

Turning Up the Heat: **Exploring a Low-cost Therapy to Combat Alzheimer's Disease**

A Behavioural Analysis

Seth Hennessy
22030943



Dr Rossana Rosa Porto
Behavioural Neuroscience Group

The Challenge:

Alzheimer's Disease (AD)

Dementia is the **second leading cause of death** in Australia

- *Leading cause of death for women¹*

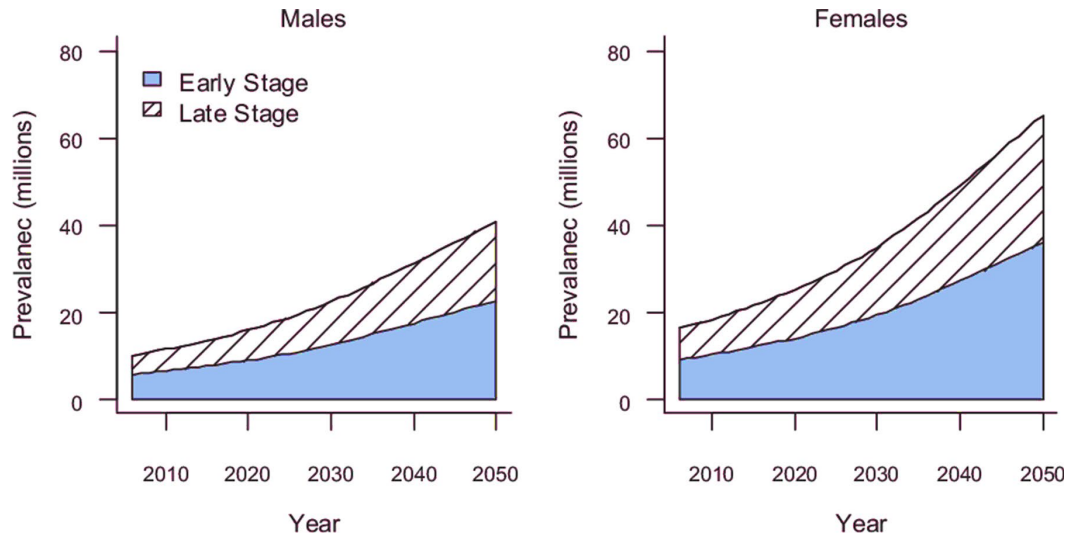


Figure 1: Worldwide projections of Alzheimer's prevalence²

Research Gaps

Lack of studies on HT in AD models

- *Underrepresentation of females in neurodegenerative research*

A Solution:

Heat Therapy (HT)

HT improves health outcomes:

- obesity
- diabetes
- cardiovascular disease
- dementia risk³



Images ©serggn / ©volkankurt via Canva.com

Aim

Investigate if heat treatment can **rescue** cognitive decline in male & female AD mouse models in later stages of the disease

Hypothesis

Heat treatment will reverse AD-like behavioural phenotype at later stages of disease

Significance

Validates heat therapy as a cost-effective & non-invasive solution



WT = Wild type
(normal mice)

TAU58/2 = Tauopathy type
(AD model)

