

# History of Peptide Chemistry in Warsaw

Aleksandra Misicka, 2017 Kierownik Pracowni Peptydów Wydział Chemii Uniwersytet Warszawski Peptide research at the University of Warsaw began in 1966 with the return of doctor **Stefania Drabarek** from her postdoctoral stay at the Cornell Medical University in New York at Vincent du Vigneaud laboratory

## **Professor Stefania Drabarek**

| 1919    | Sept. 1st born in Warsaw  |
|---------|---|
| 1937    | Graduation of a well-known high school in Warsaw  |
|         | (Liceum im. Królowej Jadwigi)   |
| 1937    | Acceptance to the University in Warsaw  |
|         | Department of Mathematics and Natural Sciences  |
| 1939    | Sept. 1st beginning of the World War II   |
| 1942-44 | Chemist in Pharmaceutical Company (ASID, previously KLAWE)  |
|         | KLAWE was the world-famous Polish pharmaceutical company, which at the beginning of the WW II, in 1940, was taken over by the German company ASID.          |
|         | Klawe's family recovered the company after the war in 1945, but only for a short time because <i>KLAWE</i> was nationalized in 1946 and became <i>POLFA</i> |
| 1944    | Aug 1st The outbreak of the Warsaw Uprising - she managed to escape from the transport to a concentration camp  |
| 1944    | Secret teaching (tajne nauczanie) in a high school in Żyrardów  |
| 1945-48 | Continuation of studies at the Uniwersity of Warsaw   |
| 1946    | Assistant at University of Warsaw, Medical Department   |
| 1948    | Master degree (July 13th)   |



#### Stefania Drabarek, student

#### maiden name Janiszewska

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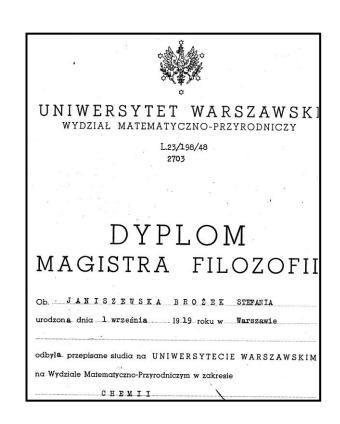


Certificate of The Józef Piłsudski University admission (20.10.1937)

Index of Warsaw University (1.10.1945)

#### Stefania Drabarek, master certificate



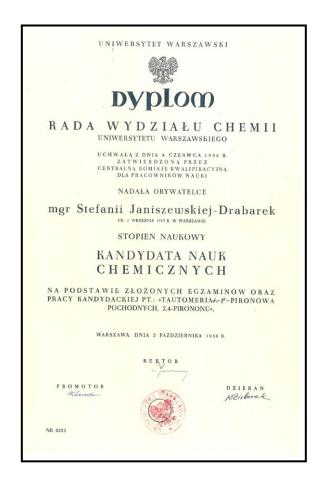


Master degree certificate

Department of Mathematics and Natural Sciences

#### Stefania Drabarek, PhD - 1956





PhD certificate (Kandydat Nauk Chemicznych) "Tautomerism of 2,4-pironon derivatives"

Promotor – Professor Lampe

#### Stefania Drabarek, PhD

1961 She received grant under the Government of the United States Educational Program and Fulbright Travel grant

1961-63 post-doc position in Vincent du Vigneaud laboratory



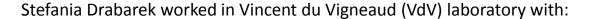
On the "Batory" – ship, which ran on the Gdynia-New York line

#### Vincent du Vigneaud

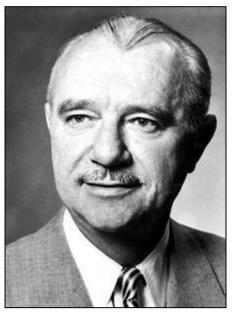
#### **Cornell Medical University in New York**

Vincent du Vigneaud (1901-78)

**Laureate of Nobel Prize in Chemistry in 1955** - "for work on biochemically important sulphur compounds, especially for the first synthesis of polypeptide hormones oxytocin and vasopressin"



Maurice Manning, Georges Flouret, Iphigenia Photaki, Victor J Hruby, Miklos Bodanszky, and others



#### Stefania Drabarek, PhD

1961-62 post-doc position in du Vigneaud laboratory results were published in 5 publications first 3 publications were presented in habilitation thesis

#### J Biol Chem. 1963 238(4):1560-1561

Effect of replacement of carboxamide group by hydrogen in glutamine or asparagine residue of oxytocin on its biological activity

Du Vigneaud V, Denning GRS, Drabarek S, Chan WY.

#### J Am Chem Soc. 1964 86(120):4477-81

Analogs of oxytocin containing glycine in place of tyrosine isoleucine or glutamine **Drabarek S**.

