

Remote control: the cancer cell-intrinsic mechanisms that dictate systemic inflammation and anti-tumor immunity Wellenstein, M.D.

Citation

Wellenstein, M. D. (2021, March 25). *Remote control: the cancer cell-intrinsic mechanisms that dictate systemic inflammation and anti-tumor immunity*. Retrieved from https://hdl.handle.net/1887/3152435

Version:	Publisher's Version
License:	<u>Licence agreement concerning inclusion of doctoral thesis in the</u> <u>Institutional Repository of the University of Leiden</u>
Downloaded from:	https://hdl.handle.net/1887/3152435

Note: To cite this publication please use the final published version (if applicable).

Cover Page



Universiteit Leiden



The handle <u>https://hdl.handle.net/1887/3152435</u> holds various files of this Leiden University dissertation.

Author: Wellenstein, M.D. Title: Remote control: the cancer cell-intrinsic mechanisms that dictate systemic inflammation and anti-tumor immunity Issue Date: 2021-03-25

Curriculum vitae

Max Daniël Wellenstein was born on December 2nd 1989 in Voorburg. He graduated preuniversity secondary education (Gymnasium) in 2008 from Dalton Voorburg, where he developed a fascination for the molecules that govern life. In 2009 he started his studies of Biology at Utrecht University (UU), obtaining his BSc. degree in 2012. During this time, he worked as a research assistant at the department of Molecular Plant Physiology (UU) with Dr. Henriette Schlüpmann, examining metabolic pathways in Arabidopsis thaliana plants. He also worked on microtubule dynamics as a research intern at the department of Cell Biology (UU) with Prof. Anna Akhmanova, and as a teaching assistant for courses on molecular biology and metabolism. Following his interest in cell biology gone awry, he enrolled in the Cancer Genomics and Developmental Biology MSc. program at UU in 2012. As part of his MSc., he studied regulators of DNA interstrand crosslink repair at the Hubrecht Institute in Utrecht under supervision of Dr. Puck Knipscheer. Subsequently, at Weill Cornell Medical College of Cornell University in New York, he studied matrix metalloproteinase signaling in the lung cancer microenvironment with Prof. Vivek Mittal, triggering his interest in the study of complex cellular interplay in tumors. This prompted him to join the lab of Prof. Karin de Visser at the Netherlands Cancer Institute in Amsterdam as a Ph.D. candidate in 2014, for further exploration of the inner workings of the tumor microenvironment and systemic inflammation in breast cancer. The results of this work are presented in this thesis. From 2020 onwards, he continues examining cancer and immune heterogeneity as a postdoctoral researcher in the lab of Prof. Alexander van Oudenaarden at the Hubrecht Institute.

Publications

In vitro assessment of cancer cell-induced polarization of macrophages

Danique E.M. Duits, <u>Max D. Wellenstein</u>, Karin E. de Visser* *Methods in Enzymology*. 2020 Feb; 632:133-154. doi: 10.1016/bs.mie.2918.06.011.

Loss of p53 triggers WNT-dependent systemic inflammation to drive breast cancer metastasis

<u>Max D. Wellenstein</u>[#], Seth B. Coffelt[#], Danique E.M. Duits, Martine H. van Miltenburg, Maarten Slagter, Iris de Rink, Linda Henneman, Sjors M. Kas, Stefan Prekovic, Cheei-Sing Hau, Kim Vrijland, Anne Paulien Drenth, Renske de Korte-Grimmerink, Eva Schut, Ingrid van der Heijden, Wilbert Zwart, Lodewyk F.A. Wessels, Ton N. Schumacher, Jos Jonkers^{*}, Karin E. de Visser^{*}

Nature. 2019 Aug; 572(7770): 538-542. doi: 10.1038/s41586-019-1450-6.

Fatty acids corrupt neutrophils in cancer

Max D. Wellenstein, Karin E. de Visser* *Cancer Cell.* 2019 Jun; 35(6):827-829. doi: 10.1016/j.ccell.2019.05.007.

Cancer-cell-intrinsic mechanisms shaping the tumor immune landscape

<u>Max D. Wellenstein</u>, Karin E. de Visser* *Immunity*. 2018 Mar; 48(3):399-416. doi: 10.1016/j.immuni.2018.03.004.

Mammary tumor-derived CCL2 enhances pro-metastatic systemic inflammation through upregulation of IL1 β in tumor-associated macrophages

Kelly Kersten, Seth B. Coffelt, Marlous Hoogstraat, Niels J.M. Verstegen, Kim Vrijland, Metamia Ciampricotti, Chris W. Doornebal, Cheei-Sing Hau, <u>Max D. Wellenstein</u>, Camilla Salvagno, Parul Doshi, Esther H. Lips, Lodewyk F.A. Wessels, Karin E. de Visser* *Oncoimmunology*. 2017 Jun; 19;6(8):e1334744. doi: 10.1080/2162402X.2017.1334744.

Matrix Metalloproteinase 14 promotes lung cancer by cleavage of Heparin-Binding EGF-like Growth Factor

Marcin Stawowczyk[#], <u>Max D. Wellenstein</u>[#], Sharrell B. Lee[#], Shira Yomtoubian, Anna Durrans, Hyejin Choi, Navneet Narula, Nasser K. Altorki, Dingcheng Gao^{*}, Vivek Mittal^{*} *Neoplasia*. 2017 Feb; 19(2):55-64. doi: 10.1016/j.neo.2016.11.005.

Neutrophils in cancer: neutral no more

Seth B. Coffelt, <u>Max D. Wellenstein</u>, Karin E. de Visser* *Nature Reviews Cancer.* 2016 Jul; 16(7):431-446. doi: 10.1038/nrc.2016.52.

