



Welcome and Updates

Pooja Khatri, MD, MSc

NIH StrokeNET National Coordinating Center

University of Cincinnati

Agenda—Morning

8:45 am – 9:15 am	Welcome/Updates NDMC NINDS	Pooja Khatri Jordan Elm Scott Janis
9:15 am – 9:30 am	Training Core Update	Randy Marshall Devin Brown
9:30 am – 9:45 am	CRP Core Update	Heena Olalde Kinga Aitken
9:45 am – 10:45 am	ARCADIA-CSI Final Results & ARCADIA Secondary Results Discussion	David Tirschwell Maarten Lansberg Ron Lazar Hooman Kamel
10:45 am – 11:30 am	Themes for priority setting conferences	Pooja Khatri Steve Cramer Karen Johnston Hooman Kamel
11:30 am – 12:00 pm	Break and grab lunch to bring back to the room	

Agenda—Afternoon

Reassemble in Salon at 12:00pm for working lunch

12:00 pm – 12:45 pm

Keynote Speaker

“Embedding pragmatic trials within emergency and critical care”

Matthew Semler, MD

Associate Professor of Medicine, Anesthesiology,
and Biomedical Informatics

Associate Director of the Medical Intensive Care Unit

Director, Center for Learning Healthcare

Vanderbilt University

****Please Note: The breakout sessions below will not be livestreamed. Remote attendees rejoin at 2:15pm****

12:45 pm – 2:00 pm

Breakout session: Small group discussion on themes for priority setting conferences. There will be 3 groups:

- Recovery Group Florida Sean Savitz, Maarten Lansberg
- Prevention Group Alabama Cheryl Bushnell, Randy Marshall
- Acute Group Salon C Enrique Leira, Mark Alberts

2:00 pm – 2:15 pm

Break

2:15 pm – 3:15 pm

Reassemble in Salon: Moderators to summarize small group discussions for larger group considerations. Each group will have 10 minutes for summary and 10 minutes for Q&A.

3:15 pm – 3:30 pm

Questions/wrap-up

Housekeeping

- **Wifi** Marriott Conference
 - Access code: encore (don't need room #)
- **Online link:** <https://tinyurl.com/4b3pw3n8>
- **Restrooms** towards lobby (two rights after exiting room)
- **Charging stations** are on left and right of room
- Can leave **luggage** in Salon B (locked during mtg)
- **Parking passes** – see Rose or Kristine if not in your name badge holder
- **Airport buses** at 4pm
- Please ask your questions and bring your ideas today!



Trial Portfolio as of Today

9 Studies Completed

- 6 Definitive (including 2 platform) and 2 Pilot Trials
- 1 Ancillary Study

1 Paused

15 Ongoing Studies

- 11 Trials, 3 Ancillary Studies, 1 Biomarker Validation Study

1 New Trial Funded

Completed Studies

PREVENTION OF STROKE (2)

ARCADIA No evidence of benefit of apixaban for stroke with evidence of atrial cardiopathy

- **ARCADIA-CSI** Cognition and silent infarcts – final results pending

PRIMARY STROKE PREVENTION IN COVID (2)

ACTIV 4C Platform Antithrombotic approach for patients discharged from hospital with COVID-19

ACTIV 4A Platform Antithrombotic approach for inpatient COVID-19 pts

ACUTE STROKE TREATMENT (4)

MISTIE 3 No evidence of benefit of minimally invasive surgery for ICH evacuation

DEFUSE 3 Large treatment benefit of EVT for imaging selected patients at 4.5-16h from onset

I-DEF Futility of deferoxamine for three-month outcomes after ICH

MOST No evidence of benefit of adjunctive epifibatide or argatroban with intravenous thrombolysis

STROKE RECOVERY & REHABILITATION (1)

TELEREHAB Noninferiority of telehealth to in-person, dose-matched post-stroke rehabilitation



Completed Studies

PREVENTION OF STROKE (2)

ARCADIA No evidence of benefit of apixaban for stroke with evidence of atrial cardiopathy

- **ARCADIA-CSI** Cognition and silent infarcts – final results pending

PRIMARY STROKE PREVENTION IN COVID (2)

ACTIV 4C Platform Antithrombotic approach for patients discharged from hospital with COVID-19

ACTIV 4A Platform Antithrombotic approach for inpatient COVID-19 pts

ARCADIA primary paper (JAMA)
Secondary analyses today
ARCADIA CSI primary results today

