





College of Human Medicine

Welcome to the Your Health Lecture Series!

Tonight's Lecture:

Exploring the Latest in Alzheimer's and Dementia Care

Event Begins @ 6:30pm

Moderator: Dr. Jean Nagelkerk

Vice Provost for Health

Grand Valley State University

Special Thanks!

alzheimer's % association®

GREATER MICHIGAN CHAPTER





The Source for Seniors

SPECTRUM HEALTH NEUROSCIENCES

Mary McPhail-Ciufo, DO

Section Chief Behavioral Neurology Division of Clinical Neuroscience MSU College of Human Medicine



Tonight's focus areas:

- Alzheimer's disease overview
- The difference between dementia and healthy brain aging
- Treatment, prevention and research

Nicholas Kanaan, PhD

Associate Professor Department of Translational Neuroscience MSU College of Human Medicine



Tonight's focus areas:

- The latest Alzheimer's disease research
- Alzheimer's diagnosis and medical interventions

Rebecca Davis, PhD, RN

Associate Dean for Research and Scholarship Professor Kirkhof College of Nursing Grand Valley State University



Tonight's focus areas:

- Caregiving and the impact on families/caregivers
- Interventions
- The future of Alzheimer's

Alzheimer's Disease,
Dementia,
and

Healthy Brain Aging

Mary McPhail-Ciufo, DO Section Chief Behavioral Neurology Division of Clinical Neuroscience MSU College of Human Medicine









An overview for today's talk

The complicated 7 questions I'm asked all the time...

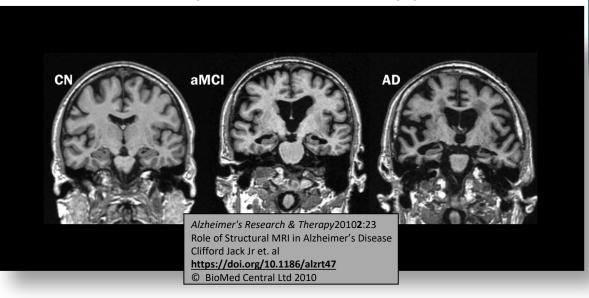
- What is Alzheimer's Disease?
- ▶ What's the difference between Alzheimer's Disease and Dementia?
- ▶ What is the difference between Alzheimer's Disease and Normal Aging?
- ▶ How do I know if my memory loss is Alzheimer's or Normal Aging?
- What are the current treatments?
- What can I do to prevent it?
- What about research?

Healthy Severe Brain Alzheimer's





Images above from National Institute of Aging



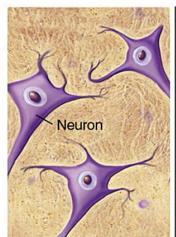
What is Alzheimer's Disease?

Plaques & Tangles

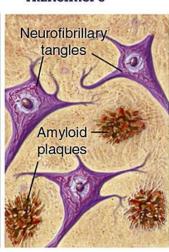
- Alzheimer's disease consist of abnormal proteins that are occurring inside and between the cells of the brain
- We call these abnormal proteins "Plaques and Tangles"
- ▶ So, what's the difference between Alzheimer's disease and Dementia?

Normal vs. Alzheimer's Diseased Brain

Normal



Alzheimer's





© 2000 by BrightFocus Foundation

What's the difference?

- Dementia Is defined as the impairment of thinking along with the loss of independent everyday functioning.
- ► Tasks that we typically perform everyday independently are tasks like driving, balancing a checkbook, cooking a meal on the stove, and performing our jobs at work.
- Dementia only defines the STAGE of impairment a person has. It does not define the WHAT or WHY a person has Dementia.
- I like to think of the stages of brain functioning as being like the dollars in a piggy bank
- I consider Alzheimer's disease to be like a thief who takes \$10 dollars or more from your piggy bank every year

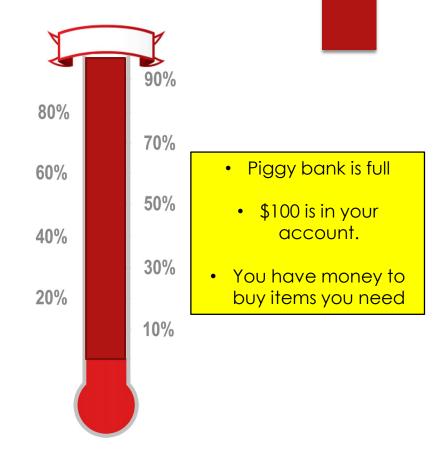




"I'll be needing one of your legs as well!"

