

# IUNS 21<sup>st</sup> ICN International Congress of Nutrition “From Sciences to Nutrition Security”



INTERNATIONAL UNION OF  
NUTRITIONAL SCIENCES

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## Conflict of Interest Disclosure

I have no personal conflict of interest to report in relation to this presentation and my data.

However, part of the presentation has been supported by the **ILSI Europe Nutrition and Mental Performance Task Force**. Industry members of this task force are listed on the ILSI Europe website at **www.ils.eu**. The opinions expressed herein and the conclusions do not necessarily represent either the views of ILSI Europe or those of its member companies.

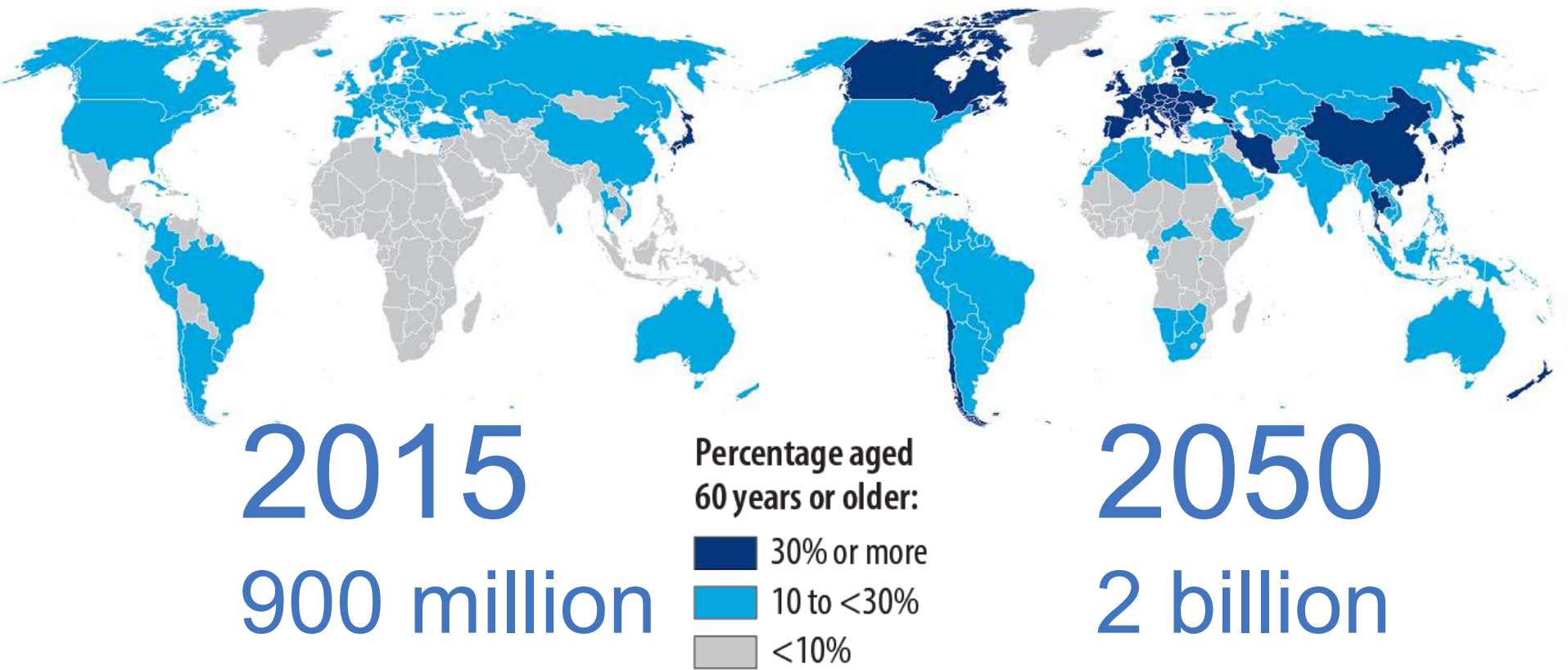
# Nutrition for the Aging Brain: Functional Aspects and Strategies



## Alteration of Neurogenesis during Ageing: Implications for Dietary Interventions



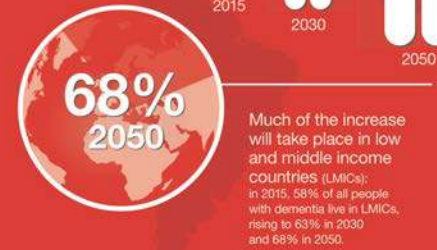
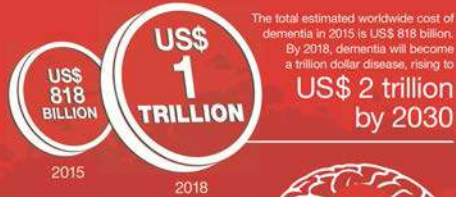
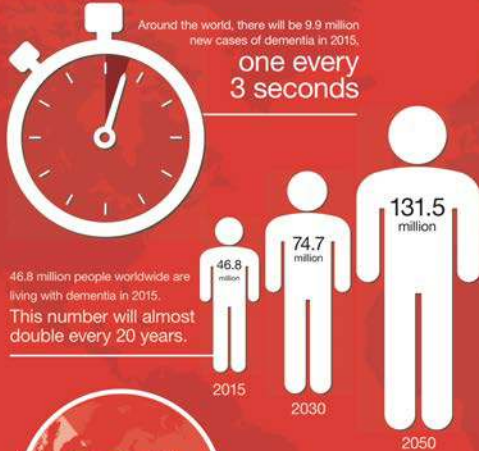
# Populations are getting older



