Curriculum vitae

A. Personal data



Name:	Igor Stagljar
Title:	Professor
Nationality:	Croatia and Canada
Languages:	Croatian (mother tongue) English (full professional proficiency) German (full professional proficiency)
LinkedIn:	https://www.linkedin.com/in/igor-stagljar-1216b82
Lab web page:	https://stagljarlab.com/

B. Personal Statement

Dr. Igor Stagljar is a Professor in the Departments of Biochemistry and Molecular Genetics at the Donnelly Centre, University of Toronto, Canada. He is also a Co-director and Laboratory Head at the Mediterranean Institute of Life Sciences, Split, Croatia. Before being recruited to the University of Toronto in 2005, Prof. Stagljar started his career at the University of Zurich (Switzerland). He received his Ph.D. from ETH Zurich (Switzerland) under the tutelage of Professors Charles Weissmann and Markus Aebi, and completed his training in the labs of Professors Walter Schaffner and Ueli Hübscher (University of Zurich, Switzerland) and Stanley Fields (University of Washington, USA).

Professor Stagliar is one of the world leaders in the field of proteomics/chemical genomics. developing high-throughput technologies to facilitate the study of the protein-protein interactions (PPIs) of various human membrane proteins involved in cancer. He has published over 140 research papers listed on PubMed, is the holder of 8 patents, and has been honored as one of the top inventors in Canada. The research in his lab has led to the development of the Membrane Yeast Two-Hybrid [MYTH; Stagljar et al. (1998) Proceedings of National Academy of Sciences] and the Mammalian Membrane Two-Hybrid [MaMTH; Petschnigg et al. (2014) Nature Methods] technologies, both of which have revolutionized membrane proteomic research. During his entire career he has been an innovator and disruptor, and his teams' recent adaptation of MaMTH into a drug screening platform [MaMTH-DS; Saraon, Snider et al. (2020) Nature Chemical Biology] and creation of the SIMPL technology [Yao et al. (2020) Nature Communications] have the potential to dramatically improve the development of therapeutics against membrane (MaMTH-DS) and soluble (SIMPL) PPI targets. The work of Stagliar lab has led to various seminal papers in the fields of genomics, proteomics, chemical biology and cancer signaling, with publications in leading scientific journals such as Science, Nature, Nature Methods, Nature Chemical Biology, Nature Cell Biology, Nature Communications, Molecular Cell, Cell Reports, Cell Systems, Proceedings of the National Academy of Sciences USA, Journal of Molecular Biology, Neuron, Genome Research, Oncogene and Molecular Systems Biology.

Professor Stagljar's lab is currently directing major proteomics/chemical genomics projects to map how integral membrane proteins interact to produce either healthy or diseased cells, to identify novel therapeutic targets, and to screen for novel drugs using artificial intelligence platforms, in particular to target lung, colon and pancreatic cancer. Notably, his group recently identified two new drugs (midostaurin and gilteritinib), suitable for potential repurposing to target EGFR-triple mutant on the surface of lung cancer cells, which are now currently the subject of prompt clinical study at The Princess Margaret Cancer Centre in Toronto, Canada and The Lung Cancer Clinic in Zagreb, Croatia. In addition, Stagljar lab is working on the further development of a novel EGFR-triple mutant inhibitor, that they named EMI1 [Saraon, Snider *et al.* (2020) *Nature Chemical Biology*; Saraon *et al.* (2021) *Journal of Molecular Biology*], which inhibits the signaling of EGFR-triple mutant lung cancer cells in a completely new way, leveraging molecular machineries to degrade mutated EGFR protein on the surface of tumor cells. They hypothesize that EMI1's complex mechanism of action will make it more difficult for lung tumors to develop resistance to it. Furthermore, Stagljar lab recently identified several potent small molecule inhibitors specific against KRAS G12V mutant and is currently analyzing these compounds in pre-clinical models of lung, colon and pancreatic cancer.

Additionally, the Stagljar lab has recently made major contributions to the fight against COVID-19. Namely, in only seven months, the Stagljar lab developed an innovative and cost-effective COVID-19 serological test named SATiN (for <u>Serological Assay</u> based on split <u>Tripart Nanoluciferase</u>). A paper describing this technology was recently published in *Nature Communications* [Yao *et al.* (2021)] and was featured in all major TV segments and news media publications in Canada and abroad. The Stagljar lab has recently further modified this split-luciferase assay so that it can be used to detect the presence and potency of neutralizing antibodies against SARS-CoV-2, including the wild-type and all major variants of concern [Kim and Yao *et al.* (2022) *in press* in *Nature Communications*]. These two papers on COVID-19 further demonstrate how productive and efficient the Stagljar lab is, able to respond quickly to current needs during this pandemic with innovative technology development.

D. Distinctions and Funding

Professor Stagljar is also a recipient of several prestigious distinctions for his outstanding contributions to biology including the Croatian Biological Society Plaque "Zdravko Lorkovic" – the highest accolade given by the Croatian Biological Society, and is a member of the Croatian Academy of Arts and Sciences. In 2015, the University of Toronto honored Dr. Stagljar with "Inventor of the Year" award for his contribution to Canada's innovation agenda and the advancement of knowledge. In 2022, Prof. Stagljar was elected to Royal Society of Canada and European Molecular Biology Organization (EMBO), which both recognize his excellence and outstanding achievements in biomedicine not only in Canada but also worldwide. In 2023, Prof. Stagljar was listed among Top 50 most influential people from the Adria region by Bloomberg.

As principal investigator, Dr. Stagljar secured over \$60 million in research funds since he moved to Canada (in 2005) from a range of national and international funding agencies, non-profit foundations and pharmaceutical companies, such as Orionis Biosciences, Genentech, Merck, Evotec and Novartis. Based on his academic successes, Dr. Stagljar has co-founded two companies, Dualsystems Biotech Inc. (<u>https://www.dualsystems.com/</u>), a world-leading company in the field of interactive proteomics, and a new Toronto-based biotech start-up company named Perturba Therapeutics (<u>https://www.perturbatx.com/</u>).

