

CHILDREN'S
TUMOR
FOUNDATION
ENDING NF
THROUGH RESEARCH



CHILDREN'S TUMOR FOUNDATION

ANNUAL
REPORT
2022

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What is NF?

NF is a term for a group of genetic conditions that cause tumors to grow on nerves throughout the body. NF includes neurofibromatosis type 1 (NF1) and all types of schwannomatosis (SWN), including NF2-related schwannomatosis (NF2), formerly known as neurofibromatosis type 2. Some type of neurofibromatosis or schwannomatosis occurs in approximately one in every 2,000 births. These conditions affect all populations equally, and may lead to blindness, deafness, bone abnormalities, disfigurement, learning challenges, disabling pain, or cancer.



The Children's Tumor Foundation

is the world's leading nonprofit dedicated to funding and driving innovative research that will result in effective treatments for the millions of people worldwide who live with NF.

Our mission

Drive research, expand knowledge, and advance care for the NF community.

Our vision

End NF.



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CTF President
Annette Bakker

Dear friends,

As I reflect on the last year, I am thrilled by our progress in advancing treatments for NF patients. At the Children's Tumor Foundation (CTF), 2022 was defined by a thoughtful, forward-looking view as we built a bold, visionary five-year strategic plan dedicated to expediting effective and tolerable treatments to patients. I could not be prouder of this transformational vision, which is summarized in this annual report. As we move forward, our unwavering focus is to lead and accelerate the innovation that ends NF. We focus on investing in high-impact projects that will accelerate treatments, strengthen clinical trials, and empower stakeholders. We will not quit until there are treatments for all manifestations of all types of NF.

Recognizing the inherent strength in collaboration, we realize this ambitious mission cannot be achieved alone, and CTF remains committed to forging fruitful partnerships and growing our community. We continue to invite pharmaceutical and biotech companies to negotiate access to additional drugs for clinical trials.

The year has landed us in an exciting moment, with many treatment options on the horizon!

Ten pharmaceutical companies are now working on NF, of which two are entirely dedicated to NF. NFlection Therapeutics is running a Phase 2b clinical trial with a topical MEK inhibitor; SpringWorks Therapeutics is showing encouraging data with a MEK inhibitor drug; and the INTUITT-NF2 clinical platform trial has shown encouraging data on Brigatinib, and has now included a second drug, Neratinib. Lastly, in collaboration with other funders, our investments in gene therapy are showing real promise.

Our NF Conference continues its reign as the most important meeting for NF in the world, and the 2022 conference in Philadelphia brought together more than 600 researchers and clinicians to discuss the most cutting-edge advancements in the field. Meanwhile, our NF Masterclasses (INFER) offer all medical professionals access to a series of educational lectures by leading NF experts. We also continue to improve our patient resources, including educational webinars, symposia, and informational brochures.

2022 saw the publication of revised diagnostic criteria and nomenclature for schwannomatosis, including NF2-related schwannomatosis (formerly called neurofibromatosis type 2). These changes will result in earlier diagnosis and better care for patients. We expanded our NF Clinic Network to include new adult clinics in Los Angeles and Little Rock, ensuring new and previously diagnosed patients have needed information and support.

Because NF knows no borders, we will continue to strengthen our global activities throughout Europe, Latin America, Asia, and beyond. To this end, we translated the NF Registry into five languages, with more to come. We are proud to inform you that 26 million euros from the European community have resulted in comprehensive platform trials for NF1 and schwannomatosis. We are ready to embark on its implementation in partnership with European colleagues.

World NF Awareness Day 2022 painted a clear picture of the reach of this community, as thousands around the world gathered virtually to watch our awareness day livestream, raising thousands of dollars for NF research. Our "Make NF Visible" and "Shine a Light on NF" awareness campaigns continued to attract more people to the cause. CTF Shine a Light Walk, NF Endurance, and Cupid's programs continue to flourish, providing more opportunities for the community to unite, grow, and raise funds for this fight.

I extend my heartfelt appreciation to everyone in this incredible community — the researchers, clinicians, regulators, industry partners, our dedicated Board, fundraisers, donors, volunteers, and our patients, their families, and caregivers. Together, we are in this fight, and only through our collective efforts will we emerge victorious. I want to say how grateful I am to all of you. Your vision of a world without NF has guided our drive to bring treatments to patients. I could not be more excited for what's to come.

Warmly and gratefully,



Annette Bakker, PhD, President

Leading the Way

At the Children’s Tumor Foundation, we’re working to better the lives of more than 4 million people who live with some type of neurofibromatosis or schwannomatosis. We envision a day when patients can live their lives free of the pain and difficulties that come with nerve tumors, and our innovative team-based approach to drug development is making that vision a reality.

NF patients are told to “watch and wait” to see if their tumors will grow, and determine later if they will impact their lives with devastating conditions such as cancer. We don’t think that’s fair, and we don’t think that’s necessary. By bringing together the brightest minds in research and industry, and revamping the systems that often slow the pathway to treatments, we can change “watch and wait” to “here’s what you can do.”

And the best part? CTF’s research model not only benefits patients with neurofibromatosis or schwannomatosis, but also the millions of patients living with cancer, or one of the 7,000 rare diseases in existence.

We’re in a rush to find treatments for all tumor types and all manifestations. Join us as we revolutionize how treatments are developed for those who need them most.

TRADITIONAL RARE DISEASE MODEL

PATIENT

TREATMENT

COST:
Hundreds of millions of dollars

Why does it take so long and cost so much?

- Patient manifestations are unclear
- Clinical trial recruitment can be slow
- Disconnect between discovery and treatment
- Data is not shared and reporting is delayed
- Experts are isolated and knowledge is lost

CTF RESEARCH MODEL

PATIENT

TREATMENT

GOAL:
Double the speed for half the cost!

Why are we faster?

- NF Patient Registry accelerates clinical trial enrollment
- Team science connects discovery to treatment
- Open NF datahub for real-time data release
- Preclinical platform accelerates drug testing
- Key opinion leader network speeds up decision-making

2023-2028 Strategic Plan

We envision a world without NF, and our strategic plan aims to make this vision a reality by driving research, expanding knowledge, and advancing care for the NF community.

Historically, the Children's Tumor Foundation has laid a firm foundation for the critical work ahead by building NF awareness, education, and community. This has complemented a robust emphasis on NF research funding and the building of essential networks. **As we move forward, our unwavering focus is to lead and accelerate the innovation that ends NF.**



INNOVATION PILLARS

Develop More Drugs - Expand the research field, develop panels of drug selection tools, double the number of R&D grants, and launch revolutionary projects.

Accelerate Treatments - Expedite the discovery of life-changing therapies with an innovative preclinical hub-and-spoke model, the ultimate all-in-one solution for pharma and biotech pioneers.

Strengthen Clinical Trials - Accelerate the development and approval of drugs being evaluated in clinical trial consortia and/or platform trials.

