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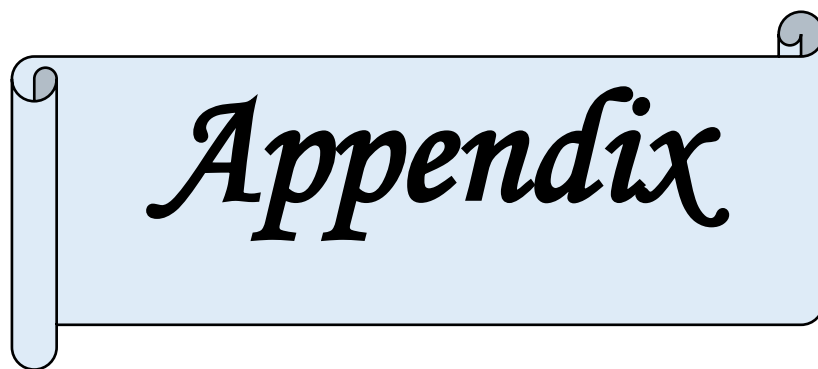
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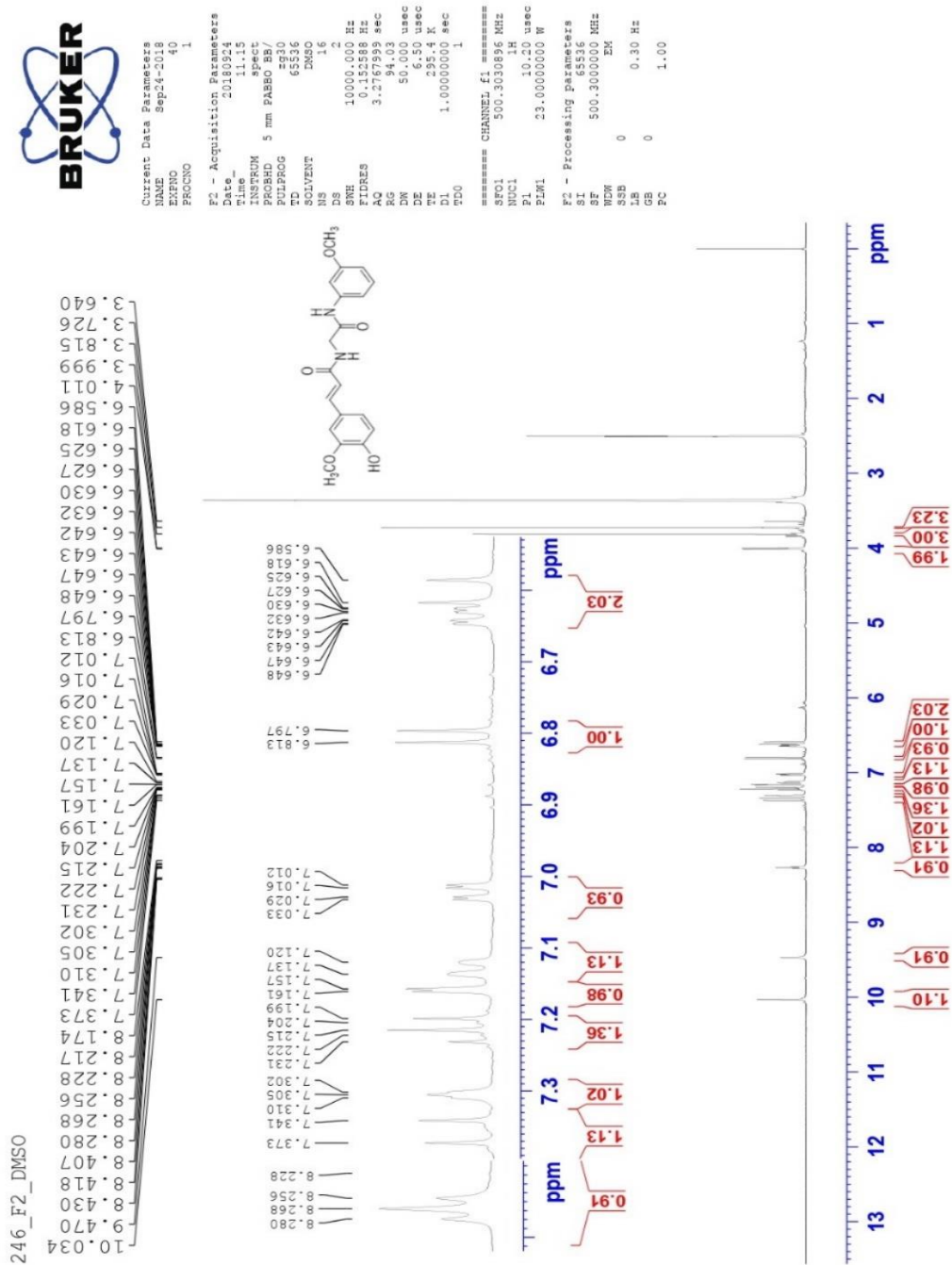