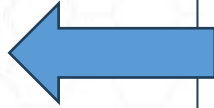
Therapeutically anticoagulated noncritically ill pts but not critically ill pts (NEJM X 2)

No benefit of P2Y12 Inhibitors, SGLT2 inhibitors, or P-Selectin-Inhibitor Crizanlizumab (JAMA, Lancet Diabetes Endocrin, Circulation)

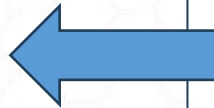


Completed Studies

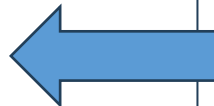
Paved for next ICH evac trials



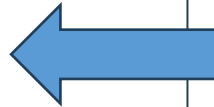
Global impact on acute stroke care



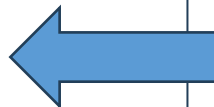
Raised awareness of 6 month outcomes for ICH



Primary paper in NEJM this month



Precursor to newly funded Telerehab-2



ACUTE STROKE TREATMENT (4)

MISTIE 3 No evidence of benefit of minimally invasive surgery for ICH evacuation

DEFUSE 3 Large treatment benefit of EVT for imaging selected patients at 4.5-16h from onset

I-DEF Futility of deferoxamine for three-month outcomes after ICH

MOST No evidence of benefit of adjunctive epifibatide or argatroban with intravenous thrombolysis

STROKE RECOVERY & REHABILITATION (1)

TELEREHAB Noninferiority of telehealth to in-person, dose-matched post-stroke rehabilitation



15 Ongoing Trials/Studies

PREVENTION OF STROKE (9)

CREST-2 Endarterectomy/stenting of asymptomatic carotid stenosis (N=2486/2480)

- **CREST-H** Cognitive outcomes in hemodynamically impaired subset (N=392/385)

Sleep-SMART Treatment of obstructive sleep apnea (Ph 3, N=1601/3062)

SATURN Statin continuation in ICH survivors (N=600/1426)

- **SATURN-MRI** Silent stroke (N=229/894)

ASPIRE Apixiban for afib after ICH (N=340/700)

CAPTIVA Anticoagulation vs antiplatelets for intracranial stenosis (N=514/1683)

- **CAPTIVA MRI** Biomarkers of recurrent stroke in intracranial atherosclerotic stenosis (N=3/300)

FOCAS Corticosteroids for pediatric stroke due to focal cerebral arteriopathy (N=8/80)

ACUTE STROKE TREATMENT (3)

FASTEST FVIIa for acute ICH (N=543/860)

SISTER Novel clot-dissolving Ab, TS23, for ischemic stroke (N=9/300)

STEP Platform Registry-supported trial platform to optimize outcomes after LVO and MVO

STROKE RECOVERY & REHABILITATION (3)

TRANSPORT-2 Transcranial direct stimulation for UE recovery (N=129/129)

I-ACQUIRE Intensive infant rehabilitation for ischemic stroke (N=215/216)

VERIFY Acute prediction of UE motor recovery and function (N=252/657)

Telerehab-2 Telehealth in home vs usual care for UE motor function (N=0/202)



Prevention Updates

PREVENTION OF STROKE (9)

CREST-2 Endarterectomy/stenting of asymptomatic carotid stenosis (N=2486/2480)

- **CREST-H** Cognitive outcomes in hemodynamically impaired subset (N=392/385)

Sleep-SMART Treatment of obstructive sleep apnea (Ph 3, N=1601/3062)

SATURN Statin continuation in ICH survivors (N=600/1426)

- **SATURN-MRI** Silent stroke (N=229/894)

ASPIRE Apixiban for afib after ICH (N=340/700)

CAPTIVA Anticoagulation vs antiplatelets for intracranial stenosis (N=514/1683)

- **CAPTIVA MRI** Biomarkers of recurrent stroke in intracranial atherosclerotic stenosis (N=3/300)

FOCAS Corticosteroids for pediatric stroke due to focal cerebral arteriopathy (N=8/80)

Completed enrollment 10/JUL/2024!
CEA-MED follow up completed SEPT/2024
and CAS-MED follow up to be completed
on 31/JUL/2025

CREST-H in follow up as well!

