

Glucose on the Ageing Mind

Healthy Brain Ageing Symposium

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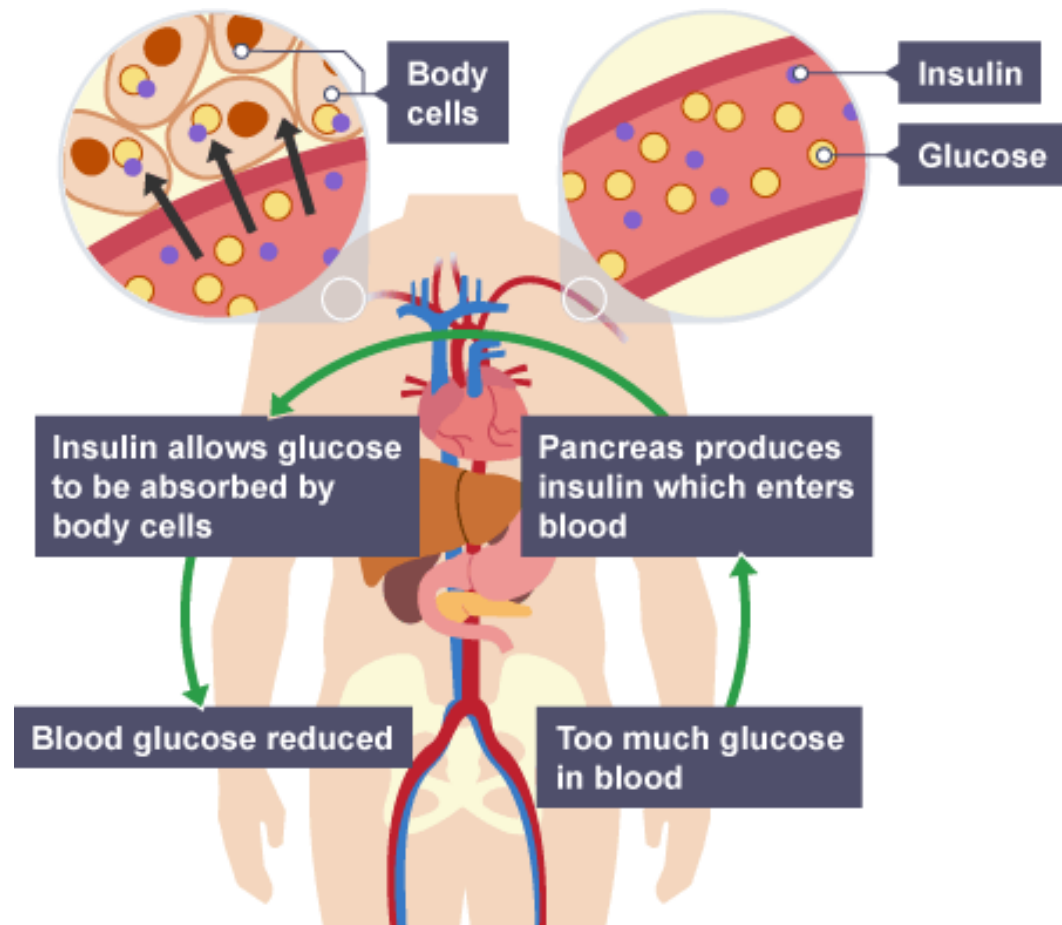
Brain Ageing

- Normal brain ageing can be affected by a wide range of factors
 - Some are fixed
 - Disease (e.g. Dementia)
 - Genetics
 - Some are modifiable
 - Personal history and environment (e.g. learning)
 - Lifestyle (e.g. physical activity, diet)

Blood glucose is the amount of glucose present in the blood plasma.

- Glucose metabolism depends on complex interaction of many factors
 - Food
 - Medication
 - Physical Activity
 - Environment
 - Genetics

Blood glucose



Blood glucose is the amount of glucose present in the blood plasma.

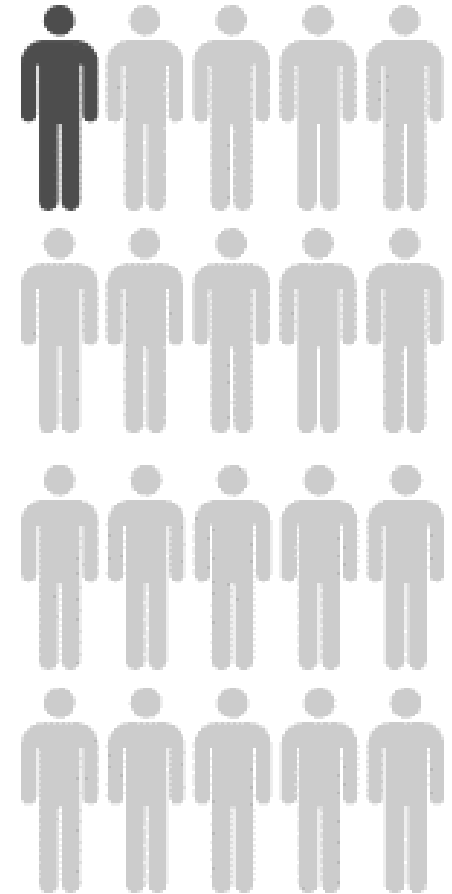
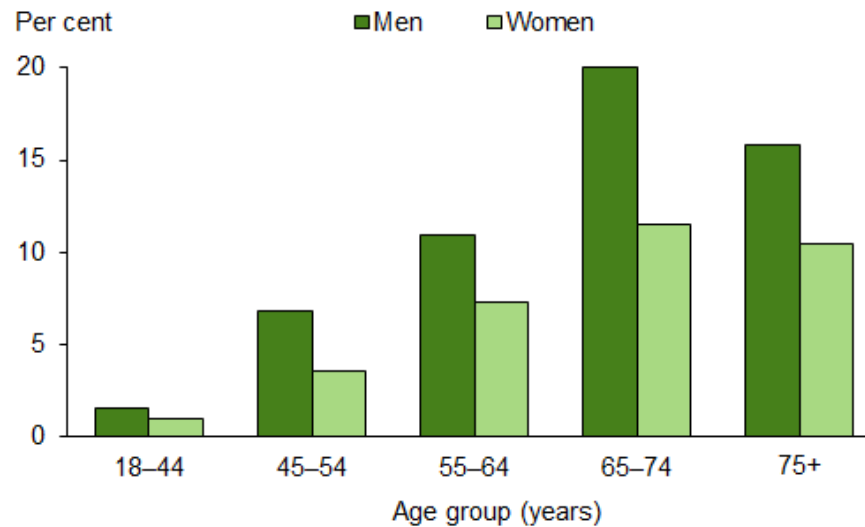
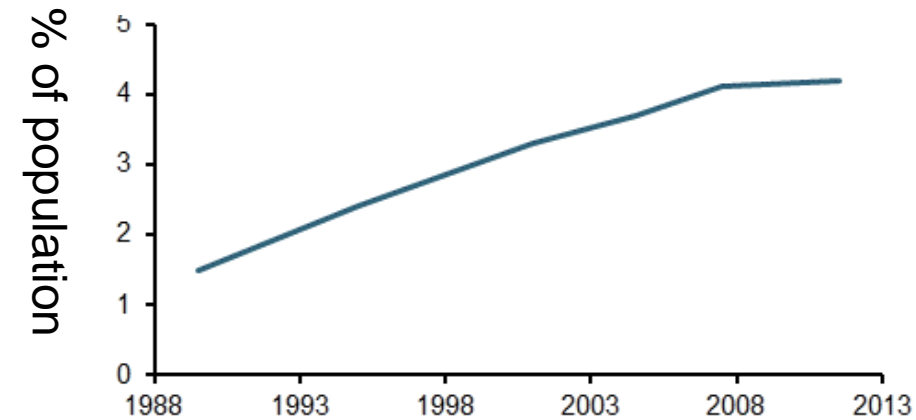
The brain needs glucose

