





Annual Report 2005





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#### MISSION

Together, we proudly serve our regional community through the provision of expert laboratory and clinical services, education and research.

#### VISION

We strive to be national leaders in advanced diagnostic services, employee success, student achievement and knowledge discovery.

#### **VALUES**

In our pursuit of excellence, we value people by practicing mutual respect, professionalism, teamwork, integrity, trust and accountability.

# Message from the Head

Photo of Ian

n 2005, growth was the theme that dominated our departmental activities. We expanded our teaching, research and clinical initiatives and, while doing so, we maintained the quality of our programs at a very high level.

In education, our graduate program continued to expand. Five-year growth in student enrollment exceeds 30% and we project further short-term growth of at least 10-20%. The ongoing growth will be spurred by an increase in the number of our graduate faculty, the investments the province of Ontario has committed to graduate education and Queen's strategic initiative to develop graduate programs. We have revised our graduate curriculum and increased its flexibility in order to better meet student needs and to optimize the duration of their programs.

The resurgence of our postgraduate program continued. In 2005, resident enrollment increased 30% over the previous year and there has been a six and a half-fold increase in the number of residents in the last three years. Total resident enrollment currently exceeds 85% of program capacity. The postgraduate research program has also been successfully reinstituted.

In 2005, we recruited two research faculty to new positions and, in parallel with the growth of our graduate program, the number of postdoctoral fellows in the department reached a historic high. Total departmental research funding exceeded \$6M for the first time in the 2004-2005 fiscal year and the quality of our scholarship continues to be excellent. This is manifested not only by the high productivity of our faculty but also by the esteem in which they are held in the academic community. Most notably, Susan Cole and Roger Deeley were honoured by the National Cancer Institute of Canada/Canadian Cancer Society as co-recipients of the 2005 Noble prize for outstanding achievement in cancer research.



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In the clinical domain, we continued to provide strong leadership in the regional delivery of medical laboratory services. Our faculty played an integral role in the establishment of a regional infection control program in southeastern Ontario and provide ongoing medical leadership for that initiative. We were instrumental in the successful implementation of a regional medical laboratory equipment plan which led to a major capital investment in both our Academic Health Sciences Centre and our regional partner health care sites.

At the provincial level in laboratory medicine, 2005 marked the implementation of the initial Laboratory Medicine Funding Framework Agreement (LMFFA) between the Laboratory Medicine Section of the Ontario Medical Association and the Ministry of Health-Long Term Care. Through this agreement, the MOH-LTC made a significant investment in the profession of laboratory medicine and, in so doing, harmonized the Ontario laboratory physician funding schedule with those of western Canada. This has restored the competitive position of both the academic and community sections of laboratory medicine in Ontario and positioned us very well to compete for and retain the best professional talent. The LMFFA also addressed the issue of laboratory physician accountability, specifically with respect to physician workload, and required that a mechanism for workload measurement be established. Although the current LMFFA does not stipulate productivity targets, this is a direction in which the province may well be heading. If so, academic laboratory medicine will face the challenge of protecting and maintaining its mission of scholarship in a funding environment which is driven largely by the clinical service domain.

Iain D. Young,
Professor and Head,
Pathology and Molecular Medicine

## People

#### **FACULTY**

### PRIMARY APPOINTED EMERITUS PROFESSORS

Sergio Bencosme
William Corbett
Alan Giles
Nathan Kaufman
George Kipkie
Robert Kisilevsky
Howard Steele
David Robertson

#### Professor And Head

Iain D. Young

#### **Professors**

Susan Cole Roger Deeley Bruce Elliott David Lillicrap Samuel Ludwin Paul Manley Lois Mulligan Sandip SenGupta Dick Zoutman

#### Associate Professors

Sandy Boag
Christine Collier
Scott Davey
David Dexter
Peter Greer
David Hurlbut
Mohamed Khalifa
David LeBrun
John Rossiter
Lorne Seargeant
Lois Shepherd
Sherryl Taylor

#### Assistant Professors

Timothy Childs
Harriet Feilotter
Karen Harrison
Tadaaki Hiruki
Phillip Isotalo
Robert Liao
Christopher Nicol
Dilys Rapson
Michael Raymond
Caroline Rowlands
Xiaolong Yang

#### Cross-Appointed

#### **Professors**

John Matthews, *Medicine*Martin Petkovich, *Biochemistry*Paul Young, *Biology* 

#### Associate Professors

Gerald Evans, Medicine
David Lee, Medicine
Donald Maurice,
Pharmacology and
Toxicology
Christopher Mueller,
Biochemistry
Leda Raptis, Microbiology

and Immunology

Lewis Tomalt, Microbiology

and Immunology

#### Assistant Professors

Cynthia Forster-Gibson, Family Medicine Paula James, Medicine Jennifer MacKenzie, Pediatrics

#### ADJUNCT FACULTY

John Ancsin
Melanie Borden
Daniel Connor
Daria Haust
Christine Hough
Lloyd Kennedy
Norma Layno
David Piper
Michael Raymond
Waheed Sangrar
Ines Sumargo
Shui Pang Tam
Greg Twemlow
Virginia Walley
Jeannette Wilkins

#### Promotions

(Effective July 1, 2005)

#### Professor

Peter Greer Sandip SenGupta

#### Associate Professor

David Hurlbut

#### NEW APPOINTMENTS

Scott Davey Norma Layno Christopher Nicol

#### Administration and Support Services

Marg Croft, Postgraduate Program Assistant John DaCosta, Electron Microscopy Service Barb Dafoe, Administrative Assistant to Managers of Clinical Laboratories

Maria Dickson, Financial Administrative Assistant
Laura Emery, Administrative Secretary
Kevin Kell, Network Coordinator
Barb Latimer, Administrative Assistant
Norma Layno, Administrative Director
Barb Saunders, Graduate Program Assistant
Shakeel Virk, Pathology Coordinator, NCIC-Clinical Trials
Group National Tissue Bank

#### CLINICAL LABORATORY SERVICES MANAGEMENT

Norma Layno, Administrative Director
Dave More, Manager, Pathology Services
Joyce deVette-McPhail, Manager, Core Laboratory Services
Susan Pugh, Manager, Microbiology and Genetics Services
Cathie Trayner, Laboratory Information System Specialist
and Manager, Point of Care Testing and Quality

#### DEPARTMENTAL HISTORIAN AND ARCHIVIST

M. Daria Haust

# Committees and Appointments

#### **DEPARTMENTAL COMMITTEES**

Promotion,

REAPPOINTMENT, TENURE

AND CONTINUING

**APPOINTMENTS** 

Sandy Boag (Chair)

Bruce Elliott

Peter Greer

Mohamed Khalifa

David LeBrun

Lois Mulligan

Cynthia Pruss

(graduate student)

John Rossiter

Andrew Schell (resident)

Sherry Taylor

GRADUATE PROGRAM

Соммітте

Peter Greer (Chair)

Harriet Feilotter

Leda Raptis

Fiona Rawle (graduate student)

John Rossiter

Barbara Saunders

Ted Wright (graduate student)

Iain Young (ex officio)

Anatomic and General Pathology Residency

TRAINING COMMITTEE

Caroline Rowlands (Chair)

Sandy Boag

Tim Childs

Christine Collier

David Dexter

Brian Gannon (resident)

Robert Liao

Marosh Manduch (resident)

John Rossiter

Andrew Schell (resident)

Lorne Seargeant

Lois Shepherd

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Greg Twemlow

Iain Young (ex officio)

Нематоратногосу

RESIDENCY TRAINING

COMMITTEE

Dilys Rapson (Chair)

Hessah Al-Sulami (resident)

David Dexter

David LeBrun

David Lee

David Lillicrap

Kristine Roland (resident)

Danny Salloum

Iain Young (ex officio)

SEAMO FINANCE

Соммітте

Iain Young (Chair)

Sandy Boag

David Dexter

Harriet Feilotter

D:1 D

Dilys Rapson

Sandip SenGupta

Sandip Sendupa

Dick Zoutman

CANADIAN COLLEGE OF

MEDICAL GENETICISTS

STANDING FELLOWSHIP

Соммітте

Karen Harrison (Chair)

Harriet Feilotter

Mohamed Khalifa

Jennifer MacKenzie

Sherry Taylor

#### INTERNAL APPOINTMENTS AND REPRESENTATION ON UNIVERSITY COMMITTEES

Susan Cole

Re-elected as Faculty Senator for the Faculty of Health Sciences Re-elected to University Senate

Member, Board of Directors, PARTEQ Innovations, Inc.