Prevention aim on hold as of
28/JUN/2024
Recovery aim ongoing (sample
size reduction)



Acute and Recovery Updates

ACUTE STROKE TREATMENT (3)

FASTEST FVIIa for acute ICH (N=543/860)

SISTER Novel clot-dissolving Ab, TS23, for ischemic stroke (N=9/300)

STEP Platform Registry-supported trial platform to optimize outcomes after LVO and MVO

STROKE RECOVERY & REHABILITATION (4)

TRANSPORT-2 Transcranial direct stimulation for UE recovery (N=129/129)

I-ACQUIRE Intensive infant rehabilitation for ischemic stroke (N=215/216)

VERIFY Acute prediction of UE motor recovery and function (N=252/657)

Telerehab-2 Telehealth in home vs usual care for UE motor function (startup, N=202)

Key interim analysis in December
FDA approved amendment to incorporate an enrichment and promising zone design

New assets under active development

Completed enrollment 24/May/2024 and last follow up last week!
Anticipate results at ISC 2025

Almost completed enrollment!
12 months follow up remaining

NEW

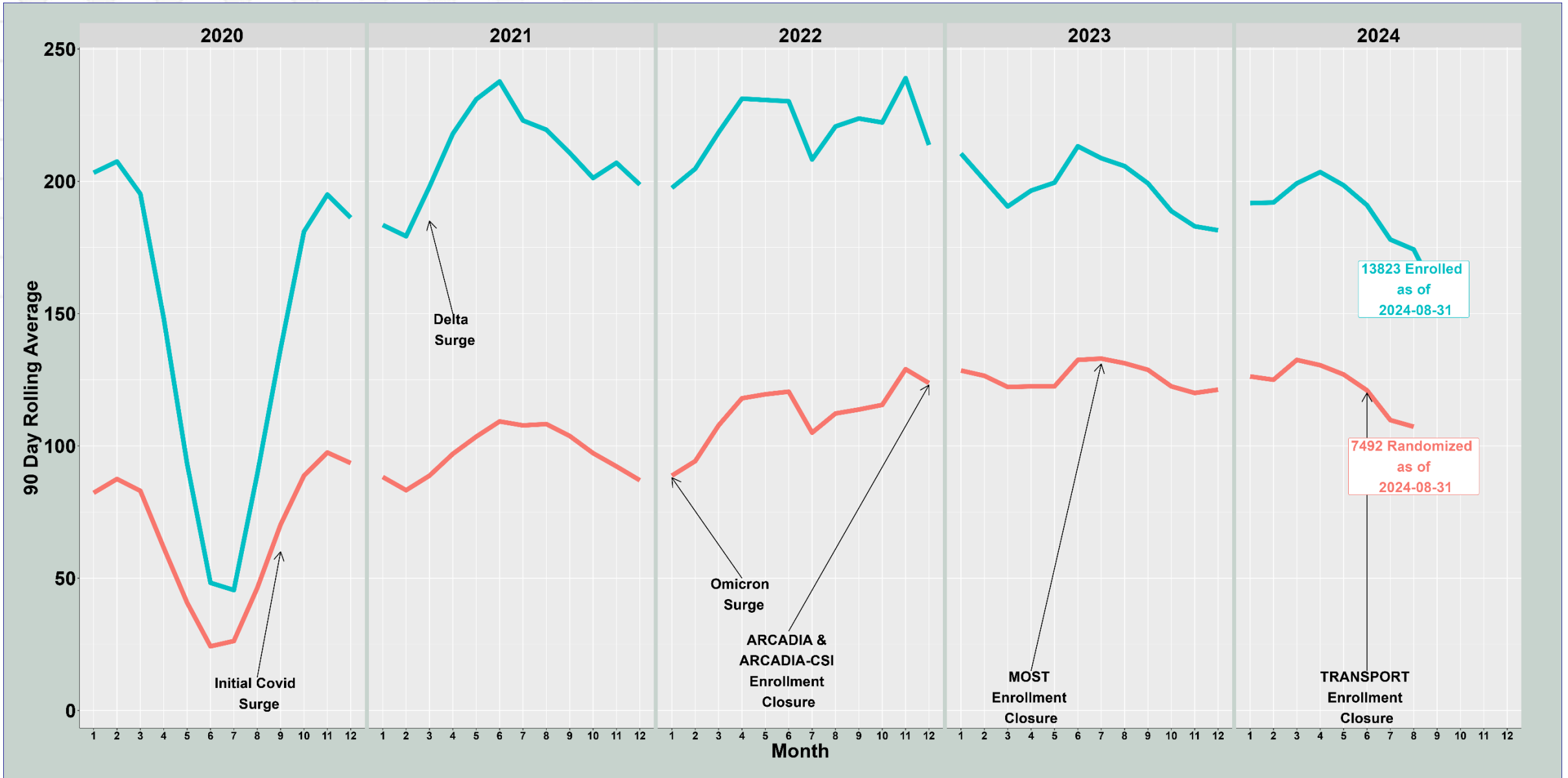


SISTER

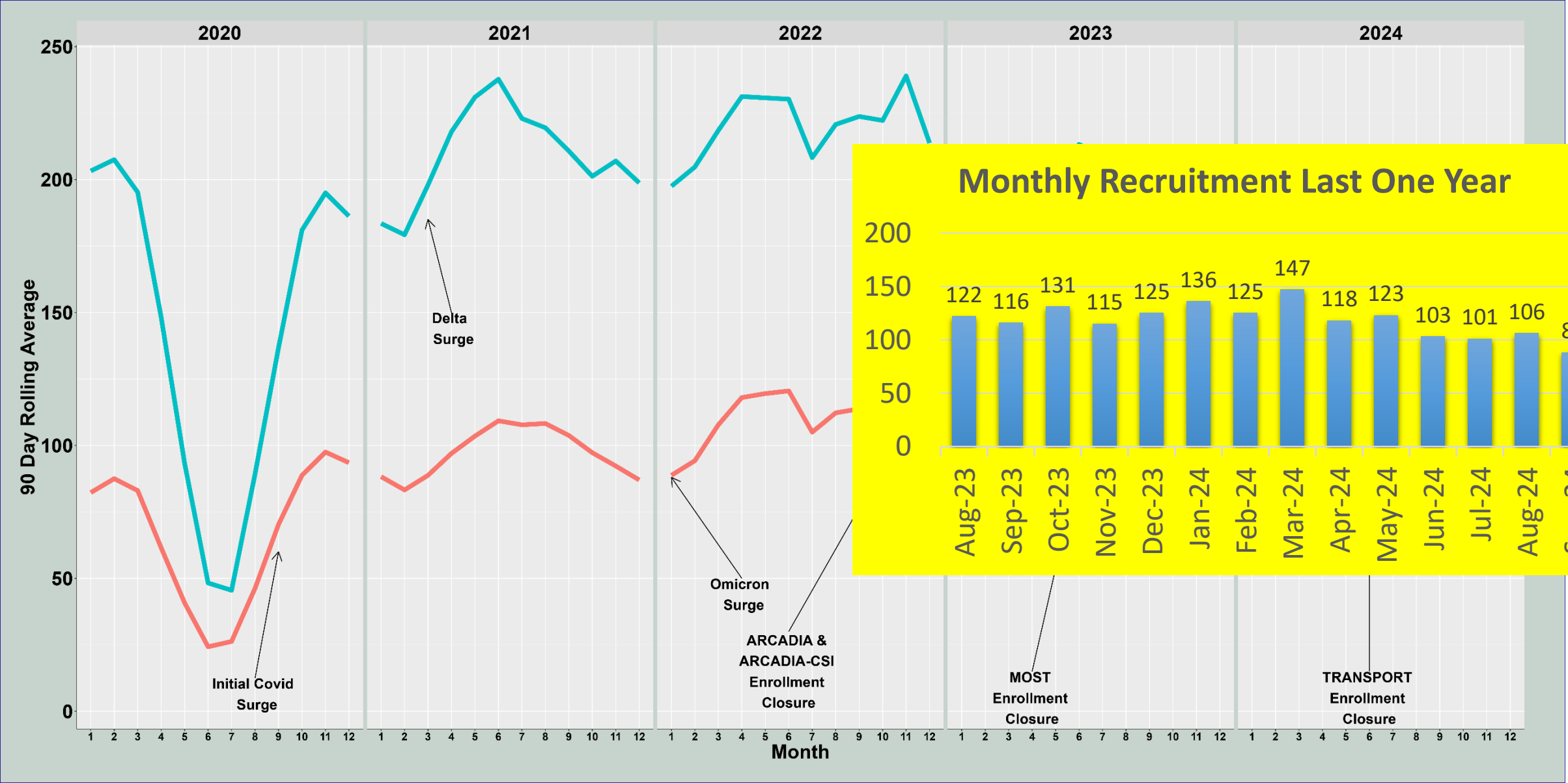


StrokeNet Thrombectomy
Endovascular Platform

>13,000 Enrolled and >7000 Randomized



>13,000 Enrolled and >7000 Randomized



Examples of Innovative Design Features

- Adaptive enrichment – DEFUSE-3
- Multi-arm, multistage – MOST, CAPTIVA
- Patient-Reported Outcome -- Utility-weighted modified Rankin – MOST, FASTEST, STEP
- Covariate adjusted randomization – Telerehab, iACQUIRE, MOST, SATURN, TRANSPORT2
- Response adaptive randomization – Sleep SMART, MOST, SISTER, STEP
