
  - Mental disorders are biologic disorders
  - Genetics, imaging, cognitive science

- RDoc will provide framework to collect data for new classification system
Future Directions

♦ Prospective field studies
  ♦ Phase II testing
  ♦ Compare DSM-5 vs. DSM-IV and DSM-5 vs. expert clinical diagnosis

♦ Define social communication disorder
DSM-5 field trial: study design

- Estimated reliability and validity of diagnoses
- Informed DSM-5 decision-making process
- Test-retest design
- Naturalistic design
- DSM-IV field trials
  - best case scenarios

DSM-5 test-retest reliability for autism

- Intraclass kappa shows very good, (0.69) reliability
- Slight decrease in prevalence of autism spectrum rates
  - Due to diagnosis of social communication disorder
Future Directions

- Search domains of social communication and repetitive, restricted behaviors
  - Isolate endophenotypes
  - Find biologic and genetic markers

- Suitable instruments to measure social communicative function
The nature of autism

❖ Lumpers
  ❖ Population distribution of social function
  ❖ Rates in twins and siblings

❖ Splitters
  ❖ Latent symptoms
  ❖ Brain development

❖ Hybrid model
  ❖ Supported by recent validity studies by Mandy et al. and Frazier et al.
FRAP!
Frenetic random acts of play

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