

CURRICULUM VITAE Leila Nematollahi

Born in September 20th 1977

Current position:

Associate Professor Department of Medical Biotechnology, Biotechnology Research Center Pasteur Institute of Iran

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Education

Ph.D. "Pharmaceutical Biotechnology"; Pasteur Institute of Iran, 2006-2012. Pharm.D.; Pharmacy School, Tehran University of Medical Science, 1996-2002. Science Diploma, Farzanegan high school, 1991-1995.

Honors

Selected as top student in Pasteur Institute of Iran, research day ceremony, 2010 Top student in Farzanegan high school, 1995.

Pharm. D. Thesis

Title: Application of a high performance liquid chromatography (HPLC) method to the analysis of Thiamine, Riboflavin and Niacin in Iranian pasta.

Supervisors: Dr.M. Hajimahmoodi, Dr. B.Jannat.

Ph. D. Thesis

Title: Expression of a novel human BMP-7 variant with enriched heparin binding site in Chinese hamster ovary (CHO) cells and evaluation of its biological activity.

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Supervisors: Dr. F.Mahboudi, Dr.V.Khalaj.

Publications

- 1. Optimization of the Expression of Recombinant Cetuximab Single-Chain Fragment Variable and Comparative its Purification with Magnetic Nanoparticles and Conventional Fast Protein Liquid Chromatography. Jalalvand M, Falahzadeh kh, Jalalvand A, Mazloumi M, Bakhshandeh H, Shahsavari G, Bayat E, Veisi Malekshahi Z, Nematollahi L, Negahdari B.Biointerface Research in Applied Chemistry; 2023; 13(6): 592.
- 2. Recombinant anti-human epidermal growth factor receptor type 2 single-chain variable fragment-alpha-luffin fusion protein as a putative immunotoxin against human epidermal growth factor receptor type 2-positive breast cancer cells: an experimental research. Damough S, Bayat E, Bakhshaiesh TO, Barkhordari F, Esmaeili R, Nematollahi L, Talebkhan Y. Annals of Medicine & Surgery; 2023; 85 (9): 4348-4354.
- 3. Expression, Purifcation, and Biological Evaluation of XTEN-GCSF in a Neutropenic Rat Model.Nikravesh FY, Gholami P, Bayat E, Talebkhan Y, Mirabzadeh E, Damough S, Aghamirza Moghim Aliabadi H, **Nematollahi L**, Hosseinzadeh Ardakani Y. Applied Biochemistry and Biotechnology; 2023.
- Identification of two neutralizing human single-chain variable fragment antibodies targeting *Staphylococcus aureus* alpha-hemolysin. Piri-Gavgani S, Ghanei M, Fateh A, Siadat D, Nematollahi L, Rahimi-Jamnani F. Iranian Journal of Basic Medical Sciences; 2022; 25 (10): 1207–1214.
- Computational analysis of fusion protein of anti-HER2 scFv and alpha luffin: A new immunotoxin protein for HER2 positive cancers. Barkhordari F, Rismani E, Tabasinezhad M, Asgari S, Nematollahi L, Talebkhan Y. Brazilian Journal of Pharmaceutical Sciences; 2022; 58:1-12.

