StrokeNet Enrollment Update

ARCADIA	494/1100	CREST 2	1709/2480
Sleep SMART	**300**/3062	CREST H	153/500
TRANSPORT2	15/129	MOST	74/1200
I-ACQUIRE	26/240	ARCADIA-CSI	69/500
ASPIRE	1/700	SATURN	13/1456

StrokeNet Trial Updates

From the CREST-2 Clinical Coordinating Center:

CREST-2 Enrollment	
CEA	897
CAS	812
Total	⭐ 1709 ⭐

WE DID IT !

CREST-2 has reached two milestones: **1700** patients randomized and **15** in August! Congratulations to Dr. Matthew Starr and team at UPMC Presbyterian for randomizing the 1700th patient! These are two incredible achievements for CREST-2 — we appreciate everyone who has contributed to each of these goals! We will need to keep up this momentum over the coming months to ensure our success. This includes **15** randomizations per month until the end of October 2020 and **25** randomizations per month starting November 2020. So far, we have had 8 randomizations in September – which averages out to more than one for each working day this month! Let's keep up the great work.

We now have 98 sites re-initiated to resume enrollment, including some of the top CREST-2 enrolling sites. If your site is ready to re-open for enrollment, please notify your site manager (if you have not done so already), so that we can make note of your site's status in CREST-2.

Site Startup and Enrollment Updates: sites: 55; enrollments: 153/350

Enrollment has picked up again with a total to 153, **44%** of our target. We are looking forward to reopening of the remaining CREST-H sites and the inclusion of **8** additional sites in the onboarding pipeline. CREST-2 is 2/3 of the way to their target, and CREST-H needs to continue the pace to meet our target as well. Keep an eye out for asymptomatic carotid patients!

In the setting of COVID, minimizing exposure of patients is critical. We currently recommend that CREST-H be discussed in the context of consenting for CREST-2, at the time other imaging is being mentioned. In order to save the patient an extra trip to your medical center, you can have the patient verbally agree to the perfusion scanning, and then have them sign the CREST-H consent when they arrive at the scanner. Please feel free to contact Jaya, Kevin, or one of the CREST-H PIs with any questions about this. Remember, either MR or CT perfusion is acceptable as long as you have submitted and have approved a test image.

Interested in becoming a CREST-H site? Contact Randy Marshall rsm2@columbia.edu, Ron Lazar rlazar@uabmc.edu, Jaya Vijayan vijayan.jaya@mayo.edu, or Kevin Slane KJS4@columbia.edu with any questions.

Congratulations!

8 sites have been approved to re-open (COVID-19)
- ❖ Ann Arbor (**3 enrolled**)
- ❖ Boston (**8 enrolled**)
- ❖ Cincinnati (**3 enrolled**)
- ❖ Columbus (**1 enrolled**)
- ❖ Houston,
- ❖ La Jolla (**5 enrolled**)
- ❖ New Haven (**2 enrolled**)
- ❖ Roanoke (**4 enrolled**)

Total enrolled as of 9/9/2020 = **26**

Save the dates: Upcoming PI/Co-I/Coordinator Webinar:

- ❖ Thursday 08 October (12-1pm ET)
- ❖ Thursday 12 November (12-1pm ET)
- ❖ Thursday 17 December (12-1pm ET)

Thank you for your participation in Sleep SMART. We truly appreciate all your hard work, especially during this difficult time.

As of September 9, 2020, 896 subjects have been enrolled and 299 subjects have been randomized.

Congratulations to Tampa General Hospital (Scott Burgin, MD, Andrea Bozeman and Corbin Hilker) for randomizing their first subject in September and Sarasota Memorial Hospital (Mauricio Concha, MD and Jeanette Wilson) for randomizing five subjects in the last two weeks including three subjects in one day!

