

# **XIV SPHINGOLIPID CLUB MEETING**

## **Pozzilli, Sept. 7-22, 2022**

### **PROGRAM**

**Wednesday, Sept. 7, 2022 – Grand Hotel Europa, Isernia**

14:00 - 20:30 *Welcoming and registration*

20:30 *Light dinner*

**Thursday, Sept. 8, 2022 – Neuromed Research Center, Pozzilli**

09:00 - 09:05 *Welcome greetings*

**Luigi Frati** *Scientific Director IRCCS Neuromed*

**Giovanni de Gaetano** *President IRCCS Neuromed*

09:05 - 09:15 *Opening remarks*

**Vittorio Maglione, Alba Di Pardo** (*IRCCS Neuromed, IT*)

09:15 - 09:50 *Keynote Lecture*

**Timothy Cox** (*University of Cambridge, UK*)

Sphingolipids – classical matters in the ancient and modern world

09:50 - 11:45 *Session 1 - Sphingolipids in health and disease (I)*

Chairs: **Marco Presta** (Brescia, IT), **Carmen Garcia-Ruiz** (Barcelona, ES)

09:50 - 10:15 **Lauren Ashley Cowart** (*Virginia Commonwealth University, US*)

Sptlc3 and novel sphingolipids in health and disease

10:15 - 10:30 **Duyen Tran** (*University of Tasmania, AU*)

Toxic 1-deoxysphinganine, compromises the functionality of skeletal myoblasts and underlies the development of type 2 diabetes mellitus

10:30 - 10:45 **Anna Kovilakath** (*Virginia Commonwealth University, US*)

The novel sphingolipid enzyme SPTLC3 ameliorates ischemic failure by reducing reactive oxygen species

10:45 - 11:00 **Marcelo Einicker-Lamas** (*Universidade Federal do Rio de Janeiro, BR*)

Sphingosine-1-phosphate prevents human embryonic stem cell death following ischemic injury

11:00 - 11:15 *Coffee break*

11:15 - 11:30 **Isabelle Seidita** (*University of Florence, IT*)

Sphingosine-1-phosphate acts as a pro-inflammatory cue in human endometrial stromal cells acting via ERK5: implication in endometriosis

11:30 - 11:45 **Zoltan Benyo** (*Semmelweis University, HU*)

Identification of signaling pathways mediating coronary flow reduction induced by sphingosine-1-phosphate

11:45 - 12:45 Session 1 - Sphingolipids in health and disease (II)

Chairs: **Lauren Ashley Cowart** (Richmond, US), **Philippe Guerre** (Toulouse, FR)

11:45 - 12:00 **Valentine Yolander** (*Virginia Commonwealth University, US*)

Sphingosine kinase 1 and sphingosine-1-phosphate receptor function in adipocyte beiging

12:00 - 12:15 **José Fernandez-Checa** (*University of Barcelona, ES*)

Dietary and genetic disruption of hepatic methionine metabolism induces acid sphingomyelinase expression to promote steatohepatitis

12:15 - 12:30 **David Montefusco** (*Virginia Commonwealth University, US*)

SPTLC3 and atypical sphingolipids as regulators of hepatic glucose production

12:30 - 12:45 **Mariana Nikolova-Karakashian** (*University of Kentucky, US*)

Role of neutral sphingomyelinase 2 in fatty liver disease and insulin resistance

12:45 - 14:20 *Lunch*

14:20 - 15:55 Session 1 - Sphingolipids in health and disease (III)

Chairs: **Thierry Levade** (Toulouse, FR), **Chiara Donati** (Florence, IT)

14:20 - 14:45 **Diego Centonze** (*IRCCS Neuromed and University of Rome Tor Vergata, IT*)

Inflammatory synaptopathy as a novel target of S1P receptor modulation in experimental multiple sclerosis

14:45 - 15:00 **Mirek Machala** (*Veterinary Research Institute, CZ*)

Sphingolipid and glycosphingolipid profiles are altered during benzo[a]pyrene-induced epithelial-mesenchymal transition of normal human bronchial epithelial cells, as well as in their exosomes

15:00 - 15:25 **Andrea Huwiler** (*University of Bern, CH*)

The role of Sphingosine kinase 2 in chronic kidney disease

15:25 - 15:40 **Christiane Mühle** (*University of Erlangen DE*)

COVID-19 and its clinical severity are associated with alterations of plasma sphingolipids and enzyme activities of sphingomyelinase and ceramidase