- 6. Extension of human GCSF serum half-life by the fusion of albumin binding domain. Nikravesh FY, Shirkhani S, Bayat E, Talebkhan Y, Mirabzadeh E, Sabzalinejad M, Aliabadi HAM, **Nematollahi L**, Ardakani YH, Sardari S. *Sci Rep*; 2022; 12 (1):667.
- Production of an Antibody Fragment (scFv) Targeting PcrV Protein of Pseudomonas aeruginosa in Fed-Batch Cultivation Mode. Karam S, Raigani M, Hassani Afshar S, Talebkhan Y, Bayat E, Komijani S, Nematollahi L, Barkhordari F, Shafiee Ardestani M, Davami F. *IBJ*; 2021; 25 (6): 390-398.
- 8. Production of soluble and functional anti-TNF-α Fab' fragment in cytoplasm of E. coli: Investigating the effect of process conditions on cellular biomass and protein yield suing response surface methodology. Talaei A, Mazaheri S, Bayat E, Bakhshandeh B, Sabzalinejad M, Damough Sh, Mahboudi F, **Nematollahi L**, Talebkhan Y. *Protein J*; 2021; 40 (5): 786-798.
- Characterization of a novel mCH3 conjugated anti-PcrV molecule. Komijani S, Bayat
 E, Rismani E, Hosseini S, Moazzami R, Nematollahi L, Sardari S, Talebkhan Y,
 Davami F, Barkhordari F, Hosseini F, Jahandar H. Sci Rep; 2021; 11: 7154.
- 10. Optimization of culture conditions for high-level expression of soluble and active tumor necrosis factor-α in E. coli. Damough Sh, Sabzalinezhad M, Talebkhan Y, **Nematollahi L**, Bayat E, Torkashvand F, Adeli A, Jahandar H, Barkhordari F, Mahboudi F. *Protein Expr Purif*; 2021; 179 (105805); 1-8.
- 11. Multilayer Alginate Microcapsules For Live Cell Microencapsulation; Is There Any Preference For Selecting Cationic Polymers? Fariba Hajifathaliha, Arash Mahboubi, Noushin Bolourchian, Elham Mohit and Leila Nematollahi, Iranian Journal of Pharmaceutical Research; 2021; 20 (2): 173-182.
- 12. Woodchuck Hepatitis Virus Post-Transcriptional Regulation Element (WPRE) Promotes Anti-CD19 BiTE Expression in Expi293 Cells, Reza Moazzami, Hasan Mirzahoseini, **Leila Nematollahi**, Farzaneh Barkhordari, Mozhgan Raigani, Fatemeh Hajari Taheri, Fereidoun Mahboudi1and Fatemeh Davami, Iranian Biomedical Journal; 2021; 25 (4): 275-283.
- 13. Different Respiratory Samples for COVID-19 Detection by Standard and Direct Quantitative RT-PCR: A Literature Review, Maryam Ahmadzadeh, Hossein Vahidi, Arash Mahboubi, Fariba Hajifathaliha, **Leila Nematollahi** and Elham Mohit, Iranian Journal of Pharmaceutical Research; 2021; 20 (3): 285-299.

- 14. Production of an Antibody Fragment (scFv) Targeting PcrV Protein of *Pseudomonas aeruginosa* in Fed-Batch Cultivation Mode. Saba Karam, Mozhgan Raigani, Sahar Hassani Afshar, Yeganeh Talebkhan, Elham Bayat, Samira Komijani, Leila Nematollahi, Farzaneh Barkhordari, Mehdi Shafiee Ardestani and Fatemeh Davami, Iranian Biomedical Journal; 2021;25 (6): 390-398.
- 15. Computational analysis of fusion protein of anti-HER2 scFv and alpha luffin: A new immunotoxin protein for HER2 positive cancers. Barkhordari F, Rismani E, Tabasinezhad M, Asgari S, Nematollahi L, Talebkhan Y. Braz J Pharm Sci; 2020; Accepted.
- 16. Improvement of Certolizumab Fab' properties by PASylation technology. Mazaheri S, Talebkhan Y, Mahboudi F, Nematollahi L, Ahangari Cohan R, Mirabzadeh Ardakani E, Bayat E, Sabzalinejad M, Sardari S, Torkashvand F. Sci Rep; 2020; 10 (1):18464.
- 17. Comparison of Linear Poly Ethylene Imine (LPEI) and Poly L-Lysine (PLL) in Fabrication of CHOK₁ Cell-Loaded Multilayer Alginate. Fariba Hajifathaliha, Arash Mahboubi, Elham Mohit, Noushin Bolourchian, Vahid Khalaj, Leila Nematollahi, Advanced Pharmaceutical Bulletin; 2020; 10(2): 290-296.
- 18. Comparison of different cationic polymers efficacy in fabrication of alginate multilayer microcapsules. Fariba Hajifathaliha, Arash Mahboubi, **Leila Nematollahi**, Elham Mohit, Noushin Bolourchian, Asian Journal of Pharmaceutical Sciences ;2020; 15: 95-103.
- 19. Optimization of HER2-based and cell-based ELISA for detection of trastuzumab biosimilar, Maryam Ahmadzadeh, Farzaneh Farhadi, Leila Nematollahi, Shayan Maleknia, Elham Mohit, koomesh journal; 2020; 22(2):341-350.
- 20. Cloning and soluble expression of mature α-luffin from *Luffa* cylindrica in *E. coli* using SUMO fusion protein, Shaghayegh Namvar, Farzaneh Barkhordari, Mozhgan Raigani, Hoda Jahandar, **Leila Nematollahi**, Fatemeh Davami. Turkish Journal of Biology; 2018;42 (1): 23-32.
- 21. Effects of peptone supplementation in different culture media on growth, metabolic pathway and productivity of CHO DG44 cells; a new insight into amino acid profiles, Davami Fatemeh, Eghbalpour Farnaz, **Nematollahi Leila**, Barkhordari Farzaneh, Mahboudi Fereidoun . Iranian Biomedical Journal;2015; 19 (4):194-205.