Science and innovation are his two greatest passions and as a member of the Editorial Board of *Molecular Systems Biology, Journal of Molecular Biology, Biochemical and Biophysical Research Communications, and BioTechniques,* Dr. Stagljar remains connected to innovative and leading-edge research.

E. Research Interests

- Proteomics (i)
- (ii) Protein interaction networks in health & disease
- (iii) Mechanism of action of drugs
- Artificial intelligence in drug discovery (iv)
- Cell signaling (v)
- (vi)
- Membrane transport COVID-19 serological methods (vii)

F. Positions and Honors

Positions

1995-2000	Post Doctoral Fellow, Department of Molecular Biology, University of Zurich, Switzerland
2000-2001	Oberassistent (research associate), Institute of Veterinary Biochemistry and Molecular Biology, University of Zurich, Switzerland
2001-2002	Senior Research Associate, Genome Sciences and Medicine, University of Washington, Washington, USA
2002-2005	Assistant Professor, Institute of Veterinary Biochemistry and Molecular Biology, University of Zurich, Switzerland
2005-2009	Associate Professor, Donnelly Centre, Department of Biochemistry, Department of Molecular Genetics, University of Toronto, Canada
2010 -	Professor, Donnelly Centre, Department Molecular Genetics, Department of Biochemistry, University of Toronto, Canada
2016	Sabbatical at the Salk Institute for Biological Sciences, La Jolla, CA (Feb-April, 2016), working in the lab of Prof. Tony Hunter
2016	Sabbatical at the German Cancer Research Center, Heidelberg, Germany (April-May 2016), working in the lab of Prof. Jörg Hoheisel
2016	Sabbatical at the Mediterranean Institute of Life Sciences, Split, Croatia (June-Aug 2016)
2023	Sabbatical at the Institute of Molecular Systems Biology at ETH Zurich, Switzerland (Jan-July 2023)

2005-	Editorial Board Member, BioTechniques
2006-	Editorial Board Member, Molecular Genetics and Genomics
2012-	Editorial Board Member, Journal of Molecular Biology
2013-	Editorial Board Member, Biochemical and Biophysical Research Communications
2014-	Editorial Board Member, Molecular Systems Biology

G. Leadership positions

- 2002-2005 Director of the Proteomics Centre at the University of Zurich, Switzerland
- 2019- Co-director and Laboratory head at the Mediterranean Institute of Life Sciences, Split, Croatia
- 2021- Co-ordinator of the Postdoctoral Association at the Department of Biochemistry and Department of Molecular Genetics, Faculty of Medicine at the University of Toronto

Journal Reviewing

Regular:

Cell, Cell Reports, Cell Systems, Science, Nature, Nature Methods, Nature Chemical Biology, Nature Reviews Drug Discovery, Nature Biotechnology, Proteomics, Nucleic Acids Research, Genome Research, Molecular Systems Biology, Molecular and Cellular Proteomics, Journal of Proteome Research, PLoS Genetics, PLoS Pathogen, Molecular Cellular Proteomics

Occasional:

Biotechniques, EMBO Journal, Genes and Development, Nature Genetics, Proceedings of National Academy of Science, Trends in Genetics, Trends in Biochemical Sciences, Current Opinion in Molecular Biology, Proteins, ACS Chemical Biology, FASEB Journal, Expert Opinion On Therapeutic Targets, Trends in Biochemical Sciences, Drug Discovery Today, Molecular and Cellular Biology, FEBS Journal

Grant Reviewing

Swiss National Foundation (2003-present)

Welcome Trust Funds (UK) (2004, 2007, 2009, 2014, 2016)

German National Genome Research Network (NGFN) (2006, 2009, 2012, 2016)

Croatian Ministry of Science (2006-present)

European Community (2007-2011)

German Scientific Society (DFG) (2007, 2008, 2010, 2015)

Canadian Cancer Society Research Institute (2007-2010, 2016, 2017, 2018)

Humbolt Foundation Germany (2008-2011)

Danish Ministry of Science (2008-2010)

National Science Foundation USA (2008, 2012, 2015, 2016)

Flanders National Foundation Belgium (2009, 2010, 2014, 2015, 2017)

Medical Research Council UK (2009, 2011, 2013)

Chilean National Commission for Scientific and Technological Research (2010-present)

European Research Council Senior Awards (ERC) (2011-present)

Canadian Institute of Health Research (CIHR) (2011-present)

European Union Horizon 2020 "Personalized Medicine" (2015)

Canadian Cancer Society Research Institute (2015-present)

Other Selected Accomplishments and Honors

2000-2011	Co-founder, Vice President and board member of the biotech	
	company Dualsystems Biotech Inc.	

- 2011- Co-founder of the biotech company Dualsystems Biotech Inc.
- 2003 EMBO Young Principal Investigator Award
- 2005 Dean's Fund Award, University of Toronto, Canada
- 2006 Leaders Technology Award, Canadian Funds for Innovation (CFI), Canada
- 2008 Featured in the article "The world renowned Protein Interaction Experts" in *The Scientist*
- 2010 Selection as one of "tomorrow's PIs" Genome Technology Magazine
- 2011 Featured in the 45 min radio interview in the Canadian Broadcasting Corporation (CBC)
- 2012 Featured in the Success Story article "The Scientist to Watch" in "The Canadian

Immigrant"

- 2014 National Award "Rudjer Boskovic" for the Scientific Achievements by University of Split, Croatia
- 2014 Member of the Croatian Academy of Arts and Science
- 2014 Featured in the Globe & Mail article "Scientists fear effect of changes in research funding"
- 2014 The "Croatian of the Year 2014" Award by *Vecernji List*, category Science
- 2015 The 2015 Inventor of the Year Award, University of Toronto
- 2016 Croatian Biological Society Plaque "Zdravko Lorkovic" for outstanding contributions to biology
- 2016 Hosted a visit of the President of Croatia, her Excellency Kolinda Grabar-Kitarovic to the University of Toronto
- 2019 Invitation to the state dinner with the Prime Minister of Canada Justin Trudeau, The Governor General of Canada Julie Payette and the President of Croatia, her Excellency Kolinda Grabar-Kitarovic at the Rideau Hall, Ottawa (May 13, 2019)
- 2021 Co-founder of the biotech company Perturba Therapeutix Inc., Toronto, ON, Canada
- 2021 Featured on the CBC's The National, CityNews TV and The Globe and Mail

