

Stanford

Carol Clayberger

Professor (Research) of Pediatrics, Emerita
Pediatrics - Immunology and Allergy

Bio

ACADEMIC APPOINTMENTS

- Emeritus Faculty, Acad Council, Pediatrics - Immunology and Allergy
- Member, Bio-X

PROFESSIONAL EDUCATION

- A.B., Mount Holyoke College , Biology (1974)
- Ph.D., Yale University , Cell Biology (1979)

Research & Scholarship

CURRENT RESEARCH AND SCHOLARLY INTERESTS

T lymphocytes play a central role in the adaptive immune response. Understanding the biology of the response of T cells to immune challenge should allow the development of new diagnostics and therapeutics to treat human disease. Research in my group is focuses on three areas:

1. Granulysin, a small molecule expressed by activated T cells and NK cells, lyses tumors and a variety of pathogens, including bacteria, fungi, and parasites. We have shown that synthetic peptides corresponding to linear regions of granulysin can recapitulate the lytic activity of the intact molecule. Substitution of critical residues resulted in mutant peptides that lyse pathogens but do not kill mammalian cells. We are currently developing granulysin derivatives as a new type of antibiotic as well as studying the mechanisms involved in granulysin induced cell death.
2. Tuberculosis kills more than 3,000,000 people each year. However, little is understood about the immune response to *Mycobacterium tuberculosis*, the bacterium that causes the disease. We are characterizing the immune response to individual antigens from *Mycobacterium tuberculosis* in lymphocytes from individuals with disease or who have been infected with *Mycobacterium tuberculosis* but do not develop tuberculosis. Correlates of protective immunity are critical to the development of an improved vaccine for tuberculosis.
3. Lymphotactin is a chemokine that is expressed in both activated and anergic T cells. Lymphocytes treated with lymphotactin are refractory to stimulation, suggesting that lymphotactin may be used to induce tolerance in vivo. Studies are ongoing to define the role of lymphotactin in the immune response in vitro and in vivo and to use this information to develop new tolerance inducing modalities.

Teaching

GRADUATE AND FELLOWSHIP PROGRAM AFFILIATIONS

- Immunology (Phd Program)

Publications

PUBLICATIONS

- Unique gene expression profiles in infants vaccinated with different strains of *Mycobacterium bovis* bacille Calmette-Guerin *INFECTION AND IMMUNITY*

