REQUEST FOR APPLICATIONS:

Fight Osteosarcoma Together Super Grant

Research for Pediatric Osteosarcoma

Purpose: This Request For Applications (RFA) seeks to support high-impact osteosarcoma research with a focus on moving quickly toward improvements in outcomes for patients. Projects can be investigator-initiated, intervention-oriented studies or translational research to create new therapeutic options. Applicants are required to propose time-dependent, measurable milestones to ensure completion of study objectives and clearly state how results at each milestone will be used in measuring research outcomes. The project budget should closely reflect the milestones. The funder will be updated through written outcomes reports every year, a phone call update every six months, and a site visit or videoconference at the end of each year thereafter. Applications may be from individuals or teams, with a preference for team science.

This RFA is to be administered by the St. Baldrick's Foundation, through the generosity and fundraising of the following partner organizations.

- Battle Osteosarcoma
- Children's Cancer Research Fund
- CureSearch for Children's Cancer
- Michael and April Egge
- The Osteosarcoma Collaborative

Requirements and Considerations:

- The focus of applications should be on creating new therapeutic options for patients as quickly as possible, aiming for a clinical trial in the next 1 to 3 years. Applications may focus on first line treatment or relapsed/metastatic disease.
- Essentially all therapeutic modalities are accepted, with a preference for molecular informed therapies, immunotherapies and combination, multi-modal therapies.
- The focus of the funded work will be treatment efficacy, but the inclusion of correlative studies to identify biomarkers for response will be well received.
- Methods that allow dynamic monitoring of disease burden and/or therapeutic efficacy will also be considered. Note, however, that the discovery of novel therapeutic interventions will be given priority.
- Both hypothesis-driven and hypothesis-neutral (e.g., drug screening and genome-wide molecular characterization) approaches will be considered, as long as the outcome of the project leads to actionable therapeutic intervention within the suggested time frame.
- Proposals without clear path to the therapeutic interventions will receive a low priority.
- Cross discipline collaboration is encouraged, with each investigator's role and responsibility clearly outlined in the proposal.
- Proposals must include proposed timeline with anticipated milestones for each quarter.
- Proposals must include a clear plan for the sharing of data.
- Grant subject to annual reporting to grant scientific review team and [Name TBD].

Award Amount:

The award amount will total $500,000/year, distributed annually, pro rata, over a three-year period.

Funding starting from the 2nd installment will be evaluated based on research progress. Continuation of funding will be dependent upon successful completion of each milestone and is contingent upon favorable review of the grantee's progress reports by the Scientific Advisory Council

Key Dates:

Letter of Intent (LOI), application and required documents must be submitted by the Principal Investigator, in English, online through ProposalCentral before 5 p.m. EASTERN TIME on the deadline.

Only applicants with LOIs deemed most meritorious and aligned with program's goals will be invited to submit full applications.

- Letter of intent due: December 9, 2022 before 5pm EASTERN TIME
- Letter of intent approval by: January 20, 2023
- Full proposal due: March 3, 2023 before 5pm EASTERN TIME

- Notification of award by: June 1, 2023
- Project start date: July 1, 2023

Reporting:

- Researchers funded will have a scheduled phone meeting update with the funding agency every 6 months.
- The funding agencies' representatives will conduct a site visit or videoconference with recipients every 12 months to monitor progress and address issues. A 45-min technical presentation followed by Q&A will be expected during the visit.
- A written report must be submitted online via ProposalCentral every 12 months.
- Annual report of expenditures must be submitted online via ProposalCentral within 60 days of the end of each grant year.
- After the grant term is complete, periodic (every 1-5 years) survey responses are required to gauge the long-term effects of investment and research outcomes.

Format of the Proposal:

Letter of Intent (LOI), application and required documents must be submitted by the Principal Investigator, in English, online through ProposalCentral (https://proposalcentral.com/) before 5 p.m. EASTERN TIME on the deadline.

Please review this helpful document for instructions on creating an LOI/Application using ProposalCentral: https://docs.proposalcentral.com/CreateApp.pdf

Specific LOI Instructions/Requirements are available in ProposalCentral upon starting the LOI.

All application instructions and templates/requirements will be available in ProposalCentral upon approval of an LOI.

- Executive summary including one graphical abstract
- Rationale
- Existing Data
- Research plan including milestones
- Potential for clinical application and commercialization
- Scientific Rigor and Transparency
- Institutional Letter of Support
- Resource Sharing Plan
- Budget Justification

Reviewing Process and Criteria:

- Invited proposals will first be reviewed by a minimum of 3 reviewers based on expertise and scored based on:
    - Potential therapeutic effect (curing being the highest; no therapeutic benefit being the lowest)
    - Scientific rationale
    - Probability of success (based on previous data, expertise of the investigator, and the environment).
    - Expected timeline to clinical benefit
- The top-rated applications from the scientific review will then be reviewed by the CureSearch Industry Advisory Council for clinical, commercial, and regulatory feasibility and to assess clinical translation and drug development potential.
- A final review discussion will take place with the scientific review scores and industry review feedback. Lay advocates will participate, after which the scientific review panel will make the final funding decision based on scientific score.
- The awardee and top rated non-funded applicants will be invited to present a lay description of their work to funder representatives and answer questions.