- Sample size re-estimation – ARCADIA, Sleep SMART, ASPIRE, SATURN
- Step forward randomization – FASTEST
- Emergency consent – FASTEST
- Infant population – iACQUIRE
- Utility function – SISTER
- Randomized, embedded, multifactorial, adaptive platform (REMAP) – STEP
- Registry-supported – STEP

International Trials

NIH StrokeNet Network Standard Operating Procedure

SOP Number: ADM 27

SOP Name: Management of StrokeNet Trials with International Sites

Effective Date: 09/24/2024



StrokeNet Trials Publications (Last One Year)

JAMA | Original Investigation

Apixaban to Prevent Recurrence After Cryptogenic Stroke in Patients With Atrial Cardiopathy

The ARCADIA Randomized Clinical Trial

JAMA Neurology | Original Investigation

Apixaban vs Aspirin in Patients With Cancer and Cryptogenic Stroke A Post Hoc Analysis of the ARCADIA Randomized Clinical Trial

PEDIATRICS®

Content ▾

Authors/Reviewers ▾

FAMILY PARTNERSHIPS | SEPTEMBER 04 2024

Parent Council for a Pediatric Stroke Rehabilitation Clinical Trial ✓

Stroke: Vascular and Interventional Neurology

ORIGINAL RESEARCH

CREST-2 Commitment to Rigorous
Assessment of Carotid Stenting for Primary
Prevention of Stroke

Circulation

Volume 148, Issue 5, 1 August 2023; Pages 381-390
<https://doi.org/10.1161/CIRCULATIONAHA.123.065190>



ORIGINAL RESEARCH ARTICLE

Effect of the P-Selectin Inhibitor Crizanlizumab on
Survival Free of Organ Support in Patients Hospitalized
for COVID

Effect of sodium–glucose co-transporter-2 inhibitors on