- Wu, B., Huang, C., Garcia, L., Ponce de Leon, A., Sifuentes Osornio, J., Bobadilla-del-Valle, M., Ferreira, L., Canizales, S., Small, P., Kato-Maeda, M., Krens, A. M., Clayberger, C.
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- **Interaction of PRP4 with Kruppel-like factor 13 regulates CCL5 transcription** *JOURNAL OF IMMUNOLOGY*
Huang, B., Ahn, Y., McPherson, L., Clayberger, C., Krensky, A. M.
2007; 178 (11): 7081-7087
 - **Messenger RNA expression of IL-8, FOXP3, and IL-12 beta differentiates latent tuberculosis infection from disease** *JOURNAL OF IMMUNOLOGY*
Wu, B., Huang, C., Kato-Maeda, M., Hopewell, P. C., Daley, C. L., Krensky, A. M., Clayberger, C.
2007; 178 (6): 3688-3694
 - **Dynamic interplay of transcriptional machinery and chromatin regulates "late" expression of the chemokine RANTES in T lymphocytes** *MOLECULAR AND CELLULAR BIOLOGY*
Ahn, Y., Huang, B., McPherson, L., Clayberger, C., Krensky, A. M.
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 - **Granulysin-mediated tumor rejection in transgenic mice** *JOURNAL OF IMMUNOLOGY*
Huang, L. P., Lyu, S., Clayberger, C., Krensky, A. M.
2007; 178 (1): 77-84
 - **Granulysin: A novel host defense molecule** *AMERICAN JOURNAL OF TRANSPLANTATION*
Krensky, A. M., CLAYBERGER, C.
2005; 5 (8): 1789-1792
 - **Glycosylated recombinant human XCL1/lymphotactin exhibits enhanced biologic activity** *JOURNAL OF IMMUNOLOGICAL METHODS*
Dong, C., Chua, A., Ganguly, S., KRENSKY, A. M., Clayberger, C.
2005; 302 (1-2): 136-144
 - **Granulysin, a cytolytic molecule, is also a chemoattractant and proinflammatory activator** *JOURNAL OF IMMUNOLOGY*
Deng, A. M., Chen, S. X., Li, Q., Lyu, S. C., CLAYBERGER, C., Krensky, A. M.
2005; 174 (9): 5243-5248
 - **A novel apoptosis pathway activated by the carboxyl terminus of p21** *BLOOD*
Dong, C., Li, Q., Lyu, S. C., KRENSKY, A. M., Clayberger, C.
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 - **Hemolysis of erythrocytes by granulysin-derived peptides but not by granulysin** *ANTIMICROBIAL AGENTS AND CHEMOTHERAPY*
Li, Q., Dong, C., Deng, A. M., Katsumata, M., Nakadai, A., Kawada, T., Okada, S., CLAYBERGER, C., Krensky, A. M.
2005; 49 (1): 388-397
 - **DQ 65-79, a peptide derived from HLA class II, mimics p21 to block T cell proliferation** *JOURNAL OF IMMUNOLOGY*
Dong, C., Lyu, S. C., KRENSKY, A. M., CLAYBERGER, C.
2003; 171 (10): 5064-5070
 - **Granulysin** *CURRENT OPINION IN IMMUNOLOGY*
CLAYBERGER, C., KRENSKY, A. M.
2003; 15 (5): 560-565
 - **Intracellular mediators of granulysin-induced cell death** *JOURNAL OF IMMUNOLOGY*
Okada, S., Li, Q., Whitin, J. C., CLAYBERGER, C., Krensky, A. M.
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 - **Protein kinase B/Akt is essential for the insulin- but not progesterone-stimulated resumption of meiosis in Xenopus oocytes** *BIOCHEMICAL JOURNAL*
Andersen, C. B., Sakaue, H., Nedachi, T., Kovacina, K. S., CLAYBERGER, C., Conti, M., Roth, R. A.
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 - **Functional domains and DNA-binding sequences of RFLAT-1/KLF13, a Kruppel-like transcription factor of activated T lymphocytes** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Song, A., Patel, A., Thamatrakoln, K., Liu, C., Feng, D. D., CLAYBERGER, C., KRENSKY, A. M.
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- **A translational rheostat for RFLAT-1 regulates RANTES expression in T lymphocytes** *JOURNAL OF CLINICAL INVESTIGATION*
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2002; 110 (1): 119-126
- **DQ 65-79, a peptide derived from HLA class II, induces I kappa B expression** *JOURNAL OF IMMUNOLOGY*
Jiang, Y., Chen, D., Lyu, S. C., Ling, X. F., KRENSKY, A. M., Clayberger, C.
2002; 168 (7): 3323-3328
- **A distinct pathway of cell-mediated apoptosis initiated by granulysin** *JOURNAL OF IMMUNOLOGY*
Kaspar, A. A., Okada, S., Kumar, J., Poulain, F. R., Drouvalakis, K. A., Kelekar, A., Hanson, D. A., Kluck, R. M., Hitoshi, Y., Johnson, D. E., Froelich, C. J., Thompson, C. B., Newmeyer, et al
2001; 167 (1): 350-356
- **A non-peptide functional antagonist of the CCR1 chemokine receptor is effective in rat heart transplant rejection** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Horuk, R., CLAYBERGER, C., KRENSKY, A. M., Wang, Z. H., Grone, H. J., Weber, C., Weber, K. S., Nelson, P. J., May, K., Rosser, M., Dunning, L., Liang, M., Buckmann, et al
2001; 276 (6): 4199-4204
- **Granulysin: a novel antimicrobial** *EXPERT OPINION ON INVESTIGATIONAL DRUGS*
Kumar, J., Okada, S., CLAYBERGER, C., KRENSKY, A. M.
2001; 10 (2): 321-329
- **Granulysin blocks replication of varicella-zoster virus and triggers apoptosis of infected cells** *VIRAL IMMUNOLOGY*
Hata, A., Zerboni, L., Sommer, M., Kaspar, A. A., CLAYBERGER, C., KRENSKY, A. M., Arvin, A. M.
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- **Granulysin expression is a marker for acute rejection and steroid resistance in human renal transplantation** *HUMAN IMMUNOLOGY*
Sarwal, M. M., Jani, A., Chang, S., Huie, P., Wang, Z., Salvatierra, O., CLAYBERGER, C., Sibley, R., KRENSKY, A. M., Pavlakis, M.
2001; 62 (1): 21-31
- **Granulysin, a T cell product, kills bacteria by altering membrane permeability** *JOURNAL OF IMMUNOLOGY*
Ernst, W. A., Thoma-Uszynski, S., Teitelbaum, P., Ko, C., Hanson, D. A., CLAYBERGER, C., KRENSKY, A. M., Leippe, M., Bloom, B. R., Ganz, T., Modlin, R. L.
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- **An immunosuppressive and anti-inflammatory HLA class I-derived peptide binds vascular cell adhesion molecule-1** *TRANSPLANTATION*
Ling, X. F., Tamaki, T., Xiao, Y., Kamangar, S., CLAYBERGER, C., Lewis, D. B., KRENSKY, A. M.
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- **Bactericidal and tumoricidal activities of synthetic peptides derived from granulysin** *JOURNAL OF IMMUNOLOGY*
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- **Proliferating cell nuclear antigen as the cell cycle sensor for an HLA-derived peptide blocking T cell proliferation** *JOURNAL OF IMMUNOLOGY*
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- **A human class II MHC-derived peptide antagonizes phosphatidylinositol 3-kinase to block IL-2 signaling** *JOURNAL OF CLINICAL INVESTIGATION*
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- **A novel method for measuring CTL and NK cell-mediated cytotoxicity using annexin V and two-color flow cytometry** *JOURNAL OF IMMUNOLOGICAL METHODS*
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- **Inhibition of cell cycle progression by a synthetic peptide corresponding to residues 65-79 of an HLA class II sequence: Functional similarities but mechanistic differences with the immunosuppressive drug rapamycin** *JOURNAL OF IMMUNOLOGY*
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- **Interactions of HLA-B*4801 with peptide and CD8** *TISSUE ANTIGENS*
MARTINEZNAVES, E., Barber, L. D., Madrigal, J. A., Vullo, C. M., CLAYBERGER, C., Lyu, S. C., Williams, R. C., Gorodezky, C., Markow, T., PETZLERLER, M. L., Parham, P.
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KRENSKY, A. M., CLAYBERGER, C.
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- **Immunologic tolerance: Tailored antigen** *1st International Cybercongress - Transplantation in the next Millennium: A Landmark in Organ Transplantation Technology*
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ELSEVIER SCIENCE INC.1996: 2075-77
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Nelson, E. L., Li, X. B., Hsu, F. J., Kwak, L. W., Levy, R., CLAYBERGER, C., KRENSKY, A. M.
1996; 88 (2): 580-589
- **HLA-derived peptides which inhibit T cell function bind to members of the heat-shock protein 70 family** *JOURNAL OF EXPERIMENTAL MEDICINE*
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1996; 183 (2): 339-348
- **Structure of HLA molecules and immunosuppressive effects of HLA derived peptides.** *International reviews of immunology*
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1996; 13 (3): 173-185
- **Human CD4(+) and CD8(+) T lymphocytes are both cytotoxic to Toxoplasma gondii-infected cells** *INFECTION AND IMMUNITY*
Montoya, J. G., Lowe, K. E., CLAYBERGER, C., Moody, D., Do, D., Remington, J. S., Talib, S., Subauste, C. S.
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- **IMMUNOSUPPRESSIVE PEPTIDES CORRESPONDING TO MHC CLASS-I SEQUENCES** *CURRENT OPINION IN IMMUNOLOGY*
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- **IN-VIVO STUDIES OF THE MAINTENANCE OF PERIPHERAL TRANSPLANT TOLERANCE AFTER CYCLOSPORINE - RADIOSENSITIVE ANTIGEN-SPECIFIC SUPPRESSOR CELLS MEDIATE LASTING GRAFT PROTECTION AGAINST PRIMED EFFECTOR-CELLS** *TRANSPLANTATION*
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- **HLA-DERIVED PEPTIDES AS NOVEL IMMUNOTHERAPEUTICS** *CLINICAL IMMUNOLOGY AND IMMUNOPATHOLOGY*
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- **IMMUNOMODULATION BY SOLUBLE HLA CLASS-I** *TRANSPLANTATION*
Buelow, R., Burlingham, W. J., CLAYBERGER, C.
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- **PROLONGATION OF ALLOGENEIC HEART GRAFT-SURVIVAL IN RATS BY ADMINISTRATION OF A PEPTIDE (AA-75-84) FROM THE ALPHA-1 HELIX OF THE FIRST DOMAIN OF HLA-B7-01** *TRANSPLANTATION*