Empower Stakeholders - Enhance the understanding of NF and the NF Registry, increase patient participation, and train ambassadors. Expand the NF Clinic Network and the number of physicians and specialists who treat NF, including adult care.

FUELED BY

- Research & Data that is Patient & Clinician Informed
- Global Action and Inclusively in all Elements of our Support
- Action Over Talk: Be Assertive & Proactive
- \$130M Over Six Years: Prioritizing Innovation Leading Investments

Selu-bration

Koselugo (selumetinib), the first approved treatment for NF

In the spring of 2020, the U.S. Food and Drug Administration (FDA) announced the approval of Koselugo (selumetinib) for use in patients with inoperable plexiform neurofibromas, a common manifestation in neurofibromatosis type one (NF1). The FDA's approval of AstraZeneca's and MSD (Merck)'s submission was a major milestone for patients living with NF. Koselugo is the first-ever approved treatment for NF, and portends the potential for the development of more treatment options for all types of NF.

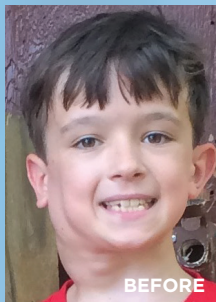
Koselugo's approval followed comprehensive clinical testing of the drug in patients at the National Cancer Institute (NCI), a division of the National Institutes of Health (NIH). In those clinical trials, over 70% of NF patients with inoperable plexiform neurofibromas saw tumor size reduction anywhere from 20–60% in size. In addition to both visible and actual tumor reduction, patients reported

higher-quality physical function, reduced pain, improved mobility, and enhanced emotional and psychological status.

Many other MEK inhibitors are also now in clinical trials, including mirdametinib from SpringWorks Therapeutics, a company which the Children's Tumor Foundation helped spin off from Pfizer. CTF is also partnering with NFlection Therapeutics on a Phase 2b clinical trial with a topical MEK inhibitor for patients with cutaneous neurofibromatosis type 1.

Since the 2020 announcement, Koselugo has been approved for the treatment of NF patients in 32 countries. The AstraZeneca group of companies acquired Alexion, a global biopharmaceutical company focused on rare diseases. Alexion now distributes Koselugo throughout the world, adding to the growing list of partners working with the Children's Tumor Foundation to end NF.

Koselugo: Stories of the Road to Approval



60%
shrinkage

"People don't ask me what is wrong with my neck anymore. The drug does make me fatigued, which is tough since most kids my age are active in sports or physical activities that are challenging for me. But I've found activities that I enjoy, and friends that enjoy being a part of my life. I enjoy reading, gaming, boy scouts, coding, and more. I'm getting ready to start high school and I'm thankful for all the donors that funded the doctors and researchers who made selumetinib possible. Now that it is FDA approved, I am thankful that others may experience what I have experienced."

— NF Hero Philip Moss

SELUMETINIB: The Path to Approval

CTF research discovery paved the way to the first FDA-approved drug for NF.

NF Preclinical Initiative

The NF Preclinical Initiative (NFPI) began in 2008 as the NF Preclinical Consortium (NFPC), a five-year, \$7 million program that concluded in 2013. The NF Therapeutic Consortium (NFTC) continued the work of the NFPC, building on its infrastructure and discoveries.

Traditionally, it takes more than 15 years and costs hundreds of millions of dollars to translate a new discovery into one clinical treatment. The impact of the NFPI was clear: these teams completed 116 preclinical trials in 8 years, at a total cost of \$11 million. The preclinical studies led to multiple clinical trials, many of which are currently underway. One of those clinical trials included the MEK inhibitor selumetinib registration trial.

MEK Inhibitor Selumetinib...the road to the first approved drug for NF

Over 70% of the patient participants in a clinical trial of the MEK inhibitor selumetinib saw a reduction of 20–60% in the size of their inoperable plexiform neurofibromas. Brigitte Widemann, MD, of the National Cancer Institute, reported at the Foundation's 2015 NF Conference that response data in this trial for children and young adults showed meaningful decreases in tumor volume. This highly successful clinical study was the result of CTF's NF Preclinical Initiative, wherein Nancy Ratner, PhD, and D. Wade Clapp, MD, demonstrated that MEK inhibitors (MEKi) have a massive result on plexiform neurofibroma tumor volume in mice. The drug was first tested in a human subject as part of a Children's Tumor Foundation Clinical Research Award granted to Michael J. Fisher, MD, in 2011.

The Path to Approval

Selumetinib, a MEK inhibitor, was granted Breakthrough Therapy Designation by the U.S. Food and Drug Administration (FDA) in April 2019. Previously, the treatment was granted Orphan Drug Designation by the US FDA in February 2018 and by the European Medicines Agency (EMA) in August 2018. The news that selumetinib was granted these designations was a giant leap toward the thrilling April 2020 announcement that selumetinib had received FDA approval.

Attracting Pharma

The Children's Tumor Foundation provides answers to the questions that pharmaceutical companies are asking in order to invest in a rare disease such as NF.

Are teams of scientists working on this problem?

CTF COLLABORATIONS AND TEAM SCIENCE INITIATIVES

The CTF research model values collaboration in every aspect of the drug discovery process. Diverse teams of experts managed by the Foundation collaborate and share their discoveries and data. By working together instead of in silos, we greatly increase the efficiency of research into problems that are too complex to be solved by individual scientists.

Are there enough care and treatment centers?

THE CTF NF CLINIC NETWORK IS CONNECTING DOCTORS AND IMPROVING CARE

A growing network of CTF-affiliated clinics is cultivating relationships between patients and doctors, and working to standardize and improve neurofibromatosis and schwannomatosis patient care.

Are there patients engaged in the drug discovery process?

CTF PATIENT ENGAGEMENT PROGRAM

Patients and caregivers are recruited to our patient engagement training program, creating a team of Patient Representatives who are knowledgeable in all aspects of NF drug discovery.

Are other investors interested in NF?

CTF AS IMPACT INVESTOR

CTF entered a new phase in its strategic NF research model by making a significant impact investment in a Phase 2b clinical trial at NFlection Therapeutics. This impact investment further establishes CTF as a co-investor with pharma/biotech in innovative NF research, expanding opportunities to increase the NF drug development pipeline.

Where do we find experts?

CTF'S KEY OPINION LEADER NETWORK

This expert network of specialists helps to guide drug discovery and development in order to increase scientific and clinical quality in decision-making.

Where can we find the patients to participate in clinical trials?

THE NF REGISTRY CONNECTS PATIENTS TO CLINICAL TRIALS

This patient-entered registry is structured to accelerate clinical trial recruitment, and fuels knowledge and understanding of the diversity of NF manifestations.

Where can we find new drug targets?

THE NF DATA PORTAL STORES OPEN DATA

Through our partnership with Sage Bionetworks, data is available and ready to use in the NF Data Portal. This centralized data repository is managed by specialists who collect, analyze, and release integrated data, accelerating the understanding of all types of neurofibromatosis and schwannomatosis, and the identification of “druggable” targets.