15:40 - 15:55 **Trushnal Waghmare** (*University of Würzburg, DE*)

Role of neutral sphingomyelinase 2 and sialophorin as virus effector in T cells

15:55 - 17:20 Session 2 - Sphingolipids and malignant diseases (I)

Chairs: **Christiane Mühle** (Erlangen, DE), **Michele Dei Cas** (Milan, IT)

15:55 - 16:20 **Marco Presta** (*University of Brescia, IT*)

Beta-Galactosylceramidase: more than a psychosine scavenger

16:20 - 16:35 **Hui-Ming Lin** (*Garvan Institute of Medical Research, AU*)

Modulation of circulating sphingolipids in metastatic prostate cancer by simvastatin

16:35 - 16:50 *Coffee break*

16:50 - 17:05 **Giovanna Chiorino** (*Fondazione Edo ed Elvo Tempia, Biella, IT*)

Targeted sphingolipid analysis to improve prostate cancer detection

17:05 - 17:20 **Lorry Carrié** (*Université Toulouse III Paul-Sabatier, FR*)

Tumor cell-derived sphingosine 1-phosphate promotes loss of adhesion to keratinocytes and contributes to primary melanoma invasion

17:20 - 18:35 Session 2 – Sphingolipids and malignant diseases (II)

Chairs: **Gehrid Van Echten-Deckert** (Bonn, DE), **Angel Gaudio** (Madrid, ES)

17:20 - 17:35 **Carine Dufau** (*Université de Toulouse III Paul-Sabatier, FR*)

Role of ceramide metabolism in TNF-induced melanoma dedifferentiation

17:35 - 17:50 **Davide Capoferri** (*University of Brescia, IT*)

Beta-galactosylceramidase modulates the pro-oncogenic features of melanoma cells

17:50 - 18:05 **Chiara Raggi** (*University of Florence, IT*)

Irregular glycosphingolipid patterns in human cholangiocarcinoma stem-like subsets

18:05 - 18:20 **Francois Paris** (*Université de Nantes, FR*)

Release of ceramide by irradiated endothelial cell inhibits tumor cell proliferation after repression of Lamin B1 expression

18:20 - 18:35 **Sumaiya Afsar** (*University of Bonn, DE*)

SGPL1 ablation affects glucose metabolism in a way that abets oncogenesis

20:30 - *Dinner*

**Friday, Sept. 9, 2022 - Neuromed Research Center, Pozzilli**

9:00 - 11:05 Session 3 - Sphingolipid biology: mechanisms and tools (I)

Chairs: **Francois Paris** (Nantes, FR), **Andrea Huwiler** (Bern, CH)

09:00 - 09:25 **Antonella De Matteis** (*Telethon Institute of Genetics and Medicine, IT*)

ER-Golgi contact sites: regulators and targets of sphingolipids

- 09:25 - 09:40 **Marco Trinchera** (*University of Insubria, IT*)  
Bi-allelic inactivating variants of Lactosylceramide Synthase beta4GALT5 responsible for a novel congenital disorder of glycosylation involving glycosphingolipids
- 09:40 - 09:55 **Roxana Manaila** (*University of Bern, CH*)  
Sphk1 and sphk2 differentially regulate erythropoietin synthesis in mouse renal interstitial fibroblast-like cells
- 09:55 - 10:10 **Tanes Imamura de Lima** (*École Polytechnique Fédérale de Lausanne, CH*)  
Inhibition of *de novo* ceramide synthesis restores mitochondrial and protein homeostasis in muscle aging
- 10:10 - 10:25 **Caterina Giovagnoni** (*Maastricht University, NL*)  
Function of ceramide transfer protein for biogenesis and sphingolipid composition of extracellular vesicles
- 10:25 - 10:50 **Webster Santos** (*Virginia Tech, US*)  
Discovery, structure-activity relationship studies, and *in vivo* activity of sphingosine-1-phosphate transporter spns2
- 10:50 - 11:05 **Michele Dei Cas** (*University of Milan, IT*)  
Mass spectrometry measurement of sphingolipids delta-4-desaturase 1 activity and its application in related genetic diseases
- 11:05 - 11:20 *Coffee break*
- 11:20 - 13:00 Session 3 – *Sphingolipid biology: mechanisms and tools (II)*
- Chairs: **Dagmar Meyer zu-Heringdorf** (Frankfurt, DE), **Massimo Aureli** (Milan, IT)
- 11:20 - 11:35 **Philippe Guerre** (*ENVT Toulouse, FR*)  
Targeted sphingolipid analysis suggests mechanisms of fumonisin toxicity in chickens differs depending on the organ studied
- 11:35 - 12:00 **Tsaffrir Zor** (*Tel Aviv University, IL*)  
Short fatty acid sulfatides are endogenous ligands for TLR4/MD-2
- 12:00 - 12:15 **Tristan Martineau** (*Université de Sherbrooke, CA*)  
Urine screening test using a tandem mass spectrometry approach for early detection of lysosphingolipidoses
- 12:15 - 12:30 **Camillo Morano** (*University of Milan, IT*)  
A new “how to” in sphingolipidomics as a tool for a common sample handling and analysis
- 12:30 - 12:45 **Essa M. Saied** (*Humboldt-Universität zu Berlin, DE*)  
Discovery of the first selective and drug-like small molecule inhibitors of Alkaline Ceramidase 3