*Appendix*

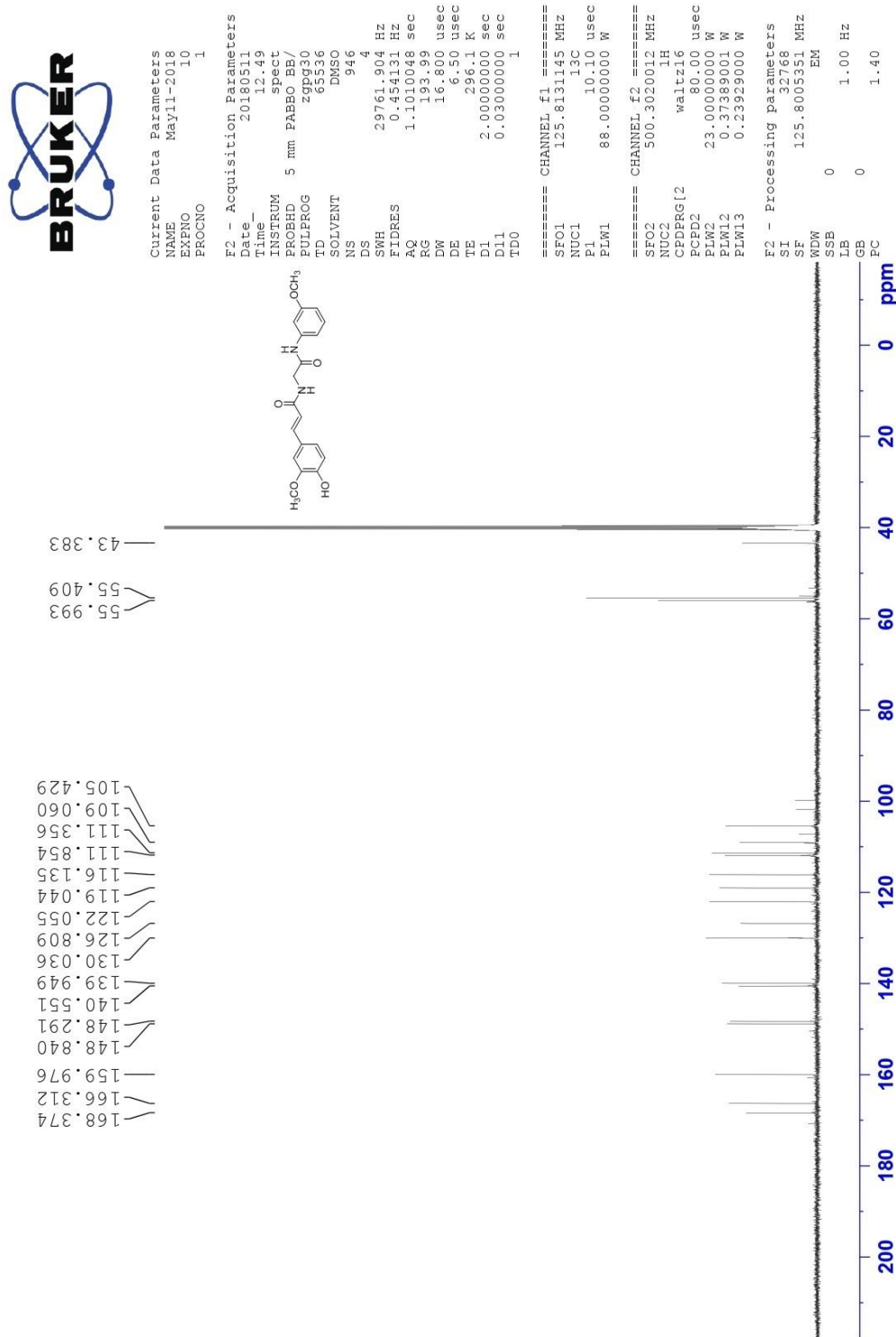
# Appendix

## Supplementary data of first series of compounds

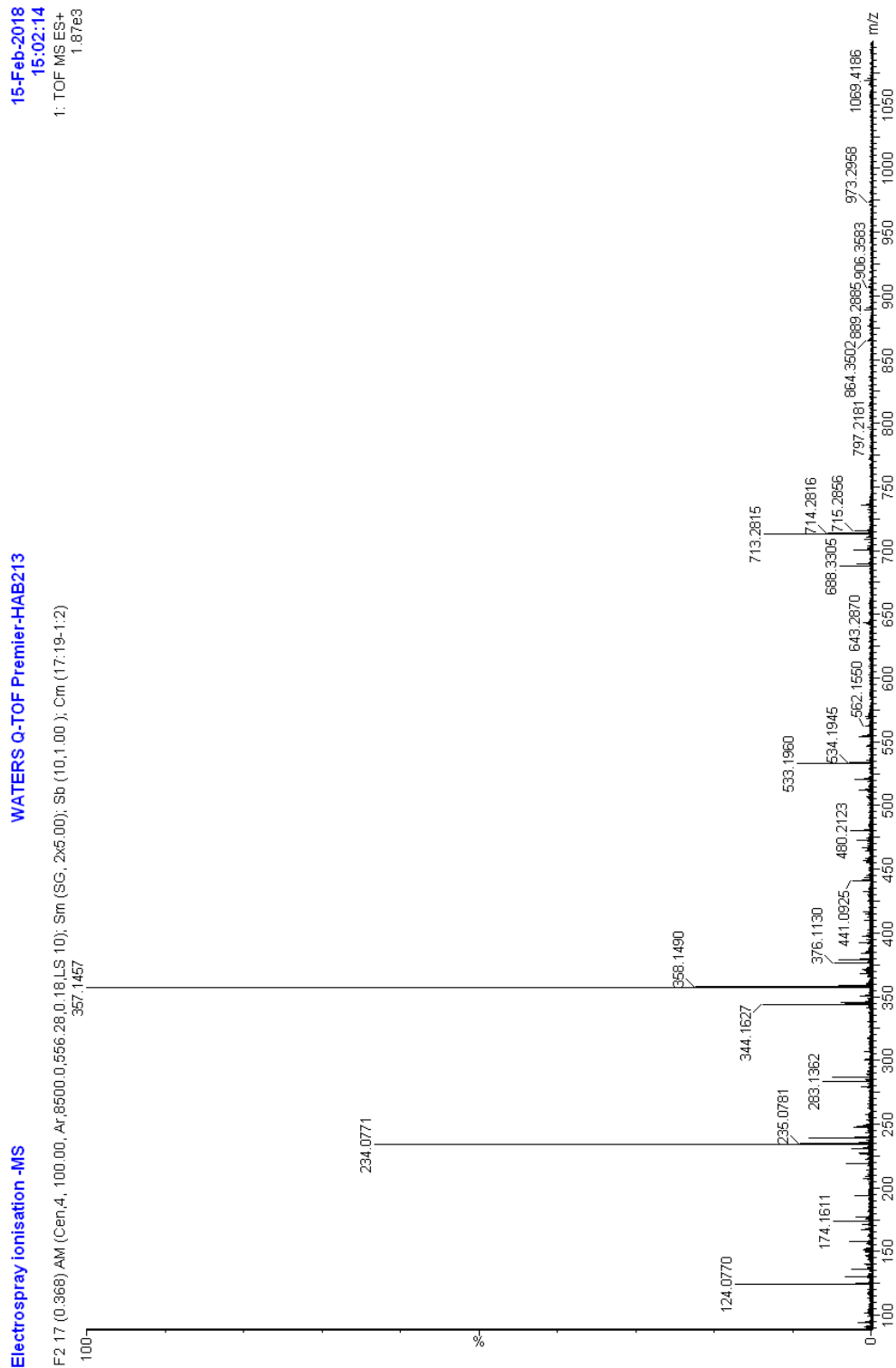
**Figure A1.**  $^1\text{H}$  NMR spectra of (*E*)-*N*-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((3-methoxyphenyl)amino)-2oxoethyl)acrylamide (**4f**).



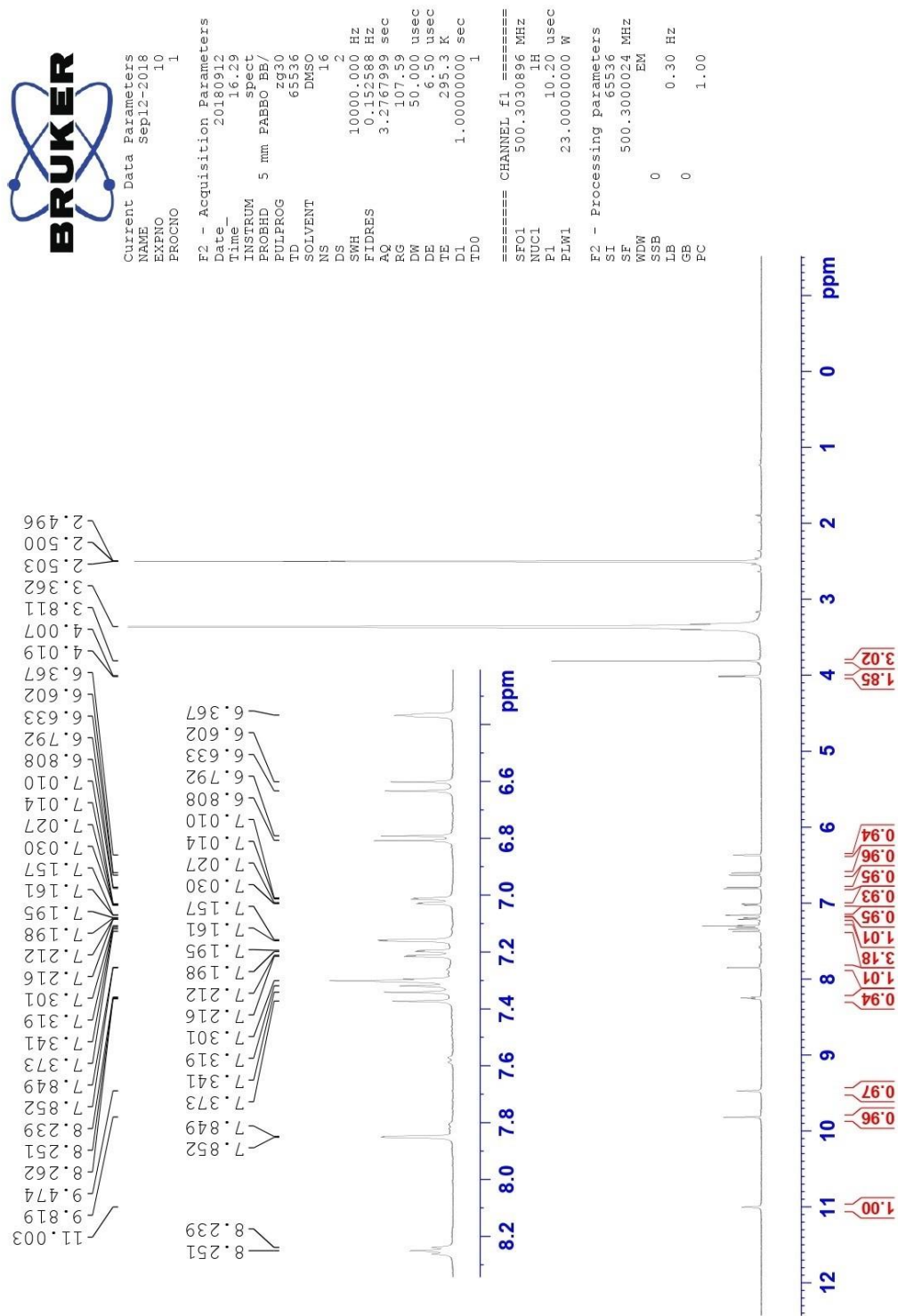
**Figure A2.**  $^{13}\text{C}$  NMR spectra of (*E*)-*N*-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((3-methoxyphenyl)amino)-2oxoethyl)acrylamide (**4f**).



