

Memphis, Tennessee

# Brain Waves

Neuroscience Institute

Winter 2016

### High-volume brain tumor program benefits patients

e Bonheur Children's Hospital has the highest brain tumor volumes when compared to a national database of other leading children's hospitals for first and second quarter 2015. This year, the neurosurgery team performed 183 brain tumor surgeries.

The brain tumor program is a partner-ship between
Le BonheurChildren's Hospital, Semmes
Murphey Clinic,
PC, University of
Tennessee Health
Science Center and
St. Jude Children's
Research Hospital.

"There are numerous papers showing that centers with higher volumes

have more reduced mortality and morbidity, complete resections and decreased length of stay. High-volume programs benefits patients with their large, experienced teams that collaborate on cases and the latest diagnostic tools," said Neuroscience Institute Co-Director and Chair of the Division of Neurosurgery Frederick Boop, MD.

Le Bonheurhas built a comprehensive program that includes the most advanced technology available, including intraoperative MRI, function MRIs, Magnetoencephalography (MEG) and Transcranial Magnetic Stimulation (TMS). Facilities include a six-bed

neurosurgical ICU, a 10-bed epilepsy monitoring unit and specialized neuroscience unit, all staffed with nurses who have specialized training.

The multidisciplinary team includes physicians representing radiology, neuropsychology, neuropathology and neuroendocrinology, plus specially trained nurses, rehabilitation

specialists, pharmacists, nutritionists and audiologists. Additionally, the weekly brain conference team includes pediatric neurosurgeons and neuroradiologists at Le Bonheur and neuro-oncologists and pediatric radiation oncologists at St. Jude.



Neuroscience Institute highlights: 2015

#### 2015 BY THE NUMBERS\*

6,240 Patient days

3,196 **EEGs** 

112 fMRIs

68 Magnetoencephalography (MEG) studies

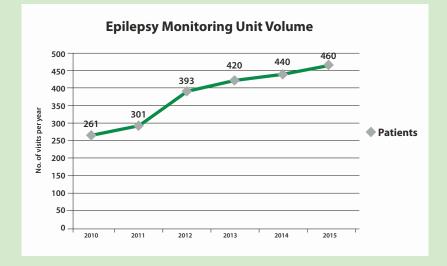
47 Transcranial Magnetic Stimulation (TMS) studies

12,454 Neurology clinic visits

183 Brain tumor cases

659 Neurosurgery cases

#### **Brain Tumor Volumes Compared to National Data Base** 150 140 130 120 110 Survival rate 100 90 Le Bonheur 80 70 60 50 40 30 20 10 2013 2014 \* Data is pulled from Children's Hospital Association's Pediatric Health Information System (PHIS) database



Referrals: 866-870-5570 www.lebonheur.org/neuroscience

A pediatric partner
with The University
of Tennessee Health
Science Center/College
of Medicine and
St. Jude Children's
Research Hospital



### Clinics cover variety of neurological conditions

### In our clinics, we treat children with:

- 15q Duplication Syndrome
- Autism
- Behavioral, attention deficit and hyperactive disorders
- Blood vessel lesions including brain aneurysm, brain and spinal arteriovenous malformation and Vein of Galen malformations
- · Brain tumors
- Cerebral palsy and related movement disorders
- · Craniofacial disorders
- Dravet Syndrome
- Developmental delays and neuromotor disorders, such as cerebral palsy
- Epilepsy
- Headaches
- · Head injuries
- Learning and language disorders
- Multiple Sclerosis
- · Muscular Dystrophy
- · Neuromuscular disease
- · Pial Synangiosis
- Seizures
- Sleep disorders
- Spinal disorders and tumors
- Spina bifida
- Stokes
- Tuberous Sclerosis Complex
- Tics, Tourette's Syndrome

### New stoke program provides clinic, emergency coverage

The Neuroscience Institute recently opened a comprehensive stroke program, which includes a multidisciplinary clinic and Emergency Department coverage. Le Bonheuris the only program in the area providing coordinated care for stroke patients.

Patients and families can see multiple specialists in one visit. The specialists discuss cases and work together to develop comprehensive plans of care. The team includes special-



Neurologist Paras Bhattarai, MD, evaluates patient Roni Gordon in the new stroke clinic.

ists from neurology, hematology, pharmacy, physical therapy, nutrition and social work.

