

Stanford



Hannah Valentine

Professor of Medicine (Cardiovascular)
Medicine - Cardiovascular Medicine

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ACADEMIC CONTACT INFORMATION

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Bio

BIO

Hannah Valentine received her M.B.B.S. degree from London University, cardiology fellowship at Stanford, and Doctor of Medicine from London University. She was appointed Assistant Professor of Medicine, rising to full Professor of Medicine in 2000, and becoming the inaugural Senior Associate Dean for Diversity and Leadership, in 2004. She pursued a data-driven transformative approach to this work, receiving the NIH director's pathfinder award. Dr. Francis Collins, NIH director, recruited her in 2014 as the inaugural NIH Chief Officer for Scientific Workforce diversity, and as a tenured investigator in the National Heart, Lung, and Blood Institute's intramural research program where she established the laboratory of transplantation genomics. Dr. Valentine is a nationally recognized pioneer in her field, with over 200 peer-reviewed publications, patents, and sustained NIH funding. She was elected to the National Academy of Medicine in 2020 for both her pioneering research in organ transplantation and workforce diversity.

CLINICAL FOCUS

- Cardiology (Heart)
- Heart Lung Transplantation
- Cardiovascular Disease

ACADEMIC APPOINTMENTS

- Professor - University Medical Line, Medicine - Cardiovascular Medicine
- Member, Cardiovascular Institute

ADMINISTRATIVE APPOINTMENTS

- Professor of Medicine, Cardiovascular Medicine, (2020- present)
- Senior Associate Dean for Diversity and Faculty Development, School of Medicine, (2005-2014)
- President, Western State Affiliation, American Heart Association, (2004-2005)

HONORS AND AWARDS

- Exceptional Contributions to Education in Medicine, Stanford Department of Medicine (2006)

- President's Awards for Excellence Through Diversity, Stanford University (2013)
- Elected to Association of American Physicians, Association of American Physicians (2018)
- NIH Director's Pathfinder Award for Diversity in the Scientific Workforce, NIH (2010)
- Elected to the National Academy of Medicine in 2020., For both her pioneering research in organ transplantation and workforce diversity. (2020)

PROFESSIONAL EDUCATION

- Internship: St George's Hospital Medical School (1979) England
- Residency: Guy's Hospital Medical School (1982) England
- Fellowship: Stanford University Cardiovascular Medicine Fellowship (1986) CA
- Fellowship: Hammersmith Hospital (1984) England
- Residency: Brompton Hospital (1982) UK
- Board Certification: Internal Medicine, Royal College of Physicians-U.K. (1981)
- Residency: St George's Hospital Medical School (1981) England
- Medical Education: St George's Hospital Medical School (1978) England
- MBBS, St Georges Hosp/London Univ, UK , Medicine (MD equivalent) (1978)
- MRCP, London University, London, UK , Internal Medicine (1982)
- Registrar, Hammersmith Hospital, London, UK , Cardiology Fellow (1984)
- Fellow, Stanford University , Cardiology (1986)
- MD, London University, London, UK , Cardiology (PhD equivalent) (1988)

Research & Scholarship

CURRENT RESEARCH AND SCHOLARLY INTERESTS

Prior to my expanded administrative role, my lab focused on understanding the mechanism mediating acute and chronic allograft failure, in particular on the role of microvascular injury in acute allograft failure and the mechanisms of mediating transplant coronary artery disease.

1. Role of microvascular injury in acute allograft failure. We observed decreased cardiac allograft function in patients to be significantly associated with loss of microvascular cell surface markers, consistent with altered biology of the vascular endothelium or injury, and with up-regulation of cytokines such as IL-6, IL-10, TGF-b, and TNF-a. Decreased cardiac allograft function, in particular diastolic dysfunction, was highly predictive of allograft vascular disease and poor outcome in long-term patients. To further characterize cellular and molecular mechanisms we developed quantitative methods to monitor allograft function and correlate it with cytokine expression in a rat heterotopic transplant model. We developed echocardiographic markers of systolic and diastolic function and found decreasing systolic and diastolic function highly correlated with up-regulation of IL-6 expression.

2. Mechanisms mediating allograft dysfunction. We observed the major risk factors for transplant atherosclerosis in patients to be metabolic (hyperglycemia, hypertriglyceridemia, and low HDL). To further study cellular and molecular mechanisms mediating this process, we recapitulated metabolic abnormalities in the rat heart transplant model and confirmed rapid development of transplant atherosclerosis.