CBC The National:

https://www.cbc.ca/player/play/1881874499884?fbclid=lwAR1o9TAekGIM7xbdnCNZR-EgDANDBuVg-E9K_5Bmxg5AtSYhH2_hb3IKRis

CityNews TV:

https://toronto.citynews.ca/video/2021/03/25/toronto-researchers-create-firefly-methodto-measure-covid-19immunity/?fbclid=IwAR1Edoud4L7v7mYU1A_HG5v9XKMjw5fCyGOfkdr00mjNICcXIcTZ d0ddVvU

The Globe and Mail:

https://www.theglobeandmail.com/canada/article-firefly-test-aims-to-shed-light-on-covid-19-vaccine-endurance/?fbclid=IwAR2XKSoux26p-G3Td3uQOWqAzAthJ5NuljLofY7LOVJZyQZWwYgohRVQExE

Global News:

https://globalnews.ca/newsletter/7861743/

Toronto Star:

https://www.thestar.com/life/health_wellness/2021/11/17/i-wonder-if-i-had-covid-when-iwas-sick-in-january-2020-why-widespread-antibody-testing-cant-come-soon-enoughfor-my-family.html?fbclid=lwAR3RFM-frL-QjCBrluTm4fASzrsCo0oyex3vdjOzFG4g9uq4A1yYE5R1gvA

2022 Featured on the CTV News, CityNews TV, Toronto Star and The Globe and Mail

CTV News:

https://www.ctvnews.ca/video?clipId=2483864&fbclid=IwAR3sj8_8eB3J-WC8UU_zsKTG2pL1WEUCFBX_CAAQObZm4zECfSrwY7vsxjQ

CityNews TV:

https://toronto.citynews.ca/2022/07/05/covid-19-antibody-test-torontoresearchers/?fbclid=IwAR2AxIpQH3kM3UQEQwyU1Xg8AS8OAV7J6TpIMm0JwmvOi_ eC2-dNw_C2LF4

Toronto Star:

https://www.thestar.com/news/canada/2022/07/10/if-you-got-covid-early-this-year-youcan-get-reinfected-now-u-of-t-studyfinds.html?fbclid=IwAR054b_T2kDofueC_dC_eyMKHWv8HHZX7jCAlv-QOa1JsxXRByzUrqDLZv4

The Globe and Mail:

https://www.theglobeandmail.com/canada/article-covid-19-summer-surge-ba5variant/?fbclid=IwAR2-C3yai6q2KWf_6yphH98Ic0gSHs_1TNrYuLibRo49j0AhDIWNiUIzOkQ#comments

- 2022 Election as the European Molecular Biology Organization (EMBO) Associate Member
- 2022 Fellow of the Royal Society of Canada (section Biomedicine)
- 2023 Listed among Top 50 most influential people in the Adria region by Bloomberg [https://ba.bloombergadria.com/businessweek-adria/glavna-tema/20114/igorstagljar/news]

Awards for non-academic accomplishments

Silver medal (2nd place) at the World Junior (U-19) Handball Championship with the Yugoslavian Junior National Handball Team, Gothenburg, Sweden (1984)

Gold medal (1st place) at the European Championship (U-19) with the Yugoslav Junior National Handball Team, Copenhagen, Denmark (1985)

H. Bibliography and Patents

Publications:	143 in PubMed, 150 publications in total
H-index	60 (Google Scholar)
Cited:	11,140
i10-index	117
Invited lectures:	> 250
Patents:	8

I. Ten most important Publications (since the move to the University of Toronto in 2005)

1. Paumi, C.M., Menendez, J., Arnoldo, A., Engels, K., Iyer, K., Thaminy, S., Georgiev, O., Barral, Y., Michaelis, S., and **Stagljar, I.** (2007) Mapping Protein-Protein Interactions for the Yeast ABC Transporter Ycf1p by Integrated Split-Ubiquitin Membrane Yeast Two-Hybrid (*i*MYTH) Analysis, *Molecular Cell* <u>26</u>, 15-25.

2. Snider, J., Hanif, A., Lee, M.E., Jin, K., Yu, A.R., Chuk, M., Damjanovic, D., Graham, C., Wierzbicka, M., Tang, P., Balderes, D., Wong, V., San Luis, B-J., Shevelev, I., Sturley, S.L., Boone, C., Babu, M., Zhang, Z., Paumi, C.M., Park, H-O., Michaelis, S., and **Stagljar, I.** (2013) A global analysis of the *Saccharomyces cerevisiae* ABC transporter interaction network: towards a greater understanding of membrane transport, *Nature Chemical Biology* <u>9</u>(9): 565-572.

3. Petschnigg, J., Groisman, B., Kotlyar, M., Taipale, M., Zheng, Y., Kurat, C.F., Sayad, A., J. Rafael Sierra, Mattiazzi Usaj, M., Snider, J., Nachman, A., Krykbaeva, I., Tsao, M-S., Moffat, J., Pawson, T., Lindquist, S., Jurisica, I. and **Stagljar, I.** (2014) The mammalian-membrane two-hybrid assay (MaMTH) for probing membrane-protein interactions in human cells, *Nature Methods* <u>11</u>, 585-592.

4. Yao, Z., Petschnigg, J., Ketteler, R., and **Stagljar, I.** (2015) Application Guide for OMICs Approaches to Cell Signaling, *Nature Chemical Biology* <u>11</u>, 387-397.

