Rheumatology



Division Data Summary

Research and Training Details

Number of Faculty	13
Number of Research Students	1
Number of Support Personnel	51
Direct Annual Grant Support	\$4,090,159
Direct Annual Industry Support	\$669,616
Peer Reviewed Publications	58
Clinical Activities and Training	
Number of Clinical Staff	3
Number of Clinical Fellows	6

Division Photo



Row 1 (seated): J Harley

Row 2: H Brunner, J Huggins, E Morgan DeWitt, T

Ting

1

5329

Row 3: E Giannini, S Thornton

Row 4: S Thompson, D Lovell, M Hendrickson

Row 5: A Grom

Significant Publications

Number of Other Students

Outpatient Encounters

Brunner HI, Huggins J, Klein-Gitelman MS. Pediatric SLE--towards a comprehensive management plan. *Nat Rev Rheumatol*. 7(4):225-33. Apr, 2011.

Significance: The many quality improvement initiatives and extensive experience of the Division physicians in caring for patients with systemic lupus erythematosus has led to better management of these patients, which is summarized in this review.

Sumegi J, Barnes MG, Nestheide SV, Molleran-Lee S, Villanueva J, Zhang K, Risma KA, Grom AA, Filipovich AH. Gene expression profiling of peripheral blood mononuclear cells from children with active hemophagocytic lymphohistiocytosis. *Blood.* 117(15), e151-160.

Significance: A deeper and much more fundamental understanding of the potentially deadly hemophagocytic lymphohisticcytosis is emerging. The division faculty is making a major contribution to the aggregate of this knowledge with the identification of critical elements in the pathophysiology of this condition. This study is the first genome wide study of gene expression in this syndrome.

Shen N, Fu Q, Deng Y, Qian X, Zhao J, Kaufman KM, Wu YL, Yu CY, Tang Y, Chen JY, Yang W, Wong M, Kawasaki A, Tsuchiya N, Sumida T, Kawaguchi Y, Howe HS, Mok MY, Bang SY, Liu FL, Chang DM, Takasaki Y, Hashimoto H, Harley JB, Guthridge JM, Grossman JM, Cantor RM, Song YW, Bae SC, Chen S, Hahn BH, Lau YL, Tsao BP. Sex-specific association of X-linked Toll-like receptor 7 (TLR7) with male systemic lupus erythematosus. *Proc Natl Acad Sci U S A.*107(36): p. 15838-43. 2010.

Significance: The toll-like receptors are an important component of the innate immune response. TLR7 is a

gene on the X chromosome. Since women and girls have systemic lupus erythematosus much more frequently than men and boys (~10-fold), we would expect that genes on the X chromosome would be more important in women than it is in men. Surprisingly, variants in TLR7 are much more strongly associated with male lupus than female lupus in Asians.

Beukelman T, Patkar NM, Saag KG, Tolleson-Rinehart S, Cron RQ, DeWitt EM, Ilowite NT, Kimura Y, Laxer RM, Lovell DJ, Martini A, Rabinovich CE, Ruperto N. 2011 American College of Rheumatology recommendations for the treatment of juvenile idiopathic arthritis: Initiation and safety monitoring of therapeutic agents for the treatment of arthritis and systemic features. *Arthritis Care & Research*. 63: 465–482. 2011.

Significance: These represent the first evidence based treatment guidelines for children with Juvenile Idiopathic Arthritis and due to the rigor of the development process they have been officially endorsed and adopted by the American College of Rheumatology. Three members of the CCHMC Division of Rheumatology (Esi Morgan Dewitt, Janalee Taylor and Daniel Lovell) were key participants in the effort.

Lovell DJ, Passo MH, Beukelman T, Bowyer SL, Gottlieb BS, Henrickson M, Ilowite NT, Kimura Y, DeWitt EM, Segerman J, Stein LD, Taylor J, Vehe RK, Giannini EH. Measuring process of arthritis care: A proposed set of quality measures for the process of care in juvenile idiopathic arthritis. *Arthritis Care & Research*. 63: 10–16. 2011.

Significance: These represent the first nationally developed quality measures for the care of children with juvenile idiopathic arthritis. Five members of the CCHMC Division of Rheumatology (Janalee Taylor and Drs. Giannini, Henrickson, Lovell and Morgan Dewitt) initiated and lead this national effort. These quality measures are being used by an international quality improvement network, PR-COIN, being led by a CCHMC Division of Rheumatology faculty member, Esi Morgan Dewitt, in collaboration with the CCHMC Anderson Center.

Division Highlights

Edward Giannini, MSc, DrPH

Dr. Giannini is the recipient of a Life-Time Achievement Award which was bestowed upon him at the Pediatric Rheumatology Symposium (PRSYM) that was held in Miami, Florida in June 2011. The Life-Time Achievement awardees are selected by a Committee representing the American Academy of Pediatrics and the American College of Rheumatology. Recipients are considered those whose career has demonstrated a sustained and lasting contribution to the field of rheumatology and rheumatology health professionals.