Is there enough tissue available for testing?

THE NF BIOBANK PROVIDES TISSUE FOR RESEARCH

In order to solve the problem of a scarcity of relevant tissue to test, CTF created a centralized library of openly available samples for biomarker discovery and development, to support all aspects of drug research. This Biobank is now managed by the Indiana University School of Medicine’s DHART SPORE program.

How can we standardize the endpoints of clinical trials?

THE REINS CONSORTIUM (Response Evaluation in Neurofibromatosis and Schwannomatosis)

This worldwide consortium of clinicians and patients develops new clinical trial designs, and works with the FDA to establish drug approval criteria.

Have these drugs been tested in animal models?

CTF’S NF PRECLINICAL INITIATIVE ESTABLISHED TREATMENTS FOR CLINICAL TRIALS

Because early testing of innovative concepts is vital, CTF has invested in teams of top laboratories with NF-relevant animal models, bringing promising drug treatments to the clinic quickly and efficiently.

Driving Collaboration

Children's Tumor Foundation's Team Science Initiatives

INTUITT for NF2

The Children's Tumor Foundation partners with Takeda Pharmaceuticals and six leading medical centers on a clinical trial called INTUITT-NF2, an innovative platform trial that evaluates multiple treatments simultaneously. This initiative is a result of the landmark work of CTF's visionary Synodos for NF2 research collaborative, its NF2 Accelerator Initiative, an investment from Takeda Pharmaceuticals, the participation of scientists at the National Center for Advancing Translational Sciences (NCATS) at the National Institutes of Health (NIH), and the vital Synodos NF2 leadership from Massachusetts General Hospital (MGH), Johns Hopkins University (JHU) and Indiana University (IU). This alliance across the academic, pharma, and patient landscapes has shortened the time from initial research to active trial, thereby bringing promising treatment options to the NF2 patients who need them.

Biomarker Project for NF1

In 2022, the Children's Tumor Foundation announced a groundbreaking three-year study, which it will fund for nearly \$2 million, to determine if a DNA-based blood test can offer better understanding and ultimately earlier diagnosis of malignant peripheral nerve sheath tumor (MPNST) predisposition in NF1 patients. CTF has committed its support to fund the study in its entirety. The project format is inspired by CTF's highly effective Synodos model, which brings top researchers together to solve complex problems in an open and collaborative environment.

Optic Pathway Glioma Multicenter Study

The Children's Tumor Foundation and the Children's Hospital of Philadelphia (CHOP) have renewed their five-year study of optic pathway glioma (OPG) in children with NF1. The study involves 25 NF clinics, and aims to provide clinicians with clear criteria that will help them decide when a patient should be treated, and when treatment (such as chemotherapy) should be avoided. OPG develops in 15–20% of children with NF1, and can cause significant health issues. In 2022, the study was renewed, and renamed to memorialize late CTF ambassador Jeffrey Owen Hanson, who suffered from OPG.

Synodos

Synodos is the premier collaborative research model of the Children's Tumor Foundation, and has represented a significant financial commitment on the part of the Foundation. In each Synodos collaboration, CTF assembles "dream teams" of doctors, scientists, and patients who work together to solve problems that are too complex for any individual lab or researcher to solve. Our goal is to speed the drug discovery process through innovative research methods, collective knowledge, and data shared openly through CTF's NF Data Portal. This data is analyzed by our partners at Sage Bionetworks, and is then made available to the rest of the world. This approach is expanding interest in NF research to additional fields, including cancer.

Building on Past Success: Synodos for NF2

The first of the Synodos models established by CTF, Synodos for NF2 launched in 2014 to provide clarity to patients about available drug options, and completed its work in 2018. This consortium of multidisciplinary investigators from 12 world-class labs and medical centers performed rigorous drug testing that laid the groundwork for further research and clinical trials currently in progress for the treatment of NF2-related schwannomatosis.

Building on Past Success: Synodos for NF1

CTF sought out the concerns of NF1 patients when establishing Synodos for NF1, a team-science initiative that began its work in 2016 and concluded in 2020. Patient concerns for better treatment options for NF1, as well as answers for low-grade glioma patients, prompted CTF to initiate three separate consortia, bringing together 24 investigators from eight leading institutions and two companies.

Building on Past Success: Synodos for Schwannomatosis

An international consortium of clinicians and scientists from multiple disciplines made up the Synodos for Schwannomatosis team. In 2021 the project completed the work of performing an extensive molecular analysis of schwannomatosis tumor samples to identify new therapeutic targets and advance the understanding and management of the disease, with a special focus on pain.

Fueling Discovery



CTF Discovery Fund

The Children's Tumor Foundation Discovery Fund for NF Research funds more than \$3 million in research grants each year, and accelerates drug discovery. This initiative is set up to attract and invest in the best and brightest minds, who will advance our goal of bringing new treatments to patients faster and more efficiently.

The Young Investigator Award (YIA)

The YIA is the Foundation's oldest research award program and serves to advance understanding of the biology of all types of neurofibromatosis and schwannomatosis, and brings young researchers into the field. This award program is one reason the understanding of NF has grown so rapidly. Many of CTF's past YIA awardees have gone on to pursue lifelong careers in the field of NF research.

Drug Discovery Initiative (DDI)

The CTF Drug Discovery Initiative (DDI) program aims to stimulate NF drug discovery by funding researchers proposing to investigate novel or repurposed therapies for NF or to develop tools that support such research. The goals of the DDI program are to support early-stage testing of therapeutic compounds for the treatment of NF, or to support the generation of new *in vitro* or *in vivo* model systems. CTF has awarded more than 70 DDI awards since the inception of the program in 2006.

Drug Discovery Initiative Registered Reports (DDI-RR)

In 2017, CTF and the scientific journal *PLOS ONE* launched a funding program called the Drug Discovery Initiative Registered Report (DDIRR) Awards, which evolved from the DDI program and was a funder-publisher partnership that guaranteed its awardees with an in-principle acceptance to publication in the journal *PLOS ONE*, regardless of study outcome.



Contract Awards

The Contract Awards are special awards that the Foundation assigns to academic researchers or for-profit entities to run specific projects. The Contract Award is not a typical award but rather an objective and task-oriented project that allows the recipient to access funding otherwise not obtainable through other grant mechanisms.

Clinical Research Award (CRA)

The Foundation's Clinical Research Award program supports early-stage NF research involving human subjects. These awards encourage studies of candidate therapeutics or other interventions and treatments; clinical-trial-enabling or ancillary studies; natural history studies; and investigations into clinical care in NF.

NF1 Gene Therapy Initiative

The NF1 Gene Therapy Initiative has the objective of exploring the feasibility of gene editing as a potential therapeutic strategy for NF1. This initiative aims to support proof of principle *in vitro* studies to investigate the feasibility of genome editing techniques, including but not limited to those based on CRISPR-Cas9, to correct pathogenic mutations in *NF1* gene.