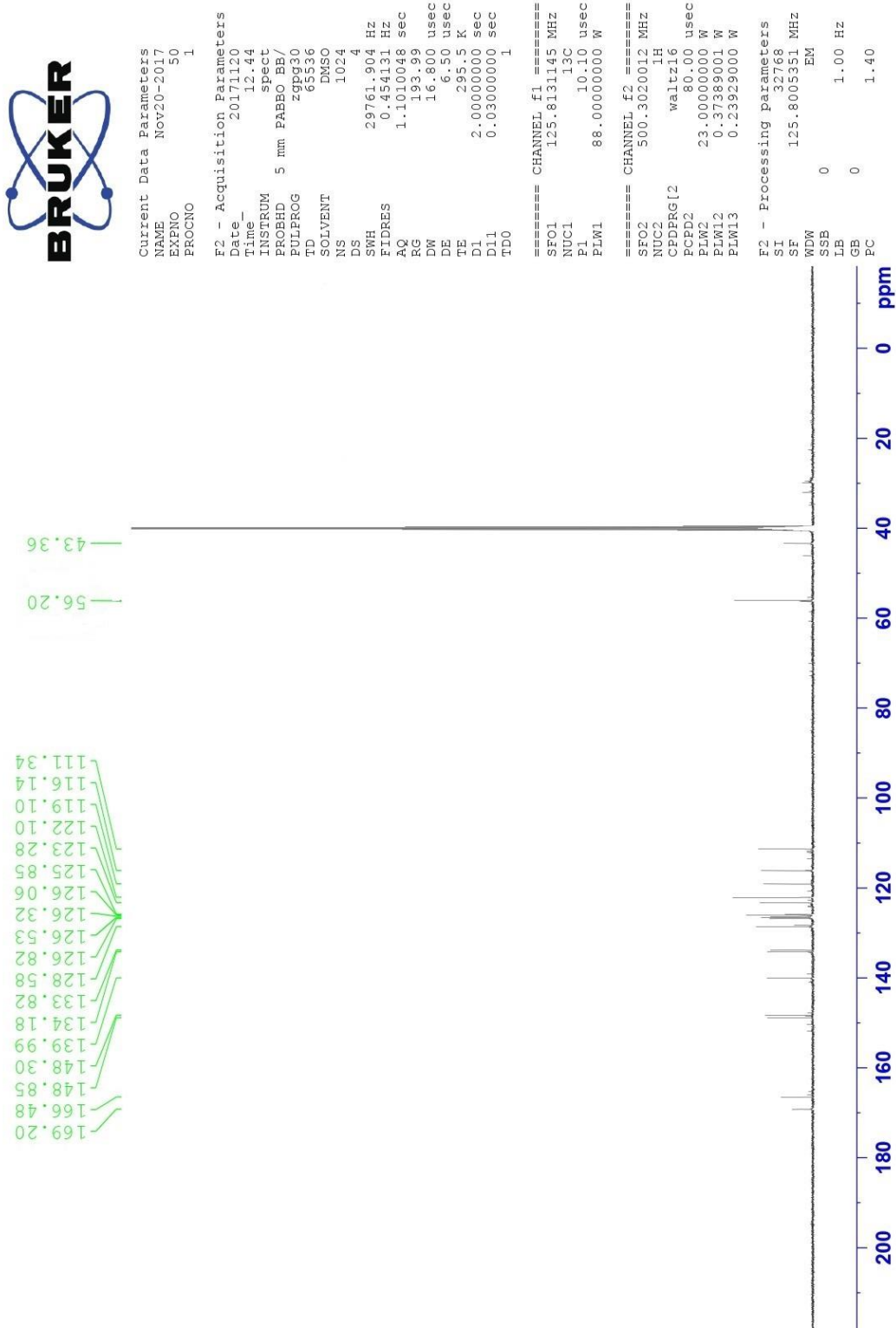
**Figure A3.** HRMS spectra of (*E*)-*N*-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((3-methoxyphenyl)amino)-2-oxoethyl)acrylamide (**4f**).



**Figure A4.**  $^1\text{H}$  NMR spectra of (*E*)-*N*-(2-((1*H*-indol-5-yl)amino)-2-oxoethyl)-3-(4-hydroxy-3-methoxyphenyl) acrylamide (**7a**).

