Learning & memory in 22q11.2 deletion syndrome

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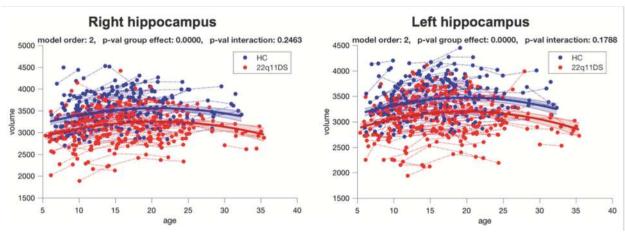


Learning & memory

Ability to acquire, store and recall information over time

Hippocampus is a key structure in the brain

Reduction of the volume have been consistently demonstrated in 22q11.2DS







Hippocampus

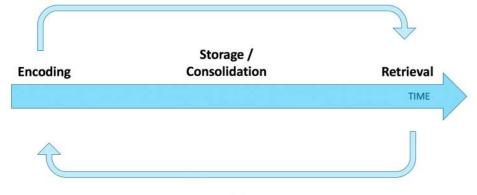
From Mancini et al. 2019, Mol. Psych.

Learning & memory in 22q11.2DS

Numerous previous studies on memory have shown verbal > non verbal

> Few have focused on the different steps leading to memory impairments

Acquisition? Storage? Recall?



Re-consolidation

Memory is tested with paradigms including immediate and differed recalls (30 minutes)

- Consolidation of memory happens over longer delays (days, weeks, months)
 - What happens after 30 minutes?

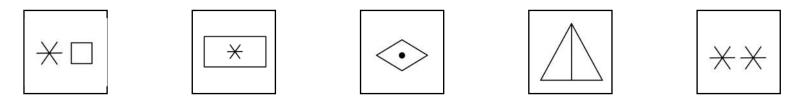
Task design

Inspired by Rey's Auditory Verbal Learning Test (RAVLT, 1958)

Verbal information = 15 common French words

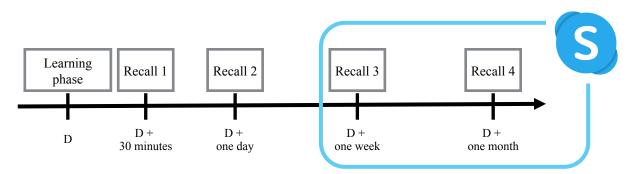
Table, cat, moon, car, ...

Non-verbal information = 15 drawings made out of 1 or 2 basic geometrical forms

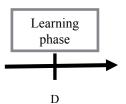


Learning criterion: 80% (12 items) or max. 6 presentations

Adapted recall design:







N= 135 (78 with 22q11.2DS) Age range: 8-25

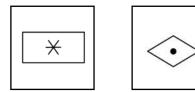
Information acquisition

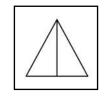
VERBAL

Table, cat, moon, car, ...

Comparable trajectories of learning over time Similar performances from trial 1 Same amount of trials necessary to reach criterion

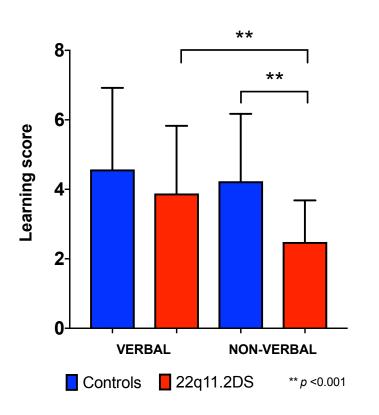
NON-VERBAL





Lower performances from trial 1 in 22q11.2DS Slower increase over time in 22q11.2DS More trials needed to reach criterion in 22q11DS

