

2026 Application Guidelines for the Crazy 8 Initiative Grant Program on Pediatric Brain Tumors

Letter of Intent Due: November 5th, 2025

Full Application Due: April 28th, 2026

Finalist Virtual Presentations: September, 2026



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About Alex's Lemonade Stand Foundation (ALSF)

Alex's Lemonade Stand Foundation (ALSF) emerged from the front yard lemonade stand of 4-year-old Alexandra "Alex" Scott, who was fighting cancer and wanted to raise money to find cures for all children with cancer. Her spirit and determination inspired others to support her cause, and when she passed away at the age of 8, she had raised \$1 million. Since then, the Foundation bearing her name has evolved into a national fundraising movement. Today, ALSF is one of the leading funders of pediatric cancer research in the U.S. and Canada, funding more than 1,500 research projects and providing programs to families affected by childhood cancer. ALSF is also the only childhood cancer research organization that has been given the NCI peer-reviewed funder designation for rigorous selection of research grants. The mission of ALSF is to change the lives of children with cancer through funding impactful research, raising awareness, supporting families, and empowering everyone to help cure childhood cancer.

The Crazy 8 Initiative Award Program

Childhood cancer is the leading cause of death in children in the United States and Canada and the Crazy 8 Initiative's aim is to change the trajectory of pediatric oncology. With this goal, the Crazy 8 Initiative kicked off in the Fall of 2018 with a meeting that brought together more than 90 scientists from around the world to help define the research landscape in eight key areas of need to tackle major obstacles impeding progress toward cures for childhood cancer. Since this meeting, ALSF has awarded 8 research teams across 28 institutions, a total of \$35 million, funding innovative research that addresses major obstacles in finding cures for Pediatric Cancer. To see the scope of projects funded in the previous round of Crazy 8 Award funding, navigate here.

In 2025, the ALSF Crazy 8 Initiative will focus on funding studies that uncover more efficacious and less toxic treatments for pediatric brain tumors. Brain and spinal cord malignancies are the second most common type of pediatric cancer, with more than 4,000 children and young adults diagnosed each year. These tumors are often very difficult to treat, as the 5-year survival rate for certain types of brain cancers is below 20%, with existing treatment options often having significant side effects. Therefore, there is a critical need to develop novel, minimally toxic therapies as alternative treatment options to current standards of care. The focus of this award will be on the development of unprecedented treatment options for brain and spinal cord tumors. Examples of potential areas of focus that are within the scope of the Crazy 8 Brain Tumor Award are as follows:

- Novel targeted therapy discovery and development.
- The uncovering of cutting-edge technologies that enhance drug penetration of the bloodbrain barrier.
- Creating and utilizing novel preclinical models that inform clinical studies.



• Identifying and incorporating methods to reduce the toxicity of novel and existing therapies.

Other areas of focus will be considered. If you have questions about if your project fits with the Crazy Brain Tumor grant goals, reach out to Margaret Poore (M.Poore@AlexsLemonade.org) Please review the Restrictions section for proposals considered out of scope for this award.

Program Description

This ALSF Crazy 8 RFA focuses on developing and enhancing pediatric brain tumor treatments. The focus must be on pediatric/adolescent brain tumors and should be a consortium of two or more institutions with different areas of research collaborating to integrate their expertise. It is strongly encouraged to have a patient advocate on the team.

The proposals will be judged on innovation, scientific soundness, significance, and the potential impact on improving the lives of children with cancer. The award selection process involves three stages: (1) a **letter of intent** to narrow the pool to the projects most in line with the scope of the Crazy 8 Initiative; (2) a **full proposal** to select finalists; and (3) a **virtual presentation** to ALSF leadership and the Crazy 8 Scientific Advisory Board to choose the award recipient(s).

Application Timeline and Review

- A letter of intent (LOI) must be submitted, and all eligibility criteria met to be considered for the full proposal stage.
- Full proposals must meet all guideline criteria and all eligibility criteria; otherwise, applications will be administratively rejected.
- LOIs and full proposals will be reviewed by an independent panel of experts according to the NIH recognized peer-review process overseen by the Crazy 8 Initiative Scientific Review Board.
- We will announce the dates for Finalist Presentations as soon as they are confirmed. If invited, teams are expected to be available at this time.

