CHEMICAL HERITAGE FOUNDATION

MANFRED FRASCH

The Pew Scholars Program in the Biomedical Sciences

Transcript of an Interview Conducted by

Andrea R. Maestrejuan

at

Mount Sinai School of Medicine New York City, New York

on

17-19 December 1998

From the Original Collection of the University of California, Los Angeles

ACKNOWLEDGEMENT

This oral history is part of a series supported by a grant from the Pew Charitable Trusts based on the Pew Scholars Program in the Biomedical Sciences. This collection is an important resource for the history of biomedicine, recording the life and careers of young, distinguished biomedical scientists and of the Pew Scholars Program in the Biomedical Sciences Advisory Committee members.

This oral history was completed under the auspices of the Oral History Project, University of California, Los Angeles (Copyright © 2000, The Regents of the University of California) and is made possible through the generosity of



From the original collection at the Center for Oral History Research, UCLA Library, UCLA.

The following oral history, originally processed at the UCLA Center for Oral History Research, has been reformatted by the Chemical Heritage Foundation. The process involved reformatting the front matter, adding a new abstract, replacing the table of contents, and replacing the index. The paragraph spacing and font of the body of the transcript were altered to conform to the standards of the Oral History Program at the Chemical Heritage Foundation. The text of the oral history remains unaltered; any inadvertent spelling or factual errors in the original manuscript have not been modified. The reformatted version and digital copies of the interview recordings are housed at the Othmer Library, Chemical Heritage Foundation. The original version and research materials remain at the Darling Library, University of California, Los Angeles and at the Bancroft Library, University of California, Berkeley.

REFORMATTING:

Marnie Berkowitz, Consultant to the Chemical Heritage Foundation. B.A., Classical Languages and Literatures, University of Minnesota; Ford Foundation Fellowship, Classical Languages and Literatures, University of Chicago.

David J. Caruso, Program Manager, Oral History, Chemical Heritage Foundation. B.A., History of Science, Medicine, and Technology, Johns Hopkins University; PhD., Science and Technology Studies, Cornell University.

UNIVERSITY OF CALIFORNIA, LOS ANGELES

Oral History Interview Agreement No. 7012699C

This Interview Agreement is made and entered into this 27th day of Setting, 1997 by and between THE REGENTS OF THE UNIVERSITY OF CALIFORNIA, a California corporation, on behalf of the Oral History Program at the UCLA campus, hereinafter called "University," and MANFRED FRASCH, having an address at Mount Sinai School of Medicine, Brookdale Center for Molecular Biology, One Gustave L. Levy Place, Box 1126, New York, New York 10029, hereinafter called "Interviewee."

Interviewee agrees to participate in a series of University-conducted tape-recorded interviews, commencing on or about December 17, 1998, and tentatively entitled "Interview with Manfred Frasch". This Agreement relates to any and all materials originating from the interviews, namely the tape recordings of the interviews and a written manuscript prepared from the tapes, hereinafter collectively called "the Work."

In consideration of the mutual covenants, conditions, and terms set forth below, the parties hereto hereby agree as follows:

- Interviewee irrevocably assigns to University all his copyright, title and interest in and to the Work. This assignment applies to University, its successors, and assigns, for and during the existence of the copyright and all renewals and extensions thereof.
- 2. By virtue of this assignment, University will have the right to use the Work for any research, educational, or other purpose, including electronic reproduction, that University may deem appropriate.
- 3. Interviewee acknowledges that he will receive no remuneration or compensation for his participation in the interviews or for the rights assigned hereunder.
- 4. Interviewee will receive from University, free of charge, one bound copy of the typewritten manuscript of the interviews.
- 5. To insure against substantive error or misquotation, Interviewee will have the right to review the manuscript before it is put into final form. University therefore will send Interviewee a copy of the edited transcript for review and comment. Interviewee will return transcript and comments to University within 30 days of receipt of the transcript. In the event that Interviewee does not respond within 30 days, University will assume that Interviewee has given full approval of the transcript.

All notices and other official correspondence concerning this 6. Agreement will be sent to the following:

Office of Research Administration If to University: University of California, Los Angeles P.O. Box 951406 Los Angeles, California 90095-1406

Attention:	

If to Interviewee:

Manfred Frasch Mount Sinai School of Medicine Brookdale Center for Molecular Biology One Gustave L. Levy Place Box 1126 New York, New York 10029

University and Interviewee have executed this Agreement on the date first written above.

INTERVIEWEE

(Signature)

Manfred Frasch (Typed Name)

10.00

Sec. Burn

Mount Sinai School of Medicine

THE REGENTS OF THE UNIVERSITY OF CALIFORNIA

(Signature)

(Typed Name)

Dale E. Treleven

(Title) Director, Oral History Program

One Gustave L. Levy Place (Address)

<u>Box 112</u>6

New York, New York 10029

Date 12/17/98

Date 1/2

Pew Scholars in the Biomedical Sciences Chemical Heritage Foundation Internet Posting Release Form

I, Manfred Frasch, Ph.D., hereby request that my wishes be followed as per the checked selection below with regards to posting portions of the digital copy of the audio-taped interview of me and the related written transcript on the internet for non-commercial, educational use only.