- Energy source
- Needed to make neurotransmitters



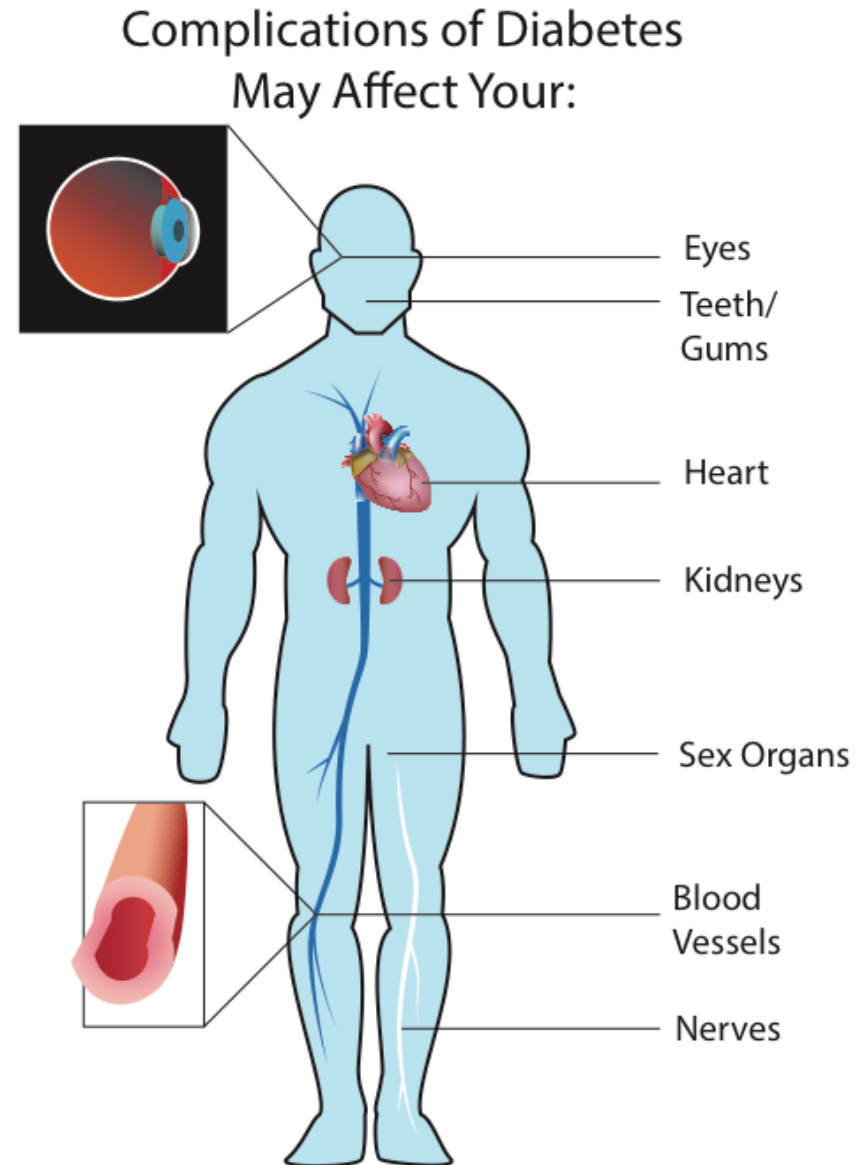
Too much of a good thing...

Diabetes in Australia



Blood glucose and diabetes

- Type 2 diabetes brain health impacts:
 - Depression
 - Dementia
 - **Cognitive decline**
- Even high blood glucose in the normal range can be harmful



Higher blood
glucose and
diabetes

Glucose → Cognition

- Effects may differ depending on...

PATH, cohort with blood glucose and cognitive test data: n=429 54% female

- Age

(Sims Wright, Levy et al. 2015; Hall, Gonder-Frederick et al. 1989)

40s cohort

3 waves – 45-60

- Diabetic status

(Meikle, Riby et al. 2004)

Full blood glucose range:
Normal / IFG/ diabetes

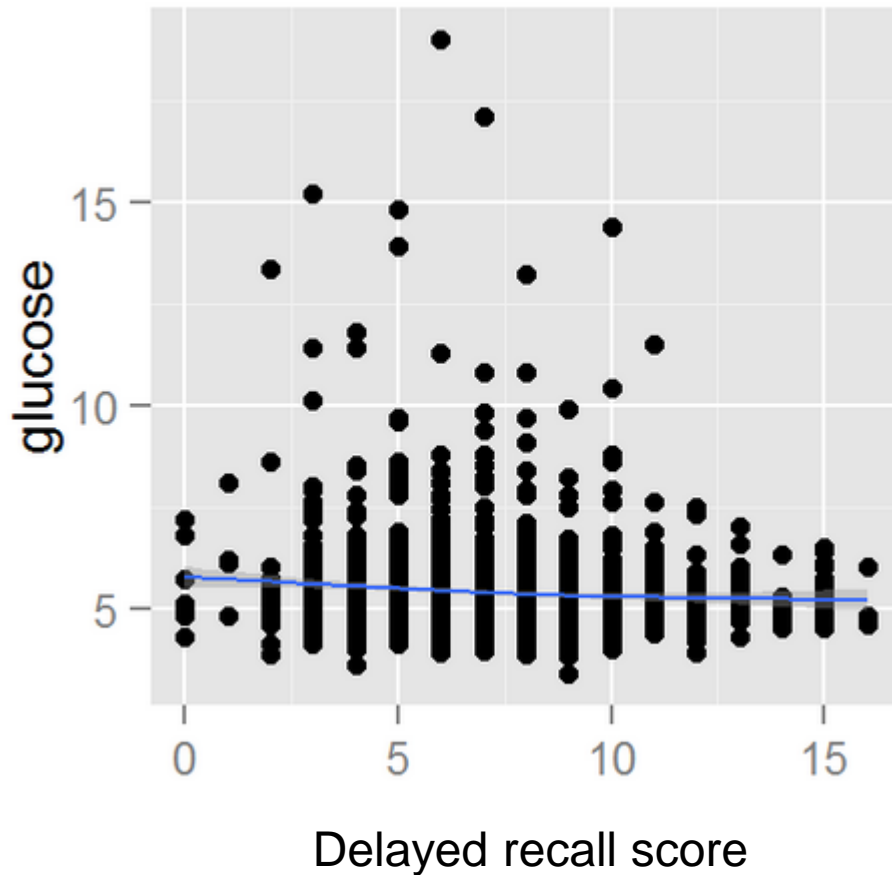
- Specific task

(Backeström, Eriksson et al. 2015; Foster, Lidder et al. 1998; Feinkohl, Keller et al. 2015)

Three cognitive tasks
from different domains

An investigative journey...