Esi Morgan DeWitt, MD, MSCE

PR-COIN - The Pediatric Rheumatology Care and Outcomes Improvement Network (PR-COIN) is a multi-center quality improvement learning collaborative formed to improve the outcomes of care of children with juvenile idiopathic arthritis and to accelerate adoption of evidence into medical practice. Our collaborative approach is based on the Institute of Healthcare Improvement (IHI) Breakthrough Series Collaborative Model and utilizes the Model for Improvement to guide performance improvement activities. Using the American College of Rheumatology Rheumatology Clinical Registry (ACR RCR), PR-COIN is designed to aggregate data from participating pediatric rheumatology practices to study performance of sites on both process measures of care delivery and measures of patient outcomes to better understand which disease management approaches are optimal. In the past year we formed our project team with leadership from the Division of Rheumatology in partnership with the Anderson Center for Health Systems Excellence with expertise in QI science and project management. After project launch, we customized and piloted the electronic data capture system. Twelve centers in the US and Canada have joined the collaborative. Initial project activities include collection of baseline data and training teams in performance improvement methodology, prior to testing change strategies

at team sites that are based on a chronic illness care model. Teams convene during three Learning Sessions over a 12 month period where QI experts offer intensive training and coaching. Sessions were held June 2011, and planned for September 2011 and March 2012. Site and aggregate data feedback reports are provided monthly to enable teams to assess progress on quality measures.

Division Collaboration

Allergy » Marc Rothenberg

John Harley and Marc Rothenberg collaboration: Candidate gene study of Eosinophilic esophagitis

Human Genetics » Greg Grabowski; Mehdi Keddache; Sarah Zimmerman

John Harley, Sarah Zimmerman, Mehdi Keddache, and Greg Grabowski collaboration: Genome wide association study of height from routine clinical samples

Molecular Immunology » Chris Karp

John Harley and Chris Karp collaboration: PXK association with lupus

Immunobiology » Marsha Wills Karp

John Harley and Marsha Wills Karp collaboration: Genome wide study of asthma in an isolated population

Bioinformatics » John Hutton; Michael Wagner

John Harley Michael Wagner and John Hutton collaboration: Genetic association analysis of systemic lupus erythematosus

Anderson Center for Health Systems Excellence » Peter Margolis; Carole Lannon

Esi Morgan DeWitt, Peter Margolis and Carole Lannon collaboration: PR-COIN

Behavioral Medicine; **Pain Management**; **Pulmonary**; **Rehabilitation** » Susmita Kashikar-Zuck; Kenneth Goldschneider; Michael Seid; Jilda Vargus-Adams

Esi Morgan DeWitt, Susmita Kashikar-Zuck, Kenneth Goldschneider, Michael Seid, and Jilda Vargus-Adams collaboration: The Patient Reported Outcomes Measurement Information System (PROMIS) - Cincinnati Children's Hospital Medical Center (CCHMC) is a research site in the NIH Roadmap PROMIS (Patient-Reported Outcomes Measurement Information System) cooperative network which mission is to use measurement science to create a state-of-the-art assessment system for self-reported health. The CCHMC PROMIS research project has two aims. The first aim it to perform longitudinal validation of current PROMIS pediatric PRO measures in the domains of physical function (upper extremity, mobility), pain interference, fatigue, anger, anxiety, depressive symptoms, and peer relationships in pediatric patients with chronic pain conditions, juvenile idiopathic arthritis, or cerebral palsy. An extensive qualitative interview study in children with these conditions was conducted as an initial step. The longitudinal portion of the study is well underway. The second aim is to develop new PRO item banks to measure pain behavior, and pain quality, including pain severity, in pediatric patients. Candidate items for testing have been created following a rigorous standard PROMIS methodology and are ready for large scale cross-sectional validation. The validation sample for the new pain assessment tools includes children with chronic and recurrent pain, including juvenile idiopathic arthritis, widespread musculoskeletal or regional pain syndromes, and sickle cell anemia. Use of IRT based PROMIS measures will allow for efficiency of PRO measurement, potential for increased sensitivity to measure change in health status over time, and facilitate comparison of an individual's results to population norms in patients with rheumatic diseases and other chronic disease cohorts.

Experimental Hematology; Human Genetics » Matthew Flick; Xiaoyang Qi

Sherry Thornton Collaboration: with Dr. Matthew Flick in Experimental Hematology and Dr. Xiaoyang Qi, Division of Human Genetics CCHMC and now Division of Hematology-Oncology-UC College of Medicine on a

project funded by the local CTSA entitled SapC-DOPS Agents: Imaging in Arthritis. This project involves assessment of the ability of SapC-DOPS fluorescently labeled agent to localize to arthritic joints. A manuscript will be submitted in the very near future describing our findings that SapC-DOPS localizes to arthritic joints. In addition a revision of an R21 has been submitted which determines whether SapC-DOPS can detect subclinical arthritis and can be used as a clinical marker to assess response to therapy in arthritis models.

