October 7, 2021 AlzeCure Pharma - Introduction & Update

Martin Jönsson - CEO



## Agenda

- Who we are
- Focus areas
- Pipeline & the science
  - Progress & goals in 2021





### AlzeCure Pharma

- ➤ Working in Alzheimer's Disease (AD) and Pain multi-billion markets / great unmet medical need
- Founded in **2016**, out of a research foundation sponsored by Alzheimerfonden
- > Experienced team with extensive background within Pharma industry
- > Based at Novum Science Park, Karolinska Institute, Stockholm, Sweden



- NeuroRestore® A novel first-in-class symptomatic treatment for cognitive disorders, e.g. AD
- Alzstatin® An innovative disease-modifying treatment for AD
- Painless Innovative projects for osteoarthritic and neuropathic pain
- Listed on Nasdaq First North Premier Growth Market since Nov. 2018 (Ticker: ALZCUR)
- Market cap: SEK 302m (October 5, 2021)
- > Cash position: **SEK 78m** (Q2 2021 report)



### Our Business Model

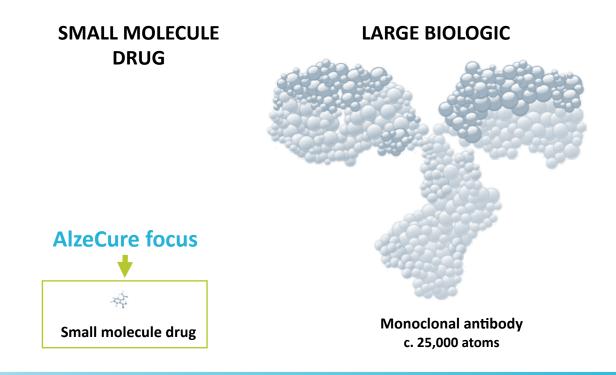
- We are a **Research & Development** company
- Research & develop through early clinical phase and then to out-license or partner on our projects
- Gain incomes through:
  - Upfront payments
  - Milestone payments
  - Royalties on sold products





## Small molecule drugs – AlzeCure's approach for increased success

#### **DIFFERENCES BETWEEN SMALL MOLECULES & BIOLOGICS\***



Smaller molecules can have increased likelihood of penetrating the Blood Brain Barrier



## A pipeline of small-molecule programs

Multiple candidates increase chance of success

Platform	Candidate	Indication	Research phase	Preclinical phase	Phase I	Phase II	Phase III
NeuroRestore®	ACD856	Alzheimer's Disease, Sleep disorders, Traumatic brain injuries Parkinson's disease				Ongoing Fully funded Ph	I study
Neuro	ACD857	Alzheimer's Disease					
Alzstatin®	ACD679	Alzheimer's Disease					
	ACD680	Alzheimer's Disease					
PainLess	ACD440	Neuropathic Pain				Finalized & positive read-out  Preparing for phase II filing	
	TrkA-NAM	Osteoarthritic Pain & other severe pain conditions					



## Our primary Focus area

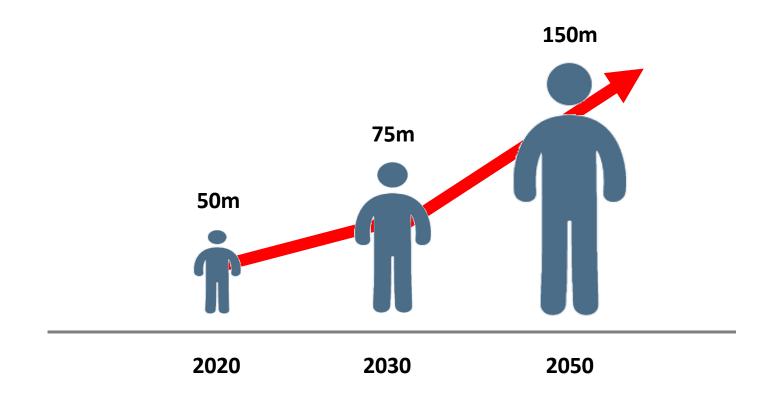
### Alzheimer's Disease

- Costs the society more than oncology and cardiovascular diseases TOGETHER
- The patient population and costs will TRIPLE in the next 30 years