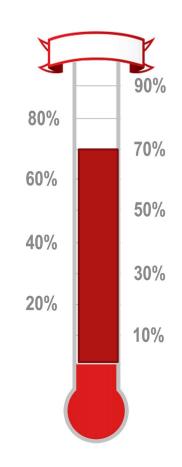
A Healthy Brain is a Healthy account

- We are functioning independently
- We have very few to no complaints about our thinking
- Those we live with and work with have not noticed any change in our personalities, mood, or ability to think



Mild Cognitive Impairment

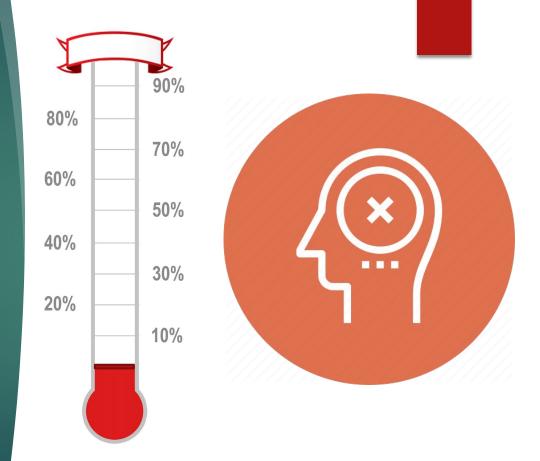
- Our piggy bank is less than what we had before but it is still enough for what we need everyday
- Our friends, family, and doctors may have noticed a change.
- Our piggy bank is at risk of being drained to zero if something should happen.
 - An example of something sudden to our brain account would be a stroke.
 - ▶ If this piggy bank has a thief stealing money, then it is very likely our piggy bank will continue to dwindle down.



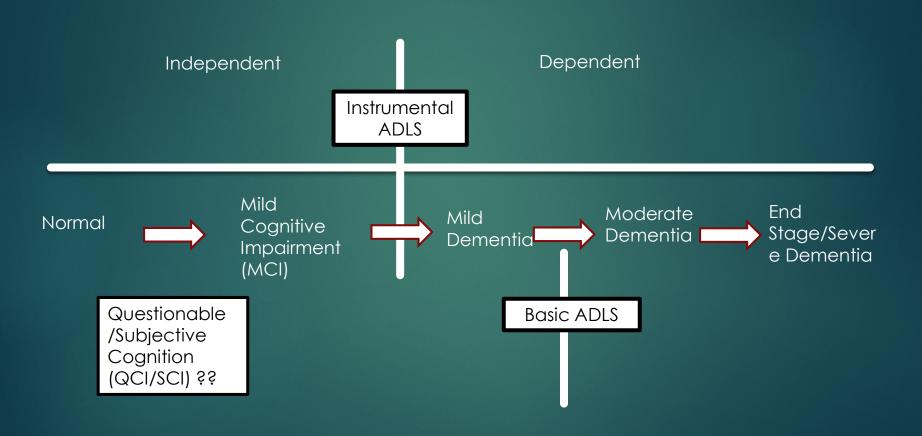
- \$70 is in your account.
- You still have money for buying items.
- We are going to be more careful with our money now
- We can't predict sudden purchases like having to buy a new car; but <u>we</u> <u>can try to prevent them</u>
- Thieves, are tricky. We can't always know for sure if they are there and how much they are taking each year.

