LifeSensors <u>TUBEs</u>: <u>TANDEM UBIQUITIN BINDING ENTITIES</u>

POLYUBIQUITIN BINDING DOMAINS



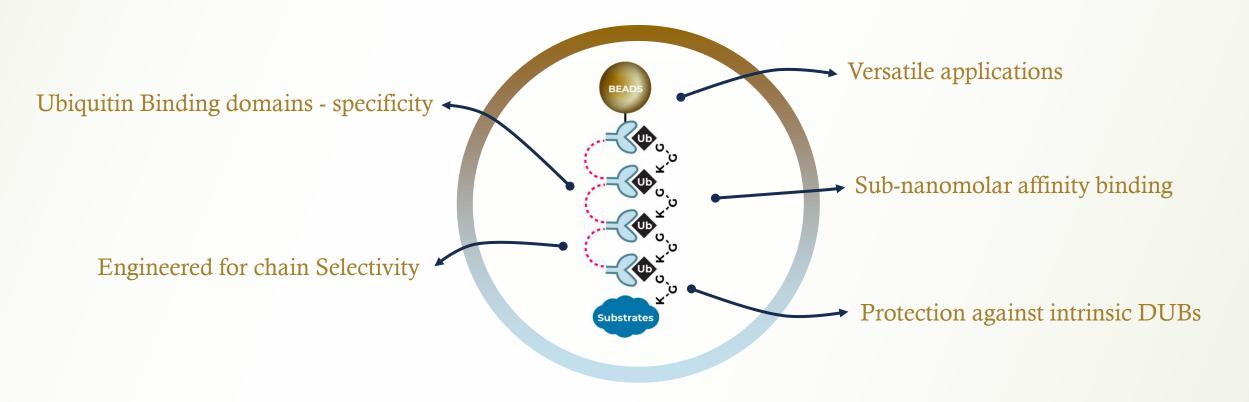
LifeSensors Inc. Mission

- Leadership in UPS, <u>PROTAC</u>, <u>DUBTAC</u>, and <u>Molecular Glues</u>
- Drug Discovery, <u>UPS Enzymes</u>, <u>DUBs</u>, <u>PROTAC Screening Services</u>
- Biomarker Development and Collaborative Research
- ~500 Products, <u>DUBs</u>, <u>E3 ligases</u>, <u>Ubiquitin Affinity Matrices (TUBEs)</u>, <u>Assay Kits</u> and Proprietary <u>Protein Expression Systems (SUMO)</u>
- Profiling Compounds Against <u>Ubiquitin Ligases</u> and <u>DUBs</u>



TUBEs:

TANDEM UBIQUITIN BINDING ENTITIES

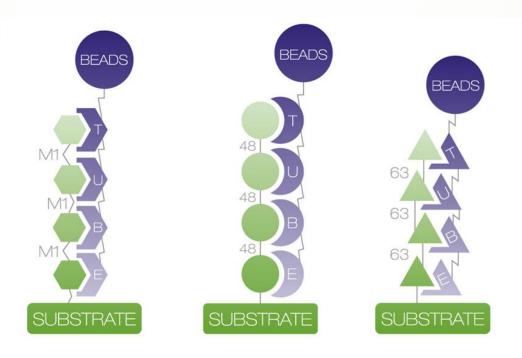






What are TUBEs?

TUBEs are high affinity 'ubiquitin traps' that capture poly-ubiquitinated proteins with low nanomolar affinity.

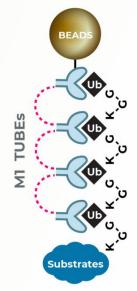


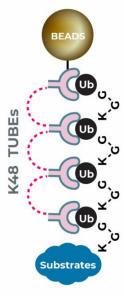
TUBEs can be used for capture or detection of total polyubiquitin or enrichment of linkage specific proteins K63-linked polyubiquitin K48-linked polyubiquitin, and M1-linked polyubiquitin.

