

**Personal details**

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**Education**

2004 – 2008 Post-doctoral training (chemical biology), Stanford University, School of Medicine (California, USA).

1999 – 2004 PhD (bio-organic chemistry), Leiden University (Leiden, The Netherlands).

1994 – 1999 M.Sc. (chemistry), *cum laude*, Leiden University (Leiden, The Netherlands). \*In the Netherlands, *cum laude* represents approximately the best 5-10% of students.

**Research/professional experience**

2008 – present Emmy Noether group leader, Technical University Munich, Chair for Chemistry of Biopolymers. Laboratory interests: chemical proteomics approaches in the study of proteases.

2004 – 2008 Post-doctoral fellow at the Department of Pathology of the Stanford University School of Medicine in the group of Dr. M. Bogoy. Research project: the development of small molecule tools for chemical proteomics, focused on cysteine proteases.

1999 – 2004 PhD research at the Leiden Institute of Chemistry in collaboration with Organon/Akzo-Nobel, with Prof. Dr. J.H. van Boom, Prof. Dr. C. A. A. van Boeckel and Dr. G. A. van der Marel. Research project: the synthesis of analogs of aminoglycoside antibiotics.

1998 – 1999 Internship at the Harvard Medical School, Department of Immunology, with Prof. Dr. H. L. Ploegh. Research project: the synthesis and evaluation of inhibitors for the proteasome and cysteine proteases.

1997 – 1998 Internship at the Leiden Institute of Chemistry, in the group of Prof. Dr. J. H. van Boom (bio-organic synthesis), under supervision of Dr. H. S. Overkleeft. Research project: the synthesis of sugar amino acids as conformational restricted dipeptide isosters. Awarded with the Unilever Research Prize 1998.

**Professional organisations**

2009 – present Center for Integrated Protein Science Munich (DFG Excellence cluster), associated member

2007 – present International Proteolysis Society, member

1994 – present Royal Dutch Chemical Society (KNCV), member

**Courses taken**

|      |  |
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| 2010 | Teaching skills – Sprachraum Weiterbildungszentrum, Ludwig-Maximilian-University                       |
| 2006 | Scientific Management – Stanford University  |
| 2005 | Instruction to Teaching and Presentation Skills – Stanford University Center for Teaching and Learning |
| 2001 | Bio-organic Synthesis – course by Holland Research School of Molecular Chemistry                       |
| 1998 | Working with Radio-isotopes – Harvard Medical School   |
| 1997 | Science & Journalism – course by the Royal Dutch Chemical Society (KNCV)                               |

**Presentations and seminars**

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| 2012 | Invited lecture, Klinikum Rechts der Isar, Munich, Germany.   |
| 2011 | Invited lecture, Drug discovery meeting, Max Planck Institute for Psychiatry.   |
| 2009 | Invited lecture, Radboud University Nijmegen, The Netherlands.  |
| 2008 | Invited lecture, Max Planck Institute for Plant Breeding, Cologne, Germany  |
| 2008 | Oral presentation, Pacific Coast Protease School, California  |
| 2007 | Seminar, Penn Center for Molecular Discovery, University of Pennsylvania.   |
| 2007 | Oral presentation, Pacific Coast Protease meeting, California.  |
| 2006 | Seminar, Medicinal Chemistry Department, Utrecht University, Utrecht, The Netherlands.  |
| 2006 | Seminar, Department of Basic Life Science, Technische Universität München, Munich, Germany.   |
| 2006 | Oral presentation, Stanford Mass Spectrometry Users Meeting, Stanford, California.  |
| 2006 | Oral presentation, Pacific Coast Protease Workshop, Palm Springs, California.   |
| 2005 | Oral presentation, Stanford Department of Pathology, Monterey, California.  |
| 2004 | Oral presentation, Pacific Protease Workshop, Half Moon Bay, California.  |
| 2003 | Seminar Bio-organic Synthesis, research group Prof. H. Redlich, Institute for Organic Chemistry, Universität Münster, Münster, Germany. |
| 2002 | Oral presentation, Leiden/Amsterdam Center for Drug Research.   |

**Other activities**

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|----------------|---|
| 2010           | Organization of the 3 <sup>rd</sup> West-European meeting on activity-based proteomics  |
| 2010           | Organization of a retreat of several research groups from the department of basic life sciences of the Technical University Munich  |
| 2009 – present | Writing blog entries for the website <a href="http://www.chemnixblog.de">www.chemnixblog.de</a> , a website where German highschoolstudents can ask questions about the chemistry in their daily life |