Please check one:

No restrictions for Internet Posting.

NOTE: Users citing this interview for purposes of publication are obliged under the terms of the Chemical Heritage Foundation Oral History Program to obtain permission from Chemical Heritage Foundation, Philadelphia, Pennsylvania.

b.

Semi-restricted Internet Postings (My review of the material intended to post is required.)

c.____

Restricted access. (Do not post.)

This constitutes my entire and complete understanding.

Share and state one of the

Manfred Frasch, Ph.D.

2/6/08

Date

This interview has been designated as Free Access.

One may view, quote from, cite, or reproduce the oral history with the permission of CHF.

Please note: Users citing this interview for purposes of publication are obliged under the terms of the Chemical Heritage Foundation Oral History Program to credit CHF using the format below:

Manfred Frasch, interview by Andrea R. Maestrejuan at the Mount Sinai School of Medicine, New York City, New York, 17-19 December 1998 (Philadelphia: Chemical Heritage Foundation, Oral History Transcript # 0442).

Chemical Heritage Foundation Oral History Program 315 Chestnut Street Philadelphia, Pennsylvania 19106	

The Chemical Heritage Foundation (CHF) serves the community of the chemical and molecular sciences, and the wider public, by treasuring the past, educating the present, and inspiring the future. CHF maintains a world-class collection of materials that document the history and heritage of the chemical and molecular sciences, technologies, and industries; encourages research in CHF collections; and carries out a program of outreach and interpretation in order to advance an understanding of the role of the chemical and molecular sciences, technologies, and industries in shaping society.

MANFRED FRASCH

1954	Born in Holzgerlingen, Germany on 15 March
	Education
1981	Diploma, University of Tübingen
1985	Ph.D., University of Tübingen
	Professional Experience
	Max Planck Institute for Developmental Biology
1981-1986	Postdoctoral Fellow, Department of Physical Biology
1988-1991	Research Fellow, Department of Genetics
1986-1988	Columbia University Postdoctoral Fellow, Department of Biology
1700 1700	rostdoetoral renow, Department of Diology
1991-1995 1996-present	Brookdale Center for Molecular Biology, Mount Sinai School of Medicine Assistant Professor Associate Professor
*	

Honors

1986-1988	Deutsche Forschungsgemeinschaft Fellowship
1989-1991	Deutsche Forschungsgemeinschaft Research Award
1993-1997	Pew Scholar in the Biomedical Sciences

Selected Publications

- Risau, W. et al., 1983. Nonpackaging and packaging proteins of hnRNA in *Drosophila melanogaster. Cell* 33:529-41.
- Frasch, M. et al., 1987. Characterization and localization of the *even-skipped* protein of *Drosophila. European Molecular Biology Organization Journal* 6:749-59.
- Frasch, M. et al., 1987. Complementary patterns of *even-skipped* and *fushi tarazu* expression involve their differential regulation by a common set of segmentation genes in *Drosophila. Genes and Development* 1:981-95.

Frasch, M. and H. Saumweber, 1989. Two proteins from Drosophila nuclei are bound to

chromatin and are detected in a series of puffs on polytene chromosomes. *Chromosoma* 97:272-81.

- Dohrmann, C. et al., 1990. A new *Drosophila* homeobox gene is expressed in mesodermal precursor cells of distinct muscles during embryogenesis. *Genes and Development* 4:2098-111.
- Frasch, M., 1991. The maternally expressed *Drosophila* gene encoding the chromatin-binding protein *BJ1* is a homolog of the vertebrate gene regulator of chromatin condensation, *RCC1*. *European Molecular Biology Organization Journal* 10:1225-36.
- Azpiazu, N. and M. Frasch, 1993. *Tinman* and *bagpipe:* two homeobox genes that determine cell fates in the dorsal mesoderm of *Drosophila*. *Genes and Development* 7:1325-40.
- Frasch, M., 1995. Induction of visceral and cardiac mesoderm by ectodermal Dpp in the early *Drosophila* embryo. *Nature* 374:464-67.
- Azpiazu, N. et al., 1996. Segmentation and specification of the *Drosophila* mesoderm. *Genes* and *Development* 10:3183-94.
- Yin, Z. et al., 1997. Regulation of the *twist* target gene *tinman* by modular cis-regulatory elements during early mesoderm development. *Development* 124:4971-82.
- Xu, X. et al., 1998. *Smad* proteins act in combination with synergistic and antagonistic regulators to target Dpp responses to the *Drosophila* mesoderm. *Genes and Development* 12:2354-70.
- Knirr, S. et al., 1999. The role of the NK-homeobox gene *slouch* (*S59*) in somatic muscle patterning. *Development* 126:4525-35.

