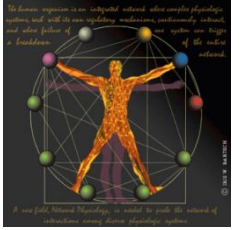




# Second International Summer Institute on Network Physiology (ISINP)

Lake Como School of Advanced Studies - 28 July - 02 August, 2019



## Developing a Data Collection System for the Injured Brain to Enable Network Physiology Research

Dick Moberg  
President  
Moberg Research, Inc.  
dick@moberg.com



- Intro to TBI and the need for big data
- MRI: Technology and Future
- Device Data Collection: Problems and solutions
- Integration of other data: EMR, Imaging, Biomarkers
- Trends: FDA panel, etc.
- Clinical examples of utility
- Future

- Intro to TBI and the need for big data
  - Sheila story, TRACK TBI, etc.
  - We need better data
- MRI: Technology and Future
  - Challenges to getting that data (devices, standards, databases, etc.)
- Device Data Collection: Problems and solutions
  - Challenges
- Integration of other data: EMR, Imaging, Biomarkers
- Trends: FDA panel, etc.
- Clinical examples of utility
  - Cases (Brandon)
- Future
  - and DOD project

- There are a couple of weeks left till the start of the International Summer Institute on Network Physiology (ISINP-2019) in Como.
- Could you please send PDFs of your lecture(s) at your earliest convenience. We would need to have them latest by July 25.
- 
- The PDFs form the package of materials for the ISINP participants and will be made available only to the participants.
- 
- Lectures duration is 30 min plus 5 min for questions.
- 
- Please note that lectures present diverse topics, and there is very little or no overlap in the background and expertise of speakers. Participants are Ph.D. and M.D. students, postdoctoral fellows, university faculty, physicians and industry researchers with diverse backgrounds in medicine, neuroscience, exercise physiology, computer science and applied mathematics, physics, and biomedical engineering.
- 
- Correspondingly, each lecture should provide clear definitions, background, general concepts, results, and vision for the future in a way that is understandable for non-specialized audiences.
- 
- Note that although the general focus of ISINP is Network Physiology, the meeting is not a narrowly specialized workshop or conference -- it is intended as an institute/school where experts across fields learn from each other. Talks given at conferences to peers in your specific fields may not be quite suitable for this meeting.

- Describe the **need** for comprehensive, high-resolution data in critical care
  - We will use the care of acute brain injury as an example
- Describe the **challenges** of collecting data in a usable form
  - We will describe the state of medical device connectivity and data interoperability
- Describe **progress** to overcome the challenges
  - We will use our work (and others) as examples
- Describe the **future** with “Smart ICUs”
  - We will show examples of what can be done

- Describe the **need** for comprehensive, high-resolution data in critical care
  - We will use the care of acute brain injury as an example
- Describe the **challenges** of collecting data in a usable form
  - We will describe the state of medical device connectivity and data interoperability
- Describe **progress** to overcome the challenges
  - We will use our work (and others) as examples
- Describe the **future** with “Smart ICUs”
  - We will show examples of what can be done



We know a lot  
But compared to other areas of medicine  
we don't even know what we don't know

# Variability in Traumatic Brain Injury



Kevin Pearce – snowboarder  
Fall on half-pipe – severe TBI  
26 days ICU, 5 months rehab  
Slight memory & visual loss



Sarah Burke – skier  
Fall on half-pipe – severe TBI  
9 days ICU  
Died

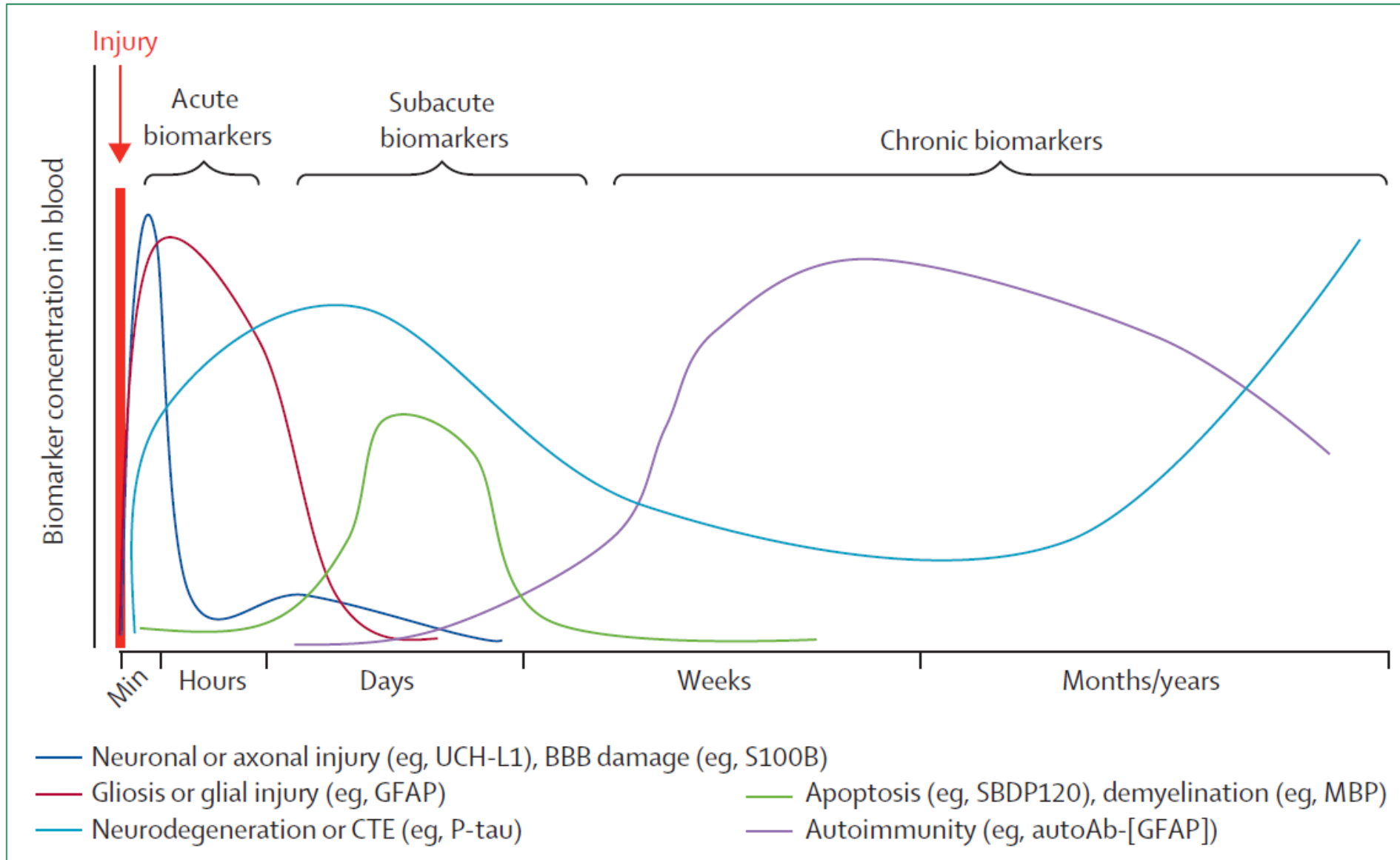


Sara Hall - arborist, runner  
Jogger vs. bicycle – mild TBI  
6 days ICU, 6 months rehab  
Slight hearing & memory loss

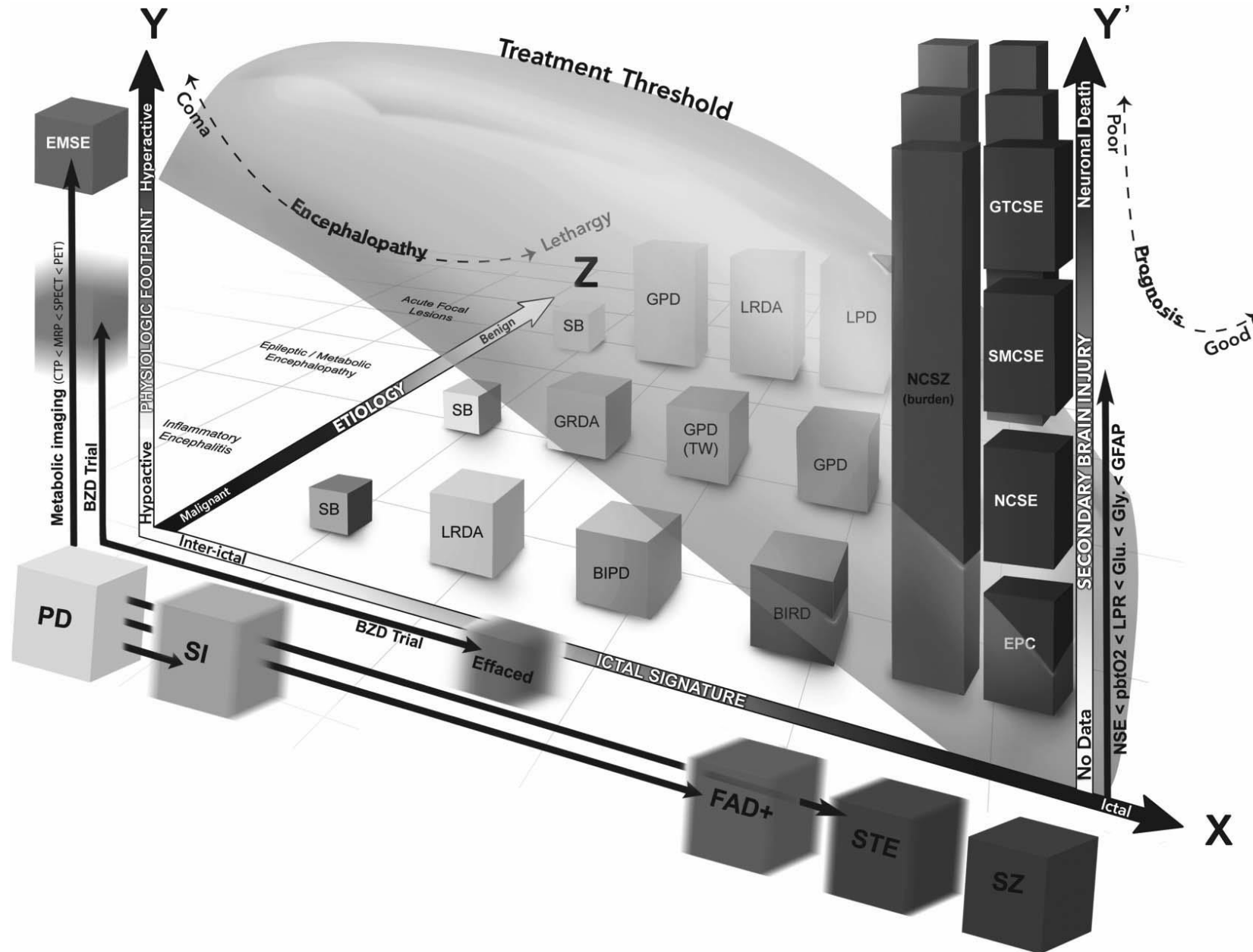
High variability in TBI outcomes due to the extreme complexity of the brain



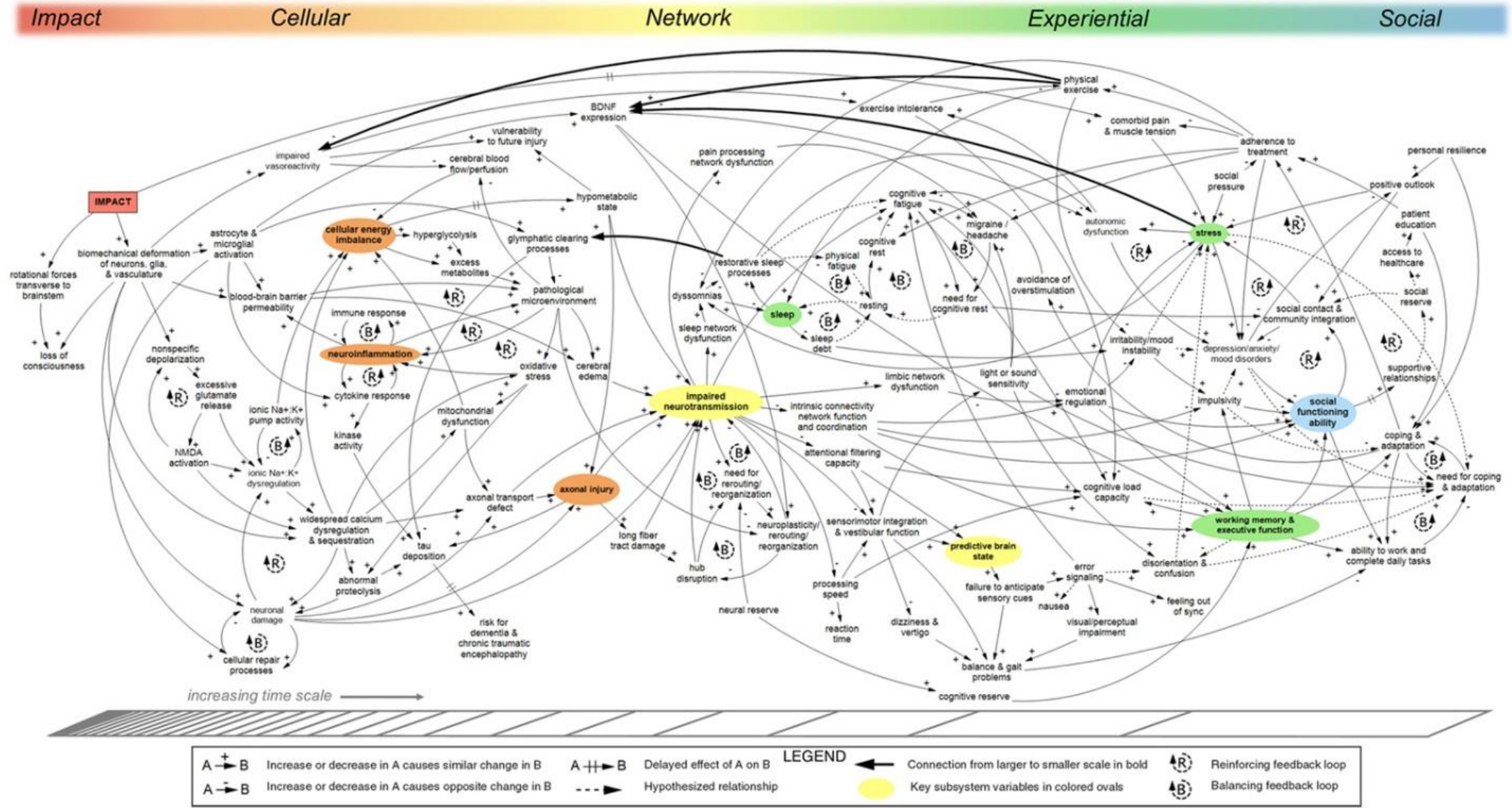
# Complexity of Traumatic Brain Injury



# Complexity of Traumatic Brain Injury



# Complexity of Traumatic Brain Injury



**After 30 years of failed trials, we realize we can't treat the brain the same way as the heart or liver or lungs...with one guideline or drug**

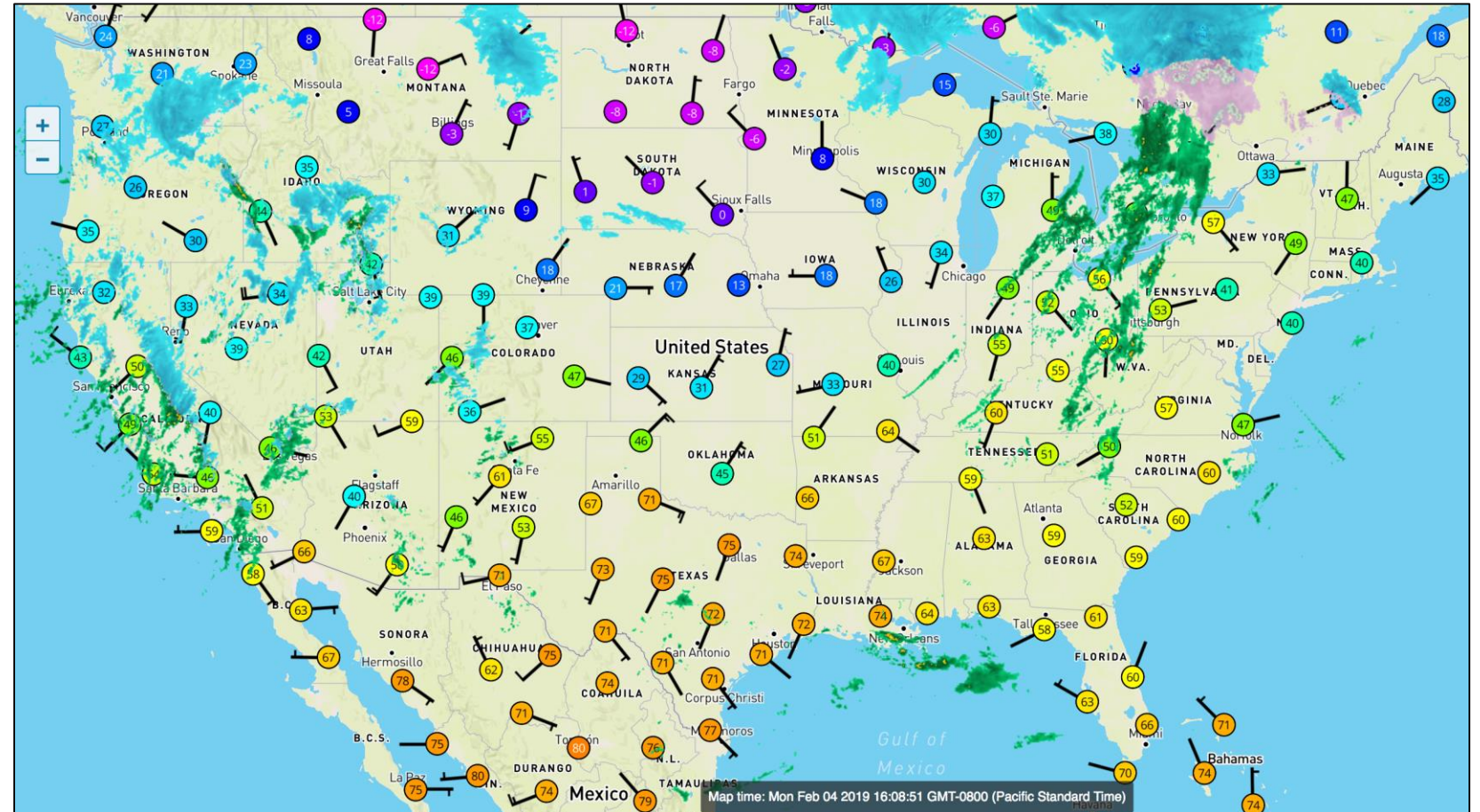


**The field is doing a “re-start” in its approach to brain injury**

- Better definition of TBI
- Better endpoints

We can predict the weather because we have massive amounts of localized data.