**List of publications**

32. Serim, S.; Haedke, U.; **Verhelst, S. H. L.** (2012) Activity-based probes for the study of proteases: recent advances and developments. *ChemMedChem.*, in press.
31. Fonovic, M.; **Verhelst, S. H. L.** (2012) Novel cleavable activity-based probes suitable for proteomics-based discovery. *Methods Mol. Biol.*, in press.
30. Haedke, U.; Götz, M.; Baer, P.; **Verhelst, S. H. L.** (2012) Alkyne derivatives of isocoumarins as clickable activity-based probes for serine proteases. *Bioorg. Med. Chem.*, 20: 633-640.
29. Vinothkumar, K. R.; Strisovsky, K.; Andreeva, A.; Christova, Y.; **Verhelst, S. H. L.**; Freeman, M. (2010) The structural basis for catalysis and substrate specificity of a rhomboid protease. *EMBO J.*, 29: 3797-3809.
28. Ravindran S.; Lodoen M. B.; **Verhelst S. H.**; Bogyo M.; Boothroyd J. C. (2009) 4-bromophenacyl bromide specifically inhibits rhoptry secretion during Toxoplasma invasion. *PLoS One.* 4: e8143.
27. Beckham, S. A.; Piedrafita, D.; Phillips, C. I.; Samarawickrema, N.; Law, R. H. P.; Smooker, P. M.; Quinsey, N. S.; Irving, J. A.; Greenwood, S.; **Verhelst, S. H. L.**; Bogyo, M.; Turk, B.; Coetzer, T. H.; Wijeyewickrema, L. C.; Spithill, T. W.; Pike R. N. (2009) A major Cathepsin B protease from the liver fluke *Fasciola hepatica* exhibits novel biochemical properties and plays a vital role in the survival of newly excysted juvenile forms *in vitro*. *Int. J. Biochem. Cell Biol.*, 41: 1601-1612.
26. Kaschani, F.; **Verhelst, S. H. L.**; Van Swieten, P. F.; Wang, Z.; Kaiser, M.; Overkleeft, H. S.; Bogyo, M.; Van der Hoorn, R. A. L. (2009) Minitags for small molecules: detecting targets of reactive small molecules in living plant tissues using click-chemistry. *The Plant Journal*, 57: 373-85.
25. Yang, Z.; Fonovic, M.; **Verhelst, S. H. L.**; Blum, G.; Bogyo, M. (2009) Evaluation of  $\alpha,\beta$ -unsaturated ketone-based probes for papain-family cysteine proteases. *Bioorg. Med. Chem.*, 17: 1071-1078.
24. Burster, T.; Marin-Esteban, V.; Boehm, B. O.; Dunn, S.; Rotzschke, O.; Falk, K.; Weber, E.; **Verhelst, S. H. L.**; Kalbacher, H.; Driessen, C. (2007) Design of protease-resistant myelin basic protein-derived peptides by cleavage site directed amino acid substitutions. *Biochem. Pharm.*, 74: 1514-1523.
23. Fonovic, M.; **Verhelst, S. H. L.**; Sorum, M. T.; Bogyo, M. (2007) Proteomic evaluation of chemically cleavable activity based probes. *Mol. Cell. Proteomics*, 6: 1761-1770.
22. Sadaghiani, A. M.; **Verhelst, S. H. L.**; Gocheva, V.; Hill, K.; Majerova, E.; Stinson, S.; Joyce, J. A.; Bogyo, M. (2007) Design, synthesis and evaluation of *in vivo* potency and selectivity of epoxysuccinyl-based inhibitors of papain family cysteine proteases. *Chem. Biol.*, 14: 499-511.
21. Cuerrier, D.; Moldoveanu, T.; Campbell, R. L.; Kelly, J.; Yoruk, B.; **Verhelst, S. H. L.**; Greenbaum, D.; Bogyo, M.; Davies, P. L. (2007) Development of calpain-specific inactivators by screening of positional-scanning epoxide libraries. *J. Biol. Chem.*, 282: 9600-9611.
20. **Verhelst, S. H. L.**; Fonovic, M.; Bogyo, M. (2007) A mild chemically cleavable linker system for functional proteomic applications. *Angew. Chem. Int. Ed.*, 46: 1284-1286.