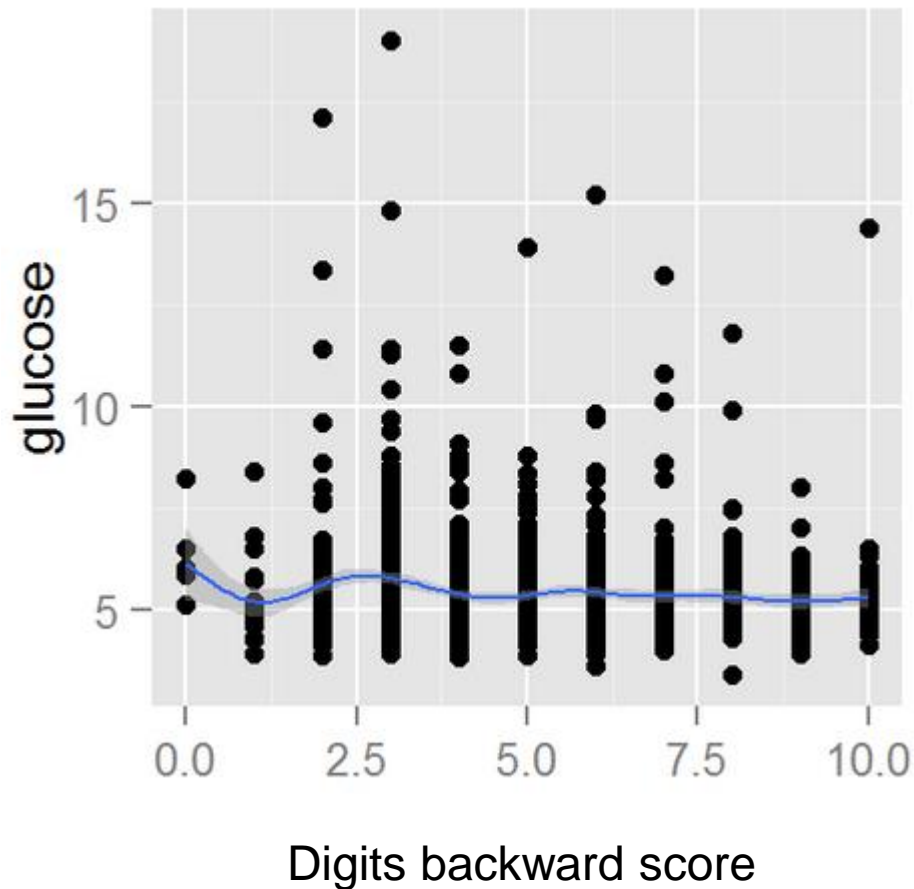
Delayed recall



	Estimate	Sig.
Delayed recall	0.01	0.8

Words selected from the California Verbal Learning Test (Delis, Kramer et al. 1987)

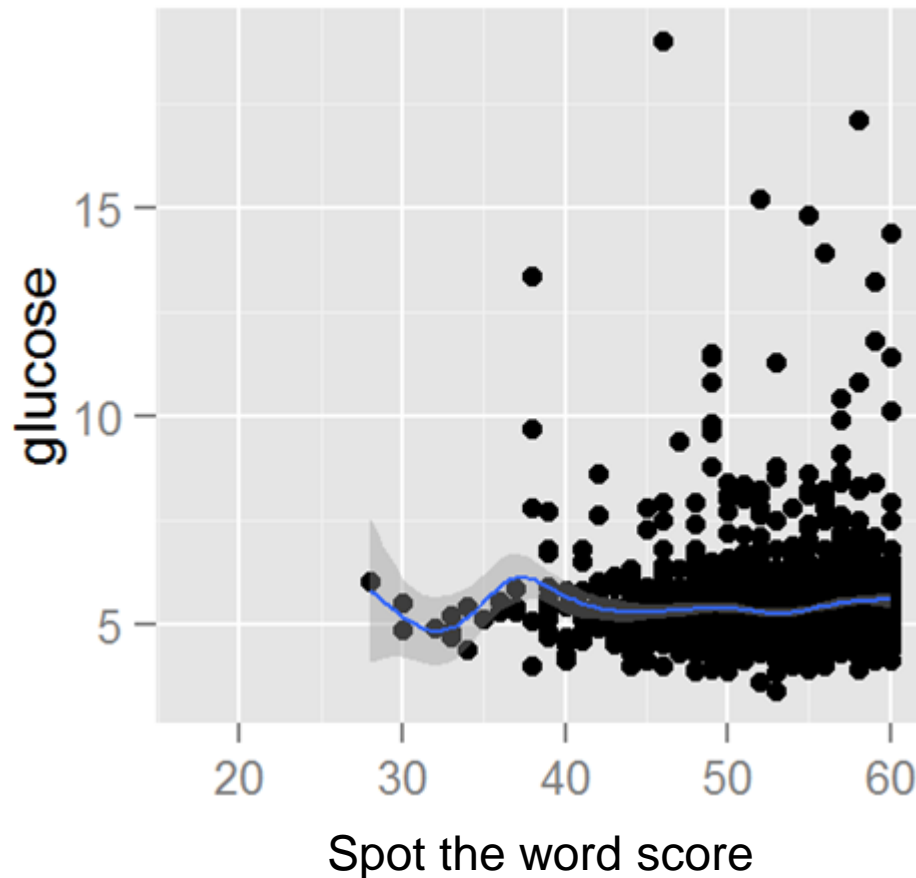
Digits backward



	Estimate	Sig.
Delayed recall	0.01	0.8
Digits backwards	-0.01	0.79

From the Wechsler Memory Scale (Wechsler 1945)

Spot the word



(Baddeley, Emslie et al. 1992)