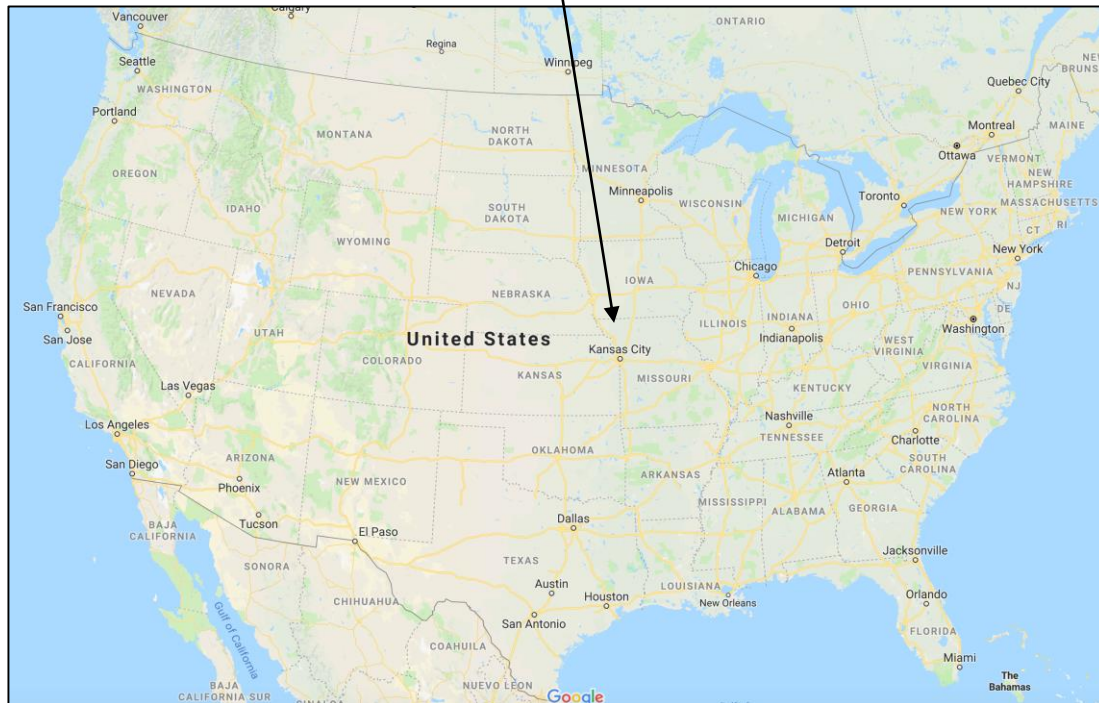
- Satellite imaging
- Localized temperatures, barometric pressure, wind speed & direction, humidity, dew point, etc.
- Sophisticated models



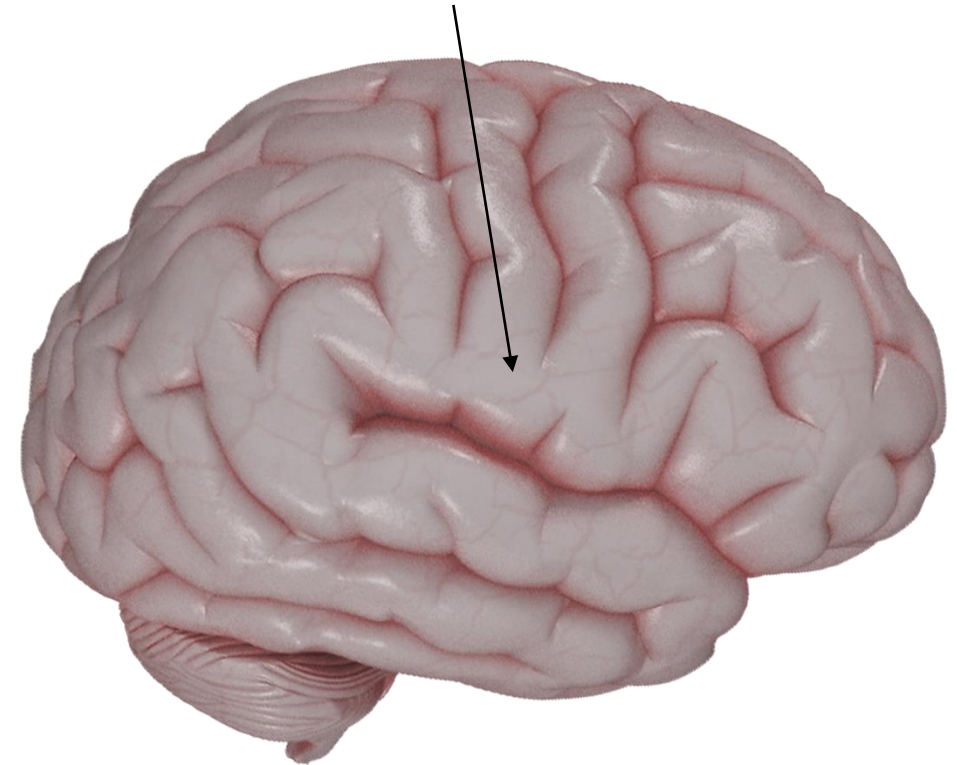
# Current State of Data in the Neuro ICU

Predicting changes in neurocritical care today is like trying to forecast the national weather with only an hourly temperature from Kansas City.

Temp = 72°



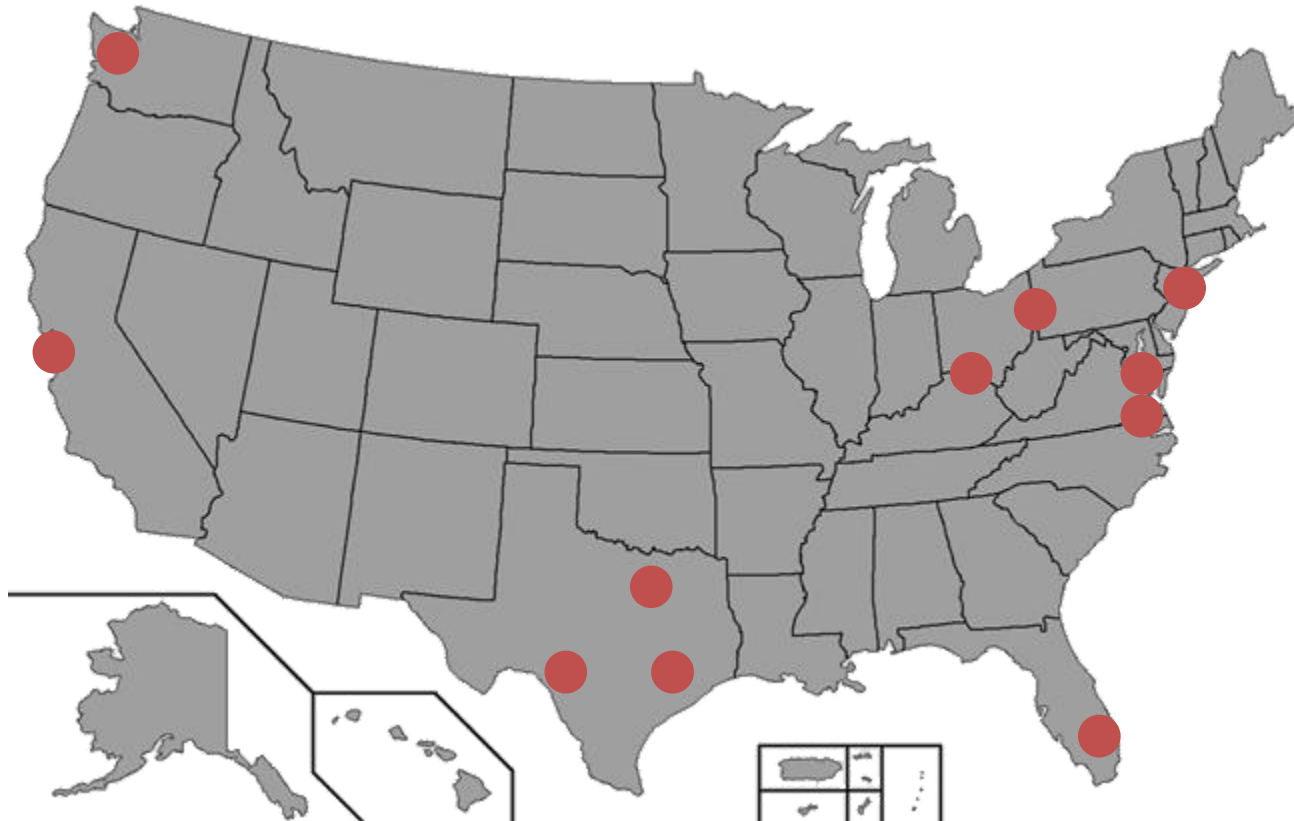
ICP = 8 mmHg



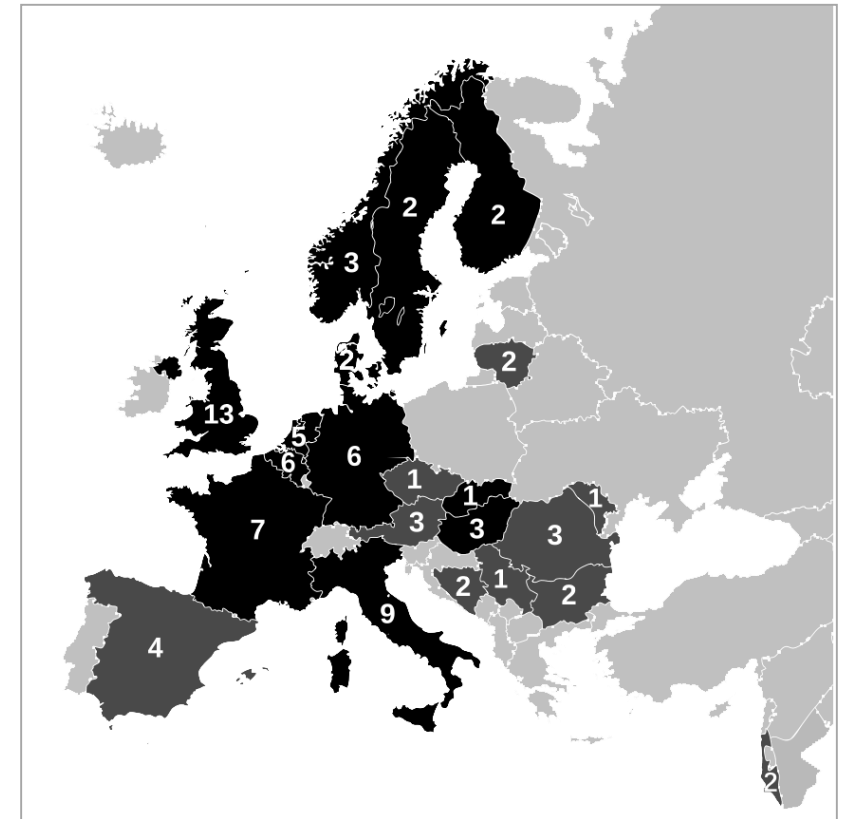
# Defining the Disease – Observational Trials



11 sites  
3,000 subjects



80 sites, 21 countries  
5,400 subjects



On Admission  
Better Definition of TBI

More Accurate  
End Points

Glasgow Coma Score

Imaging

Biomarkers

Genomics

Proteomics



Glasgow Outcome  
Score

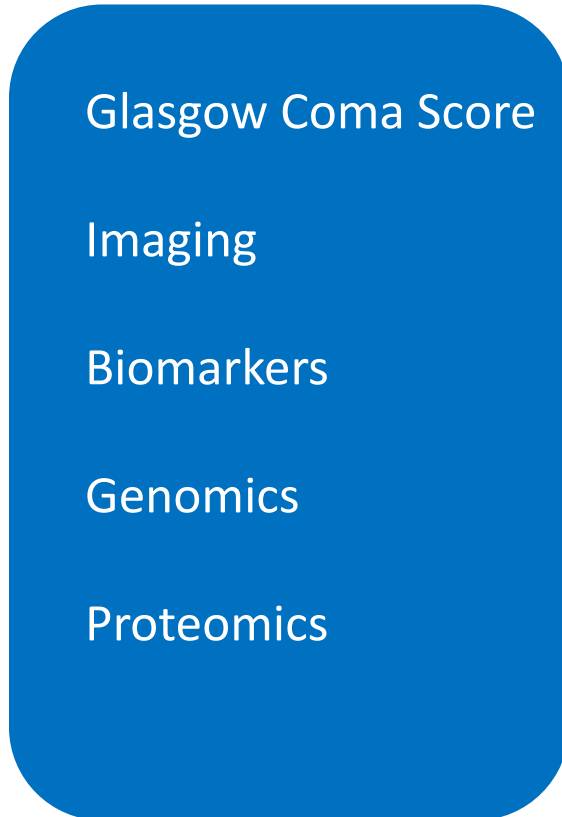
Psychometric  
Testing

Eye Tracking

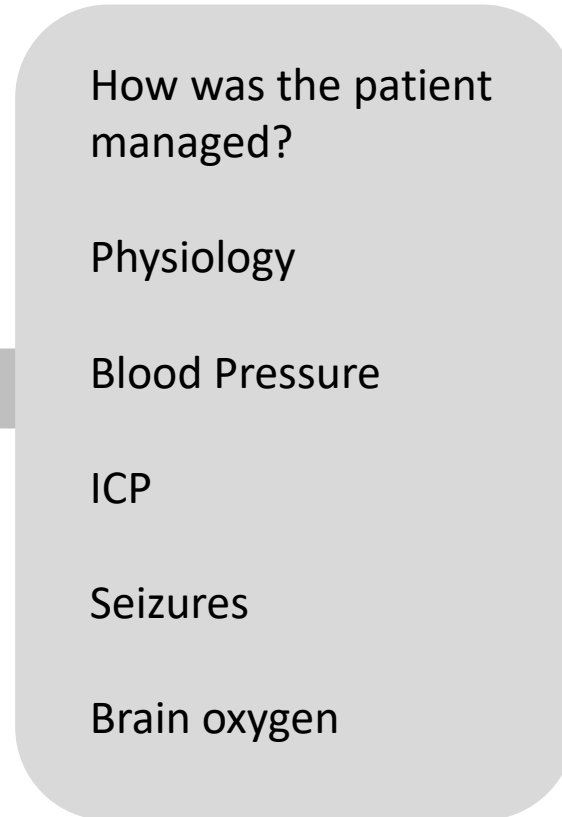
Biomarkers



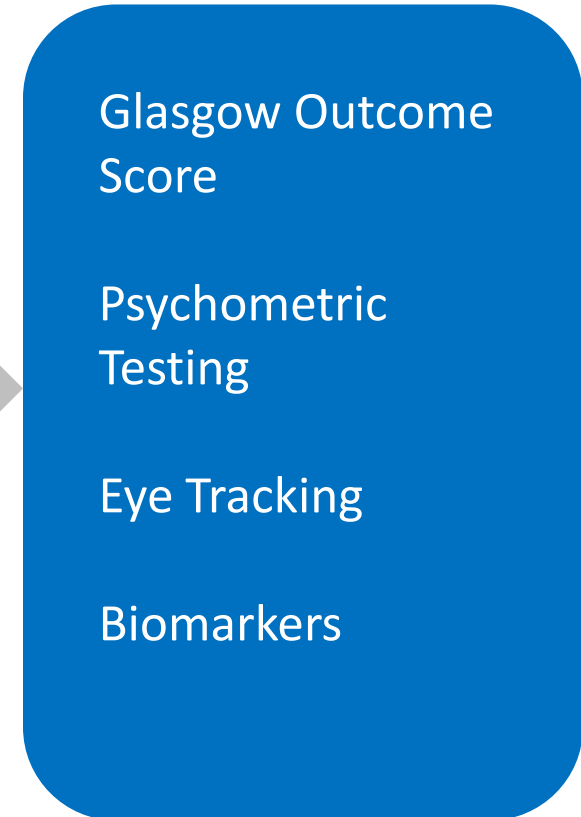
On Admission  
Better Definition of TBI



Overlooked

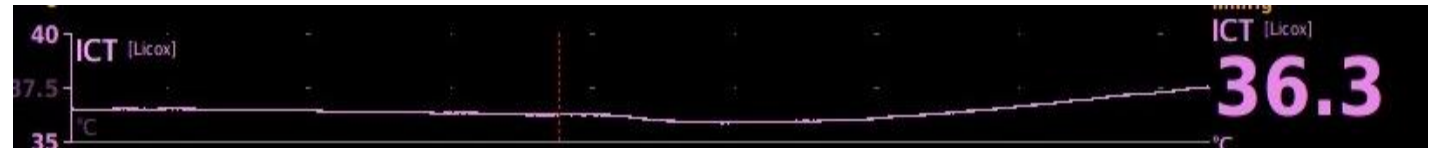
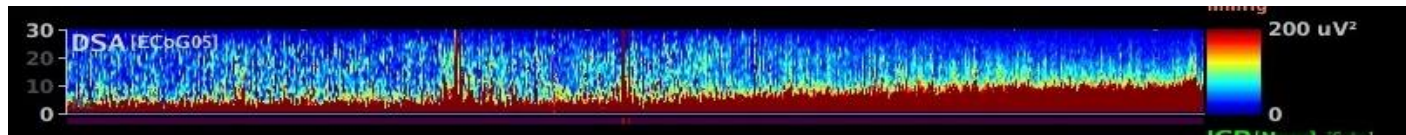
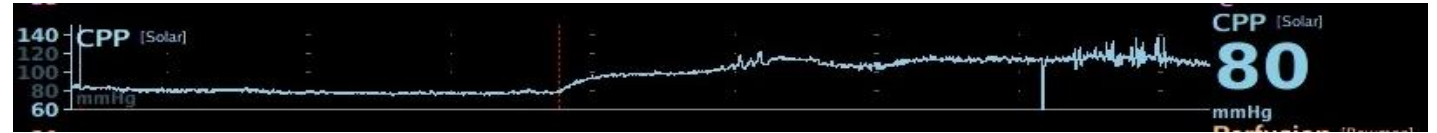
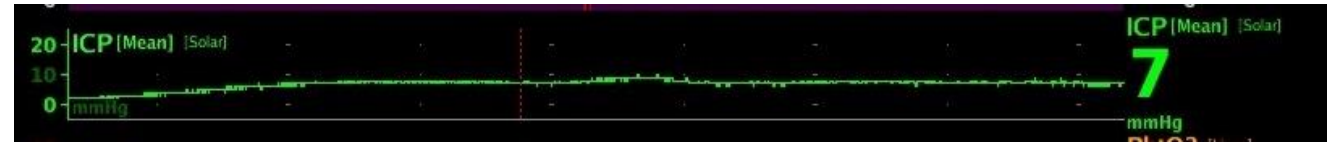
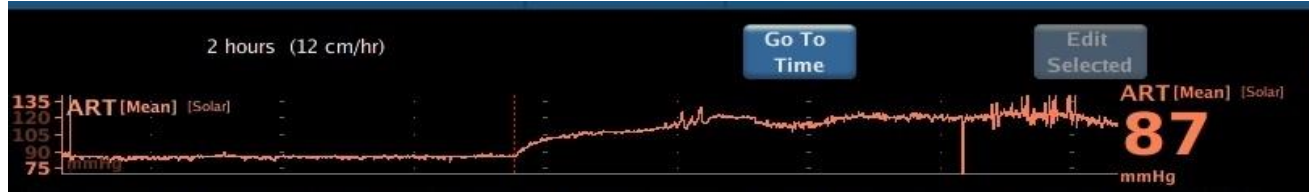