5. Yao, Z., Darowski, K., St-Denis, N., Wong, V., Offensperger, F., Villedieu, A., Amin, S., Malty, R., Aoki, H., Guo, H., Xu, Y., Iorio, C., Kotlyar, M., Emili, A., Jurisica, I., Babu, M., Neel, B.G., Gingras, A-C., and **Stagljar, I.** (2017) A global analysis of the receptor tyrosine kinase - protein phosphatase interactome, *Molecular Cell* <u>65</u>, 347-360. doi: 10.1016/j.molcel.2016.12.004. **Featured on the journal cover of the January 19, 2017 issue of** *Molecular Cell***.**

6. Saraon, P., Snider, J., Kalaidzidis, Y., Wybenga-Groot, L., E., Weiss, K., Rai, A., Radulovich, N., Drecun, L., Vučković, N., Vučetić, A., Wong, V., Thériault, B., Nhu-An Pham, Park, J.H., Datti, A., Wang, J., Pathmanathan, S., Aboualizadeh, F., Lyakisheva, A., Yao, Z., Wang, Y., Joseph, B., Aman, A., Moran, M.F., Prakesch, M., Poda, G., Marcellus, R., Uehling, D., Samaržija, M., Jakopović, M., Tsao, M-S., Shepherd, F.A., Sacher, A., Leighl, N., Akhmanova, A., Al-awar, R., Zerial, M., and **Stagljar, I.** (2020) A drug discovery platform to identify compounds that inhibit EGFR triple mutants, *Nature Chemical Biology* 16(5): 577-586. doi: 10.1038/s41589-020-0484-2

7. Yao, Z., Aboualizadeh F., Kroll, J., Akula, I., Snider, J., Lyakisheva, A. Tang, P., Kotlyar, M., Jurisica, I., Boxem, M., and **Stagljar, I.** (2020) Split Inte<u>in</u> Mediate<u>d</u> Prote<u>in</u> Ligation (SIMPL), a method for detecting protein-protein interactions and their inhibition, *Nature Communications* 2020 May 15;11(1): 2440. doi: 10.1038/s41467-020-16299-1

8. Yao, Z., Drecun, L., Aboualizadeh, F., Kim, S.J., Li, Z., Wood, H., Valcourt, E.J., Manguiat, K., Plenderleith, S., Yip, L., Li, X., Zhong, Z., Yue, F.Y., Closas, T., Snider, J., Tomic, J., Drews, S.J., Drebot, M.A., McGeer, A., Ostrowski, M., Mubareka, S., Rini, J.M., Owen, S., and **Stagljar, I.** (2021) A homogeneous split-luciferase assay for rapid and sensitive detection of anti-SARS CoV-2 antibodies, *Nature Communications* 2021 Mar 22;12(1):1806. doi: 10.1038/s41467-021-22102-6. *[Featured on the CBC's The National, CityNews TV, Global News, Toronto Star and The Globe and Mail]*

Lim, SH, Snider, J., Birimberg-Schwartz, L., Ip, W., Serralha, J.C., Botelho, H.M., Lopes-Pacheco, M., Pinto, M.C., Moutaoufik, M.T., Zilocchi, M., Laselva, O., Esmaeili, M., Kotlyar, M., Lyakisheva, A., Tang, P., López Vázquez, L., Akula, I., Aboualizadeh, F., Wong, V., Grozavu, I., Opacak- Bernardi, T., Yao, Z., Babu, M., Jurisica, I., Gonska, T., Bear, C.E, Amaral, M.D., **Stagljar, I.** (2022) CFTR Interactome Mapping Using the Mammalian Membrane Two-Hybrid High-Throughput Screening System, *Molecular Systems Biology* 18(2):e10629. doi: 10.15252/msb.202110629.
Featured on the journal cover of the February 2022 issue of *Molecular Systems Biology*.

10. Sun Jin Kim⁺, Zhong Yao⁺, Morgan C. Marsh, Debra M. Eckert, Michael S. Kay, Anna Lyakisheva, Maria Pasic, Aiyush Bansal, Chaim Birnboim, Prabhat Jha, Julio C. Delgado, Marc G. Elgort, Robert A. Campbell, Elizabeth A. Middleton, **Igor Stagljar**^{*}, Shawn C. Owen^{*} (2022) Homogeneous Surrogate Virus Neutralization Assay to Rapidly Assess Neutralization Activity of Anti-SARS-CoV-2 Antibodies, *Nature Communications* 13(1):3716 (* co-corresponding authors); DOI 10.1038/s41467-022-31300-9.

[Featured on the CTV News, CityNews TV, CP 24 News, Global News, Toronto Star and The Globe and Mail]

G. Peer-reviewed Publications

Level of Contribution Legend

SRA = Senior Responsible Author PA = Principal Author CSRA = Co-Senior Responsible Author C = Collaborator

(publications as a PI are at Reference #14 and after tenure at #27)

C 1. Amberg, D. C., Fleischmann, M., <u>Stagljar, I.</u>, Cole, C. N. and Aebi, M. (1993) Nuclear *PRP 20* protein is required for mRNA export. *EMBO J*. <u>12</u>, 233 -241.

PA 2. <u>Stagljar, I.</u>, te Heesen, S. and Aebi, M. (1994) New phenotype of mutations deficient in glucosylation of the lipid-linked oligosaccharide: cloning of the *ALG8* locus, *Proceedings of the National Academy of Sciences of the United States of America* <u>91</u>, 5977-5981.

C 3. Zufferey, R., Knauer, R., Burda, P., <u>Stagljar, I.</u>, te Heesen, S., Lehle, L. and Aebi, M. (1995) *STT3*, a highly conserved protein required for yeast oligosaccharyl transferase activity *in vivo*. *EMBO*

J. <u>14</u>, 4949-4960.

PA 4. <u>Stagljar, I.</u>, Aebi, M. and te Heesen, S. (1995) PCR-mediated cloning and sequencing of the *DmOST50* gene, a *WBP1/AvOST50/OST48* homologue, from *Drosophila melanogaster. Gene* <u>158</u>, 209-212.

PA 5. <u>Stagliar, I.</u>, Bourquin, J-P. and Schaffner, W. (1996) Use of the two-hybrid system and random sonicated DNA to identify the interaction domain of a protein. *Biotechniques* <u>21</u>, 430-432.

