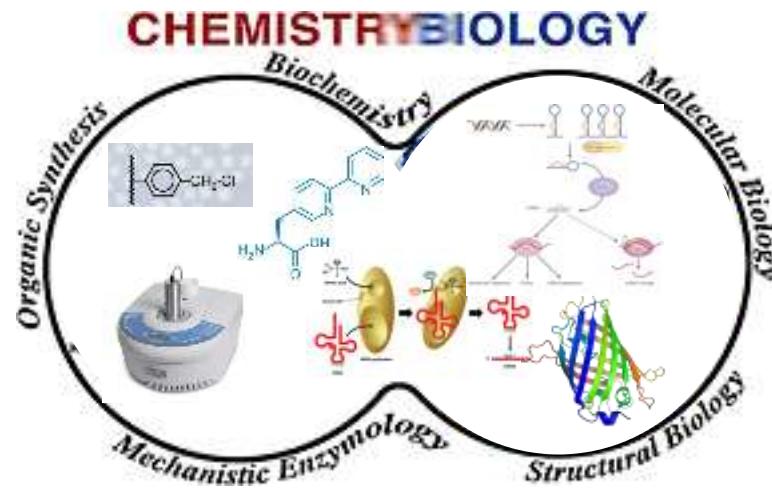




DOUGTON ABBEY

Research in the Young Lab

September 8, 2017



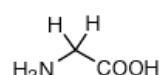
W&M

Unnatural Amino Acids

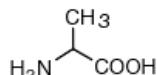
[New UAAs and Application Towards Biological Problems]