More Accurate  
End Points



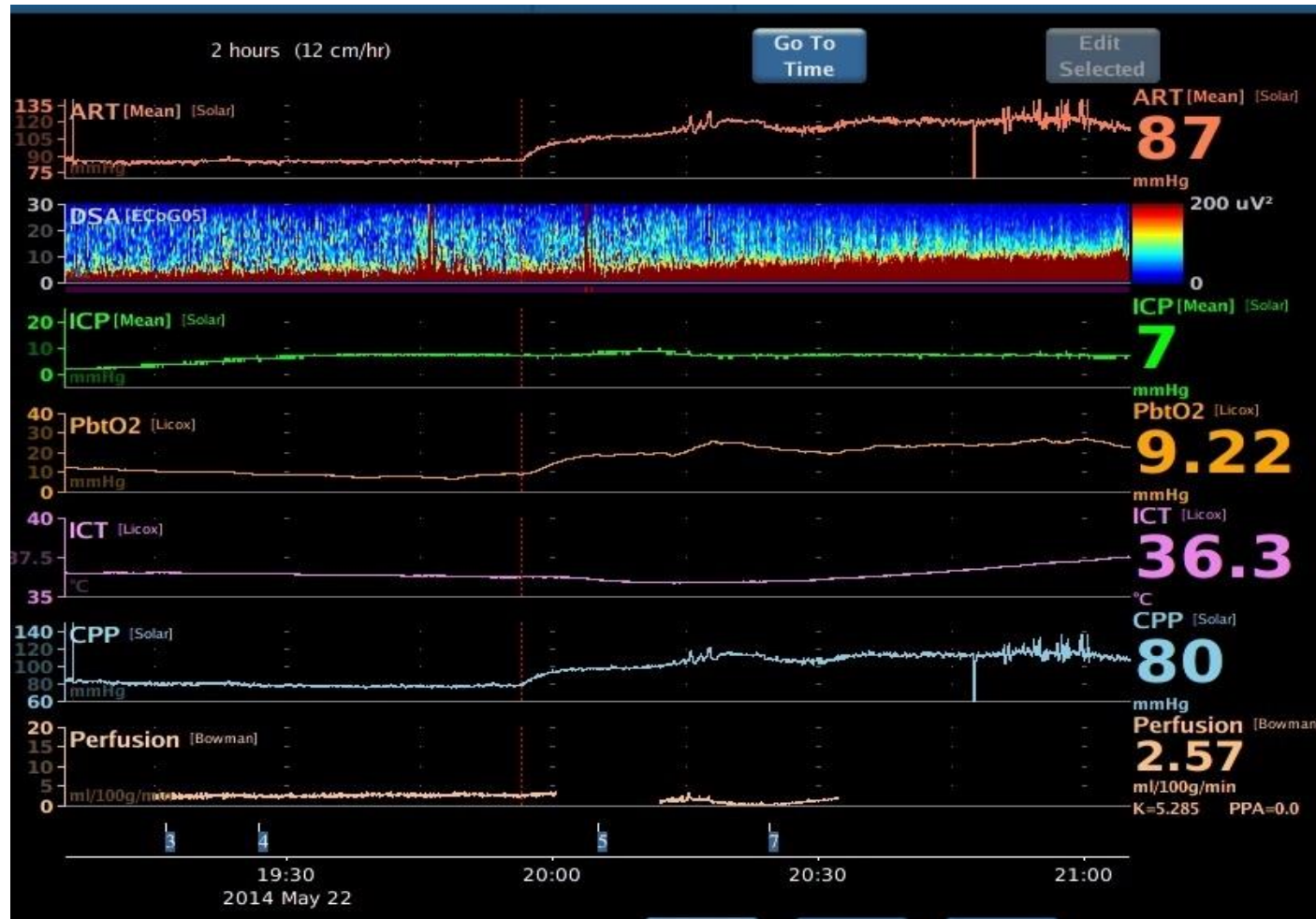
This is what we need  
But this is difficult to get

# Disparate Data



# Combined, Time-synchronized Data

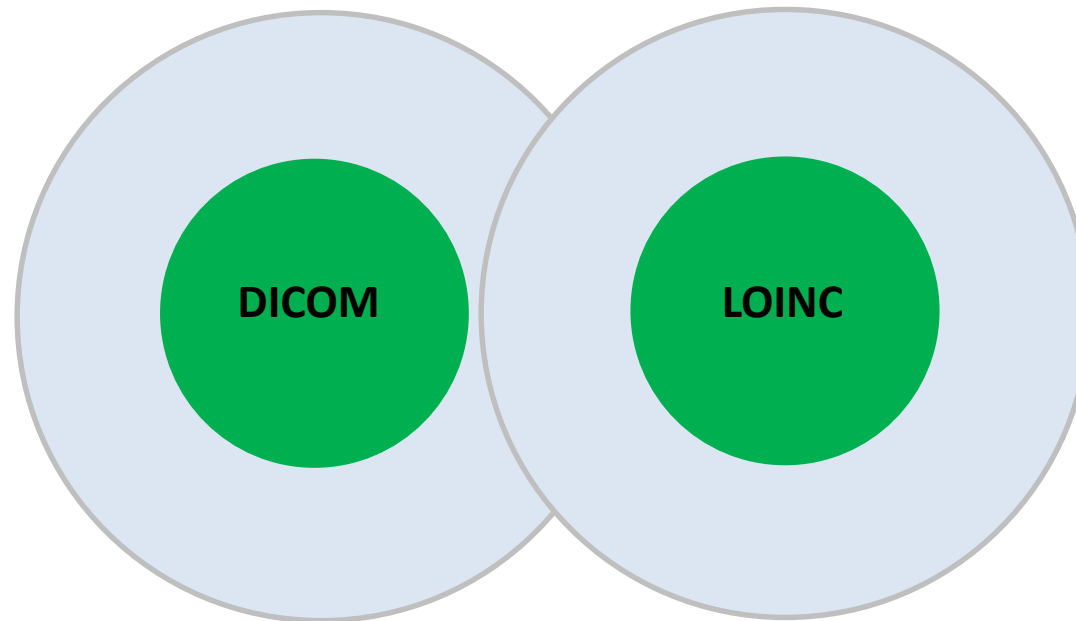
The data tells a story when consolidated and time-synchronized



- Describe the **need** for comprehensive, high-resolution data in critical care
  - We will use the care of acute brain injury as an example
- Describe the **challenges** of collecting data in a usable form
  - We will describe the state of medical device connectivity and data interoperability
- Describe **progress** to overcome the challenges
  - We will use our work (and others) as examples
- Describe the **future** with “Smart ICUs”
  - We will show examples of what can be done

- Device Communications
  - No widely adopted communication standard
  - No widely used nomenclature
  - “Quirks” in every device
    - We had to compensate for the bad designs of others
- Low Resolution Data
- Device Adapters
  - Several attempts, none widely adopted
- Systems of Devices
  - Time synchronization
  - Regulatory - Who is liable in a “closed loop system”

- Standards organizations
  - Lots of them....they all want to get into each others domains
  - DICOM (started as imaging standard) and LOINC (started as lab data standard)...each are growing into the other's territory



- Intracranial Pressure
  - What we have IEEE 11073-10101
  - What we need (location, etc. – other metadata)

## Pupillometer



Reactivity

Size

Symmetry



## Medical Record

Glucose

Blood Gases

Heart Rate

Size ??

Blood Pressure

Urine Output

Medications

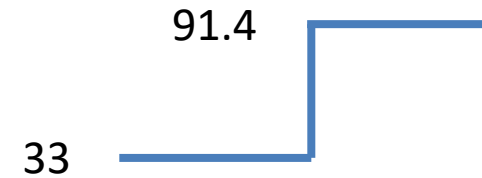
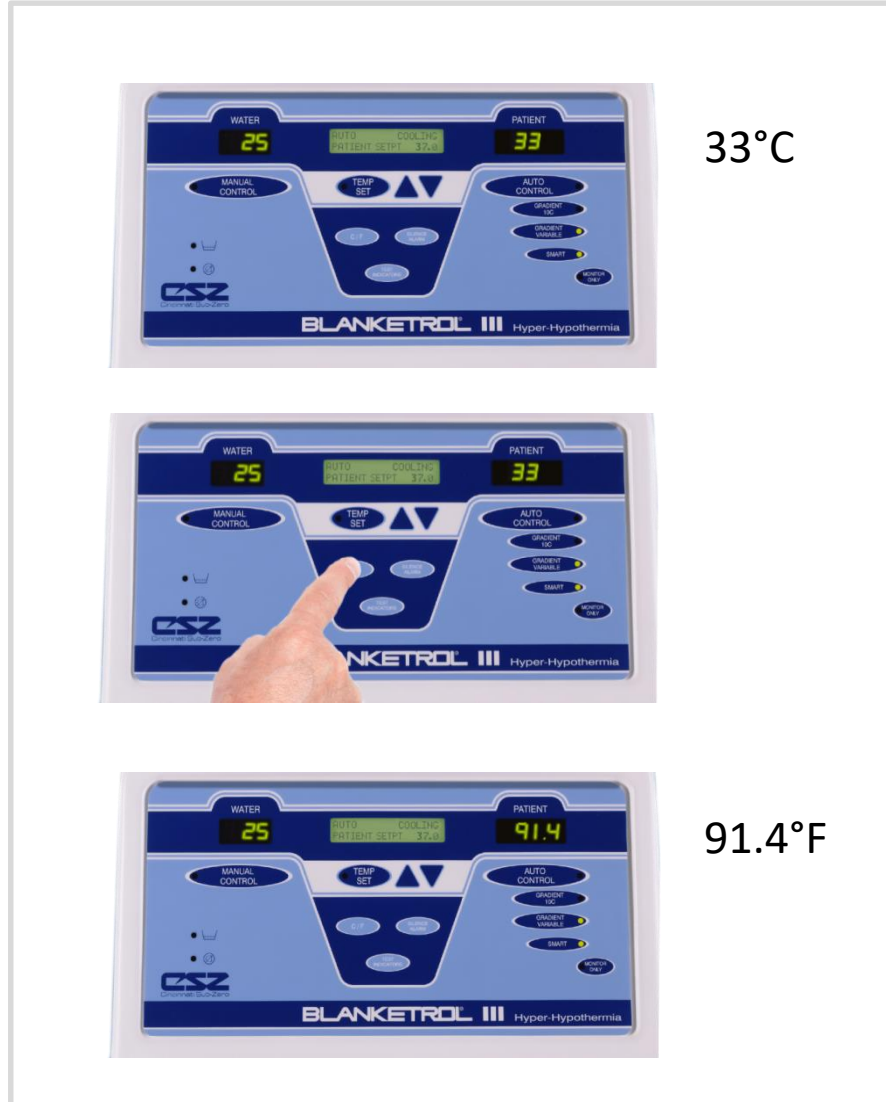


# Problem: Label Confusion



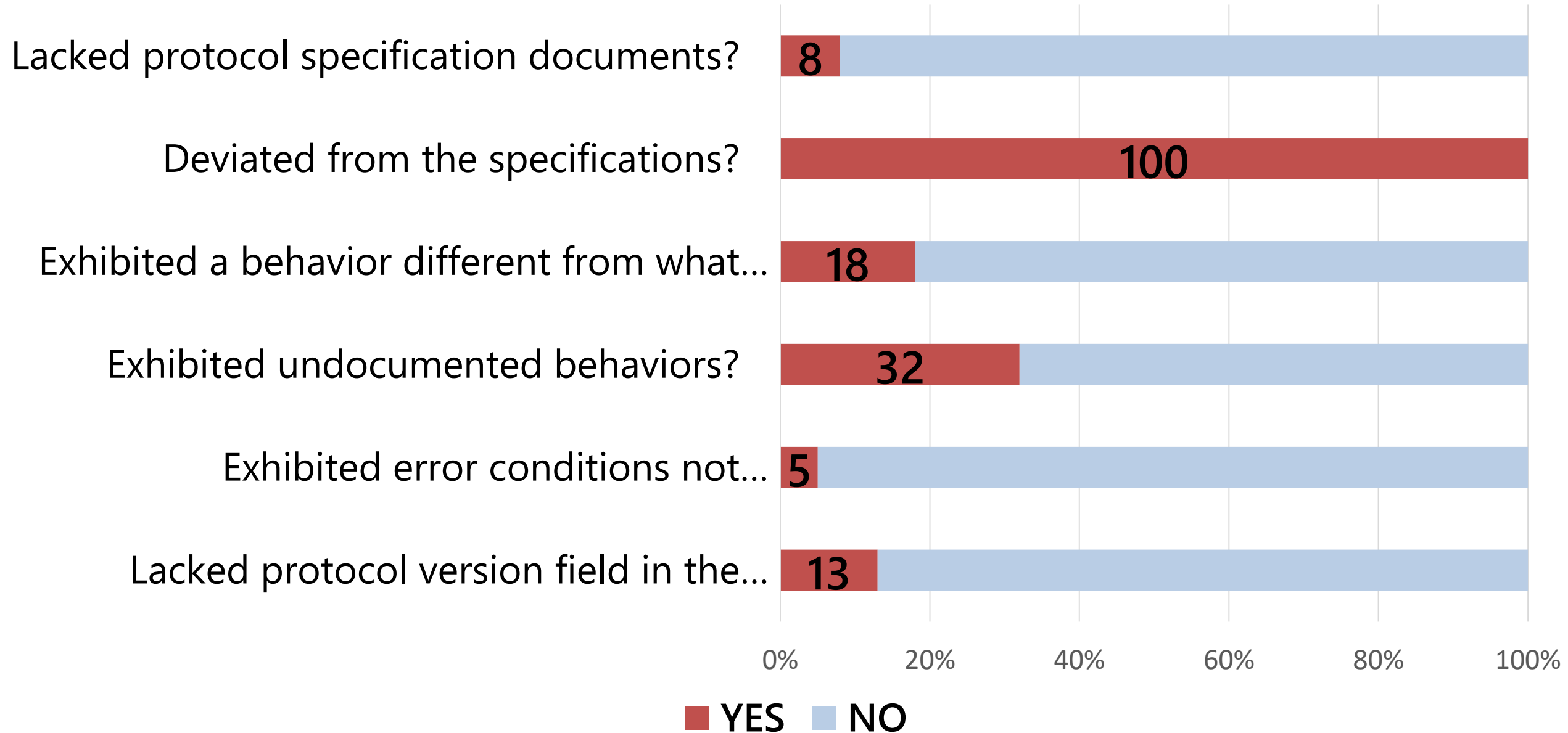
No internal  
labels for PbtO2

# Problem: Inadequate Device Protocol

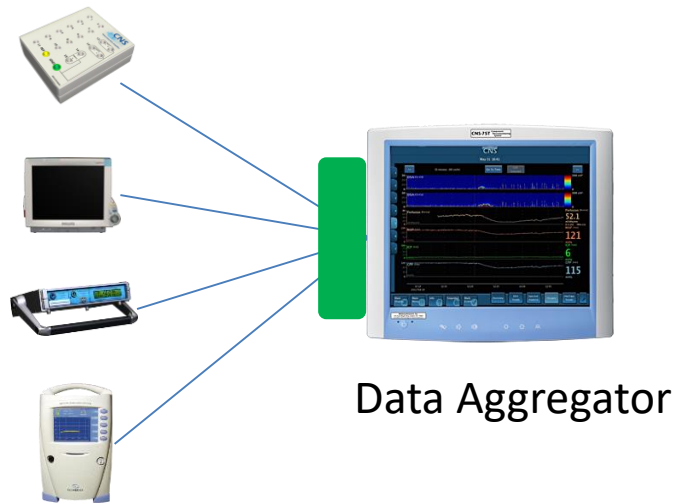


Mirrors the display  
No units sent with the data!!

# Problem: Adherence to Specifications



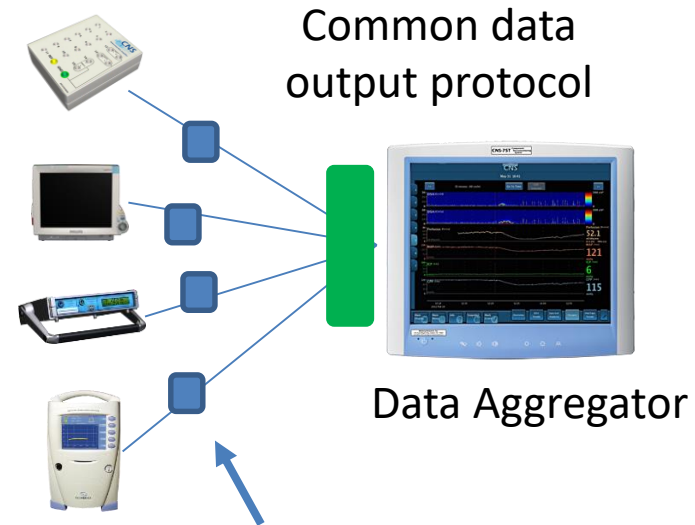
## What we want



Data Aggregator

**Plug and Play**  
Just like your PC

## One way to do this



Common data  
output protocol

Data Aggregator

Device  
Adapters



Digital Device Adapter



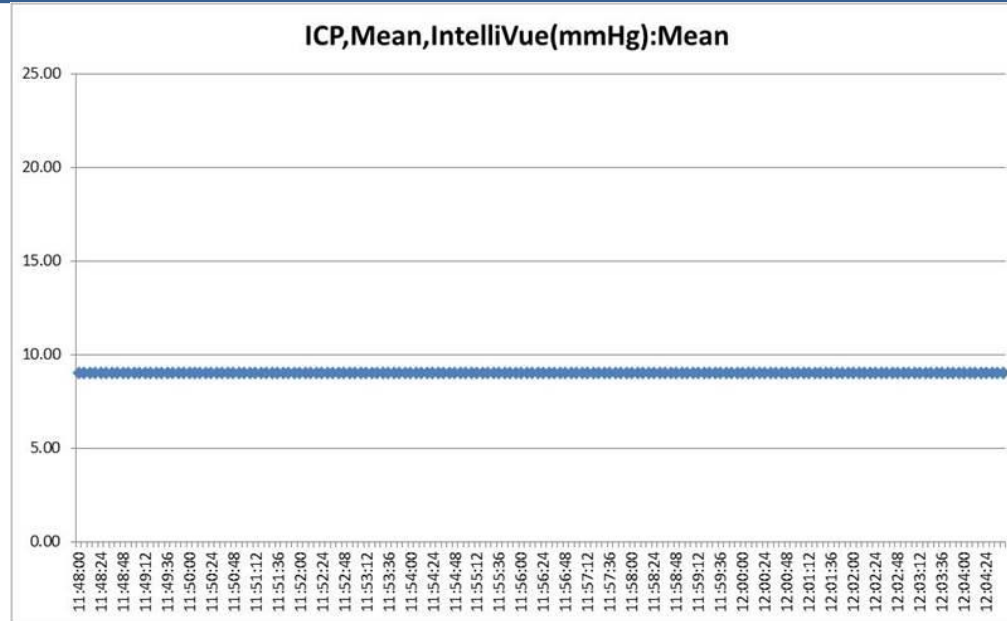
- But....there was no commercial market for an adapter
  - Unless the market scaled up
- So....it was cheaper for us and for us to write a software driver and for customers to buy just a cable



**Traumatic Brain Injury  
Management Algorithm  
Consensus Conference**  
Seattle, Washington  
April 5-7, 2019

Still debating what to monitor  
and how to manage TBI.  
RCTs hard to do.  
Lack of good evidence.  
So reverting to consensus  
conferences.

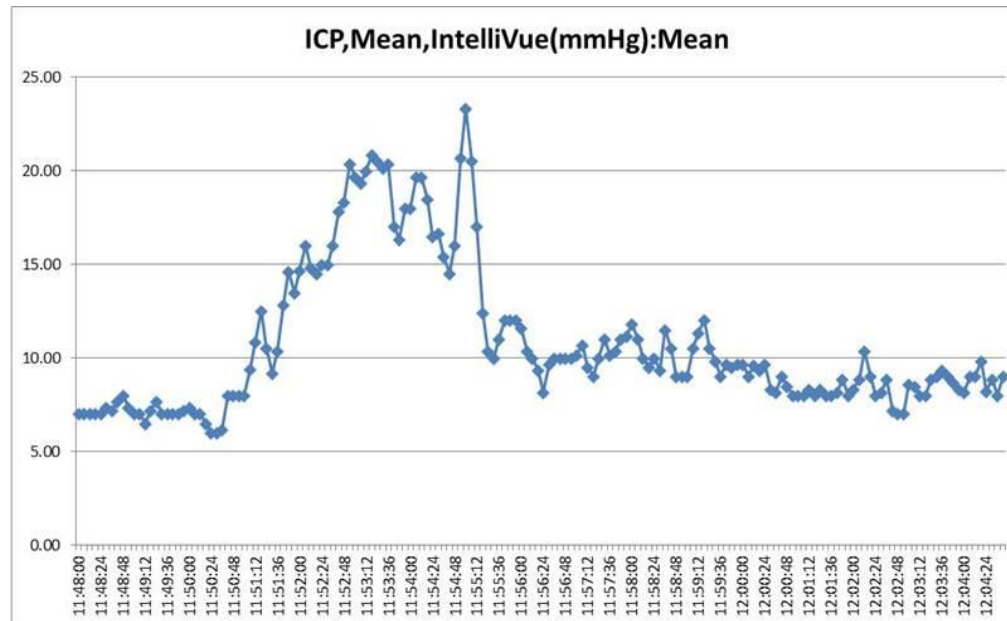
# Intracranial Pressure in the Medical Record vs. Reality



Intracranial Pressure

What the medical record showed

Medical Record



What really happened.

