## Molecular mechanisms of muscle wasting during aging and disease

Congressi Stefano Franscini, Monte Verità, Ascona, Switzerland Sunday September 23, 2018 – Friday September 28, 1018

## **Program**

| Sunday        | <b>September 23, 2018</b> |  |
|---------------|---------------------------|--|
|               |                           |  |
| 16.00 - 18.00 | Registration              |  |
| 18.00 - 19.00 | Reception                 |  |

## **Keynote Lecture**

19.00 Pura Muñoz-Cànoves

Muscle stem cell dysfunction with aging: cause and consequences

Followed by an "icebreaker"

| Monday                              | September 24, 201                         | 18 - Morning   |
|-------------------------------------|---|--|
| 08.30 - 08.50                       | Welcome address from CSF and Monte Verità |  |
|                                     | <b>Session</b><br>Chair                   | Repair of skeletal muscle<br>Simone Spuler   |
| Speakers                            | <del></del>                               |  |
| 08.50 - 09.15<br>09.15 - 09.40      | Michael Rudnicki<br>Florian Bentzinger    | Molecular regulation of muscle stem cell function Extrinsic regulation of muscle stem cell specification |
| Short talks                         | Horidii Bentember                         | Extinisie regulation of musele stem cen specification  |
|                                     |   |  |
| 09.40 - 09.55                       | Remi Mounier                              | AMPK $\alpha$ 2 as a regulator of muscle cell differentiati<br>and skeletal muscle homeostasis           |
| 09.55 - 10.10                       | Fabien Le Grand                           | High-dimensional single-cell cartography reveals novel skeletal muscle resident cell populations         |
| 10.10 - 10.40                       | Coffee Break                              |  |
| Speakers                            |   |  |
| 10.40 - 11.05                       | Andrew Brack                              | Lineage tracing reveals a subset of reserve muscle stem cells capable of clonal expansion under stres    |
| 11.05 - 11.30                       | Carmen Birchmeier                         | Oscillatory expression of Hes1 and MyoD allow activated satellite cells to maintain an                   |
|                                     |   |  |
| Short talks                         |   | undifferentiated state   |
| <b>Short talks</b><br>11.30 - 11.45 | Peter Vrtacnik                            | Somatic mutagenesis in satellite cells reduces the   |
|                                     | Peter Vrtacnik<br>Davide Gabellini        |  |

| Monday        | September 24, 2018 - Afternoon |  |
|---------------|--------------------------------|--|
| Speakers      | <b>Session</b><br><u>Chair</u> | Nerve-muscle connection David Glass  |
| 13.30 - 13.55 | Lin Mei                        | Sarcoglycan alpha mitigation of neuromuscular  |
| 13.55 - 14.20 | Markus Ruegg                   | junction decline in aged mice by stabilizing Lrp4 The role of mTOR in NMJ maintenance                                      |
| Short talks   |                                |  |
| 14.20 - 14.35 | Laure Strochlic                | Distinct branches of Wnt signaling regulate mammalian neuromuscular connectivity   |
| 14.35 - 14.50 | Daisy Proietti                 | Role of muscle interstitial cells in neurogenic muscular atrophy   |
| 14.50 - 15.20 | Coffee Break                   |  |
| Speakers      |                                |  |
| 15.20 - 15.45 | David Beeson                   | Mechanisms and treatment with β2-adrenergic receptor agonists for hereditary disorders of neuromuscular synapse            |
| 15.45 - 16.10 | Charlotte Sumner               | Failed perinatal motor axon sorting is associated with impaired myofiber growth in spinal muscular atrophy (SMA)           |
| Short talks   |                                | atrophy (Sivin)  |
| 16.10 - 16.25 | Anna Rostedt Punga             | miRNA profile in affected skeletal muscles of MuSK+<br>EAMG mice reveal novel intracellular targets for<br>muscle weakness |
| 16.25 - 16.40 | Aaron Russel                   | MicroRNA inhibition to improve skeletal muscle health in amyotrophic lateral sclerosis (ALS)                               |
| 17.00 - 19.00 | Poster Session (with o         | drinks and snacks) – <b>Posters 1 to 31</b>  |
| 19.15 - 20.45 | Dinner                         |  |