- 22. Effect of cysteamine on cell growth and IgG4 production in recombinant Sp2.0 cells. Jahandar H, Vaziri B, **Nematollahi L**, Afsharirad T, Torkashvand F, Khalaj V. Iranian Journal of Pharmaceutical Research; 2015; 14(1):177-87.
- 23. The Effects of Recombinant IgG4 Expression on the Growth of Sp2.0 Cell Lines, Authors: Hoda Jahandar, Vahid Khalaj, **Leila Nematollahi**, Fatemeh Zandi, Somayeh Enayati, Behrouz Vaziri, Modares Journal of Medical Sciences: Pathobiology ;2014; 16(4): 27-37.
- 24. Enhancement of Recombinant Human Tissue Plasminogen Activator Expression in CHO Cells using Matrix Attachment Region Containing Vectors in Combination with Promoter Activation Strategy, Authors: Azam Rahimpour, Behrouz Vaziri, Farzaneh Barkhordari, **Leila Nematollahi**, Ahmad Adeli, Fereidoun Mahboudi, Modares Journal of Medical Sciences: Pathobiology;2013; 16(1): 11-21.
- 25. Engineering the Cellular Protein Secretory Pathway for Enhancement of Recombinant Tissue Plasminogen Activator Expression in Chinese Hamster Ovary cells: Effects of CERT and XBP1s Genes, Authors: Rahimpour A, Vaziri B, Moazzami R, Nematollahi L, Barkhordari F, Kokabee L, Adeli A, Mahboudi F, J Microbiol Biotechnol;2013; 23(7): 1116-1122.
- 26. A novel human Bone Morphogenetic Protein-7 variant with enriched heparin-binding site, Authors: **Leila Nematollahi**, Fereidoun Mahboudi, Azam Rahimpour, Hoda Jahandar, and Vahid Khalaj, Molecular Biology; 2013; 47(3): 399-405.
- 27. The comparison of Bone Morphogenetic Protein-7 mutant expression in the prokaryotic and eukaryotic hosts, Authors: **Leila Nematollahi**, Fereidoun Mahboudi, Mohammad Azizi, Farzaneh Barkhordari, Ahmad Adeli, and Vahid Khalaj, Modares Journal of Medical Sciences: Pathobiology;2012; 15(3): 93-102.
- 28. Periplasmic Expression of a Novel Human Bone Morphogenetic Protein-7 Mutant in Escherichia coli , Authors: Leila Nematollahi , Vahid Khalaj , Seyedeh Maliheh Babazadeh, Azam Rahimpour, Hoda Jahandar, Fatemeh Davami , and Fereidoun Mahboudi , Avicenna J Medical Biotechnology;2012; 4(4): 178–185.