12:45 - 13:00 **Stefano Piotto** (*University of Salerno, IT*)  
Mechanistic study of the role of hydroxylation of sphingolipids

13:00 - 15:00 *Lunch and Poster viewing*

15:00 - 20:30 *Social time: Picture time and visit to Agnone*

20:30 - *Dinner*

## **Saturday, Sept. 10, 2022 – Neuromed Research Center, Pozzilli**

09:00 – 13:00 ISN Symposium: *Sphingolipids and brain: pathophysiology and therapeutics*

Chairs: **Alba Di Pardo, Vittorio Maglione** (Pozzilli, IT)

09:00 - 09:10 Introduction by Chairs

09:10 - 09:35 **Simonetta Sipione** (*University of Alberta, CA*)  
Gangliosides: old dogs with new tricks

09:35 - 10:00 **Massimo Aureli** (*University of Milan, IT*)  
Role of the lysosomal impairment in the onset of neuronal degeneration

10:00 - 10:25 **Wiebke Herzog** (*University of Erlangen, DE*)  
Regulation of blood-brain barrier properties by Sphingosine-1-phosphate signaling

10:25 - 10:50 **Eric Camerer** (*Université Paris Cité, FR*)  
Tight control of S1P signaling at the blood-brain barrier

10:50 - 11:05 *Coffee break*

11:05 - 11:30 **Pilar Martinez-Martinez** (*University of Maastricht, NL*)  
Title

11:30 - 11:45 **Angel Gaudioso** (*Centro Biología Molecular Severo Ochoa (CSIC-UAM), Madrid, ES*)  
Sphingomyelin 16:0 is a therapeutic target for brain pathology in Acid Sphingomyelinase deficiency

11:45 - 12:00 **Liubov Kalinichenko** (*University of Erlangen, DE*)  
Ceramide system contributes to learning and memory

12:00 - 12:15 **Daan Van Kruining** (*Maastricht University, NL*)  
FTY720 decreases ceramide levels in the brain and prevents memory impairments in a mouse model of familial Alzheimer's disease expressing ApoE4

12:15 - 12:30 **Sara Grassi** (*University of Milan, IT*)  
Sphingolipid-dependent membrane organization and signaling orchestrating myelin repair

12:30 - 12:45 **Giuseppe Pepe** (*IRCCS Neuromed, IT*)

Treatment with THI, an inhibitor of Sphingosine-1-Phosphate Lyase (SGPL1), modulates glycosphingolipid metabolism and results therapeutically effective in a mouse model of Huntington's disease

12:45 - 13:00 **Shah Alam** (*University of Bonn, DE*)  
S1P-lyase deficiency in the brain increases glucose breakdown evoking a P2Y1 Receptor-dependent astrogliosis

13:00 - 14:00 *Lunch*

14:00 - 15:00 Poster flash presentations (2 slides in 2 min)

Chair: **Riccardo Ghidoni** (Milan, IT)

**Gloria Grelle M.R.S.** (*Universidade Federal do Rio de Janeiro, BR*)  
LC-HRMS-based sphingolipid profiling of renal cells exposed to ischemia

**Dragana Fabris** (*University of Zagreb, HR*)  
Ceramides and sphingomyelins in ischemic stroke: a preliminary study

**Gloria Grelle M.R.S.** (*Universidade Federal do Rio de Janeiro, BR*)  
Ceramide-1-Phosphate as a potential regulator of the second sodium pump from kidney proximal tubules through different protein kinase pathways