The Genetic Code

Small

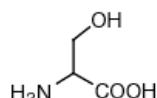


MW: 57.05

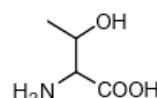


MW: 71.09

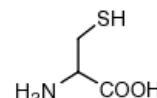
Nucleophilic



MW: 87.08, $pK_a \sim 16$

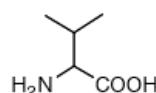


MW: 101.11, $pK_a \sim 16$

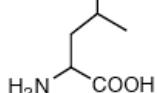


MW: 103.15, $pK_a = 8.35$

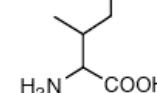
Hydrophobic



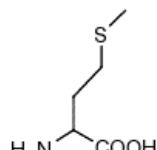
MW: 99.14



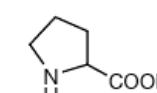
MW: 113.16



MW: 113.16

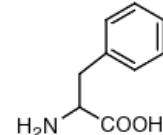


MW: 131.19

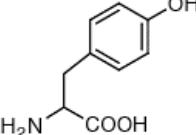


MW: 97.12

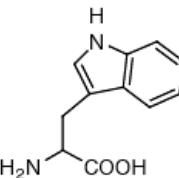
Aromatic



MW: 147.18

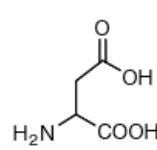


MW: 163.18

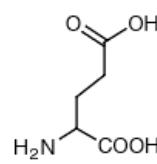


MW: 186.21

Acidic

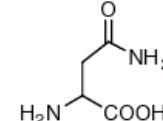


MW: 115.09, $pK_a = 3.9$