Reality

# Medical Record vs. Continuous Monitoring

Total from 34 Patients with Severe TBI

	Incidence of intracranial hypertension (SCM / EMR)	Incidence of intracranial hypertension (CNS Monitor)	Incidence of cerebral hypoperfusion (SCM / EMR)	Incidence of cerebral hypoperfusion (CNS Monitor)
Mean	23.4	63,020.4	22.2	703.0
Median	15	23,553	9.5	264



Data in the EMR can be inaccurate, missing, or difficult to obtain.



The image is a screenshot of a news article from Kaiser Health News (KHN). The article title is "Death By 1,000 Clicks: Where Electronic Health Records Went Wrong". The background of the article header features a dark, moody photograph of various medical instruments, including scalpels, forceps, and a computer mouse, arranged on a surface. The KHN logo is in the top left, and navigation links for "HEALTH LAW", "AGING", "INDUSTRY", "PHARMA", and "INVESTIGATIONS" are in the top center. Social media icons and a search bar are in the top right. The article text discusses the challenges of electronic health records in the U.S. and is dated March 18, 2019.

**KHN**  
KAISER HEALTH NEWS

CONNECT WITH US: [✉](#) [🐦](#) [f](#) [in](#) [@](#) [📡](#)

HEALTH LAW AGING INDUSTRY PHARMA INVESTIGATIONS Search 🔍

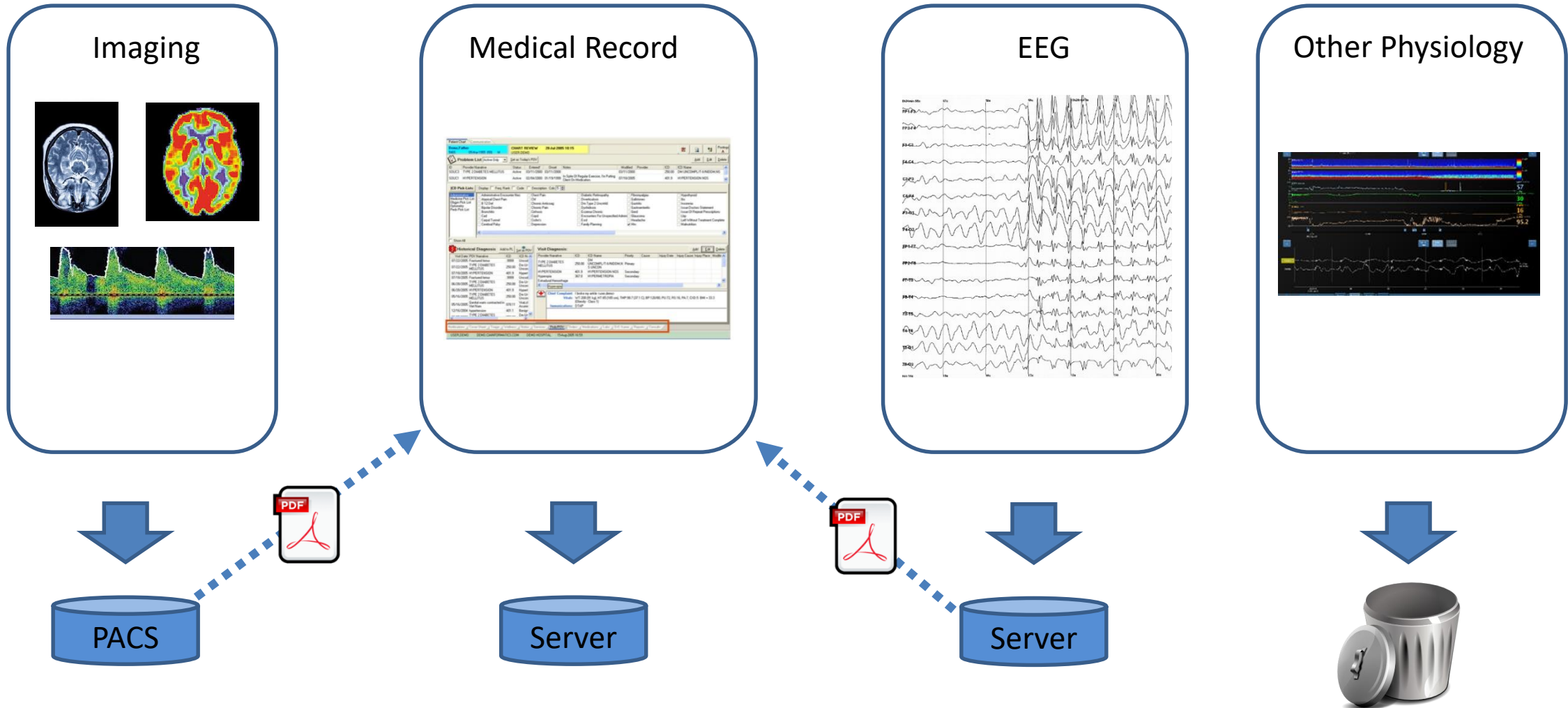
## Death By 1,000 Clicks: Where Electronic Health Records Went Wrong

The U.S. government claimed that turning American medical charts into electronic records would make health care better, safer and cheaper. Ten years and \$36 billion later, the system is an unholy mess. Inside a digital revolution that took a bad turn.

By Fred Schulte and Erika Fry, *Fortune* • MARCH 18, 2019

(The Voorhes for Fortune)

Data is not collected in a way that enables further use



The biggest problem we have is dealing with the Information Technology (IT) groups at each hospital

The biggest problem we have is dealing with the Information Technology (IT) groups at each hospital

What we need...



The Apple TV  
for Hospital IT

- Describe the **need** for comprehensive, high-resolution data in critical care
  - We will use the care of acute brain injury as an example
- Describe the **challenges** of collecting data in a usable form
  - We will describe the state of medical device connectivity and data interoperability
- Describe **progress** to overcome the challenges
  - We will use our work (and others) as examples
- Describe the **future** with “Smart ICUs”
  - We will show examples of what can be done

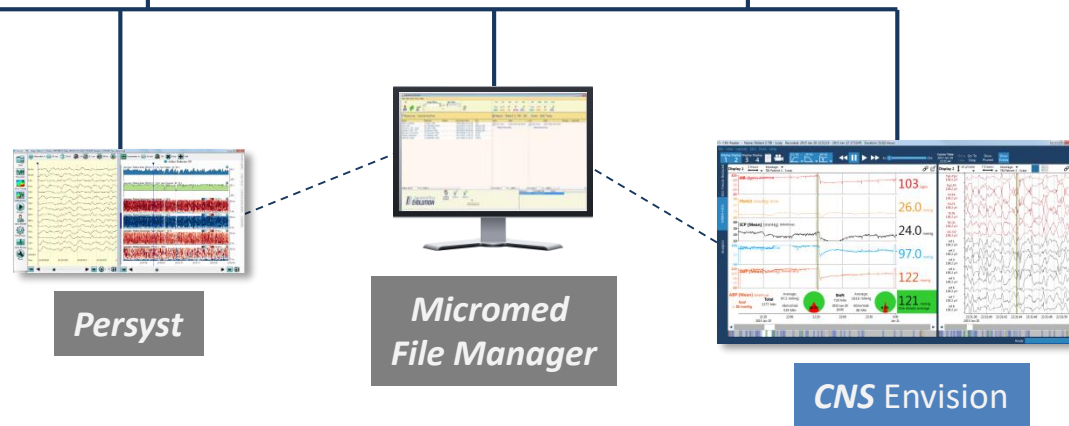
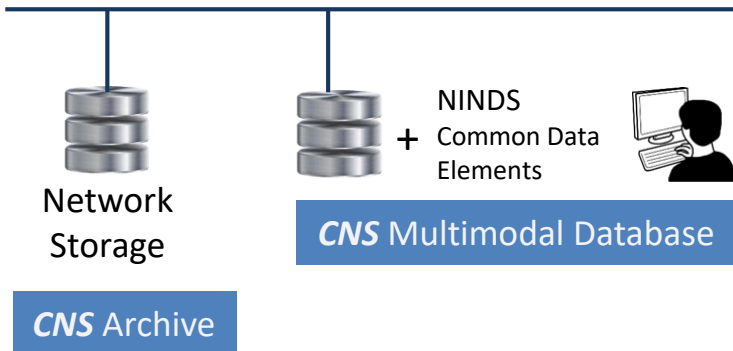
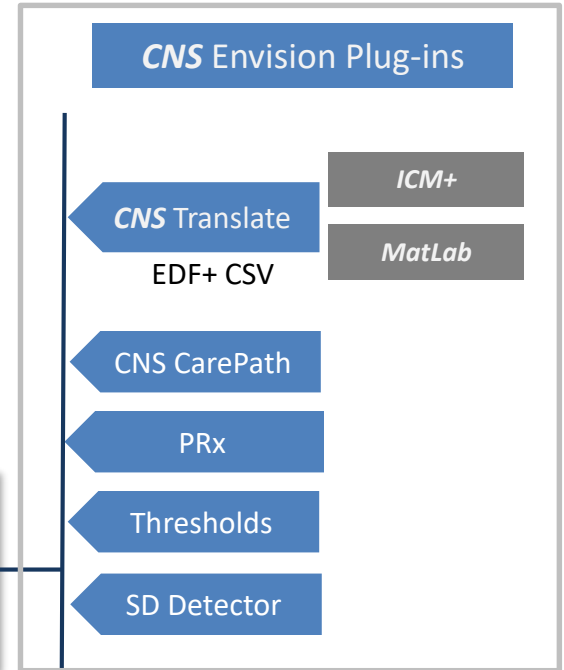
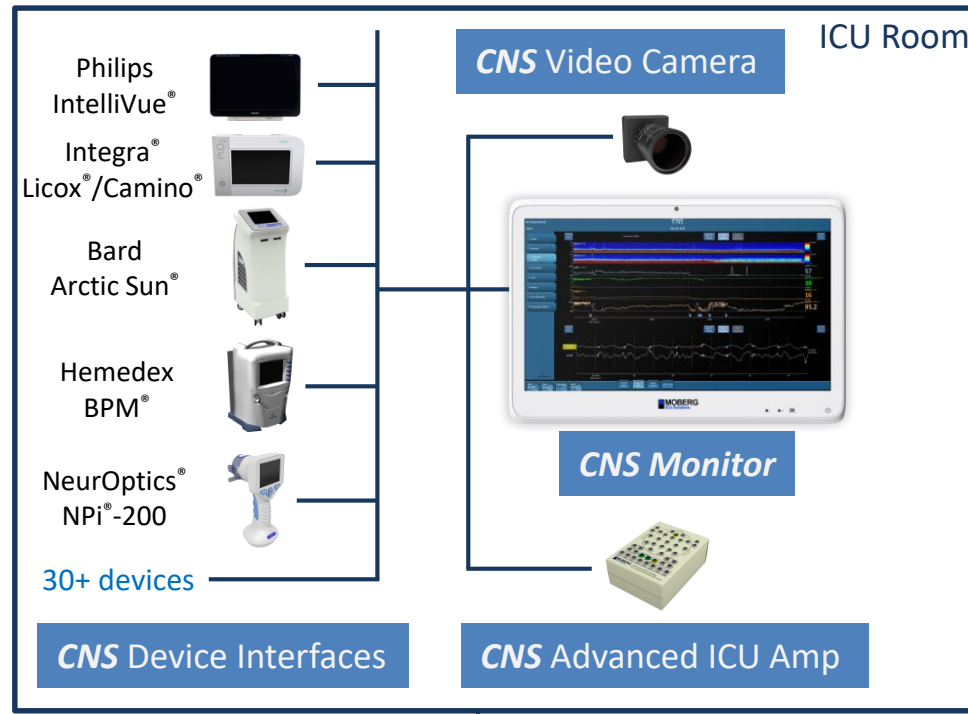
# Current Data Management Tools

## The Component Neuromonitoring System

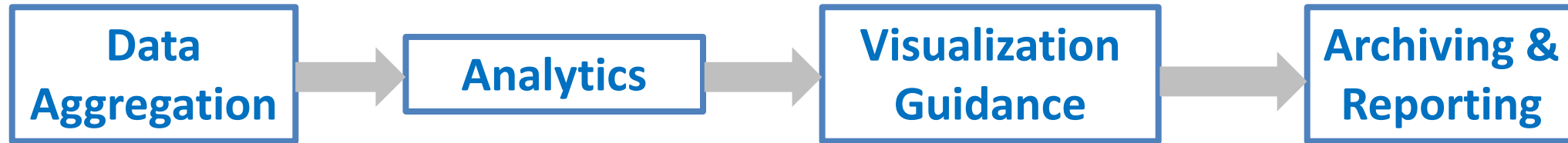
A system of components for managing data in neurocritical care