Association between blood glucose and cognition differs **by cognitive task.**

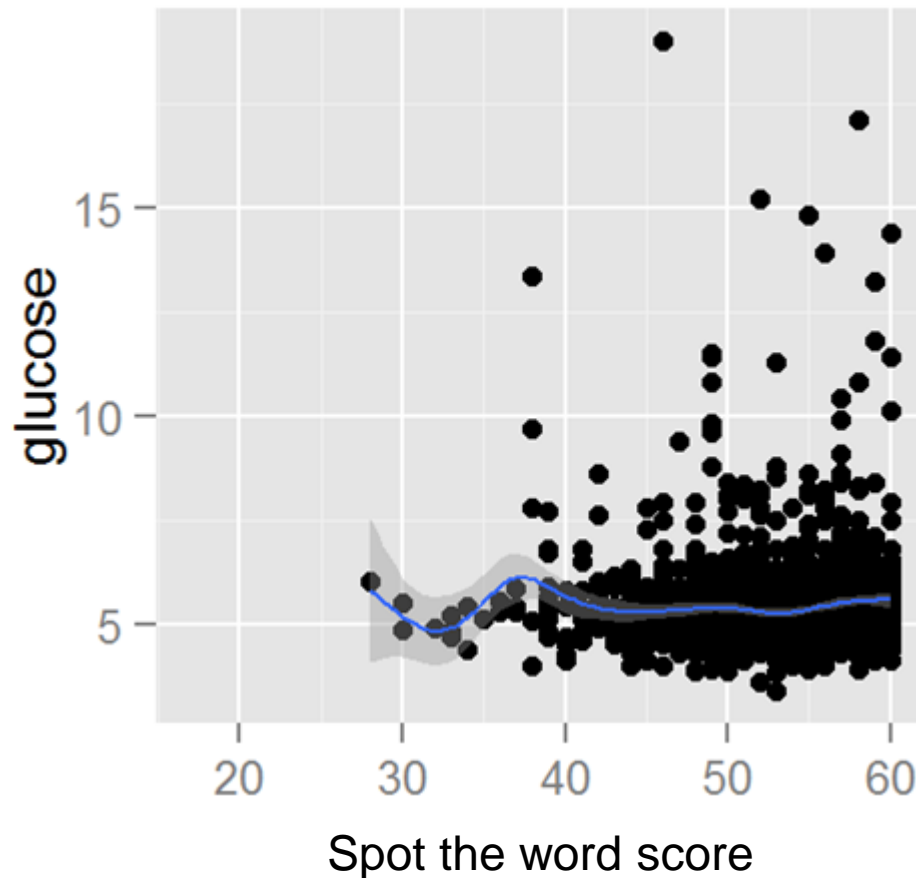
	Estimate	Sig.
Delayed recall	0.01	0.8
Digits backwards	-0.01	0.79
Spot the word	0.23	0.01

For each increase in blood glucose of approximately 4mmol/L, participants correctly identified one additional non-word.

Questions raised

- Why no decrease in performance any cognitive task?
 - Tasks too easy
 - Inappropriate choice
 - Sample too young
- Why an *increase* in performance in Spot-The-Word?
 - Task relies on verbal IQ (which increases with age)
 - Short term glucose facilitation
- Why *just* spot the word?
 - Glucose facilitation greatest for **verbal memory tasks**
 - Delayed recall and digits backwards more about short term memory
 - SDMT more about symbol correspondence

Spot the word

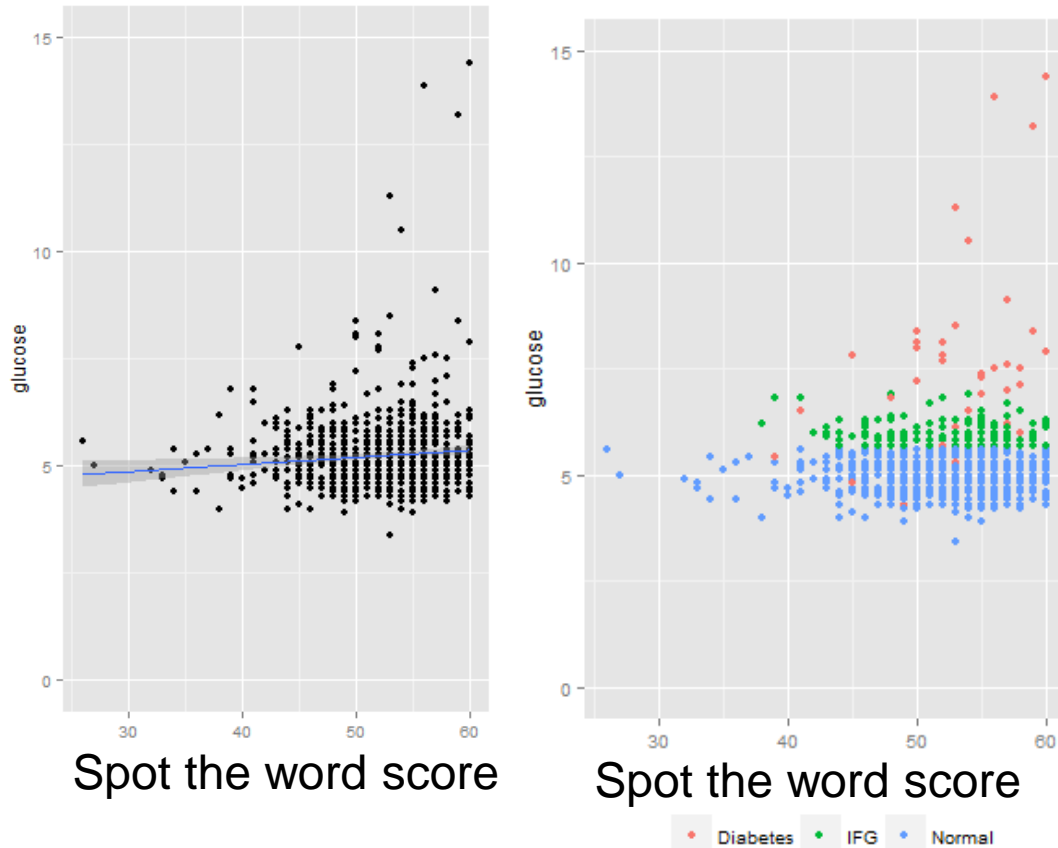


(Baddeley, Emslie et al. 1992)

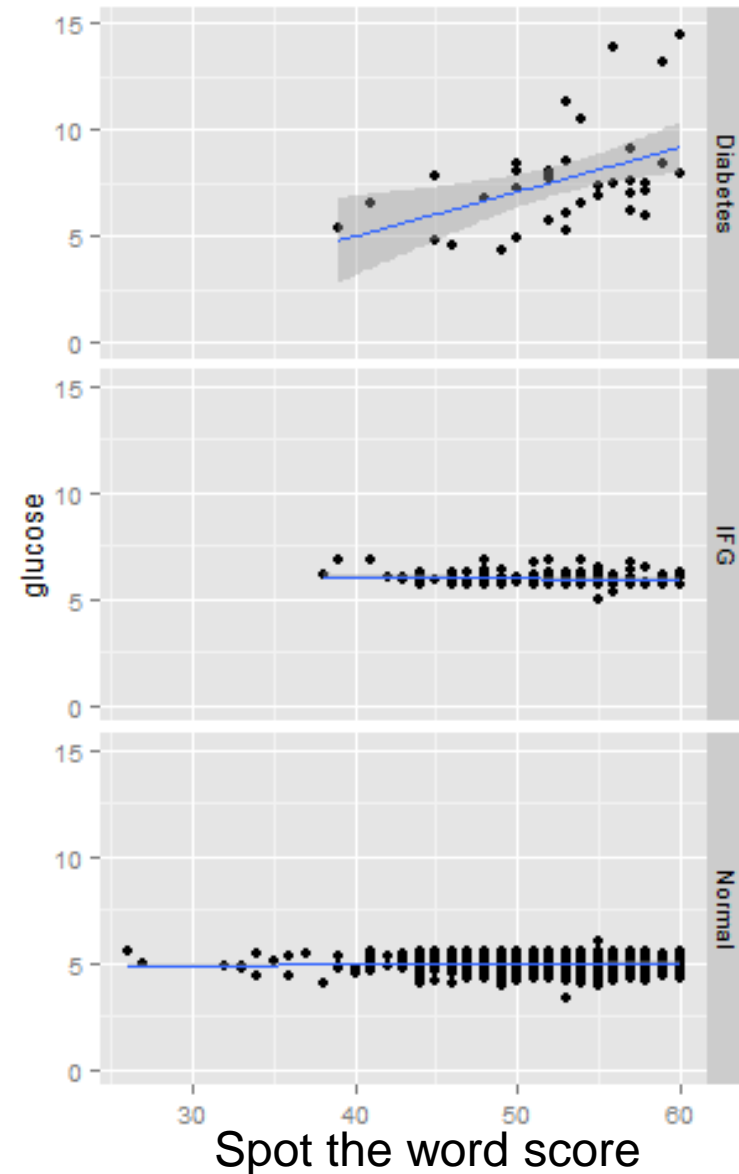
Association between blood glucose and cognition differs **by cognitive task.**