Member, Senate Promotion Advisory Committee

Elected Senate Representative to the International Centre Council

Member, Senate Search Committee on Academic Development

Elected Member, Academic Colleague for Queen's,

Council of Ontario Universities

Member, Standing Committee on Government and Community

Relations, Council of Ontario Universities

Christine Collier

Faculty Coordinator, Teaching Improvement Project System (TIPS)

Member, Council on Employment Equity

Member, Human Rights Office Advisory Council Committee

Member, Instructional Development Network Committee

Roger Deeley

Director, Queen's University Cancer Research Institute

Director, Division of Cancer Biology and Genetics,

rector, Division of Cancer Biology and Genetics Queen's University Cancer Research Institute David Lillicrap

Member, Chancellor's Research Awards Adjudicating Committee

Samuel Ludwin

Associate Dean, Research, Faculty of Health Sciences

Vice-President, Research Development, Kingston General Hospital

Lois Mulligan

Director, CIHR Transdisciplinary Training Program

in Cancer Research

Sandip SenGupta

Member, University Promotions Committee

Sherry Taylor

Director, Phase I Undergraduate Medical Curriculum

Chair, Biohazard Safety Committee

Lewis Tomalty

Assistant Dean, Continuing Professional Development

Associate Dean, Medical Education

Dick Zoutman

Appointed Chair of the Division of Infectious Diseases

in the Department of Medicine

## **Education Programs**

#### UNDERGRADUATE PROGRAM

he Department of Pathology and Molecular Medicine holds a unique position in the Faculty of Health Sciences as it serves to bridge the interface between basic medical science

and clinical medicine. The Department fulfills this role in a range of undergraduate curricula.

#### COURSES

#### MEDICINE

Phase I Pathology General Pathology Pathology 425 Current Topics in Human Genetics (Coordinator: Sherry Taylor) (Coordinator: Lois Mulligan) Medical Science Rounds Pathology 430* Molecular Basis of Disease (Coordinator: Sherry Taylor) (Coordinator: Iain Young) Phase II Pathology Integrated, systems-based curriculum Pathology 499 Research Project in Pathology (Coordinator: David Lillicrap) LIFE SCIENCES Life Sciences 422 Cellular and Molecular Neuroscience Pathology 410 General Pathology for Non-Medical Sections taught by Faculty in Pathology Students (Coordinator: John Rossiter) and Molecular Medicine

#### **GRADUATE PROGRAM**, Peter Greer, Program Coordinator, Barb Saunders, Program Assistant

ur faculty supervised 38 graduate students in 2005; more than 35% of whom were pursuing their PhD.

The last appraisal of the Program was conducted by the Ontario Council on Graduate Studies in June 2004. At that time, Drs. Paul Lasko (McGill University) and Brian Clarke (McMaster University), OCGS consultants, assessed our MSc/PhD Program as "good quality." In keeping with their suggestions, the Research Seminar Series was registered as a graduate course (Path 830/930) in 2005, enabling graduate students to include it in their course requirements.

The Department participated in two CIHR-funded Training Programs: the Protein Function Discovery Training Program,

(Coordinator: Iain Young)

and the Transdisciplinary Training Program in Cancer. In the 2005 calendar year, these Programs supplied approximately \$64,000 in full or partial funding to six Pathology students.

The Robert Kisilevsky Fund for Research Education provides a bursary for first year PhD students in Pathology and Molecular Medicine. In September 2005, Rosanne Arulanandam, Taranjit Gujral, Doug Richardson and Andrew Slot each received a \$5,000 award.

The Pathology Studentship, offered to a highly competitive applicant to graduate studies in the Department, was awarded to Mohamad Qadura who accepted a position as a Master's student in Dr. David Lillicrap's laboratory.

and Molecular Medicine

#### **COURSES**

#### **PATHOLOGY**

Pathology 822	Experimental Cancer Diagnostics and Therapeutics	Pathology 827	Research Project in Pathology (Coordinator: Peter Greer)
	(Coordinator: Bruce Elliott)	Pathology 830/930	Research Seminar Course
Pathology 823*	Cancer Biology		(Coordinator: Peter Greer)
	(Coordinator: Bruce Elliott)	* alternate year cour	se last given in 2004
Pathology 824	General Pathology for Non-Medical Students (Coordinator: John Rossiter)	Rehabilitation Therapy	
Pathology 826*	The Molecular Basis of Disease (Coordinator: Jain Young)	Physical Therapy	Master's Program, Sections in several courses taught by faculty from Pathology

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^{*} alternate year course last given in 2004

#### **GRADUATE STUDENTS**

#### PhD Students (Supervisor)

Kristin Albert (Leda Raptis) Stat3 regulation by the caveolae protein, Cav-1, in breast cancer Lilia Antonova (Christopher Mueller) Stress and breast cancer: regulation of BRCA1 expression by hydrocortisone

Taranjit Gujral (Lois Mulligan) Molecular modeling of ret: mutations and mechanisms

Brandy Hyndman (David LeBrun) Acetylation and protein – protein interactions involving the protein E2A

Glenn MacLean (Martin Petkovich) Cloning of Cyp26B1 and its role during murine embryogenesis

Cynthia Pruss (David Lillicrap) The role of von Willebrand factor in arterial thrombosis

Andrew Slot (Susan Cole) Multidrug resistance protein

Patrick Smith (Peter Greer/Tassos Anastassiades) Metabolic and gene expression modulation in cultured articular chondrocytes by N-acylation of GlcN

Peter Truesdell (Peter Greer) Understanding the diverse roles of the fps tyrosine kinase

Ted Wright (Bruce Elliott) Inhibiting hepatocyte growth factor interaction with its receptor met in breast carcinoma cells

#### MSc Students (Supervisor)

Aikaterini Anagnostopoulou (Leda Raptis) The effect of Stat3 in normal and tumor cells and in the differentiation of mammary epithelial cells

Paul Bandler (Roger Deeley) Investigation of elements involved in plasma membrane targeting of human multidrug resistance proteins, MRP1 and MRP2

Audrey Boros (Scott Davey) Characterizing the cellular defects associated with attenuated hRad9 function Crystal Chan (Susan Cole)
Pharmacogenetic analyses of MRP-related organic anion transporters

Sara Flemming (Xiaolong Yang) Isolation and characterization of both the LATS1 and LATS2 promoter regions

Cheryl Foster (David LeBrun)
Identification of prognostic biomarkers
and elucidation of molecular mechanisms
involved in low to high grade transformation in B-cell lymphomas

Morgan Jarvis (Scott Davey)
Investigation of Rad9 C-terminal tail
function in the Rad9-Rad1-Hus1 checkpoint complex in fission yeast

Jonathan Krieger (Lois Mulligan) Development of novel inducible neural models to study the ret receptor tyrosine kinase and Hirschsprung disease

Andrea Lai (Lois Mulligan) The occurrence and functionality of ret isoform heterodimers

Joe Oliver (Bruce Elliott) Role of cell matrix adhesion in an in vitro model of human breast cancer

Shawna Organ (Protein Function Discovery Program) Proteomic analysis of the Met receptor tyrosine kinase and cell adhesion signaling cascades involved in breast cancer metastasis

Jeff Ovens (Don Maurice) The role of cyclic nucleotide phosphodiesterases in endothelial cell permeability

Michelle Sam (Bruce Elliott/Chris Mueller) The regulatory effects of stat3 signaling on autocrine HGF expression in breast cancer

Brian Simo (Lois Mulligan) Differential signaling between the wild type ret tyrosine kinase (TK) receptor and the multiple endocrine neoplasia (MEN) Type 2B ret TK receptor



Jayda Sutton (Don Maurice) PDE interactions with PKA and Epac relevance to cAMP inhibition of angiogenesis

Ashley Theis (Roger Deeley) Higher order structure of MRP1: application of cysteine-based techniques

Xuefei Tian (Lois Mulligan) Characterization of newly developed ret antibodies and application in ret-related tumours

Stephanie Vander Pol (David Lillicrap) Induction of immunological tolerance to factor VIII through tolerogenic antigen presentation

Stacy Visser (Xiaolong Yang) Interaction of LATS and PAK1 and its roles in regulating cell migration

#### MSc to PhD Transfer – Mini Masters Program (Supervisor)

Rozanne Arulanandam (Leda Raptis) Mechanism of Stat3 activation by cell-tocell adhesion

Leo Mok (Roger Deeley) Elucidation of ABCA1 ATPase activity and the function of regulatory domain 1 using the Spodoptera frugiperda 21 cell line

Doug Richardson (Lois Mulligan) Ret internalization and sub-cellular localization: effects on downstream signaling

#### **GRADUATING STUDENTS**

#### MSc Students (Supervisor)

Zoya Shapovalova (Peter Greer) Involvement of fps and fer tyrosine kinases in neuronal cytoskeletal remodeling

Sara Sharifpoor (Bruce Elliott) Characterization of the function and regulation of Stat3 in autocrine HGF expression

Taranjit Gujral (Lois Mulligan) Molecular modeling of ret: mutations and mechanisms

#### PhD Students (Supervisor)

Fiona Rawle (David Lillicrap) Improving hemophilia A treatment: increasing efficacy and decreasing immunogenicity

Adina Vultur (Leda Raptis) Role of chaperones upon transformation and differentiation

Kevin Weigl (Susan Cole) Biochemical investigations of MRP1: analysis of glycosylation and topology

## GRADUATE STUDENT PRESENTATIONS

A. Anagnostopoulou, R. Arulanandam, A. Vultur, R. Jove, J. Turkson and L. Raptis. Novel pathway of Rac1, Stat3 and p53 interaction. *The American Society for Cell Biology 45th Annual Meeting*. San Francisco, California, December 2005.