**>65: Risk of dementia doubles every 5 years**

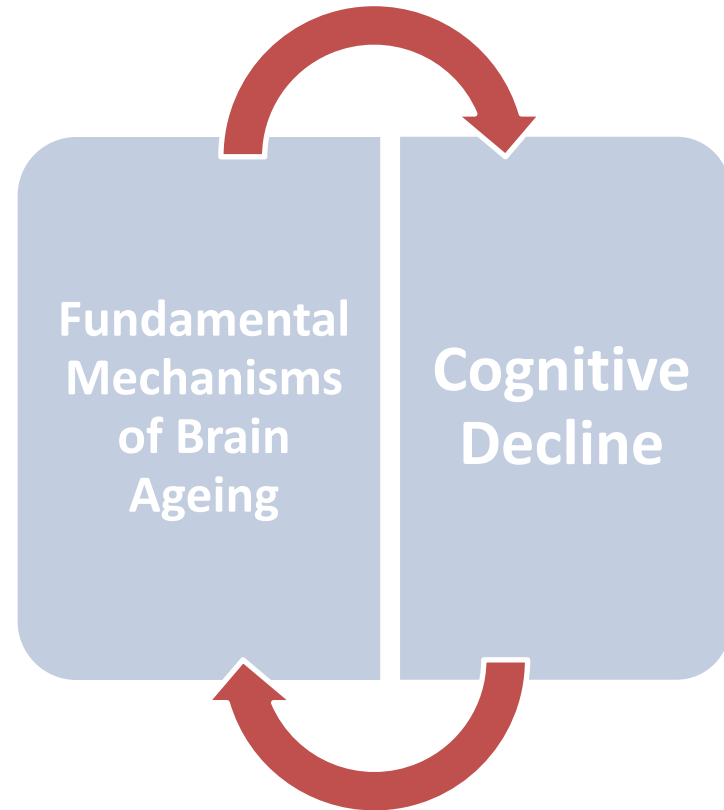
# Global Impact of Dementia

## The global impact of dementia



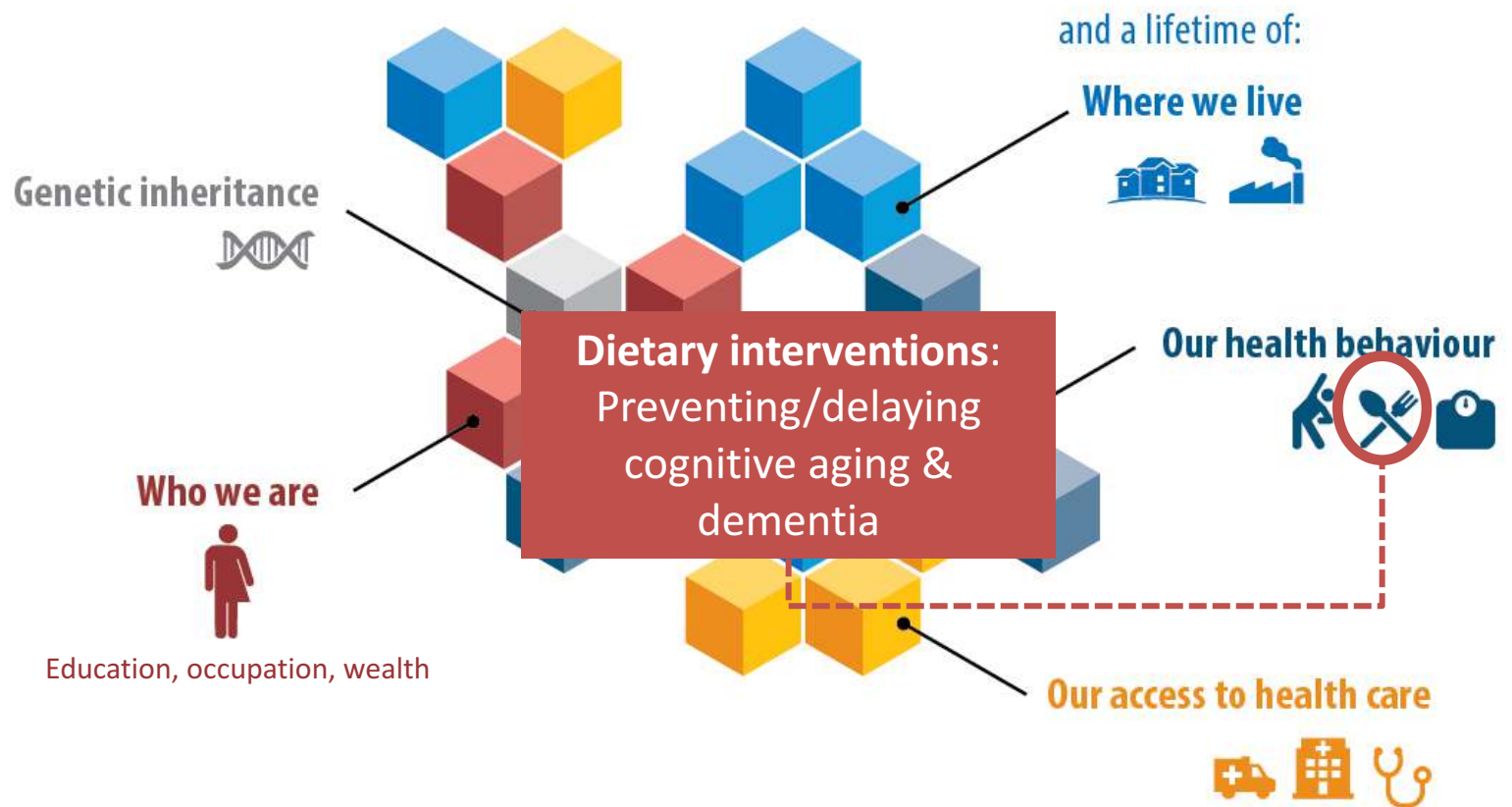
The World Alzheimer Report 2015 was independently researched by King's College London and supported by Bupa.