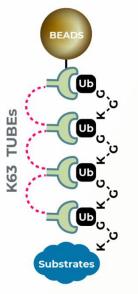


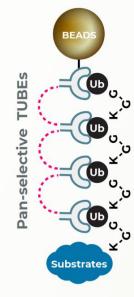


TANDEM UBIQUITIN BINDING ENTITIES (TUBEs)

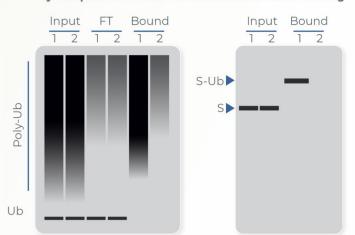




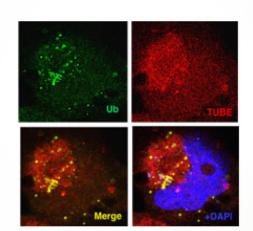




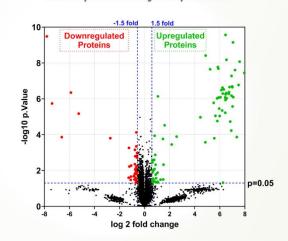
Poly-Ubiquitin Substrate Enrichment & Western Blotting



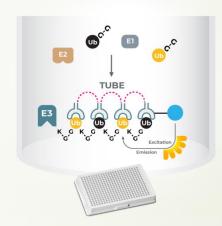
Fluorescent Microscopy



Mass Spectrometry Ubiquitomics



High-Throughput Screening







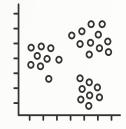
TUBEs address critical TPD challenges



Strong binders / better K_D does not guarantee better degradation



Ternary complex rigidity



Ligand dependent ubiquitination, true functional HTS



The Power of TUBE Applications

Magnetic TUBEs Pull-down of polyubiquitinated proteins

Biotin TUBEs Histochemistry/ Cytochemistry Agarose TUBEs



Biomarkers



'Far Western' blot

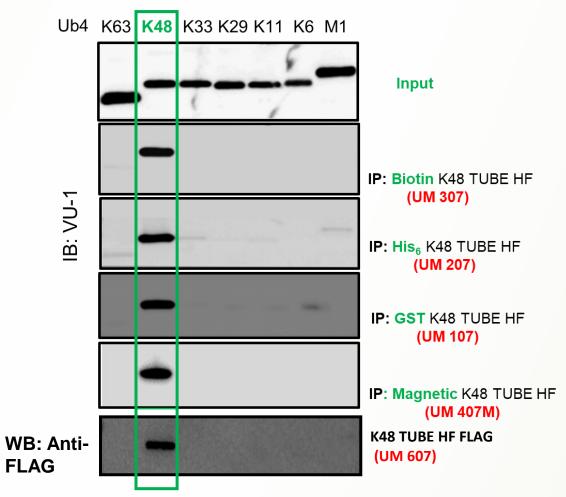


Mass Spec &
Proteomics





Lysine 48 Poly-ubiquitin TUBES are Highly Selective

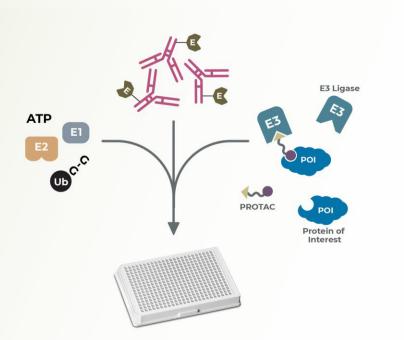


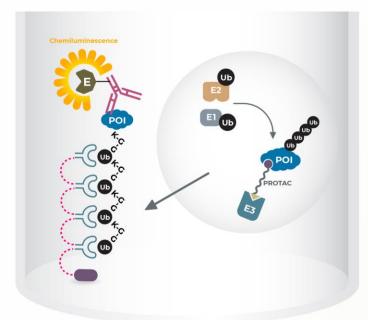
K48 TUBE HF FLAG detects only K48-linked ubiquitin chains in Western Blots

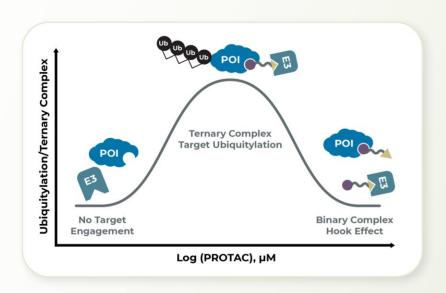


TUBE Application for HTS- In Vitro Biochemical Assay

To study functional ternary complex and 'PROTACability'