ABSTRACT

Manfred Frasch was born in Holzgerlingen, Germany, in Swabia. His father was what we would call a contractor, building mostly wooden roofs. This business was begun by Frasch's grandfather, and the Frasch family has lived in that area for many generations. Manfred lived on a farm, where his mother did the farming. His father's workshop was also on the farm. He was brought up in the Lutheran faith. He had an early curiosity about how things work, leading him to chemistry and biology. After finishing *gymnasium*, Frasch completed his compulsory military service.

He entered the University of Tübingen, where he majored in biochemistry. He also studied molecular biology at the University of Munich, with which Tübingen had an exchange program. His diploma thesis concerned gene regulation in *Drosophila*; he found *Drosophila* so fascinating that he has remained in that field.

Liking the projects, the atmosphere, and the independence of Tübingen, Frasch decided to stay there for his Ph.D., using biochemical rather than genetic techniques in his research into *Drosophila*. He eventually learned cloning techniques and decided to pursue genetic approaches rather than biochemical. He worked in Friedrich Bonhoeffer's lab, where he had a great deal of independence.

Wanting to see more of the world and wanting to expand his scientific horizons, he applied for postdocs in the United States. He accepted a position in Michael Levine's lab at Columbia University, working on the *even-skipped* gene. He had always intended to return to Germany, and he accepted a position as a research fellow in Christiane Nüsslein-Volhard's lab in the Department of Genetics at the Max Planck Institute for Developmental Biology, where his focus was on mesoderm development. There, work on *S59* led to the characterization of *tinman* and *bagpipe*. Frasch was not sanguine about his career prospects during his last year at the University of Tübingen, so he decided to return to the States, and accepted a position in the Brookdale Center for Molecular Biology at Mount Sinai School of Medicine in New York City. There he established his own lab, where he hopes to find clinical relevance for his mesoderm and heart development research. He began as an associate professor in the Brookdale but is now a tenured associate professor. He is married to Hanh Thi Nguyen, who is also a scientist.

UCLA INTERVIEW HISTORY

INTERVIEWER:

Andrea R. Maestrejuan, Interviewer, UCLA Oral History Program; B.A., History, University of California, Irvine, 1988; B.S., Biological Sciences, University of California, Irvine, 1988; C.Phil., History, University of California, Riverside.

TIME AND SETTING OF INTERVIEW:

Place: Frasch's office, Mount Sinai School of Medicine.

Dates, length of sessions: December 17, 1998 (83 minutes); December 18, 1998 (127) ; December 19, 1998 (102).

Total number of recorded hours: 5.2

Persons present during interview: Frasch and Maestrejuan.

CONDUCT OF INTERVIEW:

This interview is one in a series with Pew Scholars in the Biomedical Sciences conducted by the UCLA Oral History Program in conjunction with the Pew Charitable Trusts's Pew Scholars in the Biomedical Sciences Oral History and Archives Project. The project has been designed to document the backgrounds, education, and research of biomedical scientists awarded four-year Pew scholarships since 1988.

In preparing for this interview, Maestrejuan, in consultation with the director of the UCLA Oral History Program and three UCLA faculty project consultants, developed a topic outline to provide an overall interview framework. Maestrejuan then held a telephone pre interview conversation with Frasch to obtain extensive written background information (curriculum vitae, copies of published articles, etc.) and agree on a research and interviewing timetable.

Maestrejuan further reviewed the documentation in his file at the Pew Scholars Program office in San Francisco, including his proposal application, letters of recommendation, and reviews by Pew Scholars Program national advisory committee members. For general background on the recent history of the biological sciences, Maestrejuan consulted J.D. Watson et al., *The Molecular Biology of the Gene.* 4th ed. 2 vols. Menlo Park, CA: Benjamin/ Cummings, 1987; Lubert Stryer, *Biochemistry.* 3rd ed. New York: W.H. Freeman, 1988; H.F. Judson, *The Eighth Day of Creation: Makers of the Revolution in Biology.* New York: Simon and Schuster, 1979; and recent issues of *Science, Nature* and *Cell.*

The interview is organized chronologically, beginning with Frasch's childhood in Holzgerlingen, Germany and his undergraduate work at University of Tübingen and continuing through his graduate and postdoctoral work at Max Planck Institute for Developmental Biology, further postdoctoral work at Columbia University, and the establishment of his own lab at Mount Sinai School of Medicine. Major topics discussed include Frasch's work on *S59* in the Michael Levine lab, his characterization of the *tinman* and *bagpipe* genes, and differences between how science is conducted in Germany and in the United States.