#### J Biol Chem. 1964 239:472-78

The synthesis and pharmacological study of 4-decarboxamido-oxytocin (4-alpha-aminobutyric acid-oxytocin) and 5-decarboxamido-oxytocin (5-alanine-oxytocin).

Du Vigneaud V, Denning GRS jr, Drabarek S, Chan WY.

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#### J Am Chem Soc. 1965 87(17):3974-78.

2-D-tyrosine-oxytocin and 2-D-tyrosine-deamino-oxytocin diastereoisomers of oxytocin and deamino-oxytocin. **Drabarek S**, Du Vigneaud V.

#### J Biol Chem. 1966 241(11):2572-75.

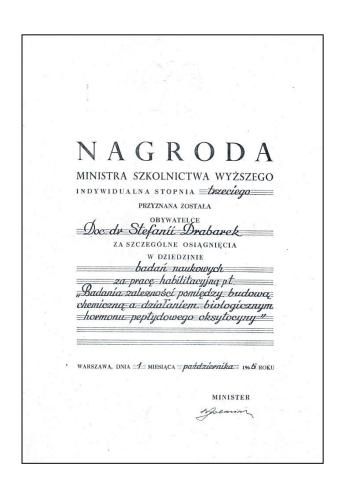
The synthesis and pharmacological properties of deamino-4-decarboxamido-oxytocin (1-beta-mercaptopropionic acid-1-alpha-aminobutyric acid-oxytocin).

Branda LA, **Drabarek S**, Du Vigneaud V.

#### Stefania Drabarek, PhD, DSc.

1964 DSc "Structure activity relationship of peptidic hormone oxytocin" 1965 Award for habilitation (Minister of Higher Education)

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## **Peptide Laboratory 1966**

The Peptide Laboratory was officially established in 1966, the first members of this group were:

Jan Izdebski, PhD Alicja Orłowska, M.Sc. Stefania Klimkiewicz, M.Sc. Jacek Pełka, M.Sc.

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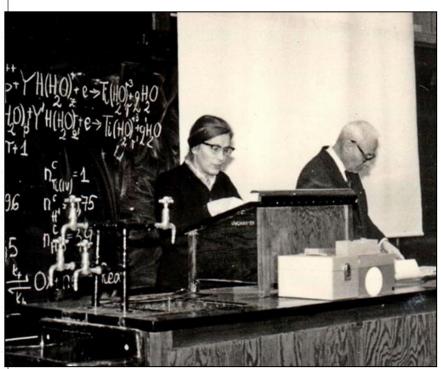
It was a difficult time, not many chemicals were available, amino acids had to be extracted from natural sources (e. g. cysteine from hair), next protecting groups had to be introduced what allowed peptide synthesis in solution, and later on solid support.

Slowly, under the supervision of Professor Drabarek members of Peptide Laboratory have gained knowledge and experience in the synthesis of peptides.

Next new members joined the peptide group:

Krzysztof Bańkowski Andrzej W. Lipkowski Teresa Wachal Teresa Kubiak and others...

#### **Professor Stefania Drabarek**





1966 – Head of the Peptide Laboratory 1969-1973, 1978 - 80 Head of the Organic Chemistry Department 1967-1972 Vice-Dean of the Chemistry Department

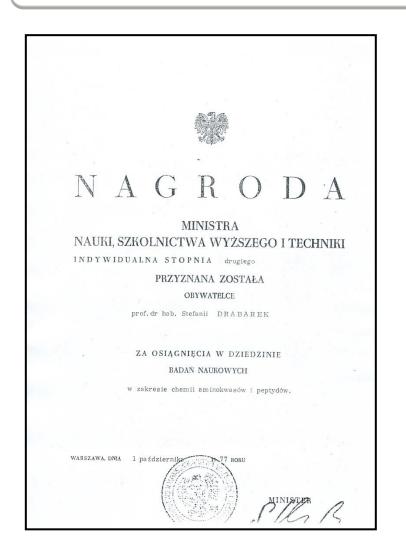
## 8th European Peptide Symposium, The Netherlands 1966



Stefania Iphigenia Drabarek Photaki

Miklos Bodanszky

## **Ministry Awards for Research and for Teaching**



## NAGRODA

MINISTRA
OŚWIATY I SZKOLNICTWA WYZSZEGO
INDYWIDUALNA STOPNIA *trzeciego* 

PRZYZNANA ZOSTAŁA

Docent dr h.Stefaníi Drabarek

ZA SZCZEGÓLNE OSIĄGNIĘCIA

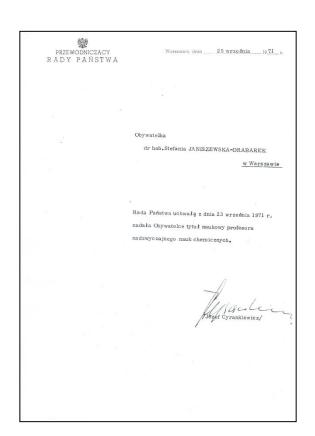
W DZIEDZINIE

dydaktyczno ~ wychowawczej, organizacji procesu dydaktycznego oraz prac związanych z kształceniem młodej kadry naukowej