Dementia

- We have no money left for our everyday needs in our piggybank
- This is "brain failure"
- There is hope however.
- All dementias are treatable to a degree although there is no cure for progressive reasons...yet
- Some treatments we have will be like adding money to our piggy bank

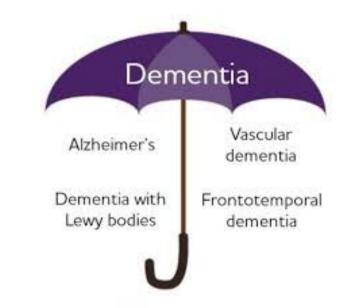


The stages of cognition



Causes of Dementia

- ► Alzheimer's disease is just ONE reason why a person might be impaired enough in their thinking to have loss of independence in these tasks.
- In other words, Alzheimer's disease is just ONE reason why a person may have Dementia.
- ► There are many other reasons why a person might be impaired enough to have Dementia.
- ► For example, they may have had several strokes or may have a progressive neurodegenerative diseases such as Lewy body disease.



What is normal aging then?

Normal aging	vs	Alzheimer's
Making bad decisions once in a while	vs	Making poor judgements and decisions a lot of the time
Losing things from time to time	vs	Misplacing things often and being unable to find them
Missing a payment	vs	Problems taking care of monthly bills
Forgetting which day it is and remembering later	vs	Losing track of the date or year
Sometimes forgetting which words to use	vs	Trouble having a conversation

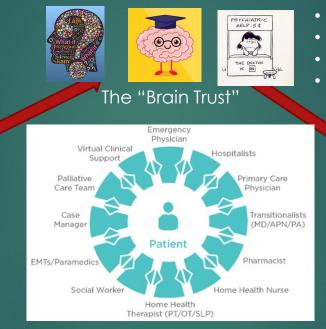
How do I know what I have?

- Establish a relationship with a Primary Care Doctor (PCP) who will..
 - ▶ Help you and your family lead a healthier life
 - ▶ Help Identify Problems
 - ▶ Help manage chronic conditions
- Part of a memory evaluation from a PCP will include a basic gathering of history and may also include a bedside cognitive screener. Based on the history and the bedside testing. Your PCP may perform the following:
 - Refer to a Neurologist
 - Refer to Neuropsychology for more in-depth testing
 - Check labs that are important for our brain
 - Order imaging of the brain
 - Order a sleep study





Your primary care doctor is the team captain of your healthcare!



- Neurology
- Psychiatry
- Neuropsychology
- Psychology (talk therapy)



- Education
- There for help to access social resources (e.g. caregiving)



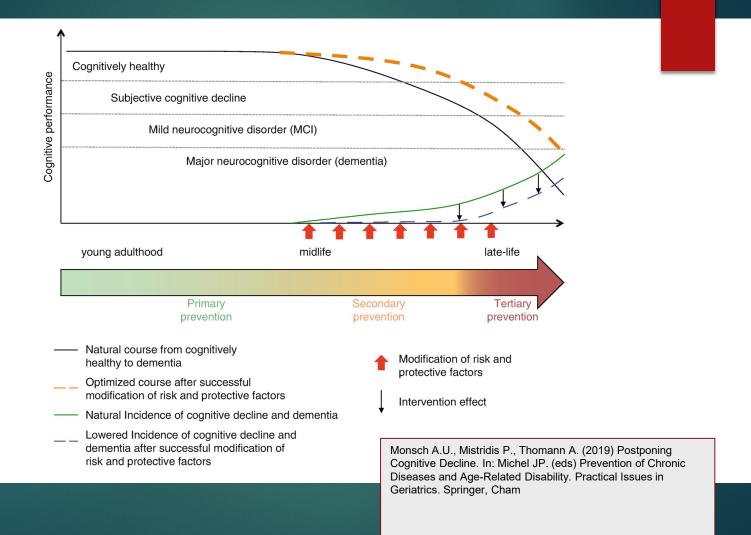
- Primary Care Providers
- Will need regular follow up
- Incredibly important

What are current treatments for Alzheimer's?

- Medications that will help the chemical nature of the brain so that healthy cells may operate better to compensate for sick cells
 - ▶ Donepezil and Memantine are two common medications used
- Supportive care
- ▶ Unfortunately, there is no cure for Alzheimer's

So what's the big picture here?