A. Anagnostopoulou, A. Vultur, R. Arulanandam, J. Turkson, R. Jove and L. Raptis. Differential effects of Stat3 inhibition in sparse vs confluent cells. *The American Society for Cell Biology 45th Annual Meeting.* San Francisco, California, December 2005.

**L.** Antonova and C.R. Mueller. The BRCA1 gene: A possible mediator of the effect of the stress hormone hydrocortisone on breast cancer development. *70th CHSL Symposium, Molecular Approaches to Controlling Cancer*, Cold Spring Harbor, New York, June 2005.

R. Arulanandam, A. Vultur, A. Anagnostopoulou, B. Elliott and L. Raptis. A novel mechanism of Stat3 activation. *American Association for Cancer Research 96th Annual Meeting*. Anaheim, California, April 2005.

R. Arulanandam, A. Vultur and L. Raptis. A novel mechanism of Stat3 activation. Signal transduction in cancer. 2005 Miami Nature Biotechnology Winter Symposium. Miami, Florida, February 2005.

**A. Boros** and S. Davey. Characterizing the cellular defects associated with attenuated HRAD9 function. 8th Annual Scientific Meeting for Health Sciences Research Trainees, Faculty of Health Sciences, Queen's University, Kingston, Ontario, May 2005.

L.M. Mulligan and **T.S. Gujral**. Ret receptor signalling: dissecting receptor function and dysfunction. *1st International Meeting on Pheochromocytomas*, Bethesda, Maryland. October 2005. *Selected for Oral Presentation* 

T.S. Gujral, J.R. Krieger, Z. Jia and L.M. Mulligan. Structural basis of ret kinase domain mutations in Hirschsprung disease. 8th Annual Meeting For Health Sciences Research Trainees. Kingston, Ontario, May 2005.

Selected for Oral Presentation

J.R. Krieger, T.S. Gujral and L.M. Mulligan. Generating a novel neural inducible model for ret in Hirschsprung disease. *Queen's University Cancer Research Institute Retreat*. Gananoque, Ontario, January 2006.

S.L. Organ, T.S. Gujral, J.R. Krieger, B.E. Elliott and L.M. Mulligan. A proteomics approach to studying receptor tyrosine kinases. *Cancer Research Across the Spectrum: National Meeting for Trainees*. Mont Tremblant, Quebec, May 2005.

S.L. Organ, T.S. Gujral, J.R. Krieger, B.E. Elliott and L.M. Mulligan. A proteomics approach to studying receptor tyrosine kinases. *Eighth Annual Meeting for Health Sciences Research Trainees*, Kingston, Ontario, May 2005.

J.D. Ovens, P. Malenfant and J.J.A. Holden. VGLUT2 as a candidate locus for autism spectrum disorders. *Queen's Health Sciences Research Trainees*Meeting, Kingston, Ontario, May, 2005.

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- M. Shibata, **F. Rawle**, A. Labelle, C. Andrews, C. Hough, A. Yoshioka and D. Lillicrap. Characterization of recombinant canine Factor VIII and quantitative determination of Factor VIII in canine plasma. *XXth Congress of the ISTH*, Sydney, Australia, August 2005.
- S.D. Andrew, **D.S. Richardson**, J.L.V. Shaw and L.M. Mulligan. Ret degradation: the role of Src and Cbl. *Australian Society for Medical Research (New South Wales) Scientific Meeting* 2005. Sydney, Australia. *Won the post-doctoral research prize for best poster presentation.*
- **D.S. Richardson**, A.Z. Lai, S. Nita, J.H. Horton and L.M. Mulligan. Characterization of ret receptor tyrosine kinase ligand-induced internalization. *Queen's Health Sciences Research Trainees Meeting*, Kingston, Ontario, May 2005. *Selected for Oral Presentation*
- **D.S. Richardson**, A.Z. Lai, S. Nita, J.H. Horton and L.M. Mulligan. Characterization of ret receptor tyrosine kinase ligand-induced internalization using a multi-disciplinary approach. *American Association of Cancer Research Annual Meeting* 2005. Anaheim, California.
- A.J. Slot, I.J. Létourneau, R.G. Deeley and S.P.C. Cole. Mutation of proline 1158 to alanine in cytoplasmic loop 7 significantly alters the substrate specificity of human MRP2(ABCC2). 2nd Annual North American ABC Genetic Workshop, NCI-Frederick, Frederick, MD, September 2005.
- **X.F. Tian**, S.D. Andrew, P. Isotalo and L.M. Mulligan. Characterization and application of newly developed ret antibodies. *Cancer Research Across the Spectrum: National Meeting for Trainees*. Mont Tremblant, Quebec, May 2005.

- H. Visram, A. Ableson, P. Truesdell, E. Eisenhauer, R.L. George, P.A. Greer and W. Sangrar. Cell signaling and cancer: a systems biology approach. 8th Annual Scientific Meeting for Health Research Trainees, Kingston, Ontario, May 2005.
- **T. Wright**, J. Tsai, A. Jia and B.E. Elliott. Inhibition by copper (II) binding of hepatocyte growth factor interaction with its receptor Met in breast carcinoma cells. *USAMRMC Era of Hope 2005 Conference*. Philadelphia, PA, June 2005.
- E.J. Wojcik, S. Sharifpoor, P. Cheung, T. Wright, R. Watering, E.A. Tremblay, C.R. Mueller and B.E. Elliott. A novel role of c-Src and Stat3 in autocrine HgF expression in invasive breast cancer. Presented at the *Annual Meeting of the American Association of Cancer Research*, Anaheim, CA, April 2005.

### GRADUATE STUDENT PUBLICATIONS

- A. Anagnostopoulou, A. Vultur, R. Arulanandam, J. Cao, J. Turkson, R. Jove, J.S. Kim, M. Glenn, A. Hamilton and L. Raptis. Role of Stat3 in normal and SV40 transformed cells (Review). *Research Trends*. Submitted July 2005.
- A. Anagnostopoulou, A. Vultur, R. Arulanandam, J. Cao, J. Turkson, R. Jove, J.S. Kim, M. Glenn, A. Hamilton.and L. Raptis. Differential effects of Stat3 inhibition in sparse vs. confluent normal and breast cancer cells. *Cancer Letters*. (in press)
- **A. Vultur, R. Arulanandam**, J. Turkson, G. Niu, R. Jove and L. Raptis. Stat3 is required for full neoplastic transformation by the Simian Virus 40 Large Tumor antigen. *Mol. Biol. Cell.* 16, 8: 3832-3846, 2005.

- J. Cao, R. Arulanandam, A. Vultur, T. Preston, E. Jaronczyk, K. Tomai, K. Zandi and L. Raptis. Adenovirus-5 E1A suppresses differentiation of 3T3 L1 preadipocytes at lower levels than required for induction of apoptosis. *Mol. Carcinog.* 43, 1: 38-50, 2005.
- **R.** Arulanandam, A. Vultur and L. Raptis. Transfection techniques affecting Stat3 activity levels. *Anal. Biochem.* 338, 1: 83-89, 2005.
- O. Loudig, **G. MacLean**, N. Dore, L. Luu and M. Petkovich. Transcriptional co-operativity between distant Retinoic acid response elements in regulation of Cyp26A1 inducibility. *Biochem. J.* 392, 241-248, 2005.
- B. Gu, **C.M. Pruss**, A.T.Gates and S.S. Khandekar. The RNA-unwinding activity of Hepatitis C virus non-structural protein 3 (NS3) is positively modulated by its protease domain. *Protein and Peptide Letters* 12(4): 315-321, 2005.
- D.H. Freed, R.H. Cunnington, A.L. Dangerfield, **J.A. Sutton** and I.M.C. Dixon. Emerging evidence for the role of Cardiotrophin-1 in cardiac repair in the infarcted heart. *Cardiovascular Research* 65(4): 782-92, 2005.
- L. Payen, M. Gao, C. Westlake, A. Theis, S.P.C. Cole and R.G. Deeley. Functional interactions between nucleotide binding domains and LTC4 binding sites of multidrug resistance protein 1 (ABCC1). *Mol. Pharmacol.* 67: 1944-1953, 2005.

#### POSTGRADUATE PROGRAM,

Caroline Rowlands, *Program Coordinator and Director, Anatomic and General Pathology Programs*, Lois Shepherd, *Director, Hematopathology Program* 

nder the umbrella of Laboratory Medicine the department offers training in Anatomical Pathology, General Pathology and Hematopathology for careers in both the academic and community sectors. Programs are approaching capacity with 13 of the 15 available residency positions now being filled. The postgraduate programs underwent the Royal College Accreditation process in 2005 and the Anatomical Pathology Program achieved full accreditation. Constructive recommendations for optimizing administrative processes within the Hematopathology Program were made by the accreditation team. These recommendations were implemented and have led to process improvements. As there were no residents in the General Pathology program in 2005, the program was deemed inactive and consequently did not undergo review. We anticipate recruiting at least two trainees into this program in 2006.

Our success in resident recruitment in the last several years has led to the re-establishment of our postgraduate research program. In 2005, residents made seven presentations of their scholarly work at international, national and provincial meetings.