Experimental Hematology » Matthew Flick

Sherry Thornton and Matthew Flick Collaboration: assessment of hemostatic factors in arthritis which generated a manuscript

Faculty Members

John Harley, MD, PhD, Professor

Division Director

Research Interests

Hermine Brunner, MD, MSc, Associate Professor

Research Interests

Edward H. Giannini, MSc, DrPH, Professor

Research Interests

David N. Glass, MD, Professor

Research Interests

Thomas Griffin, MD, PhD, Associate Professor

Research Interests

Alexei A. Grom, MD, Associate Professor

Research Interests

Michael Henrickson, MD, MPH, Associate Professor

Clinical Director

Research Interests

Jennifer Huggins, MD, Assistant Professor

Fellowship Director

Research Interests

Daniel Joe Lovell, MD, MPH, Professor

Joseph E. Levinson Endowed Chair in Pediatric Rheumatology

Research Interests

Esi Morgan DeWitt, MD, MSCE, Assistant Professor

Research Interests

Susan Thompson, PhD, Associate Professor

Associate Director

Research Interests

Sherry Thornton, PhD, Assistant Professor

Research Interests

Tracy Ting, MD, Assistant Professor

Research Interests

Clinical Staff Members

• Janalee Taylor, MSN, RN, CNP

Trainees

- Lu Pai-Yue, MD, PGY-IV, Cincinnati Children's Hospital Medical Center
- Moussa El-Hallak, MD, PGY-IV, Memorial University Medical Center
- Rina Mina, MD, PGY-VI, Downstate Medical Center New York
- Annette Lopez-Martinez, MD, PGY-VI, University of Puerto Rico Pediatric Hospital
- David Moser, DO, PGY-V, United States Army, Pediatrics
- Keith Sikora, MD, PGY-V, Johns Hopkins Hospital

Significant Accomplishments

Division and CAGE Achieve Milestones

The Division of Rheumatology and the Center for Autoimmune Genomics and Etiology (CAGE) achieved many milestones this past year.

Work led by Alexei Grom, MD, identified a pathway that is important for the development of the sometimes deadly macrophage activation syndrome. Susan Thompson, PhD, led work that provided deep genetic insight into the variations of DNA that predispose to juvenile onset idiopathic arthritis. Dan Lovell, MD, MPH, Ed Giannini, DrPH, and Hermine Brunner, MD, MSc, have shown that the new biological therapies are spectacularly successful; their critically important work has become the new standard of therapy and is helping many thousands of afflicted children avoid the life-long disability of chronic destructive arthritis.

Esi Morgan DeWitt, MD, MSc, has achieved better compliance and therapeutic outcomes in juvenile onset rheumatoid arthritis by applying quality-improvement interventions. DeWitt and Brunner are co-leading efforts to effectively align research, clinical care and quality improvement to provide high-value care locally and are leading international initiatives to develop quality indicators and benchmarks for pediatric rheumatology care with focus on systemic lupus erythematosus and juvenile idiopathic arthritis.

Biomarker development efforts are synergistic with development of clinical trial outcome measures that set the stage for testing novel medications for children with SLE. John Harley, MD, PhD, has led an effort to identify the genes that cause systemic lupus erythematosus, now numbering more than 40, followed by progress in discovering the mechanisms and pathways through which they cause lupus.

Division Publications

- Adrianto I, Wen F, Templeton A, Wiley G, King JB, Lessard CJ, Bates JS, Hu Y, Kelly JA, Kaufman KM, Guthridge JM, Alarcon-Riquelme ME, Anaya JM, Bae SC, Bang SY, Boackle SA, Brown EE, Petri MA, Gallant C, Ramsey-Goldman R, Reveille JD, Vila LM, Criswell LA, Edberg JC, Freedman BI, Gregersen PK, Gilkeson GS, Jacob CO, James JA, Kamen DL, Kimberly RP, Martin J, Merrill JT, Niewold TB, Park SY, Pons-Estel BA, Scofield RH, Stevens AM, Tsao BP, Vyse TJ, Langefeld CD, Harley JB, Moser KL, Webb CF, Humphrey MB, Montgomery CG, Gaffney PM. Association of a functional variant downstream of TNFAIP3 with systemic lupus erythematosus. Nat Genet. 2011; 43:253-8.
- 2. Antonchak MA, Saoudian M, Khan AR, Brunner HI, Luggen ME. Cognitive dysfunction in patients with systemic lupus erythematosus: a controlled study. *J Rheumatol*. 2011; 38:1020-5.