- Screen early
- Intervene earlier
- Whole person care
- Integrated multidisciplinary teams
- ▶ Integration into the community
- ▶ Hope



So what can I do to prevent it?

Life's Simple 7

- ▶ Don't smoke
- ▶ Watch your weight
- ▶ Eat a healthy diet
- Exercise
- Control your blood pressure
- Control your blood sugar
- Control your cholesterol







And an extra "7" I like to stress!

- Wear a helmet
- Reduce Falls (use your cane/walker/take a minute to stand up)
- Identify & Treat depression/anxiety
- Identify and treat Sleep problems such as Sleep Apnea
- Be a lifelong learner Challenge yourself and seek novelty
- Stay social
- ▶ Love



Dr. Marian Diamond

Social Needs and further Education

- Alzheimer's
 - ▶ Alzheimer's Association: 24/7 Helpline 800.272.3900
 - ▶ For patients, families, and even your doctors!
 - ▶ 24-7 Really? Yes Really!
 - ▶ (I called them at 3AM prepping for a conference once)

alzheimer's \bigcap association

- Parkinson's Disease/Lewy Body Disease/atypical parkinsonism (PSP/CGBD/MSA)
 - ▶ Parkinson's Disease Foundation: 1.800.4PD.INFO (1-800-473-4636
 - ▶ LBD Caregiver Link: 800.539.9767
- Stroke
 - American Heart Association And American Stroke Association
 - ▶ 1-800-AHA-USA-1
- ▶ Frontal temporal Disease: 1-866-507-7222



Further questions?

- We are a team. You're the most important member on the team. You are not alone.
- ▶ If you want to advocate for the bigger picture (e.g. changing Laws, funding research). Let the Alzheimer's Association know.

Thank you for your time listening!





Wrangling Tangled Tau: Recent Insights into Alzheimer's Disease

Nicholas M. Kanaan, PhD Associate Professor

Michigan State University
College of Human Medicine
Department of Translational Neuroscience

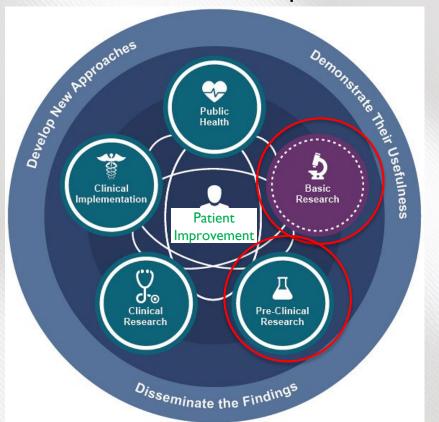






Research, The Bigger Picture and You

Translational Science Spectrum



How Can I Get Involved?

Participation in trials is key to advancing towards our goals





https://alzheimersalliance.msu.edu/index.html



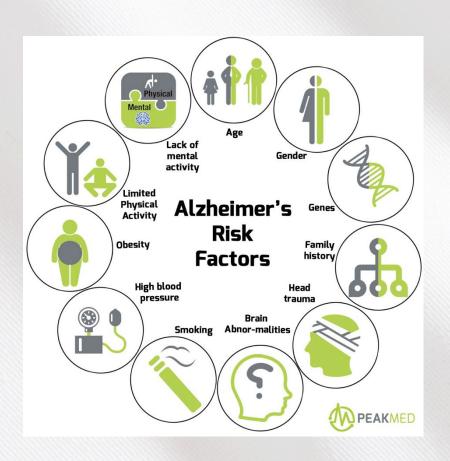
https://alzheimers.med.umich.edu/research/

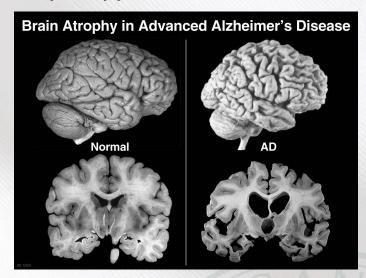




What Has Research Taught Us About Alzheimer's?

