

BIOGRAPHICAL SKETCH

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NAME Livio Luongo, Date of Birth Naples, 06-09-1979 Email address: livio.luongo@gmail.com Phone: +39(0)815667658 Mobile: +393289167523	POSITION TITLE Assistant Professor
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EDUCATION/TRAINING (Begin with baccalaureate or other initial professional education, such as nursing, and include postdoctoral training.)

INSTITUTION AND LOCATION	DEGREE (if applicable)	YEAR(s)	FIELD OF STUDY
Second University of Naples, Italy. Wiliam Harvey, Queen Mary University, London	Post-Doc	2008-2012 (2012, London)	Neuropharmacology Pharmacology
Second University of Naples, Italy, King's College, London,	PhD	2008 (2006-2007 London)	
University of Naples Federico II	Degree	2004	Pharmacy

A. Position and research interest.

Researcher: Department of Experimental Medicine, Division of Pharmacology, Second University of Naples, Italy.

Research Interests

- 1) Neuron-glia communications in the development of neuropathic pain.
- 2) Neuropathic pain pathophysiology
- 3) New pharmacological tools on microglia cells
- 4) Involvement of endocannabinoid system in the pathophysiology of the chronic neuroinflammatory diseases

Teaching:

- 1) Course of toxicology at the University of Farmacy, Naples, Italy
- 2) Course of basic pharmacology at the University of Laboratory technicians, Naples, Italy
- 3) Course of Pharmacology at University of Medicine, Naples, Italy
- 4) Seminars at the University of Medicine, Naples, Italy
- 5) Erasmus Teaching at the Complutense University, Madrid, Spain

- 6) Erasmus teaching at the University of Sevilla, Spain
- 7) Seminars at University of Brno, CZ
- 8) Seminars at the University of Galway, Ireland

Bibliometric parameters

N° Pubbs= 87

H-index= 28

Citations= 1797

B. Selected peer-reviewed publications (in chronological order).

List of publications

2018:

1. Boccella S, Guida F, Palazzo E, Marabese I, de Novellis V, Maione S, Luongo L. Spared Nerve Injury as a Long-Lasting Model of Neuropathic Pain. *Methods Mol Biol.* 2018;1727:373-378. (Book chapter)
2. Petrelli R, Scorticini M, Belardo C, Boccella S, Luongo L, Capone F, Kachler S, Vita P, Del Bello F, Maione S, Lavecchia A, Klotz KN, Cappellacci L. Structure-Based Design, Synthesis, and In Vivo Antinociceptive Effects of Selective A1 Adenosine Receptor Agonists. *J Med Chem.* 2018 ;61(1):305-318.
3. Mhillaj E, Morgese MG, Tucci P, Furiano A, Luongo L, Bove M, Maione S, Cuomo V, Schiavone S, Trabace L. Celecoxib Prevents Cognitive Impairment and Neuroinflammation in Soluble Amyloid β -treated Rats. *Neuroscience.* 2018;372:58-73.
4. Coraggio V, Guida F, Boccella S, Scafuro M, Paino S, Romano D, Maione S, Luongo L. Neuroimmune-Driven Neuropathic Pain Establishment: A Focus on Gender Differences. *Int J Mol Sci.* 2018;19(1).
5. Wahlman C, Doyle TM, Little JW, Luongo L, Janes K, Chen Z, Esposito E, Tosh DK, Cuzzocrea S, Jacobson KA, Salvemini D. Chemotherapy-induced pain is promoted by enhanced spinal adenosine kinase levels through astrocyte-dependent mechanisms. *Pain.* 2018; 159(6):1025-1034.
6. Luongo L, Salvemini D. Targeting metabotropic adenosine receptors for neuropathic pain: Focus on A2A. *Brain Behav Immun.* 2018; 69:60-61.
7. Marabese I, Boccella S, Iannotta M, Luongo L, de Novellis V, Guida F, Serra N, Farina A, Maione S, Palazzo E. Metabotropic glutamate receptor subtype 7 in the dorsal striatum oppositely modulates pain in sham and neuropathic rats. *Neuropharmacology.* 2018; 135:86-99.
8. Guida F, Turco F, Iannotta M, De Gregorio D, Palumbo I, Sarnelli G, Furiano A, Napolitano F, Boccella S, Luongo L, Mazzitelli M, Usiello A, De Filippis F, Iannotti FA, Piscitelli F, Ercolini D, de Novellis V, Di Marzo V, Cuomo R, Maione S. Antibiotic-induced microbiota perturbation causes gut endocannabinoidome changes, hippocampal neuroglial reorganization and depression in mice. *Brain Behav Immun.* 2018. pii: S0889-1591(17)30417-8. doi: 10.1016/j.bbi.2017.09.001

