

Stanford



Lawrence Leung

Maureen Lyles D'Ambrogio Professor in the School of Medicine
Medicine - Hematology

CLINICAL OFFICES

- **Hematology Clinic**

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Bio

CLINICAL FOCUS

- Cancer > Hematology
- Hematology

ACADEMIC APPOINTMENTS

- Professor, Medicine - Hematology
- Member, Cardiovascular Institute

ADMINISTRATIVE APPOINTMENTS

- Senior Associate Dean for Veterans Affairs, School of Medicine, (2006- present)
- Chief of Staff, VA Palo Alto Health Care System, (2006- present)

BOARDS, ADVISORY COMMITTEES, PROFESSIONAL ORGANIZATIONS

- Editor-in-Chief, Hematology, UpToDate (2008 - present)
- Member, Association of American Physicians (2004 - present)
- Member, American Society for Clinical Investigation (1987 - present)

PROFESSIONAL EDUCATION

- Board Certification: Hematology, American Board of Internal Medicine (1980)
- Residency: New York Presbyterian Cornell Campus Internal Medicine Residency (1978) NY

- Internship: New York Presbyterian Cornell Campus Internal Medicine Residency (1976) NY
- Fellowship: New York Presbyterian Cornell Campus Hematology Oncology Fellowship (1981) NY
- Medical Education: Columbia University Vagelos College of Physicians and Surgeons Registrar (1975) NY
- Board Certification: Internal Medicine, American Board of Internal Medicine (1978)
- Board Certification: Medical Oncology, American Board of Internal Medicine (1981)

Research & Scholarship

CURRENT RESEARCH AND SCHOLARLY INTERESTS

Our long-term interest is to have a better understanding of the natural antithrombotic pathways and the pathophysiology of vascular thrombosis. We have focused on thrombin, the key enzyme in the blood clotting cascade. Using a library of thrombin mutants generated by site-directed mutagenesis, we have mapped the critical functional sites on thrombin for its interactions with its diverse substrates. We are studying a thrombin-activatable carboxypeptidase that plays a key role in the physiologic regulation of thrombin's pro-thrombotic and pro-inflammatory properties. Our goal is to develop new antithrombotic agents and devise new diagnostic tests for vascular thrombotic disorders.

Publications

PUBLICATIONS

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- **Antibody-mediated targeting of cleavage-specific OPN-T cell interactions** *PLOS ONE*
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- **Decoding the Genomics of Abdominal Aortic Aneurysm.** *Cell*
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2018; 174 (6): 1361
- **Chemerin 156F, generated by chymase cleavage of prochemerin, is elevated in joint fluids of arthritis patients.** *Arthritis research & therapy*
Zhao, L., Yamaguchi, Y., Ge, X., Robinson, W. H., Morser, J., Leung, L. L.
2018; 20 (1): 132
- **Men and Women Differ in the Biochemical Composition of Platelet-Rich Plasma** *AMERICAN JOURNAL OF SPORTS MEDICINE*
Xiong, G., Lingampalli, N., Koltsov, J. B., Leung, L. L., Bhutani, N., Robinson, W. H., Chu, C. R.
2018; 46 (2): 409–19
- **Prochemerin cleavage by factor XIa links coagulation and inflammation** *BLOOD*
Ge, X., Yamaguchi, Y., Zhao, L., Bury, L., Gresele, P., Berube, C., Leung, L. L., Morser, J.
2018; 131 (3): 353–64
- **Dynamic and tissue-specific proteolytic processing of chemerin in obese mice.** *PloS one*
Zhao, L., Yamaguchi, Y., Shen, W., Morser, J., Leung, L. L.
2018; 13 (8): e0202780
- **Chemerin Activation in Human Obesity** *OBESITY*
Chang, S., Eisenberg, D., Zhao, L., Adams, C., Leib, R., Morser, J., Leung, L.
2016; 24 (7): 1522-1529