In order to accommodate the recent growth in our postgraduate programs, the department has made significant



#### RESIDENTS

Anatomic Pathology (AP), Hematopathology (H) (*l-r*) Reza Behjati(AP), PGY3, Jerry Chen(AP), PGY2, Christopher Davidson(AP), PGY2, Maria Abi Daoud(AP), PGY1, Patricia Farmer(AP), PGY6, Brian Gannon(AP), PGY4, Marosh Manduch(AP), PGY3, Navid Nasseri(AP), PGY3, Andrew Schell(AP), PGY3, Nafisa Ursani(AP), PGY1, Hessah Al-Sulami(H), PGY4, Kristin Roland(H), PGY5, Danny Salloum(H), PGY1

improvements to educational space, workstations and teaching equipment. The now excellent facilities also allow us to accommodate an increasing demand for electives by students and external residents.

#### **POSTGRADUATE RESIDENTS**

#### RESIDENT RESEARCH (Supervisor)

Patricia Farmer (Harriet Feilotter and David LeBrun) Development of combined clinical and biomarker prognostic factors in follicular lymphoma.

Patricia Farmer (David LeBrun) Tissue microarrays as a quality assurance tool in lymphoma immunodiagnosis.

Brian Gannon (*Tim Childs*) *Tissue* microarray analysis of protein expression patterns in leiomyomas and leiomyosarcomas.

Brian Gannon (*Tim Childs*) Analysis of the epithelial and mesenchymal components of uterine carcinosarcomas.

Kristine Roland (Dilys Rapson, David Lillicrap and Paula James) Genetic testing of Type II von Willebrand disease. Kristine Roland (Harriet Feilotter and David LeBrun) Utilization of RNA from formalin-fixed paraffin embedded tissue for gene microarray studies.

Andrew Schell (Sherry Taylor and Iain Young) Mutational analysis of the BRAF signaling pathway and mutation analysis in hormone refractory prostate carcinoma.

#### RESIDENT PRESENTATIONS

P. Farmer, S. Virk, B.F. Burns and D.P. LeBrun. Use of a tissue microarray to assess the accuracy of lymphoma classification by needle biopsy. 47th *American Society of Hematology Annual Meeting and Exposition*, Atlanta, GA, December 2005.

**B. Gannon**, A. Sharkawy, G. Swain, P. Isotalo and D. Dexter. Gastric toxoplasmosis in aplastic anemia: A case report and review of the literature. *Canadian Association of Pathologists Annual Meeting*, Victoria, BC, July 2005.

**B. Gannon**, D. Dexter, D. Petsikas and P. Isotalo. Mediastinal thymolipoma: A rare occurence with striated myoid cells. *Canadian Association of Pathologists Annual Meeting*, Victoria, BC, July 2005

M. Manduch, D. Dexter, D. Jalink, J. Louw, S. Vanner and D. Hurlbut. Osteoclast-like giant cell tumour of pancreas with osteo-chondroid differentiation: case report and literature review. *Canadian Association of Pathologists Annual Meeting*, Victoria, BC, July 2005.

K. Roland, N. Gallo, D. LeBrun and H. Feilotter. Comparison of RNA extracted from matched formalin-fixed paraffinembedded tissue and frozen tissue: use in microarray analysis. *Canadian Association of Pathologists Annual Meeting*, Victoria BC, July 2005.

A. Schell, J. Veinot, P. Isotalo, I. Young and C. Rowlands. Hypocalcemic cardiomyopathy secondary to pseudo-hypoparathyroidism Type IA. *Canadian Association of Pathologists Annual Meeting*, Victoria, BC, July 2005.

#### RESIDENT PUBLICATIONS

N. Poonai, S. van Diepen, A. Bharatha, **M. Manduch**, T. Deklaj and S.M. Tarlo. Barriers to diagnosis of occupational asthma in Ontario. *Can. J. Public Health*, 96(3): 230-3, 2005.

#### POSTDOCTORAL TRAINING PROGRAM IN CLINICAL AND LABORATORY

GENETICS, Karen Harrison, Cytogenetics Coordinator, Sherry Taylor, Molecular Genetics Coordinator

Kingston General Hospital and Queen's University are together accredited by the Canadian College of Medical Geneticists to provide postdoctoral training programs in clinical and laboratory genetics. The clinical genetics program provides training to physicians in the diagnosis and management of genetic conditions. The laboratory genetics

program provides in-depth training in the technology and application of diagnostic cytogenetics and molecular genetics to clinical service.

TRAINEE IN LABORATORY GENETICS Shulin Zhang

#### OUTREACH EDUCATION PROGRAMS, Fiona Rawle, Coordinator of Enrichment Courses

Currently, the Department of Pathology and Molecular Medicine, in cooperation with the Office of Enrichment Studies, is the only department at Queen's University that is actively involved with the organization and teaching of enrichment courses targeting three different grade levels: Grades 5-6, Grades 7-8, and Grades 11-12. These courses have been developed solely by graduate students in the Department of Pathology and Molecular Medicine and each year they are coordinated by taught by current graduate students. The programs challenge the students to higher levels of learning, expose the students to the university setting, create an opportunity for the students to meet new peers, and expose the students to new subject topics and content. All courses are advanced instructional programs that engage the students by using diverse teaching methods and hands-on activities.

The ABC program (Grades 5-6; two day duration) involves fun and interactive classes that are a direct extension of the elementary science curriculum. Also known as "That is so gross! Human Anatomy and Forensic Science", this course uses plastinated specimens and detailed computerized "body tours" to teach the students about biological systems, and how disease can occur. Students are also exposed to basic forensic techniques, such as fingerprint analysis and crime scene searching. Through interactive exercises, the students gain an understanding of how the organs, muscles, nervous system, circulatory system and skeletal system of the body work together to perform the functions needed for life. The students then use this information to solve their own forensic cases.

The SEEDS (Seven-Eight Enrichment Day Studies) program (Grades 7-8; three day duration) focuses predominantly on

disease processes and forensic pathology. The students are introduced to anatomy, genetics, and infectious disease and are also introduced to the techniques of fingerprint analysis, DNA analysis, blood detection, and detailed crime scene searching. The students then apply these techniques to solving forensic cases, and use the knowledge they've learned to engage in "CSI: FACT OR FICTION" competition wherein the students determine if what happens on the prime-time TV show is indeed possible.

During the e=mc2 program (Grades 11-12; five day duration), also known as "Hands-On Forensic Pathology", students are introduced to the multifaceted world of pathology. Students are each assigned a forensic case (to determine cause of death) and a diagnostic case (to determine the disease present) at the beginning of the program, and use the information they learn during the week to solve their cases by the week's end. Students are exposed to elements of forensic pathology, gross anatomy, human genetics, cancer pathology, viruses, gene therapy, infectious diseases, epidemiology, immunology, and ethics. Students will also conduct experiments which are commonly performed in forensic laboratories, such as isolating and analyzing DNA, blood detection, fingerprint analysis, and chromatography. Students also learn the detailed science behind the prime time CSI show and use this knowledge to participate in "CSI-FACT OR FICTION" competitions.

#### **GRADUATE STUDENT INSTRUCTORS**

Taranjit Gujral, Jon Krieger, Andrea Lai, Cyndi Pruss, Fiona Rawle, Doug Richardson, Ashley Theis, Stephanie Vander Pol, and Ted Wright

## Research

he principal themes of the research programs in the Department include drug and radiation resistance in cancer, cancer biology and genetics, molecular hemostasis, amyloidogenesis and cholesterol metabolism. An emerging focus is translational oncologic research. Recent departmental initiatives in supporting the development of translational research programs have included the establishment of laser capture microdissection and tissue microarray facilities; equipment modernization in the gene microarray facility; and, the establishment of a frozen tissue tumour bank through a partnership with the Ontario Cancer Research Network.

Departmental research funding continued to grow progressively and total funding reached almost \$6.3M in 2004/2005. Over the course of the last five years, our total funding has increased 80% and funding received from the major national agencies has gown by more than 60%.

The quality and significance of the research conducted by our faculty is very high. In 2005, Susan Cole and Roger Deeley were co-recipients of the Robert L. Noble Prize of the Canadian Cancer Society/National Cancer Institute of Canada, an annual award given in recognition of outstanding achievements in cancer research.



