Nicholas Stephanopoulos

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The Biodesign Institute, Room A120B Arizona State University 727 E. Tyler St., Tempe, AZ 85287

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EMPLOYMENT

Arizona State University, Tempe

2015 - present

Assistant Professor: School of Molecular Sciences,

The Biodesign Institute: Center for Molecular Design and Biomimetics Graduate Faculty: School of Biological and Health Systems Engineering

Graduate Faculty: Chemical Engineering (SEMTE)

Affiliate Faculty: The Biomimicry Center

Affiliate Faculty: The Global Security Initiative (GSI)

Research interests: Protein/peptide-DNA nanotechnology for novel materials, with applications in biology, medicine, energy, engineering, and nano-robotics

EDUCATION AND TRAINING

Northwestern University, Chicago

2011 - 2015

Postdoctoral research

Simpson Querrey Institute for BioNanotechnology (SQI)

Advisor: Prof. Samuel I. Stupp

Research focus: Peptide-DNA biomaterials

University of California, Berkeley

2005 - 2010

Ph.D. in Chemistry (2010)

Thesis advisor: Prof. Matthew B. Francis

Dissertation title: "Integrated Nanosystems Templated

by Self-assembled Virus Capsids"

GPA: 4.0/4.0

Massachusetts Institute of Technology

2004 - 2005

Master of Science in Chemical Engineering Practice, MSCEP (2007)

GPA: 3.9/4.0

Harvard University

2000 - 2004

A.B. in Chemistry, *summa cum laude* (2004) Undergraduate advisor: Prof. Xiaowei Zhuang

GPA: 3.9/4.0

AWARDS AND HONORS

- 2018: NIH Director's New Innovator Award
- 2018: NSF CAREER Award
- 2018: Selected as Scialog Fellow (Research Corporation for Science Advancement and the Gordon and Betty Moore Foundation)
- **2017:** Elsa U. Pardee Foundation Award for Cancer Research
- 2016: Air Force Office of Scientific Research (AFOSR) Young Investigator Program Award
- 2012: Awarded NIH Ruth L. Kirschstein NRSA Postdoctoral Fellowship
- 2011: Awarded International Institute for Nanotechnology (IIN) Postdoctoral Fellowship at Northwestern
- 2008: Teaching Effectiveness Award. One of only seven teaching assistants honored.

- 2008: Outstanding Graduate Student Instructor Award (awarded to top 10% of GSI's), UC Berkeley
- 2007: J. Edward Vivian Award for exemplary performance at the MIT Practice School
- 2006-2009: Awarded NIH Applied Bioprocess and Bioengineering Training Grant, UC Berkeley
- 2004: Graduated summa cum laude (top 4% of class) from Harvard University
- 2004: Elected to the Phi Beta Kappa Society, Harvard University
- 2001-2003: Earned prizes each year (Detur Book Prize, Harvard College Prize, John Harvard Prize) for being in top 10% of class, Harvard University

PUBLICATIONS

INDEPENDENT CAREER (* = CORRESPONDING AUTHOR):