- **Plasmin as a complement C5 convertase.** *EBioMedicine*
Leung, L. L., Morser, J.
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- **Thrombin Cleavage of Osteopontin Disrupts a Pro-chemotactic Sequence for Dendritic Cells, Which Is Compensated by the Release of Its Pro-chemotactic C-terminal Fragment** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Shao, Z., Morser, J., Leung, L. L.
2014; 289 (39): 27146-27158
- **Thrombin cleavage of osteopontin disrupts a pro-chemotactic sequence for dendritic cells, which is compensated by the release of its pro-chemotactic C-terminal fragment.** *journal of biological chemistry*
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- **Open aortic valve replacement in a patient with Glanzmann's thrombasthenia: a multidisciplinary strategy to minimize perioperative bleeding.** *Transfusion*
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- **Brief report: carboxypeptidase B serves as a protective mediator in osteoarthritis.** *Arthritis & rheumatology (Hoboken, N.J.)*
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2014; 66 (1): 101-106
- **Carboxypeptidase B Serves as a Protective Mediator in Osteoarthritis** *ARTHRITIS & RHEUMATOLOGY*
Lepus, C. M., Song, J. J., Wang, Q., Wagner, C. A., Lindstrom, T. M., Chu, C. R., Sokolove, J., Leung, L. L., Robinson, W. H.
2014; 66 (1): 101-106
- **Targeting complement component 5a promotes vascular integrity and limits airway remodeling.** *Proceedings of the National Academy of Sciences of the United States of America*
Khan, M. A., Maasch, C., Vater, A., Klussmann, S., Morser, J., Leung, L. L., Atkinson, C., Tomlinson, S., Heeger, P. S., Nicolls, M. R.
2013; 110 (15): 6061-6066
- **Thrombin-cleaved Fragments of Osteopontin Are Overexpressed in Malignant Glial Tumors and Provide a Molecular Niche with Survival Advantage** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Yamaguchi, Y., Shao, Z., Sharif, S., Du, X., Myles, T., Merchant, M., Harsh, G., Glantz, M., Recht, L., Morser, J., Leung, L. L.
2013; 288 (5): 3097-3111
- **Microfluidic impedance cytometer for platelet analysis** *LAB ON A CHIP*
Evander, M., Ricco, A. J., Morser, J., Kovacs, G. T., Leung, L. L., Giovangrandi, L.
2013; 13 (4): 722-729
- **Differential Gene Expression in Thrombomodulin (TM; CD141)(+) and TM(-) Dendritic Cell Subsets.** *PloS one*
Toda, M., Shao, Z., Yamaguchi, K. D., Takagi, T., D'Alessandro-Gabazza, C. N., Taguchi, O., Salamon, H., Leung, L. L., Gabazza, E. C., Morser, J.
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- **Chemerin Bioactivity Is Regulated by Factor XIa: A Novel Interface Linking Between Coagulation, Hemostasis and Immunity** *53rd Annual Meeting and Exposition of the American-Society-of-Hematology (ASH)*
Yamaguchi, Y., Morser, J., Leung, L. L.
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- **Chemerin158K Protein Is the Dominant Chemerin Isoform in Synovial and Cerebrospinal Fluids but Not in Plasma** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Zhao, L., Yamaguchi, Y., Sharif, S., Du, X., Song, J. J., Lee, D. M., Recht, L. D., Robinson, W. H., Morser, J., Leung, L. L.
2011; 286 (45): 39520-39527
- **Proteolytic Cleavage of Chemerin Protein Is Necessary for Activation to the Active Form, Chem157S, Which Functions as a Signaling Molecule in Glioblastoma** *JOURNAL OF BIOLOGICAL CHEMISTRY*
Yamaguchi, Y., Du, X., Zhao, L., Morser, J., Leung, L. L.
2011; 286 (45): 39510-39519
- **Plasma carboxypeptidase B downregulates inflammatory responses in autoimmune arthritis** *JOURNAL OF CLINICAL INVESTIGATION*

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- **Enhanced Abdominal Aortic Aneurysm Formation in Thrombin-Activatable Procarboxypeptidase B-Deficient Mice** *ARTERIOSCLEROSIS THROMBOSIS AND VASCULAR BIOLOGY*
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 - **Thrombospondin Type I Domain Containing 7A (THSD7A) Mediates Endothelial Cell Migration and Tube Formation** *JOURNAL OF CELLULAR PHYSIOLOGY*
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 - **Heparin-induced thrombosis without thrombocytopenia** *JOURNAL OF THORACIC AND CARDIOVASCULAR SURGERY*
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 - **Thrombin-Activatable Carboxypeptidase B Cleavage of Osteopontin Regulates Neutrophil Survival and Synovocyte Binding in Rheumatoid Arthritis** *ARTHRITIS AND RHEUMATISM*
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 - **Essential thrombin residues for inhibition by protein C inhibitor with the cofactors heparin and thrombomodulin** *JOURNAL OF THROMBOSIS AND HAEMOSTASIS*
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2007; 5 (7): 1486-1492
 - **Thrombin-activatable procarboxypeptidase B regulates activated complement C5a in vivo** *BLOOD*
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 - **Antiangiogenic antithrombin induces global changes in the gene expression profile of endothelial cells** *CANCER RESEARCH*
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 - **Perioperative evaluation of bleeding diathesis.** *Hematology / the Education Program of the American Society of Hematology. American Society of Hematology. Education Program*
Leung, L. L.
2006: 457-461

- **Thrombin-activatable fibrinolysis inhibitor (TAFI) regulates activated complement C5a in vivo.** *46th Annual Meeting of the American-Society-of-Hematology*
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- **Crystal structure of anticoagulant thrombin variant E217K provides insights into thrombin allostery** *JOURNAL OF BIOLOGICAL CHEMISTRY*
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Ishida, T., Zheng, Z., Dichek, H. L., Wang, H. J., Moreno, I., Yang, E., Kundu, R. K., Talbi, S., Hirata, K. I., Leung, L. L., Quertermous, T.
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- **Thrombin activation of factor XI on activated platelets requires the interaction of factor XI and platelet glycoprotein Ib alpha with thrombin anion-binding exosites I and II, respectively (Retracted Article. See vol 282, pg 29067, 2007)** *JOURNAL OF BIOLOGICAL CHEMISTRY*
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- **Structural requirements for the activation of human factor VIII by thrombin** *BLOOD*
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- **An extensive interaction interface between thrombin and factor V is required for factor V activation** *JOURNAL OF BIOLOGICAL CHEMISTRY*
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- **Modulation of thrombin's procoagulant and anticoagulant properties** *THROMBOSIS AND HAEMOSTASIS*
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Tsiang, M., Gibbs, C. S., Griffin, L. C., Dunn, K. E., Leung, L. L.
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