### Legend

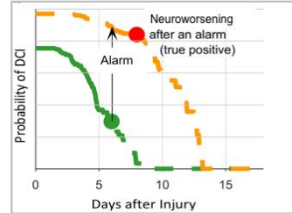
- Moberg Product
- Third Party Product



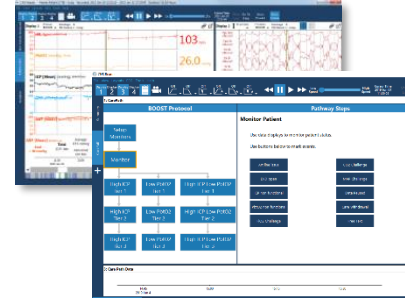
# Group the Technologies



CNS Monitor



Apps



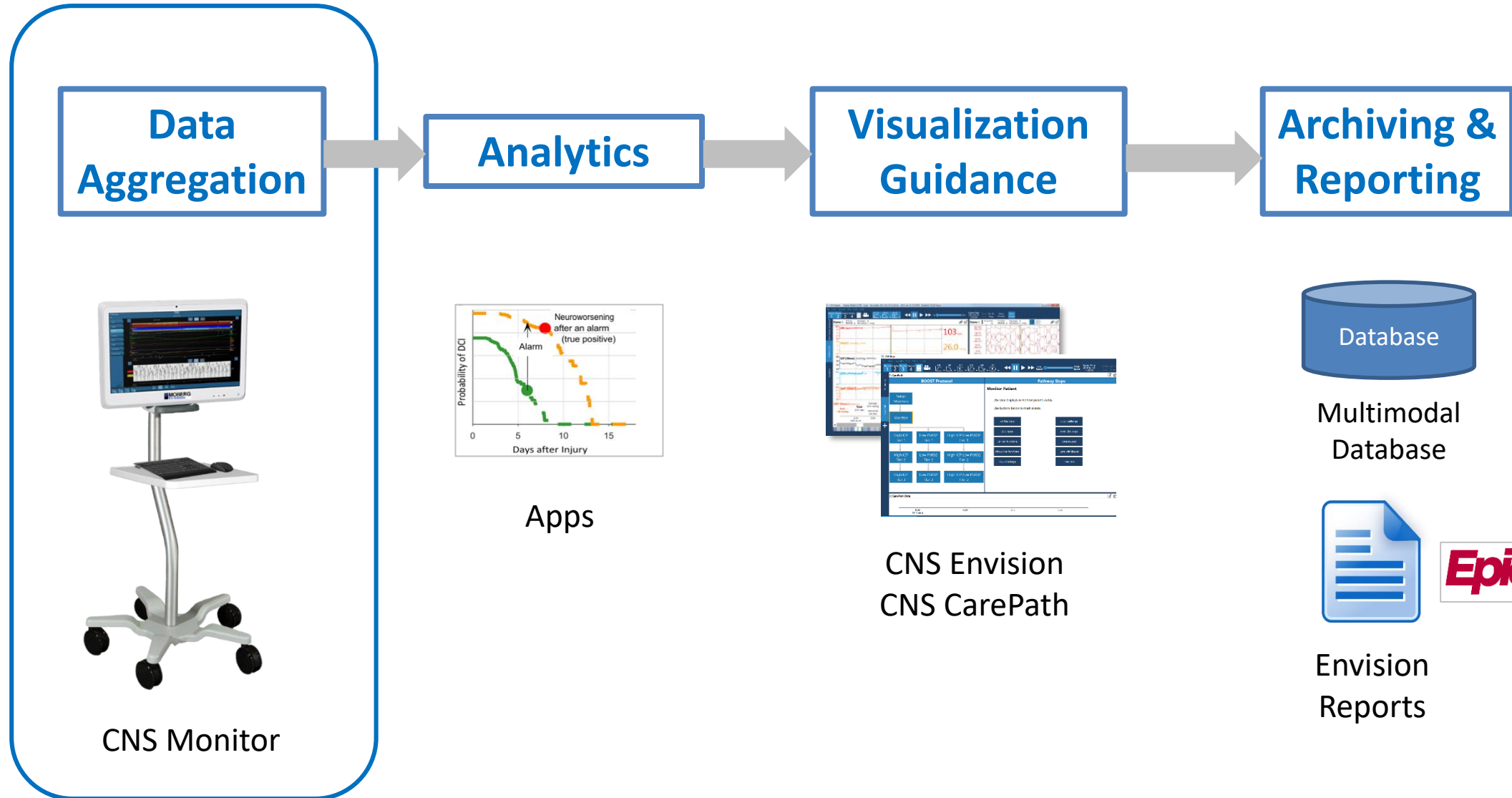
CNS Envision  
CNS CarePath



Database  
Multimodal  
Database

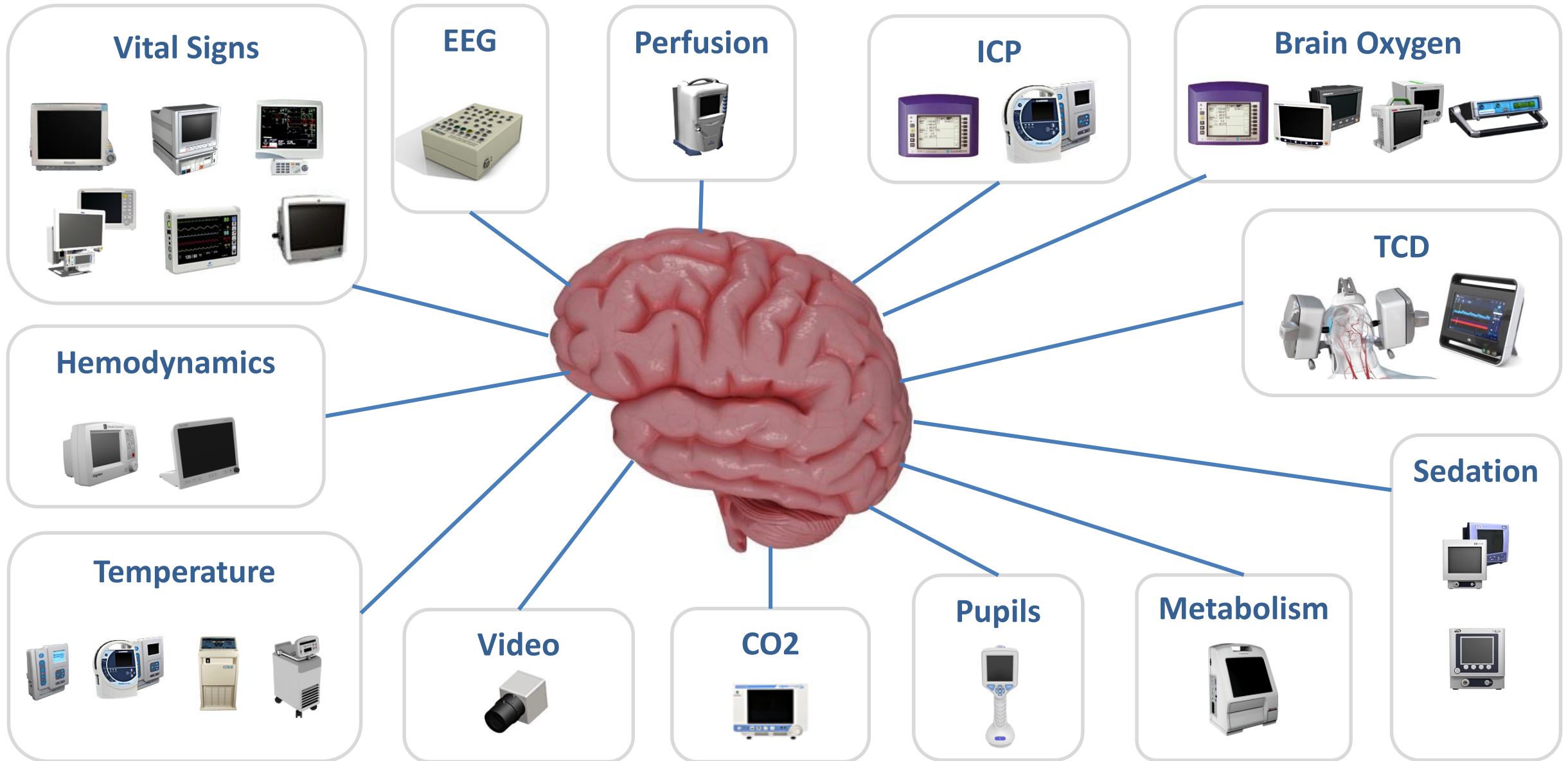


Envision  
Reports





# Strength - Device Data Collection

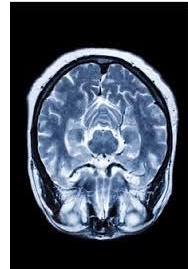


# Opportunity - Expanding Data Aggregation

Time synchronized  
physiological data



Imaging  
descriptors



Phenotypic  
data



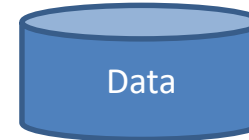
Databases  
Common Data Elements

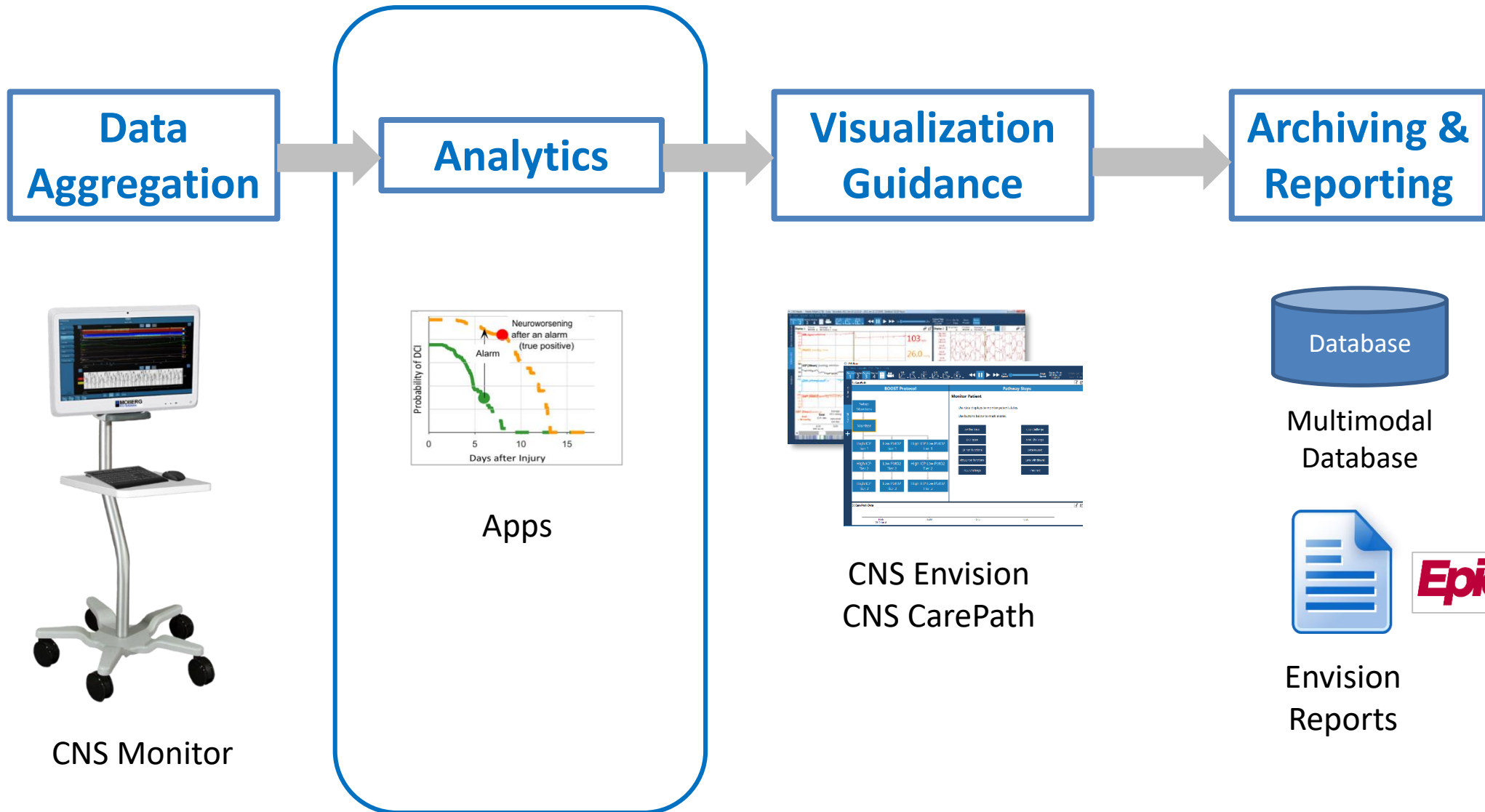


Supported by a grant from  
the Department of Defense



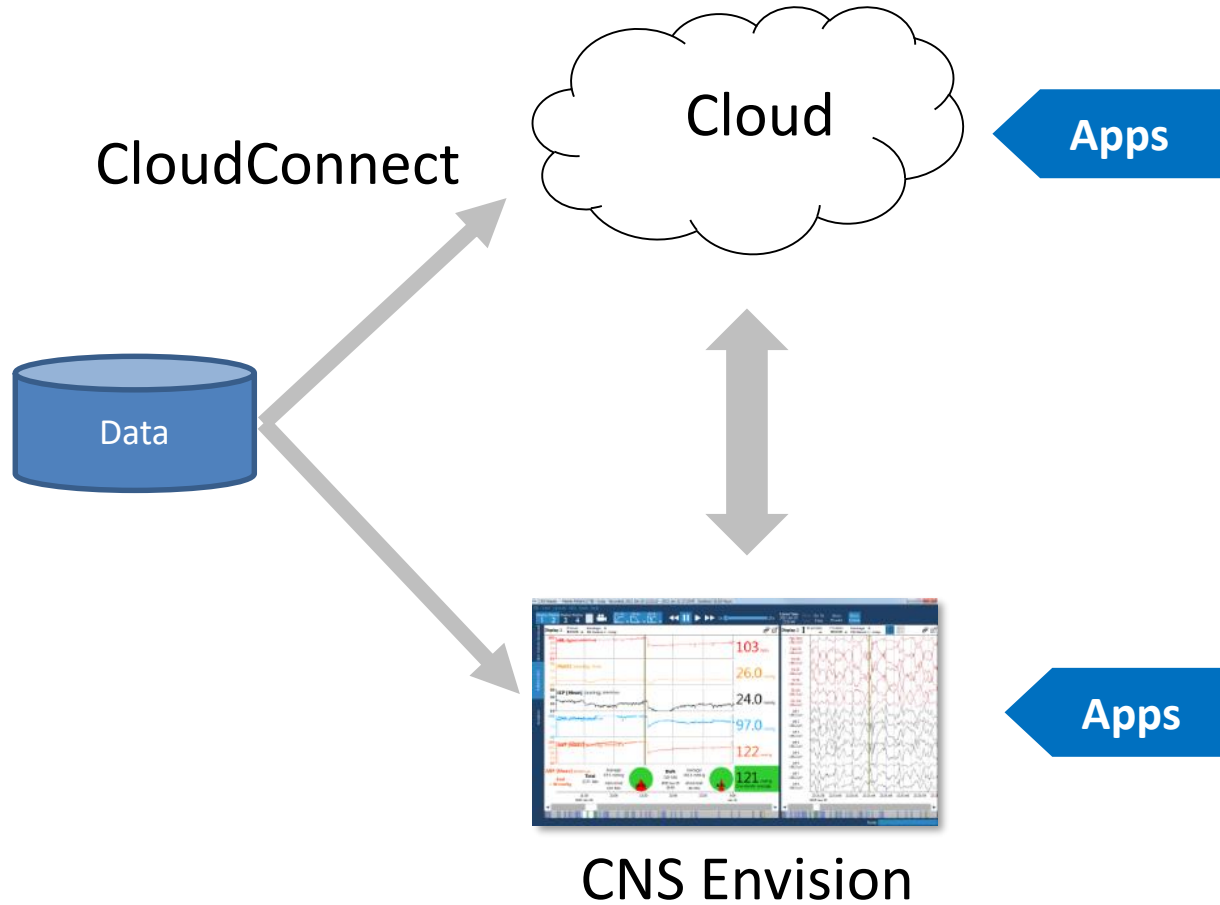
Standards  
Nomenclature  
Archiving formats  
Annotations





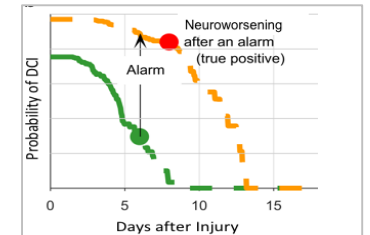
# Opportunity – Informatics/Analytics

Prediction and detection algorithms based on  
real-time and retrospective data

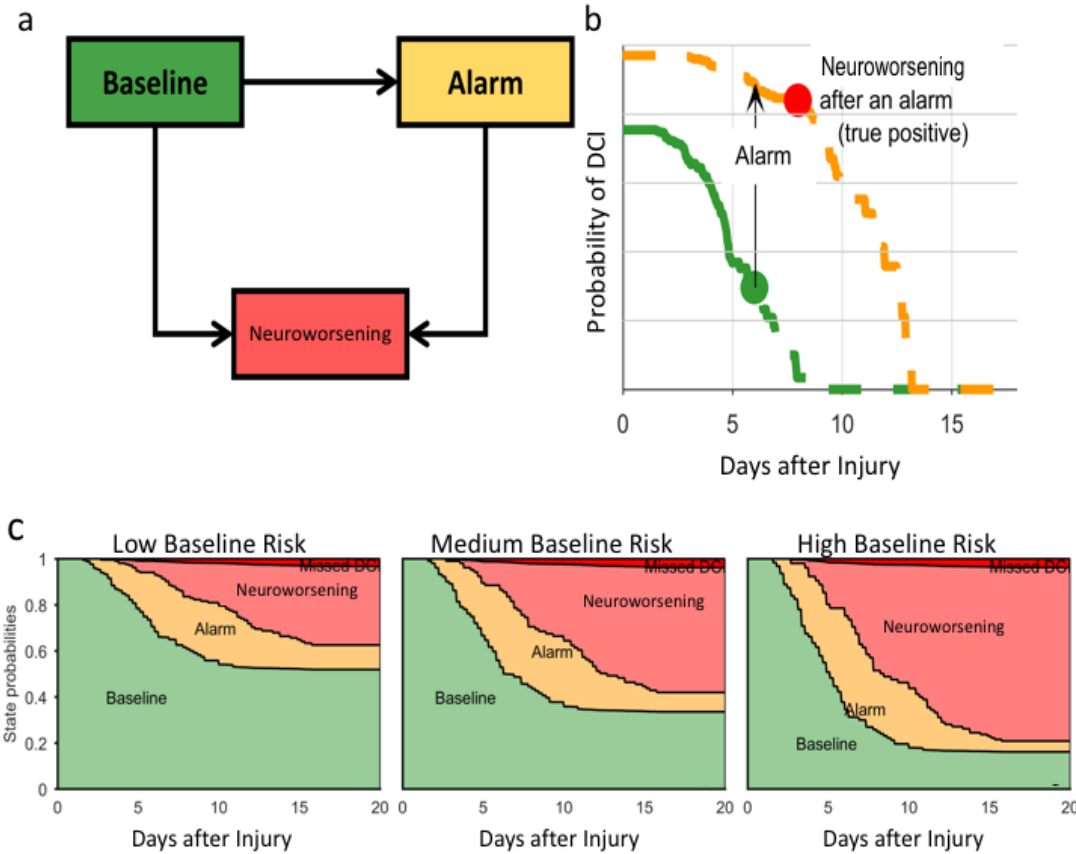


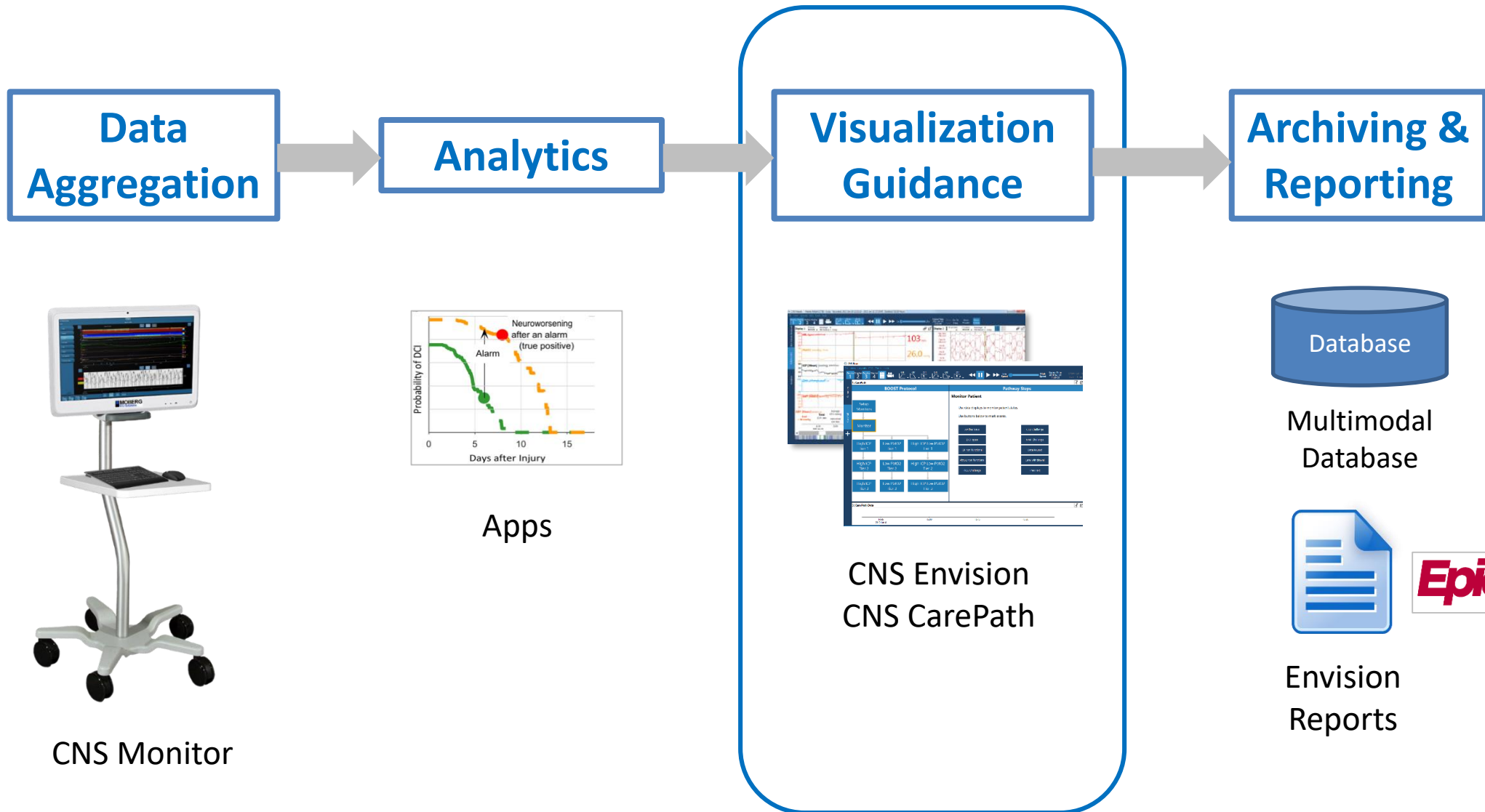
## Apps

- Data cleaning
- Event detection
- Prediction
- Classification
- Machine learning
- Data mining



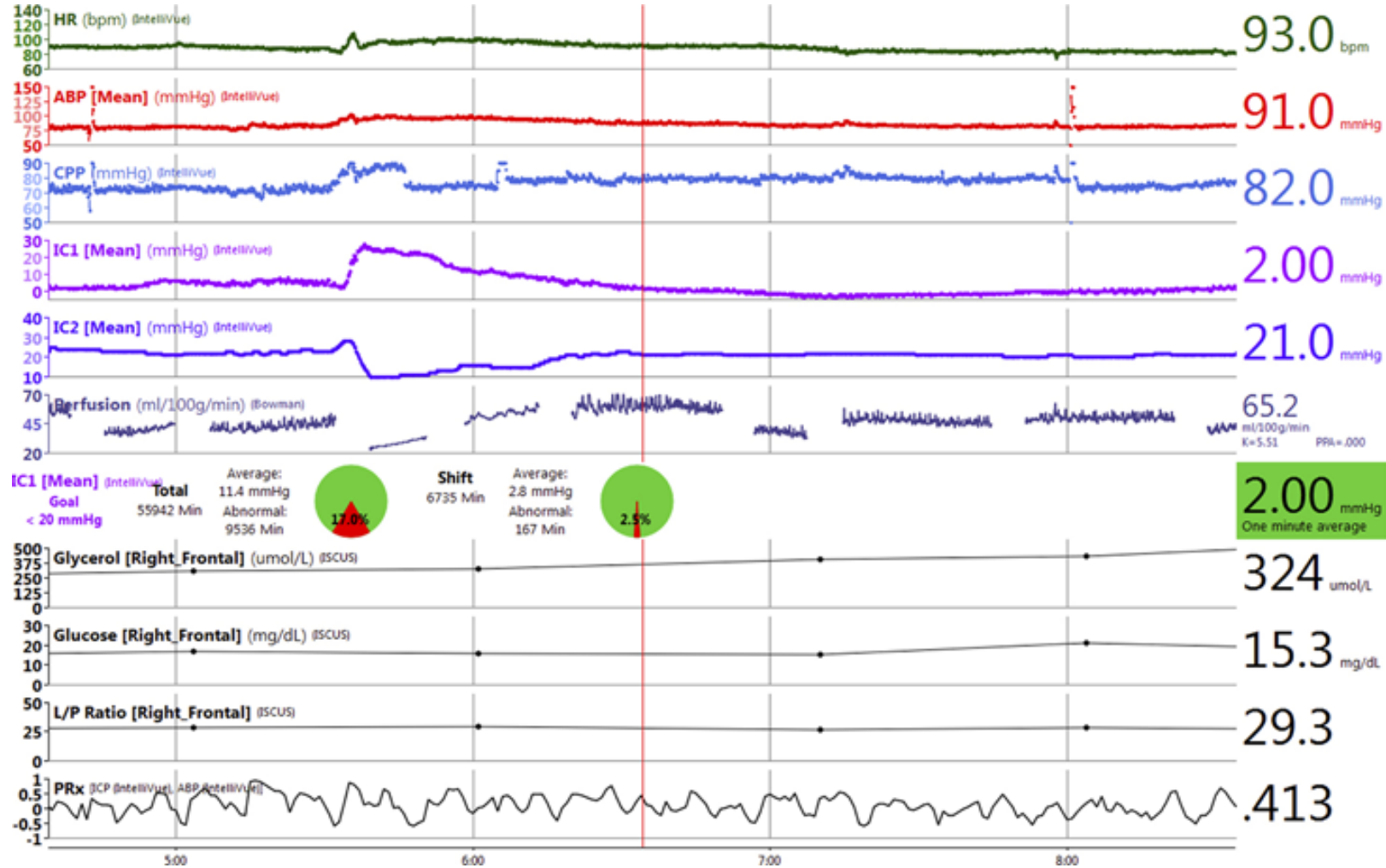
## Multi-state Predictive Models of Neuroworsening