## Tripling patient population - due to the aging population



- 50 million people worldwide live with dementia ...
- ... and doubling every 20 years
- Alzheimer's accounts for 60 70% of all dementia cases



# Alzheimer's Disease - Progressive & lethal disorder with lack of available therapies

**Pre-symptomatic Alzheimer's** *10 - 20 years prior to symptoms* 

**Symptomatic Alzheimer's** 7 - 10 years life expectancy



- In Pre-symptomatic Alzheimer's, Aβ amyloid pathology & plack is building up in the brain but there are no clinical symptoms
- There are no preventive treatments for this stage of Alzheimer's

 The stage includes dying neurons in the brain which leads to speech problems, memory loss and dementia, and symptoms start manifesting







Sales: > US\$ 2bn/year

No drugs available

Very few drugs available - associated with low efficacy and severe side effects

**HUGE UNMET MEDICAL NEED** IN BOTH CATEGORIES





**US** sales

\$17 billion



Aduhelm® projected annual sales if only 5% of the Alzheimer's patients are treated with the product.\*

=> 2<sup>nd</sup> biggest selling product in the US.



## Progression of Alzheimer's Disease



Healthy Brain



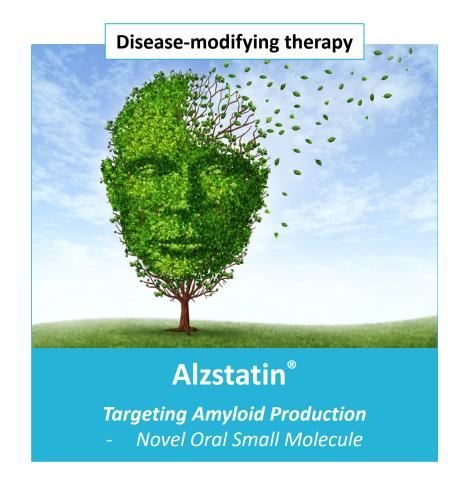
Mild Alzheimer's Disease\*