Handling
3 days

Habituation
3 days

Treatment
2x/wk - 8wks

Behaviours
2.5 weeks

Tissue
Collection

Analysis
ANOVA, T-test

Treatment

6-month-old cohort

MALE	WT	TAU58/2
CTR	5	2
HT	4	3

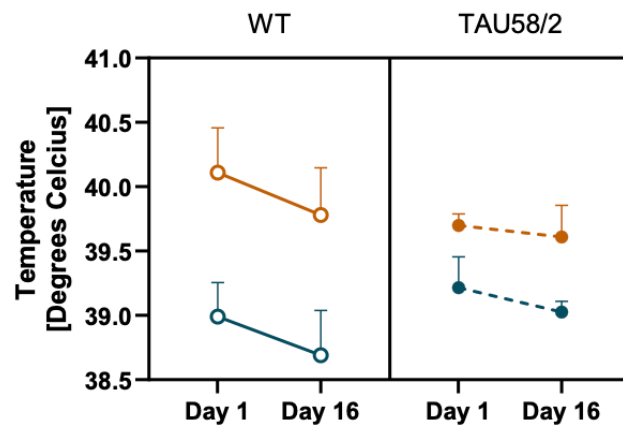
FEMALE	WT	TAU58/2
CTR	2	3
HT	1	2

Two future cohorts: all groups will have $n \geq 12$



Heat treatment

Average temperature



CTR

Control → Room temp ~24°C

HT

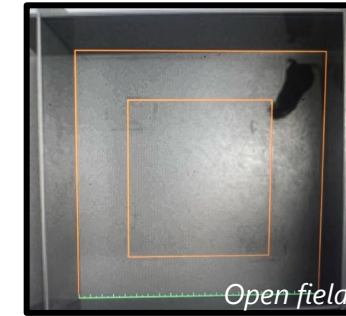
Heat therapy → 42.0±0.2°C

Effect of day $p = 0.03$
Effect of treatment $p = 0.0002$

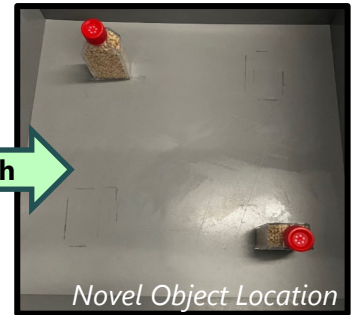
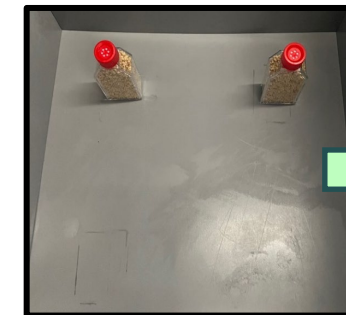
Behaviours

9-months-old

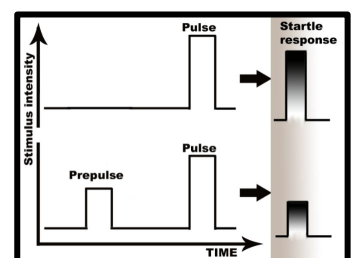
Motor



Memory



Sensorimotor
gating



WT = Wild type
(normal mice)

TAU58/2 = Tauopathy type
(AD model)

