## **5th FEBS Advanced Lecture Course**

# Matrix Pathobiology, Signaling and Molecular Targets

Rhodes, September 24-September 29, 2015

Program: Lectures (L), Selected Talks (ST), Posters (P)

## Thursday, 24 September

13:45-15:45 Registration

### Chairperson/discussion leader: John Couchman

GENERAL LECTURES / TUTORIALS (11-16)

OLNLKALL	LCTORES / TOTORIALS (LT-LO)
15:45-16:05	<b>A. Theocharis</b> (University of Patras, Greece) Overview of extracellular matrix: structure and functional properties
16:05-16:10	Discussion
16:10-16:30	J. Couchman (University of Copenhagen, Denmark) Proteoglycans: Structure, pathobiology, and signaling
16:30-16:35	Discussion
16:35-16:55	P. Heldin (University of Uppsala, Sweden) Insights into the function of glycans
16:55-17:00	Discussion
17:00-17:30	Coffee break
17:30-17:50	JO. Winberg (University of Tromsø, Norway) Matrix Metalloproteinases: biological significance in health and disease
17:50-17:55	Discussion

17.55-18.15 M. Franci (University of Bologna, Italy) Collagen and functional implications

18:15-18:20 Discussion

18.20-18.40 **D. Gullberg** (University of Bergen, Norway)

ECM cell surface receptors

18:40-18:45 Discussion

### Chairpersons: Nikos Karamanos & Renato Iozzo

18.45-19:00 Welcome Addresses by

- Chairman and members of the Organizing Committee

- Rector of the University of Patras

- Chair of the FEBS Advanced Courses Committee

- President of Hellenic Society of Biochemistry and Molecular Biology

### IUBMB LECTURE (L7)

19:00-20:00 H. Nagase (University of Oxford, UK)

The endocytic receptor LRP1 is a master regulator of extracellular

levels of ECM-degrading metalloproteinases

Honorory Medal Award conferred to Professor Hideaki Nagase

by the Rector of the University of Patras

20:00 Welcome Reception

# Friday, 25 September

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	ussion leader: Peter Friedl TUTORIALS - MATRIX PATHOBIOLOGY (L8-L10)
9:00-9:20	<b>M. Paulsson</b> (University of Cologne, Germany) How perifibrillar proteins influence collagen secretion and fibril interactions
9:20-9:25	Discussion
9:25-9:45	M. Elkin (Hadassah-Hebrew University Medical Center, Jerusalem, Israel) Tumor-host interactions in pancreatic carcinoma: role of heparanase enzyme
9:45-9:50	Discussion
9:50-10:10	P. Friedl (Radbound University, Netherlands) Tissue niches for collective cancer cell invasion and therapy resistance
10:10-10:15	Discussion
SELECTED T	ALK (ST1)
10:15-10:25	<b>M. Onisto</b> (University of Padova, Italy) Novel molecular pathways potentially involved in renal fibrosis: role of HIPK2 and Heparanase
10:25-10:30	Discussion
10:30-11:00	Coffee break
	ussion leader: Martin Götte TUTORIALS - MATRIX PATHOBIOLOGY (L11-L13)
11:00-11:20	<b>T. Wight</b> (University of Washington, USA) Versican: An enigmatic matrix component and the control of disease cell phenotype
11:20-11:25	Discussion
11:25-11:45	M. Götte (Muenster University, Germany) Interplay of syndecan-1 and heparanase in cancer stem cell function
11:45-11:50	Discussion