C 6. Fleischmann, M., <u>Stagljar, I.</u> and Aebi, M. (1996) Allele-specific suppression of a *prp20* mutation with altered protein phosphorylation by overexpression of a nuclear serine/threonine protein kinase. *Molecular Genetics and Genomics* 250, 614-625.

C 7. Tanner, S., <u>Stagljar, I.</u>, Georgiev, O., Schaffner, W. and Bourquin, J-P. (1997) A novel member of the serine-arginine domain (SR) proteins specifically interacts with the Carboxyl-terminal domain (CTD) of RNA polymerase II. *Biological Chemistry* <u>378</u>, 565-571.

C 8. Bourquin, J-P., <u>Stagljar, I.</u>, Meier, P., Moosmann, P., Silke, J., Baechi, T., Georgiev, O. and Schaffner, W. (1997) A serine/arginine-rich nuclear matrix cyclophilin interacts with the C-terminal domain of RNA polymerase II. *Nucleic Acids Research* <u>25</u>, 2055-2061.

PA 9. <u>Stagljar, I.</u>, Korostensky, C., Johnsson, N. and te Heesen, S. (1998) A new genetic system based on split-ubiquitin for the analysis of interactions between membrane proteins *in vivo*. *Proceedings of the National Academy of Sciences of the United States of America* <u>95</u>, 5187-5192.

C 10. Nayler, O., Fackelmeyer, F. O., Bourquin, J-P., <u>Stagliar, I.</u> Lindemann, L., Renz, A., Suchanek, C., Hartmann, A. M., Jasper, H. and Stamm, T. (1998) The nuclear matrix protein SAF-b couples chromatin to RNA transcription and processing, *Nucleic Acids Research* <u>26</u>, 3542-3549.

PA 11. <u>Stagljar, I.</u>, Hübscher, U. and Barberis, A. (1999) Activation of DNA replication in yeast by recruitment of the RNA polymerase II transcription complex, *Biological Chemistry* <u>380</u>, 525-530.

C 12. Stucki, M., <u>Stagljar, I.</u>, Jonsonn, Z. and Hübscher, U. (2000) A coordinated interplay: proteins with multiple functions in DNA replication, DNA repair and transcription, *Progress in Nucleic Acid Research and Molecular Biology* 65, 261-298.

PA 13. <u>Stagljar, I.</u> and te Heesen, S. (2000) Detecting interactions between membrane proteins *in vivo* using chimeras, *Methods in Enzymology* <u>327</u>, 190-198.

SRA 14. Pedrazzi, G., Perrera, C., Blaser, H., Kuster, P., Marra, G., Freire, R., Ryu, G-H., Jiricny, J. and <u>Stagliar, I.</u> (2001) Direct association of Bloom's syndrome gene product with the human mismatch repair protein MLH1, *Nucleic Acids Research* **29**, 4378-4386.

C 15. Freire, R., d'Adda di Fagagna, F., Wu, L., Pedrazzi, G., <u>Stagljar, I.</u>, Hickson, I.D., and Jackson, S.P. (2001) Cleavage of the Bloom's syndrome gene product during apoptosis by caspase-3 results in an impaired interaction with topoisomerase III alpha, *Nucleic Acids Research* **29**, 3172-3180.

C 16. Gisler, S., Stagljar, I., Traebert, M., Bacic, D., Biber, J., Murer, H. (2001) Interaction of the type

IIa Na-Pi-cotransporter with PDZ proteins, Journal of Biological Chemistry 276, 9206-9213.

C 17. Reinders, A., Schultze, W., Thaminy, S., <u>Stagljar, I.</u>, Fromer, W.B., and Ward, J.M. (2002) Intra- and intermolecular interactions in sucrose transporters at the plasma membrane detected by the split-ubiquitin system and functional assays, *Structure* **10**, 763-772.

C 18. Scheper, W., Thaminy, S., Kais, S., <u>Stagljar, I.</u>, and Römisch, K. (2003) Coordination of Nglycosylation and protein translocation across the ER membrane in yeast by Sss1 protein, *Journal of Biological Chemistry* <u>278</u>, 37998-38003.

SRA 19. Pedrazzi, G., Bachrati, C., Selak, N., Studer, I., Hickson, I.D., Jiricny, J., and <u>Stagljar, I.</u> (2003) The Bloom's syndrome helicase directly interacts with the human mismatch protein MSH6, *Biological Chemistry*, <u>384</u>, 1155-1164.

SRA 20. Thaminy, S., Auerbach, D., Arnoldo, A., and <u>Stagljar, I.</u> (2003) Identification of novel ErbB3-interacting proteins using the split-ubiquitin membrane yeast two-hybrid technology, *Genome Research* <u>13</u>, 1744-1753.

C 21. Yang, Q., Zhang, R., Wang, X.W., Linke, S.L., Sengupta, S., Hickson, I.D., Pedrazzi, G., Perrera, C., <u>Stagljar, I.</u>, Littman, S.J., Modrich, P., and Harris, C.C. (2004) The mismatch repair heterodimer, hMSH2/6, regulates BLM helicase activity in homologous DNA recombination, *Oncogene* <u>23</u>, 3749-3756.

SRA 22. Jiao, R., Bachrati, C., Pedrazzi, G., Kuster, P., Petkovic, M., Li, J-L., Egli, D., Hickson, I.D., and <u>Stagliar, I.</u> (2004) Physical and functional interaction between the Bloom's syndrome gene product and CAF-1 hp150, the largest subunit of chromatin assembly factor 1, *Molecular & Cellular Biol*ogy <u>24</u>, 4710-4719.

SRA 23. Iyer, K., Bürkle, L., Auerbach, D., Thaminy, S., Dinkel, M., Engels, K., and <u>Stagljar, I.</u> (2005) Utilizing the yeast split ubiquitin membrane yeast two-hybrid system (MbYTH) to identify interacting proteins of membrane proteins, *Science STKE* <u>275</u>, pl3.

C 24. Miller, J., Lo, R., Desmarais, C., <u>Stagljar, I.</u>, Noble, W., and Fields, S. (2005) An interaction network of yeast integral membrane proteins, *Proceedings of the National Academy of Sciences of the United States of America* <u>102</u>, 12123-12128.

