



SEPTEMBER 20-25, 2019 • VANCOUVER, CANADA



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RESTLESS LEGS SYNDROME TRACK

FULL TRACK RUNS SATURDAY - WEDNESDAY

The RLS Track at World Sleep 2019 will include some of the biggest names in the field offering cutting-edge science and information.

23 HOURS OF RLS content have been added to the final Scientific Program. Register today to learn more about RLS research and treatment options.

REGISTRATION OPTIONS:

RLS Course (C05)	\$125
IRLSSG Study Group Annual Meeting & Course	\$165
Both Courses + Congress Registration (Member).....	\$685

FOR MORE INFORMATION & ALL PRICING VISIT
worldsleepcongress.com/register

RLS TRACK OVERVIEW

TYPE	DAY	TITLE	HOURS
COURSE	Saturday	C05: Recent advances in RLS treatment	8:00am - 12:00pm
COURSE	Sunday	International Restless Legs Syndrome Study Group annual meeting & course	8:00am - 9:00pm
SYMPOSIUM	Monday	Pathophysiological insights from animal models of restless legs syndrome	3:00pm - 4:30pm
SYMPOSIUM	Tuesday	Sensory-motor network of the restless legs syndrome (RLS): Electrophysiology and imaging	10:45am - 12:15pm
KEYNOTE	Tuesday	K08: Restless legs syndrome/periodic limb movements of sleep: New insights into neurobiology and treatment	2:00pm - 2:45pm
SYMPOSIUM	Wednesday	Brain iron as a central factor in the pathophysiology of RLS: Emerging evaluation methods and therapeutic opportunities	3:00pm - 4:30pm



 **COURSE | SATURDAY, SEPTEMBER 21, 2019 | 8:00AM - 12:00PM | ROOM 120**

■ **C05 Recent advances in RLS treatment**

Saturday, September 21, 2019 | 8:00am - 12:00 pm | Room 120

Chairs Richard Allen (United States); Diego García-Borreguero (Spain)

 **ADDITIONAL REGISTRATION REQUIRED**

Summary

This course will start with a quick summary of the current practice of RLS and then move to new concepts in diagnosis and management of RLS, with a special focus on iron therapy, opioids, $\alpha 2\delta$ agents, glutamate modulation, augmentation, long term outcomes including impulse control disorders, new guidelines, and update on pathophysiology including insights from genetics and animal models.

8:00am – 8:05am

Introduction

OPIOIDS

8:05am – 8:35am

Biological differences of opioids: Low abuse potential of methadone

Sergi Ferre (United States)

8:35am – 9:15am

USA Clinical guidelines/experience with opioid use in RLS

Christopher Earley (United States)

9:15am – 9:55am

European guidelines/experience with opioid use in RLS including oxycodone/naloxone

Birgit Högl (Austria)

IRON

9:55am – 10:40am

IV iron: choices, advantages and limitations

Richard Allen (United States)

EXPERIMENTAL TREATMENTS: RATIONALE AND CLINICAL EXPERIENCE

10:40am – 11:20am

Adenosine

Diego García-Borreguero (Spain)

11:20am – 11:40am

Cannabinoid/Cannabis

Imad Ghorayeb (France)

11:40am – 12:00pm

Discussion – Questions to speakers



**SCIENTIFIC PROGRAM
NOW AVAILABLE**

To view the Scientific Program for World Sleep 2019, scan the code.





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 COURSE | SUNDAY, SEPTEMBER 22, 2019 | 9:00AM - 6:00PM | ROOM 220



**INTERNATIONAL
RESTLESS LEGS SYNDROME
STUDY GROUP**

INTERNATIONAL RESTLESS LEGS SYNDROME STUDY GROUP ANNUAL MEETING & COURSE

Summary

The International Restless Legs Syndrome Study Group (IRLSSG) will offer a full-day course on Sunday, September 22, 2019. Attendance is open to any sleep professional who is interested in RLS. A business meeting will be held after the course, which is only open to IRLSSG members. Registration includes the sessions, lunch and networking dinner.