My current focus is on clinical and translational research, expanding on my earlier laboratory-based research on the mechanisms of allograft failure and vasculopathy. I study the role of sub-clinical cytomegalovirus infection in the development of cardiac allograft vasculopathy (CAV), and the effect of antiviral agents in preventing CAV, work that has been supported by a program project grant from NIH. Translating my earlier work on gene expression profiling, I conduct clinical trials on the application of this technology for non-invasive diagnosis of transplant rejection. This work, recently published in the New England Journal of Medicine, showed that

peripheral blood leukocyte gene expression can be used safely for monitoring rejection in heart transplant recipients, with a substantial decrease in the number of endomyocardial biopsies performed. On going studies in this area will include application of the technology to diagnose CAV; monitor level of immunosuppression in a patient to guide therapy; and individualization of immunosuppression. My current research, in collaboration with bio-X faculty, uses the new technologies of whole genome scans, to detect the appearance of donor DNA in the recipient's blood as a novel, non-invasive marker of allograft damage due to acute rejection. Ongoing studies include confirmation studies of the utility, and then external validation in a randomized controlled trial.

CLINICAL TRIALS

- Heartsbreath Test for Heart Transplant Rejection, Recruiting
- Angiotensin Converting Enzyme (ACE) Inhibition and Cardiac Allograft Vasculopathy, Not Recruiting
- Clinical Trial of Pioglitazone for Prevention of Cardiac Allograft Vasculopathy After Heart Transplantation, Not Recruiting
- Genome Transplant Dynamics: Non-invasive Sequencing-based Diagnosis of Rejection, Not Recruiting
- IMAGE: A Comparison of AlloMap Molecular Testing and Traditional Biopsy-based Surveillance for Heart Transplant Rejection, Not Recruiting

Publications

PUBLICATIONS

- **Sex-Specific Patterns of Donor-Derived Cell-Free DNA in Heart Transplant Rejection: An Analysis from the Genomic Research Alliance for Transplantation (GRAFT).** *The Journal of heart and lung transplantation : the official publication of the International Society for Heart Transplantation* DeFilippis, E. M., Sweigart, B., Khush, K. K., Shah, P., Agbor-Enoh, S., Valentine, H. A., Vest, A. R. 2024
- **Racial Differences in Donor-Derived Cell-Free DNA and Mitochondrial DNA After Heart Transplantation, on Behalf of the GRAFT Investigators.** *Circulation. Heart failure* Shah, P., Agbor-Enoh, S., Lee, S., Andargie, T. E., Sinha, S. S., Kong, H., Henry, L., Park, W., McNair, E., Tchoukina, I., Shah, K. B., Najjar, S. S., Hsu, et al 2024: e011160
- **Neural responses to gender-based microaggressions in academic medicine.** *Journal of neuroscience research* Balters, S., Foland-Ross, L. C., Bruno, J., Periyakoil, V. S., Valentine, H., Reiss, A. L. 2023
- **The Time to Act Is Now: Racial Disparities After Heart Transplantation.** *Circulation* Khush, K. K., Valentine, H. A. 2023; 148 (3): 207-209
- **Time to open up NEW SCIENTIST** Elovitz, M., Quake, S., Valentine, H. 2023; 246 (3444): 21
- **Hannah A. Valentine, MD, Stanford University School of Medicine** *CELL* Valentine, H. A. 2023; 186 (12): 2524-2526
- **Applying Genomics to Unravel Health Disparities in Organ Transplantation: Paul I. Terasaki State-of-the-art Lecture; American Transplant Congress 2021.** *Transplantation* Valentine, H. A. 2022
- **Plasma Cell-Free DNA Predicts Survival and Maps Specific Sources of Injury in Pulmonary Arterial Hypertension.** *Circulation* Brusca, S. B., Elinoff, J. M., Zou, Y., Jang, M. K., Kong, H., Demirkale, C. Y., Sun, J., Seifuddin, F., Pirooznia, M., Valentine, H. A., Tanba, C., Chaturvedi, A., Graninger, et al 2022: 101161CIRCULATIONAHA121056719
- **Equity in Transplantation: A Commitment for Progress in Troubled Times** *TRANSPLANT INTERNATIONAL* Berney, T., Ulasi, I. I., Balleste, C., Martins, P. N., Bellini, M., Valentine, H. A., Potena, L.