ADHERENCE to CPAP: As in all trials that involve CPAP, we are making every effort to maximize the probability that subjects will use their CPAP nightly, all night, comfortably, and effectively from the day of randomization to the CPAP arm to the last night of participation in Sleep SMART. Since re-opening to enrollment, new subjects clearly have done best in their adherence to CPAP after successful "warm transitions" by local members of the investigator team to the subject's new contact at FusionHealth. This warm transition means asking for permission to plug the FusionHealth phone number into the subject's cell phone, and then encouraging them or helping them to phone the Fusion care team around the day before discharge from the hospital. We see with increasing clarity that when this warm transition does not occur, good adherence is much harder to establish. Please be sure that every subject is offered a "warm transition" before leaving the hospital. This could make a key difference to the success of Sleep SMART.

CONTINUING REVIEW: The annual continuing review (CR) forms were distributed to sites by email on August 13th and were expected to be returned to sicklejb@ucmail.uc.edu by September 4th. If you have not returned your forms, please take time to carefully read the email instructions and return the forms as soon as possible.

INFORMED CONSENT IMPLEMENTATION FORM: All sites need to let us know what remote consent options you want to be able to exercise. Please return your site specific Implementation form to sicklejb@ucmail.uc.edu. The Implementation form will be sent to the CIRB for review and acknowledgement.

RE-OPENING: In WebDCU, please update the COVID Impact Assessment survey to reflect the current status of research at your institution and, if you are ready to re-start enrollments, to indicate confirmation (in 'general comments') of the statements in the memo dated May 15, 2020. Email Kayla or Joelle to let them know when your site is ready to be re-released to enroll. Pending NDMC queries may delay our ability to release you, so please catch up on these. **You will not be released to enroll until you receive official notice from WebDCU.** Since our re-opening, 59 sites have been released to enroll and conduct in-person activities and 46 subjects have been randomized. If your site is not ready to enroll, but is able to complete in-person 3- and 6- month assessments, please contact Kayla or Joelle to seek permission for these limited in-person activities.

Sites not previously released to enroll:

If your site was not previously released to enroll, please continue to work toward site activation:

- CIRB submission
- Finalize the Clinical Trial Agreement and FusionHealth DUA and Consignment agreement
- Upload site and people documents to WebDCU
- Complete online trainings (https://webdcu.musc.edu/campus/)
- Complete readiness call (must have all contracts completed before this can be done)
- Please review the site to-do checklist found in the WebDCU toolbox

Save the date-Our next PI and coordinator webinar will be Monday, September 21, 2020 from 12-1 EST. Please note the time change. The KOEO platform will be reviewed.

Despite these challenging times 110 sites have completed their re-start or started new since the COVID pause. Overall, 138 of the 149 sites ever released to enroll have consented at least one subject and 105 sites have randomized at least 1 participant. The ARCADIA teams have shown commitment and perseverance despite the many challenges that have come our way – thank you!

We are currently at 1867 subjects enrolled/consented, an increase of 37 in the last two weeks, thanks to 29 sites, with many sites enrolling more than once during this period – great job!

We have randomized 494 participants, an increase of 10 in less than two weeks. A special thanks to the nine teams that provided the last two weeks' randomizations at: Barnes Jewish Hospital, St. Louis, MO; United Hospital, St. Paul, MN; Vanderbilt University Hospital, Nashville, TN; UPMC Presbyterian Hospital, Pittsburgh, PA; Central DuPage Hospital, Winfield, IL; University of Michigan University Hospital, Ann Arbor, MI; University of Mississippi Medical Center, Jackson, MS; Hartford Hospital, Hartford, CT and OU Medical Center, Oklahoma City, OK.

A special thanks to the Beth Israel Deaconess Medical Center, Boston, MA team for randomizing two subjects in one day - we wanted you to know how much we appreciate your efforts!

We would also like to welcome the Akron General Medical Center team in Ohio as they have just been released to enroll this week!

We hope to continue to re-start sites as per the ARCADIA re-start plan for the 25 sites still pending re-release to enroll. We understand many of your sites cannot yet re-start, but those that can re-start enrollment, even if only remotely, please reach out to Rebeca (ra2356@cumc.columbia.edu) to review any pending items needed for your re-release. PLEASE do not start enrolling new subjects until your site has been re-released to enroll.

