Alzheimer's Disease Advances: Leveraging the San Diego Environment

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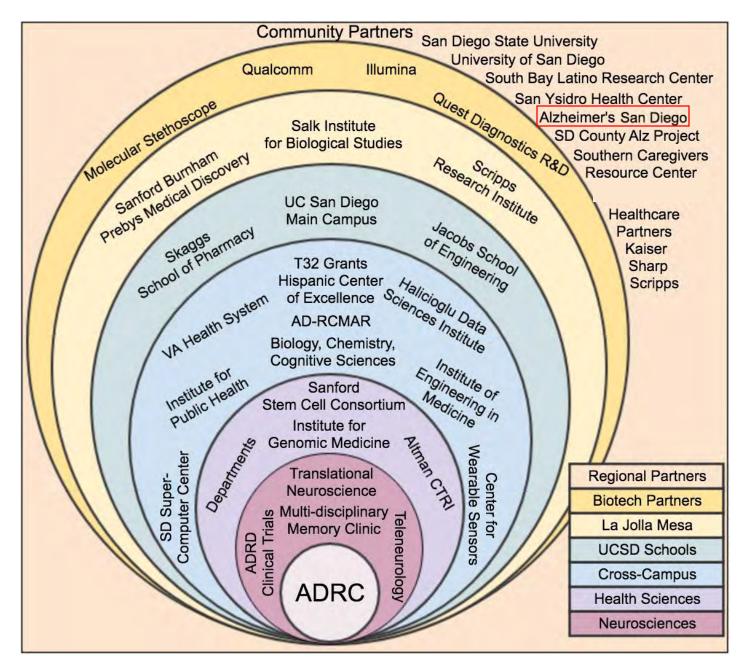
Director, Shiley-Marcos Alzheimer's Disease Research Center
Professor and Chair, Department of Neurosciences

UC San Diego

Strong Scientific Environment



Broad Collaborations



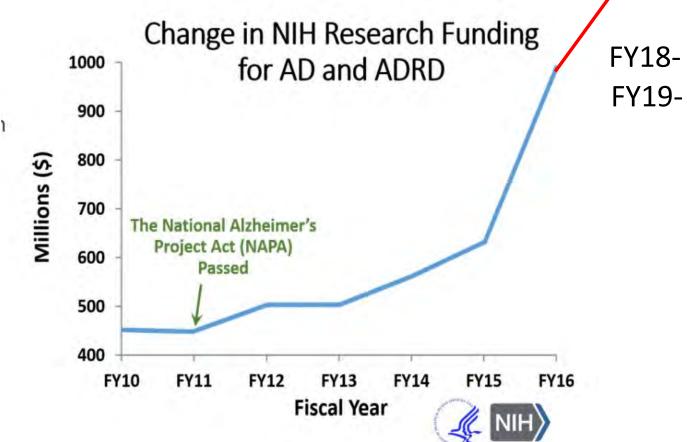
ADRC Team



NAPA

SECOND ALZHEIMER'S PROFESSIONAL JUDGMENT BUDGET REFLECTS URGENT NEED TO FIND A CURE OR TREATMENT

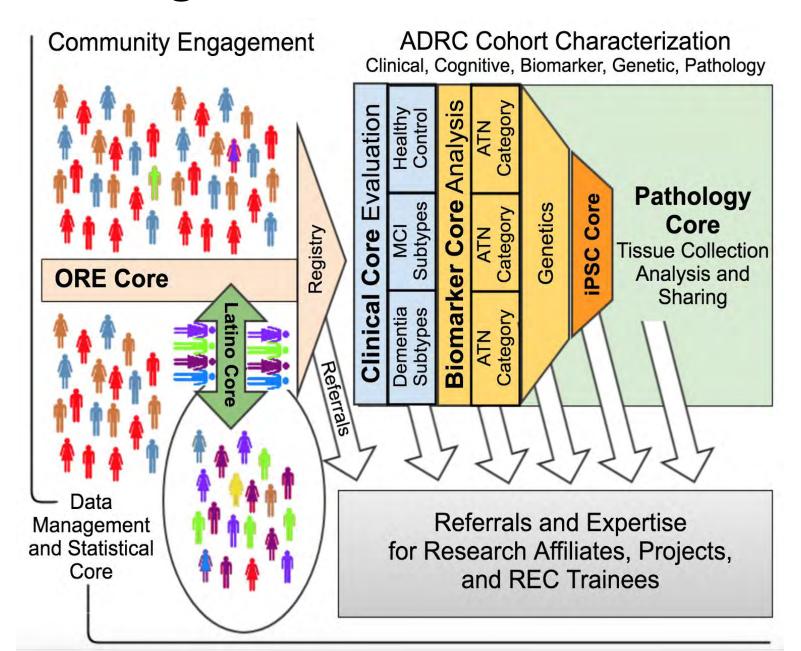
NIH Recommends \$414 Million Increase in Fiscal Year 2018 Research Funding For Our Nation's Most Expensive Disease