IT'S COMPLICATED, but this creates multiple opportunities...





Neuropathology







Plaques: Amyloid-β

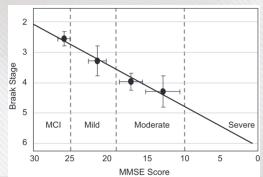
Tangles: Tau

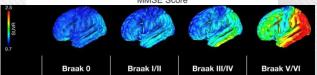
- Neuroinflammation
- Blood-Brain-Barrier Compromise
- Genetic Mutations Inherited AD (genes related to amyloid)
- Several Other Potential Causes



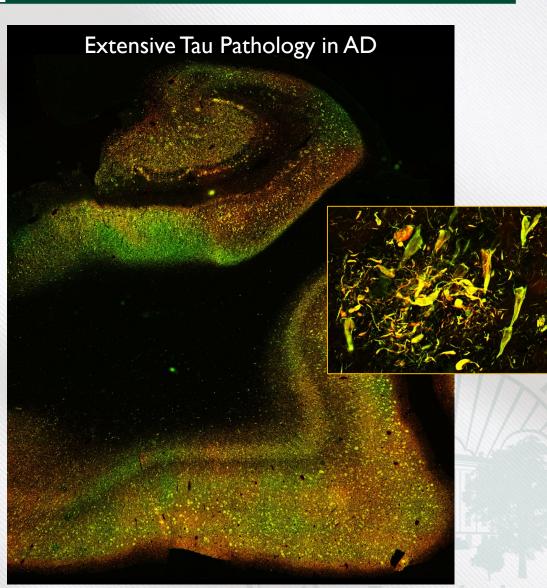
Tau as a Main Culprit in Alzheimer's Disease

- Tau is a <u>normal protein</u> that regulates microtubules
- A hallmark pathology and widespread
- Tau changes and aggregates in disease
- Tau pathology correlates with cognitive decline







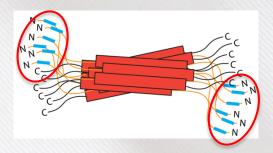


Identifying Disease Mechanisms: Toxic Tau

Tau Pathology



Misfolding Creates A Problem



Triggers Axonal Toxicity Pathway



- 1. Discovering a specific pathway!
- 2. Identifying a potential target!

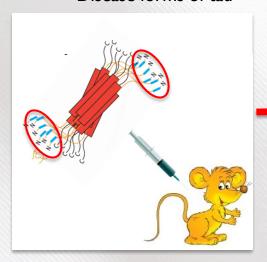


> The All Powerful Antibody: Bench to Bedside

Antibodies

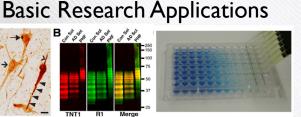
Immunization

Disease forms of tau



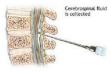






Biomarker/Diagnostic Applications

Cerebrospinal Fluid (CSF)







Therapeutic Applications

Molecular Neurodegeneration (2017) 12:33

Humanized monoclonal antibody armanezumab specific to N-terminus of pathological tau: characterization and therapeutic potency

Michael G. Agadjanyan^{1,2*}, Karen Zagorski^{1,5}, Irina Petrushina², Hayk Davtyan¹, Konstantin Kazarian¹, Maxim Antonenko¹, Joy Davis³, Charles Bon⁴, Mathew Blurton-Jones^{2,3}, David H. Cribbs² and Anahit Ghochikyan¹

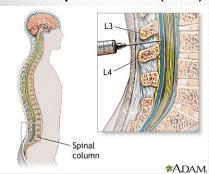




Developing Diagnostic Tools: Biomarkers

Biofluids Hold Significant Promise

Cerebrospinal Fluid (CSF)



Tau is detectable in CSF

- People who convert to AD display rapid annual increases in CSF tau
- Can Toxic Tau Be Useful?