In emergent situations, pediatric neurologists are available 24/7 to assess suspected pediatric stroke patients in Le Bonheur's Emergency Department. Expertise is also available via telemedicine.

The Neuroscience Institute has also joined the International Pediatric Stroke Study.

# Muscular dystrophy clinic brings resources to one place for families

Le Bonheur's Muscular Dystrophy Association (MDA) Clinic provides comprehensive care from a multidisciplinary team in one appointment. The team includes pediatric specialist in neurology, pulmonology, cardiol-

ogy and orthopedics. Patients also have access to nutrition/ dietitian, occupational, physical and speech therapy, palliative care and social work.



Le Bonheur's Muscular Dystrophy Association (MDA) Clinic provides comprehensive care from a multidisciplinary team, including Neurologist Namrata Shah, MD, in one appointment.

The "one-stop shop" treats children with 43 distinct diagnoses – the most common being Duchenne Muscular Dystrophy, Becker's Muscular Dystrophy and spinal muscular atrophy.

The clinic is one of 28 clinics nationwide participating in the pilot MDA registry, designed to improve the care at MDA clinics across the country.

The physician researchers are working on several studies to improve the treatment of children with muscular dystrophy including Exon skipping for Duchenne Muscular Dystrophy and participating in the Cooperative International Neuromuscular Research Group.

#### Le Bonheurforms CP Clinic

Le Bonheur's Cerebral Palsy (CP) Clinic is the only multidisciplinary center in the region that provides comprehensive care for children with CP – a group of disorders of movement and posture development. The diagnosis affects two to three in every 1,000 children.



Neurologist Swati Karmarkar, MD, works with patients in the Cerebral Palsy (CP) Clinic – the only multidisciplinary center in the region for children with CP.

In the clinic, children have access to a physiatrist, physical therapists, orthopedic surgeons and orthotic and seating/mobility experts. In

addition, specialists collaborate closely with other providers involved in the child's care, such as the child's pediatrician or other therapists, to maximize the child's potential. The clinic also offers home visits to assess any need for specific equipment or accessibility concerns at home.

"Our goal is to offer individualized treatment for each child with CP designed to meet his or her specific needs and goals," said Pediatric Neurologist Swati Karmarkar, MD.

Individualized treatment through the clinic could include medication for tone reduction, botulinum toxin injections, evaluations for surgical interventions physical therapy including bracing and equipment and treatment for associated conditions like seizures and sleep disturbances.



# Out-of-town patients find free lodging at FedExFamilyHouse

Last year, patients traveled to Le Bonheur's Neuroscience Institute in Memphis, Tenn. from 32 states. Out-of-town families who come to Le Bonheur treatment may stay at no cost at FedExFamilyHouse.

Located across the street from Le Bonheur, FedExFamilyHouse has 24-private suites, multiple dining facilities and recreational spaces and a small business center. Family-friendly amenities include outdoor spaces, living rooms, play area, teen lounge, large kitchens, dining space and laundry facilities.

For more information, visit www.fedexfamilyhouse.org.

### Study: Tailoring treatments for high-grade gliomas

new study aims to improve outcomes for patients with high-grade gliomas. Researchers are using mouse models to test the combination of surgery, chemotherapy and radiotherapy treatment for Glioblastoma multiforme (GBM), a Grade IV primary astrocytic tumor.

"Ultimately, we would like to develop a system where we could grow each individual patient's brain tumor in this model and test several different therapies to see which would work best. It's a sort of tailoring treatments based on each patient's specific tumor biology," said Neurosurgical Chief Resident Mike Decuypere, MD, PhD.

The research is led by the University of Tennessee Health Science Center departments of Neurosurgery and Pathology and the UTHSC Center for Cancer Research and funded by UT-West Cancer Center.

GBM has a poor overall prognosis, significant morbidity and limited therapeutic options. Death typically occurs within 15 months of diagnosis, even after maximal surgical and medical treatment.

The current treatment paradigm of surgical resection, chemotherapy and radiation has



Researchers are using murine models to test the combination of surgery, chemotherapy and radiotherapy treatment for improving outcomes for patients with Glioblastoma multiforme (GBM), a Grade IV primary astrocytic tumor.

poor overall results. This is due, in part, to the lack of accurate pre-clinical animal models that mimic the true disease state and are able to undergo the gamut of treatment options.

This project will compare the relative efficacies of various treatment protocols for human glioblastoma, orthotopically xenografted in a murine model. Survival intracranial resection surgery will be used in conjunction with several post-operative chemotherapy protocols using irinotecan and bevacizumab in hopes of generating data useful to human clinical trials.