Timeline	
Letter of Intent Due	November 5, 2025
Invitation to Submit Full Proposal	January 2026
Full Proposals Due	April 28, 2026
Invitation to Present	July 2026
Finalist Presentations (Virtual)	September 2026
Award Notification	November 2026



Eligibility

- Pls, Co-Pls, and Co-Is must have an MD, PhD, or MD/PhD or equivalent and be appointed as faculty (or equivalent) at an academic institution.
- PIs, Co-PIs, and Co-Is must have a track record of publication and funding productivity that demonstrates that the project can be accomplished by the investigators.
- PIs, Co-PIs, and Co-Is' institutions may be based in the United States or abroad, and applicants need <u>not</u> be United States citizens. Funds must be granted to nonprofit institutions or organizations and will be distributed in US dollars.
- Pls, Co-Pls may not have been Pls or Co-Pls on a previously funded Crazy 8 Project.
- Co-Is may not be on a currently funded Crazy 8 Project at the time of submission.
 However, if a Co-I with special expertise is needed that is already on a team, please send
 your request with a brief description of why this member is critical for the team to
 Margaret Poore before the submission deadline at M.Poore@AlexsLemonade.org and we
 will review it administratively.

Budget

The requested budget should be in proportion to the scope of the proposed project and its corresponding potential for impact. Depending on the type of project and the size of the team, the requested budget should be in the range of \$3-5 million USD over four years. Cost efficiency will be one of the major considerations in review and funding decisions. Funding through subsequent years of the grant period is contingent upon satisfactory progress as determined by a Crazy 8 Initiative Scientific Review Board.

ALSF will pay up to 10% in indirect costs; if the applicant elects to request indirect costs, the total budget including indirect costs may not exceed the range of \$3-5 million USD. The 10% indirect costs should be assessed on the Total Direct Costs. Indirect costs may be assessed on subawards' Total Direct Costs but must be excluded from the primary institution's Total Direct Costs for calculating indirect.

Restrictions:

- o ALSF adheres to the NIH salary cap for principal investigator(s).
- The grant may not be renewed; one no-cost extension request is allowed with the Final Report.
- Other budget items may include fringe, supplies, and small pieces of equipment. ALSF funds cannot be used for tuition remission. View ALSF's complete budget policy.
- Travel to the annual Crazy 8 Summit in Philadelphia must be included in the budget.
- If utilizing a subcontract or subaward, you must include this expense in the budget.
 The PI's institution is responsible for disbursing funds for subawards and/or subcontracts.



Resource and Data Sharing

Applicants should describe their track record of generating resources that are broadly re-used, the specific resources that will be generated in this proposed project, and the mechanisms by which those resources will be shared. 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. For a full description and an example, see the Resource Sharing Form. For the full Resource Sharing Policy visit our Resources for Grant Applicants page.

Resubmissions

One resubmission of an application previously reviewed by ALSF's Crazy 8 Review Board is permitted. Applicants should respond to the prior ALSF critique of the proposal in the Resubmission section of the application. The response to critique will be scored in the review process, and resubmissions compete with new applicants for funding.

Restrictions

- Proposals must fall within the scope of Alex's Lemonade Stand Foundation's mission, focused on childhood brain tumors. Proposals with a sole/primary focus on patients >19 years of age will not be considered.
- Proposals solely focused on social determinants of health or psychosocial outcomes independent of oncologic outcomes are not eligible for the Crazy 8 Brain Tumor Award.
- Projects solely focused on epidemiological aspects of pediatric cancer and/or cancer control are not eligible for the Crazy 8 Brain Tumor Award.
- ALSF does not fund proposals for research utilizing human embryonic stem cells or non-human primates. Research with human induced pluripotent stem cells is permissible.
- Investigators may submit one LOI as lead PI or Co-PI.

Grant Reporting Requirements

- Please review ALSF's Grant Policies, including Budget Expenditures, Resource Sharing and Grant Agreement with IP, found on the <u>Resources for Grant Applicants</u> page
- Annual progress reports followed by an update teleconference with ALSF.
- Attendance at a yearly Crazy 8 Summit, to be held in person.
- Each year of funding is contingent upon demonstration of satisfactory progress toward the completion of proposed research objectives and appropriate budget expenditures.



