Curriculum Vitae

ANDREW SEEBER, PhD

aseeber@fas.harvard.edu

Date of Birth: 28 October 1987 Citizenship: South Africa/Ireland Civil Status: Married to Michelle DiPietro (USA) Residency: USA PRESENT POSITION Center for Advanced Imaging Cambridge, USA Harvard University 9.2018 - present John Harvard Distinguished Science Fellow in Imaging My laboratory investigates chromatin dynamics and function **EDUCATION** Friedrich Miescher Institute for Biomedical Research Basel, CH 11.2011 - 1.2017University of Basel PhD, Genetics, summa cum laude Dissertation: "Chromatin dynamics in DNA double-strand break repair" University of Geneva Geneva/Basel, CH NCCR Frontiers in Genetics PhD rotation program 10.2010 - 11.2011Hong Kong University of Science and Technology Kowloon, HK NUIG Funded Exchange Fellowship 1.2009 - 7.2009National University of Ireland, Galway Galway, IE Bachelor of Biomedical Science, 1st Class Hons 9.2006 - 6.2010RESEARCH EXPERIENCE **Biozentrum Technology Venture** Basel, CH University of Basel 3.2018 - 7.2018SNF BRIDGE and Gebert Rüf Stiftung Independent Researcher Improving CRISPR-Cas9 efficiency by modulating chromatin structure Center for Applied Biotechnology and Molecular Medicine, Zurich, CH University of Zurich 8.2017 - 2.2018SNF BRIDGE Independent Researcher Hosted by Matthias Altmeyer Improving CRISPR-Cas9 efficiency by modulating chromatin structure Friedrich Miescher Institute for Biomedical Research Basel, CH

11.2011 - 6.2017

- Developed an imaging regime to study chromatin dynamics in living cells

- Identified the biological mechanism that drives chromatin movement

PhD student; Advisor: Susan M. Gasser

Chromatin dynamics in DNA double-strand break repair

Characterized how sister chromatids are held together at double-strand breaks

Advanced Imaging Center, Janelia Research Campus Ashburn, USA Janelia Research Campus visitor program 8.2015 - 9.2015PhD student; Advisor: Teng-Leong Chew Applying aberration-corrected multifocus microscopy to chromatin dynamics in budding yeast **Cold Spring Harbor Laboratory** NY, USA Yeast Genetics and Genomics course 7.2011 - 8.2011Advisors: Prof. Jeffery Strathern and Prof. Jeffery S. Smith Intensive training course in the use of S. cerevisiae as a model organism Friedrich Miescher Institute for Biomedical Research Basel, CH 3rd Rotation, NCCR Frontiers in Genetics 5.2011 - 11.2011Advisor: Prof. Susan M. Gasser Interaction of the MRX complex with RPA at DSBs and stalled replication forks University of Geneva Geneva, CH 2nd Rotation, NCCR Frontiers in Genetics 2.2011 - 5.2011Advisor: Prof. David Shore Sir protein interaction at an irreparable DSB Geneva, CH University of Geneva 1st Rotation, NCCR Frontiers in Genetics 10.2010 - 2.2011Advisor: Prof. Robbie Loewith Investigation of downstream targets of yeast protein kinase National University of Ireland, Galway Galway, IE Science Foundation Ireland Student Fellowship 5.2010 - 9.2010Advisor: Prof. Noel F. Lowndes Investigation of the target residues of the tudor domains of 53Bp1 and Crb2 Diaceutics, International Management and Consulting Firm Galway, IE Advisor: Patrick Considine, PhD 6.2010 - 9.2010Creation of a database of patents, products and services of diagnostic companies National University of Ireland, Galway Galway, IE Final Year Laboratory Project 9.2009 - 1.2010Advisor: Prof. Andrew Flaus Recombinant expression and purification of histone H1.2 and its effect on a DNA-bound nucleosome National University of Ireland, Galway Galway, IE Biochemical Society Summer Studentship 6.2009 - 9.2009

Creation and recombinant expression of a trimeric mutant of replication protein A, in a single vector with a

Advisor: Prof. Heinz-Peter Hasheuer

triple alanine mutation on the p32 subunit

SKILLS AND TECHNIQUES

- Genetic manipulations in budding yeast and human cells
- Chromatin immunoprecipitation and quantitative PCR
- Protein microarrays
- Co-immunoprecipitation of protein complexes

- Fluorescence microscopy
 - o LSM and spinning disk microscopy
 - o Structured illumination (SIM), PALM
 - o FRET and FRAP
 - o High content screening
- Image processing and analysis of large image datasets (ImageJ, KNIME, MATLAB, ICY)

LEADERSHIP EXPERIENCE

Friedrich Miescher Institute for Biomedical Research

Basel, CH 2014-2016

Graduate student advisor and microscopy expert

Trained multiple graduate students in a range of techniques including live cell and super resolution microscopy and the use of S. cerevisiae as a model organism