Blood/Plasma - Newer

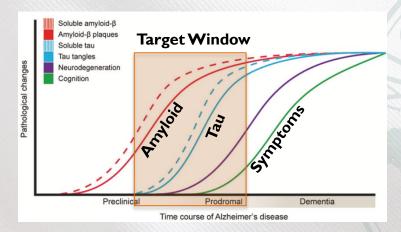


Tau is detectable in plasma

- · Less invasive
- Larger volumes
- Repeated sampling
- · Can Toxic Tau Be Useful?

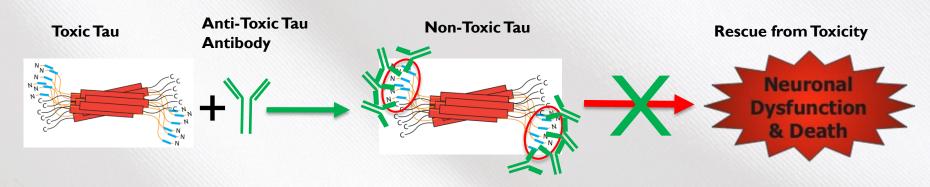
Biomarker Goals

- · Early detection
- Differential diagnosis
- Predictability
- Reproducible, safe, fast & cheap





Targets of Intervention: Blocking Toxic Tau

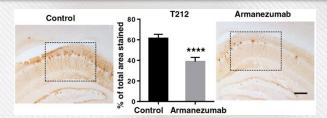


Passive Immunotherapy

Molecular Neurodegeneration (2017) 12:33

Humanized monoclonal antibody armanezumab specific to N-terminus of pathological tau: characterization and therapeutic potency

Michael G. Agadjanyan^{1,2*}, Karen Zagorski^{1,5}, Irina Petrushina², Hayk Davtyan¹, Konstantin Kazarian¹, Maxim Antonenko¹, Joy Davis³, Charles Bon⁴, Mathew Blurton-Jones^{2,3}, David H. Cribbs² and Anahit Ghochikyan¹



College of Human Medicine MICHIGAN STATE UNIVERSITY

Active Vaccination

Active immunization with tau epitope in a mouse model of tauopathy induced strong antibody response together with improvement in short memory and pSer396-tau pathology

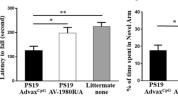
A. Joly-Amado^{a,*,1}, H. Davtyan^{b,c,1}, K. Serraneau^a, P. Jules^a, A. Zitnyar^a, E. Pressman^a, K. Zagorski^b, T. Antonyan^b, A. Hovakimyan^b, H.J. Paek^e, M.N. Gordon^{a,2}, D.H. Cribbs^c, N. Petrovsky^d, M.G. Agadjanyan^b, A. Ghochikyan^{b,3}, D. Morgan^{a,2,3}

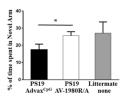
SCIENTIFIC REPORTS | (2019) 9:15455

A MultiTEP platform-based epitope vaccine targeting the phosphatase activating domain (PAD) of tau: therapeutic efficacy in PS19 mice

Armine Hovakimyan¹, Tatevik Antonyan¹, Sepideh Kiani Shabestari², Olga Svystun¹, Gor Chailyan¹, Morgan A. Coburn^{2,1}, William Carlen-Jones², Irina Petrushina⁴, Jean Paul Chadarevian^{2,4}, Karen Zagorski^{1,2}, Nikolai Petrovsky^{0,5}, Spaki H. Cribbs⁴, Michael G. Agadjanyan¹, Anahit Ghochikyan^{1,8} ⁸ Hayk Davtyan^{0,1,4,8} H.







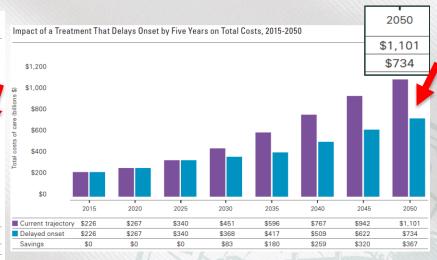
> Research Goals

Delaying disease onset by 5 years will have a huge impact!!!