# RESEARCH AIMS TO IMPROVE SLEEP QUALITY FOR PATIENTS WITH NEUROLOGICAL CONDITIONS

umans spend a third of their lives sleeping. For children with neurological disorders, achieving good sleep quality is a challenge. Research underway at Le BonheurChildren's Sleep Center aims to better understand sleep challenges and find innovative solutions.

The Sleep Center at Le Bonheuris the only pediatric center in the region accredited by the American Association of Sleep Medicine. The eight-bed unit is open seven nights a week and includes a separate but attached room for the patient's family member. The sleep center provides referring physicians with a full report within 10 days of the study.

"It's more challenging to achieve good quality sleep in patients with neurological disorders because of the disrupted neurotransmitters due to the underlying brain injury. However, studies have shown that with regular cognitive behavioral therapy for insomnia, and with the help of some medications, a decent good quality sleep can be achieved," said Ehab Dayyat, MD, Sleep Neurologist at Le BonheurChildren's Hospital.

#### **Projects underway include:**

A multi-center global clinical trial to study the efficacy and safety of Xyrem in controlling cataplexy and excessive daytime sleepiness in patients with narcolepsy/cataplexy. This study includes 41 centers in the U.S. and eight centers in Europe.

Identifying a simple way using a questionnaire of sixquestions to screen for the presence of sleep-disordered breathing, after comparing them with objective findings from sleep studies.

Studying sleep changes in epilepsy patients with Vagal Nerve Stimulator.

Evaluating changes in sleep pattern in patients with dup15q syndrome using an actigraphic measurement and sleep studies.

Establishing a sleep center data registry with the Children's Foundation Research Institute at Le Bonheur Children's Hospital.

Educating pediatrics and medical/pediatrics residents and general pediatricians about the presence of sleep-disordered breathing in patients with Down Syndrome and when to screen for it.

Studying the most specific subjective questions to evaluate for the presence of excessive daytime sleepiness.

Collaborating with the University of Chicago to identify specific microbiome — inflammatory markers in the stool, to screen for the presence of sleep-disordered breathing.

Collaborating with the University of Tennessee Health Science Center (UTHSC) College of Nursing to study Continuous Positive Airway Pressure (CPAP) treatment in cognitive function.

Teaming up with the Pediatric Obesity program and Endocrinology team at Le Bonheurand UTHSC to measure the effect of sleep duration in obesity management.

Partnering with St. Jude Children's Research Hospital to study the effect of excessive daytime sleepiness on cognition in patients with craniopharyngioma.

# Study: Brain connectivity with infantile spasms

Researchers are studying why some children experience infantile spasms and how a treatment therapy affects brain connectivity. Infants and young children with infantile spasms (IS) can experience tens to hundreds of seizures a day. Few treatment options are available, and the most common is adrenocorticotropic hormone (ACTH). The sooner the spasms are diagnosed and treated the better the chances the spasms will stop. When there is a delay in diagnosis or treatment, affected children are more likely to develop other types of seizures that are difficult to control and experience developmental delay.

"This study may allow us to understand why some children experience infantile spasms and what the process behind this type of seizure is, as well as better understanding how adrenocorticotropic hormone (ACTH)works to correct this process," said Sarah Weatherspoon, MD.

The researchers will use magnetoencephalogram (MEG) to study functional brain connectivity and brain magnetic resonance imaging (MRI) to study structural brain connectivity in these children before and after ACTH treatment. Researchers will assess if there is abnormal brain connectivity on MEG and MRI before starting treatment. The team will also evaluate whether those children whose seizures stop with ACTH will show normalized brain connectivity after finishing a course of treatment.

The study is funded by Mallinckrodt Pharmaceuticals.

#### Study: Respiratory modulation of brain activity

ontrolled deep, slow breathing is a common strategy in anger management, meditation and general stress relief? How does respiration modify cognitive and emotional states? A new study from researchers at Le BonheurChildren's Hospital, the University of Tennessee Health Science Center and University of Memphis provides the first possible explanation to for howshow breathing influences brain function. The study, headed by Detlef Heck, Ph.D., Associate Professor in the Department of Anatomy and Neurobiology, UTHSC, which bridges work with animal and human subjectsclinical and basic research and, was presented at the National International Society for Neuroscience Meeting in Chicago in October. The team first published first results in from studies in mice in *Nature Communications* in 2014.