WARSZAWA, DNIA 1 PAŹDZIERNIKA 1971. ROKU

MINISTER

#### **Professor Stefania Drabarek, 1971**



State Council certificate for Professor Nomination (title) signed by the Prime Minister (Cyrankiewicz)





# Stefania Drabarek, PhD, DSc Visiting Professor in Karl Hoffman Laboratory 1972-73

## Protein Research Laboratory, University of Pittsburgh, School of Medicine, Pittsburgh, US

She participated in the trials of the extremely ambitious project at that time – a total chemical synthesis of the protein - consisting of 104 amino acids of the ribonuclease T1, her successful synthesis was the 39-amino acid peptide of the enzyme fragment 66-104.



## Klan Kofma\_

#### Int J Pept Protein Res. 1974, 6(6):435-45.

Studies on polypeptides. LIII. Synthesis of a cysteine protected nonatricontapeptide corresponding to positions 66-104 of the peptide chain of ribonuclease T1.

Romovacek H, **Drabarek S**, Kawasaki K, Dowd SR, Obermeier R, Hofmann K.





## 19th European Peptide Symposium, Gdańsk 1978

First EPS held in Poland organized by prof. Kupryszewski



## PhD's mentored by Professor Drabarek





1974 Krzysztof Bańkowski

1974 Alicja Orłowska

1975 Stefania Klimkiewicz

1976 Marianna Kańska

1976 Andrzej W. Lipkowski

1976 Teresa Wachal

1977 Elżbieta Baran

1977 Krystyna Midura-Nowaczek

1978 Teresa Kubiak

1978 Maria Pachulska

1980 Tadeusz Majewski

1980 Aleksandra Misicka

#### First scientific experiences abroad...

Professor Drabarek had contacts with her colleagues from the du Vigneaud laboratory and with her recommendation several of her students could get post-doc positions in different US Universities.

Dr Jan Izdebski - Case Western Reserve University (Cleveland, Ohio), with Professor Miklos Bodanszky

Dr Krzysztof Bańkowski - Medical College of Ohio (Toledo, Ohio), with Professor Maurice Manning

Dr Andrzej W. Lipkowski - Northwestern University (Chicago, Illinois), with Professor Georges Flouret

**Dr Teresa Kubiak** - Texas University (Austin, Texas), with Professor Karl Folkers

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Dr Marianna Kańska - University of Arkansas (Fayetteville, Arkansas), with Professor Artur Fry

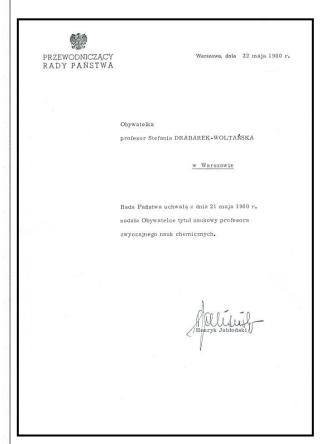
#### **Professor Stefania Drabarek and Solidarity movement**

Professor Drabarek strongly supported Solidarity movement since its very beginning (1980), and during the martial law.

As a Solidarity member, she participated in a memorable strike that took place at the Universities.

Krzysztof Bańkowski recalled that she gave us, at the beginning of the martial law, her private typewriter, which was used for typing various leaflets and Underground Solidarity newspapers.

#### **Professor Stefania Drabarek**



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State Council Certificate of Full Professor Nomination (title)

Professor Drabarek retired in 1981 and due to her personal reason she moved to Chicago.

Many of us had the opportunity to meet the hospitality of her home in Chicago, where she always waited for news from Poland and from Chemistry Department.

Sometimes she was visiting us during our stays in US.