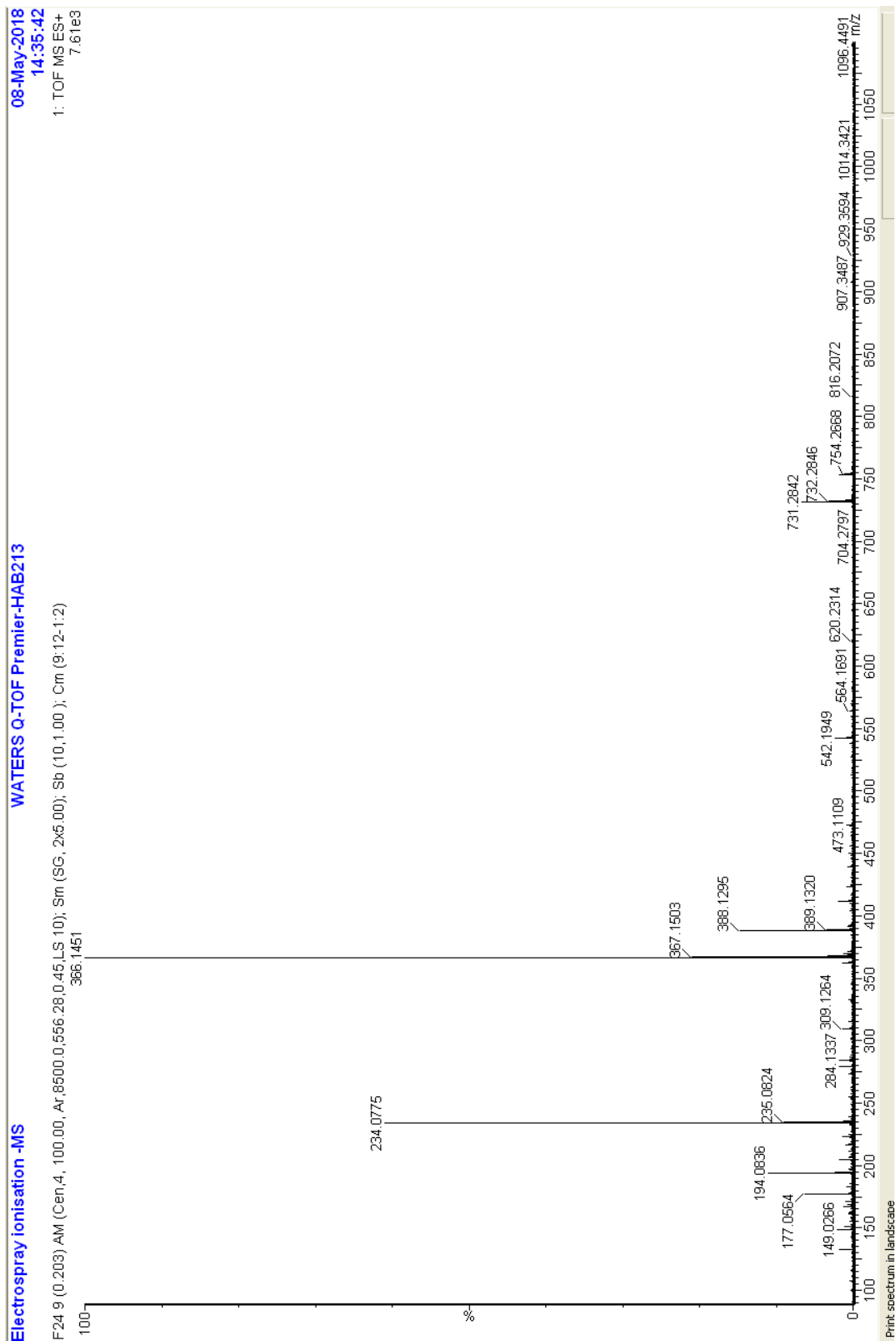


**Figure A5.**  $^{13}\text{C}$  NMR spectra of (*E*)-*N*-(2-((1*H*-indol-5-yl)amino)-2-oxoethyl)-3-(4-hydroxy-3-methoxyphenyl)acrylamide (**7a**).



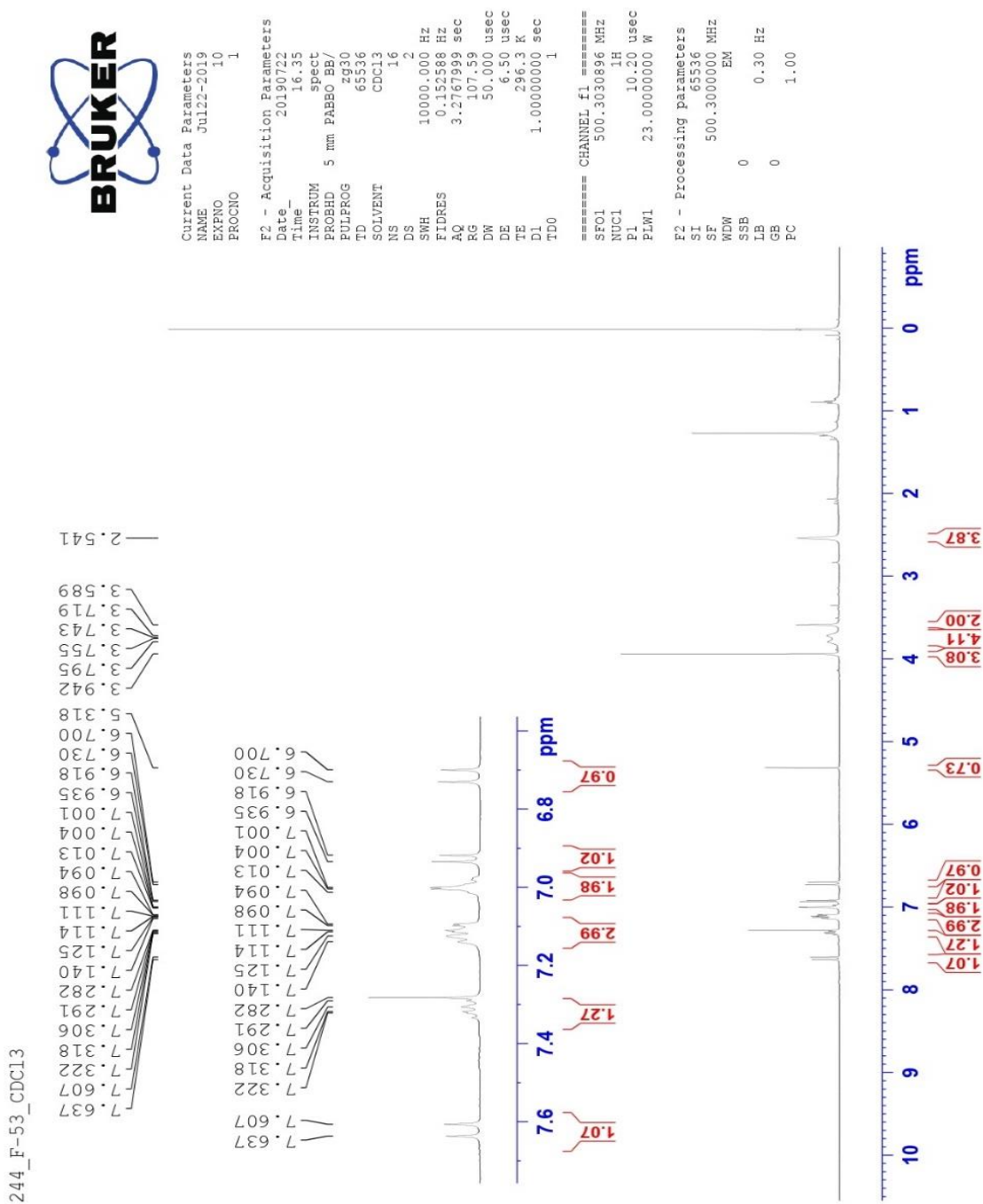


**Figure A6.** HRMS spectra of (*E*)-*N*-(2-((1*H*-indol-5-yl)amino)-2-oxoethyl)-3-(4-hydroxy-3-methoxyphenyl) acrylamide (**7a**).