* equal contribution, * corresponding author

Manuscripts in revision, under review or in preparation

Distinct p53 hotspot mutations in breast cancer regulate immunotherapy response through differential activation of autophagy

Max D. Wellenstein, Stefan Prekovic[#], Isabel Mayayo-Peralta[#], Onno B. Bleijerveld, Liesbeth Hoekman, Cheei-Sing Hau, Kevin Kos, Elisabeth A.M. Raeven, Kim Vrijland, Daphne Kaldenbach, Danique E.M. Duits, Anni Laine, Maarten Altelaar, Wilbert Zwart, Karin E. de Visser^{*}

In revision (Nature Cancer). 2020

Glucocorticoids regulate cancer cell dormancy

Stefan Prekovic[#], Karianne Schuurman[#], Anna González Manjón, Mark Buijs, Isabel Mayayo-Peralta, <u>Max D. Wellenstein</u>, Seçuk Yavuz, Alejandro Barrera, Kim Monkhorst, Anne Huber, Ben Morris, Cor Lieftink, Joana Silva, Balázs Győrffy, Liesbeth Hoekman, Bram van den Broek, Hans Teunissen, Timothy Reddy, William Faller, Roderick Beijersbergen, Jos Jonkers, Maarten Altelaar, Karin E. de Visser, Elzo de Wit, Rene Medema, and Wilbert Zwart*

Under review. Preprint at bioRxiv. 2019 Sep; doi: 10.1101/750406

Proteomic characterization of neutrophils in metastatic breast cancer reveals tissueand maturation state-specific phenotypes

<u>Max D. Wellenstein</u>, Seth B. Coffelt, Onno B. Bleijerveld, Hannah Garner, Cheei-Sing Hau, Kim Vrijland, Maarten Altelaar, Karin E. de Visser* *In preparation.*

MET controls the immune landscape of triple-negative breast cancer

Danique E.M. Duits[#], <u>Max D. Wellenstein</u>[#], Linda Henneman, Martine H. van Miltenburg, Seth B. Coffelt, Celine Sewnath, Kevin Kos, Anni Laine, Chris Doornebal, Kim Vrijland, Cheei-Sing Hau, Elisabeth A.M. Raeven, Daphne Kaldenbach, Anne Paulien Drenth, Renske de Korte-Grimmerink, Eva Schut, Jos Jonkers, Karin E. de Visser^{*} *In preparation.*

*equal contribution, *corresponding author

Acknowledgements

Here is an (in no way comprehensive) tip of the hat to the people that have shaped my thoughts and courses of action over the past 6 - 31 years. I am endlessly grateful to:

Karin, for teaching me a sharp eye, while always keeping the bigger picture in mind, the importance of conveying a narrative, of surrounding yourself with a good team and for keeping me sane.

Seth, for making me realize there are so many big things to do and to discover, and for many a post-it with scientific ideas or good music. You can't light a fire without a spark.

My paranymphs:

David, for being an inspiration and guiding light over all these years. And Kevin, who spun gold equally out of experiments and nights on the town. Thanks for our \$1e6-121s and your many overarching theorems.

My colleagues, both old and new, for creating such a fantastic and stimulating environment in the lab and far beyond:

Riccardo and Camilla. Anne, Feline, Kaspar, Lianne, Maarten, Meike, Mirjam and Ping. DD, Cheei, Kim, Kelly, Chris D, Lisanne, Daphne, ('Champ')Anni, Hannah, Antoinette, Noor, Olga, Lorenzo Spangyolo, Quinte, Claudia and Ewald. The rest of B3, both old and new: Leila, Jeremy, Luuk, Marnix, Awa, Shanna, Daan, Christel, Daniel, Serena, Marleen, Leonie, Iris, Maxime, Chris, Wietske, Paul, Karin, Hein and Heinz and their teams. Jannie, Mensink, Evert and the rest of the JB lab. Paula, Inge and Elselien. Jos, Jacco, Martine, Linda, Sjors, Anne Paulien, Eline, Eva, Koen, Stefano, Catrin and Laura. Stefan, Isabel and Wilbert. Joost, Wouter, Pia, Anastasia, Mireille, Jos, Lorenzo Fanchi, Georgi, Krijn, Salo, Chong and Ton. Lodewyk, Lorenzo Bombardelli, Luca and Robin. Marieke, Nathalie, Renske and all members of the intervention unit, Sjoerd and the pathology team, the animal caretakers, Onno, Iris, Frank, Anita and Martijn.

My dear friends, who I am fortunate enough to have by my side. Thanks for your shenanigans and always having my back; your encouragement and guidance are indispensible:

Ruben & Celine. Lins, Bennefriend, Kwark, Joep, Daan, Kimi, Jules, Aal, Pijp, Stina, Clown, Droefie and your respective ladies. Kay. Gijs, Simon. Niels, an extra big thanks for being the grand designer of this little book's layout. And of course, of the utmost importance, the bands: Rune & Bram of the renowned Tuber Solani. Populist & Biofilm Paardenfilm.

Connie, Iris and Willem, for creating a warm home for me from day one. And Marius, for your laughter and song-whistling, your razor-sharp mind and for providing advice for the writing of this book until the very end.

The fam: Ed, Saar, Noah, Jacob, Mus, Tessa, Ella, Joop, Maxime, Oma, PPM, Nana, Aad & Mary. Your love and support mean the world to me.

Pa & Ma, for magically always being there, for teaching me determination, for your love, and for actually reading my papers.

Tessa, for always saying the right things, for your songs, for our adventures, for my weak knees and for this life.