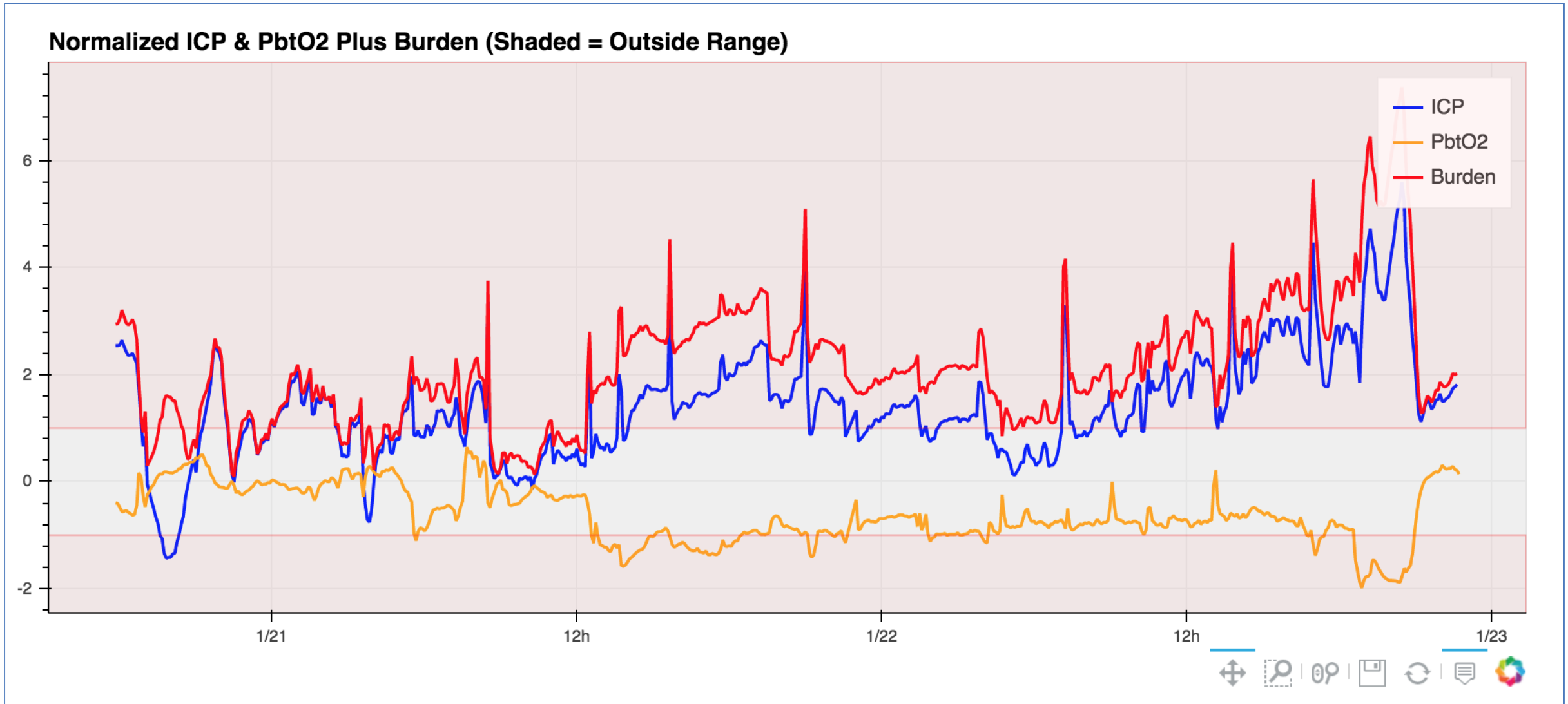
# Visualization

## Medical Record Using Comprehensive Data



Dr. Ramani Balu  
Univ. of Pennsylvania

# Threat Mitigation - Burden Scores





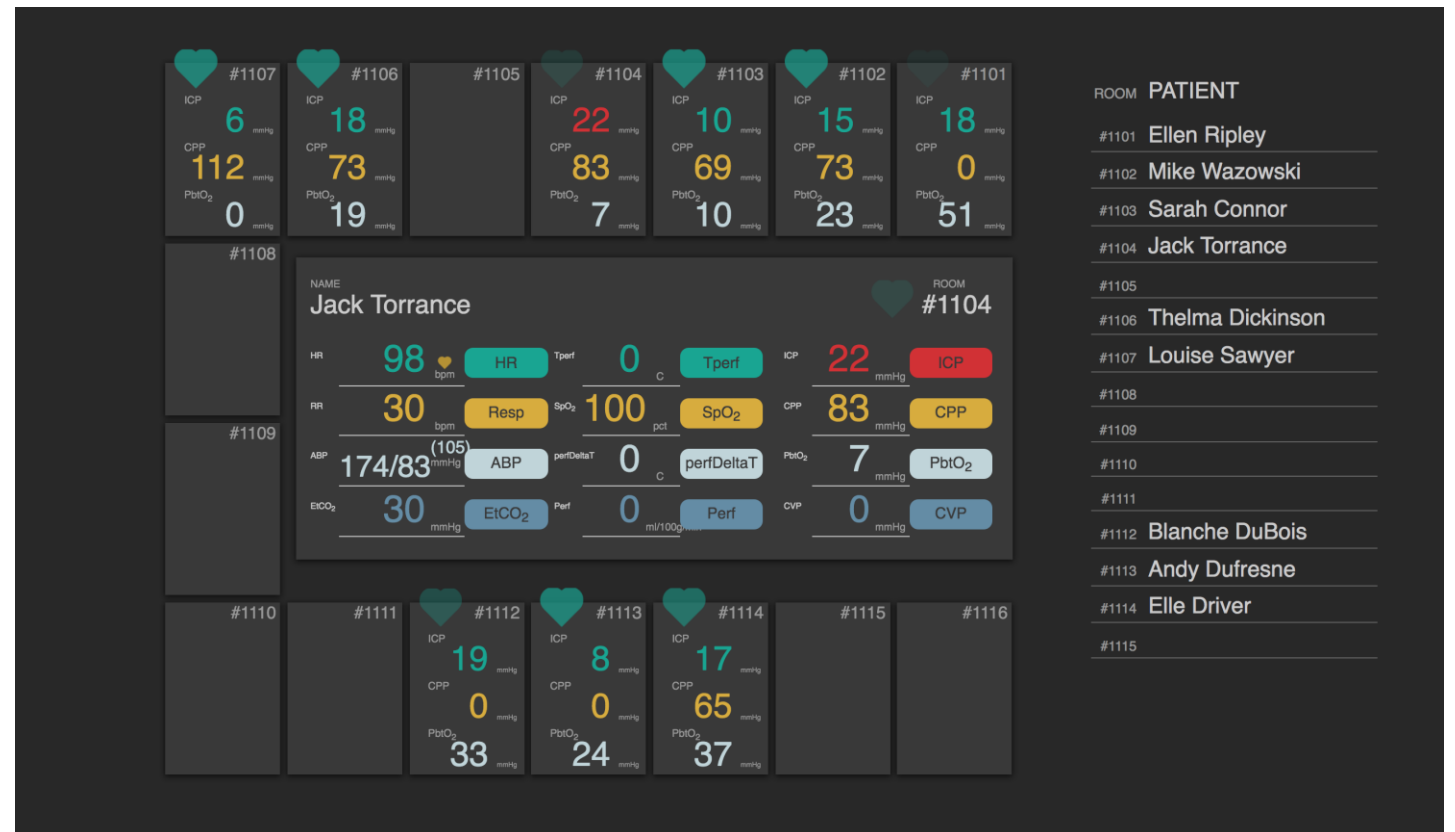
# Example Dashboard

## The Problem:

- 72 bed pediatric ICU
- 12 of those patients have neuro issues
- Pediatric neurologist has to individually click into Epic for each patient to get the status

## The Solution

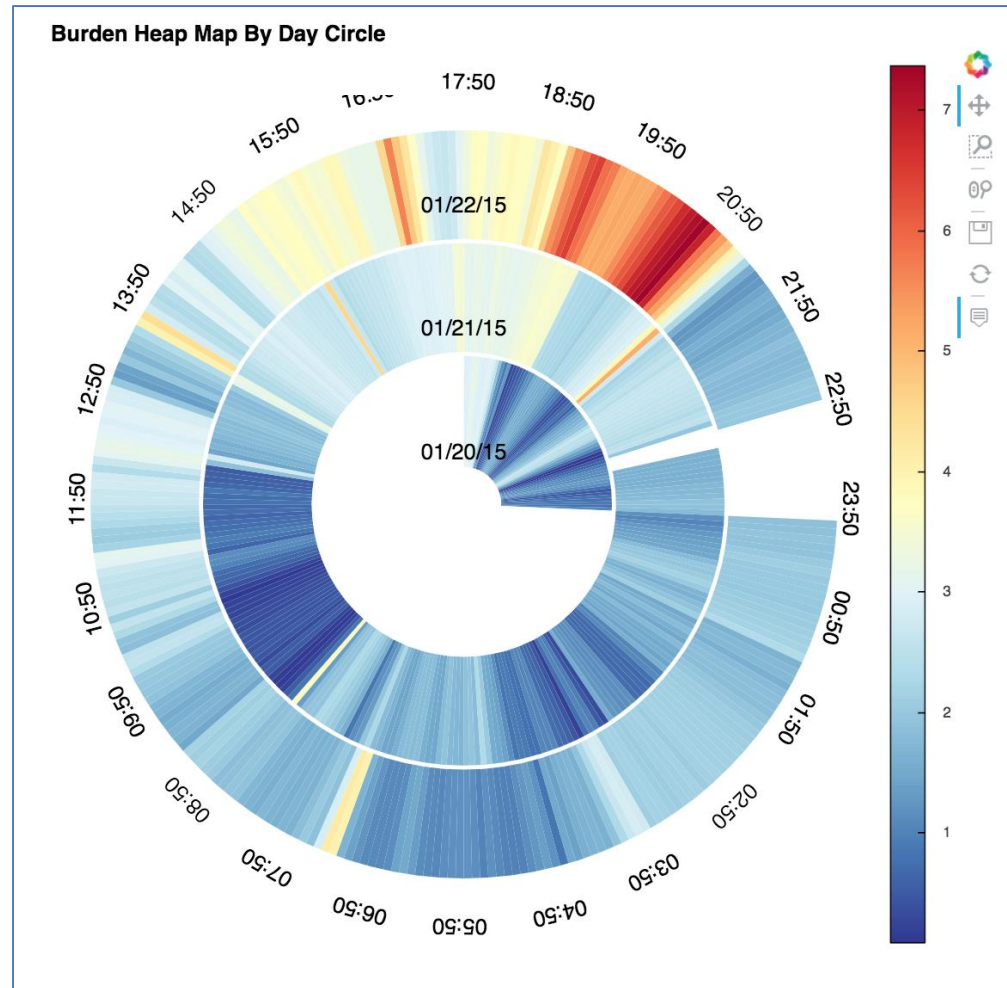
- Separate neuro dashboard for subset of patients
- Connects to Epic for some data
- Bypasses Epic for patient review



Analytics determine patient state. Circles (size and color) show those requiring urgent care.



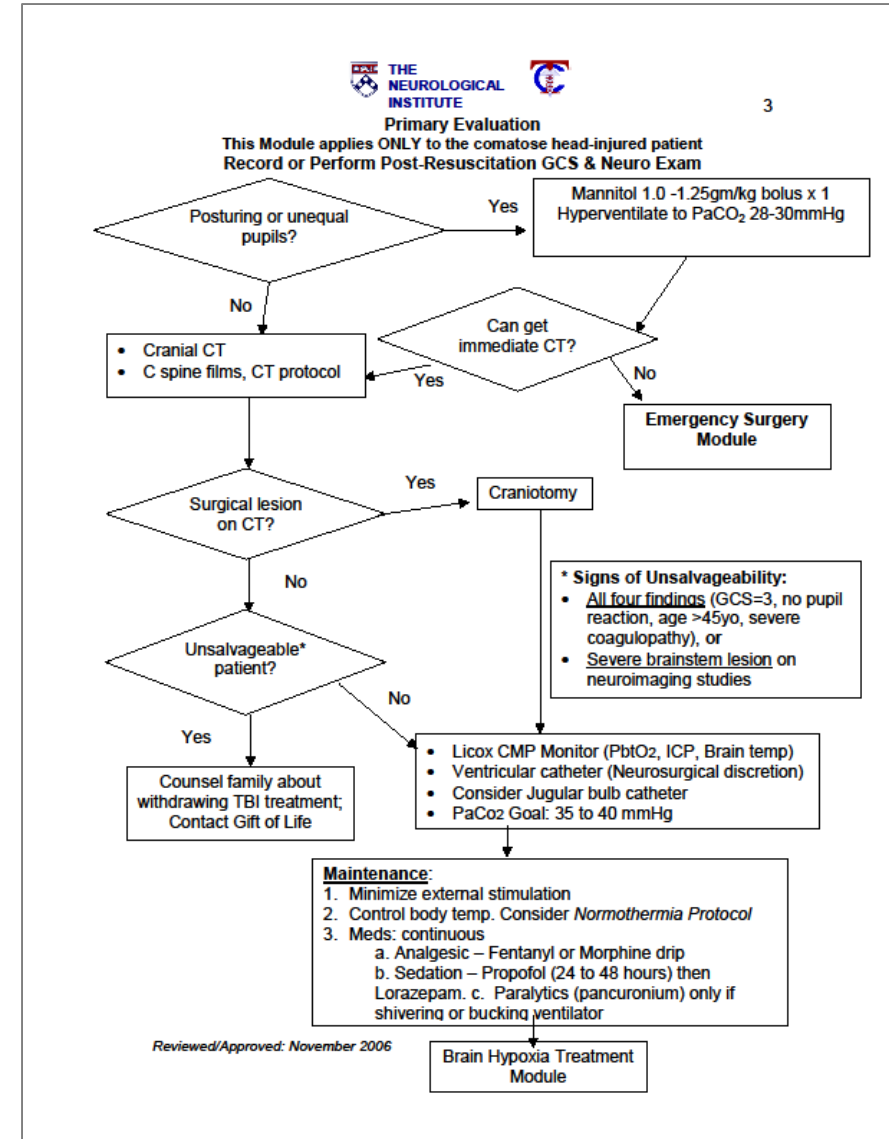
Radial heat map showing daily correlations of ICP burden.



## Clinical Pathway

### Problems Addressed by CarePath

- Compliance to Guidelines/Pathways
- Consistency of Care across Providers
- Continuous Learning in the ICU
- Compiling a Record of Actions



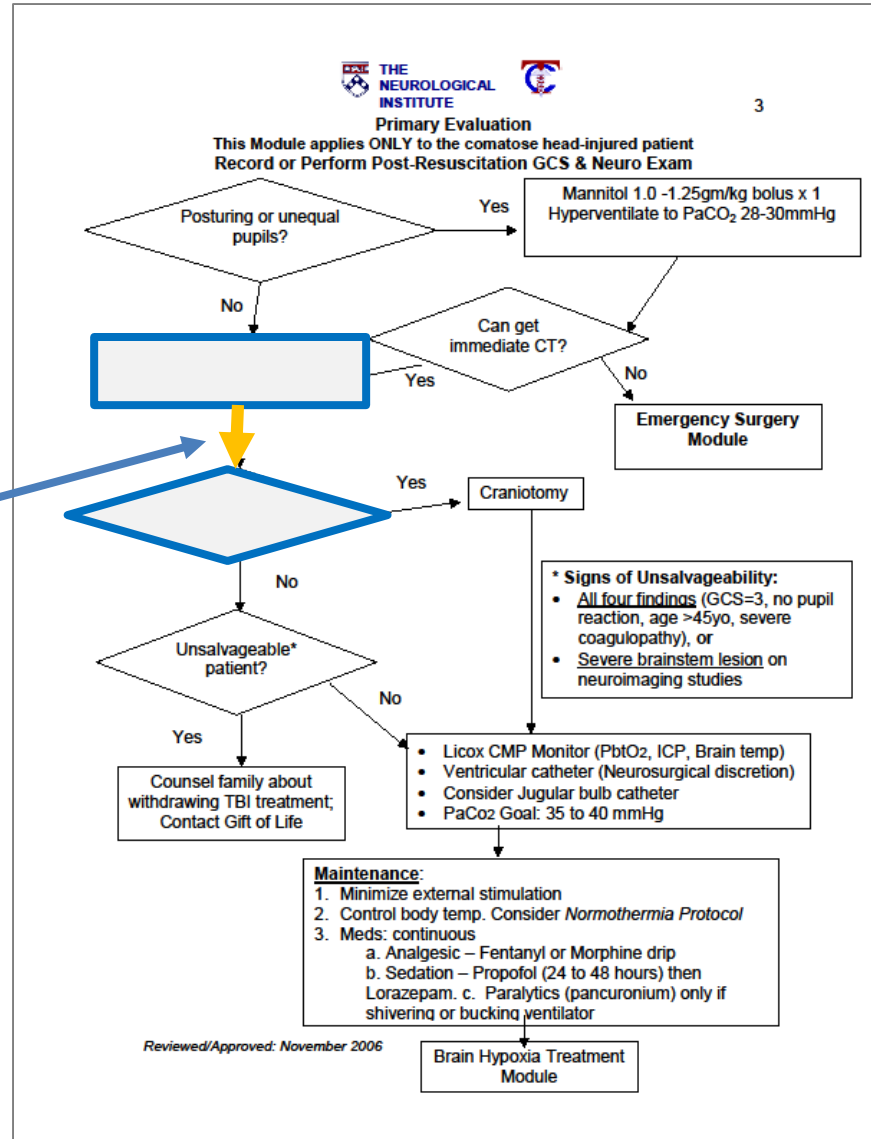
# What Does CarePath Do?

CarePath drives you through a clinical pathway based on changes in a patient's "state".