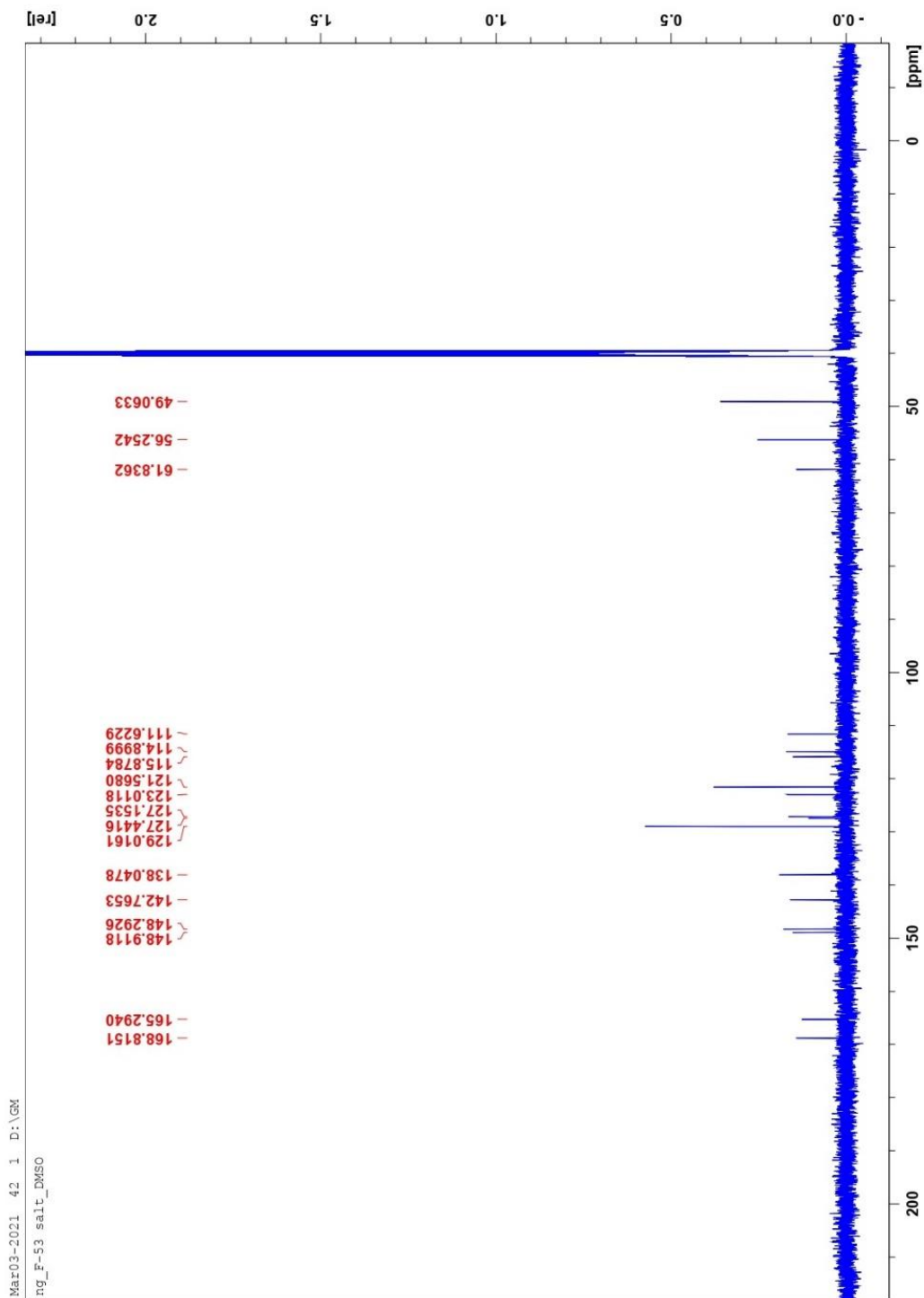


Supplementary data of second series of compounds

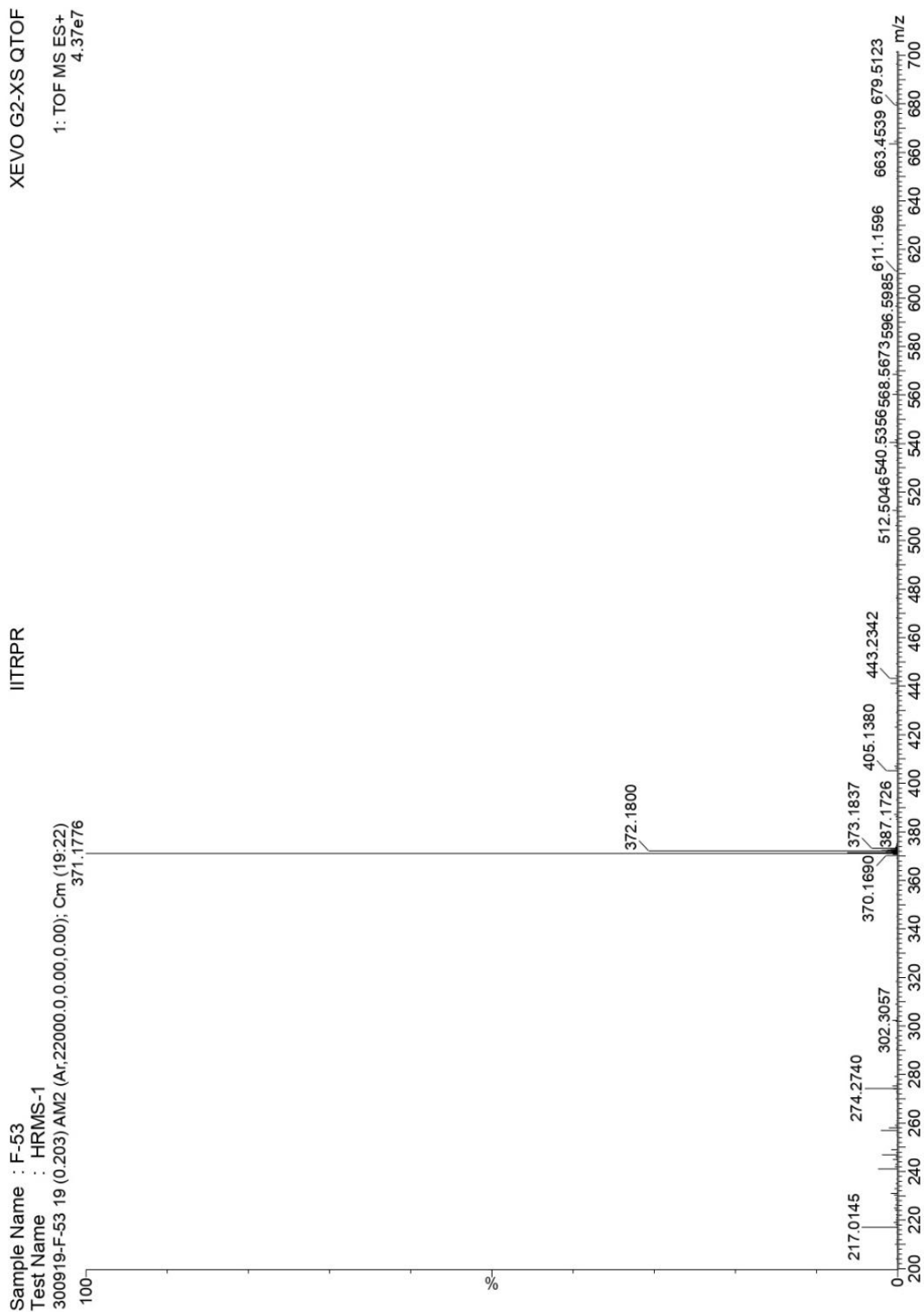
Figure A7. <sup>1</sup>H NMR spectra of (E)-1-(4-(3-fluorobenzyl)piperazin-1-yl)-3-(4-hydroxy-3-methoxyphenyl)prop-2-en-1-one (13k.HCl).



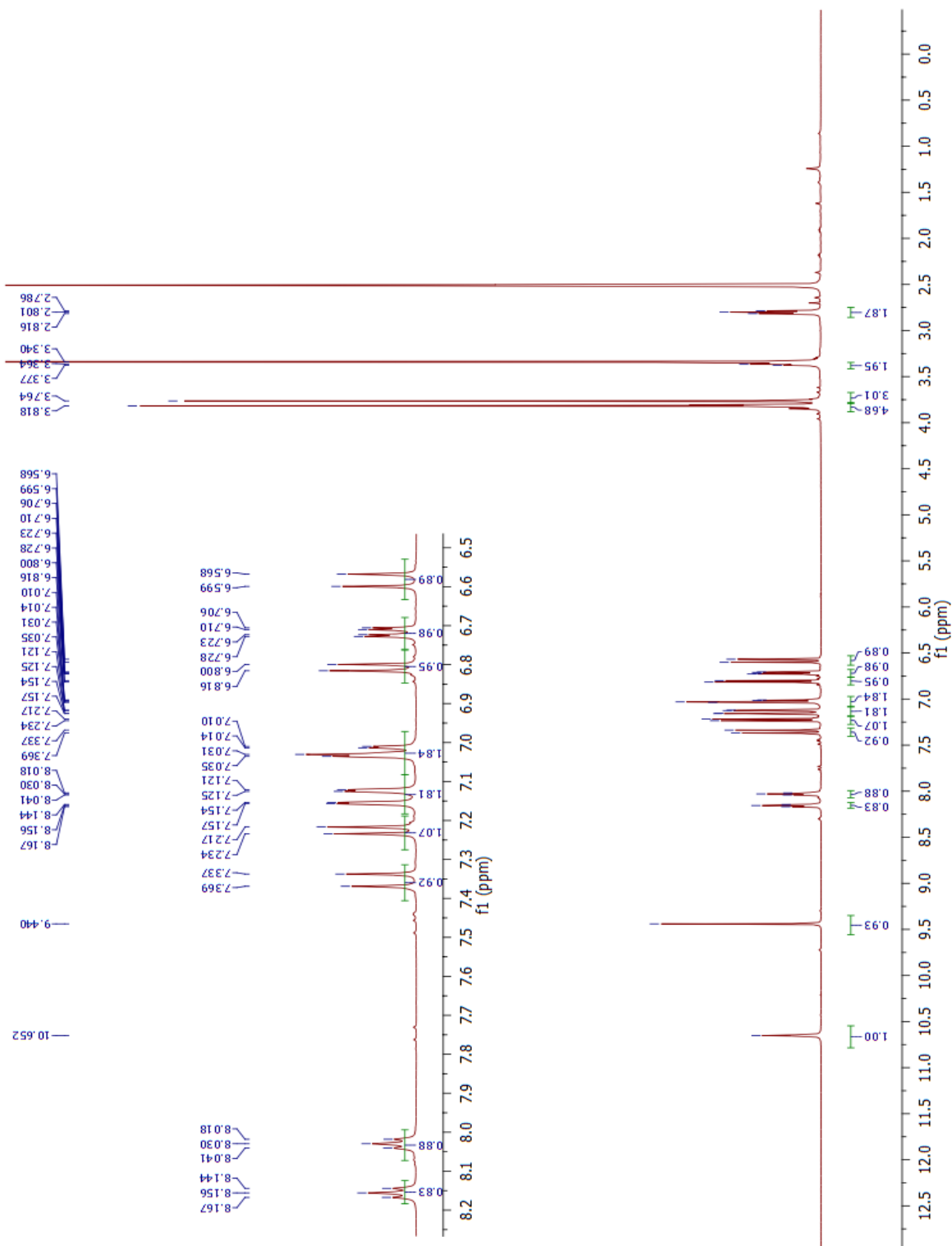
**Figure A8.**  $^{13}\text{C}$  NMR spectra of (*E*)-1-(4-(3-fluorobenzyl)piperazin-1-yl)-3-(4-hydroxy-3-methoxyphenyl)prop-2-en-1-one (**13k.HCl**).