	Estimate	Sig.
SDMT	-0.13	0.45
Delayed recall	0.01	0.8
Digits backwards	-0.01	0.79
Spot the word	0.23	0.01

	Estimate	Sig.
Diabetics only	0.31	<0.01
IFG only	-0.03	0.93
Normal only	0.2	0.4

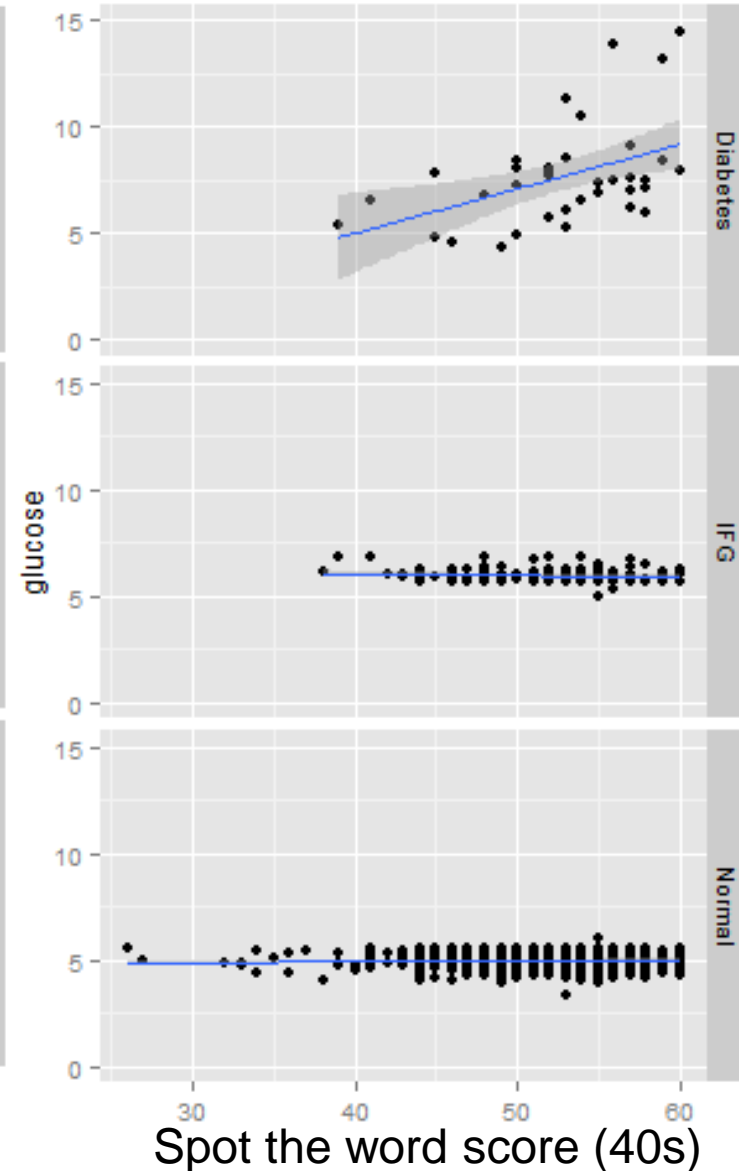
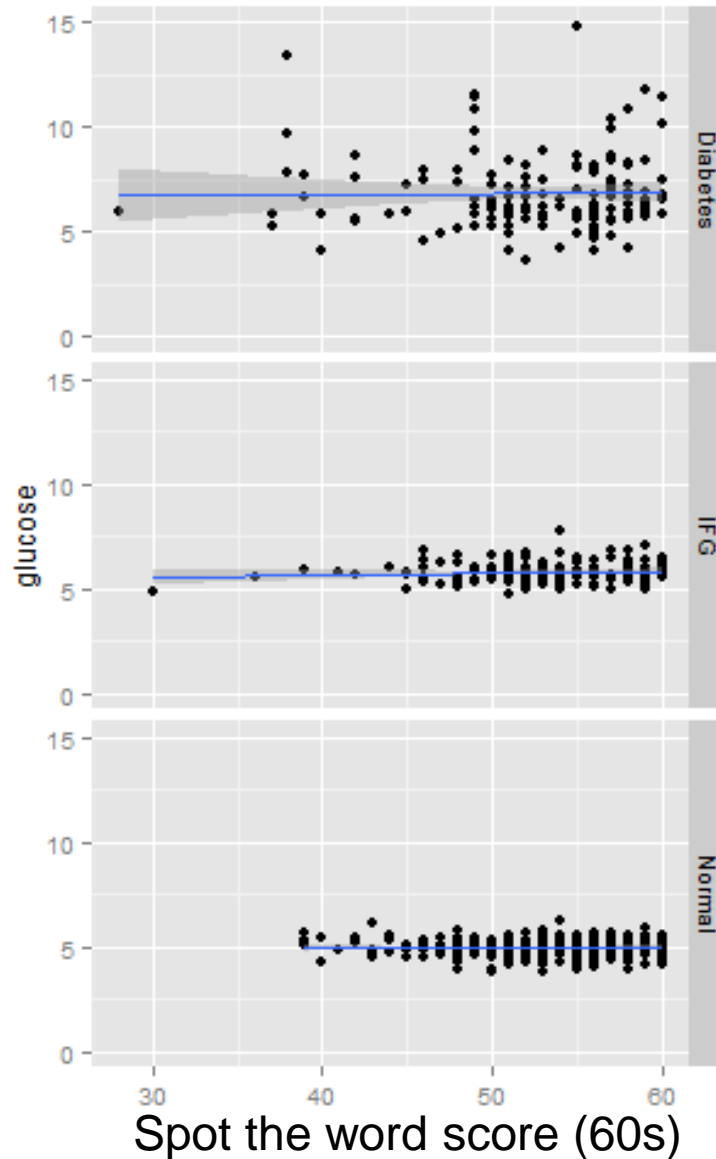