We understand that the enrollment challenges will change from site to site during this pandemic, but this is also a good time to assess performance from initial startup and see if there is anything that can be done to improve the enrollment at sites that have not been able to consent or randomize for a while prior to COVID. We will be reaching out to you and the RCCs about this in the coming weeks.

CALM lab has not moved yet, so for now please continue to send the blood sample shipments to the same address on the Lab Manual (p. 11) so that they are not missed. We will send out an email when the move has been completed.

We thank you for your continued support and effort during these challenging times, but together we can move forward in ARCADIA.

Webinar: Our next PI and coordinator webinar will be September 22nd at 2 PM Eastern--save the date! We're asking that at least 1 person from each site attend the monthly webinar and pass the information along to their team. Also, please send suggestions for topics you'd like discussed.

If you miss out on a webinar, you will soon be able to find the slides on the StrokeNet website https://www.nihstrokenet.org/intranet/minutes/trial-webinars

Thank you to all the sites for your hard work in enrolling subjects into ARCADIA-CSI!

The study team would like to welcome **Methodist University Hospital to ARCADIA-CSI!**

Our top 3 enrolling sites are:

8 PATIENTS ENROLLED
UNIVERSITY OF IOWA
ENRIQUE LEIRA & HEENA OLALDE

7 PATIENTS ENROLLLED
UNIVERSITY OF CINCINNATI
POOJA KHATRI & JENNIFER POWERS

6 PATIENTS ENROLLED
GREENVILLE HOSPITAL
PAULO ZORTEA & VICTORIA HOLT

We need your help enrolling patients into the ARCADIA-CSI substudy! We have created form 515 to make screening a breeze. When a patient is randomized in ARCADIA, form 515 is posted in the patient's ARCADIA binder in WebDCU. This form serves as the ARCADIA-CSI screening form and as a reminder to reach out to the patient to ask if (s)he would like to join ARCADIA-CSI.

Please complete the Covid19 Impact Assessment in WebDCU as soon as possible.

Please contact Tashia Harris, herndotl@ucmail.uc.edu or Stephanie Kemp, skemp@standford.edu with any questions.

- Enrollment into the ASPIRE study was paused on July 8 due to an issue with study drug stability. We expect sites will start to receive new drug by the end of September - October. Sites who have consented patients will be restocked first.
- Documents for the ASPIRE Continuing Review were sent to sites this week. Site must return the Continuing Review form, Financial Conflict of Interest form(s), and (if applicable) redacted informed consent documents to ASPIRE@yale.edu by October 1, 2020.
- All sites must complete a Remote and Electronic Informed Consent Implementation form to be submitted to the cIRB via the ASPIRE NCC Project Manager. This form is posted in WebDCU > ASPIRE Toolbox.
- Please remember to update the COVID-19 Impact Assessment v2 in WebDCU to alert us to any changes in local restrictions on screening or enrollment.
- For sites still pending activation, please:
    - Upload/waive pending regulatory documents.
    - Enter addresses for study drug and lab kit shipments.
    - If CTA and cIRB approval in place, schedule readiness call.
- The ASPIRE/SATURN Investigator Meeting in New York has been postponed indefinitely. In lieu of the meeting, protocol training is posted on the WebDCU Training Campus https://webdcu.musc.edu/campus/
- ASPIRE next Webinar is September 23, 2020 3:00p-4:00p EDT https://nihstrokenet.adobeconnect.com/trials/ To take part in the conversation dial 1 (877) 621-0220 Pass Code: 745694

SATURN has 61 sites open for enrollment and has randomized 13 patients!

SATURN held the monthly PI/Coordinator Webinar 8/27/2020, if you were unable to attend please find the recording here: https://www.nihstrokenet.org/saturn-trial/home There were good discussions about Inclusion/Exclusion criteria as well as tips from sites who have enrolled.