Projects

- 1. Preparation and characterization of niosomal nano-carriers containing Oxaliplatin targeted by Cetuximab derived scFv and evaluation of their in vitro cytotoxicity on human colorectal cancer cell line. (Co-funded by: Pasteur Institute of Iran, Tehran, Iran & Tehran University of Medical Sciences, 2021).
- Expression and characterization of single chain antibody fragments against TNF-α and IL-6 receptor alone and together as bispecific antibody in E. coli. (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2021).
- 3. Expression and characterization of single chain antibody fragments against TNF-α and IL-6 receptor alone and together as bispecific antibody in mammalian expression system. (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2021).
- 4. Expression of α-luffin immunotoxin conjugated to anti-HER2 scFv in E. coli cell and study of its inhibitory effects on breast cancer cell lines (Co-Funded by: Pasteur Institute of Iran, Tehran, Iran (2017) & Iran National Science Foundation, 2018).
- 5. Site-specific integration of GFP reporter gene in CHO cells mediated by CRISPR/Cas9 system. (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2018).
- 6. Production of an Antibody fragment (V2L2) with improved half-life against PcrV protein of Pseudomonas aeruginosa in mammalian HEK and prokaryotic E. coli host. (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2016).
- 7. Efficacy comparison of different cationic polymers in production of multilayer alginate microcapsules. (Funded by: Shahid Beheshti University of Medical Sciences Tehran, Iran, 2016).
- 8.Construction a recombinant expressing vector containing single chain variable fragment (scFv) against HER2 linked to IFN-gamma inducible protein of 10 kDa (IP-10) chemokine. (Funded by: Shahid Beheshti University of Medical Sciences Tehran, Iran, 2016).
- 9. Tracking intercellular mechanisms involved in solubility increase of anti-VEGF Fab Fragment expressed in a fed-batch cultivation mode in E. coli.(Funded by: Pasteur Institute of Iran, Tehran, Iran, 2015).

- 10. Construction of an expression vector encoding single chain variable fragment (scFv) against Human Epidermal Growth receptor 2 (HER2). (Funded by: Protein technology research center, Shahid Beheshti University of Medical Sciences Tehran, Iran, 2015).
- 11. Expression and purification of single chain variable fragment (scFv) against (Human Epidermal Growth 2 receptor (HER2), (Funded by: Shahid Beheshti University of Medical Sciences Tehran, Iran, 2015).
- 12.Expression of hBMP-7 (Human Bone Morphogenetic Protein-7) in E.coli using SUMO fusion protein and evaluation of its biological activity, (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2014).
- 13.Cloning of estrogen receptor alpha ($ER\alpha$) in GFP containing plasmid and evaluation of its expression in COS-7 cells, (Funded by: Faculty of pharmacy, Shahid Beheshti University of Medical Sciences, 2014).
- 14.Periplasmic expression of a novel human BMP-7 mutant in Escherichia coli, (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2010).
- 15. Transient and stable expression of a novel human BMP-7 variant in Sp2/0 cell lines, (Funded by: Pasteur Institute of Iran, Tehran, Iran, 2009).

> National grants:

- 1. Development of SP2/0 stable mammalian cell line producing biosimilar Golimumab anti-TNF alpha monoclonal antibody.
- 2. Development of engineered E. coli cells producing Fab antibody fragment of Certolizumab pegol.

(Funded by: Technological development department of presidential office, Tehran, Iran, 2016

Published Book:

- Monoclonal Antibodies: a novel challenge in the field of Biotechnology Authors: Dr.F.Davami, A.Rahimipour, **Dr. L.Nematollahi**, Dr.H.Jahandar, 2012.

Patent:

- Design and expression of a novel human Bone Morphogenetic Protein-7 variant with enriched heparin affinity in CHO cells, **patent No: 86057, 2015**.