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Level of Contribution for Reviews

P = Primary contributor E = Equal contributor C = Contributed some sections of review

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J. PATENTS

1. Method and kit for detecting membrane protein-protein interactions

Publication number: 20050106636

Abstract: The present invention is concerned with a method and a kit for detecting an interaction between a first membrane bound test protein or fragment thereof and a second test protein or fragment thereof which is either membrane bound or soluble with an in vivo genetic system based in yeast, bacterial or mammalian cells. The system makes use of the reconstitution of the split ubiquitin protein.

Type: grant

Filed: March 28, 2003

Publication date: May 19, 2005

Inventors: Igor Stagljar, Michael Hottiger, Daniel Auerbach

2. Method and kit for detecting membrane protein-protein interactions

Patent number: 9435055

Abstract: The present invention is concerned with a method and a kit for detecting an interaction between a first membrane bound test protein or fragment thereof and a second test protein or fragment thereof which is either membrane bound or soluble with an in vivo genetic system based in yeast, bacterial or mammalian cells. The system makes use of the reconstitution of the split ubiquitin protein.

Type: Grant

Filed: March 28, 2003

Date of Patent: September 6, 2016

Assignee: Hybrigenics Services S.A.S.

Inventors: Igor Stagljar, Michael Hottiger, Daniel Auerbach

3. Methods and compositions for inhibiting mutant EGFR signaling

Publication number: 20190091205

Abstract: Methods of inhibiting mutant EGFR and methods of treating a subject afflicted with a lung cancer having a mutant EGFR, having for example a C797 mutation, are described. The methods comprise administering to a cell or a subject in need thereof a therapeutically effective amount of a compound selected from 3-(1,3-benzoxazol-2-yl)-7-(diethylamino)-2H-chromen-2-one and a structurally related analog thereof; midostaurin; and AZD7622 and a structurally related analog thereof. Compositions and combinations comprising the compounds of the disclosure as well as uses are also provided.

Type: grant

Filed: September 28, 2018

Publication date: March 28, 2019

Inventors: Igor Stagljar, Jamie Snider, Punit Saraon

4. Detection of protein to protein interactions

Publication number: 20160124000

Abstract: The present invention relates to methods and kits for detecting the interaction between a first membrane protein and a second protein in mammalian cells as well as to identify molecules that can disrupt protein to protein interactions. The invention relies on the functional reconstitution of an active human ubiquitin by two inactive fragments upon the interaction of two proteins attached through a linker to the inactive fragments. The reconstituted ubiquitin is then cleaved by human ubiquitin proteases resulting in the release of an artificial transcription factor, which in turn activates a reporter gene transcription. Activation of the reporter gene is indicative of the interaction between the two proteins.

Type: grant

Filed: June 10, 2014

Publication date: May 5, 2016

Inventors: Igor STAGLJAR, Julia PETSCHNIGG, Bella GROISMAN

5. Detection of protein to protein interactions

Patent number: 9958461

Abstract: The present invention relates to methods and kits for detecting the interaction between a first membrane protein and a second protein in mammalian cells as well as to identify molecules that can disrupt protein to protein interactions. The invention relies on the functional reconstitution of an active human ubiquitin by two inactive fragments upon the interaction of two proteins attached through a linker to the inactive fragments. The reconstituted ubiquitin is then cleaved by human ubiquitin proteases resulting in the release of an artificial transcription factor, which in turn activates a reporter gene transcription. Activation of the reporter gene is indicative of the interaction between the two proteins.

Type: grant

Filed: June 10, 2014

Date of Patent: May 1, 2018

Assignee: THE GOVERNING COUNCIL OF THE UNIVERSITY OF TORONTO

Inventors: Igor Stagljar, Julia Petschnigg, Bella Groisman

6. <u>Detection of protein to protein interactions: new split intein-based method for detecting</u> protein-protein interactions

Patent number: 0180354.0066

Abstract: A system and method for detecting interactions between a first protein or fragment thereof (bait protein) and a second protein or fragment thereof (prey protein) comprising: (a) a bait construct comprising the bait protein, a first epitope tag and an intein N-terminal fragment (IN); and (b) a prey construct comprising the prey protein, a second epitope tag, and an intein C-terminal fragment (IC).

Type: pending

International Filing Date: June 30, 2020

Assignee: THE GOVERNING COUNCIL OF THE UNIVERSITY OF TORONTO

Inventors: Igor Stagljar and Zhong Yao

7. System and method for rapid and sensitive detection of anti-pathogen antibodies

Patent number: PCT/CA2021/051733

Type: pending

Assignee: THE GOVERNING COUNCIL OF THE UNIVERSITY OF TORONTO

Inventors: Igor Stagljar and Zhong Yao

8. Neutralization Assay to Rapidly Assess Neutralization Activity of Anti-SARS-CoV-2 Antibodies

Disclosure reference: 10004321

Type: pending (entered on 26-04-2022)

Assignee: THE GOVERNING COUNCIL OF THE UNIVERSITY OF TORONTO

Inventors: Sun Jin Kim, Shawn Owen, Igor Stagljar and Zhong Yao

K. Invited Presentations (Oral) at Meetings and Symposia (last six years)

1. 12th Annual International Bellairs Confrence "The Cell Biology Of Disease", Jan 29-Feb 6, 2016, Holetown, Barbados

2. EMBO Conference "Cellular Signaling and Cancer Therapy", May 27-31, 2016, Cavtat, Croatia

3. 8th International Conference "From Solid State to Biophysics VIII - From Basic to Life Sciences", June 5-12, 2016, Cavtat, Croatia

4. 15th Human Proteome Organization (HUPO) 2016 World Congress (*HUPO 2016*), September 18-22, 2016, Taipei, Taiwan (Igor Stagljar was the keynote speaker)

5.. International Conference organized by the EU Scientific Council "Oncogenic EGFR Signaling", Palma de Mallorca, Spain, October 1-5, 2016