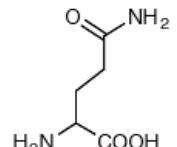


MW: 129.12, $pK_a = 4.07$

Amide

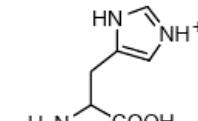


MW: 114.11

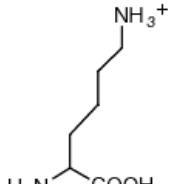


MW: 128.14

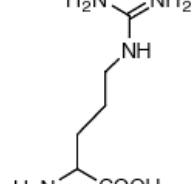
Basic



MW: 137.14, $pK_a = 6.04$



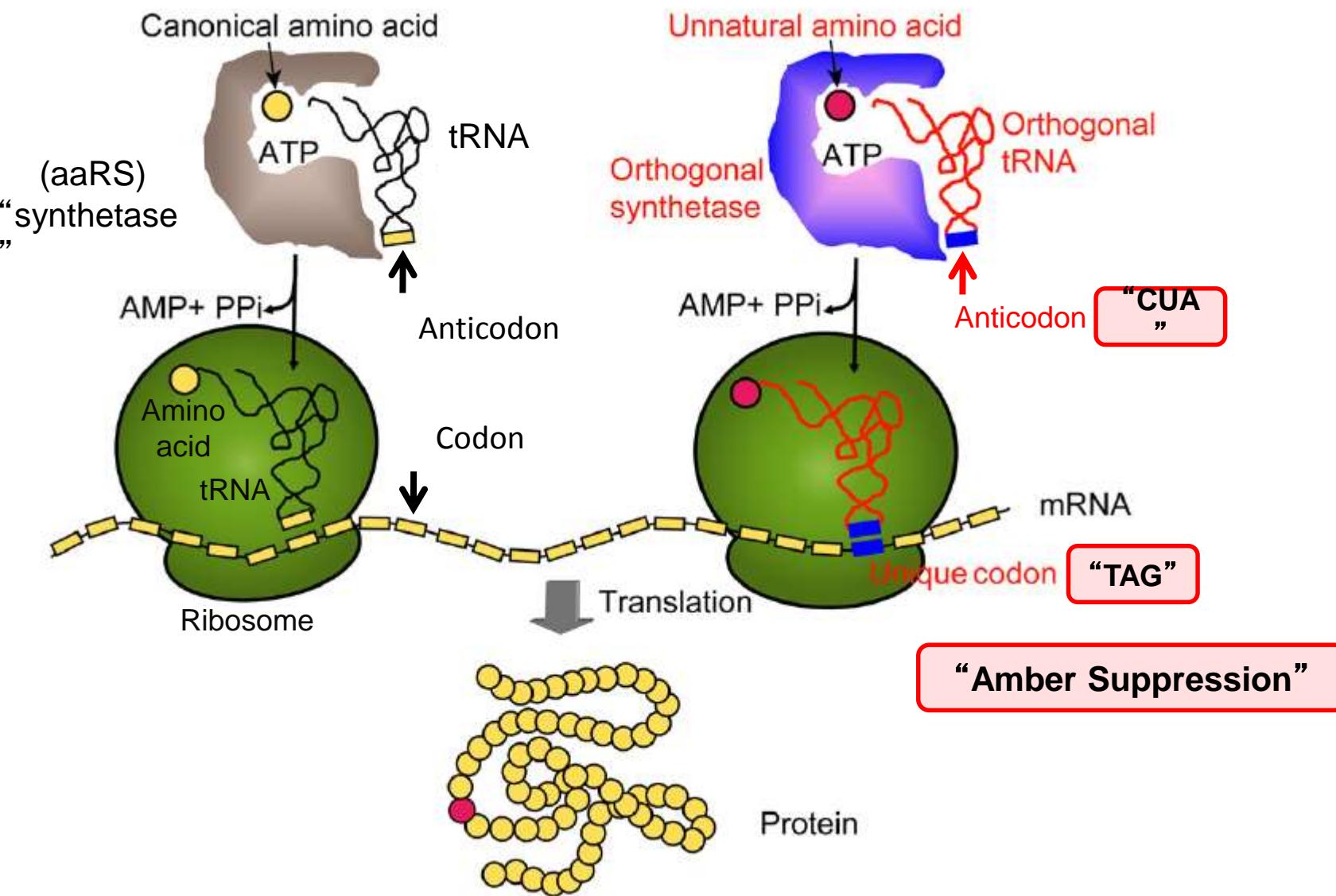
MW: 128.17, $pK_a = 10.79$



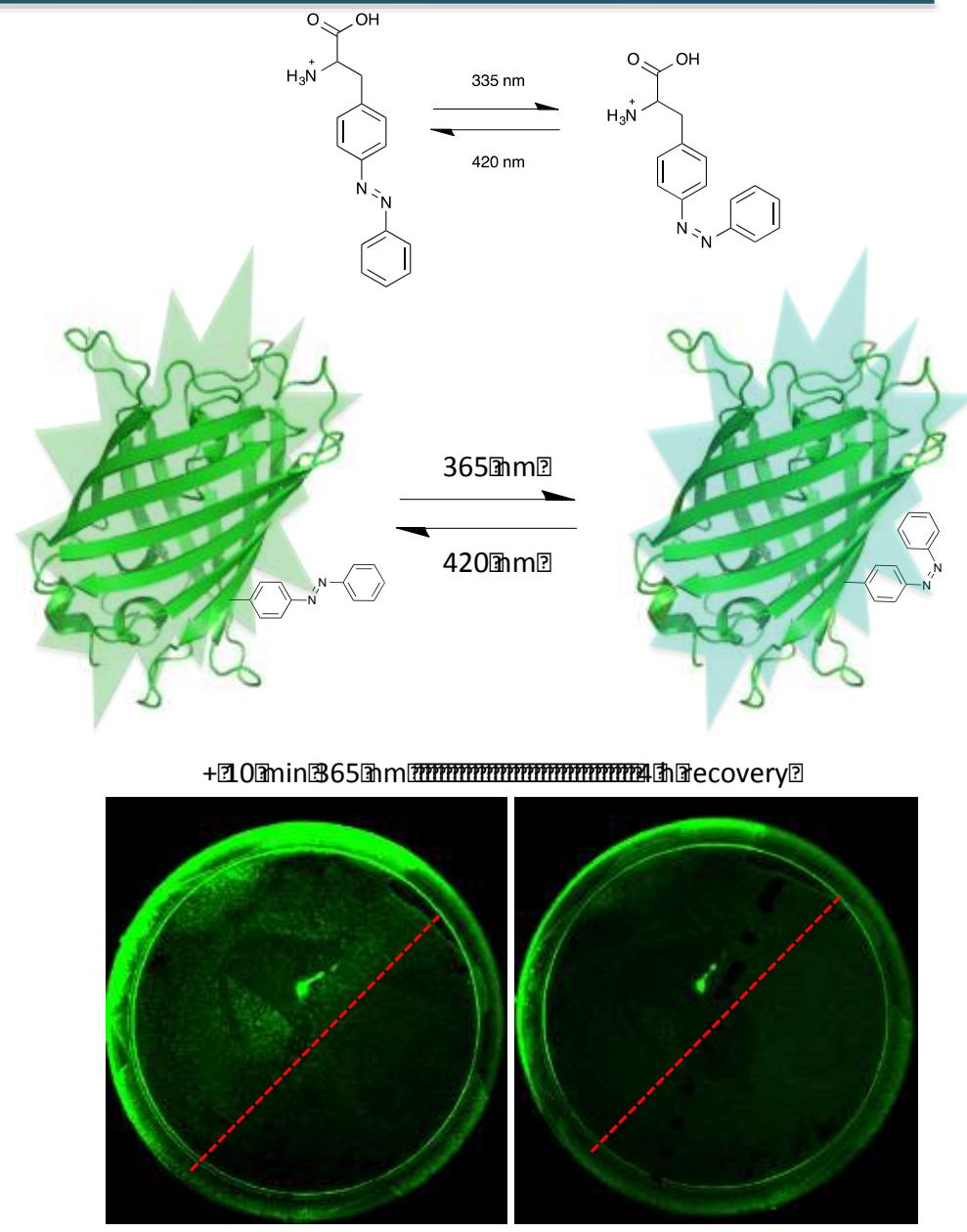
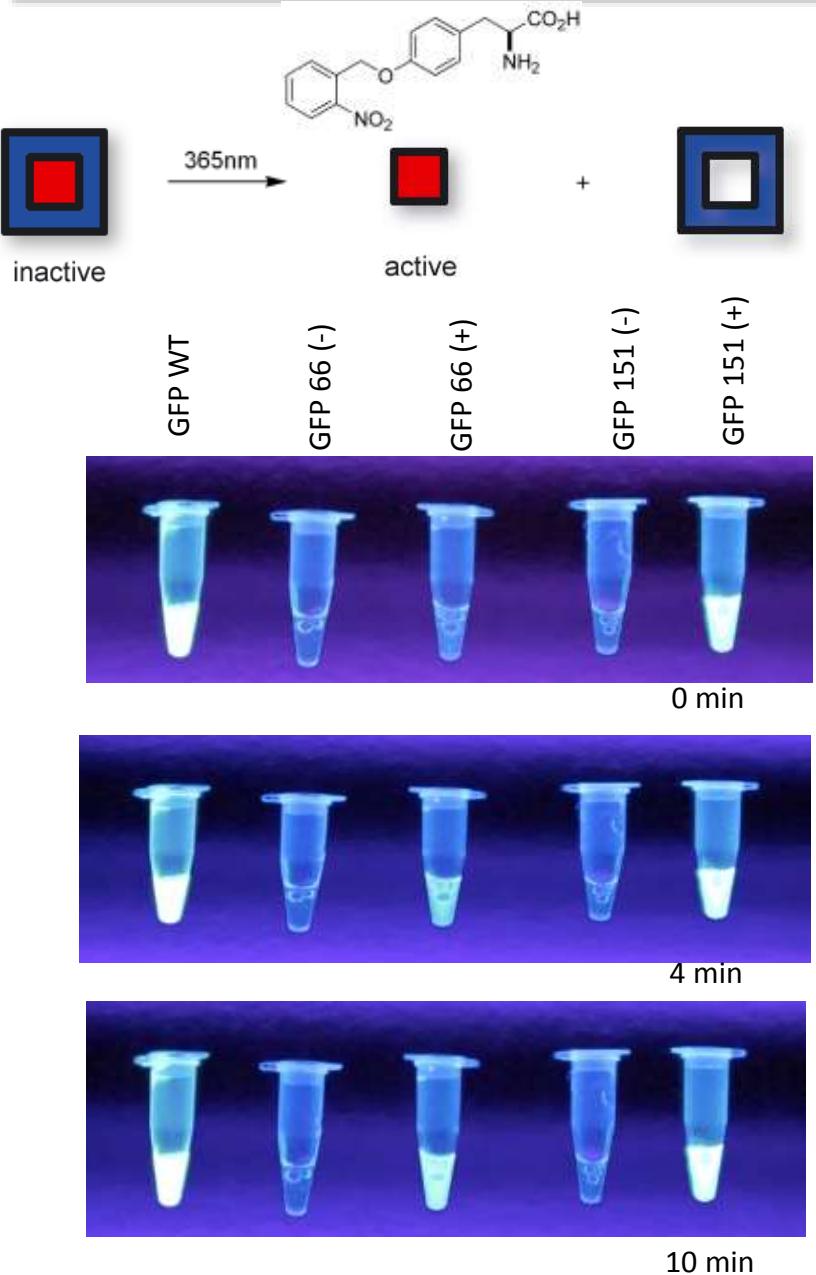
MW: 156.19, $pK_a = 12.48$