Eligibility:

- Applications will be accepted from both individual researchers/labs and from consortia, with team science preferred.
- Institutions must be located in the United States.
- Teams with existing international collaborations are eligible to apply, however clinical trial development must be available to U.S. patients.
- Applicants need not be American citizens; however, they must work at an academic, medical, or research institution within the United States.
- An institution may submit more than one LOI/Application for this RFA.
- A researcher can only apply once as the lead Principal Investigator.
- Applicants should hold at least a M.D., D.O., Ph.D., DVM, or international equivalent degree by the date the award becomes effective
- Laboratory scientists and clinical investigators must have adequate space to conduct proposed research and protected time for research, verified by a Letter of Institutional Support.
- The applicant must not hold an active grant from any partner organizations for the scope of work proposed in this application.

Grant Terms and Requirements:

- When an application is approved for funding, the transfer of funds is contingent upon all legal documents being signed and approved by the St. Baldrick's Foundation.
- St. Baldrick's funds may not be used for human embryonic stem cell research.
- Research projects must have direct applicability and relevance to pediatric osteosarcoma.
- All awards will be payable to the academic institution, non-profit research institution, or laboratory to administer for the purposes of this grant only.
- While the Principal Investigator (PI) shall own any invention, the PI shall promptly notify SBF of any invention and associated patent filing resulting from the research. If the recipient or his/her institution grants any right to the invention to a third party for commercial application or sells the invention and in either case receives any amounts for the right to use the invention, the Funding Members shall be entitled to receive a share calculated by multiplying the amounts received by a fraction, the numerator of which is the amount of the Award, and the denominator of which is the direct cost incurred by the recipient and his/her institution in developing the invention, except in no event shall the Funding Members' aggregate share of any such amount received exceed 30%. Any such amount would be remitted to SBF, to be distributed equally to the Funding Members that are registered 501c3 organizations. Such distributions would be restricted for Osteosarcoma research
- All qualified applicants will receive consideration for funding without regard to race, color, ethnicity/national origin, sex, sexual orientation, gender identity or expression, pregnancy, religion, belief and spirituality, age disability status, protected veteran status, or any other characteristic protected by law.
- Applicants can enable other users to access their proposal (e.g., department or grants administrators) in the full proposal section of ProposalCentral.
- It is the responsibility of the applicant to ensure and to verify that the application is received by the deadline and that the application is complete and correct prior to submission.
- All application evaluations are considered confidential and are available to scientific reviewers, the Foundation's Board of Directors, and the administrative personnel of the RFA partners. Reviewer comments will be made available to applicants after funding decisions are made.
- No Cost Extensions must be requested and approved by email: Grants@StBaldricks.org. Requests must be submitted, stating the amount of funds remaining, a brief report of progress, an explanation of why the extension is necessary, and the length of time requested.
- No Cost Extensions can be requested for 6 months or 1 year at a time.
- If a grant reaches an extended end date of 3 years beyond the original end date, no further No Cost Extensions will be considered, and remaining amounts of $100 and over must be returned to the Foundation.
- For multi-year awards, the Principal Investigator may request that funds be moved from a previous or current period into a future period. Carry-forward requests should be submitted in writing (email: Grants@StBaldricks.org) with an explanation for the

- unexpended balance, a plan for the use of funds, and a side-by-side comparison between the original budget and future budget categorical items.
- Re-budget requests should be submitted in writing (email: Grants@StBaldricks.org) with an explanation of each change and a side-by-side comparison between the original budget and revised budget categorical items. Rebudgeting that results in expansion of scope will be reviewed by scientific advisors for approval.
- The Institution, to the full extent legally permissible, shall hold harmless, defend, and indemnify SBF and its officers, directors, employees, scientific advisors, independent contractors, and agents against any and all liabilities, claims, and demands that relate to the research project and activities carried out in connection with this grant, including, but not limited to those for personal injury, property damage, or malpractice.
- The Institution shall be required to maintain adequate liability insurance comparable to coverage held by institutions of similar size and nature, covering the PI, employees, officers, and agents of the Institution for the duration of the Research Project.
- All awards will be payable to the research institution (or to its foundation or funding arm) to administer for the purposes of this grant only. (Consortium grant funding will be made payable to the lead institution to administer on behalf of the consortium.)
- If the lead researcher transfers to a new institution, the funding group will determine if the award may be transferred in part or in whole.
- If awarded, funds will be released upon receipt and sufficiency of all paperwork by the St. Baldrick's Foundation. Any inconsistency noted relative to the award shall be resolved by giving precedence in the following order: The signed award acceptance form; and, Fight Osteosarcoma Together Super Grant RFA and Guidelines.