2017

9. D'Aniello A, Luongo L, Romano R, Iannotta M, Marabese I, Boccella S, Belardo C, de Novellis V,

- Arra C, Barbieri A, D'Aniello B, Scandurra A, Maglizzi L, Fisher G, Guida F, Maione S. d-Aspartic acid ameliorates painful and neuropsychiatric changes and reduces β -amyloid A β 1-42 peptide in a long lasting model of neuropathic pain. *Neurosci Lett.* 2017;651:151-158.
10. Guida F, Luongo L, Boccella S, Giordano ME, Romano R, Bellini G, Manzo I, Furiano A, Rizzo A, Imperatore R, Iannotti FA, D'Aniello E, Piscitelli F, Sca Rossi F, Cristina L, Di Marzo V, de Novellis V, Maione S. Palmitoylethanolamide induces microglia changes associated with increased migration and phagocytic activity: involvement of the CB2 receptor. *Sci Rep.* 2017;7(1):375.
 11. Guida F, Boccella S, Iannotta M, De Gregorio D, Giordano C, Belardo C, Romano R, Palazzo E, Scafuro MA, Serra N, de Novellis V, Rossi F, Maione S, Luongo L. Palmitoylethanolamide Reduces Neuropsychiatric Behaviors by Restoring Cortical Electrophysiological Activity in a Mouse Model of Mild Traumatic Brain Injury. *Front Pharmacol.* 2017;8:95.
 12. Luongo L, Starowicz K, Maione S, Di Marzo V. Allodynia Lowering Induced by Cannabinoids and Endocannabinoids (ALICE). *Pharmacol Res.* 2017 ;119:272-277.
 13. Bellini G, Torella M, Manzo I, Tortora C, Luongo L, Punzo F, Colacurci N, Nobili B, Maione S, Rossi F. PKC β II-mediated cross-talk of TRPV1/CB2 modulates the glucocorticoid-induced osteoclast overactivity. *Pharmacol Res.* 2017;115:267-274.
 14. Palazzo E, Marabese I, **Luongo L**, Guida F, de Novellis V, Maione S. Nociception modulation by supraspinal group III metabotropic glutamate receptors. *J Neurochem.* 2017;141(4):507-519.

2016:

15. Rossi F, Bellini G, **Luongo L**, Manzo I, Tolone S, Tortora C, Bernardo ME, Grandone A, Conforti A, Docimo L, Nobili B, Perrone L, Locatelli F, Maione S, Del Giudice EM. Cannabinoid Receptor 2 as Antiobesity Target: Inflammation, Fat Storage, and Browning Modulation. *J Clin Endocrinol Metab.* 2016 ;101(9):3469-78.
16. Palazzo E, **Luongo L**, Guida F, Marabese I, Romano R, Iannotta M, Rossi F, D'Aniello A, Stella L, Marmo F, Usiello A, de Bartolomeis A, Maione S, de Novellis V. D-Aspartate drinking solution alleviates pain and cognitive impairment in neuropathic mice. *Amino Acids.* 2016
17. Moriello AS, **Luongo L**, Guida F, Christodoulou MS, Perdicchia D, Maione S, Passarella D, Di Marzo V, De Petrocellis L. Chalcone derivatives activate and desensitize the transient receptor potential ankyrin 1 cation channel, subfamily A, member 1 TRPA1 ion channel: structure-activity relationships in vitro and anti-nociceptive and anti-inflammatory activity in vivo. *CNS Neurol Disord Drug Targets.* 2016 *In press*
18. Punzo D, Errico F, Cristina L, Sacchi S, Keller S, Belardo C, **Luongo L**, Nuzzo T, Imperatore R, Florio E, De Novellis V, Affinito O, Migliarini S, Maddaloni G, Sisalli MJ, Pasqualetti M, Pollegioni L, Maione S, Chiariotti L, Usiello A. Age-Related Changes in d-Aspartate Oxidase Promoter Methylation Control Extracellular d-Aspartate Levels and Prevent Precocious Cell Death during Brain Aging. *J Neurosci.* 2016;36(10):3064-78.
19. Aiello F, Badolato M, Pessina F, Sticozzi C, Maestrini V, Aldinucci C, **Luongo L**, Guida F, Ligresti A, Artese A, Allarà M, Costa G, Frosini M, Schiano Moriello A, De Petrocellis L, Valacchi G, Alcaro S, Maione S, Di Marzo V, Corelli F, Brizzi A. Design and Synthesis of New Transient Receptor