11:50-12:10	<b>K. Dobra</b> (Karolinska University, Stockholm, Sweden) The role of syndecan-1 in mesenchymal tumors	
12:10-12:15	Discussion	
SELECTED T	ALKS (ST2-ST5)	
12.15-12:25	E. Regős (Semmelweis University, Hungary) Syndecan-1 in liver fibrosis and regeneration	
12:25-12:30	Discussion	
12:30-12:40	P. Bouris (University of Patras, Greece) Tumorigenic role of serglycin in breast cancer cells	
12:40-12:45	Discussion	
12:45-12:55	<b>D. Barbouri</b> (University of Patras, Greece) Syndecans as key partners in the interaction between breast cancer cells and endothelium	
12:55-13:00	Discussion	
13:00-13:10	M. Farshchian (University of Turku, Finland) AIM2 promotes progression of cutaneous squamous cell carcinoma	
13:10-13:15	Discussion	
13:15-14:45	Lunch	
14:45-16:30	Poster session (I) (P1-P26)/ Discussion groups (I)	
Chairperson/discussion leader: Renato lozzo LECTURES/TUTORIALS - MATRIX PATHOBIOLOGY (L14-L16)		
16:30-16:50	<b>R. lozzo</b> (Thomas Jefferson University, USA)  Novel proteoglycan roles in evoking autophagy independent of nutrient deprivation	
16:50-16:55	Discussion	
16:55-17:15	I. Kovalszky (Semmelweis University, Hungary) Tumor-stroma interaction of hepatomas with slow or fast proliferation rate	
17:15-17:20	Discussion	
17:20-17:40	K. Forsberg-Nilsson (University of Uppsala, Sweden) Modeling glioma and targeting the glioma niche	
17:40-17:45	Discussion	
SELECTED T	ALKS (ST6-ST8)	
17:45-17:55	T. Neill (Thomas Jefferson University, USA) A new mechanistic link between decorin-evoked Peg3 and TFEB for autophagic induction and angiostasis	
17:55-18:00	Discussion	

18:00-18:10	K. Baghy (Semmelweis University, Hungary) Decorin expression in human hepatocellular carcinoma
18:10-18:15	Discussion
18:15-18:25	<b>D. Avery</b> (University of Pennsylvania, USA) The role of fibroblast activation protein (FAP) in lung tumorigenesis
18:25-18:30	Discussion
18:30-19:00	Break
Chairpersons: Nikos Karamanos & Dimitris Kletsas HONORARY PLENARY LECTURE (L17)	
19:00-20:00	R Sanderson (University of Alahama, USA)

R. Sanderson (University of Alabama, USA) 19:00-20:00

Targeting heparanase for myeloma therapy

Honorary Medal Award conferred to Prof. R. Sanderson

by the Rector of the University of Patras

20:00 Dinner

## Saturday, 26 September

### Chairperson/discussion leader: Paraskevi Heldin LECTURES/TUTORIALS - GLYCOBIOLOGY AND METABOLIC REGULATION OF ECM MOLECULES (L18-L19) 9:00-9:20 V. Hascall (Cleveland Clinic Foundation, USA) How does heparin prevent intracellular synthesis of hyaluronan in hyperglycemic dividing cells? 9:20-9:25 Discussion 9:25-9:45 **P. Heldin** (University of Uppsala, Sweden) Role of hyaluronan-CD44 interactions during cancer progression 9:45-9:50 Discussion SELECTED TALKS (ST9-ST11) M. Mehić (Uppsala University, Sweden) 9:50-10:00 The deubiquitinating enzymes USP4 and USP17 target hyaluronan synthase 2 and affect its function 10:00-10:05 Discussion 10:05-10:15 **D. Vigetti** (University of Insubria, Italy) The long non-coding RNA HAS2-AS1 is a new regulator of hyaluronan synthesis and its involved in human vascular diseases 10:15-10:20 Discussion

