

2025 NETRF Request for Applications Mentored Research Award Application Guidelines

LOI Due: March 17, 2025

Application Due: June 23, 2025



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NETRF Summary

The Neuroendocrine Tumor Research Foundation (NETRF) is the largest private global funder of basic, translational and clinical neuroendocrine cancer research. Our mission is to accelerate scientific discovery that will help create new and more effective therapies for neuroendocrine cancers including gastrointestinal, pancreatic, lung, and adrenal-associated neuroendocrine tumors. Since its founding, NETRF has awarded more than \$38 million in research grants to investigators whose work can help provide insight into the causes of neuroendocrine cancer and/or lead to improved treatments for our patients. While we have made significant progress the past few years, we still do not have a complete understanding of the unique characteristics of these tumors.

NETRF seeks investigators and teams of investigators from around the world to study neuroendocrine cancers in new ways. NETRF supports transformative basic, translational, and clinical research studies that address critical questions to advance our understanding of tumor biology, address roadblocks to therapeutic development, and/or exploit innovative technologies or strategies.

NETRF Mentored Research Award Program

The NETRF Mentored Award is designed to encourage early career investigators to pursue neuroendocrine tumor research and make a commitment to the field. Mentored Research Awards are in the amount of \$100,000 over two years. These proposals typically have two aims that provide work requiring two years to complete. They have one PI with or without collaborators.

Application Timeline and Review

The NETRF request for applications employs a two-step process. The first step is the submission of a competitive letter of intent (LOI). One LOI per applicant, per program, may be submitted. If selected for the second stage, a full application is required.

- An LOI must be submitted and all eligibility criteria met to be considered for the full application stage; otherwise, LOIs will be administratively rejected.
- If selected, a full application must be submitted that meets all guideline criteria;
 otherwise, applications will be administratively rejected.
- LOIs and full applications will be peer reviewed by an independent panel of scientific experts.

APPLICATION TIMELINE

LETTER OF INTENT DUE	March 17, 2025 by 5PM ET
INVITATION TO SUBMIT FULL PROPOSAL	Early May 2025
FULL PROPOSALS DUE	June 23, 2025 by 5PM ET
AWARD NOTIFICATION	Oct/Nov 2025
PROJECTED START DATE	January 2026



Applicant Eligibility

- Applicants must have an MD, PhD, MD/PhD or equivalent degree and not currently be a candidate for a further doctoral or professional degree.
- Applicants must hold a full-time, mentored research position with the title of postdoctoral fellow, clinical research fellow, or equivalent. They must work under the auspices of a mentor at an academic, medical, or research institution within the US or abroad and should be focused on establishing themselves as experts in neuroendocrine tumor research.
- Eligible organizations include public or private institutions, such as universities, colleges, hospitals, and laboratories, both domestic and international. Applications from the biotech, pharmaceutical industry, or for-profit life sciences companies are **not** eligible.
- The award is transferable to the first year of a junior faculty position.

Grant Budget and Restrictions

The requested budget should be in proportion to the scope of the proposed project and should be at or under \$100,000 USD over two years. Awarded funds are directed to the institution.

- Mentored Research Awards do not cover indirect costs.
- NETRF adheres to the NIH salary cap for principal investigator(s) and personnel.
- Funds may be used for personnel salaries, supplies, small equipment, and/or research-related services only.
- Funds must not be used entirely for salaries and may not be applied to costs covered by other
- Funds should be allocated to attend the Annual NETRF Research Symposium held in Boston, MA,
 USA.

Scope

Tumor types of greatest interest

- GEP-NETs: Gastroenteropancreatic neuroendocrine tumors, including sites of metastasis.
- Lung NETs: Well-differentiated, low-grade typical carcinoids (TCs), well-differentiated, intermediate-grade atypical carcinoids (ACs), diffuse idiopathic pulmonary neuroendocrine cell hyperplasia (DIPNECH).

Tumor types in scope

- Adrenal-associated NETs: Pheochromocytoma and paraganglioma.
- **Neuroendocrine Carcinomas (NECs):** Extrapulmonary, high-grade neuroendocrine carcinomas.
- The Foundation may consider supporting work on other NET types based on allotted funding availability as long as it is not specifically listed as out of scope.

Tumor/tissue types out of scope

- Small-cell lung cancer (SCLC)
- Large-cell lung cancer (LCLC)
- Non-SCLC
- Squamous cell carcinomas
- Adenocarcinomas



- Mesotheliomas
- Poorly differentiated non–small cell lung carcinomas
- Neuroendocrine prostate cancer
- The Foundation does not support research using human embryonic or fetal tissue.