**Alzheimer's Disease International**  
The global voice on dementia

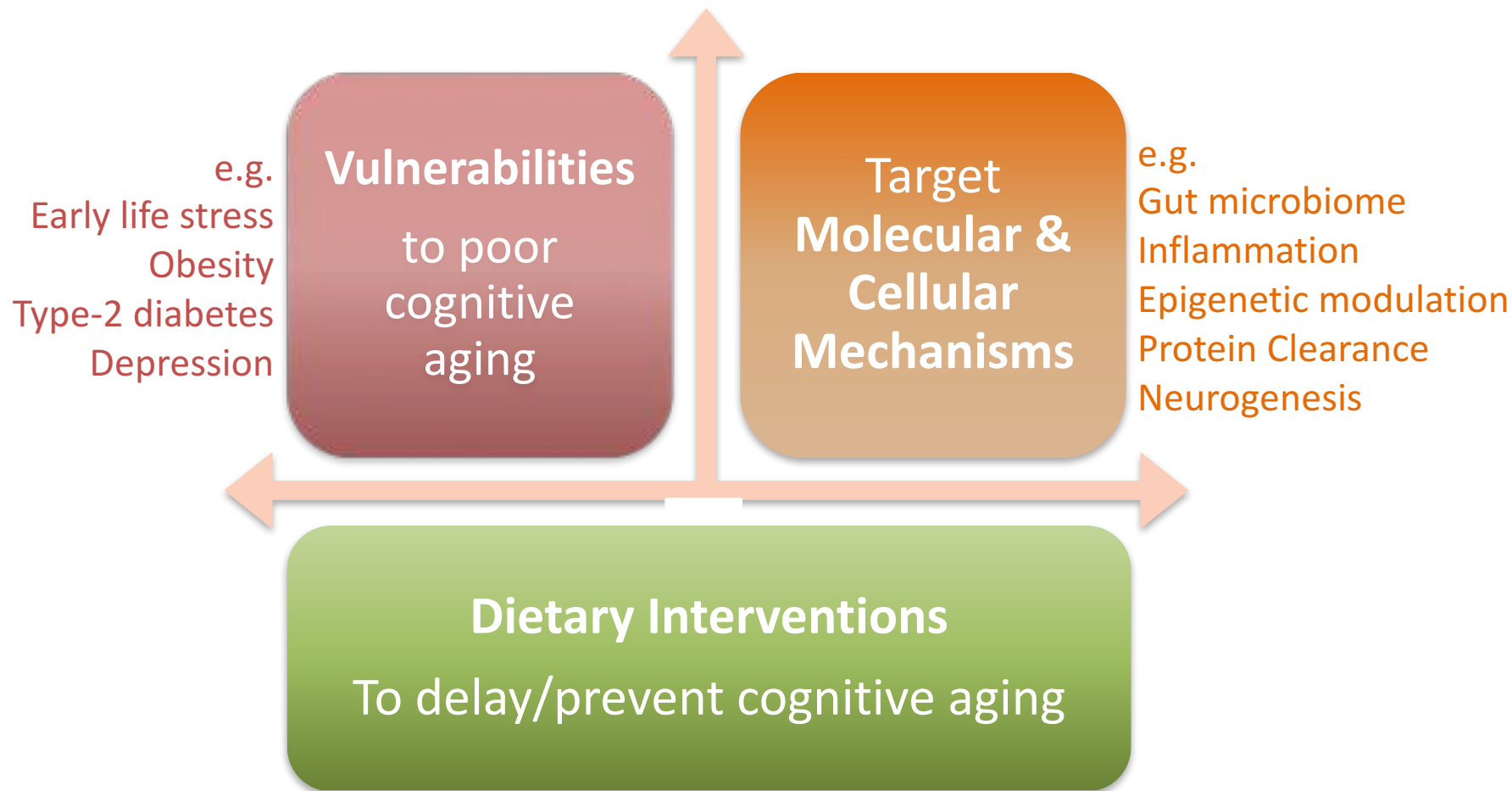


# Not all equal in front of (brain) aging

## What makes us age differently?



# Efficient Dietary Interventions Strategies: Need to identify Vulnerabilities and Mechanisms



Miquel et al. ILSI Nutrition and Mental performance task Force, in revision

*Poor cognitive ageing: Vulnerabilities, mechanisms and the impact of nutritional interventions*

# Adult Neurogenesis?

“Once development was ended, the fonts of growth and regeneration of the axons and dendrites dried up irrevocably. In the adult centers, the nerve paths are something fixed, and immutable: everything may die, nothing may be regenerated.”