We would again like to thank you and all those who donated to CTF, for believing in our projects and funding our CRAs. We are proud that our CTF awards have now produced over \$5 million in federal research grants.

— Drs. Rob Avery, Michael Fisher, and Gena Heidary

Accelerating Innovation

A woman with dark hair tied back, wearing a white lab coat over a dark top and blue nitrile gloves, is shown in profile from the chest up. She is looking towards the right side of the frame. The background is a blurred laboratory setting with various pieces of equipment and shelves.

NFX-179 Impact Investment

The Children's Tumor Foundation entered a new phase in its strategic NF research model by announcing a significant impact investment in a Phase 2b clinical trial at NFlection Therapeutics, a biotechnology company focused on the discovery and development of effective, targeted therapies for rare diseases. The trial involves NFX-179, a topical (on the skin) treatment which has successfully passed a Phase 1/2a (safety and first signs of efficacy) trial. The NFX-179 treatment is for cutaneous neurofibromas (cNF), which are tumors that grow in the skin or right underneath the skin, and result in severe, disfiguring bumps on the skin.

CTF's investment strategy in NF drug development is a diversified portfolio approach to tackling all forms of NF, from early-stage investments in young investigators, open data collaboratives such as Synodos. This first-in-its-history impact investment further establishes CTF as a co-investor with pharma/biotech in innovative NF research, expanding opportunities to increase the NF drug development pipeline.

NF2 Accelerator Initiative

This powerful initiative is bringing treatments to the clinic (and patients) by expanding the clinical drug pipeline for NF2, improving drug selection through the development of innovative testing models, and the development of gene therapy options that address the underlying genetic causes of NF2-related schwannomatosis.

As a global leader of NF research, the Children's Tumor Foundation is dedicated to developing cures for all forms of NF. The Foundation and its partners bolstered the NF2 research field through team science with Synodos for NF2, bringing together multidisciplinary scientists from 12 world-class labs. That collaborative effort resulted in the identification of promising new clinical candidates for NF2. The NF2 Accelerator Initiative leverages this new knowledge into a new and ambitious structure, with the goal of speeding up the drug discovery process.

Strategic Partnership: Deep Science Ventures and Cancer Research Horizons

The Children's Tumor Foundation has joined an alliance with Deep Science Ventures, a London-based venture creator, and Cancer Research Horizons, Cancer Research UK's new innovation organization. Together these groups will power the creation of cutting-edge ventures in oncology, with CTF acting as a strategic partner to aid efforts in identifying possible venture-creation opportunities in NF.

Focused Ultrasound Foundation and CTF Partner on NF Study

The Focused Ultrasound Foundation and the Children's Tumor Foundation have established a partnership to advance innovative, noninvasive treatments in pediatrics. For more than 15 years, the Focused Ultrasound Foundation has been dedicated to advancing the development and adoption of focused ultrasound. To launch the partnership, the organizations are co-funding an early-stage laboratory study to investigate focused ultrasound's role in addressing NF2-related schwannomatosis.

Genotype-Phenotype

Research funded by CTF's Isaac and Sadie Fuchs Genotype-Phenotype Grant found that certain NF1 genotypes can help predict phenotypes. Led by Ludwine Messiaen, PhD, of the University of Alabama at Birmingham, this discovery of genotype-phenotype correlation can impact counseling and management of more than 10% of the NF1 population.

The BRIDGE Initiative

The Children's Tumor Foundation has joined forces with the Milken Institute's FasterCures and CureSearch for Children's Cancer in a collaborative effort called the BRIDGE Initiative, which aims to convince pharmaceutical and biotech companies to release discontinued but valuable medicines. The BRIDGE Initiative is committed to unlocking these drugs for intended or new indications, such as for NF.

NF Diagnostic Criteria Workshops

In 2017, a group of NF investigators reached out to CTF to sponsor a revision of the NF diagnostic criteria, sparking a multi-year process that involved more than 90 leading NF experts from around the globe. In May of 2021, an update to the diagnostic criteria for NF1 was published in *Genetics in Medicine*, the official journal of the American College of Medical Genetics and Genomics (ACMG). An update to the diagnostic criteria and nomenclature for NF2 and schwannomatosis was announced in the same publication in early 2022. These updates reflect the tremendous increase in knowledge about these conditions, and will allow for earlier and more accurate diagnoses for patients.

NF Variant Curation Panel

The Children's Tumor Foundation has funded a ClinGen (Clinical Genome Resource) driven initiative to build a central resource that defines the clinical relevance for all NF gene variants (*NF1*, *NF2*, *SMARCB1*, *LZTR1*, *SPRED1*) for use in precision medicine and research. The expert panel is composed of 25 experts among molecular and clinical geneticists, genetic counselors, and other experts in NF from 9 countries and 18 different institutions.



NF Conference

The annual NF Conference attracts clinicians, researchers, industry representatives, and patient advocates, in person and virtually, who are interested in hearing about the most recent advancements in research and clinical care for neurofibromatosis and schwannomatosis. The event is a critical forum for consensus-building and advancing basic, translational, and clinical research in NF and related fields, while fostering collaborations within and beyond the NF community.

In 2022 the NF Conference convened in Philadelphia, Pennsylvania, and more than 600 attendees were presented with the best and latest research and clinical care practices. In addition to the core agenda, attendees were able to view poster presentations while a panel of judges selected the top three submissions from clinical and basic science entries. The meeting wrapped up with the presentation of the 2022 Friedrich von Recklinghausen Award, awarded to Jaishri Blakeley, MD, of Johns Hopkins School of Medicine.



NF Hackathon

The Children's Tumor Foundation hosted its fourth Hackathon, the HACK4NF, in late 2022. The goal of this virtual hackathon was to advance research for all types of NF by utilizing genomic datasets to develop drug targets, predict NF variants and correctly classify NF tumors. More than 300 people from diverse fields participated in the HACK4NF, including those with expertise in computational biology, predictive AI, genomics, data analytics, and medical research.



CTF in Europe

Children’s Tumor Foundation Europe is a medical research NGO that serves as a partner organization to CTF in the United States. Throughout its history, CTF has funded the best and most promising research globally, regardless of location, and as a result many European laboratories and clinicians have benefited from CTF support. Children’s Tumor Foundation Europe is further strengthening the bonds between experts and research opportunities worldwide, in line with CTF’s emphasis on open collaboration and open data.

The focus of Children’s Tumor Foundation Europe is to raise awareness of NF at the European level, and build relationships with European agencies and partners, including EFPIA (European Federation of Pharmaceutical Industries and Associations) and the EMA (European Medicines Agency, the European equivalent of the FDA), while maintaining its commitment to funding and driving innovative research worldwide that will result in effective treatments for NF.