- H. Saini[‡], K. Rahmani[‡], A. Buchberger[‡], R. Merkley, Y. Xu, J. Bernal, A. Zare, R. Ros^{*}, M. Nikkhah^{*}, N. Stephanopoulos^{*}, "Reversible control of gelatin hydrogel stiffness using DNA crosslinkers" (manuscript in preparation; [‡]co-first authors)
- C.R. Simmons[‡], T. MacCulloch[‡], F. Zhang, Y. Liu, <u>N. Stephanopoulos*</u>, H. Yan*, "Self-Assembly of a DNA Crystal Scaffold Containing Modular Cavities for the Precise Arrangement of Macromolecules" (*submitted*; [‡]co-first authors)
- F.M. Fumasi, N. Stephanopoulos, J.L. Holloway*, "Reversible Control of Biomaterial Properties for Dynamically Tuning Cell Behavior" *J. Appl. Polym. Sci.* **2020** (early view online: https://onlinelibrary.wiley.com/doi/full/10.1002/app.49058)
- N. Stephanopoulos*, "Hybrid nanostructures from the self-assembly of proteins and DNA" *Chem* **2020**, *6*, 364-405.
- A. Buchberger, C.R. Simmons, N.E. Fahmi, R. Freeman, **N. Stephanopoulos***, "Hierarchical assembly of nucleic acid/coiled-coil peptide nanostructures" *J. Am. Chem. Soc.* **2020**, *142*, 1406-1416. (selected as "ACS Editor's Choice" article)
- N. Stephanopoulos*, "Peptide-DNA hybrid molecules for bioactive nanomaterials" *Bioconjugate Chem.* **2019**, *30*, 1915-1922. (selected as "ACS Editor's Choice" article)
- N. Stephanopoulos*, "Strategies for stabilizing DNA nanostructures to biological conditions" *ChemBioChem* **2019**, *20*, 2191-2197.
- Y. Xu, S. Jiang, C. Simmons, R.P. Narayanan, F. Zhang, A.-M. Aziz, H. Yan, **N. Stephanopoulos***, "Tunable nanoscale cages from self-assembling DNA and protein building blocks" *ACS Nano* **2019**, *13*, 3545–3554.
- A. Stelson, M. Liu, C. Little, C. Long, N. Orloff, <u>N. Stephanopoulos*</u>, J. Booth*, "Label-free detection of conformational changes in switchable DNA nanostructures with microwave microfluidics" *Nat. Commun.* 2019, 10, 1174.
- T. MacCulloch[‡], A. Buchberger[‡], N. Stephanopoulos^{*}, "Emerging applications of peptide-oligonucleotide conjugates: bioactive scaffolds, self-assembling systems, and hybrid nanomaterials" *Org. Biomol. Chem.* **2019**, *17*, 1668-1682. († co-first authors)
- M. Liu, S. Jiang, O. Loza, N.E. Fahmi, P. Šulc, <u>N. Stephanopoulos*</u>, "Rapid photo-actuation of a DNA nanostructure using an internal photocaged trigger strand" *Angew. Chem. Int. Ed.* **2018**, *57*, 9341-9345. (selected as paper for Wiley's Joint Special Collection on Biopolymers, for the Murray Goodman Award Symposium at the 2019 ACS Spring Meeting: bit.ly/wileybiopolymers19)
- N. Stephanopoulos*, R. Freeman*, "DNA-based materials as self-assembling scaffolds for interfacing with cells" (invited book chapter), "Self-Assembling Biomaterials: Molecular Design, Characterization and Application in Biology and Medicine, 1st Edition" 2018, pp. 157-175. (Elsevier)
- L. Avolio, D. Sipes, <u>N. Stephanopoulos</u>, S. Sur*, "Recreating stem-cell niches using self-assembling biomaterials" (invited book chapter), "Self-Assembling Biomaterials: Molecular Design, Characterization and Application in Biology and Medicine, 1st Edition" **2018**, pp. 421-454. (Elsevier)
- C. Simmons, F. Zhang, T. MacCulloch, N.E. Fahmi, <u>N. Stephanopoulos</u>, Y. Liu, N. Seeman, H. Yan*, "Tuning the Cavity Size and Chirality of Self-Assembling 3D DNA Crystals" *J. Am. Chem. Soc.* 2017, 139, 11254-11260.
- D. Varun, G.R. Srinivaan, Y.-H. Tsai, H.-J. Kim, J. Cutts, F. Petty, R. Merkley, <u>N. Stephanopoulos</u>, D. Dolezalova, M. Marsala, D.A. Brafman*, "A Robust Vintronectin-Derived Peptide for the Scalable Long-term

Expansion and Neuronal Differentiation of Human Pluripotent Stem Cell (hPSC)-derived Neural Progenitor Cells (hNPCs)" *Acta Biomater.* **2017**, *48*, 120-130.