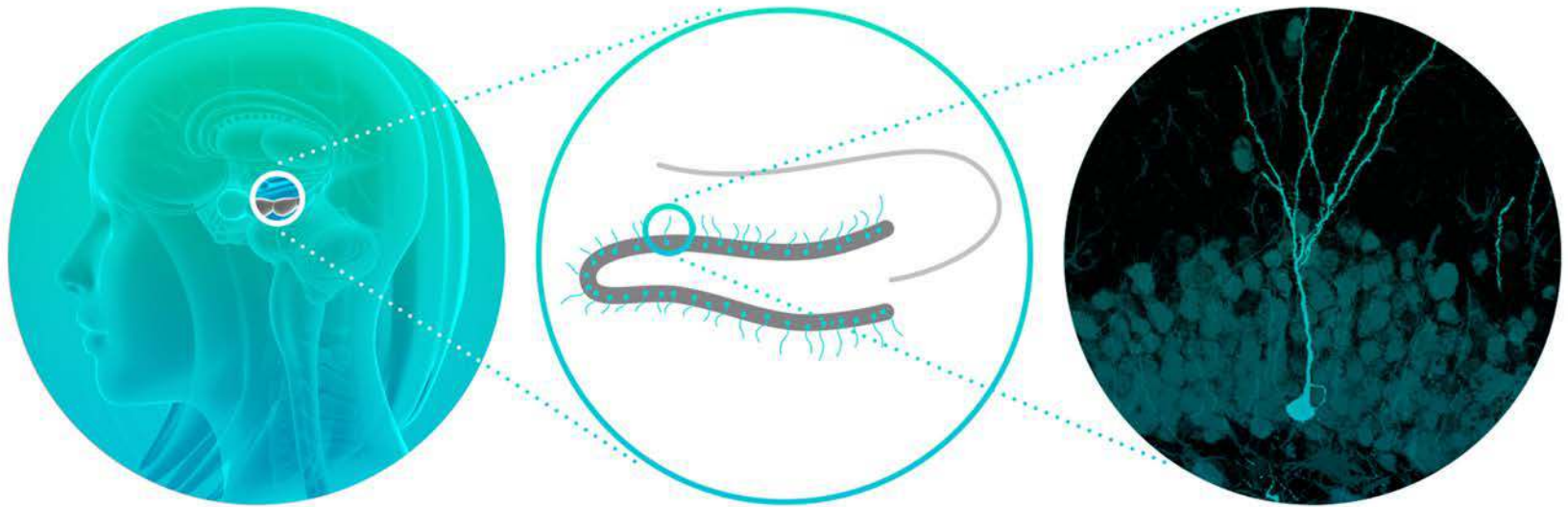
Santiago Ramon y Cajal, 1928



Altman and Das, 1965

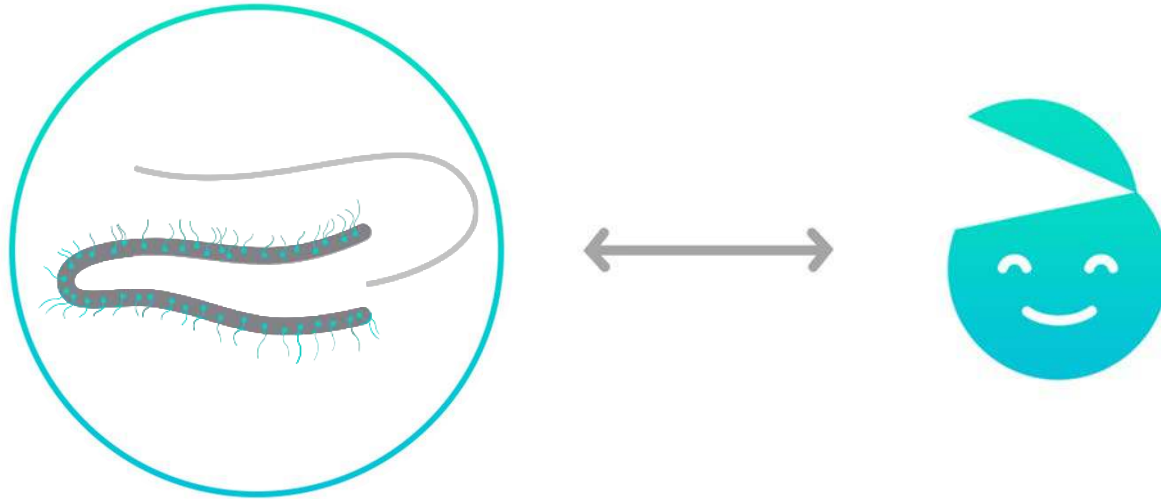


# Human Hippocampus: Adult Neurogenesis



# **Functional Relevance** of Adult Hippocampal Neurogenesis

# Functional Relevance of Adult Neurogenesis: **Learning & memory**



Level of **Neurogenesis** is positively correlated with **hippocampal dependent learning tasks**

## New hippocampal neurons:

- Increase spatial memory capacity
- Reduce interference between memories (pattern separation)
- Add information about time to new memories
- Are involved in forgetting of established context-memories.

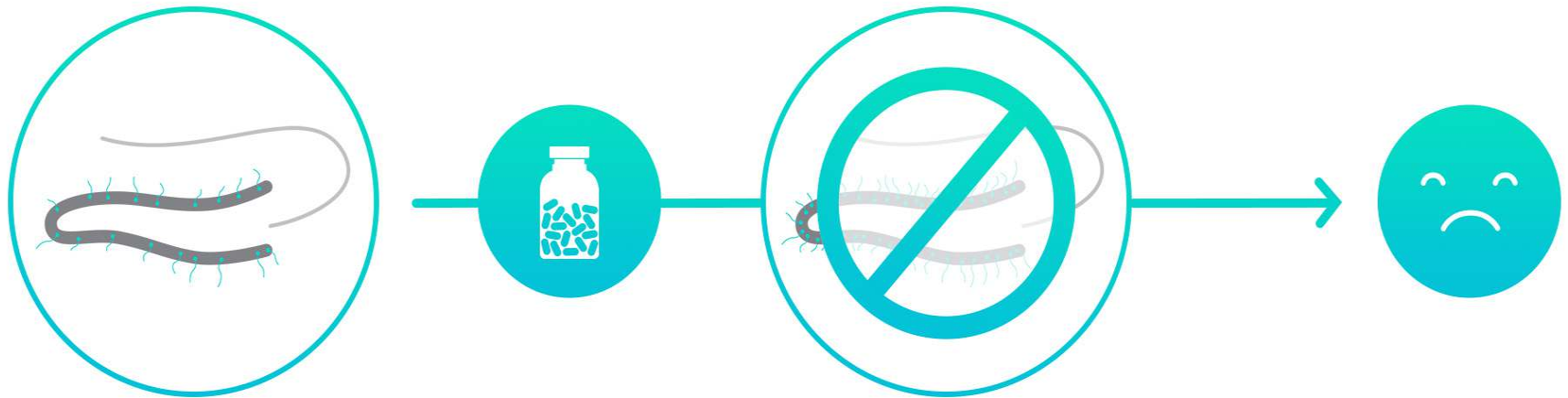


# Functional Relevance of Adult Neurogenesis: Mood & Depression

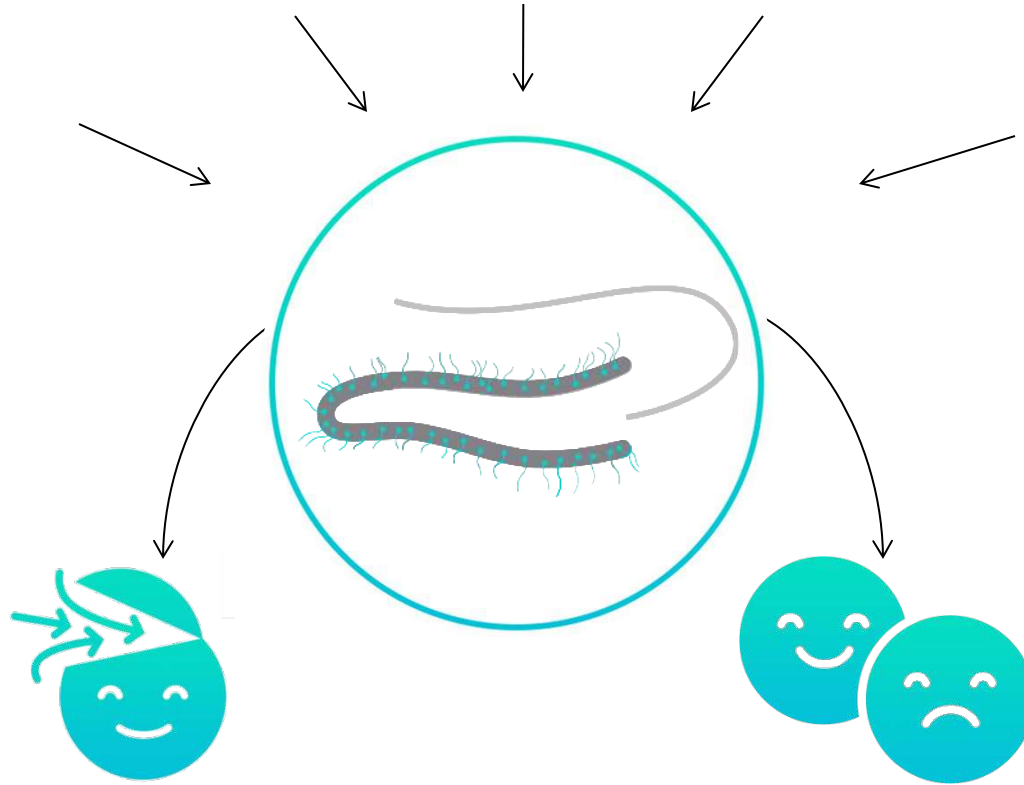
- **Adult Hippocampal Neurogenesis** is reduced in some animal models of depression.
- Many treatments for depression promote **Adult Hippocampal Neurogenesis** and are dependent on functional neurogenesis.