Research Areas of Interest

In 2024, NETRF adopted a Research Roadmap that prioritizes Early Detection, New Therapeutic Development, and Precision Medicine to drive progress in neuroendocrine cancer research. We seek applications focused on:

- Developing biomarkers, AI tools, and imaging technologies for early diagnosis
- Innovating new and improved treatments
- Leveraging molecular insights for personalized therapies

In addition to these areas, NETRF remains committed to exploring a broad range of promising research opportunities. We welcome proposals beyond these categories, including but not limited to:

- Uncovering the molecular and genetic basis of NETs: Elucidation of processes underlying NET initiation, progression, monitoring of NET stages. Understanding the mechanistic rules for treatments to overcome therapeutic resistance and toxicity.
- New/optimized experimental models: Cell lines, mouse models, optimized spheroids/organoids, patient-derived xenografts, and others. Strategies to modify model proliferation and improve experimental utility.
- Application of existing or new technologies to target NETs: Designer or nano-based drugs to create targeted therapies/treatments, drug/biologic-targeted delivery to tumors, new tumor- specific targets, oncolytic viruses/cancer vaccines, nanoparticles that promote efficient drug delivery, AI-based strategies and novel therapeutic development.
- Cancer metabolism: Understand and manipulate the way neuroendocrine cancer cells process energy in order to survive.
- Cell invasion and metastasis: Drivers of metastasis, anti-metastasis targets, small non-coding RNAs, adhesion molecules and epithelial-to-mesenchymal transition, metastasis-initiating or cancer stem cells, metastatic organotropism, triggers that stimulate invasive behavior in an indolent tumor.
- Tumor microenvironment: Understanding a tumor's interaction with its environment, immune effector functions and the metabolic landscape of the tumor microenvironment, contributing angiogenic processes, effect of microenvironment on progression, metastasis, and treatment response.
- Immunotherapy: Immunologically "cold" versus "hot" NETs, novel types of CAR T-cell therapy for solid tumors, adaptation or creation of methodologies to reprogram and activate the immune system, immune-engineering, antibody-drug conjugates.
- Diagnostics/Biomarkers: Predictive/prognostic clinical markers, biomarkers for metastasis, imaging biomarkers, new approaches to the early diagnosis of NETs, circulating tumor cells/cell free DNA/exosomes.
- Big data: Development of novel methods and use of large-scale data collections (e.g., "omics" data) to interrogate NET genetics and molecular underpinnings.



- Clinical research: Exceptional clinical and correlative studies that may build upon existing clinical trials, proof-of-concept Phase I clinical trials, innovative combination therapies, adaptive and sequencing studies.
- Nuclear medicine, Theranostics, Imaging: Non-incremental, innovative studies of novel imaging agents or therapies, new delivery/targeting systems, computational strategies.

Resource and Data Sharing

Grant recipients who create unique research resources, including but not limited to model organisms, cell lines, plasmids, protocols, software, and data using NETRF funds, are required to share such resources within the scientific community. NETRF expects that, where available, resources will be deposited and archived in standard repositories (e.g., Addgene for plasmids). Resources should be shared openly with the research community no later than the date of publication or within twelve months after the end of the grant funding, whichever comes first.

NETRF is committed to sharing research information to ensure research transparency and enable unrestricted access to research results. Recipients must submit all publications, excluding non-research articles such as review articles, that were in part or fully funded by NETRF as a preprint to bioRxiv, medRxiv, or a similar preprint sharing service prior to or at the time of initial journal submission.

Applicants must provide a resource sharing plan in the full application. To demonstrate a commitment to sharing that will be actualized, applicants should provide information in their sharing plan that clearly states the type of resource that will be shared, the method, characterization, and timing of such sharing, and the anticipated resources (budget, personnel, etc.) required by the applicant and the resource user. Reviewers will consider the extent to which the dissemination of resources produced under the award will enhance or diminish the impact of the proposed work.

Grant Reporting and Other Requirements

- Collaborative efforts are encouraged; however, a single principal investigator and institution must be selected to receive an award.
- Progress and financial reports are required at 6-month intervals throughout the duration of the
 grant. The progress reports track milestones, research progress, and use of the funds. Future
 funding is contingent upon review and approval of progress and will be paid in six-month
 installments in US dollars. A final report is required at the conclusion of the project detailing study
 findings and project expenditures.
- Post-award outcomes of the funded research are required to be reported at one, three and five years following the completion of funding.
- It is mandatory for Awardees to attend and present at the annual NETRF Research Symposium whether in person, or virtual, for the duration of their grant and upon completion. NETRF grant funds may be used to pay for NETRF conferences travel, if held in person.



Letter of Intent (LOI) Instructions

The purpose of the LOI is to ensure that the proposed research is within the scope of the Mentored Research Award and merits a full application. LOIs should outline research with the potential to transform our understanding of NETs and/or lead to improved treatments for patients. LOIs may focus on basic, translational, or clinical research. All LOIs are peer-reviewed by NETRF's Scientific Advisory Board and other leaders in the field, and a limited number of applicants whose LOIs are deemed most meritorious will be invited to submit a full proposal.