6. The University of Ulm Fall Meeting "Advances in Molecular Biology", Ulm, Germany, October 12-14, 2016

7. 13th International Conference on "Cell and Systems Biology of Disease", Holetown, Barbados, January 13-19, 2017

8. International Conference "Advances in Biomedical Research", July 3-7, 2017, Split, Croatia (Igor Stagljar was one of the co-organizers of this conference)

9. International conference "Transporters and channels in drug discovery and preclinical development", August 6-10, 2017, Lausanne, Switzerland

10. 6th International Cancer Therapeutics & Partnering Summit, September 28-29, 2017, San Diego, CA, USA

11. 14th International Conference "Cell and Systems Biology of Disease", January 20-26, 2017, Holetown, Barbados

12. International Symposium "Commercialization in science", March 8, 2018, Zagreb, Croatia (Igor Stagljar was a keynote speaker)

13. EMBL Conference "Integrating Systems Biology: From Networks to Mechanisms to Models", April 15-17, 2018, Heidelberg, Germany

14. 9th International Conference "From Solid State to Biophysics IX - From Basic to Life Sciences", June 16-23, 2018, Cavtat, Croatia

15. EMBO Conference "Cellular Signaling and Cancer Therapy", Cavtat, Croatia, September 14-18,

2018

16. International Conference 17th Ruzicka days – Today science, tomorrow Industry", Vukovar, Croatia, September 19-21, 2018 (Igor Stagljar will be a plenary speaker)

17. 3th Croatian Biological Society Congress, Poreč, Croatia, September 19-23, 2018 (Igor Stagljar will be a plenary speaker)

18. Human Proteome Organization (HUPO) conference, Orlando, FL, USA, September 30-October 3, 2018 (Igor Stagljar was the chairman of the "Systems Biology" session)

19. 15th International Conference "Cell and Systems Biology of Disease", January 20-26, 2019, Holetown, Barbados

20. International Conference "Success stories in science", March 28-30, 2019, Poreč, Croatia

21. International Conference "Advances in Biomedical Research III", Split, Croatia, June 17-21, 2019

22. 10th Congress of the Croatian Thoracic Society "Toraks 2020", Zagreb, Croatia, May 2020

23. EMBO Conference "Systems approaches in cancer", September 21-26, 2021 Split, Croatia

24. International Conference "Advances in Biomedical Research IV", Split, Croatia, October 4-9, 2021

- 25. Life Science LS2 Swiss Biomedical Society, Zurich, Switzerland, April 20-22, 2022
- 26. Gordon Research Conference "FGF Signaling: Understanding Function and Devising Therapeutic Tools", May 1-6, 2022, Lucca, Italy

27. TransCure Final Conference "Excellence in Transmembrane Protein Research", Aug 17-19, 2022, Berne, Switzerland https://www.nccr-transcure.ch/events/final-conference

mips.//www.ncci-iranscure.cn/events/mai-conference

28. EMBO New Members Symposium, October 26-28, 2022, Heidelberg, Germany https://www.embo.org/press-releases/embo-elects-67-new-members-and-associate-members/

29. International conference "Women Who are Changing Science - Driving Bold Research in Canada and Croatia" symposium, Zagreb and Split, Croatia, September 12-14, 2022 <u>https://wis.medils.hr/</u>

30. CCC-TRIO 2023 New Frontiers in Translational Oncology & Immuno-Oncology, April 27-28, 2023, Vienna, Austria

https://www.meduniwien.ac.at/web/fileadmin/content/kommunikation/events/2023/04/CCC_Trio_2023 /Einladung_CCC_Trio_2023_Web_LV5.pdf

31. 14th Annual CNPN Symposium (CNPN 2023), May 15 – 17, 2023, University of Regina, Saskatchewan (I was a keynote speaker) https://cnpn.ca/wp-cnpn/wp-content/uploads/2023/02/Preliminary-Program-for-Web-2023-02-02.pdf

32. 13th Congress of the Croatian Thoracic Society "Toraks 2023", Zagreb, Croatia, May 2023 <u>https://web.penta-pco.com/toraks2023/en/teme/</u>

33. EMBO Conference "Systems approaches in cancer", June 26-30, 2023 Split, Croatia https://meetings.embo.org/event/23-sys-approaches-cancer

34. International Conference "Advances in Biomedical Research V", Split, Croatia, September 15-19, 2023 https://conference.medils.hr/

L. Organization of Meetings and Symposia

1. International Conference "Functional Genomics in Medicine", Split, Croatia, June 2-8, 2007

2. Keystone Conference "OMICs meets cell biology: applications to health and disease", Taos, NM, USA, Feb 18-23, 2014

- 3. International Conference "Molecular Perspectives on Protein-Protein Interactions", Pillar and Post Inn, Niagara-on-the-lake, Ontario, Canada, May 29 - June 2, 2015
- 4. International Conference "OMICs in Biomedical Research", Split, Croatia, June 8-12, 2015
- 5. International Conference "From Solid State to Biophysics VIII From Basic to Life Sciences", June 5-12, 2016, Cavtat, Croatia
- 6. International BBRC Symposium "Trends in Molecular Biology", October 7-8, 2016, Toronto, Canada.
- 7. International Conference "Advances in Biomedical Research II", Split, Croatia, July 3-7, 2017
- 8. International Conference "From Solid State to Biophysics IX From Basic to Life Sciences", June 16-23, 2018, Cavtat, Croatia
- 9. International Conference "Advances in Biomedical Research III", Split, Croatia, June 17-21, 2019
- 10. EMBO Conference "Systems approaches in cancer", September 21-26, 2021 Split, Croatia
- 11. International Conference "Advances in Biomedical Research IV", Split, Croatia, October 4-9, 2021
- 12. International conference "Women Who are Changing Science Driving Bold Research in Canada and Croatia" symposium, Zagreb and Split, Croatia, September 12-14, 2022
- 13. EMBO Conference "Systems approaches in cancer", June 26-30, 2023 Split, Croatia