Total Research Funding (2005-2005)	\$ 6,285,846		
Total Number of Awards	77		
Canadian Institutes of Health Research/			
Medical Research Council of Canada	\$ 2,192,329		
National Cancer Institute of Canada	\$ 434,498		
HEART AND STROKE FOUNDATION			
of Ontario/Canada	\$ 318,864		
Industry	\$ 1,807,450		
Other	\$ 1,532,705		

#### POSTDOCTORAL RESEARCH,

Harriet Feilotter, *Director*, *Queen's Microarray Facility*, David LeBrun, *Director*, *Experimental Pathology Unit* 

#### POSTDOCTORAL RESEARCH FELLOWS (Supervisor)

Gwenaelle Conseil (Susan Cole)	Lei Qin (Roger Deeley)
Yan Gao (Peter Greer)	Alice Rothnie (Susan Cole)
Roma Gurusankar (Scott Davey)	Yinfei Tan (Peter Greer)
Tozammel Hogue (Susan Cole)	Bin Wang (Susan Cole)
Akio Nakajima (Susan Cole)	Peng Wu (Susan Cole)
Maha Othman ( <i>David Lillicrap</i> )	Jessica Wyles (Susan Cole)

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#### RESEARCH FUNDING

Researcher	Sponsor	TITLE
Tim Childs	Queen's University	Tissue microarray analysis of immunohisto- chemical profiles in leiomyosarcomas and leiomyomas
Tim Childs	Canadian Institutes of Health Research	Evaluation of natural killer cell contributions to the known elevated risk for pre-eclampsia in diabetic pregnancy
Susan Cole and David Lillicrap	Canadian Institutes of Health Research	Training grant in proteomics
Susan Cole	Canadian Institutes of Health Research	Protein function discovery resource infrastructure
Susan Cole	Canadian Institutes of Health Research	Novel aspects of nitroglycerin action: nitrate transport, apoptosis and nitrosative stress
Susan Cole	National Cancer Institute of Canada	Investigations of DNA topoisomerase II and drug resistance
Susan Cole and Roger Deeley	Canadian Institutes of Health Research	Investigations of MRP-mediated drug resistance and xenobiotic transport
Christine Collier	Queen's University	Elevated homocysteine in dementia: a randomized, double blind, placebo controlled, pilot treatment trial with B-vitamins
Christine Collier	Industry	Estimating functional testosterone: the importance of biological variation
Scott Davey and Harriet Feilotter	National Cancer Institute of Canada Canadian Breast Cancer Foundation, CBCF, Ontario Chapter	DNA damage response in fission yeast Assessment of BRCA1 and BRCA2 functional status through genomic and radiobiological analyses
Scott Davey	Canadian Institutes of Health Research	Cell cycle checkpoint control
Roger Deeley	Heart and Stroke Foundation of Ontario	Functional studies of the ATP binding cassette transporter, ABCA 1
Roger Deeley and Susan Cole	Canadian Institutes of Health Research	MRP mediated multidrug resistance
Bruce Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer, David LeBrun, and David Lillicrap	Canadian Institutes of Health Research	Multi-user maintenance and equipment for flow cytometry, confocal microscope, microinjection and intraviral microscopy facility
Person Ellion		
Bruce Elliott	Canadian Breast Cancer Research Alliance	adhesion signalling in invasive cancer.  A possible therapeutic target
Bruce Elliott  Bruce Elliott	Canadian Breast Cancer Research Alliance Canadian Institutes of Health Research	e Novel interaction between Met and cell adhesion signalling in invasive cancer.
Bruce Elliott	Canadian Institutes of Health Research  Canadian Institutes of Health Research	adhesion signalling in invasive cancer.  A possible therapeutic target  Targeting a novel activating role of Stat 3 in  HGF expression in breast cancer; a possible
Bruce Elliott  Bruce E. Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer,	Canadian Institutes of Health Research  Canadian Institutes of Health Research	adhesion signalling in invasive cancer. A possible therapeutic target Targeting a novel activating role of Stat 3 in HGF expression in breast cancer; a possible therapeutic target Maintenance grant for multi-user equipment:
Bruce Elliott  Bruce E. Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer, David LeBrun and David Lillicrap	Canadian Institutes of Health Research  Canadian Institutes of Health Research	adhesion signalling in invasive cancer. A possible therapeutic target Targeting a novel activating role of Stat 3 in HGF expression in breast cancer; a possible therapeutic target Maintenance grant for multi-user equipment: flow cytometer and confocal microscope
Bruce Elliott  Bruce E. Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer, David LeBrun and David Lillicrap Harriet Feilotter	Canadian Institutes of Health Research  Canadian Institutes of Health Research  Ontario Cancer Research Network	adhesion signalling in invasive cancer. A possible therapeutic target Targeting a novel activating role of Stat 3 in HGF expression in breast cancer; a possible therapeutic target Maintenance grant for multi-user equipment: flow cytometer and confocal microscope Ontario cancer biomarker network
Bruce Elliott  Bruce E. Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer, David LeBrun and David Lillicrap Harriet Feilotter Harriet Feilotter, Dilys Rapson, Lois Shepherd, Karen Harrison	Canadian Institutes of Health Research  Canadian Institutes of Health Research  Ontario Cancer Research Network  Ontario Association of Medical	adhesion signalling in invasive cancer. A possible therapeutic target Targeting a novel activating role of Stat 3 in HGF expression in breast cancer; a possible therapeutic target Maintenance grant for multi-user equipment: flow cytometer and confocal microscope  Ontario cancer biomarker network Investigation of non-random gene mutation associations in chronic myeloid
Bruce E. Elliott, Susan Cole, Scott Davey, Roger Deeley, Peter Greer, David LeBrun and David Lillicrap Harriet Feilotter Harriet Feilotter, Dilys Rapson, Lois Shepherd, Karen Harrison and Kristine Roland Harriet Feilotter	Canadian Institutes of Health Research  Canadian Institutes of Health Research  Ontario Cancer Research Network  Ontario Association of Medical  Laboratories	adhesion signalling in invasive cancer.  A possible therapeutic target  Targeting a novel activating role of Stat 3 in HGF expression in breast cancer; a possible therapeutic target  Maintenance grant for multi-user equipment: flow cytometer and confocal microscope  Ontario cancer biomarker network Investigation of non-random gene mutation associations in chronic myeloid disorders. Investigation of non-random gene mutation

Researcher	Sponsor	TITLE
Harriet Feilotter	Queen's University	Assessment of BRCA1 and BRCA2 functional
and Scott Davey		status through genomic and radiobiological
		analyses
Peter Greer, Susan Cole, Scott Davey, Roger Deeley, Bruce Elliott, David LeBrun, and Lois Mulligan		Multi-user multi-modal imaging system
Peter Greer	National Cancer Institute of Canada	A genetic and molecular analysis of the fps/fes proto-oncogene
Peter Greer	Canadian Institutes of Health Research	A molecular and genetic analysis of the murine fer proto-oncogene
Peter Greer	Canadian Institutes of Health Research	Genetic and cell biological analysis of calpain
Karen Harrison, Phillip Isotalo	Queen's University	Molecular cytogenetic screening of breast cancers
and Sandip SenGupta		for HER-2/neu gene amplification: comparison of fluorescence and chromogenic in situ hybridization
Karen Harrison, Harriet Feilotter, and John Rossiter	Queen's University	Molecular investigation of oligodendrogliomas using CGH microarray and molecular cytogenetic technology
Karen Harrison, Phillip Isotalo and Sandip SenGupta	Queen's University	Molecular cytogenetic screening of breast cancers for HER-2/neu gene amplification using chromogenic in situ hybridization
David Hurlbut	PSI Foundation	Sentinel lymph node mapping in colon cancer
David Hurlbut	Queen's University	Sentinel lymph node mapping in colon cancer:
		a feasibility study
Phillip Isotalo	Queen's University	Treatment variation and treatment effectiveness in differentiated thyroid cancer
Phillip Isotalo	Ontario Cancer Research Network	Tissue bank contract
Phillip Isotalo and Bruce Elliott	Kingston General Hospital	Gene expression profiles associated with met and Stat3 signaling networks in invasive human
Phillip Isotalo and Bruce Elliott	Oueen's University	breast cancer Ezrin and met expression profile in invasive
Timip isotate and prace Emote	Queens oniversity	human breast cancer
Robert Kisilevsky	Heart and Stroke Foundation of Ontario	Acute phase SAA and cholesterol metabolism during inflammation
Robert Kisilevsky	Canadian Institutes of Health Research	SAA2.1 peptide therapeutics: two lead drug development candidates for the prevention and/or regression of atherosclerosis
Robert Kisilevsky	Canadian Institutes of Health Research	Amyloidogenesis: an analysis of the causative factors in an experimental murine model
David LeBrun	National Cancer Institute of Canada	The role of acetylation in regulating the leukemogenic protein E2A-PBX1
David LeBrun	Canadian Institutes of Health Research	Biophysical characterization of the E2A-CBP protein-protein interaction in acute lymphoblastic leukemia
David LeBrun	Industry	Companion studies in a clinical trial of bevacizumab in advanced colorectal cancer
David Lillicrap	Heart and Stroke Foundation of Ontario	Influence of genotype and environment on endothelial cell expression of von Willebrand factor