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- **Circulating microRNAs in cellular and antibody-mediated heart transplant rejection.** *The Journal of heart and lung transplantation : the official publication of the International Society for Heart Transplantation*
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2022
- **Higher levels of allograft injury in black patients early after heart transplantation.** *The Journal of heart and lung transplantation : the official publication of the International Society for Heart Transplantation*
Doshi, A., Shah, K. B., Agbor-Enoh, S., Tushak, Z., Garcia, V., Kong, H., Jang, M. K., Hsu, S., Feller, E. D., Rodrigo, M. E., Najjar, S. S., Tunc, I., Yang, et al
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- **A commitment to scientific equity from a philanthropic funder.** *Nature medicine*
Bargmann, C., Claiborne, A., Valentine, H.
2021
- **Can We Predict Rejection Early After Heart Transplantation?** *Circulation*
Fearon, W. F., Valentine, H. A.
2021; 144 (18): 1473-1475
- **Ending Sexual Harassment in Science: Designing and Administering a Survey That Can Lead to an Improved Organizational Climate.** *Academic medicine : journal of the Association of American Medical Colleges*
Valentine, H. A., Le Fauve, C. E., Morris, K. A., Riley, W. T.
2021
- **Response by Shah et al to Letter Regarding Article, "Cell-Free DNA to Detect Heart Allograft Acute Rejection".** *Circulation*
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2021; 144 (10): e198-e199
- **The COVID-19 Vaccine and the Black Community: Addressing the Justified Questions.** *Journal of racial and ethnic health disparities*
Laurencin, C. T., Valentine, H., Yancy, C., Jones, C. P., Bright, C.
2021
- **A Pioneer in Transplantation Genomics, Inclusion, and Diversity A Conversation With Hannah Valentine, MBBS, MD** *CIRCULATION*
Valentine, H., Farr, M.
2021; 143 (24): 2321-2326
- **Cell-Free DNA to Detect Heart Allograft Acute Rejection.** *Circulation*
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2020
- **Everyday Heroism: Maintaining Organizational Cultures of Wellness and Inclusive Excellence Amid Simultaneous Pandemics.** *Academic medicine : journal of the Association of American Medical Colleges*
Fassiotto, M. n., Valentine, H. n., Shanafelt, T. n., Maldonado, Y. n.
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- **NHLBI-SPONSORED PROSPECTIVE STUDY TO VALIDATE THE USE OF DONOR-DERIVED CELL-FREE DNA TO DETECT LUNG-TRANSPLANT REJECTION - INTERIM RESULTS**
Agbor, S., Berry, G., Marboe, C., Tunc, I., Jackson, A., Shah, P., Fideli, U., Timofte, I., Brown, A. W., Iacono, A., Nathan, S., Jang, M., Orens, et al
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- **Early invasive assessment of the coronary microcirculation predicts subsequent acute rejection after heart transplantation** *INTERNATIONAL JOURNAL OF CARDIOLOGY*
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- **Early invasive assessment of the coronary microcirculation predicts subsequent acute rejection after heart transplantation.** *International journal of cardiology*
Okada, K., Honda, Y., Luikart, H., Yock, P. G., Fitzgerald, P. J., Yeung, A. C., Valentine, H. A., Khush, K. K., Fearon, W. F.
2019
- **Donor-derived cell-free DNA predicts allograft failure and mortality after lung transplantation.** *EBioMedicine*
Agbor-Enoh, S., Wang, Y., Tunc, I., Jang, M. K., Davis, A., De Vlaminck, I., Luikart, H., Shah, P. D., Timofte, I., Brown, A. W., Marishta, A., Bhatti, K., Gorham, et al
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- **Association of Endothelin-1 with Accelerated Cardiac Allograft Vasculopathy and Late Mortality Following Heart Transplantation.** *Journal of cardiac failure*
Parikh, R. V., Khush, K., Pargaonkar, V. S., Luikart, H., Grimm, D., Yu, M., Akada, K., Honda, Y., Yeung, A. C., Valentine, H., Fearon, W. F.
2018
- **Usefulness of Asymmetric Dimethylarginine to Predict Outcomes After Heart Transplantation** *AMERICAN JOURNAL OF CARDIOLOGY*
Parikh, R., Khush, K., Luikart, H., Sakarovich, C., Lee, J., Desai, M., Valentine, H., Yeung, A. C., Fearon, W. F.
2018; 122 (10): 1707–11
- **Late manifestation of alloantibody-associated injury and clinical pulmonary antibody-mediated rejection: Evidence from cell-free DNA analysis** *JOURNAL OF HEART AND LUNG TRANSPLANTATION*
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- **IMPACT OF ENDOTHELIN-1 ON CARDIAC ALLOGRAFT VASCULOPATHY, LATE MORTALITY AND RE-TRANSPLANTATION FOLLOWING HEART TRANSPLANTATION**
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- **The effect of negative remodeling on fractional flow reserve after cardiac transplantation** *INTERNATIONAL JOURNAL OF CARDIOLOGY*
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 - **Designing a physician leadership development program based on effective models of physician education.** *Health care management review*
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 - **Coronary Endothelial Dysfunction and the Index of Microcirculatory Resistance as a Marker of Subsequent Development of Cardiac Allograft Vasculopathy.** *Circulation*
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 - **Angiotensin-Converting Enzyme Inhibition Early After Heart Transplantation.** *Journal of the American College of Cardiology*
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 - **Impact of Asymmetric Dimethylarginine on Coronary Physiology Early After Heart Transplantation.** *The American journal of cardiology*
Parikh, R. V., Khush, K. K., Luikart, H. n., Pargaonkar, V. S., Kobayashi, Y. n., Lee, J. H., Sinha, S. n., Cohen, G. n., Valentine, H. A., Yeung, A. C., Fearon, W. F.
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