| Tuesday                        | September 25, 2018 - Morning      |   |
|--------------------------------|-----------------------------------|---|
| Speakers                       | <b>Session</b> <u>Chair</u>       | Mechanisms of muscle plasticity Markus Ruegg  |
| 08.50 - 09.15<br>09.15 - 09.40 | Nathan LeBrasseur<br>Marco Sandri | The role of myostatin in the control of muscle size A novel player in autophagy regulation and muscle wasting   |
| Short talks                    |                                   |   |
| 09.40 - 09.55                  | Hyeon-Ju Jeong                    | Skeletal muscle-specific PRMT1 deletion causes<br>muscle atrophy via deregulation of PRMT6/FoxO3a<br>axis   |
| 09.55 - 10.10                  | Gonzalo Blanco                    | Transcriptional up-regulation of Chaperone Assisted Selective Autophagy factors in animal models of KY-deficient hereditary myopathy                            |
| 10.10 - 10.40                  | Coffee Break                      |   |
| Speakers                       |                                   |   |
| 10.40 - 11.05                  | David Glass                       | Signaling required for muscle maintenance in the setting of aging   |
| 11.05 - 11.30                  | Tea Shavlakadze                   | Age-related pathway signatures — relevance for treating aging disorders   |
| Short talks                    |                                   |   |
| 11.30 - 11.45                  | Paul Gregorevic                   | A metabolic role for YAP in the regulation of skeletal muscle attributes  |
| 11.45 - 12.00                  | Shuichi Watanabe                  | Skeletal muscle specific cofactor Vgll2/3 dependent function of Teads (Tead1-4) is indispensable for mTOR signal regulation and muscle fiber type specification |
| 12.00 - 14.00                  | Lunch Break                       |   |
|                                | Free Afternoon                    |   |
| 19.15 - 20.45                  | Dinner                            |   |

| Wednesday September 26, 2018 - Morning |                                | 18 - Morning  |
|--|--------------------------------|---|
| Speakers                               | <b>Session</b><br><u>Chair</u> | Muscle and metabolism<br>Matt Kaeberlein  |
| opeaners -                             |                                |   |
| 08.50 - 09.15<br>09.15 - 09.40         | Michael Hall<br>Mara Fornaro   | mTOR signaling in growth and metabolism  IGF signaling and its effect on muscle metabolism  |
| Short talks                            |                                |   |
| 09.40 - 09.55                          | Francesca Solagna              | Treatment with soluble activin type IIB in a mouse model of polycystic kidney disease improves musc mass and strength and reduces the progression of kidney disease |
| 09.55 - 10.10                          | Anurag Singh                   | Translating Urolithin A benefits on muscle mitochondria into humans   |
| 10.10 - 10.40                          | Coffee Break                   |   |
| Speakers                               |                                |   |
| 10.40 - 11.05                          | Simone Spuler                  | Human muscle-derived CLEC14A-positive cells wit regenerative potential independent of PAX7  |
| 11.05 - 11.30                          | Rafael de Cabo                 | Caloric restriction and its effect on skeletal muscle   |
| Short talks                            |                                |   |
| 11.30 - 11.45<br>11.45 - 12.00         | Daniel Ham<br>Aaron Hinken     | Rapamycin attenuates sarcopenia in mice<br>NOPE is a novel modulator of TGF-6 family memb<br>activity and body composition  |
|  |                                | detivity and body composition   |