14. International Conference "Advances in Biomedical Research V", Split, Croatia, September 15-19, 2023

https://conference.medils.hr/

L. Invited Lectures (last seven years)

- 1. University of California San Diego, Skaggs School of Pharmacy, La Jolla, CA, USA, March 2016
- 2. The Salk Institute, La Jolla, CA, USA, March 2016
- 3. The Scripps Institute, La Jolla, CA, USA, March 2016
- 4. Genomics Institute of the Novartis Research Foundation, La Jolla, CA, March 2016
- 5. Janssen Pharmaceutical, La Jolla, CA, March 2016

6. Novartis Pharma, Basel, Switzerland, April 2016

7. University of Zurich, Department of Molecular Mechanisms of Disease, Zurich, Switzerland, April 2016

8. European Molecular Biology Laboratory (EMBL), Heidelberg, Germany, May 2016

9. University of Frankfurt, Department of Biochemistry, Frankfurt, Germany, May 2016

10. University of Zagreb, Department of Molecular Biology, Zagreb, Croatia, May 2016

11. Touro University, The School of Medicine, Las Vegas, NV, USA, November 2016

12. University Health Network (UHN), Princess Margaret Cancer Centre Seminar Series, Toronto, Canada, November 2016

13. University of Zagreb School of Medicine, Department of Pharmacology Zagreb, Croatia, December 2016

14. Memorial Sloan Kettering Cancer Center, The Molecular Pharmacology & Chemical Biology Programs Research Seminar Series, New York, NY, USA, March 2017

15. University of Montreal, Department of Immunology, April 2017

16. New York University, Department of Chemistry, New York, NY, USA, November 2017

17. University of Zagreb School of Medicine, Department of Pharmacology, Zagreb, Croatia, December 2017

18. University of California Riverside School of Medicine, Division of Biomedical Sciences, Riverside, CA, USA, February 2018

19. Genentech, San Francisco, CA, February 2018

20. Novartis, Cancer Therapeutics group, Basel, Switzerland, April 2018

21. University of Utrecht, Cancer Stem Cells & Developmental Biology, Utrecht, The Netherlands, September 2018.

22. University of Puerto Rico, Comprehensive Cancer Centre, San Juan, Puerto Rico, November 2018

- 23. McGill University, Goodman Cancer Centre, Montreal, QC, Canada, November 2018
- 24. University of Zagreb School of Medicine, Zagreb, Croatia, December 2018
- 25. Novartis Institutes for Biomedical Research, Cambridge, MA, USA, May 2019
- 26. University of Graz, Department of Genetics, Austria, July 2019
- 27. University of Ljubljana, Department of Biotechnology, July 2019
- 28. University of Zurich, Department of Pathology, July 2019
- 29. Max Delbrück Center for Medical Research, Humbolt Foundation, Berlin, September 2019
- 30. ETH Zurich, Department of Biochemistry, September 2019
- 31. Mount Sinai Hospital, The Tisch Cancer Institute, New York, NY, USA, October 2019

32. University College Dublin, Conway Institute of Biomolecular & Biomedical Research, December 2019

33. The University of Southwestern Medical Center, O'Brien Cancer Center, Dallas, TX, January 2020

- 34. The San Juan Cancer Centre, San Juan, Puerto Rico, March 2020
- 35. University of Split, Medical School, Split, Croatia, October 2021
- 36. University of Ponce Medical School, Ponce, Puerto Rico, February 2022
- 37. University of San Juan, School of Medicine, San Juan, Puerto Rico, February 2022
- 38. Swiss Federal School of Technology EPFL Lausanne, AGORA Cancer Centre, April 2022
- 39. University of Split, Medical School, Split, Croatia, October 2022

40. European Molecular Biology Organization (EMBO), inauguration talk (EMBO member), Heidelberg, Germany, October 2022

- 41. Department of Molecular and Cell Biology, University of Geneva, Switzerland, February 2023
- 42. Novartis Pharma, Basel, Switzerland, March 2023
- 43. Paul Scherrer Institute, Department of Biology, Villingen, Switzerland, March 2023

44. University of Bern, Department of Chemistry, Biochemistry and Pharmaceutical Sciences, April 2023

M. Research Grant Support

As principal investigator, I secured over US\$55 million in research funds from a range of national and international funding agencies, non-profit foundations and pharmaceutical companies, such as Orionis Biosciences, Novartis, Genentech, Merck, Evotec and Novartis.

N. Mentorship/Supervision

During his career Professor Stagljar has trained more than 25 graduate students and 15 postdoctoral fellows. Many of them are now having executive roles in pharmaceutical companies and some of them have successful careers in the academia. Below are the names and current positions of some of my previous lab members.

Dr. Tobias Dietschy received PhD in my lab 2007 – he is the Owner & Sales Director at LubioScience GmbH in Zurich, Switzerland.

Dr. Bernhard Suter worked as a postdoc in my lab from 2006-2008. He is the founder and CEO of Next Interactions (San Francisco-based biotech that employs 400 people).

Dr. Uros Petrovic worked as a postdoc in my lab from 2010-2011. He is an Associate Professor at the University of Ljubljana, Slovenia.

Dr. Saranya Kittanakom-Arnoldo worked as a postdoc in my lab from 2007-2010. She is a Clinical Biochemist at the London Health Sciences Centre, an Assistant Professor at the department of Pathology and Laboratory Medicine, Western University (London, ON, Canada), and a fellow of the Canadian Academy of Clinical Biochemistry.

Dr. Jasna Curak received PhD in my lab in 2010. She is the Global Compliance and External Collaboration Manager at Roche, Toronto, Canada

Dr. Julia Petschnigg worked as a postdoc in my lab from 2009-2014. She is the Lecturer (Assistant Professor) at the University of Munich, Germany.

Dr. Kate Sokolina received PhD in my lab in 2016. She is a senior consultant at Calico (Google's biotech company).

Dr. Punit Saraon worked as a postdoc in my lab from 2014-2019. He is a Research Scientist (group leader) at the Ontario Institute for Cancer Research (OICR) in Toronto, Canada.

Dr. SangHyun Lim received PhD in my lab in 2021 He is now a postdoctoral fellow at Genentech (Roche group) in South San Francisco, USA.