Patient State

## Clinical Pathway



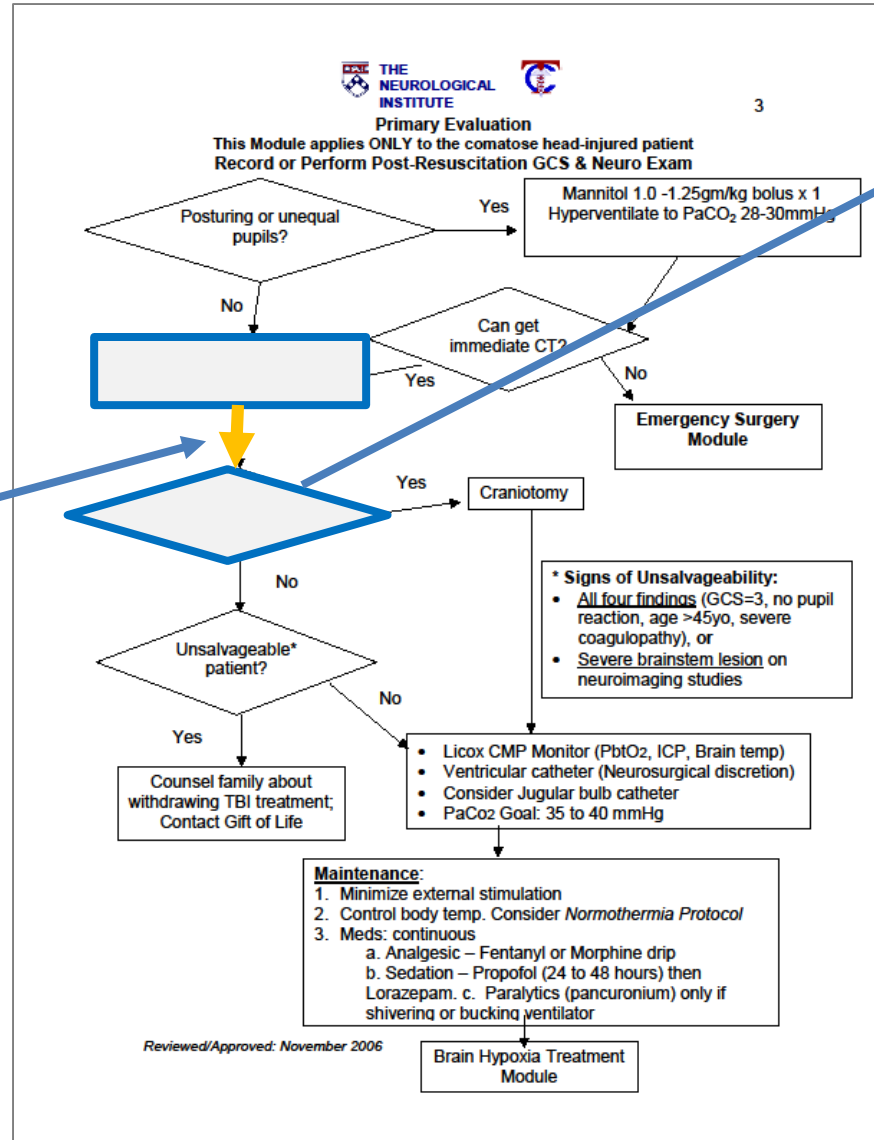
# What Does CarePath Do?

CarePath drives you through a clinical pathway based on changes in a patient's "state".



Patient State

## Clinical Pathway



It provides instructions along the way

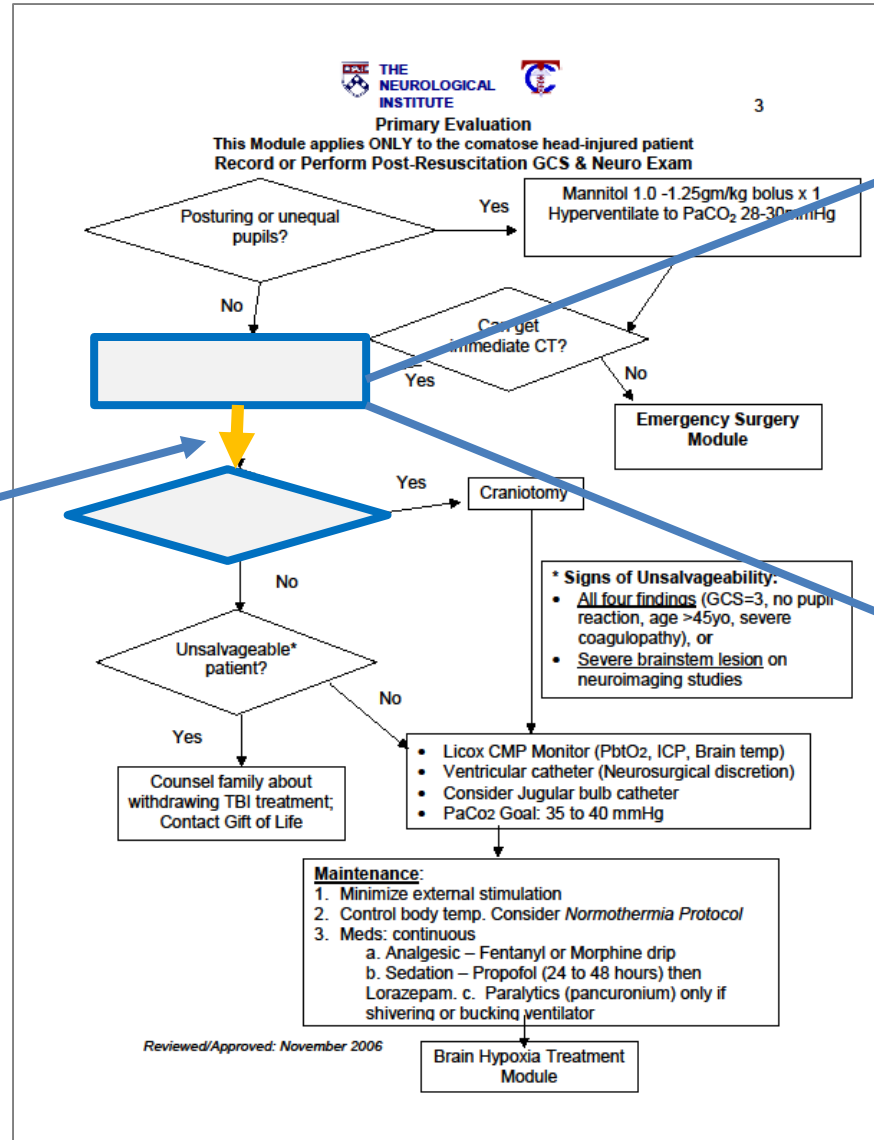
# What Does CarePath Do?

CarePath drives you through a clinical pathway based on changes in a patient's "state".



Patient State

## Clinical Pathway



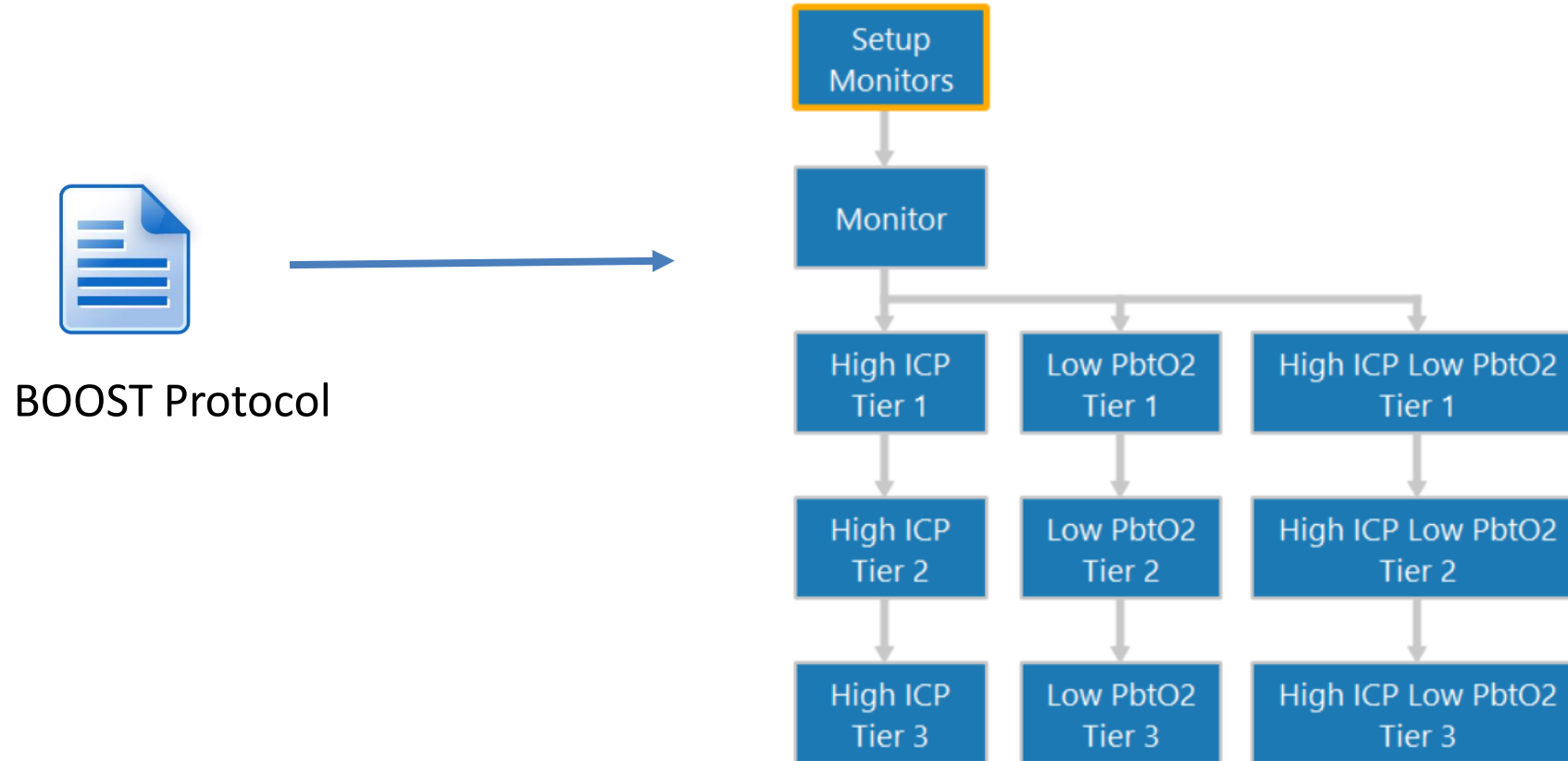
It provides instructions along the way

It provides annotations on what was done

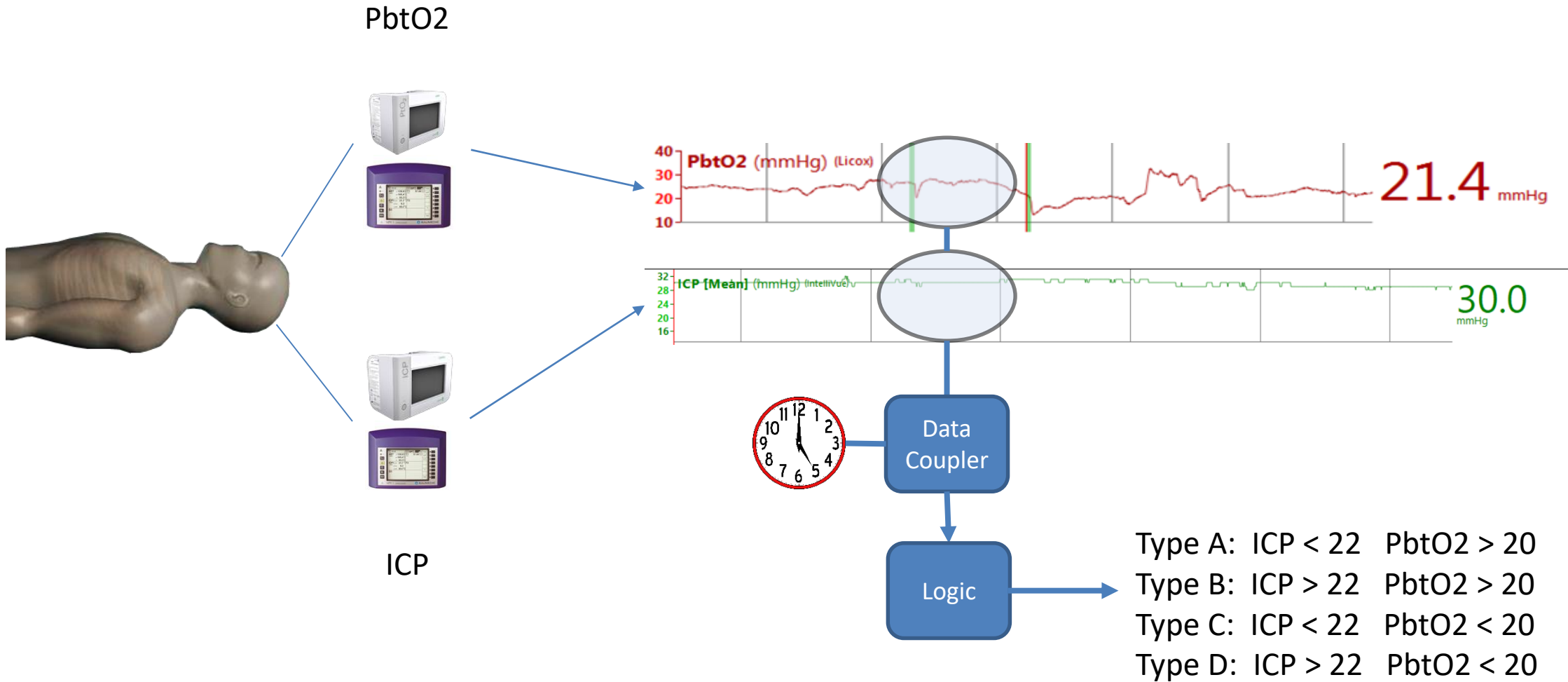
# CarePath and BOOST3



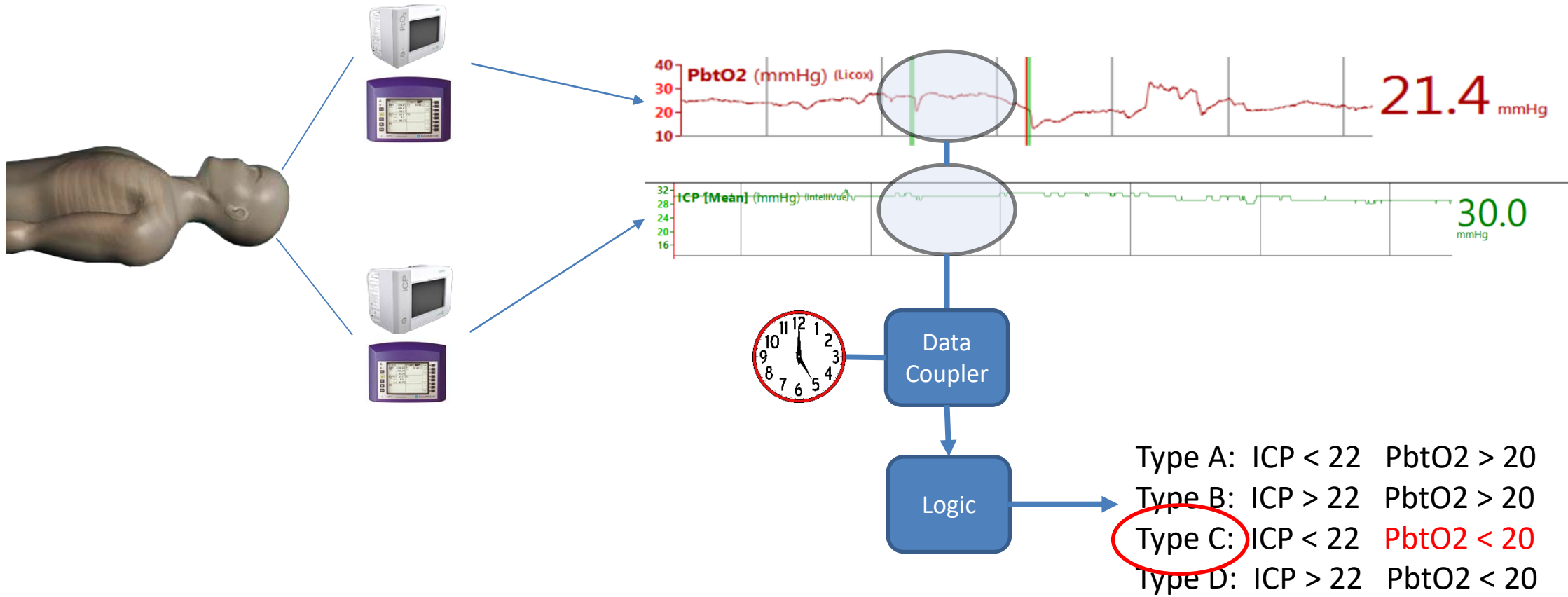
## BOOST3 Clinical Pathway



# CarePath: Detecting a Patient State



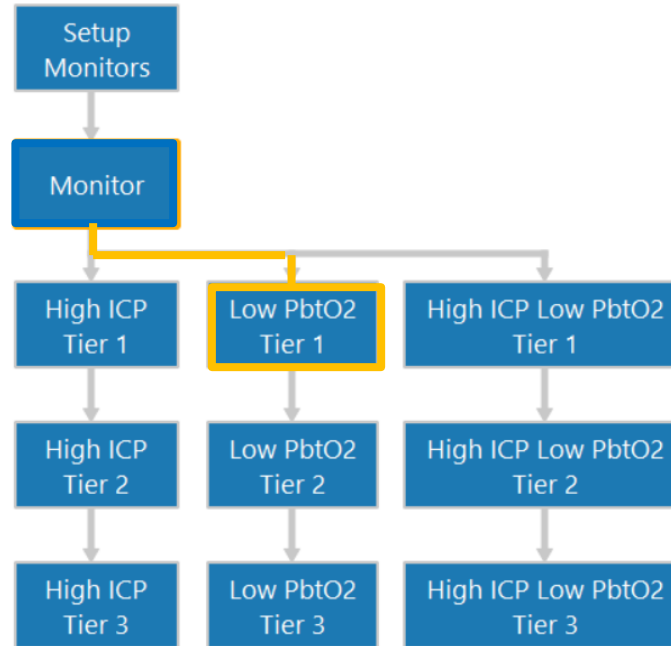
# CarePath: Detecting a Patient State



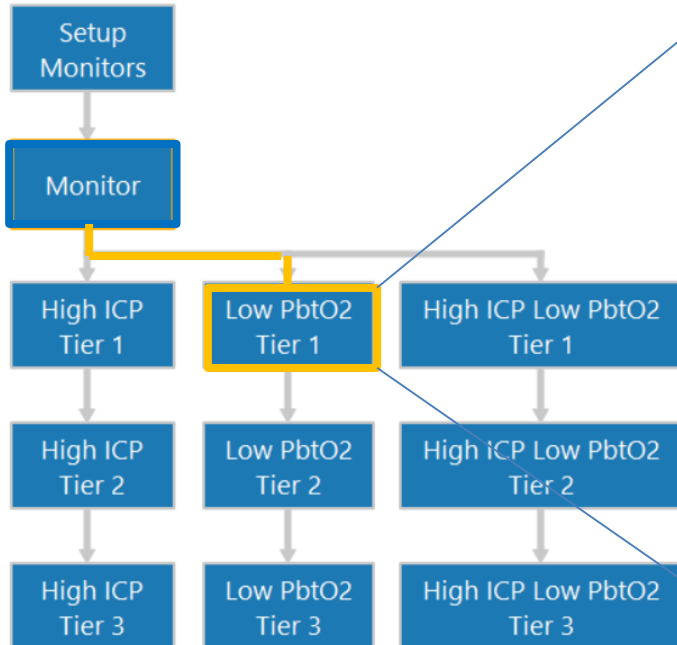
# Summary Display

The screenshot displays the CNS Reader software interface. At the top, there is a menu bar with 'File', 'View', 'Layouts', 'EEG', 'Tools', and 'Help'. Below the menu is a toolbar with various icons for display, recording, and playback. A speed control slider is set between 'Low Speed' and 'High Speed'. The 'Cursor Time' is shown as 2019 Mar 17 09:30:02. On the left, a vertical sidebar contains 'Summary' and 'CarePath' sections, with a '+' icon below 'CarePath'. The main area is divided into two panels. The top panel, titled '1: Summary', contains the following information: 'BOOST PROTOCOL', 'Goal: ICP  $\leq$  22 mmHg, PbtO2  $\geq$  20 mmHg', 'Episode PbtO2 < 20 mmHg, 5 min', and a red box labeled 'Intervention Required'. The bottom panel, titled '4: Trends', shows a 2x4 grid of trend plots for 'ICP [Mean] (mmHg) (IntelliVue)', 'PbtO2 (mmHg) (Licox)', 'ABP [Mean] (mmHg) (IntelliVue)', and 'Tcore ( $^{\circ}$ C) (IntelliVue)'. Each plot has a y-axis and a time axis from 13:30 to 14:30 on 2019 Mar 17. The plots are currently empty, and each has a question mark icon to its right. On the right side, there is an 'Events / Annotations' panel with a 'Filter by:' field and a table with columns for 'Time' and 'Event / Annotation'.