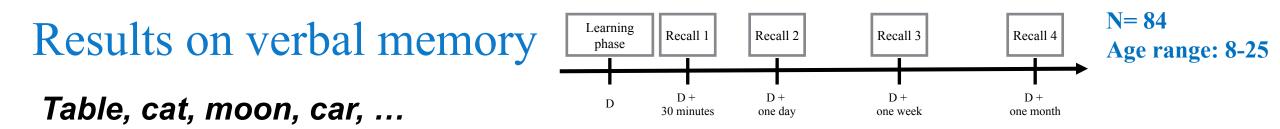
From Maeder et al. Under review

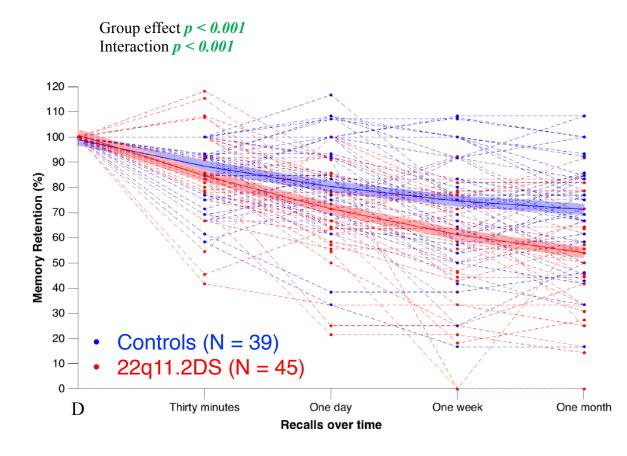


Highest. Nb. Items recalled

Learning Score =

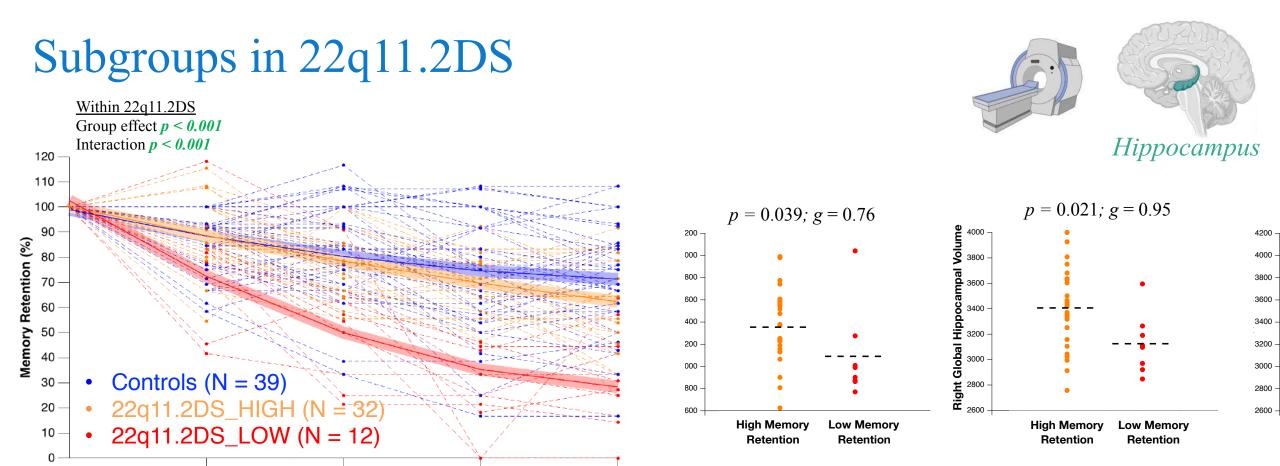
Nb. Trials necessary to reach criterion





Significant difference in shape between groups Significantly lower performance after delays of 30 minutes Evidence for accelerated long-term forgetting (ALF) (Shape with IQ as a covariate: *p* <0.001)

From Maeder et al. 2019, Child Neuropsychol.



Characteristics of the 22q11.2DS LOW memory group

One month

Thirty minutes

One day

Recalls over time

One week

2q11.2DS HIGH (N = 32)

Lower IQ Higher rates of psychotic symptoms Reduction of hippocampal volume

From Maeder et al. 2019, Child Neuropsychol.

• $22q11.2DS_LOW (N = 12)$

Conclusions

Non-verbal encoding is selectively affected in the 22q11.2DS group

> Start lower and less improvement with repetition

After 30 minutes, verbal memory performance are similar between both groups BUT after longer delay, a steeper loss of information is observed

> Evidence for long-term accelerated forgetting (ALF)

Sub-group of patients with lower verbal memory consolidation

Tips to improve learning & memory

Use the verbal channel

 \checkmark Give verbal instruction

✓ Repeat the information out loud when you learn Spilt the information in small parts

- \checkmark Learn in small chunks
- ✓ Distribute it on different days

Repetitions

- \checkmark Repeat the information several times
- \checkmark Go back to information previously learned



How does attention contribute to learning & memory?

Clinical trial in Geneva (CH)

In 22q11.2DS, attention deficit disorder (ADD) is very frequent (40%) Inattentive manifestation predominant (70%) => not always recognized

How does ADD influence learning and memory?