POSTDOCTORAL AND GRADUATE RESEARCH (* = CO-FIRST AUTHOR):

- R. Freeman, M. Han, Z. Álvarez, J.A. Lewis, J.R. Wester, <u>N. Stephanopoulos</u>, M.T. McClendon, C. Lynsky, J.M. Godbe, H. Sangji, E. Luijten, S.I. Stupp, "Reversible self-assembly of superstructured networks" *Science* 2018, 362, 808-813.
- J.J. Greene, M.T. McClendon, <u>N. Stephanopoulos</u>, Z. Alvarez, S.I. Stupp, C.-P. Richter, "Electrophysiological Assessment of a Peptide Amphiphile Nanofiber Nerve Graft for Facial Nerve Repair" *J. Tissue Eng. Regen. Med.* 2018, 12, 1389–1401.
- A.J. Matsuoka, Z.A. Sayed, <u>Nicholas Stephanopoulos</u>, E.J. Berns, A.R. Wadhwani, Z.D. Morrissey, D.M. Chadly, S. Kobayashi, A.N. Edelbrock, T. Mashimo, C.A. Miller, T.L. McGuire, S.I. Stupp, J.A. Kessler "Creating a stem cell niche in the inner ear using self-assembling peptide amphiphiles" *PLoS ONE* 2017, 12, e0190150
- R. Freeman*, N. Stephanopoulos*, Z. Álvarez, J.A. Lewis, S. Sur, C.M. Serrano, J. Boekhoven, S.S. Lee, S.I. Stupp, "Instructing cells with programmable DNA-peptide hybrids" Nat. Commun. 2017, 8, 15982.
- C. Rubert-Perez, **N. Stephanopoulos**, S.S. Lee, S. C. Newcomb, Sur, S.I. Stupp, "The Powerful Functions of Peptide-Based Bioactive Matrices for Regenerative Medicine" (invited review) *Ann. Biomed. Eng.* **2015**, *43*, 501-514.
- N. Stephanopoulos, R. Freeman, H.N. Scheler, S. Sur, S. Jeong, F. Tantakitti, J.A. Kessler, S.I. Stupp, "Bioactive DNA-Peptide Nanotubes Enhance the Differentiation of Neural Stem Cells Into Neurons" *Nano Lett.* **2015**, *15*, 603-609.
- A. Li, A. Hokugo, A. Yalom, E.J. Berns, <u>N. Stephanopoulos</u>, M.T. McClendon, L.A. Segovia, I. Spigelman, S.I. Stupp, R. Jarrahy., "A bioengineered peripheral nerve construct using aligned peptide amphiphile nanofibers" *Biomaterials* 2014, 35, 8780-8790.
- J. Sack, **N. Stephanopoulos**, D.C. Austin, M.B. Francis, J.S. Trimmer, "Antibody-guided photoablation of voltage-gated potassium channels" *J. Gen. Physiol.* **2013**, *142*, 315-324.
- N. Stephanopoulos, J.H. Ortony, S.I. Stupp, "Self-Assembly for the Synthesis of Functional Biomaterials" (invited review) *Acta Materialia* (special Diamond Jubilee Issue), **2013**, *61*, 912-930.
- N. Stephanopoulos, M.B. Francis, "Making New Materials from Viral Capsids" (invited book chapter) "Polymer Science: A Comprehensive Reference, 1st Edition" 2012, Vol. 9, pp. 247-266. (Elsevier)
- N. Stephanopoulos, M.B. Francis, "Choosing an Effective Protein Bioconjugation Strategy." (invited review) *Nat. Chem. Biol.* **2011**, *7*, 876-884.
- P.G. Holder, D.T. Finley, <u>N. Stephanopoulos</u>, R. Walton, D.S. Clark, M.B. Francis, "Dramatic Thermal Stability of Virus-Polymer Conjugates in Hydrophobic Solvents" *Langmuir* **2010**, *26*, 17383–17388.
- <u>N. Stephanopoulos</u>, G.J. Tong, S.C. Hsiao, M.B. Francis, "Dual-Surface Modified Virus Capsids for Targeted Delivery of Photodynamic Agents to Cancer Cells" *ACS Nano*, **2010**, *4*, 6014-6020.
- <u>N. Stephanopoulos*</u>, M. Liu*, G.J. Tong, Z. Li, Y. Liu, H. Yan, M.B. Francis, "Immobilization and One-Dimensional Arrangement of Virus Capsids with Nanoscale Precision Using DNA Origami" *Nano Lett.* 2010, 10, 2714-2720.
- R.A. Miller, <u>N. Stephanopoulos</u>, J.M. McFarland, A.S. Rosko, P.L. Geissler, M.B. Francis, "The Impact of Assembly State on the Defect Tolerance of TMV-based Light Harvesting Arrays" *J. Am. Chem. Soc.* 2010, 132, 6068-6074.
- N. Stephanopoulos, Z.M. Carrico, M.B. Francis, "Nanoscale Integration of Sensitizing Chromophores and Porphyrins Using Bacteriophage MS2" *Angew. Chem. Int. Ed.* **2009**, *121*, 9662-9666.
- N. Stephanopoulos, E.O.P. Solis, G. Stephanopoulos, "Nanoscale process systems engineering: Toward molecular factories, synthetic cells, and adaptive devices" (invited perspective) AIChE J. 2005, 51, 1858-1869.