Program Committee

Denise Sharon (United States); Federica Provini (Italy); Garima Shukla (Canada); Rochelle Zak (United States); Cornelius Bachman (Germany)

**COST \$165 | FREE TO IRLSSG MEMBERS
BECOME A MEMBER AT WWW.IRLSSG.ORG**

**REGISTRATION INCLUDES
SESSIONS, LUNCH & NETWORKING DINNER**

Presentation

8:00am – 08:30am

Networking and Coffee

*Denise Sharon (United States); Federica Provini (Italy);
Garima Shukla (Canada); Rochelle Zak (United States)
Cornelius Bachman (Germany)*

8:30am – 8:45am

Welcome to Vancouver & Introductions

Denise Sharon (United States); Allan O'Bryan (United States)

ANIMAL MODELS TASK FORCE

Mauro Manconi (Switzerland); Diego García-Borreguero (Spain)

8:45am – 9:00am

Behavioral Animal Models: When phenotype matters and objective markers are missing

Jerome Siegel (United States)

9:00am – 9:15am

Critical review of outcome measures of the past models: Rationale and need of consensus

Mauro Manconi (Switzerland)

9:15am – 09:30am

Expert Consensus Guideline for an animal model of RLS: How to reach a consensus on outcome measures in animal models: Methods and preliminary results

Aaro Salminen (Germany)

9:30am – 09:45am

Update on RLS animal models and iron

Richard Allen (United States)

9:45am – 10:00am

State of the research on animal model KO for BTBD9

Yuqing Li (United States)

10:00am – 10:15am

Animals models task force summary and update

Mauro Manconi (Switzerland)

10:15am – 10:30am

Coffee break

10:30am – 10:50am

RLS: Leg movements identify arousal

Richard Allen (United States)



INTERNATIONAL RESTLESS LEGS SYNDROME STUDY GROUP ANNUAL MEETING & COURSE

10:50am – 11:10am

Update on blood pressure and endothelial dysfunction in RLS

Yves Dauvilliers (France)

11:10am – 11:30am

New MRI findings in RLS

Ambra Stefani (Austria)

11:30am – 11:45am

A proteomic and system biology approach reveal novel biomarker signatures for RLS

Raffaele Ferri (Italy)

11:45am – 12:00pm

The lifespan course of short-interval, periodic and isolated leg movements during sleep

Raffaele Ferri (Italy)

12:00pm-12:15pm

Iron treatment

Richard Allen (United States)

12:15pm – 1:00pm

Lunch break

Neurologic co-morbidities of RLS

Rochelle Zak (United States)

1:00pm – 1:03pm

Introduction

Garima Shukla (Canada)

1:03pm – 1:23pm

Restless legs syndrome and Parkinson's disease - the dopaminergic connection and treatment challenges

Luigi Ferini-Strambi (Italy)

1:23pm – 1:40pm

Restless legs syndrome in acute neurological conditions - lessons from stroke and acute neuropathies

Garima Shukla (Canada)

1:40pm – 2:00pm

How RLS contributes to quality of life in Multiple Sclerosis

Mauro Manconi (Switzerland)

Young Investigators

Arthur Walters (United States); Denise Sharon (United States); Rochelle Zak (United States); John Swieca (United States)

2:00pm – 2:15pm

Young Investigator Presentation #1

2:15pm – 2:30pm

Young Investigator Presentation #2

3:15pm – 3:30pm

Break

IRLSSG Projects

Denise Sharon (United States)

3:30pm – 3:45pm

Diagnostic accuracy of RLS screening tools

Stephany Fulda (Switzerland)

3:45pm – 4:00pm

Update on PLMS scoring program certification

Stephany Fulda (Switzerland)

4:00pm – 4:15pm

National RLS Opioid Registry: 1-2 year longitudinal results

John Winkelman (United States)

4:15pm – 4:30pm

Establishing RSD as a new diagnosis

Lourdes DelRosso (Peru)

4:30pm – 4:45pm

Pediatric RLS and GP Task Force update

Arthur Walters (United States)