HUMAN INVESTIGATION

All human subject research or research on human tissue which is supported by St. Baldrick's must comply with the regulations applicable to that supported by the National Institutes of Health. As part of the application, the applicant will be asked to submit documentation of approval of the study by his or her Institutional Review Board, along with the Institution's Human Subjects Assurance Number. Approval is not required at the application stage but will be required prior to issuing funds. If the grantee's research plan changes after the award to include human subjects or tissue, the grantee must submit proper documentation of IRB approval to St. Baldrick's.

BIOHAZARDS

During the application process, the institution shall submit an acknowledgment of potential biohazards involved and provide an institutional statement of assurances regarding potential biohazards and safeguards pertaining to this aspect of the research proposed in the application to the Foundation. Projects which do not involve biohazards must so state. The St. Baldrick's Foundation assumes no responsibility or liability for any such biohazard and shall be held harmless from the results of the use of any such biohazard.

LABORATORY ANIMALS
The Foundation adheres to the most current guidelines applicable to the care and treatment of animals used in laboratory work as outlined by the National Institutes of Health. Prior to award of the grant, the institution must submit a statement that the institution meets and adheres to these policies. Failure to notify the St. Baldrick's Foundation of compliance with these guidelines or the use of laboratory animals may result in termination of the grant.

CONFLICTS OF INTEREST
Principal investigators, co-investigators, sub-contractors or collaborators submitting applications to the RFA will be excluded from serving on the scientific review committee of this grant. However, non-applicants who are invited to serve on the scientific review committee may still have a conflict of interest. When a conflict of interest is deemed to be present, the reviewer will be ineligible to review the proposal and be asked to recuse themselves from the deliberations.

Budget Guidance:

- All budgets must be justified, meet the test of reasonableness, and be consistent with institutional policy.
- No institutional overhead or indirect funding is provided under the terms of the grant.
- Fringe rates should follow institutional policy.
- Fringe benefits are generally defined as medical and dental insurance, life insurance and retirement benefits and are payable only for eligible participation in such programs.
- Personnel on the project are limited to a base salary at or below USD $203,700 per year.
- Overlap in funding is not permitted. St. Baldrick's policy on overlap applies to Scientific, Budget, and Commitment Overlap as defined by the NIH.

  o Scientific Overlap: Scientific overlap occurs when substantially the same research is proposed in more than one application; or is submitted to two or more different funding sources for review and funding consideration; or a specific research objective and the experimental design for accomplishing that objective are the same or closely related in two or more pending applications or awards, regardless of funding source.

  o Budget Overlap: Budgetary overlap occurs when duplicate or equivalent budgetary items (e.g., equipment, salary) are requested in an application but are already funded by another source.

  o Commitment Overlap: Commitment overlap occurs when any project-supported personnel have time commitments (i.e., percent effort) exceeding 100 percent, regardless of how the effort/salary is being supported or funded.

- Situations may occur when a PI has similar proposals pending with different agencies that, if all funded, will present overlap issues. If an overlap occurs before award is accepted, it must be addressed with the St. Baldrick's Foundation before the new award can be accepted, and will be considered on a case-by-case basis.

- If overlap occurs during an award, all recipients must be forthright regarding their support and notify the foundation by emailing Grants@StBaldricks.org. Adjustments will be considered on a case-by-case basis.
- Equipment costs are limited to no more than 20% of total direct costs and only if it is an integral part of the proposed project (specific approval required).
- Reasonable travel costs are allowable.
- Publication and meeting-related poster printing costs are allowable.
- All equipment purchases must be included in the original budget, rebudgets for additional equipment will not be considered.
- Unused and remaining amounts $100 and over must be returned to the Foundation at the end of each grant year, unless an approved carry forward or No Cost Extension is on file.
- All budgets and expenditure reports must be submitted in USD.
- Expenditure reports are expected to follow approved budgets. Any budget line item that changes more than 25% from approved budget to expenditure report (without an approved CF on file) will result in follow up from the foundation for explanation. Explanations will be reviewed for approval.
- This grant does not provide funds for items such as:
  - Secretarial/administrative salaries.
  - Student tuition
  - Office and laboratory furniture.
  - Office equipment and supplies.
  - Recruiting and relocation expenses.
  - Non-medical services to patients (travel to a clinical site or patient incentives are allowable expenses).
  - Construction, renovation, or maintenance of buildings/laboratories.
  - Professional association membership dues
  - Scientific publication subscriptions

CONTACTS

For grant inquiries:
St. Baldrick's Foundation
Department of Grants Administration
Grants@StBaldricks.org
(626) 792-8247, extension 236

For help using the online application system:
ProposalCentral Customer Support
pcsupport@altum.com
 (800) 875-2562

USING ProposalCentral

The St. Baldrick's Foundation uses ProposalCentral (proposalcentral.com) for electronic submission of all LOIs and Full Applications. Fax, email, or hard copy submissions will not be accepted. See "How to Create an Application using ProposalCentral" at https://docs.proposalcentral.com/CreateApp.pdf for more information.