			Second Letter	
	T	C	A	G
First Letter	TTT TTC TTA TTG } Phe	TCT TCC TCA TCG } Ser	TAT TAC TAA TAG } Tyr Stop	TGT TGC TGA TGG } Cys Stop Trp
C	CTT CTC CTA CTG } Leu	CCT CCC CCA CCG } Pro	CAT CAC CAA CAG } His Gln	CGT CGC CGA CGG } Arg
A	ATT ATC ATA ATG } Ile Met	ACT ACC ACA ACG } Thr	AAT AAC AAA AAG } Asn Lys	AGT AGC AGA AGG } Ser Arg
G	GTT GTC GTA GTG } Val	GCT GCC GCA GCG } Ala	GAT GAC GAA GAG } Asp Glu	GGT GGC GGA GGG } Gly
	T C A G	T C A G	T C A G	T C A G
Third Letter				

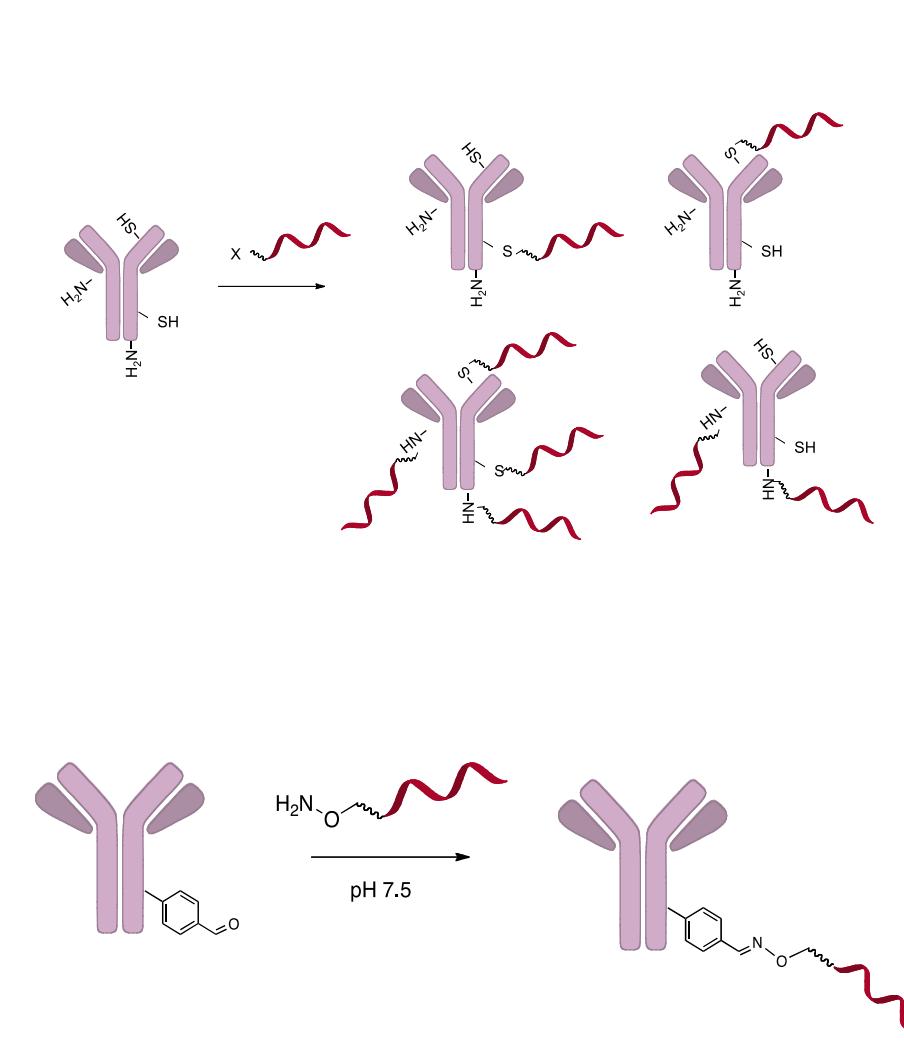
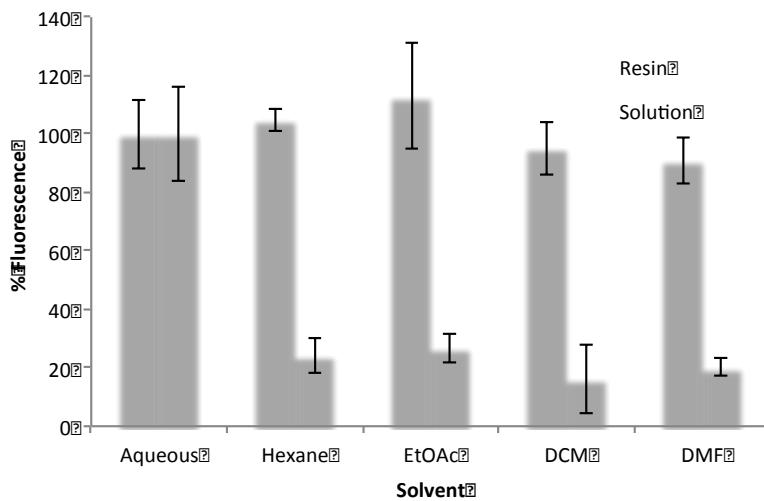
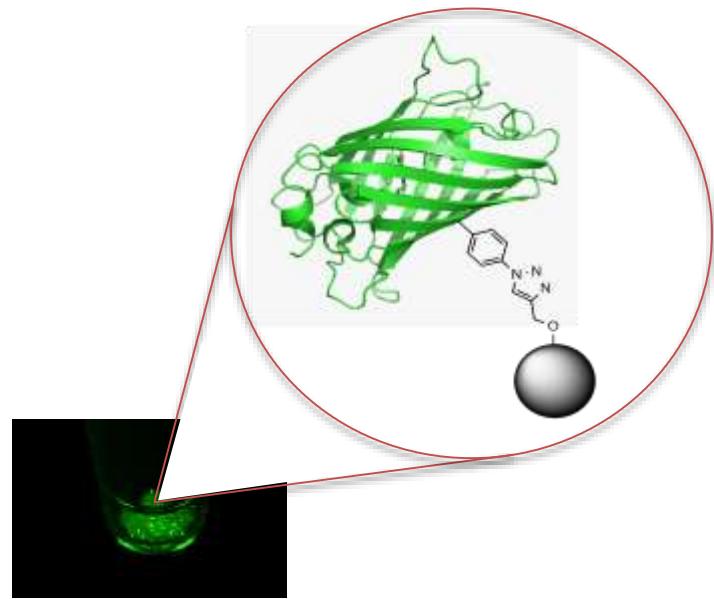
aaRS/tRNA Pairs for Amber Suppression