**Luisa Volk** (*Universität of Frankfurt am Main, DE*)  
Functional consequences of sphingosine-1-phosphate lyase deletion in mouse hepatocytes

**Samuela Cataldi** (*University of Perugia, IT*)  
Egg yolk sphingomelin facilitates the maturation of hippocampal embryonic cells

**Federico Fiorani** (*University of Perugia, IT*)  
Exosomes amplify the signal of vitamin D3-induced embryonic hippocampal cell differentiation via ceramide

**Tommaso Beccari** (*University of Perugia, IT*)  
Neuroprotective effect of Eicosapentaenoic acid and Docosahexaenoic acid on SH-SY5Y cells

**Michele Dei Cas** (*University of Milan, IT*)  
Altered sphingolipid content in extracellular vesicles derived from primary intrahepatic cholangiocarcinoma cells

**Marzia Corli** (*University of Brescia, IT*)  
Unravelling the role of  $\beta$ -galactosylceramidase in melanoma neovascularization

**Ondrej Kovac** (*Veterinary Research Institute, CZ*)  
Possible role of the aryl hydrocarbon receptor in modulation of sphingolipid and glycosphingolipid metabolism in colon cancer cell models

**Josè Luis Abad** (*CSIC Barcelona, ES*)

Fluorescent ceramide-labeled probes with bodipy and coumarin-based coupy fluorophores

**Alix Pierron** (*ENVT Toulouse, FR*)

Targeted sphingolipid analysis reveals differences on wild type mouse and SOCS2 deficient mouse in peritonitis zymosan induced

**Christiane Mühle** (*University Erlangen, DE*)

Interplay of sphingolipids and  $\alpha$ -synuclein in Parkinson's disease and major depressive disorder

**Lucia Sessa** (*University of Salerno, IT*)

The influence of hydroxylation in the mechanism of action of  $\alpha$ -hydroxy-ceramide

**Bisera Stepanovska Tanturovska** (*University of Bern, CH*)

S1P stimulates erythropoietin production in mouse renal interstitial fibroblasts by S1P<sub>1</sub> and S1P<sub>3</sub> receptor activation and HIF-2 $\alpha$  stabilization

**Karoli Liliom** (*Semmelweis University, HU*)

Apolipoprotein D is a lipocalin carrier protein for lysophospholipid mediators

**Gemma Fabrias** (*CSIC Barcelona, ES*)

A novel fluorogenic probe to monitor the activity of acid ceramidase, fatty acid amide hydrolase and N-acyl ethanol-amine acid amidase

**Salvatore Fiorinello** (*National Research Council of Italy, Napoli, IT*)

Pharmacological treatment with sphingosine analogues to correct glycosphingolipid metabolic derangements in Rett syndrome models

**Christiane Mühle** (*University Erlangen, DE*)

Sphingomyelin synthases in depression and antidepressant treatment

**Fabrizia Noro** (*IRCCS Neuromed, IT*)

The effect of gangliosides on human platelet aggregation and platelet-leukocyte mixed aggregate formation

15:00 - 15:15 *Coffee break*

15:15 – 16:30 *Laboratories guided tour  
Neuromed Research Center*

16:30 - 18:00 Session 4 – Sphingolipids and brain 2.0

Chair: **Simonetta Sipione** (Edmonton, CA), **Eric Camerer** (Paris, FR)

16:30 - 16:55 **Christian Müller** (*University of Erlangen, DE*)

Neutral Sphingomyelinase in the control of alcohol abuse and its comorbidities

16:55 - 17:10 **Floriana Della Ragione** (*National Research Council of Italy, Napoli, IT*)

Imbalance of glycosphingolipid metabolism in Rett Syndrome as a novel potential therapeutic target

17:10 - 17:35 **Thorsten Hornemann** (*USZ, Zürich, CH*)  
Motor neuron disease or sensory neuropathy? L-Serine as a modulating factor

17:35 - 18:00 **Alessandra D'Azzo** (*St. Jude Children's Research Hospital, Memphis, US*)  
Role of membrane contact sites in the neuropathogenesis of GM1-Gangliosidosis

20:30 - *Gala dinner and live entertainment*

## **Sunday, Sept. 11, 2022 – Grand Hotel Europa, Isernia**

09:00 - 10:00 *General Meeting of the SLC*

10:00 – 10:15 *Closing remarks*

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