

# Addex Pharmaceuticals

Biotech Showcase January 11, 2011



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## Vision

Goal: allosteric modulators for human health

How: proprietary discovery platform

Focus: CNS, metabolic disorders & inflammation



## Financials & Stock

- Cash through early 2013
  - CHF56.7 (US\$54/€42) million in cash as of June 30
  - CHF20 (\$20) million raised on Sep 14 from BVF
  - > \$900,000 grant from The Michael J. Fox Foundation on Sep 8
- Market cap (03 Jan): CHF64 (€51 / US\$68) million
- Symbol on SIX Swiss Exchange: <u>ADXN</u> (ISIN:CH0029850754)
- 7,835,878 shares outstanding
- Five analysts covering:

Jefferies Peter Welford & Philippa Gardner

Helvea Olav Zilian

Bank Vontobel Andrew C. Weiss & Silvia Schanz

Bank am Bellevue Bob Pooler

Edison Robin Davison

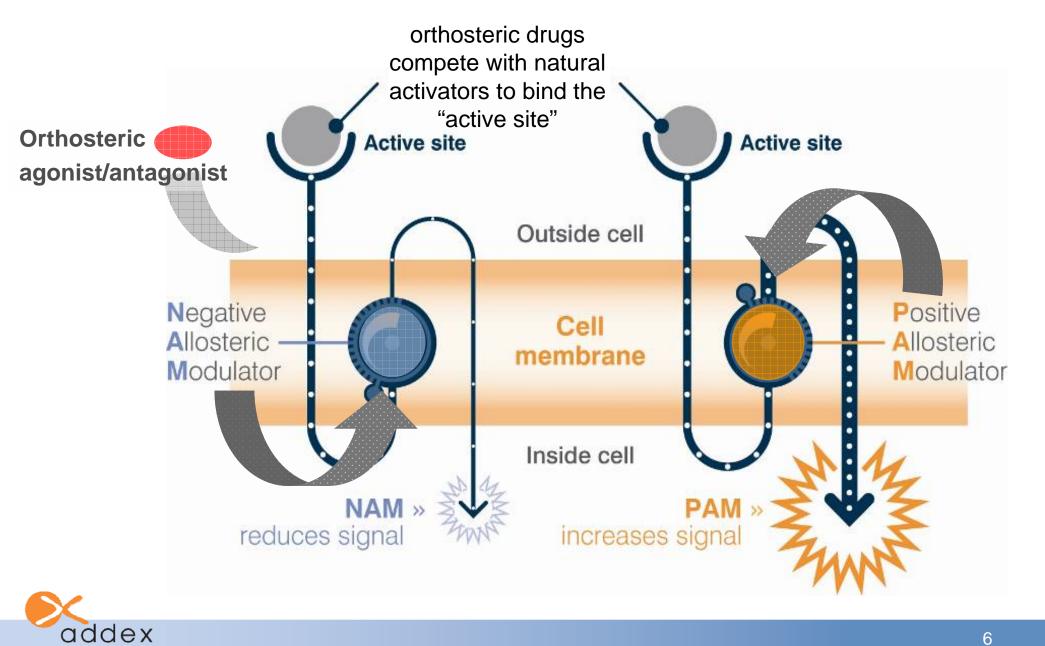


## raison d'être

- New therapeutic classes have created or promise huge markets
  - Protein / antibody / peptide therapeutics
  - Gene therapy / siRNA
  - Allosteric modulation
- Allosteric modulators (AM) are an emerging therapeutic class
  - Different from traditional "orthosteric" drugs
    - AM bind to different sites on cell surface receptors
    - AM generally are structurally different from orthosteric drugs
  - Modulatory not binary
    - Modulatory: like a dimmer switch not an on/off switch
    - Positive allosteric modulators (PAM) increase activity of cell surface receptors
    - Negative allosteric modulators (NAM) inhibit receptor activity
  - AM are proven drugs
    - Sensipar/Mimpra cinacalcet (Amgen/NPS) is a PAM of CaSR
    - Selzentry/Celsentri maraviroc (Pfizer) is a NAM of CCR5
    - But <u>AM are hard to find</u> with classical tools!

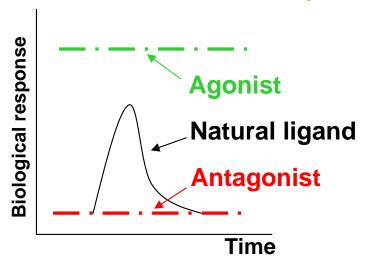


## Allosteric Modulation



# Allosteric Advantages

#### **Orthosterics are steady state**



- Better specificity/selectivity for target
  - e.g. mGluRs
- Can target receptors considered intractable for small molecules
  - e.g. GLP-1 and TNF
- PAM + natural ligand