	10:20-10:30	AJ. Deen (University of Eastern Finland, Finland) The UDP-sugar substrates of hyaluronan synthase 3 (HAS3) regulate its intracellular traffic and extracellular shedding, controlling hyaluronan synthesis and correlating with phenotypic changes in early stages of melanoma
	10:30-10:35	Discussion
	10:35-11:05	Coffee break
Chairperson/discussion leader: Alberto Passi LECTURES/TUTORIALS - I GLYCOBIOLOGY AND METAE REGULATION OF ECM MOLECULES (L20-L22)		UTORIALS - I GLYCOBIOLOGY AND METABOLIC
	11:05-11:25	<b>H. Watanabe</b> (Aichi Medical University, Japan) Role of versican, and chondroitin sulfate in development and diseases
	11:25-11:30	Discussion
	11:30-11:50	A. Passi (University of Insubria, Italy) Hyaluronan amount and size are critical in human pathology
	11:50-11:55	Discussion
	11:55-12:15	<b>L. Kjellen</b> (University of Uppsala, Sweden) Cellular design of heparan sulfate
	12:15-12:20	Discussion
	SELECTED TA	ALKS (ST12-ST15)
	12:20-12:30	<b>T. Dierker</b> (Uppsala University, Sweden) Detection of hitherto unknown chondroitin core proteins together with unexpected sulfation of chondroitin in the nematode C. elegans
	12:30-12:35	Discussion
	12:35-12:45	F. Malfait (Ghent University, Belgium) Zebrafish modeling of $\beta$ 3GalT6 and $\beta$ 4GalT7-deficient types of Ehlers-Danlos syndrome stresses the importance of glycosaminoglycans in development
	12:45-12:50	Discussion
	12:50-13.00	<b>SL. Taylor</b> (University of Liverpool, UK) The purification and characterization of heparin by-products for applications in wound healing
	13:00-13:05	Discussion
	13:05-13:15	<b>A. Abbadi</b> (Cleveland Clinic Foundation, USA) Heparin inhibits the differentiation of M1 and promotes M2 macrophages under hyperglycemic stress
	13:15-13:20	Discussion
	13:20-14:15	Speakers corner (I)
	14:15-21:30	Lunch box - Excursion tour

11:45-11:50

11:50-12:00

Discussion SELECTED TALKS (ST19-ST20)

### Chairperson/discussion leader: Donald Gullberg

LECTURES/TUTORIALS-	INTERACTIONS	AND FUNCTIONS OF
MATRIX MACROMOLECU	LES (L23-L24)	

9:00-9:20	M. Franci (University of Bologna, Italy) Collagen in tendon and ligament	
9:20-9:25	Discussion	
9:25-9:45	<b>D. Gullberg</b> (University of Bergen, Norway) Collagen receptors- static contacts or dynamic modulators?	
9:45-9:50	Discussion	
SELECTED TA	ALKS (ST16-ST18)	
9:50-10:00	<b>D. Gavriilidou</b> (Imperial College London, UK) Interactions of discoidin domain receptors with fibrillar and non-fibrillar collagen forms: ligand binding versus receptor activation	
10:00-10:05	Discussion	
10:05-10:15	<b>B. Pilecki</b> (University of Southern Denmark, Denmark) MFAP4 is a novel regulator of elastic fiber assembly and integrin-dependent cellular signaling	
10:15-10:20	Discussion	
10:20-10:30	<b>S. Molon</b> (University of Padova, Italy) Role of collagen VI in intestine homeostasis	
10:30-10:35	Discussion	
10:35-11:05	Coffee break	
Chairperson/discussion leader: François-Xavier Maquart LECTURES/TUTORIALS - INTERACTIONS AND FUNCTIONS OF MATRIX MACROMOLECULES (L25-L26)		
11:00-11:20	<b>FX. Maquart</b> (University of Reims, France) Basement membrane collagens in the control of tumor cell invasion	
11:20-11:25	Discussion	
11:25-11:45	A. Blom (Lund University, Malmo, Sweden) Cartilage oligomeric matrix protein (COMP) contributes to the development and metastasis of breast cancer	

Mesenchymal stem cells adhesive behavior to different extracellular matrix proteins alters during osteogenic differentiation