# Functional Relevance of Adult Neurogenesis: **Mood & Depression**



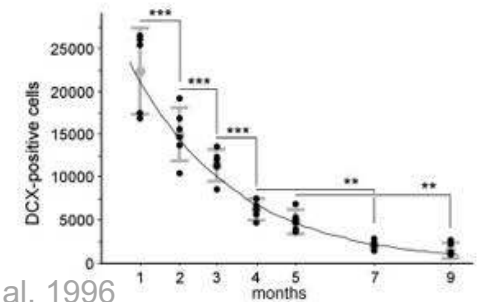
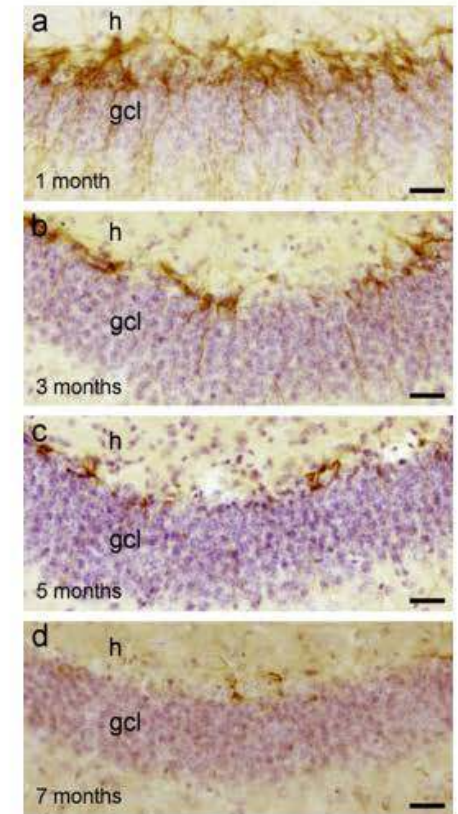
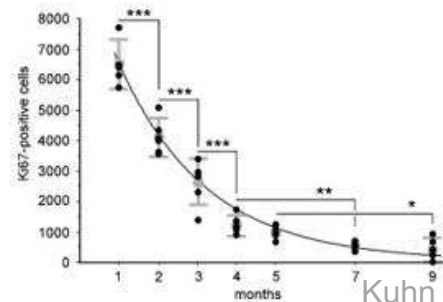
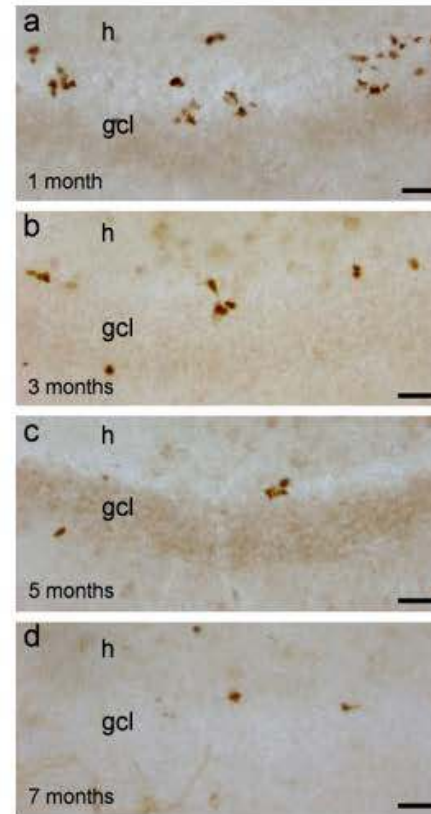
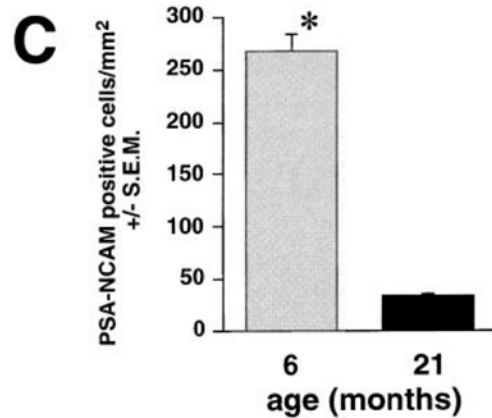
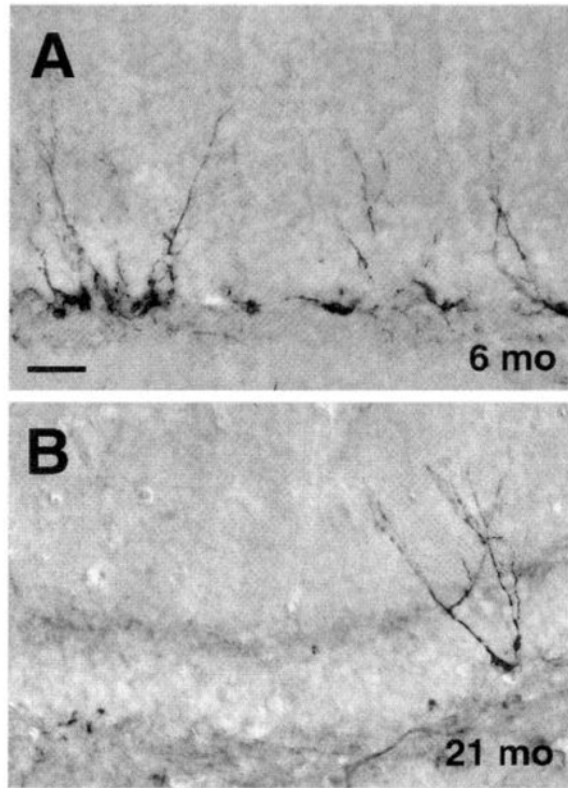
# Adult Hippocampal Neurogenesis emerging as **Target of choice**?