The University of Hong Kong	Pok Fu Lam, HK
Anatomy demonstrator and prepared prosections	1.2009 - 5.2009
Assisted in gross anatomical and histological laboratory practical sessions	

National University of Ireland, Galway	Galway, IE
Vice Auditor of the Galway Film Society	2008 - 2009

National University of Ireland, Galway	Galway, IE
Student Support Worker	7.2007 - 5.2010

Tutoring students with learning disabilities (dyslexia) in biochemistry

GRANTS AND AWARDS

Gebert Rüf Foundation (388,649 CHF)	10.2017
SNF BRIDGE fellowship (130,000 CHF)	8.2017
Carl Singer Foundation award	8.2014
Biochemical Society travel grant	9.2011
Swiss Society for Biochemistry grant for junior scientists	8.2011
NCCR – Frontiers in Genetics, travel grant	8.2011
Science Foundation Ireland student fellowship	5.2010 - 9.2010
Biochemical Society Summer studentship	6.2009 - 9.2009
NUIG fellowship to Hong Kong University of Science and Technology	1.2009 - 7.2009

PROFESSIONAL ASSOCIATIONS

Member of the Biochemical Society (UK) 2009 – present

INVITED CONFERENCE TALKS

Multiscale analysis and reconstruction of chromatin and nuclear organization, Pisa	10.2018
"Chromosome Dynamics in Response to DNA Damage"	

'Using single-

Research Center for the Mathematics on Chromatin Live Dynamics, Hiroshima

particle trajectory statistics and polymer simulations to analyze and predict changes in chromatin structure"	11.2016
39th Annual Meeting of the Molecular Biology Society of Japan "Change in local chromatin structure during homology search"	11.2016
At the Intersection of DNA Replication and Genome Maintenance, ICGEB Trieste "RPA recruits MRX to forks and breaks to hold replicated sister chromatids together"	6.2016
International Centre for Theoretical Physics Chromatin Workshop, Trieste "Chromatin (modifiers) and the dynamic chromatin response to DNA damage"	9.2014
FASEB, Yeast Chromosome Structure, Replication and Segregation "Stable interaction between MRX and RPA is required for stalled fork recovery and maintenance of DSB structure for repair"	7.2014
TriRhena transcription and chromatin club "Checkpoint kinases regulate global chromatin mobility after DSB induction"	2.2013
Nucleosome 4D Barcelona Retreat "How checkpoint kinases influence global chromatin dynamics"	10.2012
Image DDR meeting, Sussex "How checkpoint kinases influence global chromatin dynamics"	9.2012
NCCR – Frontiers in Genetics Annual Meeting "How checkpoint kinases influence global chromatin dynamics"	6.2012

PATENTS

Inventors: Seeber, A., Hauer, M., Gasser, S.M.,

"Methods for increasing the frequency of gene targeting by chromatin modification"

PEER-REVIEWED PUBLICATIONS

- 1) Chen, J., Young, S.M., Allen, C., **Seeber, A.**, Péli-Gulli, M.-P., Panchaud, N., Waller, A., Ursu, O., Yao, T., and Golden, J.E. (2012). Identification of a small molecule yeast TORC1 inhibitor with a multiplex screen based on flow cytometry. <u>ACS Chemical Biology</u> 7, 715-722.
- 2) Shimada, K., Filipuzzi, I., Stahl, M., Helliwell, S.B., Studer, C., Hoepfner, D., **Seeber, A.**, Loewith, R., Movva, N.R., and Gasser, S.M. (2013). TORC2 signaling pathway guarantees genome stability in the face of DNA strand breaks. <u>Molecular Cell</u> 51, 829-839.
- 3) Dion, V., Kalck, V., **Seeber, A.**, Schleker, T., and Gasser, S.M. (2013). Cohesin and the nucleolus constrain the mobility of spontaneous repair foci. EMBO reports 14, 984-991.
- 4) **Seeber, A.**, Dion, V., and Gasser, S.M. (2013). Checkpoint kinases and the INO80 nucleosome remodeling complex enhance global chromatin mobility in response to DNA damage. <u>Genes & Development</u> 27, 1999-2008.