| Wednesday     | September 26, 2018 - Afternoon |   |
|---------------|--------------------------------|---|
|               | <b>Session</b><br><u>Chair</u> | <b>Epigenetics and non-coding RNA</b> Alessandra Sacco  |
| Speakers      |                                |   |
| 13.30 - 13.55 | •                              | Micropeptides encoded by IncRNA in muscle   |
| 13.55 - 14.20 | Thomas Braun                   | Posttranscriptional processes regulating metabolic switch during differentiation of skel muscle stem cells                                    |
| Short talks   |                                |   |
| 14.20 – 14.35 | Alyson Fiorillo                | miR-146a inhibits dystrophin production and dire<br>activates innate immune signaling to promote<br>inflammation in multiple muscle disorders |
| 14.35 - 14.50 | Jozef Dulak                    | Silencing of miR-378a attenuates dystrophic phenotype in mdx mice   |
| 14.50 - 15.20 | Coffee Break                   |   |
| Speakers      |                                |   |
| 15.20 - 15.45 | Vittorio Sartorelli            | A muscle-specific enhancer RNA mediates Cohesi<br>recruitment and regulates transcription in trans  |
| 15.45 - 16.10 | Robert Krauss                  | Regulation of satellite cell quiescence and activate by niche adhesive junctions  |
| Short talks   |                                |   |
| 16.10 - 16.25 | Barbara Franke                 | Mechanosensing in the kinase region of titin  |
| 16.25 - 16.40 | Giorgia Careccia               | HMGB1 as novel target in Duchenne Muscular<br>Dystrophy   |
| 17.00 – 19.00 | Poster Session (with d         | rinks and snacks) – <b>Posters 32 to 62</b>   |
|               |                                |   |

| Thursday      | September 27, 2018 - Morning   |   |
|---------------|--------------------------------|---|
|               | <b>Session</b><br><u>Chair</u> | <b>Mechanisms of aging</b> Michael Rudnicki   |
| Speakers      |                                |   |
| 08.50 - 09.15 | Peter de Keizer                | Targeted apoptosis of senescent cells against aging and cancer  |
| 09.15 - 09.40 | James Kirkland                 | Age-related dysfunction, cellular senescence, and senolytic agents: The path toward translation                         |
| Short talks   |                                |   |
| 09.40 - 09.55 | Akiyoshi Uezumi                | Roles of interstitial mesenchymal progenitors in the maintenance of skeletal muscle and its implications for sarcopenia |
| 09.55 - 10.10 | Thomas Vogler                  | Amyloid-like TDP-43 myo-granules associate with sarcomeric RNAs during skeletal muscle formation                        |
| 10.10 - 10.40 | Coffee Break                   |   |
| Speakers      |                                |   |
| 10.45 - 11.10 | Matt Kaeberlein                | Targeting mTORC1 signaling to promote healthy longevity   |
| 11.10 - 11.35 | Luigi Ferrucci                 | The aging of skeletal muscle  |
| Short talks   |                                |   |
| 11.35 - 11.50 | Marco De Cecco                 | Endogenous Retroelements become de-repressed and active in skeletal muscle of aging mice                                |
| 12.00 - 13.30 | Lunch Break                    |   |
|               |                                |   |

| Thursday                       | September 27, 2018 - Afternoon      |   |
|--------------------------------|-------------------------------------|---|
| Smarkana                       | <b>Session</b> <u>Chair</u>         | Muscle wasting diseases<br>Rhonda Bassel-Duby   |
| Speakers                       |                                     |   |
| 13.30 - 13.55<br>13.55 - 14.20 | Alessandra Sacco<br>Denis Guttridge | Stem cell-based strategies to treat muscle wasting<br>Insights in rhabdomyosarcoma pathogenesis                         |
| Short talks                    |                                     |   |
| 14.20 - 14.35                  | Domagoj Cikes                       | Changes of mitochondrial mechanics trigger metabolic crisis and cachexia  |
| 14.35 - 14.50                  | Foteini Mourkioti                   | A robust Pax7EGFP mouse that enables the visualization of muscle stem cell dynamics                                     |
| 14.50 – 15.05                  | Brad Olwin                          | Cellular responses during regeneration of skeletal muscle   |
| 15.10 - 15.30                  | Coffee Break                        |   |
| Speakers                       |                                     |   |
| 15.30 - 15.55                  | Kevin Campbell                      | Structural studies of dystroglycan extracellular matrix receptor function in skeletal muscle                            |
| 15.55 - 16.20                  | Jeff Chamberlain                    | Augmented muscle function via systemic gene delivery with AAV   |
| Short talks                    |                                     |   |
| 16.20 - 16.35                  | Stefano Previtali                   | Jab1 in the pathogenesis of Merosin deficient congenital muscular dystrophy (MDC1A)                                     |
| 16.35 - 16.55                  | Gordon Lynch                        | Therapeutic potential of slow muscle programming by low-frequency stimulation for muscle wasting and muscular dystrophy |
| 16.55 - 17.10                  | Mattia Quattrocelli                 | Weekly and daily glucocorticoid steroids divergently remodel muscle metabolism and function                             |
| 17.10 -                        | CSF Award Ceremon                   | У   |
| 19.15 -                        | Gala Dinner                         |   |
| Friday                         | September 28, 202                   | 18  |
|                                | Departure                           |   |
|                                |                                     |   |

## **Posters**

List of poster titles and presenting author. Refer to the number of your poster to locate the board where your poster can be displayed. The list is sorted alphabetically by presenting author's last name.