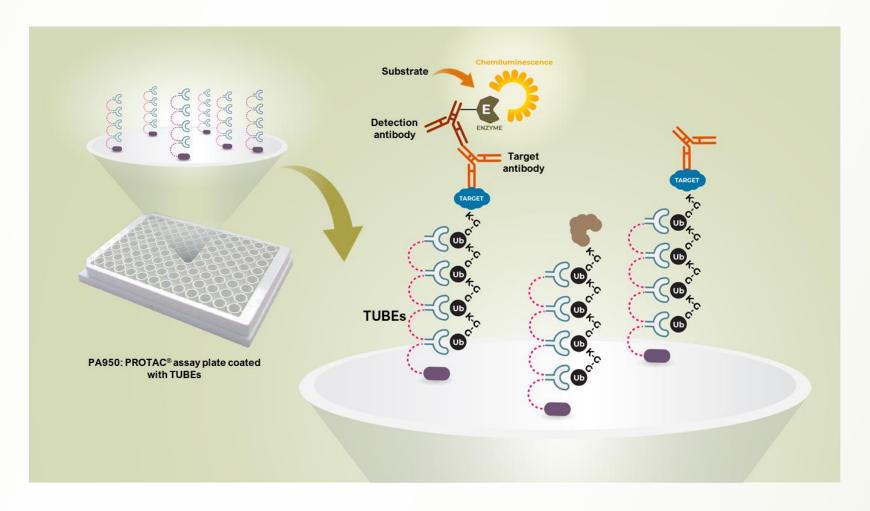




TUBE Capture & PROTAC Mediated Ubiquitination of POI Detection

Cell-Based PROTAC Assay using TUBE-coated plate

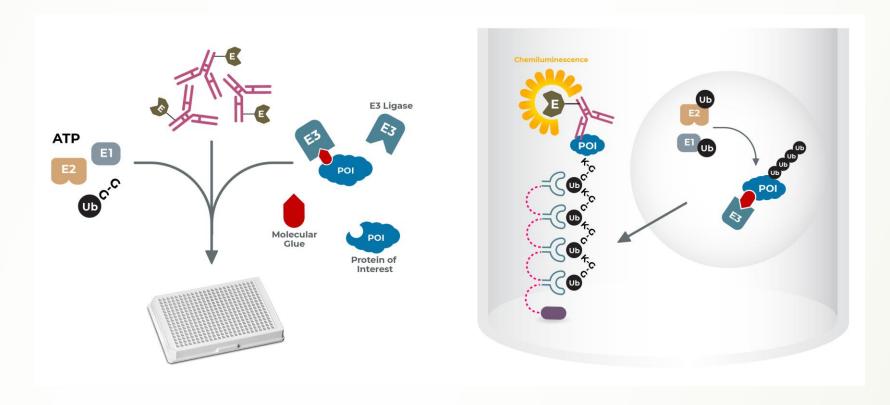
Directly monitor PROTAC-mediated ubiquitination and degradation of target protein





TUBEs' Role in Molecular Glue Discovery

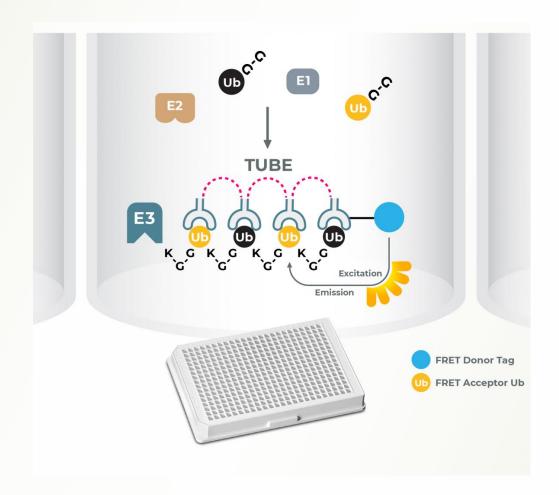
Study Molecular Glue mediated ubiquitination and degradation simultaneously





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TUBEs Facilitate Discovery of Novel E3 Ligase for KRAS



- ✓ High throughput screening for E3 ligase activators
- ✓ Homogenous assay for library screening
- ✓ Identification of novel E3 ligands
- ✓ SPR / TSA based confirmation and PROTACability





Novel Application of TUBES, in addition to Ubiquitination Studies

- ✓ Remarkable tool for monitoring PROTAC and molecular glue function
- ✓ HTS of in vivo ubiquitylated proteins
- ✓ Isolation of PROTAC/Mol Glue mediated ubiquitylated substrates from cell lysates
- ✓ Ubiquitin mass spec proteomics bypassing SILAC
- ✓ Perform E3 ligase assays using TR-FRET assays
- ✓ Superior to antibodies, detection by Western blot
- ✓ Imaging tools for In situ detection with fluorescence



Thank You

We are your partner in UPS, TUBEs, DUBs, E3s, PROTAC, Mol Glue, Protein Expression, CAR-T/Gene therapy and vaccine development

Contact Us!

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