- 5) Hustedt, N., **Seeber, A.**, Sack, R., Tsai-Pflugfelder, M., Bhullar, B., Vlaming, H., van Leeuwen, F., Guénolé, A., van Attikum, H., and Srivas, R. (2015). Yeast PP4 interacts with ATR homolog Ddc2-Mec1 and regulates checkpoint signaling. <u>Molecular Cell</u> 57, 273-289.
- 6) Poli, J., Gerhold, C.-B., Tosi, A., Hustedt, N., **Seeber, A.**, Sack, R., Herzog, F., Pasero, P., Shimada, K., and Hopfner, K.-P. (2016). Mec1, INO80, and the PAF1 complex cooperate to limit transcription replication conflicts through RNAPII removal during replication stress. <u>Genes & Development</u> 30, 337-354.
- 7) **Seeber, A.**, Hegnauer, A.M., Hustedt, N., Deshpande, I., Poli, J., Eglinger, J., Pasero, P., Gut, H., Shinohara, M., Hopfner, K.P., Shimada, K., Gasser, S.M. (2016) RPA Mediates Recruitment of MRX to Forks and Double-Strand Breaks to Hold Sister Chromatids Together. <u>Molecular Cell</u> 64, 951-966.
- 8) Hauer, M., **Seeber, A.**, Singh, V., Thierry, R., Amitai, A., Kryzhanovska, M., Eglinger, J., Holcman, D., Owen-Hughes, T. (2017) Histone degradation in response to DNA damage enhances chromatin dynamics and recombination rates. <u>Nature Structural and Molecular biology</u> 24, 99-107.
- 9) Amitai, A.*, **Seeber, A.***, Gasser, S.M., Holcman, D. (2017) Single particle trajectory statistics and polymer simulations model and accurately predict changes in chromatin structure at DNA breaks. <u>Cell Rep</u> 18, 1200-1214.

 *equal contribution
- 10) Deshpande, I., **Seeber, A.**, Shimada, K., Keusch. J., Gut, H., Gasser, S.M. (2017) Structural basis of Mec1-Ddc2-RPA assembly and activation on single-stranded DNA at sites of damage. Mol Cell, 68, 431-445
- 11) Marcomini, I, Shimada, K, Delgoshaie, N, Yamamoto, I, **Seeber, A**, Cheblal, A, Horigome, C, Naumann, U, Gasser, S.M. (2018) Asymmetric Processing of DNA Ends at a Double-Strand Break Leads to Unconstrained Dynamics and Ectopic Translocation. <u>Cell Rep</u>, 24, 2614-2628
- 12) Cheblal, A., **Seeber, A.**, Gasser, S.M. Kinetochore detachment or declustering is not responsible for damage induced chromatin movement. *In preparation*.
- 13) Shimda, K., Gerhold, C.B., Yamasaki, S., Hurst, V., **Seeber, A.**, Filipuzzi, I., Stahl, M., Bodenmiller, B., Helliwell, S.B., Knapp, B., Loewith, R., Movva, R., Harata, M., Gasser, S.M. Modulation of actin distribution and polymerization dynamics impairs DNA damage repair. *In preparation*.
- 14) Poli, J., Barthe, A., Tittel-Elmer, M., Soutourina, J., **Seeber, A.**, Dubrana, K., Wery, M., Werner, M., Cobb, J., Gasser, S.M., Pasero, P., Legronne, A. MRX and mediator limit pervasive transcription through chromatin compaction at the nuclear pore complex. *In preparation*.

 *equal contribution

REVIEW ARTICLES AND BOOK CHAPTERS

- 1) Chen, J., Young, S.M., Allen, C., Waller, A., Ursu, O., Strouse, J.J., Yao, T., Golden, J.E., Peterson, B.R., Foutz, T.D.,... Seeber, A., et al. (2010). Profiling a Selective Probe for RTG Branch of Yeast TORC1 Signaling Pathway. Probe Reports from the NIH Molecular Libraries Program
- 2) **Seeber, A.**, Hauer, M., and Gasser, S.M. (2013). Nucleosome remodelers in double-strand break repair. <u>Current Opinion in Genetics & Development</u> 23, 174-184.
- 3) **Seeber, A.**, Dion, V., and Gasser, S.M. (2014). Remodelers move chromatin in response to DNA damage. <u>Cell Cycle</u> 13, 877-878.
- 4) Horigome, C., Dion, V., **Seeber, A.**, Gehlen, L.R., and Gasser, S.M. (2015). Visualizing the spatiotemporal dynamics of DNA damage in budding yeast. <u>Methods in Molecular Biology</u> (Clifton, NJ) 1292, 77-96.
- 5) **Seeber, A.**, and Gasser, S.M. (2017). Chromatin organization and dynamics in double-strand break repair. <u>Current Opinion in Genetics & Development</u> 43, 9-16.
- 6) **Seeber, A.**, Hauer, M., Gasser, S.M., (2018). Chromosome Dynamics in Response to DNA Damage. <u>Annual Review of Genetics</u> 52.

REFERENCES

Susan M. Gasser, PhD Director of FMI Professor of Molecular Biology Quantitative Biology FMI Maulbeerstrasse 66 4058 Basel Switzerland susan.gasser@fmi.ch Noel F. Lowndes, PhD Professor of Biochemistry Biochemistry NUI, Galway Galway Ireland noel.lowndes@nuigalway.ie Vincent Dion, PhD SNF Professor CIG University of Lausanne Lausanne Switzerland vincent.dion@unil.ch