Potential Vanilloid Type-1 (TRPV1) Channel Modulators: Identification, Molecular Modeling Analysis, and Pharmacological Characterization of the N-(4-Hydroxy-3-methoxybenzyl)-4-(thiophen-2-yl)butanamide, a Small Molecule Endowed with Agonist TRPV1 Activity and Protective Effects against Oxidative Stress. *ACS Chem Neurosci.* 2016 *In press.*

20. Brindisi M, Maramai S, Gemma S, Brogi S, Grillo A, Di Cesare Mannelli L, Gabellieri E, Lamponi S, Saponara S, Gorelli B, Tedesco D, Bonfiglio T, Landry C, Jung KM, Armirotti A, **Luongo L**, Ligresti A, Piscitelli F, Bertucci C, Dehouck MP, Campiani G, Maione S, Ghelardini C, Pittaluga A, Piomelli D, Di Marzo V, Butini S. Development and Pharmacological Characterization of Selective Blockers of 2-Arachidonoyl Glycerol Degradation with Efficacy in Rodent Models of Multiple Sclerosis and Pain. *J Med Chem.* 2016;59(6):2612-32.
21. Cristino L*, **Luongo L***, Imperatore R, Boccella S, Becker T, Morello G, Piscitelli F, Busetto G, Maione S, Di Marzo V. Orexin-A and Endocannabinoid Activation of the Descending Antinociceptive Pathway Underlies Altered Pain Perception in Leptin Signalling Deficiency. *Neuropsychopharmacology.* 2016 Jan;41(2):508-20

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22. Rinaldi B, Guida F, Furiano A, Donniacuo M, **Luongo L**, Gritti G, Urbanek K, Messina G, Maione S, Rossi F, de Novellis V. Effect of Prolonged Moderate Exercise on the Changes of Nonneuronal Cells in Early Myocardial Infarction. *Neural Plast.* 2015;2015:265967. doi: 10.1155/2015/265967
23. Guida F*, **Luongo L***, Marmo F, Romano R, Iannotta M, Napolitano F, Belardo C, Marabese I, D'Aniello A, De Gregorio D, Rossi F, Piscitelli F, Lattanzi R, de Bartolomeis A, Usiello A, Di Marzo V, de Novellis V, Maione S. Palmitoylethanolamide reduces pain-related behaviors and restores glutamatergic synapses homeostasis in the medial prefrontal cortex of neuropathic mice. *Mol Brain.* 2015 Aug 12;8:47. doi: 10.1186/s13041-015-0139-5
24. Imperatore R, Morello G, **Luongo L**, Taschler U, Romano R, De Gregorio D, Belardo C, Maione S, Di Marzo V, Cristino L. Genetic deletion of monoacylglycerol lipase leads to impaired cannabinoid receptor CB1 R signaling and anxiety-like behavior. *J Neurochem.* 2015. doi: 10.1111/jnc.13267
25. Rossi F, Bellini G, Tortora C, Bernardo ME, **Luongo L**, Conforti A, Starc N, Manzo I, Nobili B, Locatelli F, Maione S. CB2 and TRPV1 receptors oppositely modulate in vitro human osteoblast activity. *Pharmacol Res.* 2015. pii: S1043-6618(15)00124-3. doi: 10.1016/j.phrs.2015.06.010
26. Lorenzo Di Cesare Mannelli, Alessandra Pacini, Francesca Corti, Serena Boccella, **Livio Luongo**, Emanuela Esposito, Salvatore Cuzzocrea, Sabatino Maione, Antonio Calignano, Carla Ghelardini Antineuropathic profile of N-Palmitoylethanolamine in a rat model of oxaliplatin-induced neurotoxicity, *Plos One* *in press*
27. Calzetta L, **Luongo L**, Cazzola M, Page C, Rogliani P, Facciolo F, Maione S, Capuano A, Rinaldi B, Matera MG. Contribution of sensory nerves to LPS-induced hyperresponsiveness of human isolated bronchi. *Life Sci.* 2015;131:44-50.
28. **Luongo L**, Malcangio M, Salvemini D, Starowicz K. Chronic pain: new insights in molecular and

cellular mechanisms. Biomed Res Int.; 2015:676725.