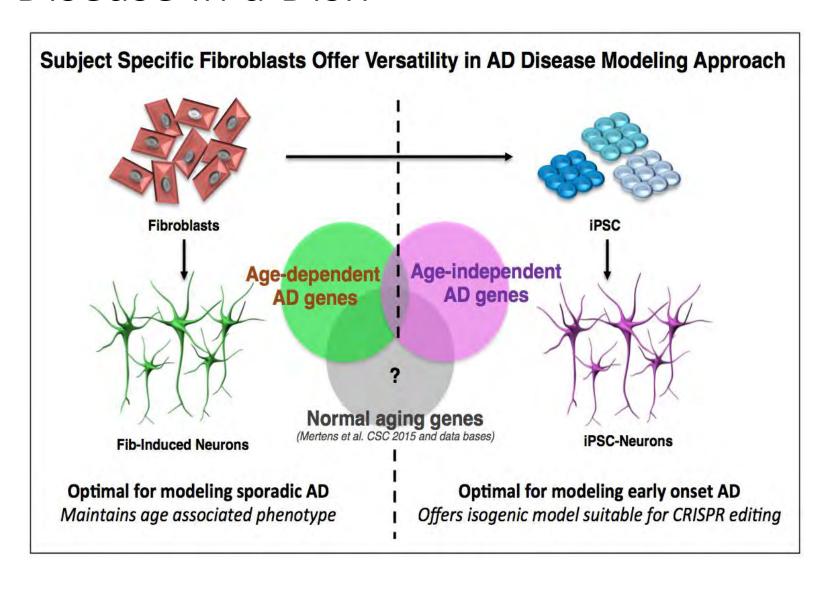
FY18- 1.9 Billion

FY19- 2.3 Billion

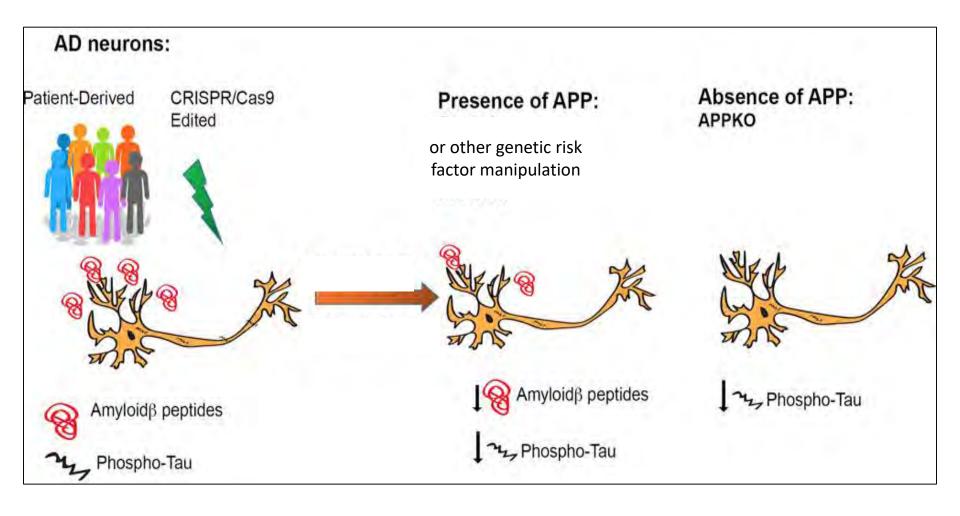
Fostering ADRD Science



Induced Pluripotent Stem Cells "Disease in a Dish"



Induced Pluripotent Stem Cells "Disease in a Dish"