Please continue to update the COVID Impact Assessment with information on anticipated timelines for resuming research activities. Please reach out to the study team when your site is able to resume research activities. We will assist you with opening for enrollment when local restrictions allow.

Sites that have not yet been released to enroll should complete the COVID Impact Assessment once they have been activated.

MGH is shipping lab kits out to sites who have completed a readiness call. Please remember to confirm receipt of the lab kits in WebDCU upon receipt.

SATURN is actively looking to add sites; please reach out for further details.

If you have any questions regarding SATURN and reopening enrollment at your site please contact:
Kimberlee Bernstein NCC Project Manager gammk@ucmail.uc.edu
Sarah Marchina Prime Project Manager (BIDMC) smarchin@bidmc.harvard.edu

MOST Enrollment Update:
Total randomizations: **74**
Randomizations between 01Sep2020 and 08Sep2020: **2**
Sites released to enroll with at least one patient consented: **27**
There are now **69** sites that have been released to enroll, **56** of which are actively recruiting!
- Congratulations to the following sites that have randomized 1 subject in the past week!
    - Wake Forest Baptist Medical Center – Dr. Bushnell and Karin Haski **(5 total subjects!)**
    - UVA Medical Center – Dr. Chapman and Sonya Gunter **(1st subject!)**

- Congratulations to Santa Barbara Cottage Hospital who has been released to receive study drug and to Stanford University Medical Center who has been re-released to enroll!

In order to reach our enrollment, target each site must enroll **one patient every 3 months**. Thank you for keeping MOST a priority!

Mark your calendars for the **MOST Trial Investigator Call** on 21Sep2020 at 2:00 PM ET. All primary and secondary study pharmacists have been invited to join the call to contribute to the discussion regarding pharmacy processes.

Please remember to update your **Delegation of Authority logs** to reflect current and active team members. The email distribution list for study-wide correspondence is generated through site DOA lists.

We are pleased to announce that Melissa Hoffman has joined the NIH StrokeNet National Coordinating Center team as the MOST Clinical Research Project Manager. Melissa joins the team with 18 years of clinical research experience.

The PI Hotline is available 24/7 for any questions: **1-833-229-MOST.**

Our next PI and Coordinator call will be on **Monday, 9/28/2020 at 11am ET**. If anyone has topics they would like to discuss during the call, please send them to Kristina Balderson (Kristina.balderson@duke.edu) or Julia Gonzalez, (Julia.jackson@duke.edu).

As sites are working to reopen to enrollment, please take inventory of your supplies and contact either Julia Gonzalez or Kristina Balderson for restock. When reaching out, please indicate the number of supplies needed, contact, and shipping location with confirmation that someone will be able to receive these shipments. Home addresses are approved for receiving shipments during this time of remote working or limited time at the site. For other questions or concerns, please contact Julia or Kristina.

Please make sure you have completed your BlueCloud recertification training by logging into the website: http://duke-transport2.trainingcampus.net. Individuals who have expired or about to expire training have been sent reminder emails. **Note this is for the scoring of the Fugl Meyer, not the video training.** Annual renewal is now being tracked in WebDCU, which will notify coordinators directly when any of their staff have outdated training.

Seven sites have reopened to enrollment! Congratulations to Baystate, Duke, Medstar, Kentucky, UAB, USC, and Moss for being re-open to enrollment amidst the COVID-19 restrictions. Cleveland VA is under IRB review and UPMC is finalizing study start-up activities. We are hopeful that our remaining sites will be reopened to enrollment soon!! There are 15 subjects randomized in the trial, and seven have completed the study.

Thank you for your continued effort and flexibility during this time!

FREE NIH Educational Opportunities

Ethical and Regulatory Aspects of Clinical Research - 7 Week Course

Register for the course Ethical and Regulatory Aspects of Clinical Research. This 7-week course covers essential topics in the ethics of clinical research and is designed for researchers and research teams, clinical staff, IRB members, and others interested in the ethics of clinical research. Presentations are given by experts from both within and outside the NIH.