Projected # of People with AD

Impact of a Treatment That Delays Onset by Five Years on the Number of Americans Age 65 and Older Living with Alzheimer's Disease, 2015-2050 14.0 12.0 10.0 8.0 6.0 2050 4.0 13.5 2.0 7.8 0.0 2015 2020 2025 2030 2035 2040 2045 2050 8.2 9.9 11.6 12.8 13.5 — Current trajectory 5.1 - - Delayed onset 5.8 6.8 5.8 5.9 7.3 7.8 Decrease 0.0 0.0 2.5 4.0 5.0 5.5 5.7

Projected Total Costs for AD





Alzheimer's Disease Research at MSU (in GR)

Department of Translational Neuroscience



Nicholas M. Kanaan, Ph.D. Associate Professor Molecular Mechanisms of AD



Scott E. Counts, Ph.D. Associate Professor Selective vulnerability in AD



Marcia Gordon, Ph.D.
Professor
Role of brain inflammation in AD



Irving E.Vega, Ph.D.
Associate Professor
Novel proteins associated with
tau toxicity in AD



Dave Morgan, Ph.D.
Professor
AD therapeutics and novel biomarkers for disease







Website: http://translationalscience.msu.edu

Alzheimer's Disease: A Look into the Future

Incidence and Impact

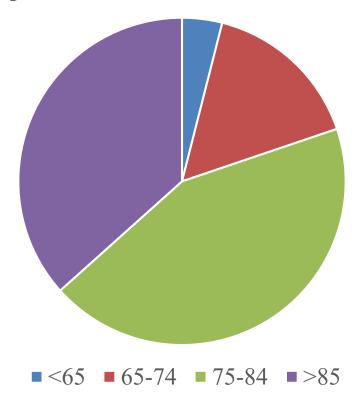
Rebecca Davis, PhD, RN
Professor, Associate Dean for Research
and Practice
Kirkhof College of Nursing





Who does Alzheimer's Disease Affect?

Ages of People with Alzheimer's Disease in the US, 2018

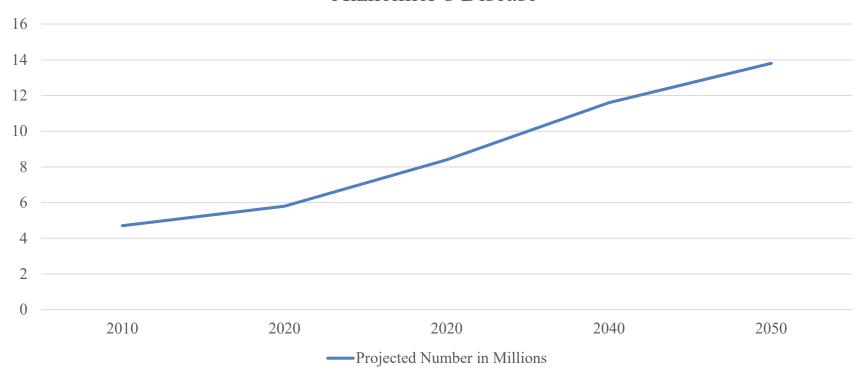


Alzheimer's Association, 2018



Projection into the Future

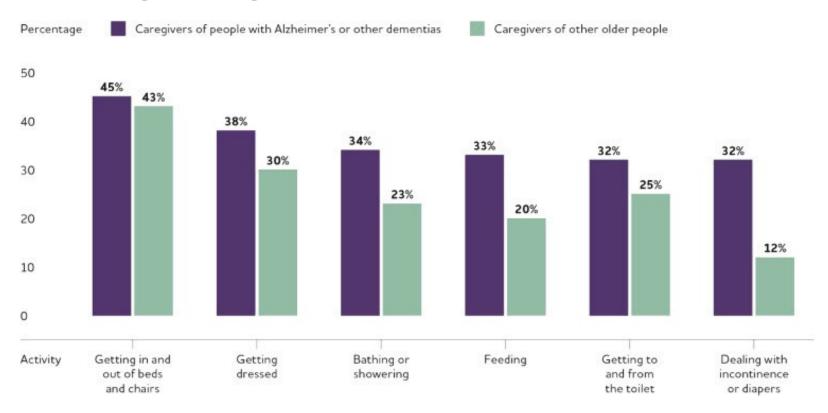
Projected Number of People (over age 65) in US with Alzheimer's Disease



