The neural basis of the phonological deficit in developmental dyslexia

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Backward Reading

- 30% of school leavers cannot read better than an eleven year old
- 75% of young offenders in the UK are illiterate
- Whether or not we call these ‘dyslexic’, the same teaching methods will help any poor reader
- Developmental dyslexia: reading significantly below that expected from age and intelligence, despite good health, teaching and family support
- Dyslexia is more than a reading problem- the brain is different - a **neurological syndrome**
- Genetic, immunological, nutritional
- Slower processing, inattention, poor sequencing in all domains, poor short term memory, incoordination
- ? caused by impaired development of ‘magnocellular’ timing neurones
Many Dyslexics Experience Visual Confusions When Reading

- Letters reverse, blur, move across, in and out of page, change places, superimpose, distort, change colour
- At age 8 still getting letters and figures backwards & in wrong order
- Spelling remains very bad even into adulthood
- Poor visual attention
- Short visual memory
- Loses place easily
- ‘Careless’ mistakes
- Dreadful handwriting
- Mixes up capitals with lower case letters
- Glare - visual discomfort

“We were shattered to learn he was dyslexic. We thought he was learning Bulgarian.”
Reading is primarily a visual process
10% are large magnocellular cells (100x p- cells in area) - for timing visual events: fast responses, high contrast sensitivity, motion, flicker, control attention & eye movements

Most retinal ganglion cells are parvocellular (small): for colour, fine detail, high contrast (less vulnerable)
Visual magnocellular system dominates dorsal visuomotor pathway - directs visual attention & eye movements.
The visual magnocellular system is impaired in poor readers

- 30% smaller LGN magnocells *post mortem*
- Reduced and delayed evoked brain waves
- Unstable eye control
- Reduced visual motion sensitivity
- Lower sensitivity to flicker
- Lower sensitivity to low spatial, high temporal frequency contrast gratings
- Reduced activation of cortical visual motion areas (FMRI)
- Lower stereoacuity
- Poor visual sequential attention - slower visual search

- In the last 10 years 90% of new research papers have supported the magnocellular deficit theory
- But it is still controversial:
  - Definition of magno- system - only completely separate in periphery
  - Selectivity of stimuli
  - Mild deficit requires highly sensitive test to reveal it
Abnormal magnocells in dyslexic visual relay - LGN
DTI - Fewer large axons in left angular gyrus in dyslexics. Successful remediation increases their size.
Delayed Brain Potentials Evoked by Moving Visual Stimulus (10 hz component)
The visual magnocellular system stabilises the eyes to avoid visual wobble.

- Unwanted image motion, ‘retinal slip’
- Locks eyes on target
- Detected by M-system
- Feedback to eye muscle control system
- Visual stability
- Identify letter order
- Orthographic skill
- Phonological skill

Orthographic skill and Phonological skill are related.
Weak magnocellular system causes unstable vision - oscillopsia

“The letters go all blurry”
“The letters move over each other, so I can’t tell which is which”
“The letters seem to float all over the page”
“The letters move in and out of the page”
“The letters split and go double”
“The c moved over the r, so it looked like another c”
“The p joined up with the c”
“d’s and b’s sort of get the wrong way round”
“The page goes all glary and hurts my eyes”
“I keep on losing my place”
Interventions that improve magno-function and eye control often improve reading

In half of all dyslexic children, simply reading through deep blue or yellow filters can rebalance input to the visual M-system. Reading improves by av. 6 months in 3 months.

In older children, fixation exercises can stabilise binocular fixation and greatly improve reading.
Auditory/phonological errors

- Whisk/wisp, deaf/death, effect/affect
- Ears (eves) dropping; sort (saute) of potatoes
- They came down on the food like a flock of vouchers
- The gossip spread like wild flies
- Concord is so noisy – too many decimals
- After a time I syphoned (deciphered) it
- Its just like flogging a brick wall
- I torned and tussed all night
- This strike will bring the country to a Stanstead
- Dry as a door nail; dead as rust; a soup opera
- Endearment (endowment) policy; customs and exile
The visual/orthographic and auditory/phonological pathways
Auditory m-cells?

2nd and 3rd formants ascend in frequency for ‘b’; but descend for ‘d’.

Auditory m-cell impairment may reduce sensitivity to these changes in sound frequency.
Developmental Dyslexics are less sensitive to changes in sound frequency and intensity.