The course runs from September 23, 2020 - November 4, 2020 and will be broadcast live. The course meets Wednesday mornings from 8:30 -11:30 am Eastern. This year, the course will be delivered on-line. In addition to lectures, we will host conversations with experts and engage in case discussions.

Information about the course and how to register as well as the schedule and syllabus are available at https://bioethics.nih.gov/courses/ethical-regulatory-aspects.shtml For Nursing CEU* or Core Curriculum (for NIH employees only) credits you must be registered for the course and complete course requirements.

This course is available to anyone via live video-cast live at http://videocast.nih.gov. **There is no charge.** If you want access to additional course materials and receive a Certificate of Completion for the course you must register. We will be using CANVAS to coordinate course activities. Once you register you will receive an invitation to join the course site.

For best performance, CANVAS should be used on the current or first previous major release of Chrome, Firefox, Edge, or Safari. Because it's built using web standards, Canvas runs on Windows, Mac, Linux, iOS, Android, or any other device with a modern web browser.

If your institution or group has participated from a satellite location in the past we are not organizing proctored groups. Each individual from your institution will register directly. For more information click https://bioethics.nih.gov/courses/ethical-regulatory-aspects.shtml.

The course textbook is "Ethical and Regulatory Aspects of Clinical Research: Readings and Commentaries (JHU Press, ISBN 9780801878138)." You can order the book on-line from the FAES Bookstore @ NIH https://faes.org/content/faes-bookstore-nih or other on-line purveyor of books. Additional readings will be recommended.

Individuals with disabilities who need sign language interpreters and/or reasonable accommodation should contact 301-496-2429 or bioethics-inquiries@mail.nih.gov. Federal TTY Relay number is 1-800/877-8339. Requests should be made at least 5 days in advance of the event.

-Nurses will receive 22 contact hours of CEU credits. Credits are provided through the Maryland Nurses Association, an accredited approver of continuing nursing education by the American Nurses Credentialing Center's Commission on Accreditation.

Successful completion of this course is required for the Clinical Research Curriculum Certificate. For more information, see http://intranet.cc.nih.gov/clinicalresearchtraining/curriculumcert.

2020-2021 NIH Introduction to the Principles and Practice

of Clinical Research (IPPCR) course

Interested in expanding your clinical research knowledge base? Registration for the 2020-2021 NIH Introduction to the Principles and Practice of Clinical Research (IPPCR) course is now open.

This *free*, self-paced, online course runs from September 1, 2020 to August 1, 2021. Graduate students, post-doctoral scholars, medical, dental, and pharmacy students, scientists, nurses, and other health professionals are encouraged to enroll now.

The IPPCR course is a lecture series from thought-leaders around the world covering: study designs; measurement and statistics; ethical, legal, monitoring, and regulatory considerations; preparing and implementing clinical studies; communicating research findings and other topics.

To register, please visit the IPPCR website at https://ocr.od.nih.gov/courses/ippcr.html. If you have any questions, please contact the course coordinator at ippcr2@mail.nih.gov.

2020-2021 NIH Principles of Clinical Pharmacology (PCP) Course

Registration is now open for the 2020-2021 NIH Principles of Clinical Pharmacology (PCP) Course.

The PCP course is a *free* online lecture series covering the fundamentals of clinical pharmacology as a translational scientific discipline focused on rational drug development and utilization in therapeutics.