M. Keremidarska (Bulgarian Academy of Sciences, Bulgaria)

12:00-12:05	Discussion
12:05-12:15	AC. Silva (Universidade do Porto, Porto, Portugal) The versatility of fetal and adult ECM-derived microenvironments: The impact of ontogeny on cardiac cells
12:15-12:20	Discussion
12:30-14:15	Lunch
14:15-16:15	Poster session (II) P27-P53/ Discussion groups (II)
16:15-17:00	BG. Vertessy (Budapest University of Technology and Economics, Hungary) Career Planning
CELL RECEPT	ession leader: Liliana Schaefer FORS SIGNALING AND ECM BASED OLOGY (L27-L30)
17:00-17:20	L. Schaefer (Goethe-Universität Frankfurt am Main, Germany) SLRP signaling in inflammation
17:20-17:25	Discussion
17:25-17:45	J. Couchman (University of Copenhagen, Denmark) Syndecans: transmembrane proteoglycans controlling the cell adhesion phenotype
17:45-17:50	Discussion
17:50-18:10	M. Pavao (University of Rio de Janeiro, Brasil) Inhibition of manganese-mediated tumor cell migration by heparin analogs
18:10-18:15	Discussion
18:15-18:45	Break
18:45-19:05	<b>RC. Savani</b> (University of Texas Southwestern Medical Center, USA) Molecular mechanism of RHAMM activation of Src kinase and the signaling pathway for nitric oxide production in the endothelium
19:05-19:10	Discussion
SELECTED TA	ALKS (ST21-ST23)
19:10-19:20	FCOB. Teixeira (Universidade Federal do Rio de Janeiro, Brasil) Sulfated fucans and sulfated galactans from sea urchins as potent inhibitors of selectin-dependent hematogenous metastasis. Are there any structural requirements?
19:20-19:25	Discussion
19:25-19:35	<b>AO. Melleby</b> (University of Oslo, Norway) Glypican-6 is increased in experimental and clinical heart failure and might play a role in cardiac fibrosis through BMP4 signaling
19:35-19:40	Discussion

19:40-19:50	<b>H. Pratsinis</b> (NCSR "Demokritos", Greece)  Three-dimensional culture systems for the study of growth factors on intervertebral disc cells
19:50-19:55	Discussion
20:00	Dinner

# Monday, 28 September

Monday, 20 ocptember		
Chairperson/discussion leader: Carl-Henrik Heldin SIGNALING AND DISEASE MOLECULAR TARGETING (L31-L33)		
9:00-9:20	CH. Heldin (Ludwig Cancer Institute, Sweden) Smad and non-Smad signaling via TGF- $\beta$ receptors – possible targets in tumor therapy	
9:20-9:25	Discussion	
9:25-9:45	N. Karamanos (University of Patras, Greece) The regulatory roles of estrogen receptors and syndecans in breast cancer cell properties and functions	
9:45-9:50	Discussion	
9:50-10:10	<b>D. Nikitovic</b> (University of Crete, Greece) IGF-I/ EGF signaling affect breast cancer cell adhesion through cytoskeleton reorganization	
10:10-10:15	Discussion	
SELECTED T	ALKS (ST24-ST28)	
10:15-10:25	N. Afratis (University of Patras, Greece) Syndecan-4 as a switch of epithelial to mesenchymal transition in breast cancer cells	
10:25-10:30	Discussion	
10:30-10:40	<b>SK. Jha</b> ( <i>University of Helsinki, Finland</i> ) CCBE1 enhances lymphangiogenesis by regulating VEGF-C activation	
10:40-10:45	Discussion	
10:45-11:15	Coffee break	
11:15-11:25	J. Nuechel (University of Cologne, Germany) TGF-β release by fibroblasts requires regulated secretion via autophagosomal intermediates	
11:25-11:30	Discussion	
11:30-11:40	C. Kolliopoulos (University of Uppsala, Sweden) HMGA2 regulates HAS2 and its natural antisense transcript during TGF-β-mediated EMT	
11:40-11:45	Discussion	