29. Spaziano G°, **Luongo L°**, Guida F, Petrosino S, Matteis M, Palazzo E, Sullo N, de Novellis V, Di Marzo V, Rossi F, Maione S, D'Agostino B. Exposure to Allergen Causes Changes in NTS Neural Activities after Intratracheal Capsaicin Application, in Endocannabinoid Levels and in the Glia Morphology of NTS. Biomed Res Int.; 2015:980983.
30. Cristino L°, **Luongo L°**, Squillace M, Paolone G, Mango D, Piccinin S, Zianni E, Imperatore R, Iannotta M, Longo F, Errico F, Vescovi AL, Morari M, Maione S, Gardoni F, Nisticò R, Usiello A. d-Aspartate oxidase influences glutamatergic system homeostasis in mammalian brain. Neurobiol Aging. 2015; 36(5):1890-902.
31. Palazzo E, Romano R, **Luongo L**, Boccella S, De Gregorio D, Giordano ME, Rossi F, Marabese I, Scafuro MA, de Novellis V, Maione S. MMPIP, an mGluR7-selective negative allosteric modulator, alleviates pain and normalizes affective and cognitive behavior in neuropathic mice. Pain. 2015;156(6):1060-73.
32. Petrelli R, Torquati I, Kachler S, **Luongo L**, Maione S, Franchetti P, Grifantini M, Novellino E, Lavecchia A, Klotz KN, Cappellacci L. 5'-C-Ethyl-tetrazolyl-N(6)-substituted adenosine and 2-chloro-adenosine derivatives as highly potent dual acting A1 adenosine receptor agonists and A3 adenosine receptor antagonists. J Med Chem. 2015;58(5):2560-6.
33. Lopez-Canul M, Palazzo E, Dominguez-Lopez S, **Luongo L**, Lacoste B, Comai S, Angeloni D, Fraschini F, Boccella S, Spadoni G, Bedini A, Tarzia G, Maione S, Granados-Soto V, Gobbi G. Selective melatonin MT2 receptor ligands relieve neuropathic pain through modulation of brainstem descending antinociceptive pathways. Pain. 2015;156(2):305-17.
34. Maione S, Radanova L, De Gregorio D, **Luongo L**, De Petrocellis L, Di Marzo V, Imming P. Effects of metabolites of the analgesic agent dipyrone (metamizol) on rostral ventromedial medulla cell activity in mice. Eur J Pharmacol. 2015;748:115-22.
35. Guida F, Lattanzi R, Boccella S, Maftei D, Romano R, Marconi V, Balboni G, Salvadori S, Scafuro MA, de Novellis V, Negri L, Maione S, **Luongo L**. PC1, a non-peptide PKR1-preferring antagonist, reduces pain behavior and spinal neuronal sensitization in neuropathic mice. Pharmacol Res. 2015;91:36-46.
36. Little JW, Ford A, Symons-Liguori AM, Chen Z, Janes K, Doyle T, Xie J, **Luongo L**, Tosh DK, Maione S, Bannister K, Dickenson AH, Vanderah TW, Porreca F, Jacobson KA, Salvemini D. Endogenous adenosine A3 receptor activation selectively alleviates persistent pain states. Brain. 2015;138(Pt 1):28-35.