- 3. Ardoin SP, Schanberg LE, Sandborg C, Yow E, Barnhart HX, Mieszkalski K, Ilowite NT, von Scheven E, Eberhard A, Levy DM, Kimura Y, Silverman E, Bowyer SL, Punaro L, Singer NG, Sherry DD, McCurdy D, Klein-Gitelman M, Wallace C, Silver R, Wagner-Weiner L, Higgins GC, Brunner HI, Jung LK, Imundo L, Soep JB, Reed AM. Laboratory markers of cardiovascular risk in pediatric SLE: the APPLE baseline cohort. *Lupus*. 2010; 19:1315-25.
- 4. Barnes MG, Grom AA, Thompson SD, Griffin TA, Luyrink LK, Colbert RA, Glass DN. Biologic similarities based on age at onset in oligoarticular and polyarticular subtypes of juvenile idiopathic arthritis. *Arthritis Rheum.* 2010; 62:3249-58.
- 5. Beukelman T, Patkar NM, Saag KG, Tolleson-Rinehart S, Cron RQ, DeWitt EM, Ilowite NT, Kimura Y, Laxer RM, Lovell DJ, Martini A, Rabinovich CE, Ruperto N. **2011 American College of Rheumatology recommendations for the treatment of juvenile idiopathic arthritis: initiation and safety monitoring of therapeutic agents for the treatment of arthritis and systemic features.** *Arthritis Care Res (Hoboken)***. 2011; 63:465-82.**
- 6. Bruner BF, Vista ES, Wynn DM, Harley JB, James JA. **Anti-neutrophil cytoplasmic antibodies target** sequential functional proteinase 3 epitopes in the sera of patients with Wegener's granulomatosis. *Clin Exp Immunol.* 2010; 162:262-70.
- 7. Brunner HI, Higgins GC, Klein-Gitelman MS, Lapidus SK, Olson JC, Onel K, Punaro M, Ying J, Giannini EH. Minimal clinically important differences of disease activity indices in childhood-onset systemic lupus erythematosus. *Arthritis Care Res (Hoboken)*. 2010; 62:950-9.
- 8. Brunner HI, Huggins J, Klein-Gitelman MS. Pediatric SLE--towards a comprehensive management plan. *Nat Rev Rheumatol.* 2011; 7:225-33.
- 9. Chung SA, Taylor KE, Graham RR, Nititham J, Lee AT, Ortmann WA, Jacob CO, Alarcon-Riquelme ME, Tsao BP, Harley JB, Gaffney PM, Moser KL, Petri M, Demirci FY, Kamboh MI, Manzi S, Gregersen PK, Langefeld CD, Behrens TW, Criswell LA. **Differential genetic associations for systemic lupus erythematosus based on anti-dsDNA autoantibody production**. *PLoS Genet*. 2011; 7:e1001323.
- 10. Clancy RM, Marion MC, Kaufman KM, Ramos PS, Adler A, Harley JB, Langefeld CD, Buyon JP. Identification of candidate loci at 6p21 and 21q22 in a genome-wide association study of cardiac manifestations of neonatal lupus. *Arthritis Rheum*. 2010; 62:3415-24.
- 11. Clericuzio C, Harutyunyan K, Jin W, Erickson RP, Irvine AD, McLean WH, Wen Y, Bagatell R, Griffin TA, Shwayder TA, Plon SE, Wang LL. **Identification of a novel C16orf57 mutation in Athabaskan patients with Poikiloderma with Neutropenia**. *Am J Med Genet A*. 2011; 155A:337-42.
- 12. Crowe SR, Ash LL, Engler RJ, Ballard JD, Harley JB, Farris AD, James JA. Select human anthrax protective antigen epitope-specific antibodies provide protection from lethal toxin challenge. *J Infect Dis.* 2010; 202:251-60.
- 13. Crowe SR, Garman L, Engler RJ, Farris AD, Ballard JD, Harley JB, James JA. **Anthrax vaccination induced anti-lethal factor IgG: fine specificity and neutralizing capacity**. *Vaccine*. 2011; 29:3670-8.
- 14. Dillon S, Aggarwal R, Harding JW, Li LJ, Weissman MH, Li S, Cavett JW, Sevier ST, Ojwang JW, D'Souza A, Harley JB, Scofield RH. **Klinefelter's syndrome (47,XXY) among men with systemic lupus erythematosus**. *Acta Paediatr*. 2011; 100:819-23.
- Dinan MA, Compton KL, Dhillon JK, Hammill BG, Dewitt EM, Weinfurt KP, Schulman KA. Use of patient-reported outcomes in randomized, double-blind, placebo-controlled clinical trials. *Med Care*. 2011; 49:415-9.
- 16. Flick MJ, Chauhan AK, Frederick M, Talmage KE, Kombrinck KW, Miller W, Mullins ES, Palumbo JS, Zheng X, Esmon NL, Esmon CT, Thornton S, Becker A, Pelc LA, Di Cera E, Wagner DD, Degen JL. The development of inflammatory joint disease is attenuated in mice expressing the anticoagulant prothrombin mutant W215A/E217A. *Blood*. 2011; 117:6326-37.