  Natural ligand

  NAM + natural ligand

Time

Allostery preserves natural rhythm

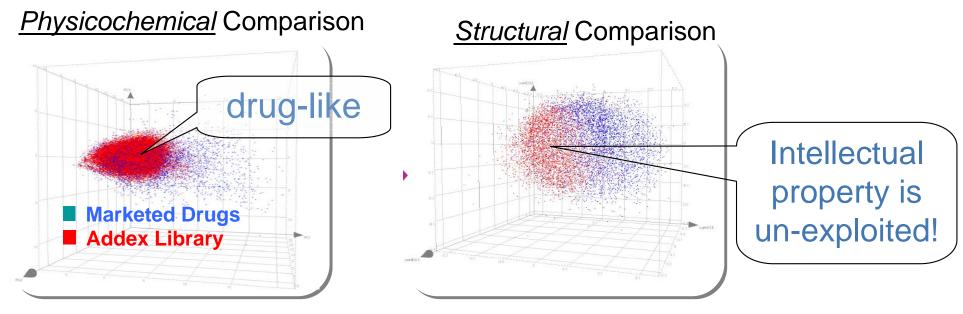
- Acts like a dimmer not "on/off" switch
  - better control = better drugs





## **Platform**

70,000+ compound allostery-biased library



- Proprietary high throughput screening tools
  - ProxyLite
  - Phoenyx
  - AddeLite



## **PIPELINE**

Partner	Molecule / Mechanism	Assay Development & Screening	Hit-to- Lead	Lead Optimization	Preclinical	Phase I	Phase II	Milestone
	ADX48621 mGluR5 NAM	Parkinson's		evodopa In			LID)	Start PhII 1Q11
		Dystonia						Start Phlla 2H11
Ortho-McNeil- Janssen	ADX71149 mGluR2 PAM	Schizophre	enia	funded &	& developed	by OMJPI*		Start Phlla 1Q11
		Anxiety		funded &	& developed l	oy OMJPI*		
	ADX68692 FSHR NAM	Endometric	osis					
	<b>ADX71943</b> GABA-B PAM	Osteoarthr	itic Pain					
Merck & Co.	ADX63365 mGluR5 PAM	Schizophre		developed by I	Merck			



**NAM** = negative allosteric modulator (an inhibitor) **PAM** = positive allosteric modulator (an activator)

<sup>&</sup>lt;sup>‡</sup> and undisclosed additional indications

<sup>\*</sup>Ortho-McNeil-Janssen Pharmaceuticals, Inc., a Johnson & Johnson subsidiary

## DISCOVERY

Partner	Molecule / Mechanism	Assay Development & Screening	Hit-to- Lead	Lead Optimization	Preclinical	Phase I	Phase II	Milestone
	mGluR2 NAM	Alzheimer's	s / Depress	ion				
Merck & Co.	mGluR4 PAM	Parkinson'	s Disease ‡ funded	d by Merck				CNC
	mGluR7 NAM	Depression Post Traun	n natic Stress	Disorder				CNS
	Orexin 2R NAM	Sleep Diso	orders					
	GLP1 PAM	Type II Dia	betes		D.A.	atabali	o Dios	w d o w o
	GIPR PAM	Type II Dia	betes		IVI	etaboli	ic Disc	raers
	TNFR1 NAM (CD120a)		i <mark>d Arthr</mark> itis, I s, Multiple S	Psoriasis, In Sclerosis	flammatory	Bowel Dise	ease	
	A2A PAM	Psoriasis,	Osteoarthrit	is		Ir	ıflamı	nation
	IL1R1 NAM (CD121a)	Gout, Type	II Diabetes	5				



**NAM** = negative allosteric modulator (an inhibitor) **PAM** = positive allosteric modulator (an activator)

<sup>‡</sup> and undisclosed additional indications

## Platform Performance

- Addex has received partnering revenue every year since 2004
- Cash inflows generated to date: CHF44 (US\$45) million
- All three partnerships are fully funded by our partners
- Addex is eligible for up to about \$1 billion in milestones plus royalties

Summary of Partnerships									
Partner	Product	Indication(s)	Status at signing	Upfront Cash	Revenues to date	Total Milestones	Royalty		
Ortho-McNeil- Janssen	mGluR2 PAM ADX71149	Anxiety & schizophrenia*	Hit-to-Lead (Dec 2004)	€3	€5.2	€112	low double-digit		
Merck & Co., Inc.	mGluR4 PAM	Parkinson's disease*	Hit-to-Lead (Dec 2007)	\$3	\$2.5	\$167.5	ND		
Merck & Co., Inc.	mGluR5 PAM ADX63365	Schizophrenia*	Clinical Candidate (Jan 2008)	\$22	-	\$680	ND		





# Partnering Priorities

- mGluR5 NAM (ADX48621 & backups)
  - PD-LID / dystonia depression
  - fragile X / autismGERD
  - anxietypain
- FSH receptor NAM (ADX68692)
  - endometriosis
  - prostate cancer
- mGluR2 NAM
  - Alzheimer's disease
  - depression



### allosteric modulators for human health

www.addexpharma.com