2014:

37. Rossi F, Perrotta S, Bellini G, **Luongo L**, Tortora C, Siniscalco D, Francese M, Torella M, Nobili B, Di Marzo V, Maione S. Iron overload causes osteoporosis in Thalassemia Major patients through interaction with TRPV1 channels. Haematologica. 2014; 99(12):1876-84
38. Maftei D, Marconi V, Florenzano F, Giancotti LA, Castelli M, Moretti S, Borsani E, Rodella LF,

- Balboni G, **Luongo L**, Maione S, Sacerdoti P, Negri L, Lattanzi R. Controlling the activation of the Bv8/Prokineticin system reduces neuroinflammation and abolishes thermal and tactile hyperalgesia in neuropathic animals. *Br J Pharmacol.* 2014; 171(21):4850-65
39. Brizzi A, Aiello F, Marini P, Cascio MG, Corelli F, Brizzi V, De Petrocellis L, Ligresti A, **Luongo L**, Lamponi S, Maione S, Pertwee RG, Di Marzo V. Structure-affinity relationships and pharmacological characterization of new alkyl-resorcinol cannabinoid receptor ligands: Identification of a dual cannabinoid receptor/TRPA1 channel agonist. *Bioorg Med Chem.* 2014; 22(17):4770-83.
40. **Luongo L**, Maione S, Di Marzo V. Endocannabinoids and neuropathic pain: focus on neuron-glia and endocannabinoid-neurotrophin interactions. *Eur J Neurosci.* 2014; 39(3):401-8
41. Rossi F, Bellini G, Torella M, Tortora C, Manzo I, Giordano C, Guida F, **Luongo L**, Papale F, Rosso F, Nobili B, Maione S. The genetic ablation or pharmacological inhibition of TRPV1 signalling is beneficial for the restoration of quiescent osteoclast activity in ovariectomized mice. *Br J Pharmacol.* 2014; 171(10):2621-30
42. Rossi F, Marabese I, De Chiaro M, Boccella S, **Luongo L**, Guida F, De Gregorio D, Giordano C, de Novellis V, Palazzo E, Maione S. Dorsal striatum metabotropic glutamate receptor 8 affects nocifensive responses and rostral ventromedial medulla cell activity in neuropathic pain conditions. *J Neurophysiol.* 2014; 111(11):2196-209
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- 2013:**
45. Rossi F, Bernardo ME, Bellini G, **Luongo L**, Conforti A, Manzo I, Guida F, Cristino L, Imperatore R, Petrosino S, Nobili B, Di Marzo V, Locatelli F, Maione S. The cannabinoid receptor type 2 as mediator of mesenchymal stromal cell immunosuppressive properties. *PLoS One.* 2013; 8(11):e80022
46. Palazzo E, Marabese I, **Luongo L**, Boccella S, Bellini G, Giordano ME, Rossi F, Scafuro M, Novellis Vd, Maione S. Effects of a metabotropic glutamate receptor subtype 7 negative allosteric modulator in the periaqueductal grey on pain responses and rostral ventromedial medulla cell activity in rat. *Mol Pain.* 2013; 9:44.
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glial/microglial phenotypical changes in mice. *CNS Neurol Disord Drug Targets*. 2013;12(1):45-54.

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50. **Luongo L**, Petrelli R, Gatta L, Giordano C, Guida F, Vita P, Franchetti P, Grifantini M, de Novellis V, Cappellacci L, Maione S. 5'-Chloro-5'-deoxy-(\pm)-ENBA, a potent and selective adenosine A(1) receptor agonist, alleviates neuropathic pain in mice through functional glial and microglial changes without affecting motor or cardiovascular functions. *Molecules*. 2012;17(12):13712-26.
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54. **Luongo L**, Costa B, D'Agostino B, Guida F, Comelli F, Gatta L, Matteis M, Sullo N, De Petrocellis L, de Novellis V, Maione S, Di Marzo V. Palvanil, a non-pungent capsaicin analogue, inhibits inflammatory and neuropathic pain with little effects on bronchopulmonary function and body temperature. *Pharmacol Res*. 2012;66(3):243-50.
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66. G Aviello, F Borrelli, F Guida, B Romano, K Lewellyn, M De Chiaro, **L Luongo**, J K Zjawiony, S Maione, A A Izzo, R Capasso Ultrapotent effects of salvinorin A, a hallucinogenic compound from *Salvia divinorum*, on LPS-stimulated murine macrophages and its anti-inflammatory action in vivo *J Mol Med*. 2011 89(9):891-902.
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amide hydrolase decreases symptoms and central sequelae in the medial prefrontal cortex of neuropathic rats. Mol Pain. 2011;7:7

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C. Current Research Support.