19. Sexton, K. B.; Kato, D.; Berger, A.; Fonovic, M.; **Verhelst, S. H. L.**; Bogyo, M. (2007) Specificity of aza-peptide electrophile activity-based probes of caspases. *Cell Death Differ.*, 14: 727-732.
18. Sadaghiani, A. M.; **Verhelst, S. H. L.**; Bogyo, M. (2007) Tagging and detection strategies for activity-based proteomics. *Curr. Opin. Chem. Biol.*, 11: 20-28.
17. Sadaghiani, A. M.; **Verhelst, S. H. L.**; Bogyo, M. (2006) Solid phase methods for the preparation of epoxysuccinate-based inhibitors of cysteine proteases. *J. Comb. Chem.*, 8: 802-804.
16. Yuan, F.; **Verhelst, S. H. L.**; Blum, G.; Coussens, L. M.; Bogyo, M. (2006) A selective activity-based probe for the papain family cysteine protease dipeptidyl peptidase I/cathepsin C. *J. Am. Chem. Soc.*, 128: 5616-5617.
15. **Verhelst, S. H. L.**; Witte, M. D.; Arastu-Kapur, S.; Fonovic, M.; Bogyo, M. (2006) Novel aza peptide inhibitors and active site probes of papain family cysteine proteases. *ChemBioChem*, 7: 943-950.
14. Kato, D.; **Verhelst, S. H. L.**; Sexton, K. B.; Bogyo, M. (2005) A general solid phase method for the preparation of diverse azapeptide probes directed against cysteine proteases. *Org. Lett.*, 7: 5649-5652. \*equal authorship.
13. Timmer, M. S. M.; **Verhelst, S. H. L.**; Grotenbreg, G. M.; Overhand, M.; Overkleeft, H. S. (2005) Carbohydrates as versatile platforms in the construction of small compound libraries. *Pure & Applied Chem.*, 77: 1173-1182.
12. **Verhelst, S. H. L.**; Bogyo, M. (2005) Solid phase synthesis of double headed epoxysuccinyl activity based probes for selective targeting of papain family cysteine proteases. *ChemBioChem*, 6: 824-827.
11. **Verhelst, S. H. L.**; Bogyo, M. (2005) Dissecting protein function using chemical proteomic methods. *QSAR Comb. Sci.*, 24: 261-269.
10. **Verhelst, S. H. L.**; Bogyo, M. (2005) Chemical proteomics applied to target identification and drug discovery. *Biotechniques*, 38: 175-177.
9. Chehade, K. A. H.; Baruch, A.; **Verhelst, S. H. L.**; Bogyo, M. (2005) An improved preparation of the activity-based probe JPM-OEt and in situ applications. *Synthesis*, 240-244.
8. **Verhelst, S. H. L.**; Magnée, L.; Wennekes, T.; Wiedenhof, W.; Van der Marel, G. A.; Overkleeft, H. S.; Van Boeckel, C. A. A.; Van Boom, J. H. (2004) Glycosylation of cyclitols: novel analogs of aminoglycoside antibiotics. *Eur. J. Org. Chem.* 11: 2404-2410.
7. **Verhelst, S. H. L.**; Michiels, P. J. A.; Van der Marel, G. A.; Van Boeckel, C. A. A.; Van Boom, J. H. (2004) Surface plasmon resonance evaluation of various aminoglycoside – RNA hairpin interactions reveals low degree of selectivity. *ChemBioChem*, 5: 937-942.
6. **Verhelst, S. H. L.**; Wennekes, T.; Van der Marel, G. A.; Overkleeft, H. S.; Van Boeckel, C. A. A.; Van Boom, J. H. (2004) Synthesis of orthogonally protected 2-deoxystreptamine stereoisomers. *Tetrahedron* 60: 2813-2822.
5. **Verhelst, S. H. L.**; Paez Martinez, B.; Timmer, M. S. M.; Lodder, G.; Van der Marel, G. A.; Overkleeft, H. S.; Van Boom, J. H. (2003) A short chiral route towards polyhydroxylated indolizidines and quinolizidines. *J. Org. Chem.* 68: 9598-9603.

4. **Verhelst, S. H. L.**; Wiedenhof, W.; Ovaa, H.; Van der Marel, G. A.; Overkleeft, H. S.; Van Boeckel, C. A. A.; Van Boom, J. H. (2002) A stereoselective route towards highly functionalized 4,6-diaminocyclohexene derivatives. *Tetrahedron Lett.* 43: 6451-6455.
3. Shi, G. P.; Bryant, R. A. R.; Riese, R.; **Verhelst, S.**; Driessen, C.; Li, Z. Q.; Brömme, D.; Ploegh, H. L.; Chapman, H. A. (2000) Role for cathepsin F in invariant chain processing and major histocompatibility complex class II peptide loading by macrophages. *J. Exp. Med.* 191: 1177-1185.
2. Bogyo, M.; **Verhelst, S.**; Bellingard-Dubouchaud, V.; Toba, S.; Greenbaum, D. (2000) Selective targeting of lysosomal cysteine proteases with radiolabeled electrophilic substrate analogs. *Chem. Biol.* 7: 27-38.
1. Overkleeft, H. S.; **Verhelst, S. H. L.**; Pieterman, E.; Meeuwenoord, N. J.; Overhand, M.; Cohen, L. H.; Van der Marel, G. A.; Van Boom, J. H. (1999) Design and synthesis of a protein: farnesyltransferase inhibitor based on sugar amino acids. *Tetrahedron Lett.* 40: 4103-4106.

### Patents

Bogyo, M.; **Verhelst, S. H. L.**; Fonovic, M. A mild chemically cleavable linker for use in affinity purification and proteomics applications. Int. Patent PCT/US2007/017233.

Bogyo, M.; Sadaghiani, A. M.; **Verhelst, S. H. L.** Design and Synthesis of epoxysuccinyl inhibitors of cysteine cathepsins for cancer chemotherapy. US Patent, no US 11/762,735; Int. Patent PCT/2007/071145.

Bogyo, M.; **Verhelst, S. H. L.** Solid-Phase Synthesis of Small Molecule Inhibitors of Cathepsin Cysteine Proteases. US patent, no US 11/; Int. patent, PCT/US2006/000869.

Overkleeft, H. S.; **Verhelst, S. H. L.**; Meeuwenoord, N. J.; Pieterman, E. J.; Cohen, L. H.; Overhand, M.; Van der Marel, G. A.; Van Boom, J. H. Novel protein: prenyl transferase inhibitors. European patent, no EP1028117.