AlzeCure work to preserve the healthy brain



## Two Alzheimer's platforms - 1st-in-class properties & potential game-changers

#### TARGET TWO KEY AREAS WITH A HIGH UNMET MEDICAL NEED





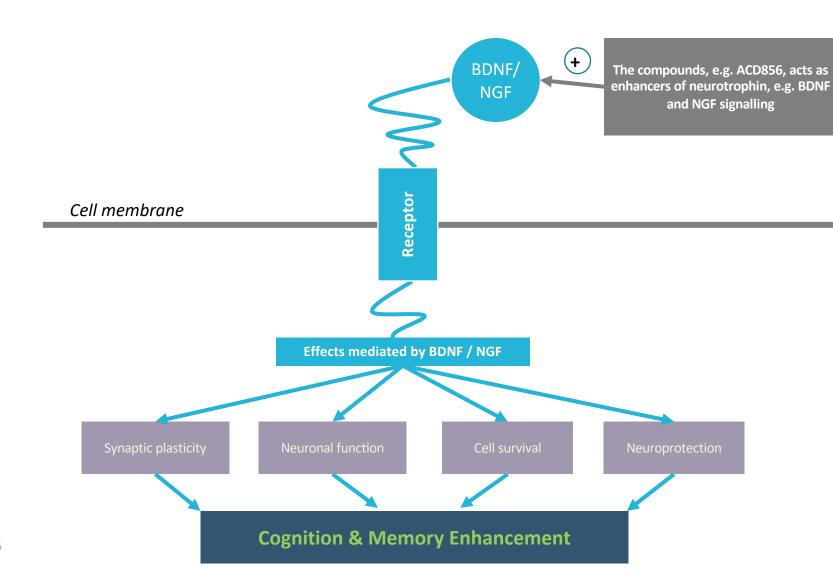


## **NeuroRestore®**

- A cognitive enhancer



### NeuroRestore® enhances neuronal function & cognitive capabilities



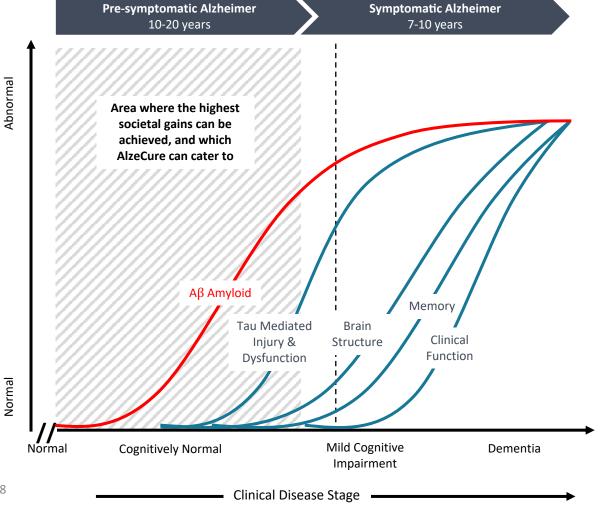
 AlzeCure's compounds act as enhancers of neurotrophin, e.g. BDNF/NGF signalling, and the broad effect profile in this specific biological pathway implies multiple possible indications, including Alzheimer's disease, but also e.g. Parkinson's disease, Traumatic Brain injury and Sleep disorders.



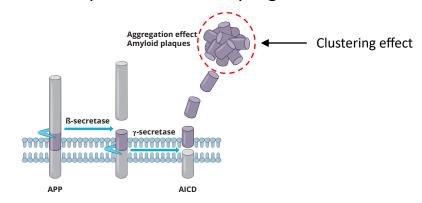


### Alzheimer's Disease Modifier – Preventing or delay disease progression

#### **ALZHEIMER'S DISEASE PROGRESSION**

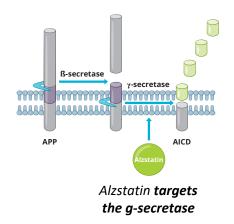


Ab-42 - main culprit in Alzheimer progression



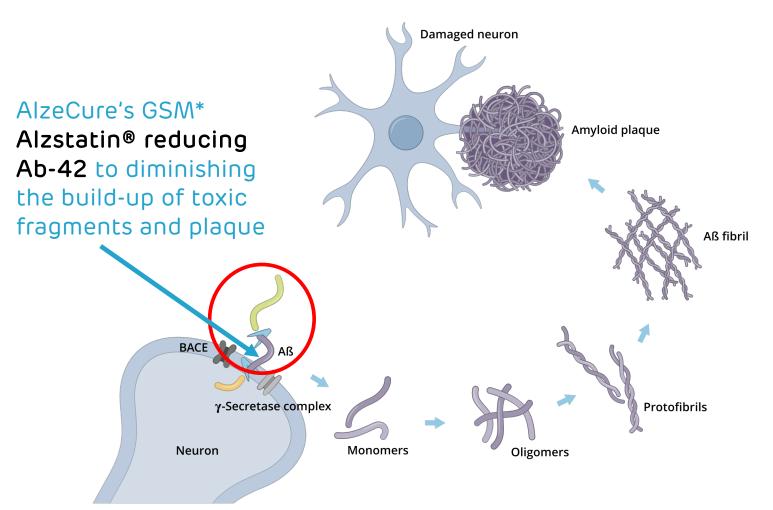
found a way to limit Ab-42 clustering

## Alzstatin modulates the enzyme and thereby limits the clustering effect





# The Amyloid cascade - Generating toxic and damaging fragments, including plaques, damaging neurons and brain structures