Registration
First-time users must register and fill out a Professional Profile in ProposalCentral to begin the LOI/Application process (see "How to Register as a ProposalCentral User" at https://docs.proposalcentral.com/RegUser.pdf for more information).

Submitting the LOI/Application
1. LOG-IN & BEGIN AN LOI/APPLICATION. To start a new LOI/application, login as an Applicant, and select the Grant Opportunities tab (grey tab on the top right). Find St. Baldrick's Foundation in the drop-down list and select "Filter List by Grant Maker." Find the program for which you wish to apply, and click "Apply Now" to begin the proposal.
2. COMPLETE AN APPLICATION AFTER LOI APPROVAL. Login as an Applicant, and select the Proposals tab. Select the "In Progress" status from the drop-down menu. Find the LOI that was approved, and click the "Edit" button to access and complete your application.
3. ENABLE ACCESS FOR OTHER USERS. The Principal Investigator must start the LOI or Application. However, once an application is begun, access for application submittal can be given to other users, such as collaborators, assistants, or grants and contracts staff.
4. SAVE. Applicants do not need to complete the LOI/Application in one session; a partially completed application can be saved and completed at any time prior to the deadline.
5. NOTIFICATION OF LOI APPROVAL. The system will notify you when the LOI has been approved, and you will be asked to login to the ProposalCentral website to access and submit the full application by the application deadline.
6. VALIDATE. Applications that have not been validated cannot be submitted. "Validate" checks for required items and attachments, and you will not be able to submit if required items and/or attachments are missing.
7. SUBMIT. After successfully passing the validate check and uploading your signature page, click the "Submit" link. An email will confirm your submission. Once your application is submitted, you may view it by accessing the "Submitted" status under the "Proposals" tab.

It is the responsibility of the applicant to ensure and verify that the application is received by the deadline date, before 5 p.m. ET and that the application is complete and correct prior to submission.

Instructions for Submitting Deliverables

The ProposalCentral system is designed to collect and manage all grant information. Grantees must keep their ProposalCentral profile current for the duration of the grant. ProposalCentral should be used to upload all required reports (deliverables) and publications related to the grant. Report templates and deadlines can be found on the site, as well. Grantees may provide access to others at their institution (e.g. grants officers) to access and upload deliverables.

See "Instructions for Navigating Your Awarded Grant" at https://docs.proposalcentral.com/Instructions_Award_Info.pdf for more information.

Uploading Award Deliverables

1. Login to ProposalCentral under the "Applicant Login" section: https://proposalcentral.com/.
2. Click the "Awards" tab.
3. Click the blue Identifier number link.
4. Click the blue "Deliverables" link.
5. On this page you will see a table which lists all of the deliverables that are scheduled for your grant. The templates for these deliverables are at the bottom of this page.
6. To upload completed deliverables, click the blue "Upload" link next to the appropriate deliverable. If you have a pop-up blocker turned on, make sure you disable it for this site as the deliverable upload feature is a pop-up.
7. In the deliverable pop-up window, click the "Browse" button to select the file. You can add a description in the "Deliverable Description" if you choose. When finished, click the "Save" button, and it will upload your deliverable.
    o When completing the Publications Webform, click the blue plus icon to enter information on publications related to your SBF award
        ▪ If there are no publications to report, click the "No publications to report" checkbox
    o When completing the Budget Webform, click the blue plus icon to be able to enter your expenses in the Expenditure column on the right
        ▪ Please note, only contacts who have Administrative and/or Edit access to the award are able to electronically sign and submit this webform. If there are others who may need to sign and submit the budget, please let us know so we can grant them access to do so.
        ▪ Instructions on using the budget webform can be found at the ProposalCentral link, starting at the bottom of Page 11: https://docs.proposalcentral.com/Instructions_Award_Info.pdf

Useful Resources for ProposalCentral

• How to register your institution (grants and contracts personnel only): https://docs.proposalcentral.com/RegInst.pdf
• How to register as a ProposalCentral user: https://docs.proposalcentral.com/RegUser.pdf
• How to create an application using ProposalCentral: https://docs.proposalcentral.com/CreateApp.pdf

- Grantee instructions to access award information:
https://docs.proposalcentral.com/Instructions_Award_Info.pdf

Contact
For help using the online application system:
ProposalCentral Customer Support
pcsupport@altum.com
800.875.2562