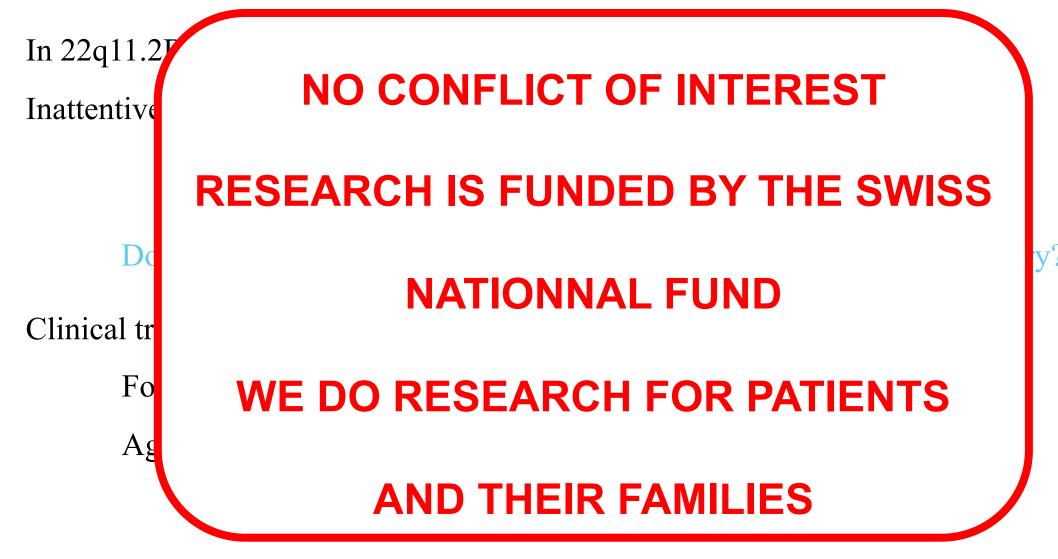
Does ADD medication (methylphenidate) improve learning and memory?

Clinical trial in Geneva:

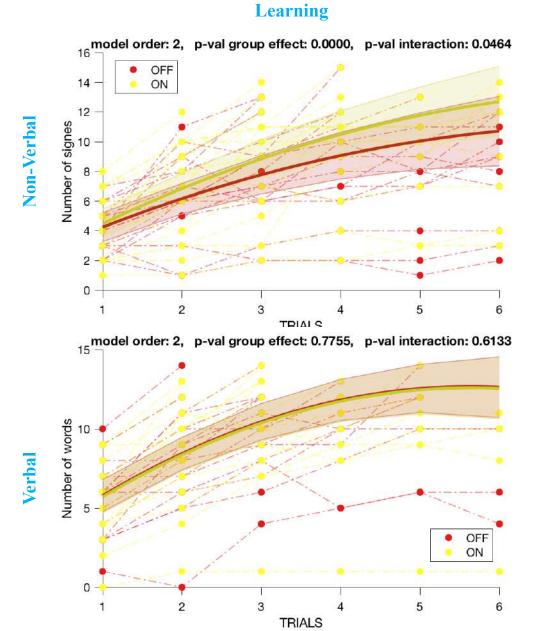
For people with 22q11.2DS

Age range 8-25 years old

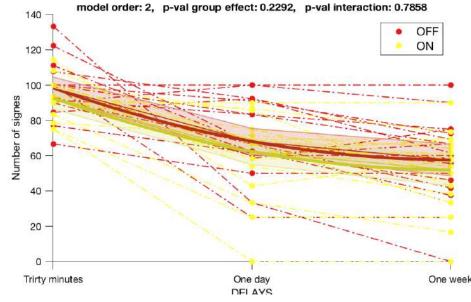
Clinical trial in Geneva (CH)



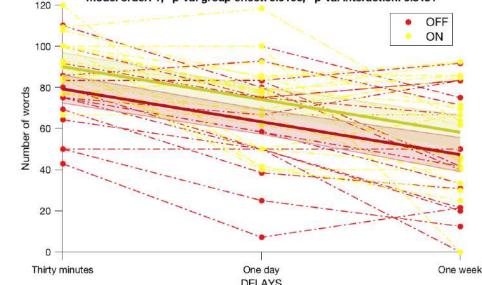
Preliminary results on learning & memory



Memory retention over time



model order: 1, p-val group effect: 0.0103, p-val interaction: 0.5151



More efficient learning of visual stimuli with medication, but similar decline over time

For **verbal stimuli**, similar learning with medication, but remember more information over time

From Maeder et al. In prep



















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