INVITED LECTURES AND SEMINARS

- "Protein-DNA nanotechnology" 1st Symposium on Protein and DNA Nanotechnology, National Autonomous University of Mexico, Mexico City, Mexico; August 2020 (invited; exact date TBA)
- "Supramolecular polymerization of DNA origami nanostructures with peptides, proteins, and small molecules" ACS National Meeting, Philadelphia PA; March 22, 2020
- "Protein-DNA nanotechnology" Institute for Protein Design, Seattle WA; March 12, 2020
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" California Institute of Technology, Pasadena CA; March 9, 2020
- "Rapid photo-actuation of a DNA nanostructure using an internal photocaged trigger strand" APS National Meeting, Denver CO; March 2, 2020
- Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Technische Universität München (Technical University of Munich), Munich, Germany; January 10, 2020
- Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Fyzikální Ústav AV ČR, (FZU; Institute of Physics of the Czech Academy of Sciences), Prague, Czech Republic; January 7, 2020
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Wyss Institute (Harvard University),
 Cambridge MA; December 2, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Memorial Sloan-Kettering Cancer Center, New York NY; November 26, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Johns Hopkins University, Baltimore MD; October 31, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" University of California, San Diego,
 San Diego CA; October 28, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" University of North Carolina at Chapel Hill, Chapel Hill NC; October 22, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Institute for Molecular Engineering,
 Chicago IL; October 18, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" McGill University, Montreal Canada; October 1, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" The Ohio State University, Columbus OH; September 20, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Department of Chemical Engineering, Massachusetts Institute of Technology, Cambridge MA; September 13, 2019
- "Hybrid self-assembled nanomaterials from proteins, peptides, and DNA" Macromolecules Innovation Institute, Virginia Tech, Blacksburg VA; September 11, 2019
- "Hybrid peptide/protein-DNA nanomaterials for medicine and biology" 10th International Nanomedicine Conference, Sydney, Australia; June 24, 2019
- "Hybrid nanomaterials through the self-assembly of coiled-coil peptides and DNA nanostructures" ACS National Meeting, Orlando, FL; April 3, 2019
- "DNA nanoscaffolds for molecular machines, structures, and biomaterials," ASU BME Seminar, Tempe AZ;
 October 12, 2018
- "Light-triggered self-assembly and actuation of DNA nanostructures using photocaged nucleotides," ACS National Meeting, San Francisco CA; April 5, 2017
- "Peptide-DNA Hybrids for Dynamic, Programmable Control of Biomaterials," ASU Molecular, Cellular, and Tissue Bioengineering (MCTB) Symposium, Tempe AZ; April 2, 2016
- "Instructing cells with programmable peptide-DNA extracellular matrices," University of Science and Technology of China (USTC), Hefei, China; December 7, 2015