- 17. Fu Q, Zhao J, Qian X, Wong JL, Kaufman KM, Yu CY, Mok MY, Harley JB, Guthridge JM, Song YW, Cho SK, Bae SC, Grossman JM, Hahn BH, Arnett FC, Shen N, Tsao BP. **Association of a functional IRF7 variant with systemic lupus erythematosus**. *Arthritis Rheum*. 2011; 63:749-54.
- 18. Furnrohr BG, Wach S, Kelly JA, Haslbeck M, Weber CK, Stach CM, Hueber AJ, Graef D, Spriewald BM, Manger K, Herrmann M, Kaufman KM, Frank SG, Goodmon E, James JA, Schett G, Winkler TH, Harley JB, Voll RE. Polymorphisms in the Hsp70 gene locus are genetically associated with systemic lupus erythematosus. *Ann Rheum Dis.* 2010; 69:1983-9.
- 19. Giannini EH, Ilowite NT, Lovell DJ, Wallace CA, Rabinovich CE, Reiff A, Higgins G, Gottlieb B, Chon Y, Zhang N, Baumgartner SW. Effects of long-term etanercept treatment on growth in children with selected categories of juvenile idiopathic arthritis. *Arthritis Rheum.* 2010; 62:3259-64.
- 20. Gottlieb AB, Gordon K, Giannini EH, Mease P, Li J, Chon Y, Maddox J, Weng HH, Wajdula J, Lin SL, Baumgartner SW. Clinical trial safety and mortality analyses in patients receiving etanercept across approved indications. *J Drugs Dermatol*. 2011; 10:289-300.
- 21. Grom AA, Mellins ED. **Macrophage activation syndrome: advances towards understanding pathogenesis**. *Curr Opin Rheumatol*. 2010; 22:561-6.
- 22. Heinlen LD, Ritterhouse LL, McClain MT, Keith MP, Neas BR, Harley JB, James JA. Ribosomal P autoantibodies are present before SLE onset and are directed against non-C-terminal peptides. *J Mol Med.* 2010; 88:719-27.
- 23. Hinks A, Martin P, Flynn E, Eyre S, Packham J, Barton A, Worthington J, Thomson W. Subtype specific genetic associations for juvenile idiopathic arthritis: ERAP1 with the enthesitis related arthritis subtype and IL23R with juvenile psoriatic arthritis. *Arthritis Res Ther.* 2011; 13:R12.
- 24. Hughes T, Kim-Howard X, Kelly JA, Kaufman KM, Langefeld CD, Ziegler J, Sanchez E, Kimberly RP, Edberg JC, Ramsey-Goldman R, Petri M, Reveille JD, Martin J, Brown EE, Vila LM, Alarcon GS, James JA, Gilkeson GS, Moser KL, Gaffney PM, Merrill JT, Vyse TJ, Alarcon-Riquelme ME, Nath SK, John BH, Sawalha AH. Finemapping and transethnic genotyping establish IL2/IL21 genetic association with lupus and localize this genetic effect to IL21. Arthritis Rheum. 2011; 63:1689-97.
- 25. Kashikar-Zuck S, Flowers SR, Verkamp E, Ting TV, Lynch-Jordan AM, Graham TB, Passo M, Schikler KN, Hashkes PJ, Spalding S, Banez G, Richards MM, Powers SW, Arnold LM, Lovell D. **Actigraphy-based physical activity monitoring in adolescents with juvenile primary fibromyalgia syndrome**. *J Pain*. 2010; 11:885-93.
- 26. Kashikar-Zuck S, Johnston M, Ting TV, Graham BT, Lynch-Jordan AM, Verkamp E, Passo M, Schikler KN, Hashkes PJ, Spalding S, Banez G, Richards MM, Powers SW, Arnold LM, Lovell D. Relationship between school absenteeism and depressive symptoms among adolescents with juvenile fibromyalgia. *J Pediatr Psychol.* 2010; 35:996-1004.
- 27. Kashikar-Zuck S, Parkins IS, Ting TV, Verkamp E, Lynch-Jordan A, Passo M, Graham TB. Controlled follow-up study of physical and psychosocial functioning of adolescents with juvenile primary fibromyalgia syndrome. *Rheumatology (Oxford)*. 2010; 49:2204-9.
- 28. Kim-Howard X, Maiti AK, Anaya JM, Bruner GR, Brown E, Merrill JT, Edberg JC, Petri MA, Reveille JD, Ramsey-Goldman R, Alarcon GS, Vyse TJ, Gilkeson G, Kimberly RP, James JA, Guthridge JM, Harley JB, Nath SK. ITGAM coding variant (rs1143679) influences the risk of renal disease, discoid rash and immunological manifestations in patients with systemic lupus erythematosus with European ancestry. *Ann Rheum Dis.* 2010; 69:1329-32.
- 29. Lessard CJ, Adrianto I, Kelly JA, Kaufman KM, Grundahl KM, Adler A, Williams AH, Gallant CJ, Anaya JM, Bae SC, Boackle SA, Brown EE, Chang DM, Criswell LA, Edberg JC, Freedman BI, Gregersen PK, Gilkeson GS, Jacob CO, James JA, Kamen DL, Kimberly RP, Martin J, Merrill JT, Niewold TB, Park SY, Petri MA, Pons-