11:45-11:55	<b>Z. Piperigkou</b> (University of Patras, Greece) The role of ER $\beta$ in regulation of functional properties and gene expression of matrix macromolecules in aggressive breast cancer cells
11:55-12:00	Discussion
Chairperson/discu	ssion leader: Jerry Turnbull
S I G N A L I N G / ( L 3 4 - L 3 5 )	AND DISEASE MOLECULAR TARGETING
12:00-12:20	J. Turnbull (University of Liverpool, UK) Proteoglycans as drug development targets: chemical biology routes to new therapeutics
12:20-12:25	Discussion
12:25-12:45	<b>D. Kletsas</b> (NCSR "Demokritos", Greece) Senescence of stromal fibroblasts triggered by anticancer treatments: implications in tumor progression
12:45-12:50	Discussion
SELECTED TA	ALK (ST29)
12:50-13:00	<b>T. Brown</b> (Monash University, Australia)  Development of hyaluronan as a CD44-targeted drug delivery vehicle in the treatment of cancer
13:00-13:05	Discussion
13:05-14:30	Lunch
14:30-16:00	Poster Session (III) P54-P79 / Discussion groups (III)
16:00-17:00	Speakers corner (II)
	ussion leader: Jan-Olof Winberg ULATION IN HEALTH AND DISEASE (L36-L38)
17:00-17:20	JO. Winberg (University of Tromsø, Norway) The core protein of serglycin is a matrix metalloprotease-9 substrate
17:20-17:25	Discussion
17:25-17:45	<b>S. Brezillion</b> (University of Reims, France) Lumican: A new inhibitor of matrix metalloproteinase-14 activity
17:45-17:50	Discussion
17:50-18:10	<b>S. Heymans</b> (KU Leuven, Belgium) Osteoglycin (mimecan) protects against ischemic heart disease but increases adverse cardiac inflammation in viral myocarditis
18:10-18:15	Discussion
18:15-18:45	Break

SELECTED	TALKS (ST30-ST33)
18:45-18:55	<b>E. Karousou</b> (University of Insubria, Italy) Cross talk between breast cancer cells and fibroblasts: effect of the uncharacterized protein q7z3e2 on hyaluronan synthesis
18:55-19:00	Discussion
19:00-19:10	<b>B. Deschrevel</b> (University of Rouen, France) An innovative biomaterial for tridimensional eukaryotic cell development: applications in tissue engineering and in tumor engineering
19:10-19:15	Discussion
19:15-19:25	X. Stachtea (Lund University, Sweden) Loss of dermatan sulfate results in neonatal lethality in mice despite normal lymphoid and non-lymphoid organogenesis
19:25-19:30	Discussion
19:30-19:40	ME. Strand (University of Oslo, Norway) Shedding of syndecan-4 promotes immune cell recruitment and preserves heart function after lipopolysaccharide challenge
19:40-19:45	Discussion
20:00	Dinner

# Tuesday, 29 September

Chairperson/discussion leader: Boris Turk  MATRIX REGULATION IN HEALTH AND DISEASE (L39 - L41)	
9:00-9:20	<b>B. Turk</b> ( <i>J. Stefan Institute, Slovenia</i> ) Extracellular cysteine cathepsins: from signalling to matrix degradation
9:20-9:25	Discussion
9:25-9:45	VM. Kähäri (University of Turku, Finland) Proteolytic control of skin cancer progression
9:45-9:50	Discussion
9:50-10:10	<b>D. Vynios</b> (University of Patras, Greece) ADAMTS proteinases in health and disease
10:10-10:15	Discussion
SELECTED TA	ALKS (ST34-ST37)
10:15-10:25	K. Andenæs (University of Oslo, Norway) The extracellular matrix proteoglycan fibromodulin is up-regulated in experimental and clinical heart failure, and might attenuate development of myocardial fibrosis by interacting with TGF-β signaling

10:25-10:30	Discussion
10:30-10:40	<b>B. Bluhm</b> (University of Cologne, Germany) MicroRNAs: posttranscriptional modulators of cellular and extracellular cartilage compartments
10:40-10:45	Discussion
10:45-10:55	<b>L. Monti</b> (University of Pavia, Italy) Animal models of desbuquois dysplasia type 1 to study the role of CANT1 in proteoglycan metabolism
10:55-11:00	Discussion
10:55-11:00 11:00-11:10	Discussion  AV. Suhovskih (Novosibirsk State University, Russia)  Prostate cancer cells stably change proteoglycans expression in normal fibroblasts in cell culture model in vitro
	AV. Suhovskih (Novosibirsk State University, Russia) Prostate cancer cells stably change proteoglycans expression
11:00-11:10	AV. Suhovskih (Novosibirsk State University, Russia) Prostate cancer cells stably change proteoglycans expression in normal fibroblasts in cell culture model in vitro
11:00-11:10 11:10-11:15	AV. Suhovskih (Novosibirsk State University, Russia) Prostate cancer cells stably change proteoglycans expression in normal fibroblasts in cell culture model in vitro Discussion