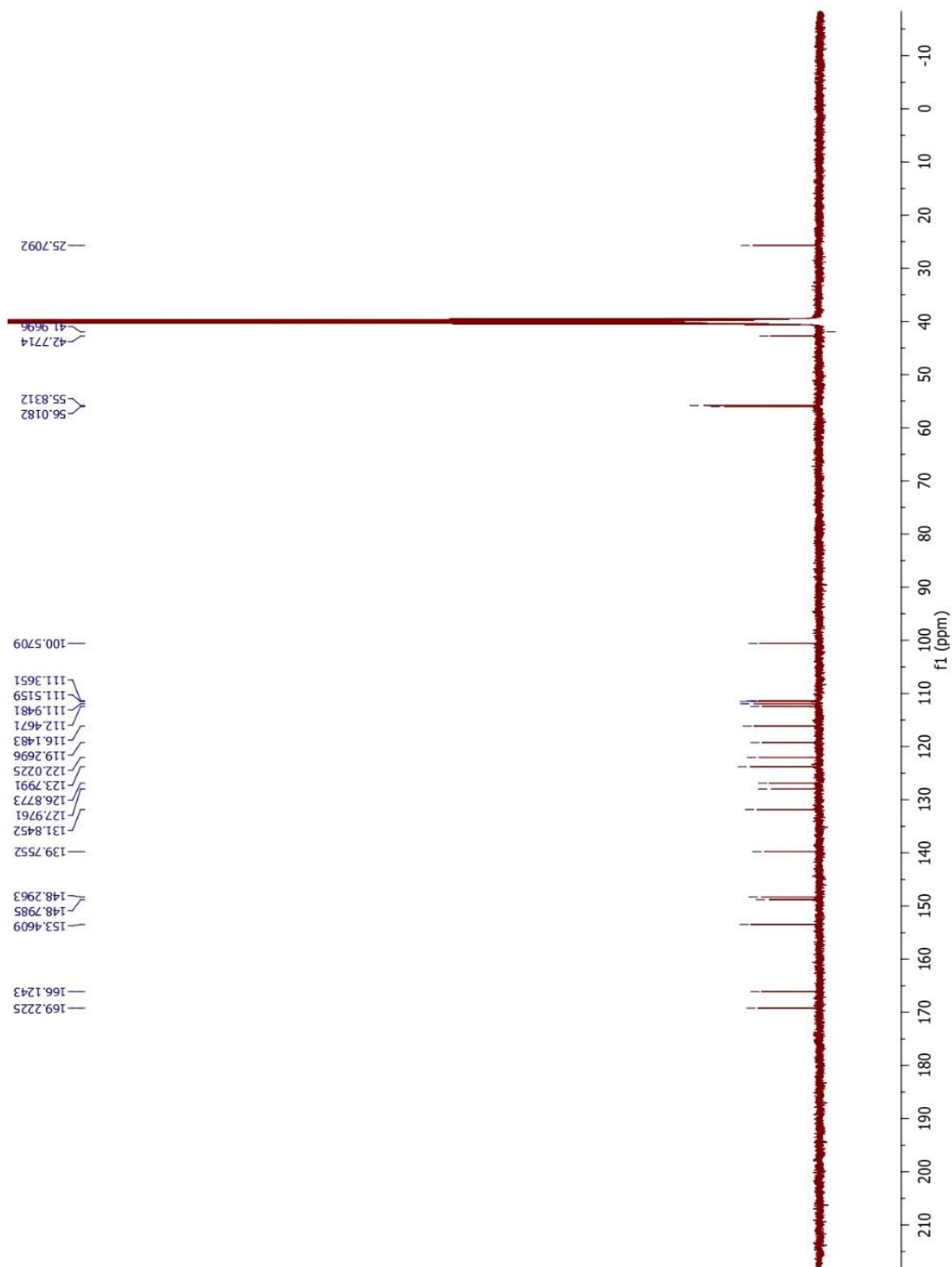
**Figure A9.** HRMS spectra of (*E*)-1-(4-(3-fluorobenzyl)piperazin-1-yl)-3-(4-hydroxy-3-methoxyphenyl)prop-2-en-1-one (**13k.HCl**).



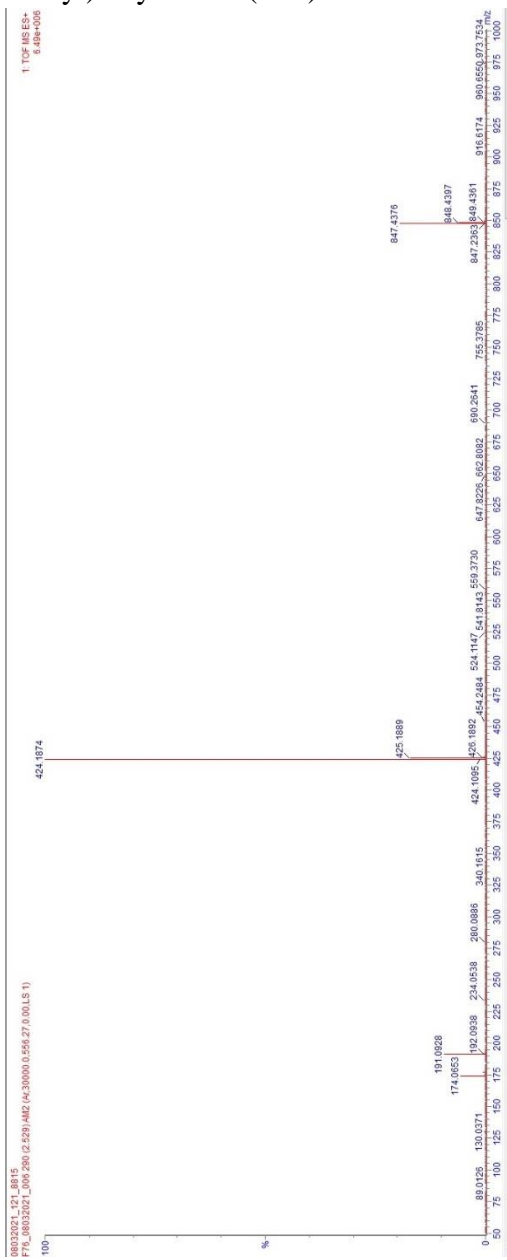
**Figure A10.**  $^1\text{H}$  NMR spectra of (*E*)-3-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((2-(5-methoxy-1*H*-indol-3-yl)ethyl)amino)-2-oxoethyl)acrylamide (**23b**).