Topics covered in the course include pharmacokinetics, drug therapy in special populations, drug discovery and development, and pharmacogenomics.
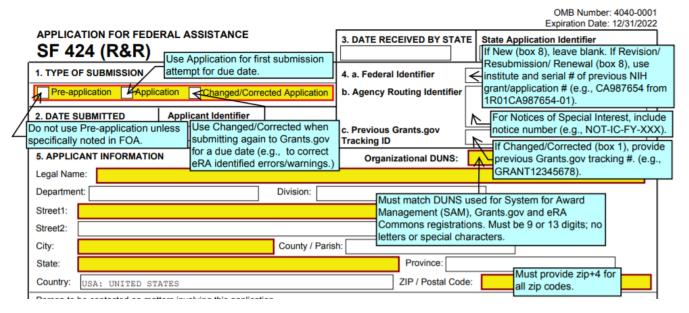
The course is free, self-paced, and entirely online through the PCP website: https://ocr.od.nih.gov/courses/principles-clinical-pharmacology.html.

Registration is now open and course materials are available through August 1, 2021. A certificate of completion is awarded to participants who achieve a passing score on the final exam.

The course will be of interest to graduate students, post-doctoral scholars, medical and pharmacy students, scientists, and health professionals interested in expanding their pharmacology knowledge base. For additional information on the course, please visit the website above or contact the course coordinator at odpcp@mail.nih.gov. Please feel free to share the course information with any internal or external colleagues that you feel may benefit from the course.

New Annotated Form Sets: Forms F-Series

For those who may be preparing an NIH grant application, keep these annotated form sets handy for key tips on filling out each section. These documents are a great visual resource for understanding many of the business rule checks we will run against your submitted application. Of course, you MUST follow the instructions in your funding opportunity announcement (FOA) and application guide, but these documents are helpful for those times when you just need a quick reference.

Please visit https://grants.nih.gov/grants/ElectronicReceipt/files/Annotated_Forms_General_FORMSF.pdf for more information.

Coordinator Webinar

Coordinator Webinars are a requirement for the NIH StrokeNet RCC Coordinators/Managers however, all are welcome to attend

Wednesday, 23-September, 1:30pm ET

Topic: Roundtable Discussion/Trial Project Manager Updates
Moderator: Dave Haney, Case Western Reserve University

To join the meeting: https://nihstrokenet.adobeconnect.com/coordinator/
To take part in the conversation you MUST dial in. 1 (877) 621-0220 Passcode: 434578

Steering Committee Call

Steering Committee Calls are a requirement for all NIH StrokeNet RCCs; Please invite satellite sites to attend

The next Steering Committee call is the Fall Network Webinar, scheduled for **Wednesday, 14-October, 2020, 12pm – 3pm ET.** Calendar invitations have been sent.

To join the meeting please log in to the webinar and also dial in in order to speak:

https://nihstrokenet.adobeconnect.com/network/

1-877-621-0220 – Passcode 631088 - Please enter as a guest and then your first and last name.

Grand Rounds

Grand Rounds are a requirement for the NIH StrokeNet Trainees, however all are welcome to participate.

Thursday, September 24, 2020
4:00 pm ET

Topic: Management of Malignant MCA Stroke

Presenter: **Kevin Sheth, MD,** Yale University

Moderator: Cemal Sozener, MD, University of Michigan

To join the meeting: *https://nihstrokenet.adobeconnect.com/grandrounds/.* Please enter as a guest, then your email address or your first and last name.

To take part in the conversation you MUST dial-in. (877) 621-0220 Passcode: 190825

Professional Development Webinar

Professional Development Webinars are a requirement for the NIH StrokeNet Trainees
Tuesday, September 29, 2020
12:00 pm ET

Presenter: **Dan Woo, MD,** University of Cincinnati

Presentation: Grant Writing

Moderator: Randy Marshall, MD, Columbia University

To join the meeting: *https://nihstrokenet.adobeconnect.com/pdw/*. Please enter as a guest, then your email address or your first and last name.

To take part in the conversation you <u>MUST</u> dial-in. (877) 621-0220 Passcode: 190825

NIH StrokeNet Employment Opportunities

For more information about StrokeNet employment opportunities, please visit
http://nihstrokenet.org/education/employment-opportunities

Please share your satellites and study teams!
Contact: Jamey Frasure, PhD, RN, Administrative Director · frasurjs@ucmail.uc.edu https://www.nihstrokenet.org