# How is Adult Hippocampal Neurogenesis altered during **Aging**?



# Adult Hippocampal Neurogenesis decreases with age in rodents

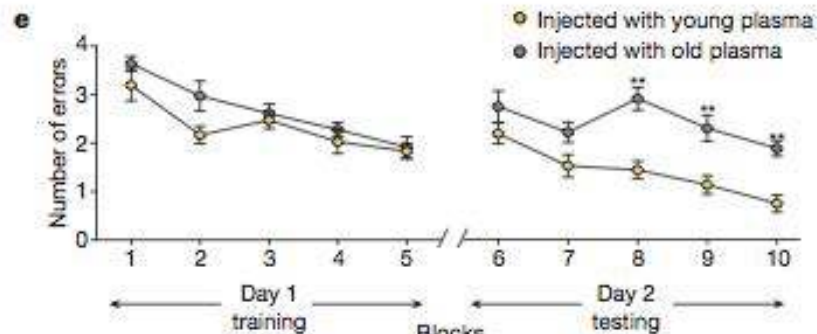
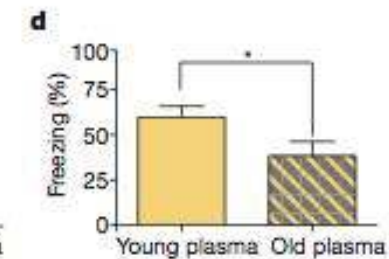
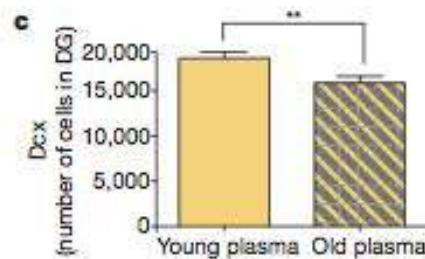
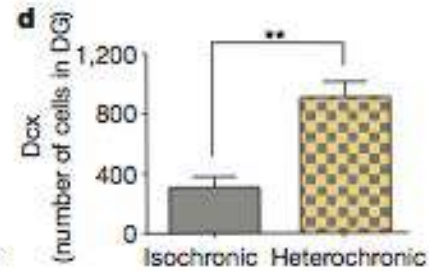
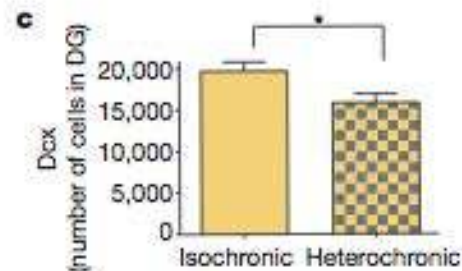
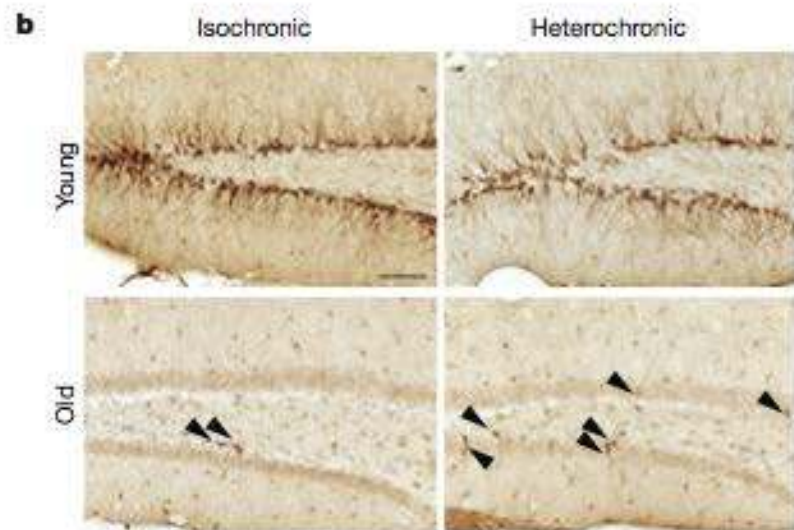
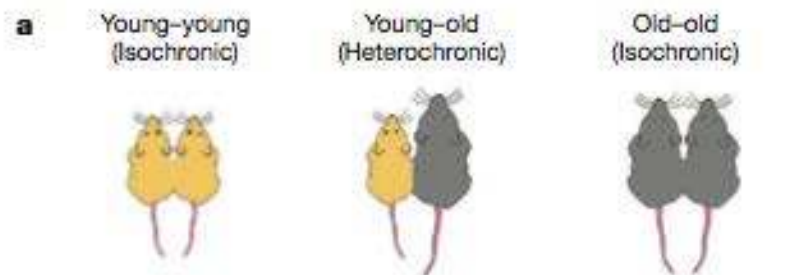


Kuhn et al. 1996

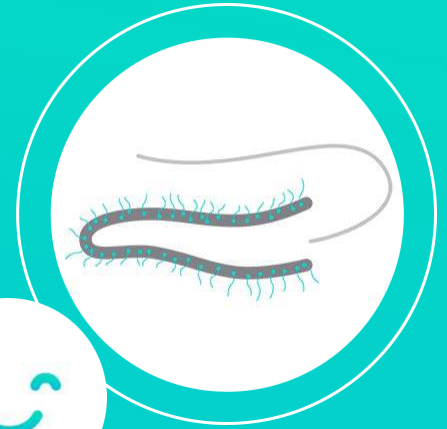
Ben Abdallah et al., 2010

For Review, Klemplin and Kempermann 2007

# Neurogenesis decline with age: Parabiosis and the Systemic Milieu



# **Can life style modulates Adult Hippocampal Neurogenesis?**



# Diet modulates Adult Hippocampal Neurogenesis

DIET	EFFECT ON AHN	STUDY MODELS	REFERENCES
<b>Calorie Restriction</b>	Increased survival	Mouse	68-70, 125-127
<b>Omega 3 fatty acids</b>	Increased	Rat	77, 87, 128-130
<b>Flavonoids</b>	Increased proliferation	Rat chronically stressed	88
	Increased proliferation	Mice	131, 132
<b>Blueberry</b>	Increased proliferation	Rat	133
<b>Curcumin low concentrations</b>	Increased proliferation	Mouse	91, 93-95
<b>Retinoic acid excess</b>	Decreased proliferation	Mouse	106
<b>Vitamin A deficiency</b>	Decreased proliferation (rescued with retinoic acid)	Rat	108
<b>Thiamine deficiency</b>	Decreased proliferation/ survival	Mouse	134, 135
<b>Zinc deficiency</b>	Decreased proliferation/survival	Rat male	100, 103, 136
<b>Folic acid</b>	Increased proliferation	Rat	137
<b>Folate deficiency</b>	Inhibited proliferation	Mouse	138
<b>Increased Homocystein</b>	Inhibited proliferation	Mouse	139, 140
<b>High fat</b>	Decreased proliferation	Rat male	115, 116
	No change	Rat female	
<b>Soft diet</b>	Decreased proliferation	Rat	73, 76
<b>Caffeine</b> at physiologically relevant doses	Decreased proliferation	Mouse	114
<b>Caffeine</b> at supraphysiological doses	Increased proliferation/ Decreased survival	Mouse	113
<b>Caffeine</b> low doses chronically	Decreased proliferation	Rat	
<b>Ethanol</b>	Decreased proliferation	Rat	141, 142
	Decreased proliferation	Mouse	143
<b>Capsaicin</b>	Decreased proliferation	Mice	144
<b>Resveratrol</b>	Increased proliferation	Mice	97, 98
<b>High sugar (fructose)</b>	Decreased proliferation	Male Rat	117
<b>Vitamin E deficiency</b>	Increased proliferation	Rat	145-147
<b>Vitamin E high doses</b>	Increased survival	Rat	148