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Researcher	Sponsor	TITLE
David Lillicrap	Canadian Institutes of Health Research	Gene therapy for hemophilia A: studies of efficacy and the host immune response
David Lillicrap	Canadian Institutes of Health Research	Pathophysiologic mechanisms of von Willebrand factor biosynthesis
David Lillicrap	National Centers of Excellence	Genetically modified, autologous stem cell populations for hemophilia A
David Lillicrap	Heart and Stroke Foundation of Ontario	Collaborative studies of hemostasis, fibrinolysis and vascular cell growth and function
David Lillicrap	Canadian Hemophilia Society	Impact of von Willebrand disease in the primary care setting
David Lillicrap	Heart and Stroke Foundation of Ontario	The role of von Willebrand factor in arterial thrombosis
David Lillicrap	Canadian Institutes for Health Research	Evaluation of novel therapies in a unique canine model of hemophilia A
David Lillicrap	National Centers of Excellence, Stem Cell Network	Stem cell-based gene therapy for hemophilia A
David Lillicrap	US National Institutes of Health	The molecular genetic pathology of Type 1 von Willebrand disease
David Lillicrap	Association of Hemophilia Clinic Directors of Canada	National hemophilia mutation testing laboratory
Lois Mulligan Susan Cole, Roger Deeley, Bruce Elliott, Peter Greer, D. LeBrun and Scott Davey	Canadian Institutes of Health Research	Transdisciplinary training program in cancer research
Lois Mulligan	Canadian Institutes of Health Research	Developing and characterizing models for RET receptor function
Lois Mulligan	SWEP	Molecular biology research assistant
Lois Mulligan	SWEP	Research assistant: Pathology and Molecular Medicine
Lois Mulligan, Susan Cole, Roger Deeley and Peter Greer	Canadian Institutes of Health Research	Transdisciplinary training program in cancer research
Christopher Nicol	Kingston General Hospital and Dering Cancer Research Grant	Mechanisms of peroxisome proliferator-activated receptors influence on breast cancer
Christopher Nicol	Cancer Care Ontario	PPARs and cancer
Christopher Nicol	Queen's University	Biochemical and molecular mechanisms of peroxisome proliferator-activated receptors (PPARs) influence on carcinogenesis
Christopher Nicol	Canada Foundation for Innovation	Biochemical and molecular mechanisms of peroxisome proliferator-activated receptors (PPARs) influence on carcinogenesis
Caroline Rowlands	Queen's University	Department of Pathology and Molecular Medicine
Dilys Rapson	Industry	Waveform analysis of the aPTT in critically ill hospitalized patients
Dilys Rapson	Canadian Intensive Care Foundation	The relationship between insulin resistance, coagulation, inflammation, and clinical outcomes
Dilys Rapson and David Lillicrap	Queen's University	Prospective investigation of non-VWD bleeders
Lorne Seargeant	Queen's University	Glycosylation disorders

Researcher	Sponsor	TITLE
Lorne Seargeant and	Queen's University	Disorders of glycosylation in patients with
David Lillicrap	,	unexplained disorders of hemostasis
Lorne Seargeant	Garrod Association of Canada	Glycan analysis in patients with bleeding disorders
Lois Shepherd	CTRNet	CTRNet data interface project
Lois Shepherd	Industry	Multiple Phase I/II/II clinical trials of
		chemotherapeutic agents
Lois Shepherd	Industry	Tumour bank data project
Lois Shepherd	Industry	Fellowship funding
Sherry Taylor	Ontario Women's Health Council	Genetics education for health care provinces
Sherry Taylor	Ontario Women's Health Council	Research on a new genetic mutation screening
		technique for hereditary breast and ovarian
		cancer
Sherry Taylor	Canadian Institutes of Health Research	Transdisciplinary cancer research training grant
Sherry Taylor	Queen's University	A phase II study of BAY 43-9006 (NSC 724772) in
		patients with hormone refractory prostate cancer
Xiaolong Yang	Canadian Institutes of Health Research	Molecular and cellular functions of tumor
		suppressor gene LATS
Xiaolong Yang	Queen's University	Tumor suppressors and cancer
Xiaolong Yang	Ontario Research Fund	Molecular and cellular functions of tumor
		suppressor genes
Xiaolong Yang	Canadian Foundation for Innovation	Molecular and cellular functions of tumor
		suppressor genes
Xiaolong Yang	Queen's University	Identification of tumour suppressor LATS
		kinase substrates
Iain Young	National Cancer Institute of Canada	Tumour bank contract
	Clinical Trials Group	
Dick Zoutman	Health Canada	Canadian surgical wound surveillance
Dick Zoutman	Health Canada	Clostridium difficile: national surveillance project
Dick Zoutman	Health Canada	Epidemiology of febrile respiratory infection
		in acute hospitals
Dick Zoutman	Community and Hospital Infection	Resources for infection control in long-term
	Control Association of Canada	care facilities



#### VISITING LECTURERS

Michael Bookman

Fox Chase Cancer Center,

Sean Egan

Hospital for Sick Children, Toronto, ON

Richard Fahlman

University of British Columbia,

Vancouver, BC

Nathalie Lamarche

McGill University, Montreal, QC

Peter McPherson

McGill University, Montreal, QC

Farah Moid

Ohio State University, Columbus, OH

William Muller

McGill University, Montreal, QC

Linda Penn

Ontario Cancer Institute, Toronto, ON

Peter Siegel

McGill University, Montreal, QC

Timothy Triche

University of Southern

California/Childrens Hospital Los Angeles,

Los Angeles, CA

William Trimble

University of Toronto, Toronto, ON

James Turkson

University of South Florida, Tampa, FL

John Veinot

University of Ottawa, Ottawa, ON

#### THE NATHAN KAUFMAN LECTURE

athan Kaufman was born in Lachine, Quebec and educated at McGill University graduating with a medical degree in 1941. He interned at the *Royal Victoria Hospital* and then served as a Medical Officer to a tank battalion in Western Europe and was honoured with an MBE. After 18 months as a pathology resident at the *Jewish General Hospital* in Montreal he moved with his wife Rita to the *Cleveland Metropolitan General Hospital* to complete his residency. He then joined the Faculty at *Case Western* and quickly began a successful and satisfying career in iron metabolism research, medical education and laboratory administration. In 1967 after seven years as a Professor at *Duke University*, he was recruited by Dean Harry Botterell to succeed Bob More as the Head of Pathology at Queen's.

His accomplishments at Queen's were numerous. They included the development of the NCIC Cancer Research Unit, recruitment and nurturing of many senior faculty, distinguished service to senior committees of the Hospital, University and the Medical Research Council of Canada, and expansion of our research and training programs. During his twelve years at Queen's, Dr. Kaufman became internationally recognized for his distinguished leadership as Editor of

Laboratory Investigation, President of the US-Canadian Academy of Pathology and the International Academy of Pathology. On leaving Queen's he moved to Augusta as the first full-time Secretary/Treasurer of the USCAP. He has been recognized by the USCAP for his numerous contributions and was honoured with the establishment of the Nathan Kaufman Timely Topics Lecture which is delivered annually at the USCAP meeting.

Through this lectureship the Department honours Nathan Kaufman's extraordinary influence in shaping the scholarly life of our department and his international contributions to academic pathology.

2004 Dietrich Keppler, German Cancer Centre, University of Heidelberg, Heidelberg, Germany

2003 Ulf Lindahl, Uppsala University, Uppsala, Sweden

2002 Janet Rossant, Samuel Lunenfeld Research Institute, Mount Sinai Hospital, Toronto, ON

2000 Errol Friedberg, Southwestern Medical School, TX

#### THE M. DARIA HAUST LECTURE

. Daria Haust was born in Poland, and graduated summa cum laude from medical school of Heidelberg University in 1951. After emigrating to Canada with her husband in 1952, she entered a rotating internship at the Kingston General Hospital. She undertook a year of atherosclerosis research with Dr. Robert More and then enrolled in the general pathology residency program at Queen's. In 1959 she achieved specialty certification from the Royal College of Physicians and Surgeons and also obtained an MSc-degree from Queen's. Following a postdoctoral fellowship in pediatric pathology with Dr. Benjamin Landing at the Cincinnati Children's Hospital, Dr. Haust joined the Faculty at Queen's in 1960. She successfully nurtured two sons, embarked on a career in experimental atherosclerosis and pediatric pathology and, in 1967, moved to the University of Western Ontario where she remains active as Professor Emeritus.

Dr. Haust has had a distinguished career in basic research in several areas including atherosclerosis, the process of elastogenesis, and the pathogenesis of several genetic diseases. She is a highly regarded educator and has played important roles in the establishment of national and international scientific societies. She also served on a number of Editorial Boards of scientific journals and was invited to lecture around the globe on countless occasions. Her scientific contributions have been honoured by a multiplicity of distinctions (e.g., the *Canada Council Killam Prize in Medicine*; Gold Medal Award from the *International Atherosclerosis Society*; Honorary membership of the *Academy of Science of Heidelberg*; Best Teacher Awards at the *University of Western Ontario*; the William Boyd Lectureship of the *Canadian* 

Association of Pathology; the Andreas Vesalius Medal by the University of Padua; Honorary Professorship of Medicine from the University of Chile, and many others). Dr. Haust's biography, as one of the Founders of Pediatric Pathology, appeared in 2001, and a Festschrift in her honour with contributions from 14 international scientists was published in a scientific journal in 2002. In 2004 she received the Distinguished Pathologist Award from the US/Canadian Academy of Pathology. She is the recipient of honorary degrees in medicine from three mediaeval Universities (Jagiellonian University, Krakow; Charles University, Prague; Havana University, Havana).

Through this lectureship the Department honours Daria Haust's scholarly achievements, her contributions to her profession, and her continuing passionate devotion to our department and to Queen's University.