Association between blood glucose and cognition differs **by diabetic status.**



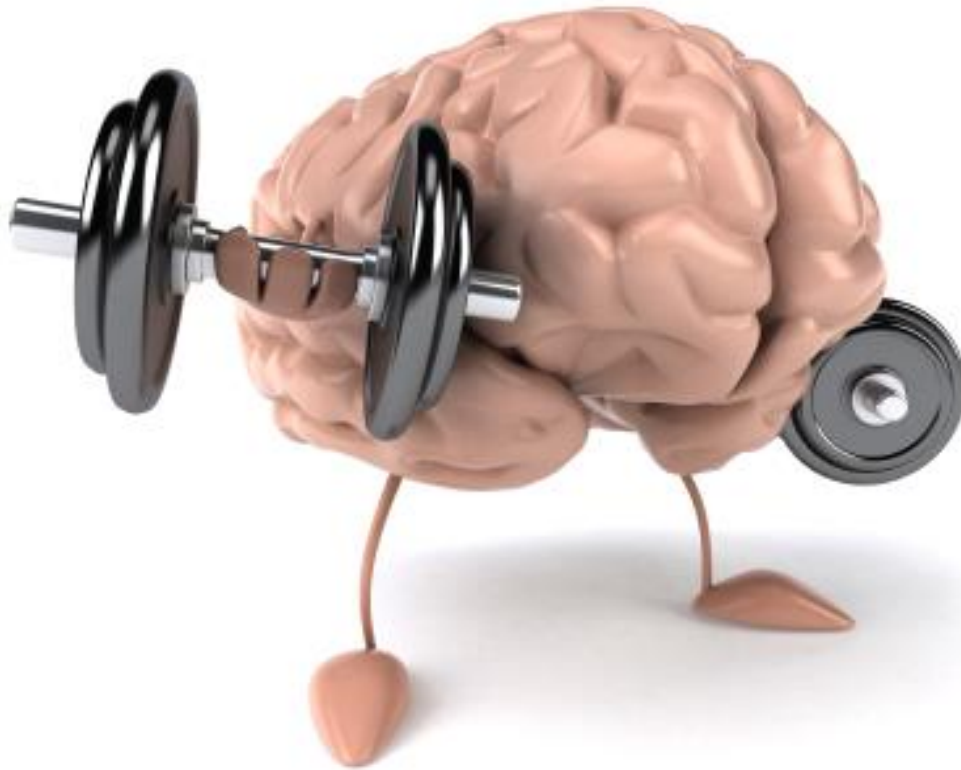
Association between blood glucose and cognition differs **by** **age.**

60s cohort $n=357$,
aged 60 at
recruitment, 47%
female



Outlook

- Growing evidence that chronically high blood glucose is associated with poor cognitive outcomes
- Blood glucose is associated with age
- Investigation needs to be sensitive to different effects across tasks, ages and diabetic categories
- **Potential for prevention of one of the contributing factors to cognitive decline in ageing**



Acknowledgements

Kaarin Anstey, Nicolas Cherbuin and the PATH team
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Awad, N., M. Gagnon and C. Messier (2004). "The relationship between impaired glucose tolerance, type 2 diabetes, and cognitive function." *Journal of Clinical and Experimental Neuropsychology* 26(8): 1044-1080. Backström, A., S. Eriksson, L.-G. Nilsson, T. Olsson and O. Rolandsson (2015). "Glucose but not insulin or insulin resistance is associated with memory performance in middle-aged non-diabetic women: a cross sectional study." *Diabetology and Metabolic Syndrome* 7. Baddeley, A., H. Emslie and I. Nimmo-Smith (1992). *The Spot-the-Word Test*. Bury St Edmunds, England, Thames Valley Test Company. Best, T., J. Bryan and N. Burns (2008). "An investigation of the effects of saccharides on the memory performance of middle-aged adults." *The journal of nutrition, health & aging* 12(9): 657-662. Cherbuin, N., P. Sachdev and K. J. Anstey (2012). "Higher normal fasting plasma glucose is associated with hippocampal atrophy: The PATH Study." *Neurology* 79(10): 1019-1026. Delis, D. C., J. H. Kramer, E. Kaplan and B. A. Ober (1987). *CVLT, California Verbal Learning Test: Adult Version: Manual*, Psychological Corporation. Feinkohl, I., M. Keller, C. M. Robertson, J. R. Morling, S. McLachlan, B. M. Frier, I. J. Deary, M. W. Strachan and J. F. Price (2015). "Cardiovascular risk factors and cognitive decline in older people with type 2 diabetes." *Diabetologia*: 1-9. Foster, J., P. Lidder and S. Sünram (1998). "Glucose and memory: fractionation of enhancement effects?" *Psychopharmacology* 137(3): 259-270. Goldberg, D., K. Bridges, P. Duncan-Jones and D. Grayson (1988). "Detecting anxiety and depression in general medical settings." *Bmj* 297(6653): 897-899. Gonder-Frederick, L., J. Hall, J. Vogt, D. Cox, J. Green and P. Gold (1987). "Memory enhancement in elderly humans: effects of glucose ingestion." *Physiology & behavior* 41(5): 503-504. Hall, J. L., L. Gonder-Frederick, W. Cheung, J. Silveira and P. Gold (1989). "Glucose enhancement of performance of memory tests in young and aged humans." *Neuropsychologia* 27(9): 1129-1138. Kumar, R., K. J. Anstey, N. Cherbuin, W. Wen and P. S. Sachdev (2008). "Association of type 2 diabetes with depression, brain atrophy, and reduced fine motor speed in a 60-to 64-year-old community sample." *The American Journal of Geriatric Psychiatry* 16(12): 989-998. Meikle, A., L. M. Riby and B. Stollery (2004). "The impact of glucose ingestion and gluco-regulatory control on cognitive performance: a comparison of younger and middle aged adults." *Human Psychopharmacology: Clinical and Experimental* 19(8): 523-535. Sims Wright, R., S.-A. T. Levy, L. I. Katzel, W. F. Rosenberger, Z. Manukyan, K. E. Whitfield and S. R. Waldstein (2015). "Fasting glucose and glucose tolerance as potential predictors of neurocognitive function among nondiabetic older adults." *Journal of clinical and experimental neuropsychology* 37(1): 49-60. Smith, A. (1982). *Symbol Digit Modalities Test (SDMT) Manual (Revised)*. Los Angeles, Western Psychological Services. Wechsler, D. (1945). *Wechsler Memory Scale*, The Psychological Corporation.

Thank you!