ORIGINAL EDITING:

Ji Young Kwon, editorial assistant, edited the interview. She checked the verbatim transcript of the interview against the original tape recordings, edited for punctuation, paragraphing, and spelling, and verified proper names. Words and phrases inserted by the editor have been bracketed.

Frasch reviewed the transcript. He verified proper names and made a number of corrections and additions.

William Van Benschoten, editor, prepared the table of contents and index. Kwon compiled the biographical summary and interview history.

TABLE OF CONTENTS

Early Years Childhood in Holzgerlingen, Germany. Family background. Parents' expectations. Frasch's early education. Interest in chemistry and biology. Religious upbringing in the Lutheran church. His military service in the German army.	1
Undergraduate Years Decision to study biochemistry at the University of Tübingen. Studies molecular biology at the University of Munich. Frasch conducts research on gene regulation in <i>Drosophila</i> for his diploma thesis. His decision to pursue a Ph.D. degree. Reasons for remaining at University of Tübingen.	23
Graduate Years Frasch's first publication. Transition from undergraduate to graduate student. Learns cloning techniques. Decision to pursue genetic rather than biochemical approachesDecision to remain in the Friedrich Bonhoeffer lab. Frasch's independence in the Bonhoeffer lab.	34
Postgraduate Years in United States Decides to seek a postdoc in the U.S. Enters the Michael Levine lab at Columbia University. His research on the <i>even-skipped</i> gene. Frasch's publications in the Levine lab.	42
 Return to Germany Accepts position as a research fellow in the Christiane Nüsslein-Volhard lab. Discovery of <i>S59</i> His focus on mesoderm development. Work on <i>S59</i> leads to the characterization of <i>tinman</i> and <i>bagpipe</i>. Frasch's career prospects during his last year at the University of Tübingen. 	50
Return to United States Problems with the German scientific system and funding. Accepts a position at Mount Sinai School of Medicine in New York City. Establishes own lab in the U.S. Future research plans. Collaboration with fellow scientists. Clinical relevance of Frasch's mesoderm and heart development research. Differences between U.S. and German scientific institutions. Frasch's wife, Hanh Thi Nguyen. Status of women and minorities in science.	58
Index	88

INDEX

Α

Abitur, 9, 12

B

BASF Group, 23 Bavaria, 14 Böblingen, Germany, 4, 14 Bonhoeffer, Friedrich, 36, 41, 42, 48 Buck, Jochen, 1

D

Daimler-Benz Aktiengesellschaft, 3 Diplom, 36, 37, 40 Diplomarbeit, 25, 34, 39 DNA, 23, 25, 27, 78 Drosophila, 26, 27, 29, 34, 39, 40, 41, 50, 56, 58, 66, 69, 71, 72, 73, 75, 77, 78, 79, 83, 86

E

EMBL. *See* European Molecular Biology Laboratory European Molecular Biology Laboratory, 27, 61

F

Fliessband, 16 Frasch, Johanna Maurer (mother), 3 Frasch, Martin (brother), 6 Frasch, Otto (father), 3 Frasch, Thomas (paternal grandfather), 3 Frasch, Walter (brother), 6, 18, 43

G

Grundschule, 5, 8 *Gymnasium*, 8, 9, 10, 11, 16, 17, 19, 23, 31

H

Harvard University, 21

Hauptschule, 5 Heidelberg, Germany, 27, 28, 61 Holzgerlingen, Germany, 1, 2, 4, 7

J

Jäckle, Herbert, 38, 39, 44

K

Krüppel, 38

L

Lehre, 6, 7 Lutheran, 19

М

Maurer, Otto (maternal grandfather), 5 Max Planck Institute, 36, 37 Max Planck Society, 37 Max-Planck Institut, 29, 38 Max-Planck Institut für Entwicklungsbiologie, 37 Max-Planck Institute for Biochemistry, 24, 26 *Meister*, 5 Miescher Institute, 28 Miró, Joan, 32 Munich, Germany, 28

Ν

Nüsslein-Volhard, Christiane, 27, 36, 37, 38, 39, 42, 52, 85

0

Oberschule, 5

Р

Pew Scholars in the Biomedical Sciences, 1 polytene, 27

R

Ribonucleic Acid hnRNA, 40 Risau, Werner, 34, 35

S

Stuttgart, Germany, 1, 4, 22 Swabia/Swabian, 1, 14, 63

Т

Tjian, Robert T., 39, 40, 43 Tübingen, Germany, 1, 28

U

Ulm, Germany, 14

Universität Konstanz, 22
Universität Tübingen. *See* University of Tübingen
University of California at Berkeley, 39
University of California at Irvine, 36
University of Munich, 24, 32
University of Tübingen, 1, 22, 24, 25, 26, 27, 28, 34, 38, 74

V

Volksschule, 5 *Vordiplom*, 24, 25

W

Wieschaus, Eric, 27