**Figure A11.**  $^{13}\text{C}$  NMR spectra of (*E*)-3-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((2-(5-methoxy-1*H*-indol-3-yl)ethyl)amino)-2-oxoethyl)acrylamide (**23b**).



**Figure A12.** HRMS spectra of (*E*)-3-(4-hydroxy-3-methoxyphenyl)-*N*-(2-((2-(5-methoxy-1*H*-indol-3-yl)ethyl)amino)-2-oxoethyl)acrylamide (**23b**)



## List of Publications

### Publications from the research work

1. **Y.P. Singh**, G.N.V.C. Tej, A. Pandey, K. Priya, P. Pandey, G. Shankar, P.K. Nayak, G. Rai, A. G.Chittiboyina, R. J. Doerksen, S. Vishwakarma, G. Modi. Design, synthesis and biological evaluation of novel naturally-inspired multifunctional molecules for the management of Alzheimer's disease. *European journal of medicinal chemistry* (IF 5.57, 2019). 2020;198:112257.
2. **Y. P. Singh**, G. Shankar, S. Jahan, G. Singh, N. Kumar, A. Barik, P. Upadhyay, L. Singh, K. Kamble, G. K. Singh, N. Ranjan, P. Garg, S. Gupta, G. Modi. Further SAR Studies on Natural Template Based Neuroprotective Molecules for the Treatment of Alzheimer's Disease. *Bioorganic and medicinal chemistry* (IF 3.64, 2021, 2021).
3. **Y.P. Singh**, H. Rai, G. Singh, G.K. Singh, S. Mishra, S. Kumar, S. Srikrishna, G. Modi, A Review on Ferulic Acid and Analogs Based Scaffolds for the Management of Alzheimer's Disease, *European Journal of Medicinal Chemistry* (IF 5.57,2021) 113278.
4. **Y.P. Singh**, A. Pandey, S. Vishwakarma, G. Modi. A review on iron chelators as potential therapeutic agents for the treatment of Alzheimer's and Parkinson's diseases. *Molecular Diversity*. 2018 (IF 2.0, 2018).

### Publication under review

1. **Y. P. Singh**, N. Kumar, K. Priya, B. S. Chauhan, G. Shankar, S. Kumar, G. K. Singh, S. Srikrishna, P. Garg, G. Rai, G. Modi. Exploration of Neuroprotective Properties of Naturally Inspired Multifunctional Molecule (F24) Against Oxidative Stress and Amyloid  $\beta$  Induced Neurotoxicity in Alzheimer's Disease Models. *ACS Chemical Neuroscience* (Revised article under review).

### Other Publications

1. N. Kumar, A. Gahlawat, R.N. Kumar, **Y.P. Singh**, G. Modi, P. Garg. Drug repurposing for Alzheimer's disease: in silico and in vitro investigation of FDA-approved drugs as acetylcholinesterase inhibitors, *Journal of Biomolecular Structure and Dynamics*, (2020) 1-15.
2. H. Rai, A. Barik, **Y.P. Singh**, A. Suresh, L. Singh, G. Singh, U.Y. Nayak, V.K. Dubey, G. Modi, Molecular docking, binding mode analysis, molecular dynamics, and prediction of



ADMET/toxicity properties of selective potential antiviral agents against SARS-CoV-2 main protease: an effort toward drug repurposing to combat COVID-19, *Molecular Diversity*, 25 (IF 2.0, 2019) 1905-1927.

#### **List of Patents filed**

1. G. Modi, **Y.P. Singh**, A. Pandey, Development of multifunctional hybrid 3-(4-hydroxy-3-methoxy-phenyl)prop-2-enoic acid and substituted amine /piperazine derivatives for the treatment of neurodegenerative disorders (Patent Application # **201911000362**).
2. G. Modi, **Y.P. Singh**, C. P. Kumar, M. Yadav, G. Shankar, G. Singh, S. Kumar, S Srikrishna. A Multifunctional diarylurea-hydroxyamidine based compound for treatment of Alzheimer disease, **TEMP/E-1/1680/2021-DEL**, Jan, 2021.
3. G. Modi, **Y.P. Singh**, G. Shankr, G. Singh, A. Barik, L. Singh A natural template based anticholinesterase inhibitors and antioxidants for the treatment of Alzheimer's disease, Indian patent application # **202111016470**, Filled on April 07, 2021.

#### **List of conferences attended**

1. Design, synthesis and biological evaluation of novel molecules for the management of Alzheimer's disease, Meenu Yadav\*, Yash Pal Singh, Prince Kumar, Prem Chopra, Amruta Pandey, Gyan Modi, **Indian Academy of Neuroscience, 2018**.
2. Design, Synthesis and Biological Evaluation of Novel Molecules for The Management of Alzheimer's Disease, C.S. Praveen Kumar, Yash Pal Singh, Khusbu Priya, Meenu Yadav, Geeta Rai, Gyan Modi, **ISCB, 2018**.
3. Development of Novel Dual Acting Neuroprotective Agents for the Treatment of Alzheimer's Disease. Yash Pal Singh, Amruta Pandey, Swati Vishwakarma, Gyan Modi\* (**EMBO workshop** to be held from August 29- September 01, 2018).
4. Design, synthesis and characterization of novel molecules used in the treatment of Alzheimer disease" Yash Pal Singh, Amruta Pandey, Swati Vishwakarma and Gyan Modi (**ETDDD, 2018, IIT (BHU), Varanasi**).
5. Participated in seminar on "Recent Applications of Carbohydrates in Chemistry and Biology (RACCB-2017)" organized by **IIT BHU, Varanasi in Feb 14-16, 2017**
6. Participated in workshop on "Emerging Trends in Drug Design and Molecular Modelling" organized by DIC, **IIT BHU, Varanasi** in 16-21 July 2017.

## Curriculum Vitae

**YASH PAL SINGH**

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Indian Institute of Technology (BHU), Varanasi, India,**

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**Email: yashps.rs.phe16@itbhu.ac.in**

### Academic Qualification

Year	Degree	Institute	CGPA/% AGE
2016-21	Ph.D. thesis submitted (Pharmaceutical Chemistry)	IIT (BHU), Varanasi	8.75/10
2016	M.S. (Medicinal Chemistry)	NIPER-Kolkata	8.96/10
2013	B.Pharmacy	GPC, Rohru	69.75%
2009	CLASS XII	HPBOSE, Dharmshala	69.56%
2007	CLASS X	HPBOSE, Dharmshala	70.62%

### RESEARCH EXPERIENCE

Doctor of Philosophy (PhD) .....July 2016-Sep. 2021

Supervisor: Dr. Gyan Prakash Modi

Dept. of Pharma. Engg. & Tech. IIT (BHU), Varanasi, INDIA

Thesis: "Design, Synthesis, and Biological Evaluation of Ferulic Acid Template Based Naturally Inspired Novel Neuroprotective Multifunctional Molecules for the Treatment of Alzheimer's Disease"

Master of Science (Pharma).....June 2014-July 2016

Supervisor: Prof. Uday Bandyopadhyay (IICB-India), Director Bose Institute, Kolkata (Project work), INDIA.

Supervisor: Prof. V. Ravichandiran, Director NIPER-Kolkata (course work), INDIA.

Thesis: "A novel synthetic route towards the synthesis of 11*H* Benzo[a] Carbazole derivatives by using Palladium catalyzed C-H activation".