## Visit in Maurice Manning laboratory in Toledo, USA (1985)







#### Scientific experience of some peptide laboratory members

**Teresa Kubiak** – **PhD 1978**, post-doc at Austin University with Professor Folkers (1980-83), next Rockerfeller University with Professor Merriefield (1983-87), next scientist in The Upjohn Company, Farmacia, Pfizer, now retired

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**Krystyna Midura-Nowaczek** – **PhD 1977**, post-doc at Texas University with Professor Folkers, she next moved to the branch of Warsaw University in Białystok, **1988 DSc**, **2011 Professor Nomination**, now a Head of Department of Organic Chemistry at Białystok Medical University

**Elżbieta Baran** – **PhD 1979**, post-doc at Texas University with Professor Folkers, next was a chemistry teacher in Botswana

Marianna Kańska PhD-1977 post-doc at University of Arkansas with Professor Fry, 1992 DSc, 2004 Profesor Nomination, now retired

**Tadeusz Majewski – PhD 1979**, post-doc at Northwestern University in Chicago with Professor Flouret, now at The University of Texas MD Anderson Cancer Center, Houston, Texas, US as a MS specialist

Maria Pachulska – PhD 1980, post-doc at Austin University with Professor Folkers, now at the Didactic Laboratory at Warsaw University

Aleksandra Misicka, PhD 1981, post-doc at Medical College of Ohio with Professor Manning, next research associate at University of Arizona with prof. Victor J. Hruby, at Vrije Universiteit Brussels with Professor Georges Van Binst, and at Kyoto University with Professor Masaaki Yoshikawa, 1999 DSc, 2013 Profesor Nomination,

- Recently invited professor at Universite Paris 13, Paris, France with Professor Gerard Perret visiting scholar with Professor Manning at University of Toledo
- Now: Head of The Peptide Laboratory at Warsaw University (from 2008) and Head of The Neuropeptide Laboratory in Medical Research Centre Polish Academy of Sciences (from 2015)

#### Professor Jan Izdebski

#### **Head of Peptide Laboratory 1981-2008**

1965, PhD – Promotor Professor Jan Świderski, 1979 DSc, Professor Nomination 1991



1980-81 Research associate at University California, San Francisco with Professor Cho H. Li

1994-95 Visiting professor at Tulane University, New Orleans with Professor

Andrew W. Schally

#### **Research Interests:**

Synthesis of growth hormone-releasing hormone (GH-RH) agonist

Cyclic analogs of opioid peptides containing ureido bridge

Studies on DCC coupling method with the use of different additives –

ethyl 2-hydroximino-2- cyanoacetate (now OXYMA)



#### Professor Andrzej W. Lipkowski (1946-2014)

PhD 1976, DSc 1984, Professor Nomination 2001

1977-78 Post-doc at Northwestern University (Chicago, USA) with Professor George Flouret

1983 Research fellow at the Max Planck Institute for Psychiatry (Munich, Germany) with

**Professor Albert Herz** 

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\_\_0 \_\_0 1986-88 Research fellow at University of Minnesota with Professor Philip Portoghese (Minneapolis,

1988 Research fellow at University of Arizona (Tucson, USA) with Professor Victor Hruby

1990 Visiting scholar at Tufts University School of Medicine (Boston, USA) with anestesiologist Dan Carr

Andrzej designed and synthesized "biphalin" - a dimeric enkephalin analogue, which is 1000 times more potent than morphine when administrated intrathecally.

While working in Portoghese's lab, Andrzej synthesized the famous norbinaltorphimine (norNBI), a  $\kappa$ -selective antagonist used in most laboratories involved in opioid research to the present day.

**1993-2010** Head of the Biotechnology Department at the **Industrial Chemistry Research Institute** in Warsaw **1991** Head of the Department of Neuropeptides in **Medical Research Centre Polish Academy of Science** and from **2010** a **Director** of this Institute.

Andrzej had extensive knowledge of chemistry, pharmacology and medicine and was an extremely kind and modest person with an outstanding mind.

We miss Him a lot...

#### Dr Krzysztof Bańkowski (1945-2017)

1974 PhD

**1977-78** post-doc in Medical College of Ohio, Toledo with Professor **Maurice Manning**Working in synthesis of oxytocin and vasopressin analogs

After returning to Poland, Krzysztof shared with all of us his experience and knowledge about solid phase peptide synthesis, which was at that time new for us.

Krzysztof returned to Professor Manning several times what resulted in over a dozen publications. The last one (2015) concerned the future perspective of peptide therapeutics.



Development of production technologies for several active drugs, including Leuprolid, an anticancer drug.

Krzysztof had been an excellent teacher and scientist, for many years involved in the work of the Chemistry Olympiad, always friendly and helpful in solving chemical problems...

We will miss Him a lot...



#### **Professor Stefania Drabarek (1919-1999)**

Krzysztof Bańkowski
Alicja Orłowska
Stefania Klimkiewicz
Andrzej Lipkowski
Teresa Wachal
Marianna Kańska
Krystyna Midura-Nowaczek
Elżbieta Baran
Teresa Kubiak
Maria Pachulska
Tadeusz Majewski
Aleksandra Misicka
Ewa Witkowska



Professor Drabarek was a pioner of peptide chemistry in Warsaw, but she will be remembered not only because of her peptide achievements but because she was a good boss, always taking care of her collaborators, always elegant and charming, until the end...