PRESS COVERAGE AND ARTICLES

- Press release for AFOSR Young Investigator Award:
 - https://biodesign.asu.edu/news/asu-top-recipient-prestigious-air-force-young-investigator-awards
- Northwestern University press release for Nat. Commun. manuscript: https://news.northwestern.edu/stories/2017/iulv/manipulate-cells-treat-diseases/
- ASU press release highlighting work in regenerative medicine: https://biodesign.asu.edu/news/restoring-loss-bio-inspired-materials-boost-regenerative-medicine
- Biodesign Institute article about publication and NSF CAREER Award: https://biodesign.asu.edu/news/useful-pinch-nanoscale-tweezers-are-triggered-light
- School of Molecular Sciences article about NSF CAREER Award: https://asunow.asu.edu/20180719-asu-molecular-sciences-professor-receives-nsf-career-award
- ASU article about NIH New Innovator Award: https://asunow.asu.edu/20181002-two-asu-professors-receive-2018-nih-new-innovator-award-0
- SMS article about research program:
 https://asunow.asu.edu/20190108-asu-assistant-professor-exemplifies-cutting-edge-molecular-science-research
- Biodesign article about tweezer sensor:
 https://biodesign.asu.edu/news/plucky-science-researchers%E2%80%99-nanotweezers-used-detection-conformational-changes
- ASU article about protein-DNA nano-cages: https://asunow.asu.edu/20190319-programmable-legos-dna-and-protein-building-blocks-create-novel-3d-cages
- Podcast with "Common Science" to explain work to non-scientists: https://www.youtube.com/watch?v=zGdhXp4nDFA
- Profile on "Ask A Biologist" website: https://askabiologist.asu.edu/explore/building-nanoscale

CURRENT AND PAST FUNDING

- **DOD-AFOSR (Young Investigator Program) –** "Peptide-DNA Tiles as Building Blocks for the Synthesis of Complex Nanostructures" (PI: N. Stephanopoulos): \$359,541 (12/1/2016 11/30/2019)
- **NSF CAREER –** "Hybrid protein-DNA nanostructures and devices" (PI: N. Stephanopoulos): \$539,785 (7/1/2018 6/30/2023)
- NIH DP2 (New Innovator Award) "Chemical synthesis of G protein-coupled
- receptors using sequential DNA-templated reactions (PI: N. Stephanopoulos): \$2,230,906 (9/30/2018 5/31/2023)
- NIH R21 "DNA Hyaluronic Acid Platform for Spatiotemporally Probing the Role of Adhesion Ligands and Growth Factors" (Co-Pls: J. Holloway, N. Stephanopoulos): \$347,738 (4/1/2019 1/31/2021)
- **NSF Elements** "Models and tools for on-line design And simulations for DNA and RNA Nanotechnology" (PI: P. Sulc; Co-PIs: H. Yan, N. Stephanopoulos): \$436,407 (10/1/2019 9/30/2022)
- DOD-NAVY-ONR (DUPIP instrument grant) "Mass Spectrometry Instrument for Mass Determination of Protein and Nucleic Acid Conjugates" (PI: H. Yan; Co-PIs: N. Stephanopoulos, J. Mills): \$300,000 (8/15/2016 8/14/2017)
- Elsa U. Pardee Foundation "Synthetic Peptide-DNA Antibodies for Targeting Cancer Cells" (PI: N. Stephanopoulos): \$186,374 (1/1/2018 12/31/2018)

POSTDOCTORAL SCHOLARS & RESEARCH SCIENTISTS

- Dr. Minghui Liu (co-advised with Prof. Hao Yan; recently promoted to Research Assistant Professor)
- Dr. Yang Xu
- Dr. Nour Eddine Fahmi (co-advised with Prof. Hao Yan)

GRADUATE STUDENTS MENTORED

- Tara MacCulloch (5th year)
- Alex Buchberger (5th year)
- Raghu Pradeep Narayanan (5th year; co-advised with Prof. Hao Yan)
- Kirstie Swingle (4th year; co-advised with Profs. Neal Woodbury, Alexander Green
- Julio Bernal (2nd year)
- Md Al-Amin (2nd year)
- Jonah Procyk (2nd year; co-advised with Prof. Petr Šulc)
- Skyler Henry (1st year)