For the full authors list of each poster, check the corresponding abstract.

Authors of **posters nr. 1 to 31** please attend your poster in Session 1 (Monday). Authors of **posters nr. 32 to 62** please attend your poster in Session 2 (Wednesday).

- 1. Redox remodeling causes "leakiness" in Ryanodine Receptor (RyR1): New Insight into the molecular etiology of chronic hypobaric hypoxia induced skeletal muscle atrophy Akanksha Agrawal
- **2.** Activating the MasR/Ang1-7 Pathway Reduces Muscle Atrophy and Function Loss Following Denervation

Hind M. Albadrani

- **3.** Identification of a novel TFEB-exercise dependent gene Andrea Armani
- **4.** Implantation of MCK-PGC-1α myogenic progenitors after muscle damage results in enhanced oxidative phenotype
  Marc Beltrà
- **5.** Identification of novel molecular targets to manipulate satellite cell function Anna Benedetti
- **6.** The role of Raptor/mTORC1 in adult skeletal muscle Bert Blaauw
- **7.** Sarcopenia chronicles: dynamics of gene expression during aging of rat skeletal muscles Anastasiya Börsch
- 8. Vivo-morpholinos targeting dynamin 2 ameliorate pathological signs of myotubular myopathy

Caroline Bogni

**9.** Rev-erb- $\alpha$  exacerbates endoplasmic reticulum stress-induced apoptosis in mouse skeletal muscle

Alexis Boulinguiez

**10.** Supplementation of Schisandra Fructus ethanol extract increased insulin sensitivity and muscle mass in aged mice

Hojung Choi

11. Investigating anti-cachectic role of Unacylated Ghrelin

Sara Clerici

12. Genes differentially expressed during reversion of androgend dependent skeletal muscle atrophy

Priscila O. Coelho

13. Improving muscle wasting in cancer cachexia

Paola Costelli

**14.** IGFN1 interacts with the actin nucleator COBL and is required for myoblast fusion Tobias Cracknell

- 15. Opposing effects of 25-hydroxy- and  $1\alpha$ ,25-dihydroxy-vitamin  $D_3$  on pro-cachectic cytokine- and cancer conditioned medium-induced atrophy in C2C12 myotubes Marilisa De Feudis
- **16.** Role of mito $K_{ATP}$  in skeletal muscle

Giulia Di Marco

- **17.** CaVβ1: The missing link from voltage sensing to muscle mass homeostasis Sestina Falcone
- **18.** Physiopathological characterization of the role of MCUb in skeletal muscle regeneratin Simona Feno
- **19.** Acylated ghrelin, in contrast to unacylated ghrelin, fails to directly counteract muscle wasting *in vivo*

Michele Ferrara

**20.** Exercise prevents cancer-induced fatigue by affecting skeletal muscle, systemic metabolism and anemia

Regula Furrer

- **21.** Characterization of the D2/mdx mouse as a clinically relevant model of Duchenne Muscular Dystrophy and demonstration that treatment with mRK35, a murine anti-myostatin antibody, induces functional improvements in D2/mdx and aged Bl10/mdx mice Sweta Girgenrath
- **22.** Modulating bone morphogenetic signalling in cancer cachexia Adam Hagg
- **23.** Effects of metformin and N-acetyl-L-cysteine on congenital muscular dystrophy type 1A disease progression in mice

Vahid M. Harandi

- 24. Aging associated reduction in skeletal muscle stem cell proliferation rate is accompanied by reduced focal adhesion formation, and increased YAP signaling Mohammad Haroon
- **25.** Collagen XIII secures development and maturation of the neuromuscular synapse Anne Heikkinen
- **26.** TGF- $\beta$  signalling differentially affects myogenic and fibrotic gene expression in muscle stem cells and myotubes