Handling
3 days

Habituation
3 days

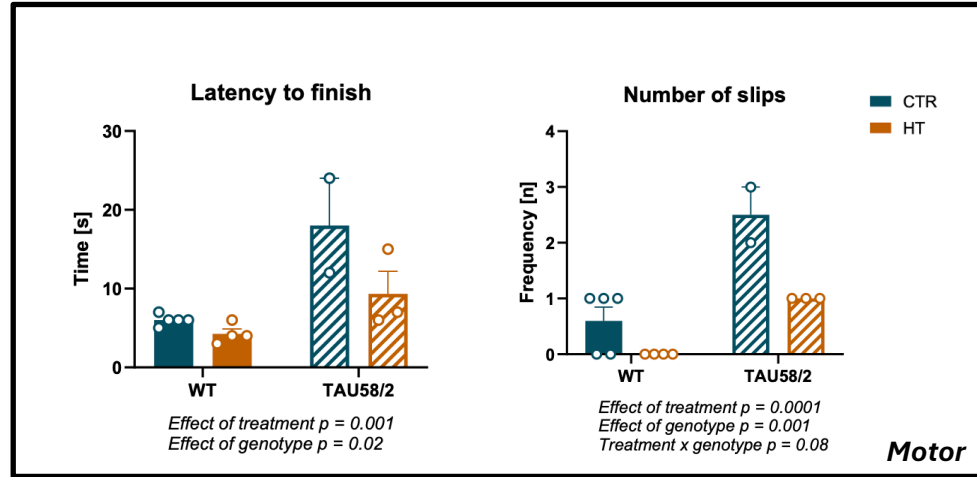
Treatment
2x/wk - 8wks

Behaviours
2.5 weeks

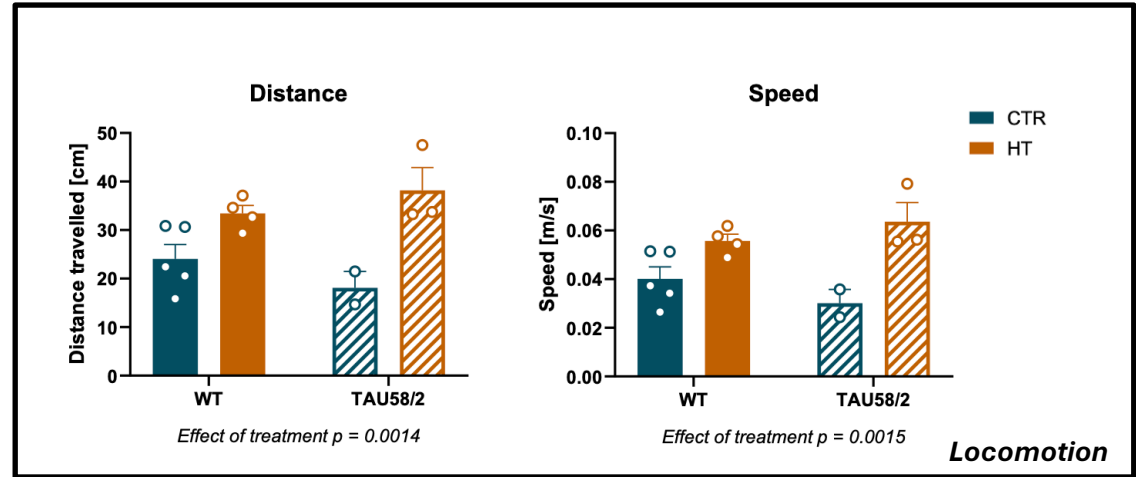
Tissue
Collection

Analysis
ANOVA, T-test

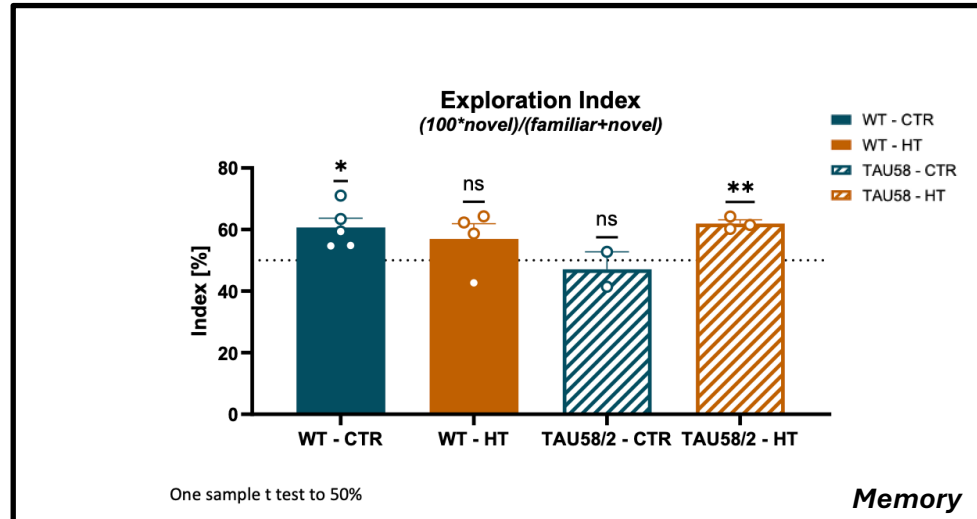
BEAM WALK



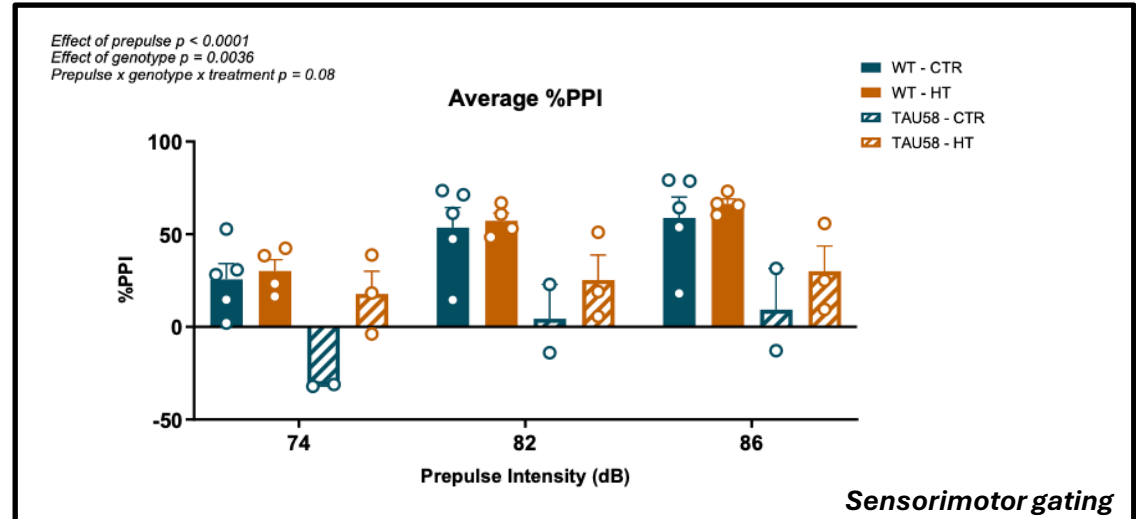
OPEN FIELD



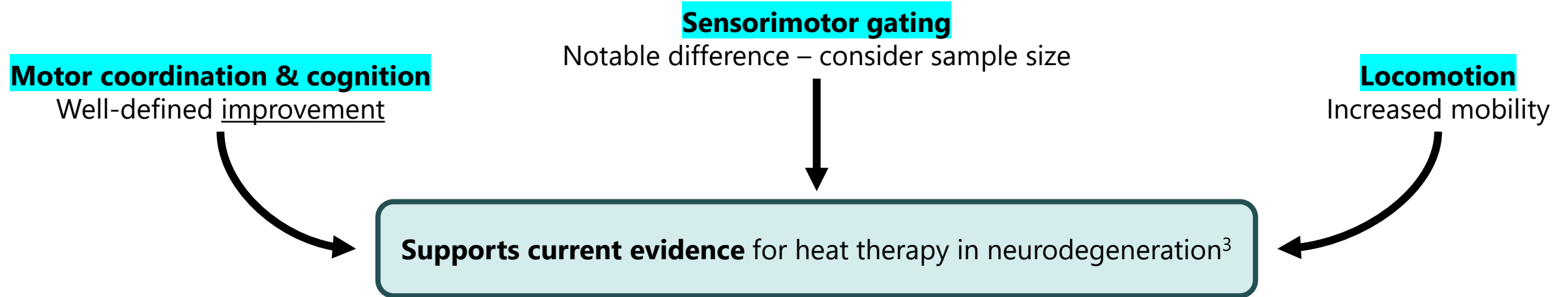
NOVEL OBJECT LOCATION



PREPULSE INHIBITION



Heat therapy has the **potential** to improve behavioural responses in AD mouse models



Strengths

Comprehensive behavioural assessment

Easily translatable

Safe protocol

Limitations

Small sample size*

Future Considerations

Finish all cohorts – six & two months

Western Blot for heat shock pathway

Immunofluorescence (microglia, astrocytes & tau)

**Two future cohorts: all groups will have $n \geq 12$*

*Special thanks to **Dr. Rossana Rosa Porto** for their invaluable guidance & support throughout this project.*

I also appreciate the contributions of:

- Ali Sultanzadeh
- Benjamin Smits
- Isabella Jajjo
- Jack Coleman

Their assistance & insightful contributions were instrumental in the success of this project!

Thanks to Professor Tim Karl, the members of Behavioural Neuroscience group & the animal facility staff for their additional support!

References

1. Australian Institute of Health and Welfare. Dementia in Australia [Internet]. Canberra: Australian Institute of Health and Welfare; 2024 [cited 2024 Nov 13]. Available from: <https://www.aihw.gov.au/reports/dementia/dementia-in-aus>.
2. Brookmeyer R, Johnson E, Ziegler-Graham K, Arrighi HM. **Forecasting the global burden of Alzheimer's disease.** Alzheimers Dement [Internet]. 2007 Jul [cited 2024 Nov 15];3(3):186–91. Available from: <https://pubmed.ncbi.nlm.nih.gov/19595937/>
3. Von Schulze AT, Deng F, Morris JK, Geiger PC. **Heat therapy: possible benefits for cognitive function and the aging brain.** J Appl Physiol [Internet]. 2020 Dec 1 [cited 2025 Mar 18];129(6):1468–1476. Available from: <https://doi.org/10.1152/japplphysiol.00168.2020>.



MY CONTRIBUTIONS

Mice handling & ethics, protein assay, western blotting, statistical & data analysis, tissue collection, heat treatment, behavioural tests