MASTERS STUDENTS MENTORED

- Ryan Merkley (graduated 2016)
- Ann-Marie Aziz (graduated 2017)
- Timothy Griswold (graduated 2018)
- Omar Loza (graduated 2018)
- Brittany Torczynski (graduated 2019)
- Tyler Rockwood (graduated 2020)

UNDERGRADUATE STUDENTS MENTORED

- Brittany Torczynski (stayed in lab for Masters)
- Will Tuli
- Matthew Nam
- Omar Loza (stayed in lab for Masters)
- Jonah Procyk (stayed in lab for PhD)
- Ilyssa Farmer
- Tyler Rockwood (stayed in lab for Masters)
- Rifat Adam Akkad
- Grant Severson
- Jessica Guido
- Anna Hostal
- Brooke Lovell

MEMBER OF PHD STUDENT THESIS COMMITTEE

- Abhishek Debnath
- Swarup Dey
- Patrick Gleason
- Lan Zhu
- Soma Chaudhary
- Sanchari Saha
- Pritha Bisarad
- Subhadeep Dutta
- Sthitadhi Maiti
- Towshif Rabbani
- Chang Liu
- Mahasish Shome
- Zina Al-Sahouri
- Roslyn Dermody (School of Life Sciences)
- Zaoqing Yan

SERVICE AND OUTREACH ACTIVITIES

- Have served as reviewer for the following journals: Nature Communications, Journal of the American Chemical Society, Biomacromolecules, Acta Biomaterialia, Nano Research, ACS Applied Materials & Interfaces, Nucleic Acids Research, Trends in Biochemical Sciences, Science Advances, Chem, Accounts of Chemical Research, Angewandte Chemie, International Journal of Molecular Sciences, Chemical Communications, Bioconjugate Chemistry, Scientific Reports, ACS Nano, ChemBioChem, Advanced Materials; served as reviewer for an Elsevier textbook proposal on self-assembled materials and a book chapter for the Karty organic chemistry textbook
- Chaired the "Molecular, Cellular, and Tissue Bioengineering Symposium", a meeting at ASU covering various aspects of bioengineering, with speakers from both within and outside the university. My co-chair (Prof. Julianne Holloway) and I expanded the symposium from one day to two, and had ~100 students, postdocs, and faculty attendees.
- Served on Committee for Undergraduate Education and Awards (since Fall 2016)
- Traveled to China for two weeks (Dec. 2015) to recruit talented students from top universities for the SMS PhD program. Made 20 offers, 5 of which were accepted
- Participated in Association of College and University Educators (ACUE) pilot program (Fall 2016) for more effective teaching techniques; one of two SMS representatives (and only junior faculty) chosen to evaluate the effectiveness of this program for future SMS participation.
- Served on Served on Goldwater Scholarship selection committee (Fall 2018)
- Served on departmental search committees for ASU:
 - Junior faculty candidate for bioengineering (SBHSE); Spring 2018
 - Junior faculty candidate in Chemical Engineering for SEMTE; Spring 2018
 - o Senior faculty candidate for Director for the School of Molecular Sciences; Spring 2020
- Initiated outreach with Kyrene del Pueblo Middle School (Chandler) and taught basic concepts of DNA nanotechnology to 8th grade science class
- Working with Ask A Biologist initiative at ASU (https://askabiologist.asu.edu/) to develop an interactive game for K-12 students on DNA self-assembly, with concomitant teacher training workshop
- Volunteered as judge for the Intel Science and Engineering Fair (ISEF) in Phoenix (May 2016)
- School of Molecular Sciences liaison to Regenerative Medicine Core at ASU (director: Prof. David Brafman)
- Served on grant review panels and forward-looking workshops:
 - NSF FDA panel (July 2018)
 - Ad hoc review for NSF FDA (July 2019)
 - NSF Square Table 2 workshop on the interface between synthetic biology and biomaterials science (Oct. 2019)
 - o ARPA-E workshop (Dec. 2019)