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- **PROLONGATION OF SKIN ALLOGRAFT SURVIVAL IN MICE FOLLOWING ADMINISTRATION OF ALLOTRAP** *13th Annual Meeting of the American-Society-of-Transplant-Physicians*
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 - **GAMMA-DELTA T-CELL RECOGNITION OF TUMOR IG PEPTIDE** *JOURNAL OF IMMUNOLOGY*
Kim, H. T., Nelson, E. L., CLAYBERGER, C., SANJANWALA, M., Sklar, J., KRENSKY, A. M.
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 - **NOVEL TUMOR-ASSOCIATED ACCESSORY MOLECULES INVOLVED IN THE GAMMA/DELTA CYTOTOXIC T-LYMPHOCYTE BURKITT'S-LYMPHOMA INTERACTION** *CANCER*
Nelson, E. L., Kim, H. T., MAR, N. D., Goralski, T. J., MCINTYRE, B. W., CLAYBERGER, C., KRENSKY, A. M.
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 - **OVERLAP IN THE REPERTOIRES OF PEPTIDES BOUND IN-VIVO BY A GROUP OF RELATED CLASS-I HLA-B ALLOTYPES** *CURRENT BIOLOGY*
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 - **DECAMER PEPTIDE DERIVED FROM THE ALPHA-1-HELIX OF THE FIRST DOMAIN OF HLA-B7-01 PROLONGS ALLOGRAFT SURVIVAL IN RATS WITH AN INHIBITION OF GRAFT INFILTRATING CELL CYTOTOXICITY** *XVth World Congress of the Transplantation-Society*
Cuturi, M. C., Josien, R., Cantarovich, D., Douillard, P., Pannetier, C., SMIT, H., Menoret, S., Pouletty, P., CLAYBERGER, C., Soullillou, J. P.
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 - **ANALYSIS OF BRONCHOALVEOLAR LAVAGE FROM HUMAN LUNG-TRANSPLANT RECIPIENTS BY FLOW-CYTOMETRY** *RESPIRATORY MEDICINE*
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