- Estel BA, Ramsey-Goldman R, Reveille JD, Song YW, Stevens AM, Tsao BP, Vila LM, Vyse TJ, Yu CY, Guthridge JM, Bruner GR, Langefeld CD, Montgomery C, Harley JB, Scofield RH, Gaffney PM, Moser KL. Identification of a systemic lupus erythematosus susceptibility locus at 11p13 between PDHX and CD44 in a multiethnic study. *Am J Hum Genet*. 2011; 88:83-91.
- 30. Lovell DJ, Passo MH, Beukelman T, Bowyer SL, Gottlieb BS, Henrickson M, Ilowite NT, Kimura Y, DeWitt EM, Segerman J, Stein LD, Taylor J, Vehe RK, Giannini EH. **Measuring process of arthritis care: a proposed set of quality measures for the process of care in juvenile idiopathic arthritis.** *Arthritis Care Res* (*Hoboken*). 2011; 63:10-6.
- 31. Martini A, Lovell DJ. **Juvenile idiopathic arthritis: state of the art and future perspectives**. *Ann Rheum Dis*. 2010; 69:1260-3.
- 32. Mellins ED, Macaubas C, Grom AA. Pathogenesis of systemic juvenile idiopathic arthritis: some answers, more questions. *Nat Rev Rheumatol.* 2011; 7:416-26.
- 33. Moncrieffe H, Hinks A, Ursu S, Kassoumeri L, Etheridge A, Hubank M, Martin P, Weiler T, Glass DN, Thompson SD, Thomson W, Wedderburn LR. **Generation of novel pharmacogenomic candidates in response to methotrexate in juvenile idiopathic arthritis: correlation between gene expression and genotype**. *Pharmacogenet Genomics*. 2010; 20:665-76.
- 34. Namjou B, Kothari PH, Kelly JA, Glenn SB, Ojwang JO, Adler A, Alarcon-Riquelme ME, Gallant CJ, Boackle SA, Criswell LA, Kimberly RP, Brown E, Edberg J, Stevens AM, Jacob CO, Tsao BP, Gilkeson GS, Kamen DL, Merrill JT, Petri M, Goldman RR, Vila LM, Anaya JM, Niewold TB, Martin J, Pons-Estel BA, Sabio JM, Callejas JL, Vyse TJ, Bae SC, Perrino FW, Freedman BI, Scofield RH, Moser KL, Gaffney PM, James JA, Langefeld CD, Kaufman KM, Harley JB, Atkinson JP. Evaluation of the TREX1 gene in a large multi-ancestral lupus cohort. *Genes Immun*. 2011; 12:270-9.
- 35. Ojwang JO, Adrianto I, Gray-McGuire C, Nath SK, Sun C, Kaufman KM, Harley JB, Rayan GM. **Genome-wide** association scan of Dupuytren's disease. *J Hand Surg Am.* 2010; 35:2039-45.
- 36. Orozco G, Eyre S, Hinks A, Bowes J, Morgan AW, Wilson AG, Wordsworth P, Steer S, Hocking L, Thomson W, Worthington J, Barton A. **Study of the common genetic background for rheumatoid arthritis and systemic lupus erythematosus**. *Ann Rheum Dis*. 2011; 70:463-8.
- 37. Rasmussen A, Sevier S, Kelly JA, Glenn SB, Aberle T, Cooney CM, Grether A, James E, Ning J, Tesiram J, Morrisey J, Powe T, Drexel M, Daniel W, Namjou B, Ojwang JO, Nguyen KL, Cavett JW, Te JL, James JA, Scofield RH, Moser K, Gilkeson GS, Kamen DL, Carson CW, Quintero-del-Rio AI, del Carmen Ballesteros M, Punaro MG, Karp DR, Wallace DJ, Weisman M, Merrill JT, Rivera R, Petri MA, Albert DA, Espinoza LR, Utset TO, Shaver TS, Arthur E, Anaya JM, Bruner GR, Harley JB. **The lupus family registry and repository**. *Rheumatology (Oxford)*. 2011; 50:47-59.
- 38. Ross GS, Zelko F, Klein-Gitelman M, Levy DM, Muscal E, Schanberg LE, Anthony K, Brunner HI. A proposed framework to standardize the neurocognitive assessment of patients with pediatric systemic lupus erythematosus. *Arthritis Care Res (Hoboken)*. 2010; 62:1029-33.
- 39. Ruperto N, Giannini EH, Pistorio A, Brunner HI, Martini A, Lovell DJ. Is it time to move to active comparator trials in juvenile idiopathic arthritis?: a review of current study designs. *Arthritis Rheum*. 2010; 62:3131-9.
- 40. Ruperto N, Lovell DJ, Li T, Sztajnbok F, Goldenstein-Schainberg C, Scheinberg M, Penades IC, Fischbach M, Alcala JO, Hashkes PJ, Hom C, Jung L, Lepore L, Oliveira S, Wallace C, Alessio M, Quartier P, Cortis E, Eberhard A, Simonini G, Lemelle I, Chalom EC, Sigal LH, Block A, Covucci A, Nys M, Martini A, Giannini EH. Abatacept improves health-related quality of life, pain, sleep quality, and daily participation in subjects with juvenile idiopathic arthritis. *Arthritis Care Res (Hoboken)*. 2010; 62:1542-51.
- 41. Ruperto N, Pistorio A, Ravelli A, Rider LG, Pilkington C, Oliveira S, Wulffraat N, Espada G, Garay S, Cuttica R, Hofer M, Quartier P, Melo-Gomes J, Reed AM, Wierzbowska M, Feldman BM, Harjacek M, Huppertz HI,