- 2005 Timothy Triche, Professor of Pathology and Pediatrics,
   University of Southern California, and Head,
   Department of Pathology, Childrens Hospital Los
   Angeles, Los Angeles, CA
- 2003 Luc Oligny, Pediatric and Molecular Pathologist, Department of Pathology and Cellular Biology, Université de Montréal and Hôpital Sainte-Justine, Montréal,P Q
- 2001 Kurt Benirschke, Emeritus Professor of Pathology and Reproductive Medicine, University Medical Center, San Diego, CA

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#### INVITED LECTURES AND PRESENTATIONS BY FACULTY

#### Susan Cole

Markey Cancer Center, University of Kentucky, Lexington, KY
School of Pharmacy, University of North Carolina, Chapel Hill, NC
13th Meeting on Pollution Responses in Marine Organisms, Alessandria, Italy John Roder 25th Anniversary Symposium, Mount Sinai Hospital, Toronto, ON NCIC/CCS Robert L. Noble Prize Lecture, Queen's University, Kingston, ON

12th Annual MDIBL Environmental Health Sciences Symposium, Salisbury Cove. ME

2nd Annual North American ABC Genetic Workshop, Frederick, MD

#### **Christine Collier**

Canadian Society of Nephrologists, Calgary, AB

#### **Roger Deeley**

2005 FASEB Summer Research Conference, FASEB, Saxtons River, VT NCIC/CCS Robert L. Noble Prize Lecture, Queen's University, Kingston, ON

#### David Dexter

Ontario Coroners' Meeting, Niagara Falls, ON

#### **Bruce Elliott**

le Centre de Recherche en Cancerologie de l'Universite Laval, Quebec City, QC

#### Peter Greer

The Hospital for Sick Children, Toronto, ON

The Blood Centre of Southeastern Wisconsin, WI

Ottawa Health Research Institute, Center for Cancer Therapeutics, Ottawa, ON

#### **Karen Harrison**

Michener Institute, Toronto, ON

#### **Robert Kisilevsky**

Council of Academic Hospitals of Ontario, Toronto, ON

#### David LeBrun

Ottawa Hospital Research Institute, Ottawa, ON

#### David Lillicrap

Hemophilia Products: Safety, Function and Alternative Administration, London, UK

Department of Genetics, McGill University, Montreal, QC

Department of Pathology and Molecular Medicine, McMaster University, Hamilton, ON

Fifth Bari International Conference on Hemophilia and Allied Disorders, Foggia, Italy

Mayo Clinic, Rochester, MN Imperial College School of Medicine, London, UK

Sir John Dacie Lecture, 37th Annual Advances in Haematology Course, Imperial College, London, UK

International Society on Thrombosis and Hemostasis, Sydney, Australia

Hemophilia Foundation of Australia Annual Conference, Melbourne, Australia

Baxter Bioscience, Toronto, ON US National Hemophilia Foundation 57th Annual Meeting, San Diego, CA

University of Nevada, Department of Pediatrics, Las Vegas, NV Genzyme Inc, Boston, MA Canadian Stem Cell Network Annual Meeting, Calgary, AB

#### Sam Ludwin

Multiple Sclerosis Update, Philadelphia PA
19th Annual Meeting of the Consortium of
Multiple Sclerosis Centres, Orlando, FL
40th Meeting of the Canadian Congress
of Neurological Sciences, Ottawa, ON
Symposium on the Role of T Cells
in Pathological and Clinical Events
in Multiple Sclerosis, Carolinas
Healthcare System, Charlotte, NC

#### Lois Mulligan

AACR, Anaheim, CA

First International Symposium on

Pheochromocytoma,

Pheochromocytoma Research

Support Organization (PRESSOR),

Bethesda, MD

#### Lois Shepherd

Canadian Society of Transfusion Medicine, St. John's, NFLD

#### **Sherry Taylor**

Faculty of Law, Health Law Day, University of Toronto, Toronto, ON

#### **Iain Young**

University of Toronto, Toronto, ON St. Michael's Hospital, Toronto, ON

#### Dick Zoutman

SEO Symposium on Emergency Preparedness, Healthcare Network of SEO, Kingston, ON

Ontario Hospital Association Symposium on Hospital Infections, Ontario Hospital Association, Toronto, ON

Community & Hospital Infection Control Association-Canada Annual Meeting, Winnipeg, MB

#### PATENTS GRANTED

**Susan Cole** and **Roger Deeley** Multidrug Resistance Protein: Japanese Patent

#### **Robert Kisilevsky**

Method for Treating Amyloidosis: Japanese Patent Method for Treating Amyloidosis: European Patents (2)

#### David Lillicrap

Canine Factor VIII cDNA Gene Therapy – US Patent

#### LICENSES GRANTED

Susan Cole and Roger Deeley Solvo Biotechnologies

#### **PUBLICATIONS**

- R. Leong, B.R. Gannon, **T.J. Childs**, **P.A. Isotalo**, H. Abdollah. Aspergillus fumigatus pacemaker endocarditis: a case report and review of the literature. *Canadian Journal of Cardiology* (in press).
- C.J. Westlake, **S.P.C. Cole** and **R.G. Deeley**. Role of the NH2-terminal membrane spanning domain of MRP1/ABCC1 in protein processing and trafficking. *Mol. Biol. Cell* 16: 2483-2492, 2005.
- L. Payen, M. Gao, C. Westlake, A. Theis, **S.P.C. Cole** and **R.G. Deeley**. Functional interactions between nucleotide binding domains and LTC4 binding sites of multidrug resistance protein 1 (ABCC1). *Mol. Pharmacol*. 67: 1944-1953, 2005.
- D-W. Zhang, K. Nunoya, M. Vasa, H-M. Gu, **S.P.C. Cole** and **R.G. Deeley**. Mutational analysis of polar amino acid residues within predicted transmembrane helices 10 and 16 of multidrug resistance protein 1 (ABCC1): effect on substrate specificity. *Drug Metab. Disp.* doi.10.1124.dmd. 105.007740.
- G. Conseil, **R.G. Deeley** and **S.P.C. Cole**. Polymorphisms of MRP1 (ABCC1) and related ATP-dependent drug transporters. *Pharmacogenet*. *Genomics* 15: 523-533, 2005.
- I.J. Létourneau, R.G. Deeley and S.P.C. Cole. Functional characterization of non-synonymous single nucleotide polymorphisms in the gene encoding human multidrug resistance protein 1 (MRP1/ABCC1). *Pharmacogenet*. *Genomics* 15: 647-657, 2005.
- G. Conseil, **R.G. Deeley** and **S.P.C. Cole**. Role of two adjacent cytoplasmic tyrosine residues in MRP1 (ABCC1) transport activity and sensitivity to sulfonylureas. *Biochem. Pharmacol.* 69: 451-461, 2005.

- E.M. Leslie, R.G. Deeley and S.P.C. Cole. Multidrug resistance proteins in toxicology: role of P-glycoprotein, MRP1, MRP2 and BCRP (ABCG2) in tissue defense. *Toxicol. Appl. Pharmacol.* 204: 216-237, 2005.
- I.J. Létourneau, R.J. Bowers, R.G. Deeley and S.P.C. Cole. Limited modulation of the transport activity of the human multidrug resistance proteins MRP1, MRP2 and MRP3 by nicotine glucuronide metabolites. *Toxicol. Lett.* 157: 9-19, 2005.
- P. Wu, C.J. Oleschuk, Q. Mao, B.O. Keller, R.G. Deeley and S.P.C. Cole. Analysis of human multidrug resistance protein 1 (ABCC1) by matrix-assisted laser desorption ionization/time of flight mass spectrometry: toward identification of leukotriene C4 binding sites. *Mol. Pharmacol.* 68: 1455-1465, 2005.
- **R.G. Deeley** and **S.P.C. Cole**. Substrate recognition and transport by multidrug resistance protein (MRP) 1 (ABCC1). *FEBS Lett.* doi.10.1016/j.febslet.2005.12.036.
- J.A. Simpson, R. Labugger, **C.P. Collier**, R.J. Brison, S. Iscoe, and J.E. Van Eyk. Fast and slow skeletal troponin I in serum from patients with various skeletal muscle disorders: a pilot study. *Clin. Chem.* 51(6): 966-72, 2005.
- **C. Collier**. Estimating glomerular filtration rate (eGFR). *CSCC News* August 2005.
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#### **BOOK CHAPTERS**

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# Laboratory Medicine and Clinical Services

Through its divisions of Anatomic Pathology, Hematopathology, Clinical Chemistry, Microbiology, and Genetics, the department provides services in Laboratory Medicine, Transfusion Medicine, Infection Control and Clinical Genetics to both the *Academic Health Sciences Centre* and the broader region of southeastern Ontario.

In 2005, there were a number of exciting changes and developments in the Clinical Laboratories, most notably involving new staff and new equipment. Due to a departure and some retirements at the management level, we started the year with a new Administrative Director and new Managers in Core Lab and Microbiology/Infection Control. We created several new senior and charge technologist positions throughout the Labs and continued our focus on having front-line staff participate through Continuous Quality Improvement teams to address some process and structural issues in the Labs. Our department has been congratulated by the hospital for our excellent quality assurance program and we continue to be a model for others to strive towards.