In 2022, the Children’s Tumor Foundation Europe further focused on the consolidation of European clinic networks, the organization and planning of INFER (International NF Educational Resources) masterclasses for healthcare providers, and provided travel grants to medical professionals for their training in European NF excellence centers, as well as training to European researchers to allow them to

successfully apply for grants issued by US funding bodies. CTF also continued to work at the expansion of the NF Registry in Europe in partnership with national patient associations.

CTF and 35 other organizations joined together in a project called EU Patient-centric clinical tRIal pLatforms (EU-PEARL), a unique public-private strategic partnership funded by the Innovative Medicines Initiative to conceptualize and lead the design of integrated research platforms, enabling a more efficient and patient-centric drug development in Europe. CTF is the co-leader of the neurofibromatosis package of this exciting project, where NF is serving as a test case for rare diseases in general.



INFER

INTERNATIONAL
NF
EDUCATIONAL
RESOURCES

A series of online educational lectures for medical professionals by leading NF experts

Sustaining Hope

NF Summit

The 2022 inaugural NF Summit was an educational and networking conference that evolved from CTF's annual NF Forum and Volunteer Leadership Conferences, which have taken place for over a decade. Those annual meetings built the necessary foundation for a new, dynamic, inclusive event intended for anyone with a connection to NF. The NF Summit brought together over 50 speakers covering a wide range of content, including chronic pain, hearing loss, and using social media to raise awareness and build community. Of those speakers, 36 had never before attended or spoken to the NF community at a CTF event. Community Recognition Awards (The Make NF Visible Community Awards) were presented to volunteers, clinicians, researchers, corporate partners and community members who strive to make NF visible in various ways and who are true champions of the Foundation and the NF cause.

NF Clinic Network (NFCN)

The NF Clinic Network was established by the Children's Tumor Foundation to standardize and raise the level of NF clinical care, and to integrate research into clinical care practices. The NF Clinic Network (NFCN) has grown to 69 clinics in the US and Canada that serve almost 20,000 patients each year. Clinics may apply to join the NFCN and are evaluated based on several factors, including NF expertise, patient volume, multidisciplinary approach, commitment to NF education and training, research activities, and connections with the Foundation.

Patient Engagement

The Foundation's Patient Engagement initiative is designed to prepare individuals with NF and their families to participate as advocates in NF research. Patient Representatives are trained to work with stakeholders such as researchers, the pharmaceutical industry, and the U.S. Food and Drug Administration to add their perspectives during all phases of the research process. In 2022, the Children's Tumor Foundation took the opportunity to examine the broader potential for patient engagement in the field. As we move into 2023, CTF is excited to build on the success of the past several years with a newly designed patient engagement initiative called CTF Engage, which promises to elevate the patient voice in research in even more impactful ways.



NF Camp

More than 100 teens and young adults attend CTF's annual NF Camp each year. NF Camp takes place at the beautiful Camp Kostopulos in Emigration Canyon, Utah, with ropes courses, horseback riding, and more. Attendees participated in games, kitchen takeovers, arts and crafts, and laughter. Campers were able to come together to talk freely about their NF journeys, share their experiences, and make lasting friendships.

NF Registry

More than 11,000 individuals have joined the Foundation's NF Registry, making it the largest patient-entered database of people with NF, and the only one designed to be available to all interested investigators. The Registry has proven to be a valid and useful tool for both patients and researchers, according to an analysis published in the journal *PLOS ONE*. Not only are thousands of patients from all over the world contributing their data online at nfregistry.org, but the data is being actively used—and appreciated—by researchers working on all forms of NF, as well as attracting the attention of pharmaceutical companies. The Registry is now on the OpenApp platform, which has made the site easier to use and allowed it to be translated into multiple languages, further increasing participation.

“

Over the decades, this organization has done more to advance care of families with NF than any other U.S. organization, and it has not lost sight of the importance of interactions between individuals with these conditions, as exemplified at NF Camp. CTF truly enhances my care of families dealing with all issues related to NF.

— David Viskochil, MD, PhD, University of Utah

”

Spreading Awareness

CTF's marketing, communications, and advocacy efforts support patients and their families no matter where they are on their NF journey. Whether newly diagnosed, in the midst of a treatment regimen, or engaging with the broader community to improve awareness and understanding, patients and families can rely on CTF for the latest information about all types of NF.

All too often the patient journey starts with an online search that leads to inaccurate or outdated information, and so the Children's Tumor Foundation prides itself on being a safe haven for all who need direction and support. We provide up-to-date NF knowledge on our website and in our patient brochures, newsletters, webinars, and videos. We also drive a dynamic and engaging presence on social media, connecting patients and families worldwide, and ensuring that those

who don't have NF support in their community can find it through the global CTF family.

In a world that runs 24/7, the Foundation breaks through with impressive media outreach and public relations efforts that spread the message that NF is important and that NF patients deserve support for bettered lives. Our multichannel approach in print, digital, TV, and radio results in hundreds of media pickups each year, and brings the NF story to hundreds of millions of people worldwide.

While NF is a serious condition, the Foundation creates inspirational and engaging tools that patients can use to share their NF story in a personal way. From "I Know a Fighter" to "Shine a Light on NF" to "Make NF Visible," we make sure that everyone knows our driving passion: to END NF.

The Foundation's "Make NF Visible" campaign draws attention across the globe to those living with NF.