Ph.D thesis supervisor:

- ➤ Preparation and characterization of niosomal nanocarriers containing Oxaliplatin targeted by Cetuximab derived scFv and evaluation of their in vitro cytotoxicity on human colorectal cancer cell line
- Fabrication of different alginate-based polyelectrolyte complexes (PECs) and evaluation of their characteristics in cell microencapsulation
- ➤ Identification and characterization of Fully Human scFv Antibody Inhibiting Staphylococcus aureus Alpha-Hemolysin
- Epression of PASylated form of anti-tumor necrosis factor alpha Fab' antibody fragment and evaluation of its physicochemical and biological properties.

Master thesis supervisor:

- > Preparation and characterization of polysaccharide nanoparticles containing recombinant alfa luffin and its effect on breast cancer cell line
- Expression of hBMP-7 (Human Bone Morphogenetic Protein-7) in *E.coli* using SUMO fusion protein and evaluation of its biological activity.
- ➤ Cloning and Expression of hBMP-7 (Human Bone Morphogenetic Protein-7) in Shuffle: a novel *E. coli* protein expression strain.
- ➤ Cloning and expression of an Antibody fragment (V2L2) with improved half-life against PcrV protein of Pseudomonas aeruginosa in prokaryotic E. coli host.
- > Cloning and comparison expression of a humanized anti-TNF-α antibody fab fragment (Certolizumab) in *E.Coli* hosts

Ph.D thesis advisor:

Synthesis of Oxaliplatin-PEG-PLGA nanopolymer particles incorporated with Cetuximab monoclonal antibody derived scFv against epidermal grwoth factor

- receptor (EGFR) and evaluation of its *in vitro* cytotoxicity on human colorectal cancer cell line.
- ➤ Expression of an CD3/CD19 bispecific antibody in HEK 293 cell line and evaluation of the effect of WPRE element on expression level.
- Expression of a fusion protein containing scFv against HER2 linked to IP-10 and *in vitro* evaluation of its biological activity
- ➤ Optimization of expression and pharmacokinetic and pharmacodynamic studies of fusion derivatives of GCSF proteins in a Rat model

Pharm.D & Master thesis advisor:

- ➤ Production of an Antibody fragment (V2L2) with improved half-life against PcrV protein of Pseudomonas aeruginosa in prokaryotic E. coli host by using SUMO-tag in fed-batch cultivation system mode.
- ➤ Production of an antibody fragment (ScFV) against amyloid-beta for Alzheimer's disease in prokaryotic E.coli host by using SUMO-tag in fed batch cultivation system mode.
- ➤ Cloning and soluble expression of mature a-luffin from Luffa cylindrical in *E.coli* using SUMO fusion protein and rehamnose tunable promoter system.
- \triangleright Construction of recombinant plasmid encoding estrogen receptor alpha (ER α) and evaluation of its expression in COS-7 cells

Teaching Experiences

(2017-until now) " Novel Drug Delivery" for Ph.D students of Medical Biotechnology (Pasteur Institute of Iran)

(2016-until now) " Cell Biology " for Ph.D students of Medical Biotechnology and Pharmaceutical Biotechnology (Pasteur Institute of Iran)

(2015-until now) " Protein Chemistry" for Ph.D students of Pharmaceutical Biotechnology (Pasteur Institute of Iran)

(2015-until now) "protein engineering" for Ph.D students of Medical Biotechnology (Pasteur Institute of Iran)

(2014-until now) "Advanced genetic engineering" for Ph.D students of Pharmaceutical Biotechnology (Pasteur Institute of Iran)

(2018) "Introduction to Biotechnology" for Master students of Pharmaceutical Chemistry, Islamic Azad University

(2003-2005) "Workshop on Rational Drug Use" for Medical Students:

- Tehran University of Medical Sciences & Health Services
- Iran University of Medical Sciences & Health Services
- Birjand University of Medical Sciences & Health Services
- Ardabil University of Medical Sciences & Health Services

(1996-1999) "Chemistry" for the university entrance exam

Professional experiences:

- Stable and Transient Expression in mammalian cells (CHO, Sp2/0)
- Expression in prokaryote systems (*E.coli*)
- Cloning
- Expert in Cell culture techniques
- Purification of Recombinant Proteins