LOI Section Descriptions

Letters of intent must be submitted through the <u>Proposal Central</u> platform. Once you create or log into your account, click on the "Grant Opportunities" tab and search for the Neuroendocrine Tumor Research Foundation. Click on the "Apply Now" button to begin.

The LOI content will be entered directly into Proposal Central. You will be asked for the following information:

- 1. Title Page
- 2. Enable Other Users to Access This Proposal Optional. Enter an email address for an individual if you would like to grant them access to your LOI.
- 3. Applicant/PI Information
- 4. Organization/Institution Information
- 5. Mentor Information
- 6. Abstract & Keywords
 - a. Scientific/Clinical Impact Statement (1,500 characters max)
 - b. Scientific Abstract (3,000 characters max)
 - c. Lay Summary (2,000 characters max)

The LOI Submission Deadline is Monday, March 17, 2025, at 5PM ET.



Full Application Instructions

Full applications are by invitation only. Full applications are due by **Monday**, **June 23**, **2025**, **by 5PM ET**. All of the information submitted for the LOI will be copied to the full application and available for editing. To access the full application, log in to <u>Proposal Central</u>, click on your proposals tab to find the full application, and click edit to begin.

The full application content will be entered directly into Proposal Central. You will be asked for the following information:

- 1. Title Page
- 2. **Download Templates and Instructions** download instructions and templates for required uploads. Use the Research Proposal Template to create your proposal.
- 3. **Enable Other Users to Access this Proposal** optional. Enter an email address for an individual if you would like to grant them access to your proposal.
- 4. Applicant/PI Information
- 5. PI Data Sheet
- 6. Organization/Institution
- 7. Key Personnel
- 8. Milestones and LOI Recommendations
- 9. Abstract & Keywords
 - a. Scientific/Clinical Impact Statement (1,500 characters max)
 - b. Scientific Abstract (3,000 characters max)
 - c. Lay Summary (2,000 characters max)
- 10. Budget Period Detail
- 11. Budget Summary
- 12. Organization Assurances human subjects and vertebrate animal approvals
- 13. Upload Attachments these items will be uploaded directly to Proposal Central
 - a. **Research Proposal**. *Download and use the Research Proposal Template in "Download Templates and Instructions" for this.* The proposal sections include:
 - i. Title
 - ii. Proposal Narrative (limit six pages)
 - Introductory Statement include objectives as they relate to the NETRF mission
 - 2. Background and Preliminary Data
 - 3. Rationale and Hypothesis
 - 4. Specific Aims
 - 5. Research Design and Methods
 - 6. Potential Problems and Contingencies
 - iii. Significance (limit one page)
 - Significance: Describe the key question or therapeutic roadblock being addressed by your proposal and why it is important to our understanding of NETs and/or its treatment.



- 2. Statement of Innovation: describe how the proposed study employs a new idea and/or innovative approach.
- 3. Potential for translational application and patient benefit.
- iv. Facilities (limit two pages)
- v. Plan for Sharing Research Data and Resources
- vi. Plan for Access to Patient Tumor Samples, if applicable
- vii. **References Cited** use Vancouver or NIH style (numbered citations within text) format
- b. PI Biosketch(es) limited to five pages, including references
- c. Collaboration Letter(s) and Biosketches if applicable
- d. Mentor Letter of Support the letter must be written by the applicant's mentor on behalf of the applicant. The letter should confirm that the applicant holds a full-time, mentored research position with the title of postdoctoral fellow, clinical research fellow, or the equivalent as well as the mentored environment and laboratory support necessary to complete the project. The letter must critically address the scientific merit and novelty of the proposed research, the scientific expertise and independence demonstrated by the applicant in previous work, and the dedication of the applicant to NET research.
- e. Letter of Institutional Commitment the letter(s) must be written by the department head, dean, or other senior member of the institution on behalf of the applicant, on institution letterhead. The letter(s) should confirm that the institution has the infrastructure required to support the project. The letter(s) must critically address the scientific merit and novelty of the proposed research, the requisite scientific expertise demonstrated by the applicant in previous work, and the dedication of the applicant to NET research.
- f. **Other Support** list other sources of support and amounts, including funds that may be contributed by the Sponsoring Institution.
- 14. **Signature** the PI and an institutional signing official must sign before the application can be submitted.

Formatting Instructions

Applicants must adhere to the following instructions for the research proposal:

- Must use 12-point Times New Roman for the text, and no smaller than 9-point type for figures, legends, and tables.
- Single-spacing is acceptable, and space between paragraphs is recommended.
- The page margins must be no less than 0.75 inches on each side.
- The Research Proposal must be numbered consecutively.

Contact

If you have any questions regarding this grant mechanism, contact Anna Greene, PhD, Director of Research at grants@netrf.org.