Make NF Visible
Children's Tumor Foundation

BORN A FIGHTER

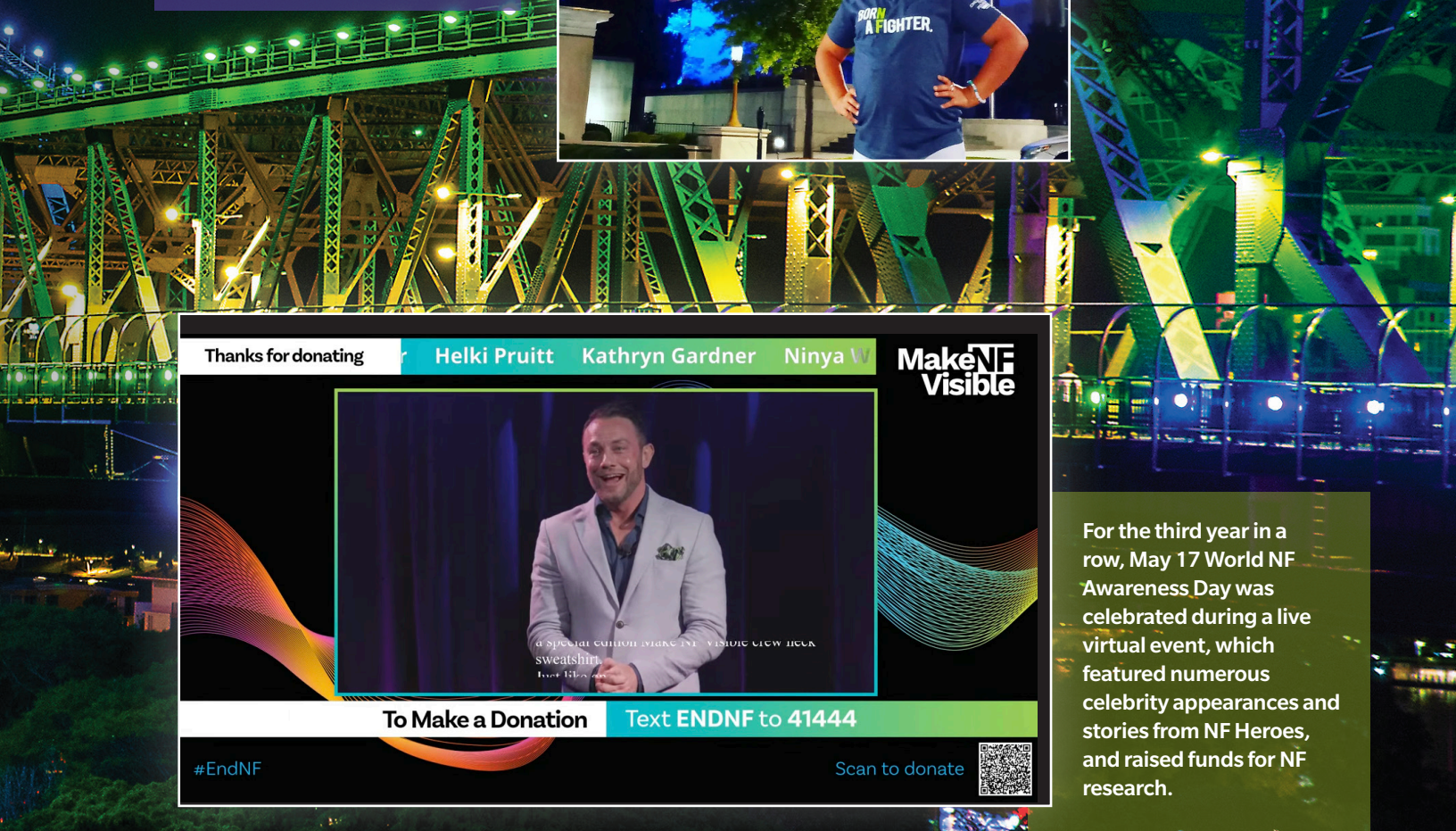
“

Making NF visible cultivates community, connection, and deep friendships with others just like you. . . . By ourselves, we can only go so far, but together we can go even further.

”

— Nissa Novas, who lives with NF2

The "Shine a Light on NF" initiative cast a blue-and-green glow on hundreds of buildings, bridges, monuments, and family residences across the country and around the globe.



Thanks for donating Helki Pruitt Kathryn Gardner Ninya W

MakeNF Visible

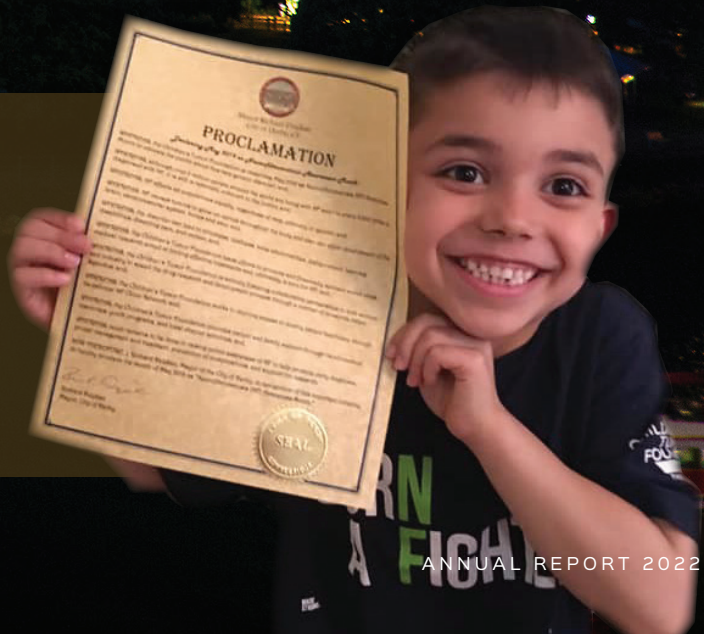
a special edition MAKE NF VISIBLE crew neck sweatshirt

To Make a Donation Text ENDNF to 41444

#EndNF Scan to donate

For the third year in a row, May 17 World NF Awareness Day was celebrated during a live virtual event, which featured numerous celebrity appearances and stories from NF Heroes, and raised funds for NF research.

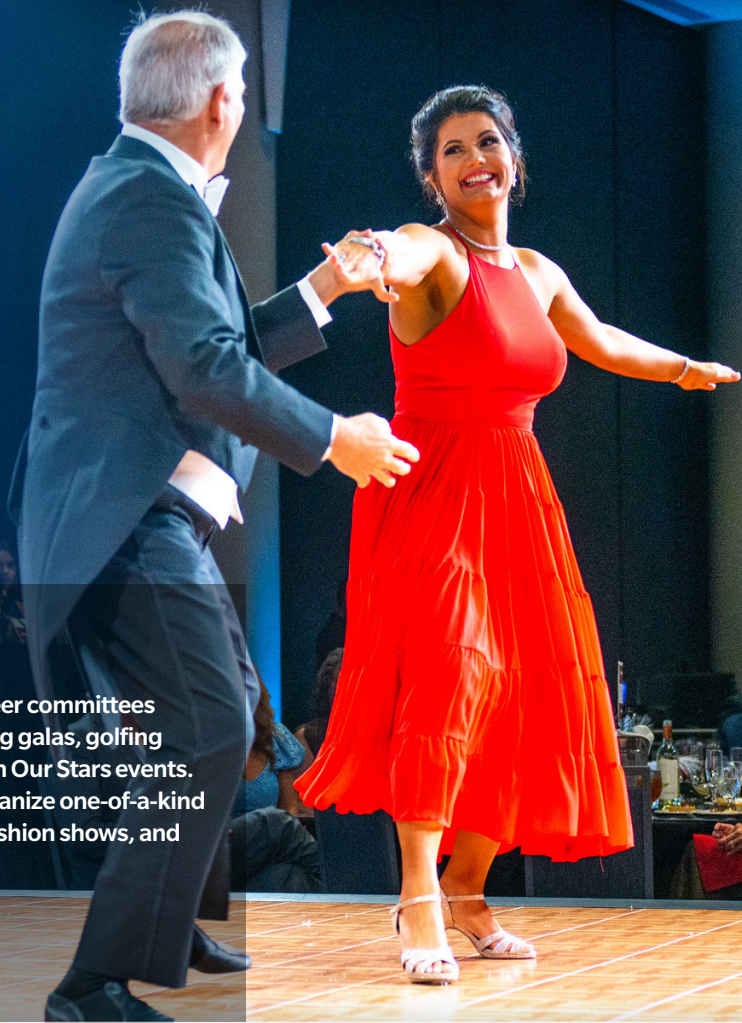
Proclamations were issued in state houses and city halls across the country recognizing "NF Awareness Month." CTF volunteers engaged with their local leaders and helped ensure that more people in office know about NF and how it affects their constituents.