Michèle M. G. Hillege

**27.** Effects of aberrant accumulation of MFG-E8 at neuromuscular junction on skeletal muscle aging

Madoka Ikemoto-Uezumi

- **28.** The MuSK-BMP pathway is required for biphasic BMP-regulated myogenesis Diego Jaime
- **29.** The role of Cdon in the generation of embryonic stem cell-derived motor neuron Jong Sun Kang
- **30.** Time of day and exercise: a new role for muscle contraction as time cue Denise Kemler

**31.** Expression of a slow myosin motor ( $\beta$ -MyHC) drives physiology and metabolism in skeletal muscle

Genevieve C. Kerr

**32.** Ex2 suppresses muscle wasting through PDK1/Akt activation augmenting muscle mass increase

Hye Been Kim

**33.** Acute sleep deprivation impairs skeletal muscle protein synthesis

Severine Lamon

**34.** Wnt inhibitory factor-1 biding to a promyogenic receptor Cdo is critical for reciprocal modulation of Wnt signal and promyogenic kinases

Sang-Jin Lee

**35.** GLP-1 analogue, exendin-4, ameliorates dexamethasone-induced muscle atrophy through regulating atrophy-related gene expression

Jong Han Lee

**36** Inducing mitophagy in muscle stem cells with Urolithin A restores muscle function in muscular dystrophy

Peiling Luan

**37.** The interference with IL-6 trans-signaling modulates secondary mechanisms of dystrophic muscle

Carmen Miano

**38.** Pax7/MyoD high-content screening of myogenic fate in human primary myoblasts Joris Michaud

**39.** Role of the AKT/mTOR/ FoxO pathway in muscle protein homeostasis

Giulia Milan

**40.** MicroRNA regulation of the Ndrg2 gene in skeletal muscle cells

Bilal A. Mir

**41.** Development of screening system for identification of novel factors influencing muscle fiber types

Norio Motohashi

42. SIRT1 inhibition restores PABPN1-dependent muscle wasting

Cyriel S. Olie

**43.** Group I Paks support muscle regeneration and counteract cancer-associated muscle atrophy

Rosanna Piccirillo

44. Data-driven single-myofibertyping reveals age-associated muscle-specific patterns

Vered Raz

45. Role of ghrelin peptides in the onset of sarcopenia

Simone Reano

**46.** HDAC4 mediates the responses to multiple stimuli in skeletal muscle

Alessandra Renzini

47. FGF21 controls mitophagy and muscle mass

Vanina Romanello

**48.** HIF prolyl hydroxylase inhibition protects skeletal muscle from eccentric contraction-induced injury

Alan J. Russell

**49.** Muscle wasting and recovery in a Drosophila gut tumor model

Pedro Saavedra

**50.** Manipulating BMP pathway to prevent NMJ's dismantling and denervation in Cancer Cachexia

Roberta Sartori

**51.** Platelet releasate as a key driver for skeletal myogenesis

**David Scully** 

**52.** *C.elegans* as a model to decipher muscle aging mechanisms

Florence Solari

**53.** The TGF- $\beta$ -signalling inhibitor, follistatin concomitantly increases muscle mass and insulin sensitivity in skeletal muscle

Lykke Sylow

**54.** Generation of a new mouse to model pancreatic cancer-induced muscle wasting and cachexia

Erin E. Talbert

55. Stem Cell regulation in muscular dystrophy

Elisia D. Tichy

- **56.** Identification of age-related modulators of protein synthesis in the muscle Lionel Tintignac
- **57.** Nature and role of interstitial non myogenic cells in human fibrotic muscles Capucine Trollet
- **58.** Disuse atrophy is characterized by marked decrease in mitochondrial content and function

Rick B. Vega

- **59.** CD36 deficiency restores impaired satellite cell proliferation in response to high-fat diet Sandrine Verpoorten
- **60.** Muscular improvement to physical exercise in Myasthenia Gravis patients Elisabeth Westerberg
- **61.** Iron metabolism regulates cancer related skeletal muscle wasting Elisabeth Wyart
- **62.** Expression of Swedish mutant APP in skeletal muscles results in muscle wasting-like deficits

Wen-Cheng Xiong