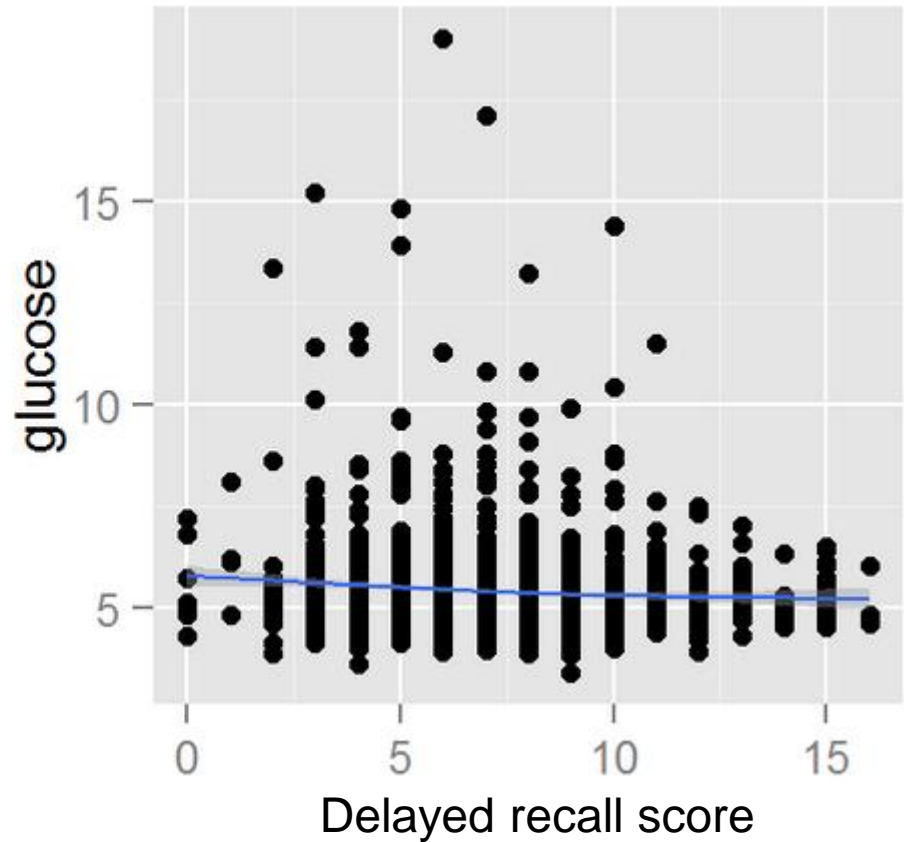
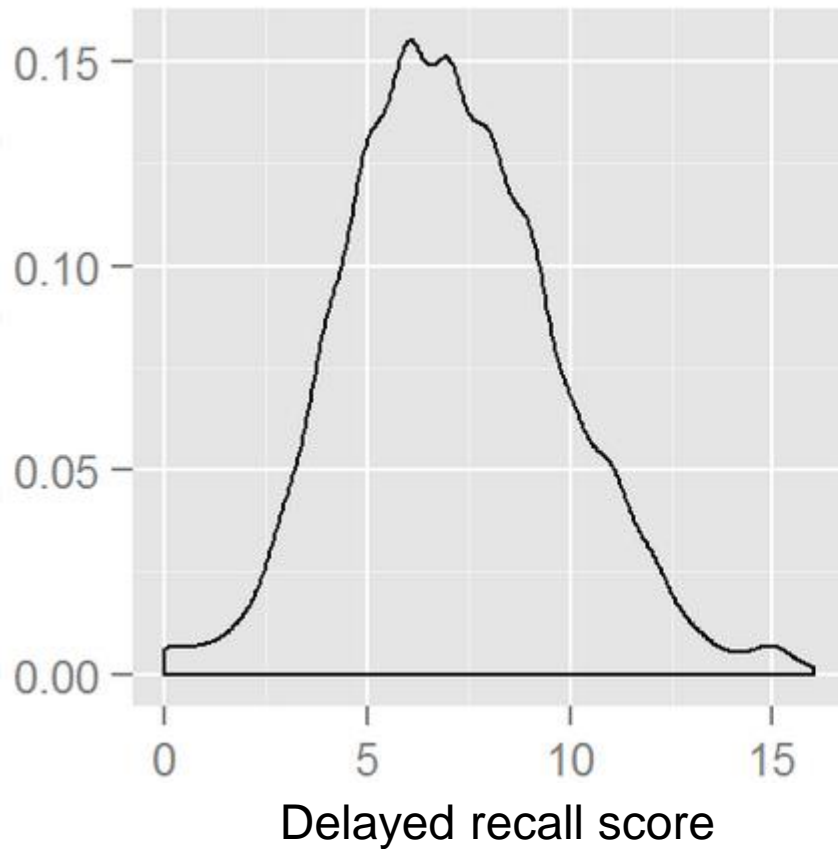
Statistics

	Estimate	Std. Error	t	Sig.	95% CI	
Spot the word	0.23	0.08	2.81	0.01	0.07	0.4
Delayed recall	0.01	0.05	0.26	0.8	-0.09	0.12
Digits backwards	-0.01	0.05	-0.26	0.79	-0.11	0.09

Spot the word

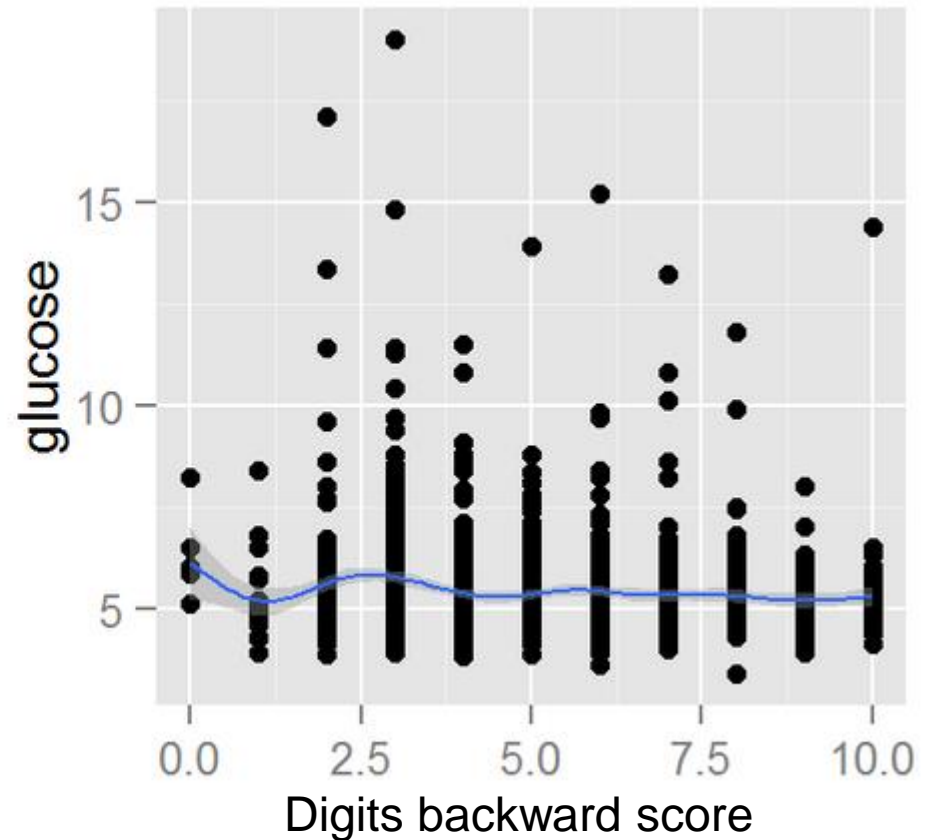
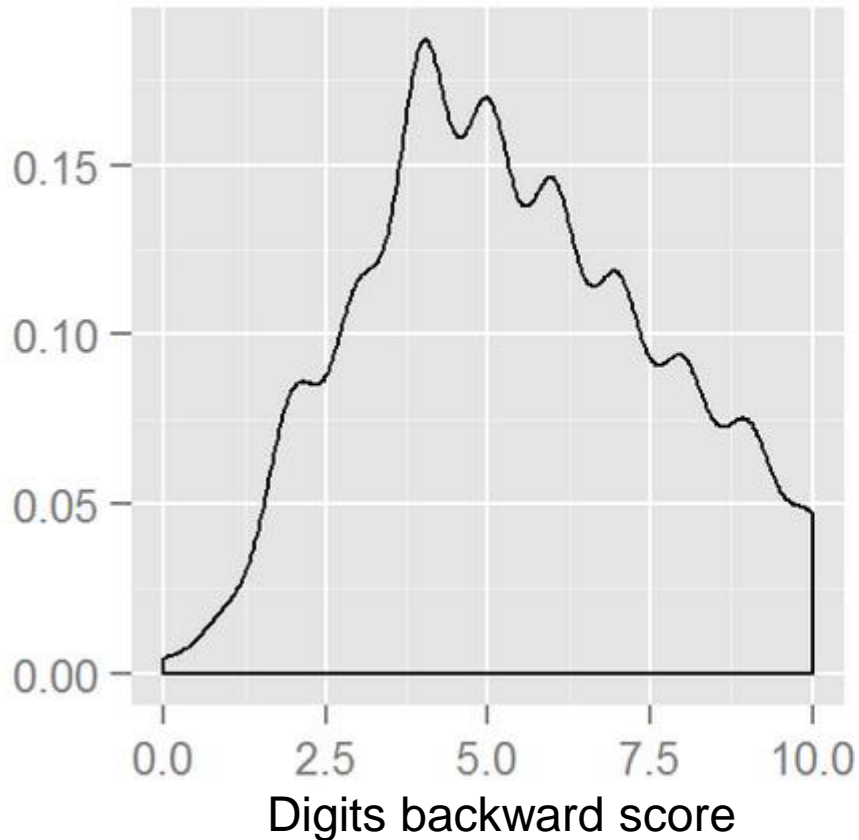
	Estimate	Std. Error	t	Sig.	95% CI	
Diabetics only	0.31	0.1	3.04	<0.01	0.11	0.52
IFG only	-0.03	0.34	-0.09	0.93	-0.69	0.63
Normal only	0.2	0.24	0.85	0.4	-0.26	0.66