Raising Funds

Special Events

The CTF Special Events team works with volunteer committees across the country to organize festive fundraising galas, golfing tournaments, cocktail parties, and Dancing With Our Stars events. Meanwhile, “Fight NF Your Way” volunteers organize one-of-a-kind fundraising events, including comedy nights, fashion shows, and sweet tea stands that help fund NF research.



Shine a Light NF Walk

Shine a Light NF Walk is the signature fundraising event of the Children’s Tumor Foundation, bringing NF out of the shadows and inspiring the community to come together to raise critical funds for NF research. At these fun and inspirational events held across the country, communities rally around local families affected by NF. Frequent appearances of costumed characters, balloon animals, and face painting bring bubbling energy from start to finish. The 2022 Shine a Light Walk season included 4,600 participants and raised \$1.53 Million to further NF research.

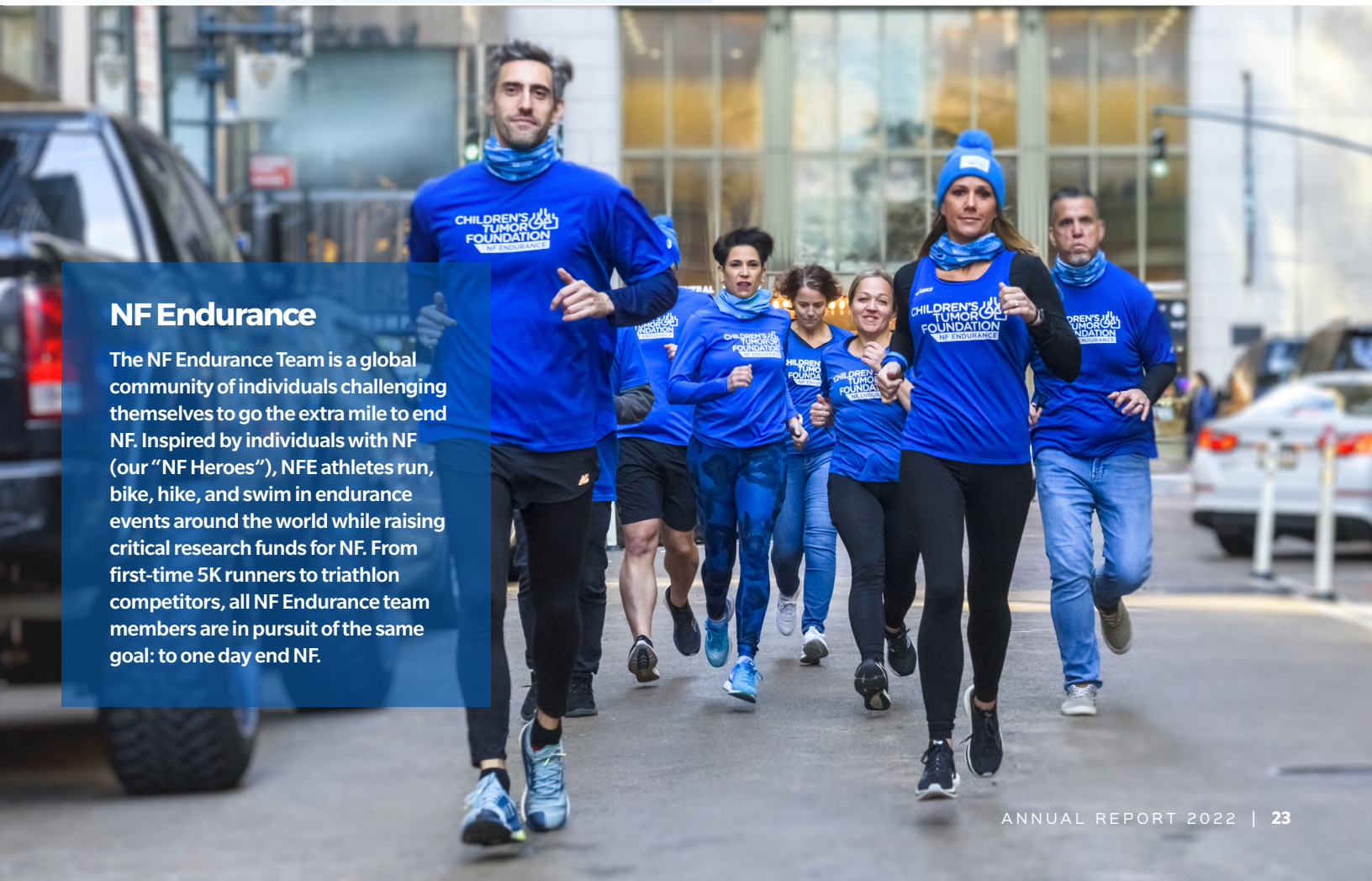
Cupid's Undie Run

Cupid's Undie Run is a mile(ish) fun run in which brightly adorned, underwear-clad team members run outdoors for NF awareness and to raise funds for NF research. The first Cupid's Undie Run took place in 2010 in Washington, D.C. What started as a unique "let's put hilarity into charity" twist on traditional charity walks/runs turned into a national phenomenon attracting wide attention and increased funding for NF. In the decade since, Cupid's Undie Run has raised millions of dollars for CTF-funded NF research.



NF Endurance

The NF Endurance Team is a global community of individuals challenging themselves to go the extra mile to end NF. Inspired by individuals with NF (our "NF Heroes"), NFE athletes run, bike, hike, and swim in endurance events around the world while raising critical research funds for NF. From first-time 5K runners to triathlon competitors, all NF Endurance team members are in pursuit of the same goal: to one day end NF.





Advocacy

From the earliest days of the Foundation, CTF staff and volunteers have advocated relentlessly for continual federal funding of NF research, with frequent and highly strategic visits to Capitol Hill and Member District Offices. The CTF Government Affairs Team continually expands the breadth of its advocacy and profile-building efforts with the guidance of outside counsel Squire Patton Boggs. We are also actively engaged with the Defense Health Research Consortium. This Consortium is composed of over 50 organizations dedicated to the preservation of annual funding levels for Congressionally Directed Medical Research Programs (CDMRP) within the Department of Defense.

Our voices are being heard! In 2022, because of continued strong advocacy work from the Children's Tumor Foundation and the NF community, bipartisan leadership in Washington, D.C., included \$25 million for NF research through the CDMRP for Fiscal Year 2023. This funding supports strategic research resulting in new discoveries and better outcomes for NF patients and families, and will allow us to further develop scientific data, break through barriers, and forge a pathway to end NF.