### PUBLICATIONS

1. **Y. P. Singh**, N. Kumar, K. Priya, B. S. Chauhan, G. Shankar, S. Kumar, G. K. Singh, S. Srikrishna, P. Garg, G. Rai, G. Modi. Exploration of Neuroprotective Properties of Naturally Inspired Multifunctional Molecule (F24) Against Oxidative Stress and Amyloid  $\beta$  Induced Neurotoxicity in Alzheimer's Disease Models, *ACS Chemical Neuroscience*, 2021 (Under minor revision)

2. **Y. P. Singh**, G. Shankar, S. Jahan, G. Singh, N. Kumar, A. Barik, P. Upadhyay, L. Singh, K. Kamble, G. K. Singh, N. Ranjan, P. Garg, S. Gupta, G. Modi. Further SAR Studies on Natural Template Based Neuroprotective Molecules for the Treatment of Alzheimer's Disease, *Bioorganic and Medicinal Chemistry*, 2021, 116385.
3. **Y.P. Singh**, H. Rai, G. Singh, G.K. Singh, S. Mishra, S. Kumar, S. Srikrishna, G. Modi, A Review on Ferulic Acid and Analogs Based Scaffolds for the Management of Alzheimer's Disease, *European Journal of Medicinal Chemistry*, (2021) 113278.
4. H. Rai, A. Barik, **Y. P. Singh**, A. Suresh, L. Singh, G. Singh, U. Y. Nayak, V. K. Dubey, G. Modi. Molecular docking, binding mode analysis, molecular dynamics, and prediction of ADMET/toxicity properties of selective potential antiviral agents against SARS-CoV-2 main protease: an effort toward drug repurposing to combat COVID-19, *Molecular Diversity*, 2021.
5. **Y.P. Singh**, G.N.V.C. Tej, A. Pandey, K. Priya, P. Pandey, G. Shankar, P.K. Nayak, G. Rai, A.G. Chittiboyina, R.J. Doerksen, S. Vishwakarma, G. Modi, Design, synthesis and biological evaluation of novel naturally-inspired multifunctional molecules for the management of Alzheimer's disease, *European Journal of Medicinal Chemistry*, 198 (2020) 112257.
6. N. Kumar, A. Gahlawat, R.N. Kumar, **Y.P. Singh**, G. Modi, P. Garg, Drug repurposing for Alzheimer's disease: in silico and in vitro investigation of FDA-approved drugs as acetylcholinesterase inhibitors, *J Biomol Struct Dyn*, (2020) 1-15.
7. **Y.P. Singh**, A. Pandey, S. Vishwakarma, G. Modi, A review on iron chelators as potential therapeutic agents for the treatment of Alzheimer's and Parkinson's diseases, *Molecular Diversity*, 2018,.

#### PATENT FILED

1. A multifunctional diarylurea-hydroxyamidine based compound for treatment of Alzheimer disease, G. modi, Y. P. Singh, C. P. Kumar, M. Yadav, Gauri Shankar, Gourav Singh, Saroj Kumar, S Srikrishna. temp-1/1680/2021-del, 12 jan, 2021.
2. Development of multifunctional hybrid 3-(4-hydroxy-3-methoxy-phenyl)prop-2-enoic acid and substituted amine/piperazine derivatives for the treatment of neurodegenerative disorders G. Modi, Y. P. Singh, and A. Pandey (Patent Application # 201911000362).

3. G. Modi, Y.P. Singh, G. Shankr, G. Singh, A. Barik, L. Singh A natural template based anticholinesterase inhibitors and antioxidants for the treatment of Alzheimer's disease, Indian patent application # 202111016470, Filled on April 07, 2021.

### AREA OF INTEREST

Medicinal Chemistry, Process Chemistry.

Chemistry of Natural Products, Pharmacognosy.

### KEY SKILLS AND KNOWLEDGE

- Synthesis, purification and characterization of novel intermediates.
- Ability to deliver small to large quantities of novel chemical entities for drug discovery.
- Knowledge of functional group conversion and multiple step synthesis.
- Knowledge of retro- synthetic analysis and handling exposure of various pyrophoric reactions.
- Exposure to TLC, Column Purification, Extraction and Analysis of NMR, IR and Mass Spectrometry Data.
- Good understanding of Microsoft Office and Structure drawing (Chem Draw/ISIS) and literature search using SCI Finder, etc.
- Good understanding of GraphPad prism, Kinetica, EndNote, and Autodock software.
- Animal dissection skills: Organ isolation (Brain, liver, kidney)
- Behavioral assessment (Y maze, Morris water maze, Passive avoidance)

### SCHOLASTIC ACHIEVEMENTS

- GPAT qualified in 2013 and 2014
- NIPER JEE qualified 2014 (M.S. pharma admission)
- NIPER JEE qualified 2016 (Ph.D. admission)
- IIT BHU entrance qualified in 2016 (Ph.D. admission)
- Received M.H.R.D. fellowship from AICTE, New Delhi.
- Received M.H.R.D. fellowship during Ph.D. (2016-2021)

### WORK EXPERIENCE

Worked in Smilex Pharma. for one-month industrial training.

### SEMINARS & WORKSHOPS ATTENDED:

- Design, synthesis and biological evaluation of novel molecules for the management of Alzheimer's disease, Meenu Yadav\*, Y. P. Singh, Prince Kumar, Prem Chopra, Amruta Pandey, Gyan Modi, **Indian Academy of Neuroscience**, 2018.

- Design, Synthesis and Biological Evaluation of Novel Molecules for The Management of Alzheimer’s Disease, C.S. Praveen Kumar, Y. P. Singh, Khusbu Priya, Meenu Yadav, Geeta Rai, Gyan Modi, **ISCB**, 2018.
- Development of Novel Dual Acting Neuroprotective Agents for the Treatment of Alzheimer’s Disease Y. P. Singh, Amruta Pandey, Swati Vishwakarma, Gyan Modi\* (**EMBO** workshop to be held from August 29- September 01, 2018)
- Design, synthesis and characterization of novel molecules used in the treatment of Alzheimer disease” Amruta Pandey, Swati Vishwakarma, Y. P. Singh and Gyan Modi (ETDDD, 2018, **IIT (BHU)**, Varanasi).
- Participated in seminar on “Recent Applications of Carbohydrates in Chemistry and Biology (RACCB-2017)” organized by IIT BHU, Varanasi in Feb 14-16, 2017
- Participated in workshop on “Emerging Trends in Drug Design and Molecular Modelling” organized by DIC, IIT BHU, Varanasi in 16-21 July 2017
- Participated in workshop on “Metals in Genetics Chemical Biology and therapeutics” organized by Indian Institute of Science Bangalore in Feb 17-20, 2016.
- Participated in seminar on “Molecules from Laboratory to Clinics” Organised by NIPER-Kolkata in February 2015.
- Participated in seminar on “Integrated approach for promotion and development of herbal medicine” organized by School of Natural Product Studies, JU-Kolkata, Nov 24, 2015.
- Participated in workshop on “Laboratory Safety, Radiation Safety, Chemical Safety and Bio Safety” organized by IICB-Kolkata, Kolkata in September, 2014.

#### LANGUAGES KNOWN

- English
- Hindi
- Pahari

#### HOBBIES

- Playing Cricket, Volleyball
- Traveling

#### PERSONAL DETAILS

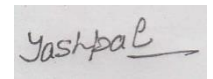
Father’s name : Om Prakash  
 Mother’s name : Urmila Devi  
 Dob : 01/01/1992

Gender : Male  
Religion : Hindu  
Blood group : A+  
Address- Vill-Samnos, PO-Jachh, Teh- Chachyot Mandi (HP), 175039

DECLARATION

I hereby declare that the above said particulars are true to the best of my knowledge and belief.

Date: 05/08/2021  
Place: Varanasi, U.P.

A rectangular box containing a handwritten signature in black ink that reads "Yashpal" with a horizontal line extending to the right.

(Yash Pal Singh)