- Slow frequency changes in speech are tracked in *real time* by large magnocells in the auditory system.

\[ \text{Detection Threshold (Mod. Index)} \]

<table>
<thead>
<tr>
<th></th>
<th>2 Hz FM</th>
<th>40 Hz FM</th>
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<tbody>
<tr>
<td>Controls</td>
<td>N=23</td>
<td></td>
</tr>
<tr>
<td>Dyslexics</td>
<td>N=21</td>
<td></td>
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- **2 Hz FM**
  - *p* < 0.001

- **40 Hz FM**
  - *p* = 0.027

- **500 Hz**
  - Pure tone

- **240 Hz FM**
  - *p* = 0.360, N.S.

Witton, Talcott, Hansen, Richardson, Griffiths, Rees, Stein & Green, 1998
Nonsense Word Reading
Nonwords

‘tegwop’, ‘blint’, ‘plomt’, ‘peltip’, ‘visht’ can only be read if letters can be translated into their sounds quickly and accurately; hence they test phonological ability.
2 Hz FM Thresholds in 10 year-olds (n = 54)

$r = 0.54, p < 0.001$
Spearman rank-order Correlation Coefficient

240 Hz FM
2 Hz FM

p < 0.05
p < 0.01
p < 0.001

Reading
Spelling
Nonwords
Exception words
Spoonerisms
Pseudohomophones

Spearman rank-order Correlation Coefficient
Norwegian 10 year olds

- Low sensitivity to auditory frequency change is found not only in English dyslexics, but also in more regular Norwegian.
Auditory brainstem responses more sensitive (AM)
Psychophysical sensitivity to 20 Hz FM was normal in 10 adult dyslexics. But their mismatch negativity was significantly decreased compared to normal readers.
Auditory Brainstem Responses (speech in noise)
Is there an Auditory Magnocellular System?

• Large neurones in the auditory brainstem (DCN, ICX, mMGN) signal start, finish, changes in amplitude, frequency and phase of sounds.

• FM and AM sensitivity in both good and poor readers determines their phonological ability.

• Poor readers have:

  reduced pure tone discrimination,
  reduced AM & FM sensitivity,
  smaller brainstem auditory evoked potentials (ABR)
  smaller magnocellular neurones in MGN.

• Thus poor readers may have impaired development of their auditory magnocells
Auditory and visual magnocellular sensitivity determines over half of variability in children’s phonological ability. Thus the most important determinant of reading ability appears to be low level magnocellular sensitivity. Encouraging because this can be improved by training.
Sensorimotor Basis of Dyslexia

Low **visual** magnocellular sensitivity - orthographic weakness

Low **auditory** magnocellular sensitivity - phonological problems

Lower **motor** magnocellular sensitivity – in coordination, poor balance

Lower **kinaesthetic** magnocellular sensitivity
Magnocellular Neurones

- A system of large neurones specialised for temporal processing – tracking changes in light, sound, position etc. for direction of attention
- Large, fast conduction, fast transmission, high anisotropy
- All express same surface antigen, CAT 301
- Found throughout the whole brain: visual, auditory, skin, muscle proprioceptors, cerebral cortex, hippocampus, cerebellum, brainstem

- Very vulnerable. Impaired m-cell development has been found in prematurity, foetal alcohol syndrome, developmental dyslexia, dyspraxia, dysphasia, ADHD, ASD, Williams syndrome, schizophrenia, depression
- M-cell high dynamic sensitivity requires high membrane flexibility provided by local environment of essential fatty acids, particularly omega 3s, found in fish oils
- Hence vulnerable to omega-3 deficiency
What causes this general magnocellular impairment?

Genetic
Immune System
Nutrition
Genetic linkage/association

- Are particular chromosomal markers/sites associated with poor reading?
- Analyse the DNA of father, mother and their dyslexic and normally reading children
- >400 Oxford; 100 Boulder (US) families
- EU consortium; 1000 families, 2000 cases, 2000 controls, 50,000 markers per case
C6 KIAA 0319 controls neuronal migration during early brain development in utero. Downregulation in dyslexics may explain ectopias and other mismigrations of magnocellular neurones.
Abnormal magnocells in dyslexic visual relay - LGN
2nd trimester ectopias in dyslexic brain specifically affect dorsal visuomotor stream
Left hemisphere language areas that activate less in dyslexics.
KIAA 0319 is strongly expressed in visual magnocellular pathway.
C6p KIAA mutation?

- KIAA underexpression disrupts neuronal migration early in brain development
- Controls expression of cell surface recognition molecules
- Fail to respond to developmental signals, enable successful contact with other magnocells
- Could explain why the development of magnocellular neurones is impaired in dyslexia
What causes phonological problems?

- Impaired visual magnocellular function
- Impaired auditory magnocells
- Due to genetic vulnerability
- This knowledge is exciting because these weaknesses can be remedied