- Nielsen S, Flato B, Lahdenne P, Michels H, Murray KJ, Punaro L, Rennebohm R, Russo R, Balogh Z, Rooney M, Pachman LM, Wallace C, Hashkes P, Lovell DJ, Giannini EH, Gare BA, Martini A. **The Paediatric**Rheumatology International Trials Organisation provisional criteria for the evaluation of response to therapy in juvenile dermatomyositis. *Arthritis Care Res (Hoboken)*. 2010; 62:1533-41.
- 42. Sagcal-Gironella AC, Fukuda T, Wiers K, Cox S, Nelson S, Dina B, Sherwin CM, Klein-Gitelman MS, Vinks AA, Brunner HI. **Pharmacokinetics and pharmacodynamics of mycophenolic acid and their relation to response to therapy of childhood-onset systemic lupus erythematosus**. *Semin Arthritis Rheum*. 2011; 40:307-13.
- 43. Sanchez E, Webb RD, Rasmussen A, Kelly JA, Riba L, Kaufman KM, Garcia-de la Torre I, Moctezuma JF, Maradiaga-Cecena MA, Cardiel-Rios MH, Acevedo E, Cucho-Venegas M, Garcia MA, Gamron S, Pons-Estel BA, Vasconcelos C, Martin J, Tusie-Luna T, Harley JB, Richardson B, Sawalha AH, Alarcon-Riquelme ME. Genetically determined Amerindian ancestry correlates with increased frequency of risk alleles for systemic lupus erythematosus. *Arthritis Rheum*. 2010; 62:3722-9.
- 44. Sestak AL, Furnrohr BG, Harley JB, Merrill JT, Namjou B. **The genetics of systemic lupus erythematosus** and implications for targeted therapy. *Ann Rheum Dis.* 2011; 70 Suppl 1:i37-43.
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Grants, Contracts, and Industry Agreements

Grant and Contract Awards

Annual Direct / Project Period Direct

BRUNNER, H		
Early Aggressive Therapy in Ju	venile Idiopathic Arthritis	
National Institutes of Health(Child	ren's Hosp & Reg Med Ct-Seattle)	
R01 AR 049762	07/01/10-06/30/11	\$12,859
Early Aggressive Therapy in Ju	venile Idiopathic Arthritis	
National Institutes of Health(Child	ren's Hosp & Reg Med Ct-Seattle)	
R01 AR 049762	07/01/10-06/30/11	\$4,372
Family Study of Pediatric Autol	mmunity	
Children's Hosp & Reg Med Ct-Se	eattle	
	08/01/09-07/31/11	\$2,200
Myositis Registry		
Centers for Disease Control and F	Prevention(The Myositis Association)	
H75 DP 001743	09/01/10-08/31/11	\$78,536
Towards Measures of Lupus Ne	phritis Activity & Damage in Children	
National Institutes of Health		
U01 AR 059509	08/08/10-05/31/13	\$135,002