Delayed recall



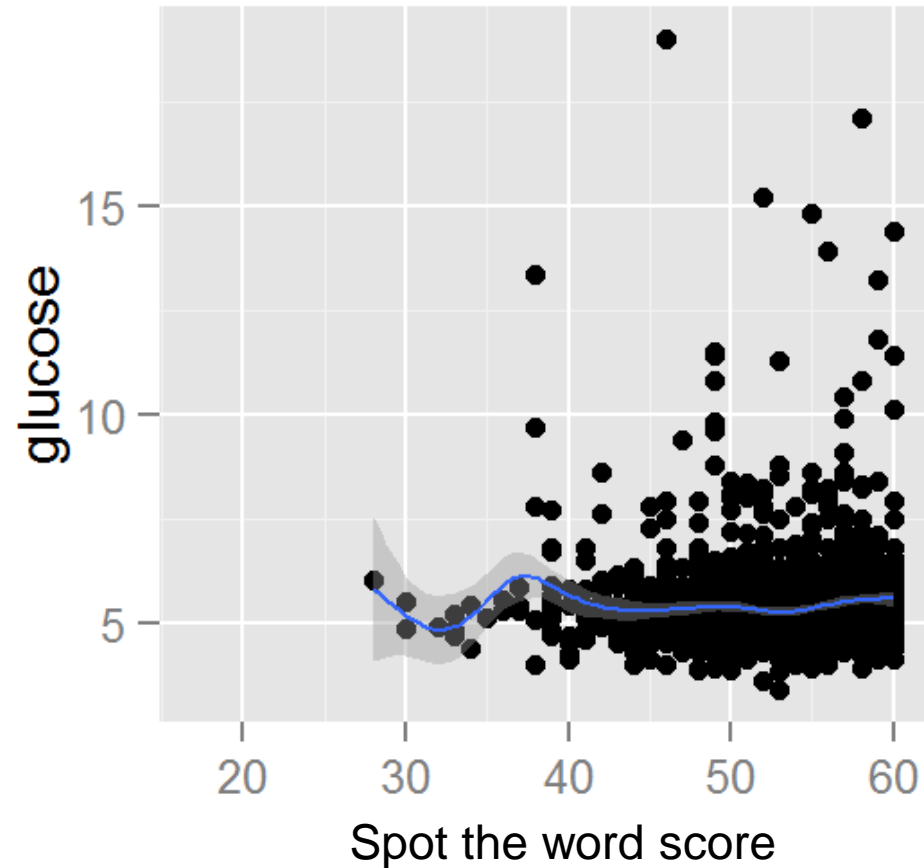
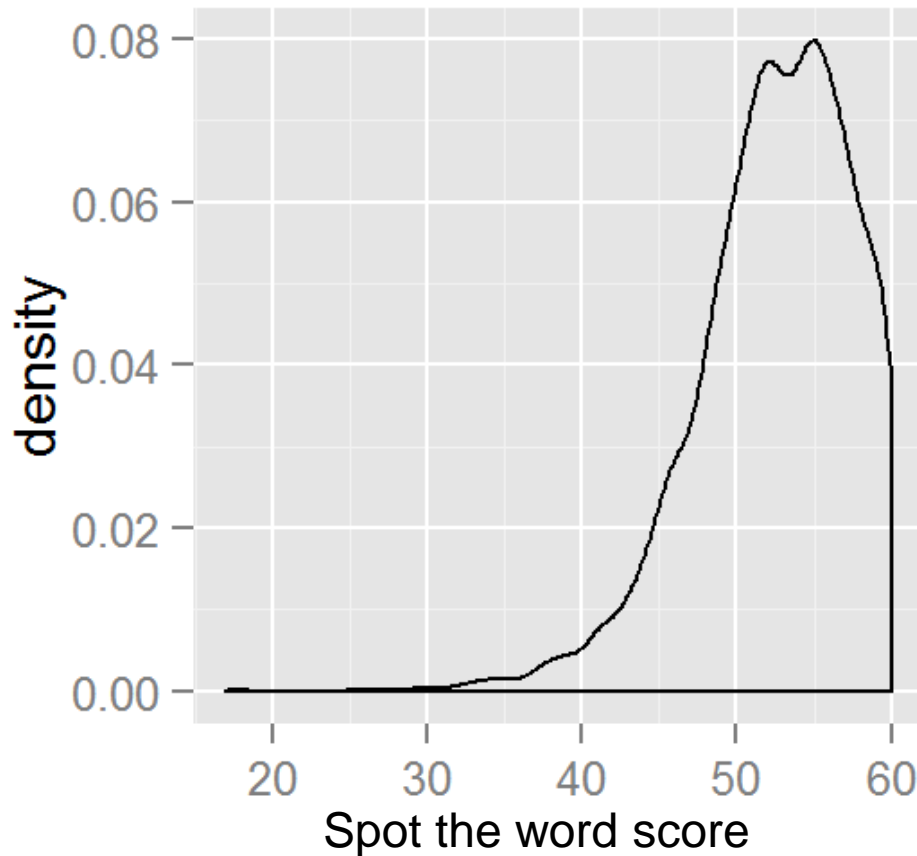
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