The provincial government provided \$3.48 million to our region to purchase diagnostic medical equipment and, as a result, we obtained a total of 46 new instruments for our region's hospital laboratories. Locally, this included coagulation analyzers, blood gas analyzers, a variant hemoglobin analyzer, automated bacterial identification and sensitivity system, automated blood bank system, automated immunohistochemistry stainers, histology slide stainer, labeler and an embedding centre. PCR instrumentation was introduced to the Microbiology Laboratory, where its initial application has been the identification of methicillin resistant Staph aureus, and expanded in diagnostic molecular genetics. With funding from the OCRN, we began operating a Tumour Bank which harvests fresh tissue from the Operating Room for the cancer research. Renovations to the Autopsy Suite, including the installation of x-ray equipment, were virtually completed in 2005, permitting us to improve the efficiency and quality of the work done in our Regional Forensic Unit. Our Infection Control Program continues to grow: we added a fourth Infection Control Practitioner and, at a regional level, we played a lead role in the development of a regional Infection Control program which has received significant new funding from the provincial government.

#### CLINICAL LEADERSHIP

Iain Young, Department Head
Sandip SenGupta, Deputy Head and
Medical Director, Clinical Laboratories

#### Division of Anatomic Pathology

Sandy Boag, Service Chief
David Dexter, Clinical Director, Regional
Forensic Unit and Autopsy Service
Caroline Rowlands, Clinical Director,
Cytology

#### DIVISION OF HEMATOPATHOLOGY

Dilys Rapson, Service Chief Lois Shepherd, Clinical Director, Blood Bank and Transfusion Medicine

#### Division of Clinical Chemistry Lorne Seargeant, Service Chief

#### Division of Microbiology

Dick Zoutman, Service Chief and Clinical Director, Infection Control

#### **DIVISION OF GENETICS**

David Lillicrap, Service Chief
Karen Harrison, Laboratory Coordinator
and Clinical Director, Cytogenetics
Sherry Taylor, Clinical Director,
DNA Diagnostics

#### REGIONAL LABORATORY

#### OUTREACH PROGRAM

Sandip SenGupta, Clinical Director
David Dexter, Laboratory Director,
Kingston MDS Laboratories
Dilys Rapson, Laboratory Director, Lennox
and Addington Hospital
Mike Raymond, Laboratory Director,
Weeneebayko Regional Hospital
Sandip SenGupta, Laboratory Director,
Perth/Smiths Falls Hospital and Ongwanada
Dick Zoutman, Laboratory Director and
Medical Director of Infection Control,

Providence Continuing Care Centre

# Awards, Honours and Achievements

#### **FACULTY**

#### PERSONNEL AWARDS

Susan Cole, Canada Research Chair, Tier I, Cancer Biology.

Scott Davey, Cancer Care Ontario Scientist Salary Award.

Roger Deeley, Stauffer Research Professor, Faculty of Health Sciences, Queen's University.

David Lillicrap, Canada Research Chair, Tier I, Molecular Hemostasis.

David Lillicrap, Career Investigator Award, Ontario Heart and Stroke Foundation.

#### International

#### Susan Cole

Elected to the Nominating Committee of the American Association for Cancer Research.

Elected Co-Chairperson, 2007 Gordon Research Conference on "Multi-Drug Efflux Systems".

Member, Scientific Advisory Board, Federation of European Biological Societies.

Co-organizer, FEBS Special Meeting on ABC Proteins -ABC 2006, Federation of European Biological Societies.

Member, External Advisory Board, Roswell Park Cancer Institute.

#### **Christine Collier**

Program Chair (President Elect), Upstate NY-American Association of Clinical Chemists.

Councillor, International Association of Therapeutic Drug Monitoring and Clinical Toxicology.

#### **Robert Kisilevsky**

Board of Director, Member at Large, International Society of Amyloidosis.

Ad Hoc Consultant, Project Program Review, National Institute of Health.

Member, Advisory Committee, XIth International Symposium on Amyloidosis.

Member, Scientific Advisory Board, Institute for the Study of Aging.

Member, Editorial Board, Amyloid: The Journal of Protein Folding Disorders.



#### ROBERT L. NOBLE PRIZE PRESENTATIONS AND LECTURE

Wednesday, November 16, 2005, 5:00 P.M. **BOTTERELL HALL, ROOM B139** 

RECIPIENTS OF THE 2005 ROBERT L. NOBLE PRIZE



Roger G. Deeley Ph.D.

Lecture Title

"Multidrug Resistance Protein 1...and Friends"

The Robert L. Noble Prize is sponsored by the Canadian Cancer Society and awarded by the National Cancer Institute of Canada for outstanding achievements in cancer research



Canadian Société
Cancer canadienne
Sociéty du cancer
Of Canada du Canada
du Canada
du Canada



#### David Lillicrap

Chairperson, Gene Therapy and Novel Technology Committee, World Federation of Hemophilia.

Co-Chairperson, Scientific Standardization Committee Subcommittee on Von Willebrand Factor, International Society for Thrombosis and Hemostasis.

Presented the Inaugural Sir John Dacie Lecture at the Imperial College School of Medicine, London, UK

Presented the Arosenius Lecture "Genetic and Cellular Therapies for Hemophilia" to the Swedish Hemophilia Society.

#### Samuel Ludwin

Delivered the lecture, "Oligodendrocytes and Tissue Repair in MS" at the 2005 MS Update Meeting, Philadelphia, PA.

Delivered the lecture, "Understanding MS through Pathology", at the 19th Annual Meeting of the Consortium of MS Centres, Navigating the World of MS, Orlando, Florida.

Delivered the lecture, "Current Understanding of the Evolution of Cellular Pathology in MS", at the Symposium on the Role of T Cells in Pathological and Clinical Events in MS by the Carolinas HealthCare System, Charlotte, NC.

Invited Chair/Organizer, Ectrims/Actrims Annual Meeting, Thessaloniki, Greece.

#### Lois Shepherd

Member, Group Banking Committee, US National Cancer Institute.

Member, International Development Project, Breast International Group.

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#### NATIONAL

#### Susan Cole

Co-recipient of the Robert L. Noble Prize of the Canadian Cancer Society/National Cancer Institute of Canada.

Elected Researcher of the Week (February 16, 2005) by Canadians for Health Research.

Member, College of Reviewers, Canada Research Chair Program.

Member, Research Advisory Committee, Cancer Care Ontario.

#### **Christine Collier**

Received the 2005 Canadian Association of Medical Education (CAME) Certificate of Merit.

President, Ontario Society of Clinical Chemists.

#### **Roger Deeley**

Director, Division of Research, Cancer Care Ontario.

Co-recipient of the Robert L. Noble Prize of the Canadian

Cancer Society/National Cancer Institute of Canada.

#### Phillip Isotalo

Received the award for Best Poster Presentation at the 2005 *Canadian Surgery Forum*.

#### **Robert Kisilevsky**

Recipient of the Award of Excellence – Emerging Technology (AtheroChem) of the Kingston Technology Council.

#### David Lillicrap

Chair, von Willebrand Disease Scientific Subcommittee of the Association of Hemophilia Clinic Directors of Canada.

#### Samuel Ludwin

Chair, Medical Advisory Committee, Canadian Multiple Sclerosis Society.

#### Dilys Rapson

Vice Chair, Examination Board for Hematological Pathology, RCPSC.

#### John Rossiter

Vice-Chair, Neuropathology Fellowship Examination Board, RCPSC.

#### Lois Shepherd

Vice-President, Canadian Society of Transfusion Medicine.
Physician Coordinator of Multicentered Phase III Clinical
Trials, National Cancer Institute of Canada – Clinical Trials
Group.

Member, Management Committee, Canadian Tumour Repository Network.

Member, Clinical Advisory Committee in Transfusion Medicine, Canadian Blood Services.

#### **Sherry Taylor**

Chair, Laboratory Genetics Fellowship Review Committee, Ontario Ministry of Health.

#### Dick Zoutman

Co-Chair, Provincial Infectious Diseases Advisory Committee.

#### University

#### **David Hurlbut**

Received the Department of Pathology and Molecular Medicine R.S.A. Prentice Award for Excellence in Teaching.

#### **Lewis Tomalty**

Received the Reddick Award for excellence in nursing education from the School of Nursing

#### **RESIDENTS**

R.S.A. PRENTICE AWARD

Kristine Roland, for best presentation

#### **GRADUATE STUDENTS**

ACADEMIC HONOURS AND AWARDS

Queen's University Governor General's

ACADEMIC GOLD MEDAL

Adina Vultur, received upon graduation

#### Daria Haust Award

Zoya Shapovalova

#### ROBERT KISILEVSKY AWARD

Rozanne Arulanandam

Taranjit Gujral

Doug Richardson

Andrew Slot

## POSTDOCTORAL FELLOWS AND TRAINEES

#### GEORGE CHRISTIAN HOFFMAN FELLOWSHIP

Takayuki Murase

Akio Nakajima

Maha Othman

#### NOVO Nordisk Canada

INC-CHS-AHCDC FELLOWSHIP

Maha Othman, for congenital and acquired bleeding disorders.

#### MOH-LTC Laboratory Genetics Fellowship

Shulin Zhang