# Diet, cognition and mood



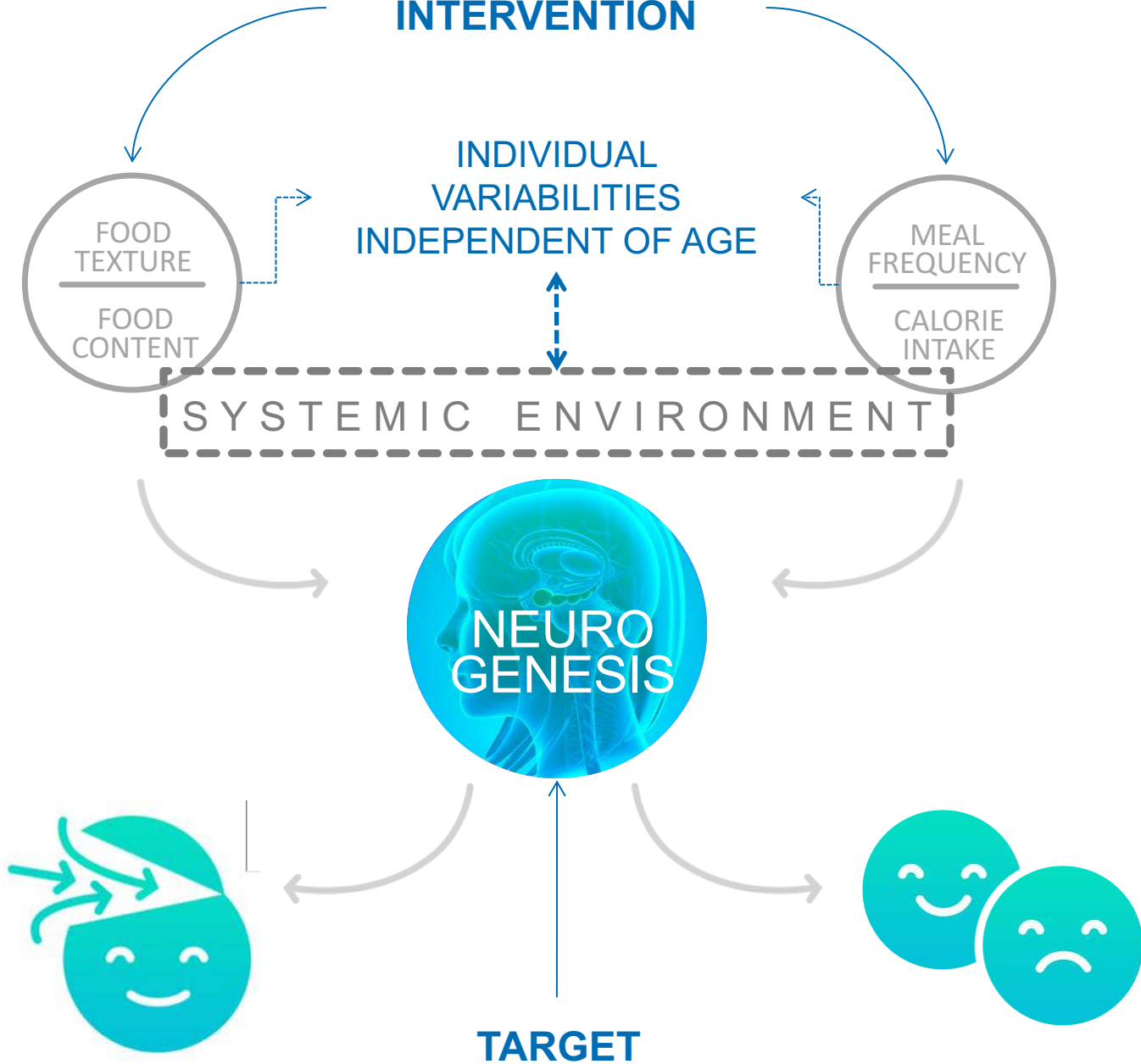
DIET	EFFECT ON DEPRESSIVE BEHAVIOUR	EFFECT ON LEARNING AND MEMORY	STUDY MODELS	REFERENCES
Caloric/Dietary Restriction Intermittent Fasting	Improved	Enhanced spatial learning	Rat (aged)	149
		Increased learning and motor performance	Mouse	150
		Increased learning consolidation	Mouse	151
		Enhanced verbal memory	Human (Healthy Elderly)	71
		Improved spatial learning	Rat (traumatic brain injury)	152
Omega 3 fatty acids	Improved		Human (bipolar)	85, 86
	Delayed onset		Human (bipolar)	154
	Improved		Human (bipolar)	155
	No benefit		Human (bipolar)	156
		Improved spatial memory	Mouse (Alzheimer model)	157
		Improved acquisition and retention	Mouse (Aged)	158
		Improved learning performance	Rat (Diabetic)	159
	Small effect	No effect	Human (recovered from depression)	160
Flavonoids	Improved		Rat	89
		Improved	Various Species	For review <sup>161</sup> , 162
		improved	Rodent Species	For review 162-164
Blueberry		Improved	Rat	165-169
		Increased spatial memory	Rat	90
		Improved	Human (old age)	170
Curcumin		Improved	Mice	171
		Improved cognitive performance	Human	92
		Improved cognitive performance	Rat	94, 172
		Improve spatial memory and learning	Rat	173, 174
		Improved	Mice	175, 176
	Improved	Rat (Alzheimer's disease)	177	
	Improved		Rat	96, 178-180
Improved		Mice	181	