Alzheimer's Association, 2018



Caregiving



Alzheimer's Disease Fact and Figures 2019; Alzheimer's Association



Impact on Families

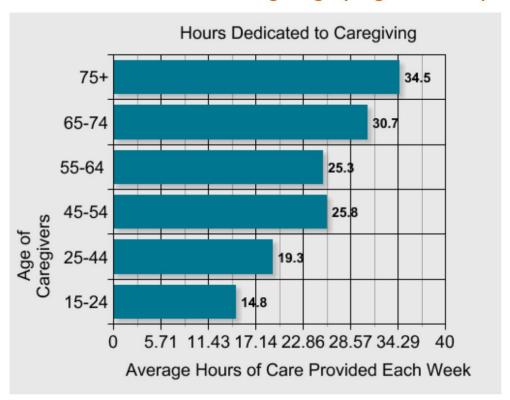
- 18.4 billion hours of care provided by unpaid/family caregivers
 - \$232 billion in care
- Average age of caregiver is almost 70 years
- Most persons with dementia live in the community

Family Caregiver Alliance (2016)



Hours spent caregiving

Number of Hours Dedicated to Caregiving by Age of Family Caregiver



Family Caregiver Alliance (2016)



Caregiving Challenges

Most Challenging Behaviors	Biggest Caregiving Challenges
Agitation/Aggression	Dealing with Memory loss and impact of disease on loved ones
Repetitive speech	Handling stress and emotional toll on self
Wandering	Having patience
Incontinence	Handing mood swings/behaviors
Late day confusion	Keeping loved one positive and motivated

Source: Center to Advance Palliative Care—Gary and Mary West Health Institute National Online Survey: Adults who care for loved ones with Alzheimer's/Dementia.



Decision Making

- Independence/Dependence?
- Driving?
- Living arrangements?
- Medication safety?
- Home safety?
- Health?
- Values?





Research Based Interventions

- Non-pharmacological interventions
 - Functional Based (ABC's)
 - Music therapy
 - Reminiscence
 - Case Management
 - Support Groups
- Pharmacological Therapies

Dyer, S. M., Harrison, S. L., Laver, K., Whitehead, C., & Crotty, M. (2018).



Help for Caregivers

- Acceptance of disease
- Reduce stigma (tell people)
- Seeking support and help
 - Alzheimer's Association
 - Support groups
 - Family/Friends



shutterstock.com · 513803629

• Making a plan (i.e. SHARE intervention)

Gillhooly et al., 2016; Whitlatch et al., 2016



Into the Future

- Earlier, better diagnoses
- More comprehensive care planning
- More people with dementia living alone in the community
- More long-distance caregiving
- "Dementia friendly" communities
- More technological interventions
- Better long term care options



Wayfinding Study





Funded by the National Institute on Aging, National Institutes of Health1R01AG056384-01A1; A Randomized Controlled Trial of Visual Cues, Signage, and Spaced-Retrieval Education within Long Term Care Communities to Assist with Wayfinding



References

Alzheimer's Association. (2018). 2018 Alzheimer's disease facts and figures. *Alzheimer's & Dementia*, 14(3), 367-429.

Family Caregiving Alliance (2016). *Demographics: Statistics*. Retrieved from https://www.caregiver.org/

Gilhooly, K. J., Gilhooly, M. L. M., Sullivan, M. P., McIntyre, A., Wilson, L., Harding, E., ... & Crutch, S. (2016). A meta-review of stress, coping and interventions in dementia and dementia caregiving. *BMC geriatrics*, *16*(1), 106.

Whitlatch, Carol J., et al. "The Support, Health, Activities, Resources, and Education program for early stage dementia: Results from a randomized controlled trial." *Dementia* (2017): 1471301217743033.



Questions?







Thank you for attending the Your Health Lecture Series!