- Minor carry-over of funds (25% or less) is permitted each year with justification. Each year's budget will be approved subsequent to the review of project progress. If carry-over is excessive, ALSF may elect to partially fund or hold new funds.
- A final report is required at the conclusion of funding. Report must state findings, expenditures, as well as publications and presentations which acknowledge ALSF funding. The grant may not be renewed; one no-cost extension may be requested in the final report.
- Publications, presentations, and posters featuring results of the experiments funded by this grant mechanism must acknowledge "Alex's Lemonade Stand Foundation (Grant #)".
 Copies should be sent via email to Grants@AlexsLemonade.org.

Letter of Intent (LOI) Instructions (Due November 2025)

A Letter of Intent is required to ensure that the proposed research is within the scope of the Crazy 8 Initiative. LOIs will be reviewed by a review board prior to advancing to the full proposal stage.

Format Instructions

- PAGE HEADER: All pages of the LOI should be numbered; the name of the principal investigator should appear in the upper right-hand corner of each page.
- FORMAT: Follow NIH format guidelines: Arial, Helvetica, Palatino Linotype, or Georgia fonts with a font size of 11 points or larger with a minimum of ½ inch margins.
- ORDER & LENGTH: The order of the LOI should be followed, adhering to the maximum number of pages allowed for each subsection indicated in parentheses, not to exceed 2 pages.
- No appendices allowed.
- Create a **single PDF** for all sections and bio sketch(es) and upload to the ALSF online application form (see Application Submission Instructions).

LOI Section Descriptions

- **Scientific Abstract** (500 words) Address the big problem and Crazy 8 focus areas that you will tackle. Must also include: Rationale, Hypothesis, Specific Aims, Design.
- Investigators and Institutions List all key personnel, disciplines, their affiliated institutions, and any unique attributes of your team. It is understood that this may change, and the list is not considered binding.
- **Budget** (2-4 sentences) List the total requested budget and short justification (PI and Co-I support, supplies and contracts).



- Impact Statement (~4 sentences) State how this project will impact children with brain tumors.
- Innovation Statement (~4 sentences) State how this project is a novel approach to childhood cancer research and how specifically the innovation will help you address the big problem you are tackling.
- Resource Sharing (~4 sentences) Describe how outputs from the project will be shared. State how this project's impact will be enhanced by sharing outputs.
- **Literature References** (1 page) Use Vancouver style: numeric references within the text.
- Biosketch(es) (not included in page count) Use the NIH 5-page bio sketch form for PI and Co-PIs.

Full Proposal Instructions (By invitation only with an approved LOI)

Application Package

- All sections described below should be combined into one PDF (max 20 MB) and uploaded to the ALSF online application form (see Application Submission Instructions).
- All templates mentioned can be found on ALSF's Resources for Grant Applicants page.

Format Instructions

- PAGE HEADER: All pages of the application should be numbered; the name of the principal investigator should appear in the upper right-hand corner of each page.
- FORMAT: Follow NIH format guidelines: Arial, Helvetica, Palatino Linotype, or Georgia fonts with a font size of 11 points or larger with a minimum of ½ inch margins.
- ORDER & LENGTH: The order of the application should be followed, adhering to the maximum number of pages allowed for each subsection indicated in parentheses.

Section Descriptions

1. Project Information

- a. Cover Page (1 page): Download and complete the Cover Page Template.
- b. **Table of Contents** (1 page): Provide a Table of Contents with page numbers to the corresponding sections.
- c. **Scientific Abstract** (0.5 page): Summarize the research objectives and rationale.
- d. **Impact Statement** (0.5 page): How will this project impact children with brain tumors? Describe the timeline for translation to the clinic.
- 2. Budget/Justification (3 pages):



- a. **Budget Template** (1.5 pages): Complete the <u>ALSF Crazy 8 budget template</u>. The signature from an institutional representative on the cover page of this grant application specifically acknowledges and accepts this budget.
 - i. The award amount is between \$3-5 million over 4 years.
 - ii. ALSF adheres to the NIH salary cap for principal investigator(s).
 - iii. Indirect costs are allowed up to 10% and should be assessed on the Total Direct Costs.
 - iv. If utilizing a subaward, you must include their budget. Indirect costs may be assessed on subawards' Total Direct Costs but must be excluded from the primary institution's Total Direct Costs for calculating indirect. The contact PI's institution is responsible for disbursing funds for subawards/subcontracts.
 - v. Reasonable travel expenses to national/international research meetings to disseminate findings may be budgeted. Travel to the ALSF annual Crazy 8 Summit will be incorporated into the yearly budget
- b. **Budget Justification** (1.5 pages): Include a narrative for the following. Use N/A in sections as needed:
 - i. Personnel
 - ii. Subcontractors/Subawards/Consultants
 - iii. Equipment (for equipment costs above \$5000)
 - iv. Travel
 - v. Computer and Software
 - vi. Other
- **3. Biographical Sketch(es):** Use the NIH five-page biographical sketch (SF424) format for the principal investigator and all key personnel.