	herapy in Juvenile Idio	-		
National Institutes of R01 AR 049762	Health(Children's Hosp	05/01/10-09/22/11		\$113,924
		00/01/10/00/22/11		Ψ110,021
GRIFFIN, T				
HLA-B27 Misfolding National Institutes of	g an the UPR in Spond	lyloarthritis		
R01 AR 046177	пеаш	09/01/06-06/30/11		\$273,473
HARLEY, J				
Genetic Linkage in	Lupus			
National Institutes of	Health			
R37 AI 024717		09/07/10-02/28/15		\$207,939
Genetic Linkage in	-			
National Institutes of	Health	00/46/40 00/24/44		¢452.000
R37 Al 024718 Genetic Linkage in	Lunus	09/16/10-08/31/11		\$153,900
National Institutes of				
R37 AI 024719		09/07/10-02/28/15		\$258,680
Illumina iScan Syst	tem for the OMRF Micr	oarray Research Facility		
National Institutes of	Health			
S10 RR 027190		06/10/10-06/09/11		\$413,350
	•	an-Americans with Systemic Lup	ous Erythematosus	
Department of Defer	ise	09/01/10-08/31/13		\$269,095
		00/01/10 00/01/10		Ψ200,000
HUGGINS, J				
_	-	ellowship Training Award		
American College of	Rheumatology Research	ch & Education Foundation		405.000
		07/01/10-06/30/11		\$25,000
LOVELL, D				
Multidisciplinary C	linical Research Cente	r		
National Institutes of	Health			
P60 AR 047784		08/18/08-07/31/13		\$847,406
Lovell, D	Administrativ		\$70,938	
Giannini, E	Methodology	Core	\$112,710	
Brunner, H	Project 1		\$179,060	
Lovell, D	Project 2		\$178593	
Grom, A	Project 3		\$163,409	
Seid, M	Project 4		\$142,696	
MORGAN DEWITT, E				
	S in Pediatric Pain, Rho	eumatology, and Rehabilitation F	Research	
National Institutes of				
U01 AR 057940		02/01/11-07/31/13		\$72,262

Enhancing PROMIS in Pediatric F	Pain, Rheumatology, and Rehabilitation Research	
National Institutes of Health		
U01 AR 057940	02/01/11-07/31/13	\$72,262
Enhancing PROMIS in Pediatric F	Pain, Rheumatology, and Rehabilitation Research	
National Institutes of Health		
U01 AR 057940	09/30/09-07/31/13	\$367,410
Juvenile Arthritis Improvement N	etwork for Clinical Excellence & Safety	
Arthritis Foundation		
	07/01/10-06/30/12	\$92,593

National Institutes of Health RC2 AR 058934	09/30/09-08/31/11		\$19,634
Cincinnati Rheumatic Dis	ease Core Center		
National Institutes of Health	1		
P30 AR 047363	09/01/06-06/30/11		\$388,400
Thompson, S	Administrative Core	\$45,464	
Thompson, S	Core 1 - Tissue	\$30,791	
Degen, J	Core 2 - Animal Models of Arthritis/Inflammatory Disease	\$66,736	
Thornton, S	Core 3 - Phenotyping	\$85,332	
Wagner, M	Core 4 - Informatics	\$57,743	
UC subcontract	Pilot & Feasibility	\$78,500	
Strait. R	Pilot & Feasbility	\$50,000	
-	pathic Arthritis and Subtypes		
Genetics of Juvenile Idiop National Institutes of Health R01 AR 057106			\$47,093
National Institutes of Health	n(Wake Forest University)	Current Year Direct	\$47,093 \$4,090,159
National Institutes of Health R01 AR 057106	n(Wake Forest University)		
National Institutes of Health R01 AR 057106 ndustry Contracts	n(Wake Forest University)		
National Institutes of Health R01 AR 057106 ndustry Contracts	n(Wake Forest University)		
National Institutes of Health R01 AR 057106 ndustry Contracts BRUNNER	n(Wake Forest University)		\$4,090,159
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc	n(Wake Forest University)		\$4,090,159 \$47,356
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc.	n(Wake Forest University)		\$4,090,159 \$47,356 \$30,389
National Institutes of Health R01 AR 057106 Industry Contracts RUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc.	n(Wake Forest University)		\$4,090,159 \$47,356 \$30,389
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc.	n(Wake Forest University)		\$4,090,159 \$47,356 \$30,389 \$44,617
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc. OVELL Pfizer, Inc Roche Laboratories, Inc. Novartis Pharmaceuticals	n(Wake Forest University)		\$4,090,159 \$47,356 \$30,389 \$44,617 \$127,777 \$230,781 \$181,363
National Institutes of Health R01 AR 057106 ndustry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc. OVELL Pfizer, Inc Roche Laboratories, Inc.	n(Wake Forest University)	Current Year Direct	\$4,090,159 \$47,356 \$30,389 \$44,617 \$127,777 \$230,781
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc. LOVELL Pfizer, Inc Roche Laboratories, Inc. Novartis Pharmaceuticals	n(Wake Forest University)		\$4,090,159 \$47,356 \$30,389 \$44,617 \$127,777 \$230,781 \$181,363
National Institutes of Health R01 AR 057106 Industry Contracts BRUNNER Pfizer, Inc Abbott Laboratories Centocor, Inc. LOVELL Pfizer, Inc Roche Laboratories, Inc. Novartis Pharmaceuticals	n(Wake Forest University)	Current Year Direct	\$4,090,159 \$47,356 \$30,389 \$44,617 \$127,777 \$230,781 \$181,363 \$7,333