4:45pm – 4:50pm

Ideas for projects from the attendees

4:50pm – 5:00pm

Outgoing chair summary

Diego Garcia-Borreguero (Spain)

5:00pm – 5:15pm

Break

5:15pm – 6:00pm

Business meeting

Diego Garcia-Borreguero (Spain)

6:00pm – 9:00pm

Dinner

2:30pm – 2:45Ppm

Young Investigator Presentation #3

2:45pm – 3:00pm

Young Investigator Presentation #4

3:00pm – 3:15pm

Young Investigator Presentation #5



■ Pathophysiological insights from animal models of restless legs syndrome

3:00pm – 4:30pm | Room 219

Chair

Yuqing Li (United States)

Summary

Iron deficiency, which produces changes in dopaminergic neurons and receptors in the substantia nigra and putamen, has been reported to correlate with restless legs syndrome (RLS). Iron Deficient rats have insomnia and severe PLM in wake and in Slow Wave Sleep. The sleep pattern and symptoms of putamen-lesioned rats and ID rats resemble human RLS patients. Using neurotoxic lesion, in vivo microdialysis HPLC analysis, microinfusion of GABAA receptor agonists and antagonists, systemic injection of histamine receptor agonist and antagonist, Western blotting, and EEG spectral analysis techniques, a comprehensive understanding of RLS pathophysiology has emerged.

Recently, genome-wide association studies were performed, and 19 genetic loci were found to impart varying increased risk of developing RLS. Among these loci, genetic regions containing the genes MEIS1 and BTBD9 represent the top two hits and have been replicated in multiple independent genetic studies. The identification of these RLS candidate genes paved the way for making genetic animal model of RLS that could potentially be more relevant in elucidating the pathophysiology of RLS and developing therapeutic treatments.

The speakers are established scientists in the RLS pathophysiology and published extensively in this and related topics.

3:00pm – 3:02pm

Introduction

3:02pm – 3:22pm

Pathophysiological insights from the iron deficient rats

Yuan-Yang Lai (United States)

3:22pm – 3:42pm

Pathophysiological studies of RLS using BTBD9 mutant animal models

Yuqing Li (United States)

3:42pm – 4:02pm

MEIS1-based animal models and the pathophysiology of RLS

Aaro Salminen (Germany)

4:02pm – 4:22pm

Use of animal models for the pathophysiological study of RLS

Mauro Manconi (Switzerland)

4:22pm – 4:30pm

Conclusion

■ Sensory–motor network of the restless legs syndrome (RLS): Electrophysiology and imaging

10:45am – 12:15pm | Room 119

Chair

Richard Allen (United States)

Summary

In recent years, there are most progress on pathophysiology of restless legs syndrome (RLS), especially electrophysiological and neuroimaging researches in sensory-motor disorder of RLS.

Novel imaging techniques such as functional MRI and diffusion tractography imaging have demonstrated activation or connectivity changes in the sensory–motor network. The cortex, basal ganglia, cerebellum, thalamus, and their connections seem to play a key part in abnormalities of sensory–motor processing in RLS. Also, RLS patients exhibit increased excitability of the sensorimotor cortex, a remarkable abnormality existing in early somatosensory gating control and an attenuated inhibitory interneuron network by electrophysiological magnetoencephalography. But in vivo excitability studies on motor and sensory axons of the median nerve provide evidence that the increased excitability of peripheral motoneurons but not sensory axons contributes to the pathophysiology of RLS. And RLS like tics in Tourette's syndrome, the movement disorders are modulated by internal and external sensory signals and that abnormal sensorimotor integration might alter normal motor control. Reduced short-latency afferent inhibition, a marker for sensorimotor integration, has been shown with transcranial magnetic stimulation (TMS) in RLS patients. Further, low-frequency rTMS on S1-M1 connectivity alleviated the sensory–motor complaints of RLS patients by modulating cortical excitability and inducing short-term synaptic plasticity.

This symposium will provide the electrophysiological and imaging evidence for the abnormality of sensory–motor network and gain novel insight into physiopathologic mechanism of RLS in order to better guide the treatment.