Title: Role of glial and immune-inflammatory cells in chronic pain: identification of novel drug targets in the peripheral and central analgesia and neuroinflammation

PRIN 2015 Project of the Italian Ministry of University and Research, Import **48.000,00 euro**

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GW Pharmaceuticals, London, UK, import **68.000,00 euro (Principal Investigator)**

International Conferences:

-**Luongo L**. "the 10th Congress of the European Pain Federation, EFIC® 2017, 6-9 september 2017, Copenhagen relazione dal titolo "N-acylethanolamines in pain modulation and plasticity". (**Invited Speaker**)

Luongo L. **DIMETHYL FUMARATE REDUCES TACTILE ALLODYNIA IN A HCAR2-MEDIATED MECHANISM IN TWO MODELS OF PERIPHERAL NEUROPATHIC PAIN** "13th World Congress on Inflammation", July 8-12 2017, London

Luongo L "Palmitoylethanolamide Induces Microglia Changes Associated with Increased Migration and Phagocytic Activity: Involvement of the CB2 Receptor"; PCS International Conference of neuroscience (ICN-2017), 7-8 Aprile 2017, Lisbona. (**Invited Speaker**)

Luongo L. Palmitoylethanolamide systemic treatment reduces spinal and supraspinal formalin-induced neuroinflammation and allodynia . **XIV Congress of the European Shock Society**, Taormina Giardini Naxos, 2011. **(Invited Speaker)**

- **Livio Luongo** TRPV1 channel is involved in symptoms and central sequelae in the medial prefrontal cortex in a model of neuropathic pain in rats., **14th World Congress on Pain** August 27 – August 31, 2012 – Milan, Italy **(Invited Speaker)**

- Maione S, Palazzo E., **Luongo L**, Rossi F. and de Novellis V. (2011). Role of metabotropic glutamate receptors in the basolateral amygdala-driven prefrontal cortical deactivation or over-activation in models of inflammatory or neuropathic pain in the rat . **7th International Meeting on Metabotropic Glutamate Receptors**. Taormina, October 2-7, 2011

- **Luongo L**, Giordano C, Maione S (2011). The role of glial NO in the oxidative stress associated neuropathic pain. **7th Congress of the European Federation of ISAP Chapters, EFIC 2011**, Hamburg, Germany **(Invited Speaker)**.

- **Livio Luongo**, Phytocannabinoid modulation of neuropathic pain-associated neuroinflammation "Cannabinoid Function in the CNS" **Gordon Research Conference** Les Diablerets Conference Center, Les Diablerets, Switzerland, May 22-27, 2011

- **Livio Luongo**, Leptin-controlled orexin/endocannabinoid interactions in the mouse periaqueductal grey: role in the regulation of the descending antinociceptive pathway. **XV Congress Of The Italian Society Of Neuroscience (S.I.N.S.)**, Rome 3-5 October 2013 **(Invited Speaker)**.

- **Livio Luongo**, A1 adenosine receptor regulate ATP-mediated microglial activation in vivo and in vitro: role in the spinal neuronal plasticity in neuropathic mice. **31st Camerino-Cyprus-Noordwijkerhout Symposium, Camerino**, May 19-23, 2013

- **Livio Luongo** Microglia activation is regulated by cannabinoid system, **Naples Pain Conference (NPC): Research and Therapy for human and animal suffering** MAY 16/19, 2010

- **Livio Luongo** TRPV1 channel is involved in symptoms and central sequelae in the medial prefrontal cortex of neuropathic rats Ewcbr 2012 – Villars Sur Ollon – Switzerland **(Invited Speaker)**.

Applicant Name (Last, first, middle):

- **Livio Luongo**, Luigia Cristino, Roberta Imperatore, Serena Boccella, Stefania Petrosino, Francesca Guida, Piero Orlando, Vincenzo Di Marzo and Sabatino Maione. Leptin-controlled orexin/endocannabinoid interactions in the mouse periaqueductal grey: role in the regulation of the descending antinociceptive pathway **ICRS2014 - The 24th International Cannabinoid Research**, 28 june-3 july, 2014, Baveno, Italy

- **Livio Luongo**, Palmitoylethanolamide chronic treatment reduces sensory and cognitive dysfunction associated with mild traumatic brain injury, Symposium "Endocannabinoids and related endogenous lipid signaling molecules" Mediterranean Neuroscience Society Meeting, 12-15 June 2015, Cagliari (**Invited Speaker**)