Congress Presentations:

- Comparison of linear poly ethylene imine (LPEI) and poly l-lysine (PLL) in fabrication of CHOK1 cell-loaded multilayer alginate microcapsules . 18th Annual Congress on Pharmaceutics & Drug Delivery Systems. 2019. Amsterdam, Netherlands.
- Preparation and Characterization of Different Multilayer Alginate Microcapsules.
 Pharmacy Updates, Shahid Beheshti University of Medical Sciences. 2018.Iran.
- Cloning and expression of V2L2 antibody fragment against *Pseudomonas aeruginosa* PcrV protein in *E. coli* prokaryotic host. **7**th **FEMS**. 2017. Spain.
- Purification and in vitro biological assessment of V2L2 antibody fragment against
 Pseudomonas aeruginosa PcrV protein. 10th National Biotechnology Congress.

 2017. Iran.

- Improving soluble recombinant proteins formation by expression at reduced temperature in specialized *E. coli* strain. **2**nd **international and 10**th **national Biotechnology Congress**. 2017. Iran.
- SUMO fusion protein effect on α-luffin expression in E. coli host. **The 20th Iranian Pharmacy Students' Seminar.** 2017. Iran.
- Influence of the *Escherichia coli* 's host on the solubility of recombinant proteins. 17th

 International and Iranian congress of Microbiology. 2016.Iran.
- Expression of recombinant α-luffin in Escheriachia coli. The 17th International & Iranian Congress of microbiology. 2016. Iran.
- Temperature effect on αluffin expression and solubility rate in Escheriachia coli bacteria as a host. **The 17th International & Iranian Congress of microbiology**. 2016. Iran.
- Anti-HER2 scFv expression in SHuffle host. 12th International Congress on Breast Cancer. 2016.Iran.
- Expression and Purification of hBMP-7 (Human Bone Morphogenetic Protein-7) in
 E.coli using SUMO fusion partner. The 16th International and Iranian Congress of
 Microbiology. 2015. Iran.
- Cloning and Expression of hBMP-7 (Human Bone Morphogenetic Protein-7) in Shuffle: a novel *E. coli* protein expression strain. The 16th International and Iranian Congress of Microbiology. 2015. Iran.
- Application of a liquid chromatography method to the analysis of thiamin, riboflavin and niacin in Iranian pasta. Hajimahmoodi Mannan, 10th Iranian pharmaceutical Sciences Conferece. 2006. Iran.
- Study on Adverse Drug Reactions, due to antibiotics in the cases reported to ADR monitoring center from 1998 to 2001. **The 8th Iranian Seminar of Pharmaceutical Sciences (ISPS8)**. 2002.Iran.

Work Experiences

2014-until now

Assistant Professor

Department of Medical Biotechnology, Biotechnology Research Center

Pasteur Institute of Iran

2005 - 2006

Sobhan Pharmaceutical Co.

Main duties and responsibilities:

- Finding new products for local and under license production
- Feasibility study of new product developments specially for anti-cancer products

2004-2005

RDU Committee, Food and Drug administration Main duties and responsibilities:

- Research on Irrational Use of Drugs
- Statistical analysis

2007-2004

Adverse Drug reaction (ADR) monitoring center, Food and Drug administration, Tehran,Iran

Workshop:

- > Scientific director and lecturer of "Workshop in targeted drug delivery systems", 2027, Pasteur Institute of Iran.
- Scientific director and lecturer of "Workshop in strain engineering for improved expression of recombinant proteins in Escherichia coli", 2021, Pasteur Institute of Iran.
- > Scientific director and lecturer of "Workshop in post-translational modifications of protein biopharmaceuticals", 2021, Pasteur Institute of Iran.
- > Lecturer of "Workshop in Production of therapeutic antibodies by phase display technology", 2021, Pasteur Institute of Iran.
- Lecturer of "Workshop in expression of monoclonal antibodies in eukaryotic expression systems", 2016, Pasteur Institute of Iran.