National Ambassador

Eddie Purtell



To quote the great Vince Lombardi, 'It's not whether you get knocked down, it's whether you get up.' Anyone with NF will have to deal with some difficult situations, so no matter how many times you get knocked down, always remember to get up and keep fighting.



— Eddie Purtell



Eddie Purtell was diagnosed with NF at just one-month-old. By the time Eddie understood what it meant to have NF, he had already undergone several surgeries and had his left eye removed due to an optic nerve glioma behind the eye.

Since then, he has had over 20 procedures but has never let that stop him from achieving all he has. His parents recall that Eddie has always been fearless through all that he's experienced. He lives his life to the fullest, determined to prove that having NF does not have to stop him from enjoying life. From skiing to soccer to lacrosse, Eddie enjoys being active and playing sports. His love of sports led him to start running marathons, and he has since run 14 half marathons, 11 of which have been to raise money for the Children's Tumor Foundation. With these efforts, he

has raised more than \$180,000 to find a cure for NF and is still going strong.

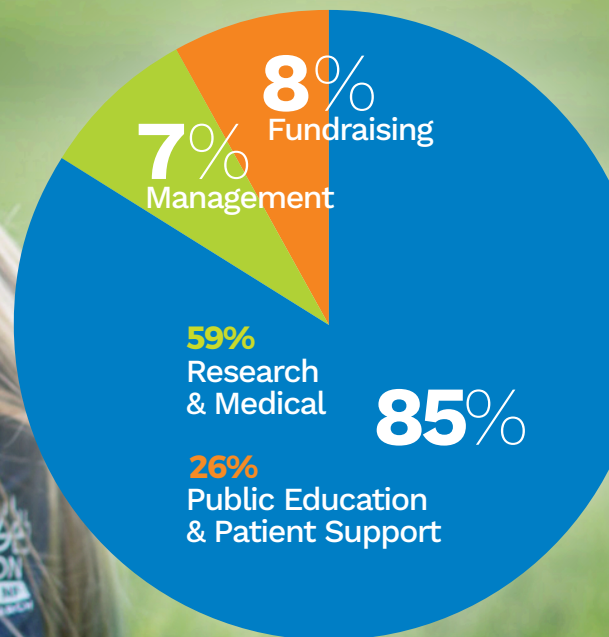
On top of that, he is also involved in the Children's Tumor Foundation's Volunteer Leadership Council and Junior Board.

Eddie was honored as the 2022 National Ambassador at the CTF National Gala, an award recognizing courage living with NF and personal efforts to further the Foundation's goals of research, public awareness, and patient support.

Despite the challenges of NF, Eddie has graduated college, is working a great job, has run multiple marathons, and serves as a role model for others. He hopes that his contributions will help make life easier for young kids with NF and that he can inspire those children.

Financial Summary

Expenses 2022



Operating support and revenue			
	2022	2021	2020
Contributions: individuals	\$8,136,795	\$8,071,808	\$5,943,186
Contributions: corporations and foundations	\$3,951,038	\$4,035,129	\$3,310,653
Bequests	\$306,122	\$185,000	\$580,538
Contributed goods	\$226,605	\$214,955	\$17,684
Other income	\$1,252,318	\$1,539,709	\$995,628
Government grants	\$299,188	\$1,077,026	\$691,400
Loss from disposition of property and equipment	(\$9,445)	0	0
CONTRIBUTIONS AND OTHER INCOME	\$14,162,621	\$15,123,627	\$11,539,089
	2022	2021	2020
Special event revenue	\$2,970,370	\$1,818,650	\$1,084,549
Less: direct benefits to donors	(\$222,968)	(\$40,000)	(0)
Special event revenue, net	\$2,747,402	\$1,778,650	\$1,084,549
TOTAL OPERATING & SUPPORT REVENUE	\$16,910,023	\$16,902,277	\$12,623,638

Operating expenses			
	2022	2021	2020
Program Services			
Research and medical	\$9,468,472	\$6,032,140	\$6,326,821
Public education and patient support	\$4,122,574	\$2,720,523	\$2,337,711
TOTAL PROGRAM SERVICES	\$13,591,046	\$8,752,663	\$8,664,532
Supporting Services			
Management and general	\$1,144,982	\$945,320	\$904,881
Fundraising	\$1,266,163	\$930,453	\$815,402
TOTAL SUPPORT SERVICES	\$2,411,145	\$1,875,773	\$1,720,283
TOTAL OPERATING EXPENSES	\$16,002,191	\$10,628,436	\$10,384,815

Change in Net Assets from Operations	\$907,832	\$6,273,841	\$2,238,823
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Other changes	2022	2021	2020
NON-OPERATING REVENUE	(651,650)	\$561,848	\$532,959

Change in Net Assets			
	2022	2021	2020
Net Assets, beginning of year	\$27,295,800	\$20,460,111	\$17,688,329
Net Assets, end of year	\$27,551,982	\$27,295,800	\$20,460,111

With Thanks

The Children's Tumor Foundation is grateful for the continued support of the many individuals, corporations, foundations, and communities who have joined us in the fight against NF. Thank you for your help in advancing the Foundation's mission.

Architects of Impact

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As a supporter and friend of the Children's Tumor Foundation, you have been vital in building us up from a grassroots group with just a few members into the leading organization we are today, fully committed to finding treatments for neurofibromatosis. Make no mistake about it — the progress that has been made in the fight against NF is because of people like you, who are working to improve the lives of those with NF. It is a legacy of which you can be proud.

Our vision is to end NF. We owe it to future generations of NF patients and families to see that vision become a reality. And as long as there is the Children's Tumor Foundation, there will always be an advocate fighting hard for the NF community.

By making a special legacy gift to the Children's Tumor Foundation, you will play an important role in ensuring that this work continues. Your planned gift is an investment in the long-term future of the organization, ensuring that the Children's Tumor Foundation will continue to lead the way in the fight to end NF.

The Children's Tumor Foundation NF Legacy Society consists of individuals who have taken the extra initiative to ensure the future of NF research by including the Children's Tumor Foundation in their estate plans.

To learn more about leaving a legacy and making a planned gift, please visit freewill.com/CTF, contact the Foundation at info@ctf.org, or call us directly at 1-800-323-7938.



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TOP TEN ADVANCEMENTS IN NF RESEARCH 2022



At the Children’s Tumor Foundation, we’re dedicated to bettering the lives of millions of people living with NF: all forms of neurofibromatosis and schwannomatosis. We want to end NF as fast as we can by connecting the unconnected, leading the way with a strategy that applies innovative and inventive approaches to scientific advancement and improved patient care.

This pioneering approach attracts the brightest minds in research and industry to NF, revamps systems to accelerate the pathway from discovery to treatment, and includes the voice of the patient every step of the way.

Read about the important things that happened in NF research in 2022, which are laying the groundwork for even greater scientific advancements in the near future.

Stay informed with NF research updates throughout the year at [ctf.org/news](https://www.ctf.org/news).

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