Combination of Genes with Greatest Effect on 'Sporadic' AD Age of Onset

	Chr	Position	Gene	IGAP marginal p in –log ₁₀	β	(95% CI)	Conditional p in –log ₁₀	1								
ε2 allele	19		APOE		-0.45	(-0.55,-0.35)	> 17	IKI	Time .	The state of the s				1-1-01/0	-11	
ε4 allele	19		APOE		0.98	(0.94,1.01)	> 17		Control of the last	The state of	The state of			1st PHS Qua		
rs61822977	1	207796065	CR1	7.1	-0.12	(-0.16, -0.09)	10.5	0.9 -			3			2nd PHS Qu		
rs6733839	2	127892810	BIN1	43.2	-0.16	(-0.20, -0.12)	16.1							3rd PHS Qua		
rs73048831	5	14721590	ANKH	5.5	-0.12	(-0.19, -0.05)	3.4							4th PHS Qua		
rs2282802	5	139712239	HBEGF	5.2	-0.07	(-0.10, -0.03)	3.4	0.8	£					Uncorrected	ADGC base	eline
rs11645714 6	6	32578632	HLA-DRB1	7.1	0.09	(0.05, 0.13)	4.2			- Jal		4	4			
rs14528102 5	6	32674252	HLA-DQA2	6.4	-0.10	(-0.16, -0.05)	3.4	O 0.7 -				1 Ro				
rs9394766	6	41159801	TREML2	5.9	0.08	(0.04, 0.12)	3.9				\ <u> </u>	17	-			
rs9296567	6	47553402	CD2AP	9.5	-0.08	(-0.12, -0.04)	4.2	.⊆ 0.6 -		į:	1 5	(E)	¥	¥-1		
rs6957573	7	37611912	BC043356	5.7	-0.07	(-0.10, -0.03)	3.5	no		٠.		E	1			
rs1859788	7	99971834	PILRA	9.2	0.11	(0.07, 0.15)	7.3	Ę				F1 -3	1			
rs78571833	7	143122924	EPHA1-AS1	6.4	0.15	(0.09, 0.22)	5.3	Proportion 5.0					1 1			
rs17057043	8	27220310	PTK2B	11	-0.08	(-0.12, -0.04)	4.3	0					1		ر-ني	
rs2741342	8	27330096	CHRNA2	7.4	0.08	(0.03, 0.12)	3.1	4			-	1.17	1.3	/		
rs1532277	8	27466181	CLU	22.9	0.08	(0.04, 0.12)	4.6	₹ 0.4 F			!=\			77	1	
rs12358692	10	11721102	LOC105376412	5.3	0.06	(0.02, 0.10)	2.8	Survival - 6.0			17		3	- T-		
rs3740688	11	47380340	SPI1	7.8	0.07	(0.04, 0.11)	4.3	200				\		1		in .
rs1286170	11	59831930	MS4A3	5.9	-0.07	(-0.12, -0.03)	2.7	S 0.3				1	ing	1		
rs611418	11	60011013	MS4A4E	12.3	0.12	(0.08, 0.17)	6.7					1	\		\	-
rs471470	11	85831541	PICALM	23.5	0.09	(0.05, 0.13)	5.6	0.2 -					t!	1		
rs1952090	14	53342992	FERMT2	6.3	-0.10	(-0.16, -0.05)	3.6	0.2					.,		Z	
rs12590273	14	92934120	SLC24A4	7.2	0.12	(0.07, 0.17)	6					-			V	L
rs7145100	14	107160690	abParts	6.1	0.10	(0.05, 0.15)	3.7	0.1						1		
rs2526378	17	56404349	BZRAP1-AS1	6.1	0.08	(0.04, 0.11)	4.6	0.1					-			140
rs3795065	19	1039444	ABCA7	8.8	-0.06	(-0.10, -0.02)	2.3							==-	i	-
rs4147916	19	1057137	ABCA7	5.1	0.26	(0.15, 0.36)	6	0	1	i	1	i	i			227
rs2074442	19	1074000	HMHA1	8.7	-0.07	(-0.11, -0.03)	3	60	65	70	75	80	85	90	95	10
rs6024870	20	54997568	CASS4	7.1	0.12	(0.05, 0.18)	3.3				255	Age	24.5	- 54	17.5	
rs2092563	22	41592677	EP300-AS1	5.1	-0.06	(-0.10, -0.02)	2.5					1.90				

ADGC – 28 SNPs from Cox forward regression model