survival free of organ support in patients hospitalised for
COVID-19 (ACTIV-4a): a pragmatic, multicentre, open-label,
randomised, controlled, platform trial

The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Adjunctive Intravenous Argatroban
or Eptifibatide for Ischemic Stroke

Committee Updates

CHAIRS		
Working Groups	Acute	Karen Johnston, MD & Jeff Saver, MD
	Prevention	Hooman Kamel, MD & Scott Kasner MD
	Recovery	Steve Cramer, MD & Steven Wolf, PT PhD
Cores	Fellow Education/Training	Randy Marshall, MD & Devin Brown, MD
	CRP Education/Training (new)	Heena Olalde, RN, MSN & Kinga Aitken, MPH, CCRP
	Diversity/Equity/Inclusion (new)	B. Boden-Albala, MPH, DrPH & L. Skolarus, MD
Advisory	Patient Rep/Advocacy (new)	Flannery O'Neil, MPH & Aqualyn Kennedy, MBA
	Pediatrics	Heather Fullerton, MD
	Preclinical (new)	Lauren Sansing, MD
	Telestroke	Chris Streib, MD & Abbey Staugaitis, RN, MSN, CCRC

Committee Updates

Working Groups	Acute	Membership rotated; Trial development; Scientific themes for priority setting conferences
	Prevention	
	Recovery	
Cores	Fellow Education/Training	Stay tuned today
	CRP Education/Training (new)	
	Diversity/Equity/Inclusion (new)	Formed group—attending WGs—how to collect baseline demographics and analysing demographics of ongoing trials
Advisory	Patient Rep/Advocacy (new)	Expanded involvement, STEP ICF
	Pediatrics	Formed group –attdg WGs – STEP protocol
	Preclinical (new)	Formed group – attdg WGs
	Telestroke	Remote Consent Practices Survey, MOST eCONSENT pub (higher enrollment, fewer ICF PVs)