# Diet, cognition and mood



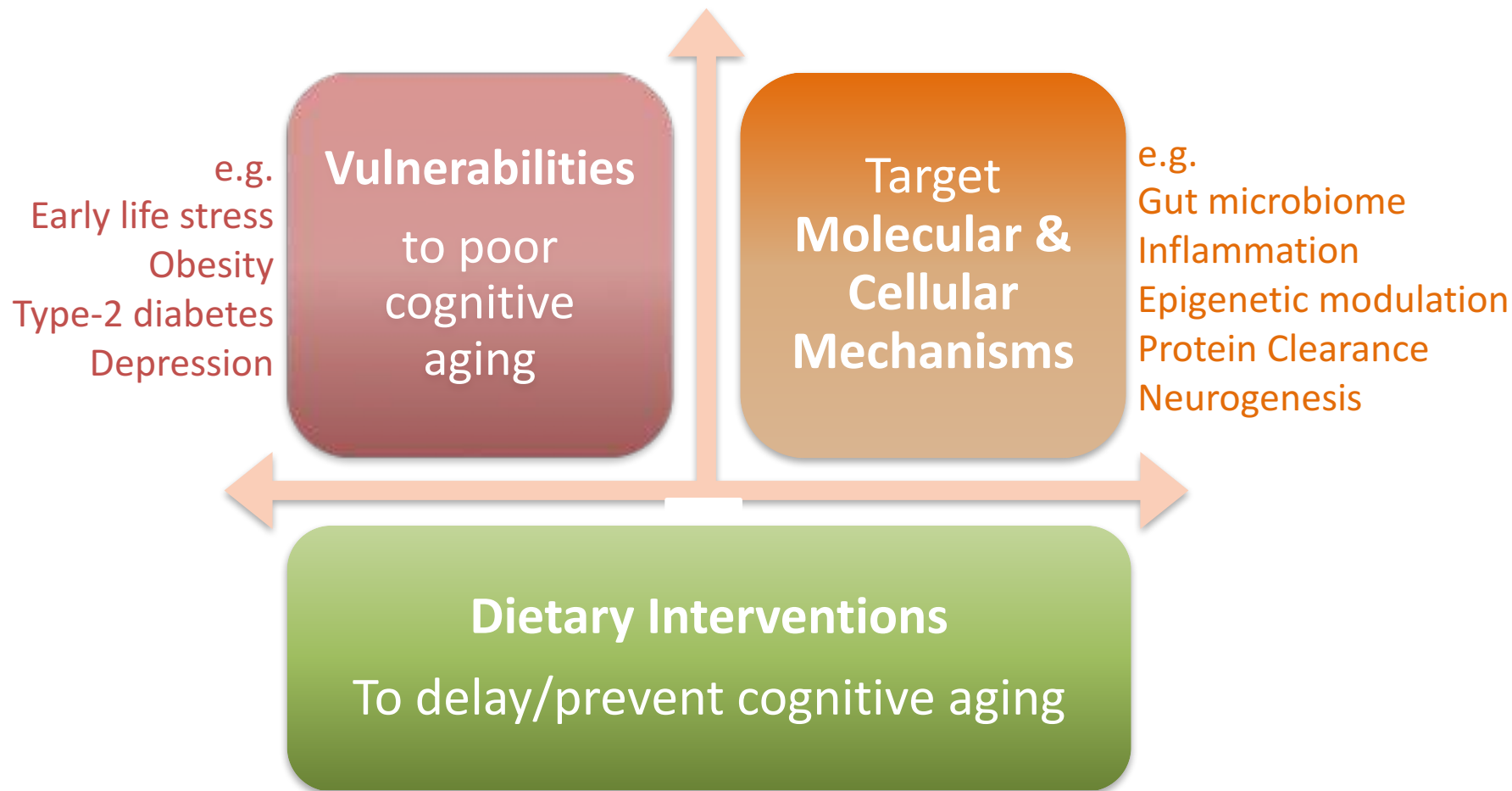
<b>Retinoic acid excess</b>	Improved		Mouse	107
<b>Vitamin A/retinoid deficiency</b>		Impaired spatial learning and memory	Rat	182
		Impaired relational memory	Mouse	183
<b>Zinc</b>	Improved		Rodent, Human	For review <sup>102</sup>
<b>High fat</b>	Exacerbates	Decreased spatial learning	Rat	184
		Decreased learning and memory and increased risk for dementia	Rat	185
		Impair spatial learning	Mice	186
		Increased susceptibility to spatial learning deficit	Rat	188, 189
		Impair memory	Rat	190
<b>High sugar</b>		Impaired spatial learning	Rat	191
		Impaired spatial learning	Rat	192
		Impaired	Rat	193
<b>Low glucose (extracellular)</b>		Impaired memory	Rat (Aged)	194
<b>Soft diet</b>		Impaired	Rat (Alzheimer's model)	74
		Impaired learning and memory	Aged animals	For review <sup>195</sup>
		Impaired spatial learning and memory	Mice (female albino)	196
<b>Caffeine</b>		Improved object recognition	Mouse	197
	Reduced risk		Human	198
	Reduced risk		Human	199
		Improved cognitive function	Rat	200
<b>Ethanol</b>		Improved associative learning with moderate chronic consumption	Mouse	201
		Impaired	Human	202
	Induced depressive behaviour		Rat	203
<b>Capsaicin</b>		No effect	Mice (young)	144
<b>Resveratrol</b>	Improved		Mice (despair test)	99
		Improved cognitive function	Mice	98
<b>Vitamin E deficiency</b>	Associated risk		Human (depression)	204
	Associated risk		Human (depression)	205
	No association		Human (aged adult with depression)	206
<b>Vitamin E</b>		Protective effect	Rat (brain injury)	207
		Delayed memory loss	Mouse	99

# MODES OF INTERVENTION





# Efficient Dietary Interventions Strategies: Need to identify Vulnerabilities and Mechanisms



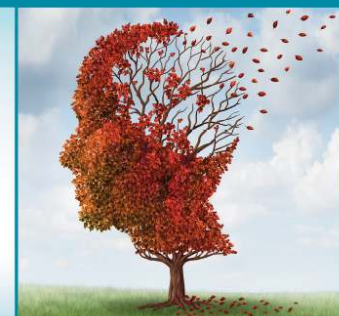
Miquel et al. ILSI Nutrition and Mental performance task Force, in revision

*Poor cognitive ageing: Vulnerabilities, mechanisms and the impact of nutritional interventions*



3<sup>rd</sup> WORKSHOP  
Nutrition for the Ageing Brain:  
Moving Towards Clinical Applications

Save the Date: 30-31 August 2018  
Madrid, Spain



**Main themes to be addressed:**

- Impact of nutrition on brain functions using neuroimaging technologies;
- Microbiome and immune status: impact on brain function;
- Biomarkers of food intake and cognitive health;
- Sleep deprivation: effects on diet and cognitive performance;
- New methodologies applied to dementia and how nutrition could play a role.

Organised by the ILSI Europe  
Nutrition & Mental Performance Task Force  
More information can be found on [www.ils.eu](http://www.ils.eu).

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The **P**sychiatry **R**esearch **T**rust

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