4. Research Plan

- a. **Specific Aims** (1 page): List the goals, long-term objectives and what the specific research proposed in this application is intended to accomplish. State the hypothesis to be tested and relevance to childhood cancer research.
- b. Significance (1 page): Describe the relevant background for the current research plan. State the significance and importance of your proposed project with respect to childhood cancer research. Relate the specific aims to the goals and long-term objectives.
- c. **Innovation** (0.5-1 page): Describe how the proposed research challenges and shifts paradigms or introduces a novel concept, approach, or technology.



- d. **Approach** (10 pages): Describe the experimental approach to the research question and how the research will be realistically accomplished within the proposed funding period. This section should include *but is not limited to:*
 - i. Feasibility of the approach to reach project goals; include preliminary studies pertinent to the project.
 - ii. Anticipated potential problems and plans to address these issues.
 - iii. Timeline and deliverables
- **5. Resource Sharing (1-2 pages)**: Use the <u>Resource Sharing Form</u> to complete this section of the application to describe the outputs from the proposed project and how they will be shared. Reviewers will be asked to consider the manner in which outputs from this project will be shared and the extent to which this plan will increase or decrease the impact of the proposed project.
- **6.** Literature Cited: Use Vancouver or NIH style (numbered citations within text) format.
- **7. Human Subjects** (1 page): If approved, include the IRB approval letter or equivalent. If approval is pending, indicate the expected approval date. If IRB approval is not applicable, include a note in this section.
- **8. Vertebrate Animals** (1 page): If approved, include the IACUC approval letter or equivalent. If approval is pending indicate the expected approval date. If IACUC approval is not applicable, include a note in this section.
- **9. Letters of Support**: Include any appropriate letters from individuals confirming their roles in the project. Institutional letters of support are not required, but can be included, especially if there are issues of feasibility that can be addressed.
- **10. Appendix:** A brief appendix is allowed for Crazy 8 Initiative application. Appendices should be included only if essential to the understanding of the application. Appendices are limited to one accepted but not yet published manuscript OR two pages of additional information such as a summary of the protocol and supplementary figures. (Reminder: tables and figures noted in the text should be embedded in the body of the proposal.) **Reviewers are not required to read the appendix.**

Application Submission Instructions

- 1. To start an application, navigate to <u>Proposal Central</u> and select I am an: Applicant or Awardee
 - a. <u>ORCID Registrants:</u> you can login using your ORCID. If you don't have one, please obtain one by registering through the link https://orcid.org/register. An ORCID is required for this application.
- 2. After you've logged in to Proposal Central, navigate to the Grant Opportunities tab and you can search Alex's Lemonade Stand Foundation in the search bar to find our active



- grant application cycles, or click <u>here</u> to see the ALSF's Grant Opportunities currently open. Click the "Apply Now" button to start an application.
- 3. The application document must be uploaded as **one PDF** (maximum of 20 MB), in the Attachments tab. Please see the guidelines for specific format and section instructions.
- 4. You may save your application to finish later. Just click "Save".
 - a. When you return to Proposal Central, click the "Proposals" tab at the top to navigate to your applications. You can filter your application based on Proposal Status in the upper right-hand corner. Click "In Progress" to return to your existing applications.
- 5. Once completed, click "Submit". Within 10 minutes you will receive a confirmation email. As long as the deadline has not passed, you may unsubmit your application to make changes by clicking the "Unsubmit" button on the "Proposals" tab.

If you have any questions regarding your Proposal Central account, please contact pcsupport@altum.com.

Contact

• If you have any questions regarding this grant mechanism, reach out to Margaret Poore, Grant Coordinator by email at M.Poore@AlexsLemonade.org.