Photoregulated Protein Function



Protein Immobilization



Glaser Hay Reactions

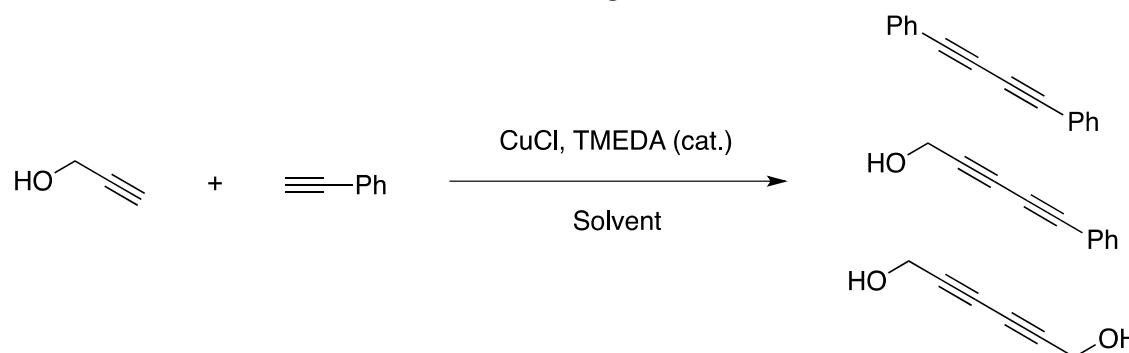
[Developing New Methodologies]

Development of Small Molecule Libraries

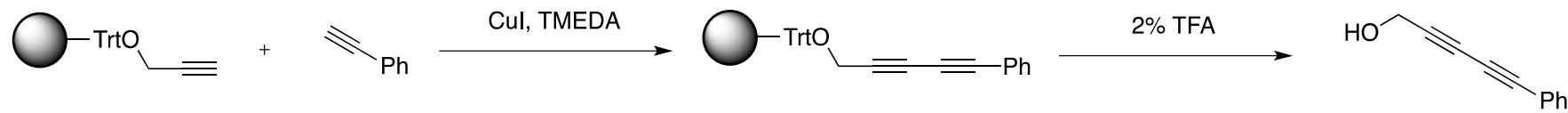
Synthetic Issues

Lack of chemoselectivity

12 - 48 h

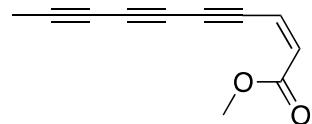


Solution



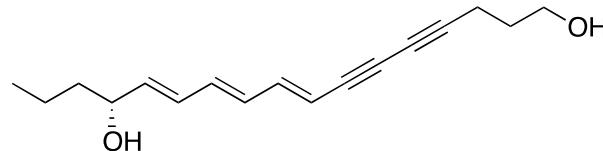
Development of Small Molecule Libraries

Polyyne Cores as Therapeutics



Dehydromatricaria ester

1826, *Artemisia*



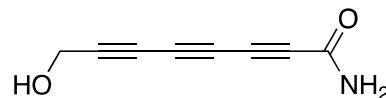
Circutoxin

Convulsions, respiratory paralysis



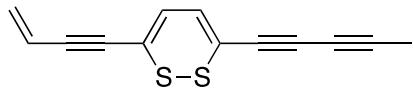
Monitporyne A

Cytotoxic to solid tumor cell lines



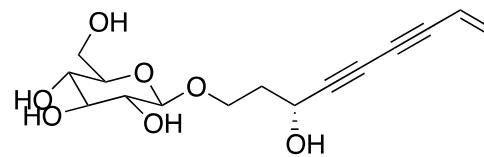
Agrocybin

Anti-fungal



Thiaurbine B

Anti-parasite

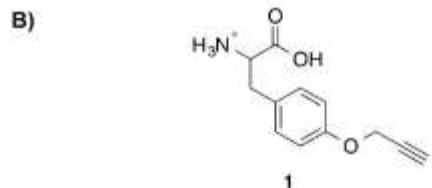
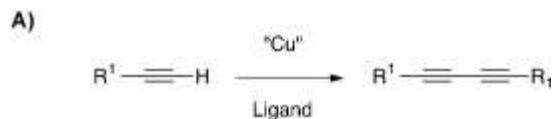