This Results from this current translational study project presents a radically new view of respiration – breathing not only provides oxygen, but also directly influences cognitive brain function.

"The brain generates oscillatory activity in a frequency range between 30 to 100 cycles per second whenever it performs cognitive tasks, such as thinking of a word associated with a picture, trying to remember something where we left our keys or simply focusing on a specific task. We found that The the strength of those oscillations increases and decreases with every breath. That is why we believe that respiration affects cognitive function. We are designing experiments to further test this hypothesis," said Detlef HeckDetlef Heck, PhD, associate professor in the Department of Anatomy and Neurobiology in the College of Medicine, UTHSC.

The Clinical Neurosciences and Neurology departments at Le Bonheurand UTHSC participated in this study by conducting series of experiments using magnetoencephalography (MEG) and electrodes placed directly onto the surface of the cortex. The team continues this line of study and will be publishing the research shortly.

#### IN BRIEF

# Oral appliance for Tourette syndrome

The Neuroscience Institute and the University of Tennesee Health Science Center College of Dentistry are part of a multi-site study to test the effectiveness of a new oral appliance for patients with Tourette syndrome.

The appliance, much like a mouth guard, is designed to reposition the lower jaw to lessen the severity and frequency of motor and vocal tics associated with Tourette's.

The study is funded by TicTocStop, Inc., an organization founded by Craig Carton, who is also a host of the "Boomer and Carton" show on sports radio station WFAN in New York. Carton and two of his children have Tourette syndrome.

# Nurse practitioners share at CNS meeting

Four Le Bonheurnurse practitioners shared research at the Child Neurology Society Nursing Meeting in October.

- "Febrile seizures, to treat or not to treat...that is the question!" – Lauren Siebrase, MSN, FNP-C
- "Evaluation and treatment management of childhood epilepsy syndromes" – Lai Brooks, DNP, FNP-BC, and Siebrase
- "Glut1 transporter deficiency presenting as alternating hemiplegia of childhood" – Kelli Ward, DESIGNATION, and Davi Ledet, BSN, MSN, MBA

### Neurosurgeons present at AANS/CNS meeting

Le Bonheurand Semmes Murphey neurosurgeons presented research on brain stem gliomas and pineoblastomas at the American Association of Neurological Surgeons/Congress of Neurological Surgeons Section on pediatric neurological surgery meeting in December. Paul Klimo Jr., MD, also served as a panelist for the clinical symposia on Hydrocephalus: Shunting vs. ETV.

# Spring Neurology Symposium set for April

Save the date for the 10th Annual Mac Armour Pediatric Neurology Symposium April 29-30 at the Big Cypress Hotel in Memphis, Tenn. Doug Nordli, MD, will be guest speaker. He serves as professor of neurology and pediatrics at Ann & Robert Lurie Children's Hospital and Northwestern University. For more information or to register, visit www.methodistmd.org or call

# Neuroradiologist's stories included in new book

Le BonheurDirector of Neuroradiology Asim Choudhri, MD, is one of 20 radiologists who shared case studies in a new book, "Radiologists at Work: Saving Lives with the Lights Off" by Carolyn Jourdan.



Asim Choudhri, MD



Brain Waves is a quarterly publication of the Neuroscience Institute at Le Bonheur Children's Hospital. The institute is a nationally recognized center for evaluation and treatment of nervous system disorders in children and adolescents, ranging from birth defects and learning and behavioral disorders to brain tumors, epilepsy and traumatic injuries.

Frederick A. Boop, MD Andrew C. Papanicolaou, PhD James W. Wheless, MD

Abbas Babajani-Feremi, PhD Paras Bhattarai, MD Elena Caron, MD Asim F. Choudhri, MD Ehab Dayyat, MD Stephanie Einhaus, MD Lucas Elijovich, MD Stephen Fulton, MD Masanori Igarashi, MD Swati Karmarkar, MD Paul Klimo, MD

Diana Lebron, MD Amy McGregor, MD Kathryn McVicar, MD Robin Morgan, MD Basan Mudigoudar, MD Michael S. Muhlbauer, MD Shalini Narayana, MBBS, PhD Brian Potter, Psy D Roozbeh Rezaie, PhD Namrata Shah, MD Nicole Shay, PhD Adeel Siddiqui, MD Sarah Weatherspoon, MD

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