10:45am – 10:47am

Introduction

10:47am – 11:03am

fMRI: Connectivity and sensory-motor systems in RLS

Yong Won Cho (Republic of Korea)

11:03am – 11:19am

The mechanism of sensory disorder in RLS based on MEG

Yuping Wang (China)

11:19am – 11:35am

Non-invasive brain stimulation and RLS: Clinical, electrophysiological and neuroplastic effects

Giuseppe Lanza (Italy)



11:35am – 11:51am

The sensory experience of RLS and its relationship to pain, itch and Tourette's

John Winkelman (United States)

11:51am – 12:07pm

Peripheral mechanisms in restless legs syndrome

Dirk Czesnik (Germany)

12:07pm – 12:15pm

Conclusion

KEYNOTE SPEAKER



■ K08: Restless legs syndrome/periodic limb movements of sleep: New insights into neurobiology and treatment

2:00pm - 2:45pm | Room 212

Keynote

Diego García-Borreguero (Spain)

Summary

Restless legs syndrome (RLS) is a common chronic neurological disorder that manifests through sensorimotor symptoms that interfere with rest and sleep. It has a wide spectrum of symptom severity affecting not only quality of life but also possibly increasing cardiovascular risk.

Our knowledge on the causes and mechanisms of RLS is still limited: several susceptible single nucleotide polymorphisms such as BTBD9 and MEIS1, which are thought to be involved in embryonic neuronal development, have been reported to be associated with RLS. An increasing number of studies have suggested an important role of brain iron deficiency in the pathophysiology of RLS. Moreover, a number of recent preclinical and clinical studies suggest a hypoadenosinergic state leading to hypersensitive cortico-striatal input and leading to a striatal presynaptic hyperglutamatergic and hyperdopaminergic neurotransmission. Understanding the interplay between these dysfunctional striatal circuitries might be crucial to develop new therapeutic targets.

2:00pm – 2:02pm

Introduction

2:00pm – 2:45pm

Restless legs syndrome/periodic limb movements of sleep: New insights into neurobiology and treatment

■ Brain iron as a central factor in the pathophysiology of RLS: Emerging evaluation methods and therapeutic opportunities

3:00pm – 4:30pm | Ballroom A

Chair

Diego García-Borreguero (Spain)

Summary

A number of epidemiological and clinical studies support the notion that a brain iron dysregulation, despite normal peripheral iron, plays a key role in the pathophysiology of RLS. Such a concept is also supported by an increasing number of experimental and animal data. In addition, new, large multicentric studies show a complete, long-lasting remission of RLS symptoms for some patients when this brain iron deficit is addressed by treatment with intravenous iron.

The present symposium will discuss the latest concepts on brain iron homeostasis, along with very recent studies that show how a brain iron deficit causes an increased corticostriatal hyperexcitability by means of changes in extracellular adenosine, leading to a hyperdopaminergic and hyperglutamatergic state. It will also discuss methods to evaluate brain iron homeostasis in RLS. The Symposium will discuss most recent neuroimaging data (3 and 7 Tesla MRI), identification of critical brain regions, and the goals and safety of iron treatments. Preliminary data will be presented on transcranial sonography of the substantia nigra which demonstrate its potential as a new clinical tool predicting benefit from intravenous iron treatment.

3:00pm – 3:02pm

Introduction

3:02pm – 3:22pm

Brain iron deficiency relation to dopamine dysfunction and augmentation in RLS

Christopher Earley (United States)

3:22pm – 3:42pm

Brain iron dysregulation in RLS relation to brain adenosine and glutamate

Sergi Ferre (United States)

3:42pm – 4:02pm

MRI evaluation of regional brain iron relation to RLS symptoms and iron treatments

Richard Allen (United States)

4:02pm – 4:22pm

Transcranial sonography evaluation of substantia nigra iron: A potential clinical tool to predict IV iron treatment outcome

Celia Garcia Malo (Spain)

4:22pm – 4:30pm

Conclusion