Bidensyneoside A₂

Anti-histamine
Inhibit NO formation

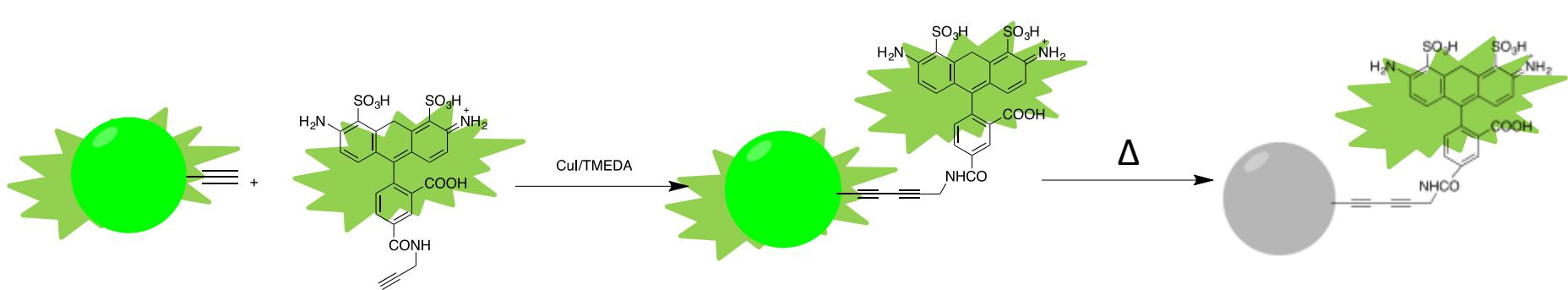
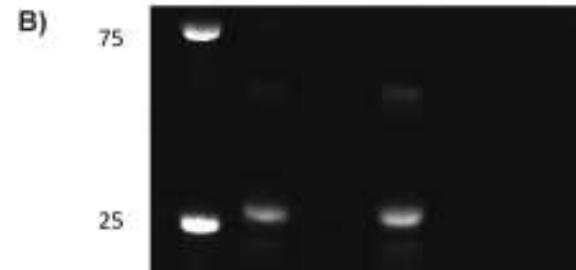
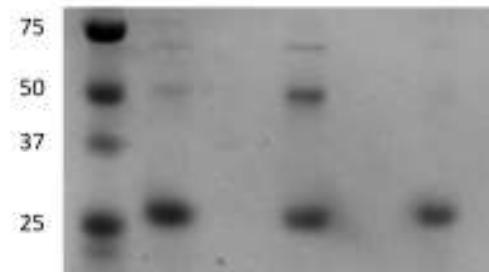
Translation to Biological Systems