### Our second Focus area

### **Chronic Pain**

- Suicide due to chronic pain is as common as due to depression
- Most common cause for sick leaves, creating misery & high societal costs
- Opioid crisis in the US is huge & reversing the mean average lifespan of Americans



**Huge need** for more efficacious and safer treatments



## Our platform PAINLESS - Targeting unmet medical needs within pain



**Neuropathic pain\*** 

25 – 30 million patients

Project: ACD440



Osteoarthritis & severe pain conditions

> 300 million patients

Project: TrkA-NAM



## TrkA-NAM - none-opioid treatment of severe pain conditions

Attractive Target population

E.g. **osteoarthritis** in patients who have experienced insufficient pain relief from 1st line standard of care or unacceptable side effects

Clinical validation

Mechanism with **strong validation** – step-changing clinical efficacy with novel anti-NGF mAbs - sets the standard for future therapies

Blockbuster opportunities

**Blockbuster opportunities** for NGF therapies that would avoid adverse events and allow non parenteral routes of administration



#### Differentiation factors for TrkA-NAM

- TrkA selective MoA vs anti-NGF antibodies also targeting p75 signaling
  - Maintain potent clinical efficacy
  - > Improved side-effect profile
- > Convenient oral administration small molecule compound
- > No addiction compared to opioids





### **Professor David Julius**

Nobel prize medicine laureate 2021

University of California, San Francisco, USA



Prize motivation: ... for the discoveries of receptor (TRPV1) for temperature and touch.

The identification of and knowledge about the TRPV1 receptor is central for the mediation of neuropathic pain, which 7-8% of the adult population is affected by. AlzeCure Pharma is developing a TRPV1 antagonist (ACD440), which now is in clinical phase, for treatment of this pain condition.



## ACD440 - Novel TRPV1 antagonist in clinical phase for neuropathic pain



#### **PROJECT OVERVIEW**

Emanates from Big Pharma

- Approximately SEK 200m spent on project development
- > Mode of action confirmed in several Phase 1 clinical trials
- Synthesized compound and formulation developed



VR1 – optimized for local delivery

- > The vanilloid receptor subtype 1 (TRPV1) is expressed in nociceptive sensory neurons
- > TRPV1 is upregulated in the skin of patients with neuropathic pain
- > Strong scientific support for peripheral/local treatment with TRPV1 antagonists

Phase 1b trial fully funded

- The trial was fully funded with existing funds
- > Trial with topical formulation has now ended
- > Phase 1b study addressed both tolerability & efficacy POSITIVE OUTCOMES

Now preparing filling for a phase II clinical trial



## Neuropathic pain - Fast growing market

- The most valuable segment within the pain indications
- Poorly served patients
- Huge demand for better drugs

2020 **\$11 billions** 

CARG to 2027 12.9% => **\$25 billions** 

The Neuropathic Pain market was valued at \$10,8 billion in 2020 globally and is forecast to reach \$25,2 billions by 2027, at a Compound Annual Growth Rate (CARG) of 12,9%



### Key short-term milestones for 2021



Read-out of clinical 1b study Painless ACD440 in summer 2021



**Read-out** of clinical 1a SAD study **NeuroRestore**® **ACD856** summer 2021



Start NeuroRestore® ACD856 1a MAD clinical study in second half 2021



Select CD for our pain project Painless TrkA-NAM in second half 2021



File phase 2a clinical study for Painless ACD440 second half 2021



## Key investment highlights in AlzeCure



Targeting areas of huge unmet medical needs



Strong team with extensive experience and track record



Platforms with first-in-class properties and potential game-changers



Parallel investments in several candidates and potent follow-up programs



Multi-billion dollar market opportunities



Evolving into a phase II company