Using UAAs as for a bioorthogonal reaction

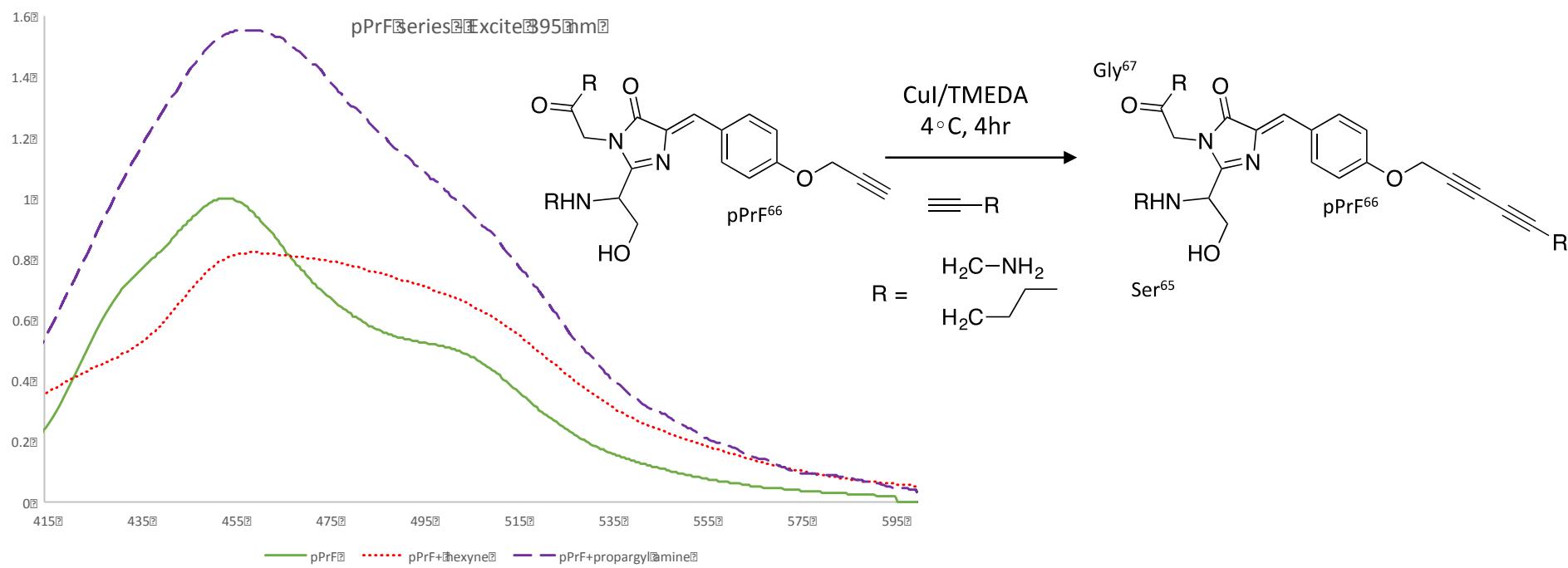
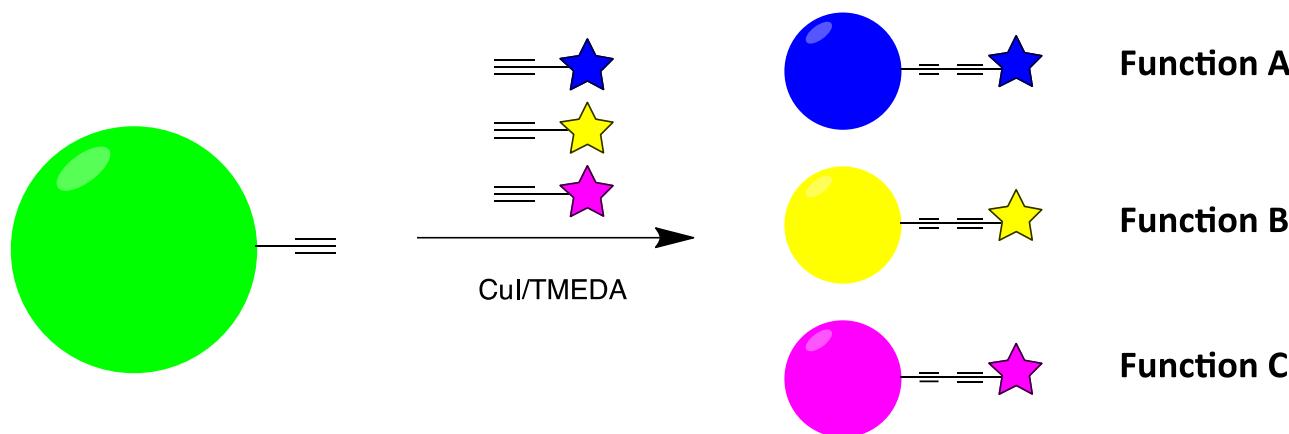


A)

	1	2	3	4	5	6	7
Temp (°C)	L	37	37	4	4	37	37
CuI/TMEDA	+	+	+	+	-	-	-



Chemically Programmed Protein Function



The Lab

- Current Students
 - Graduate Student : John Halonski
 - Seniors: Zack Nimmo, Ryan Kelley, Melanie Becher, Somya Sankar
- Juniors: Chris Travis, Anna Martin, Brent Russell, Srijan Bhasin
- Sophomores: Becca Deitch
- Interested in taking:
 - Sophomores (1-2)
 - Freshmen (1-2)