## BOOST3 Clinical Pathway



From the BOOST Protocol

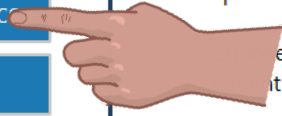


**1: CarePath**

Tier 1 Steps to Raise PbtO2: Select in any order	Pathway Steps
<div style="display: flex; flex-direction: column; gap: 10px;"> <div style="border: 2px solid orange; padding: 5px; display: inline-block;">Tier 1</div> <div style="border: 1px solid gray; border-radius: 15px; padding: 5px; display: inline-block;">Go To Tier 2</div> <div style="border: 1px solid gray; border-radius: 15px; padding: 5px; display: inline-block;">RETURN</div> </div>	<div style="background-color: #0056b3; color: white; padding: 5px; text-align: center; font-weight: bold;">Tier 1 Raise PbtO2</div> <p style="font-size: small; margin-top: 10px;">Treatment must be initiated within 15 minutes of the start of the episode, as detected by the continuous ICP recording.</p> <p style="font-size: small; margin-top: 10px;">Choose therapies from the list in any order, based on individual patient characteristics and local protocols.</p> <p style="font-size: small; margin-top: 10px;">The bedside treatment team has the option to progress to higher tiers as rapidly as they feel is clinically indicated.</p> <p style="font-size: small; margin-top: 10px;">Move to Tier 2 therapies (press button) if desired and at least one intervention from Tier 1 has been used.</p>
<div style="display: flex; flex-direction: column; gap: 5px;"> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Adjust Head of Bed</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Active Cooling</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Optimize CPP</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Optimize Hemodynamics</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Adjust FiO2</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Adjust PEEP</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Pulmonary Toileting</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Adjust Cardiac Output</div> <div style="background-color: #0056b3; color: white; padding: 5px; text-align: center;">Seizure Prophylaxis</div> </div>	

1: CarePath ✎ ↗

Tier 1 Steps to Raise PbtO2: Select in any order	Pathway Steps
<div data-bbox="122 514 343 578" style="border: 2px solid orange; padding: 5px; margin-bottom: 10px;">Tier 1</div> <div data-bbox="122 656 343 721" style="border-radius: 15px; padding: 5px; margin-bottom: 10px;">Go To Tier 2</div> <div data-bbox="122 799 343 863" style="border-radius: 15px; padding: 5px;">RETURN</div> <div data-bbox="407 514 777 578" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Adjust Head of Bed</div> <div data-bbox="407 585 777 649" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Active Cooling</div> <div data-bbox="407 656 777 721" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Optimize CPP</div> <div data-bbox="407 728 777 792" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Optimize Hemodynamics</div> <div data-bbox="407 799 777 863" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Adjust FiO2</div> <div data-bbox="407 871 777 935" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Adjust PEEP</div> <div data-bbox="407 942 777 1006" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Pulmonary Toileting</div> <div data-bbox="407 1013 777 1078" style="border: 1px solid gray; padding: 5px; margin-bottom: 5px;">Adjust Cardiac Output</div> <div data-bbox="407 1085 777 1149" style="border: 1px solid gray; padding: 5px;">Seizure Prophylaxis</div>	<h3 data-bbox="840 471 1108 506">Tier 1 Raise PbtO2</h3> <p data-bbox="930 692 1579 756">Treatment must be initiated within 15 minutes of the start of the episode, as detected by the continuous ICP recording.</p> <p data-bbox="980 785 1579 849">Use the therapies from the list in any order, based on individual patient characteristics and local protocols.</p> <p data-bbox="930 878 1579 942">The bedside treatment team has the option to progress to higher tiers as rapidly as they feel is clinically indicated.</p> <p data-bbox="930 971 1579 1035">Move to Tier 2 therapies (press button) if desired and at least one intervention from Tier 1 has been used.</p>



1: CarePath



**Tier 1 Steps to Raise PbtO2: Select in any order**

Tier 1

Adjust Head of Bed

Active Cooling

Go To Tier 2

Optimize CPP

Optimize Hemodynamics

RETURN

Adjust FiO2

Adjust PEEP

Pulmonary Toileting

Adjust Cardiac Output

Seizure Prophylaxis

**Pathway Steps**

**Optimize Hemodynamics**

Optimize hemodynamics through any of these options:

- Resuscitation: Address hypovolemia to achieve clinical euvolemia with volume augmentation per local protocol.
- Diuresis: Avoid hypervolemia, consider furosemide or other agent for diuresis.

Press button if you optimized hemodynamics during this step.

Volume Augmentation

Diuretics

From the  
BOOST Protocol



1: CarePath



Tier 1 Steps to Raise PbtO2: Select in any order

Tier 1

Adjust Head of Bed

Active Cooling

Go To Tier 2

Optimize CPP

Optimize Hemodynamics

RETURN

Adjust FiO2

Adjust PEEP

Pulmonary Toileting

Adjust Cardiac Output

Seizure Prophylaxis

Pathway Steps

Optimize Hemodynamics

Optimize hemodynamics through any of these options:

- Resuscitation: Address hypovolemia to achieve clinical euvolemia with volume augmentation per local protocol.
- Diuresis: Avoid hypervolemia, consider furosemide or other agent for diuresis.

Press button if you optimized hemodynamics during this step.

Volume Augmentation

Diuretics



1: CarePath



Events / Annotations

Filter by:

Time ^	Event / Annotation ^
2019 Mar 14 13:59:39	[CP] Volume augmentation

Tier 1 Steps to Raise PbtO2: Select in any order

Tier 1

Adjust Head of Bed

Go To Tier 2

Active Cooling

Optimize CPP

Optimize Hemodynamics

RETURN

Adjust FiO2

Adjust PEEP

Pulmonary Toileting

Adjust Cardiac Output

Seizure Prophylaxis

Pathway Steps

Optimize Hemodynamics

Optimize hemodynamics through any of these options:

- Resuscitation: Address hypovolemia to achieve clinical euvolemia with volume augmentation per local protocol.
- Diuresis: Avoid hypervolemia, consider furosemide or other agent for diuresis.

Press button if you optimized hemodynamics during this step.

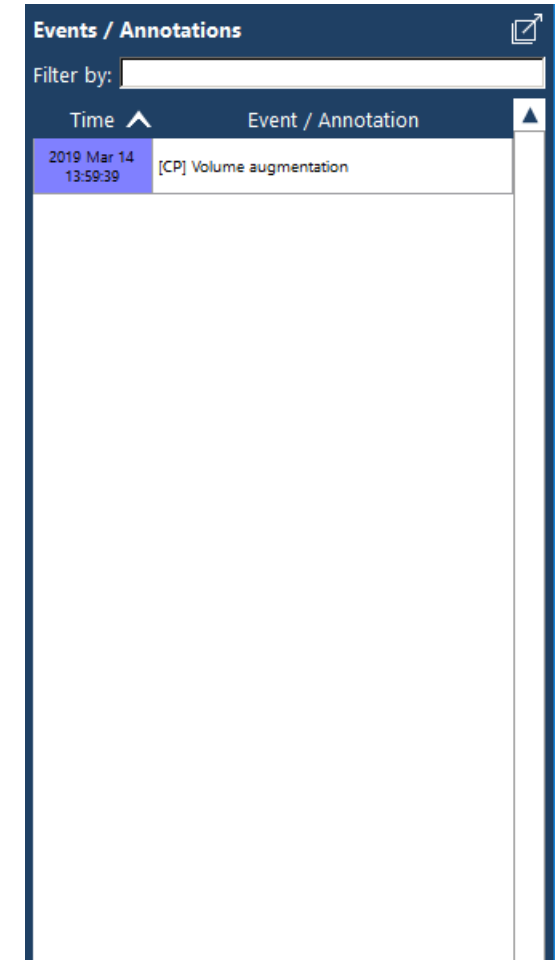
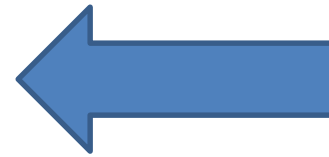
Volume Augmentation

Diuretics

A “high resolution” nursing record

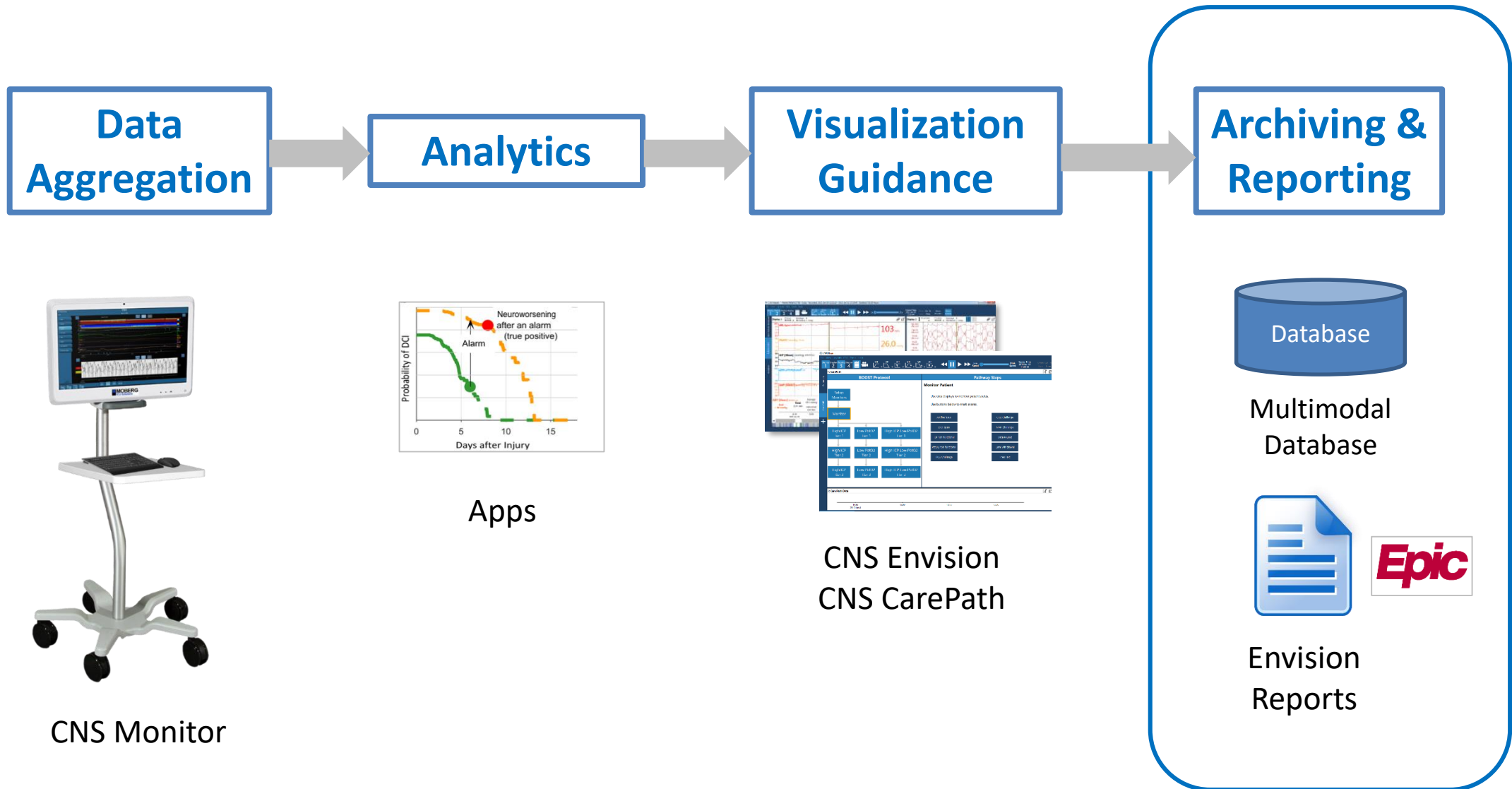


Machine learning  
Clinical trials

A screenshot of a software interface titled "Events / Annotations". It features a "Filter by:" search bar at the top. Below it is a table with two columns: "Time" and "Event / Annotation". The "Time" column has a small upward-pointing arrow next to it. The table contains one row of data: "2019 Mar 14 13:59:39" in the Time column and "[CP] Volume augmentation" in the Event / Annotation column. There is also a small upward-pointing arrow in the top right corner of the table area.

Time	Event / Annotation
2019 Mar 14 13:59:39	[CP] Volume augmentation

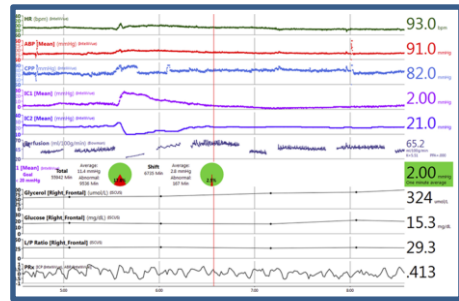
**Compliance** to Guidelines/Pathways  
**Consistency** of Care across Providers  
**Continuous Learning** in the ICU  
Compiling a **Record of Actions**



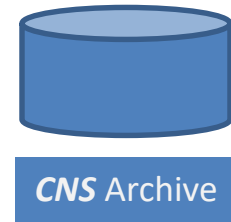
# Threat Mitigation - Multimodal Database



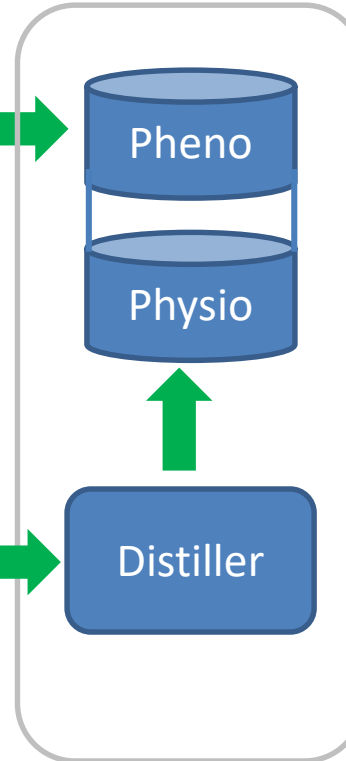
Phenotypic data  
In NINDS Common Data  
Elements format



Collect Comprehensive  
Data from CNS Monitor



High Resolution  
Data from Multiple  
Patients



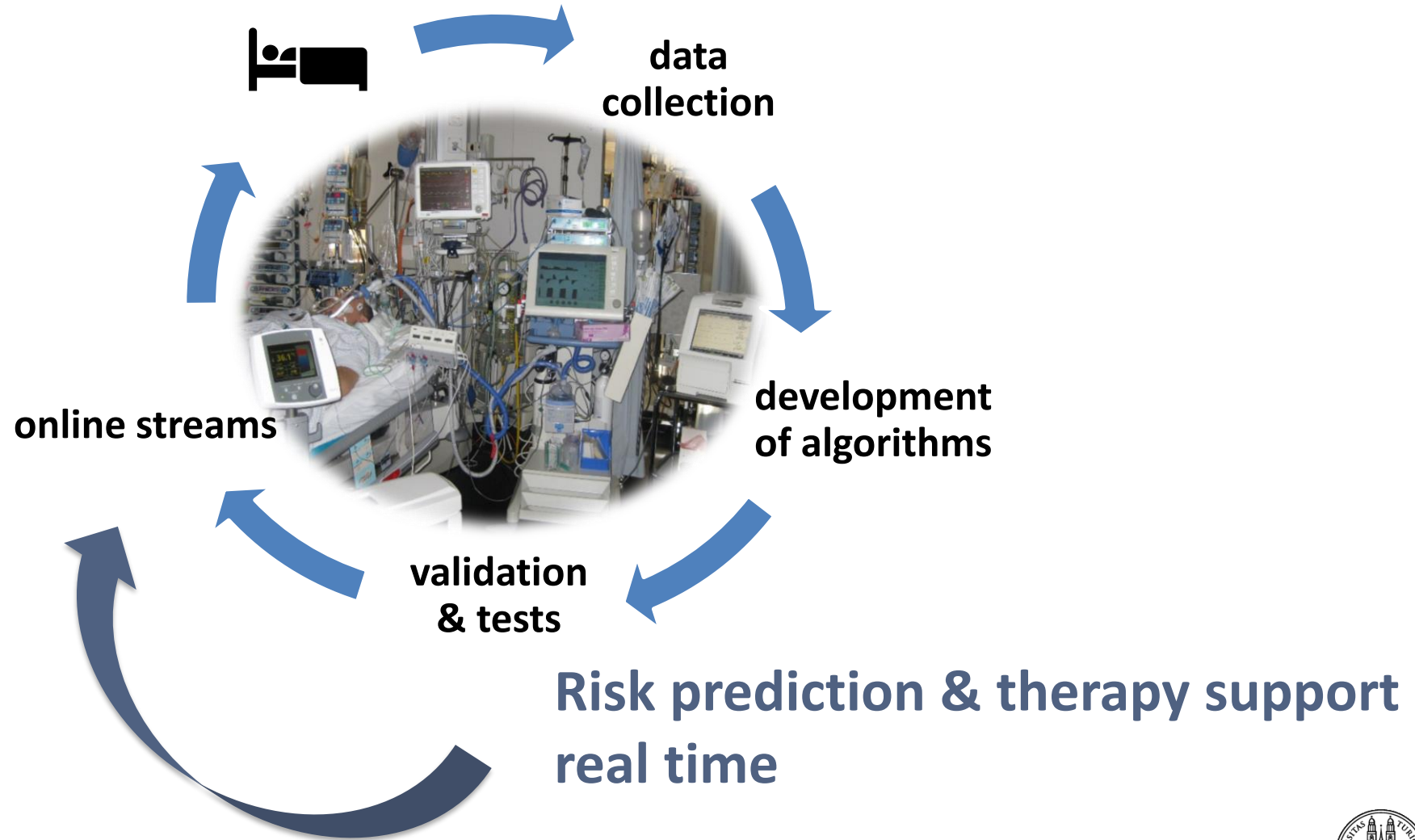
CNS Multimodal  
Database



Powerful Queries

- Describe the **need** for comprehensive, high-resolution data in critical care
  - We will use the care of acute brain injury as an example
- Describe the **challenges** of collecting data in a usable form
  - We will describe the state of medical device connectivity and data interoperability
- Describe **progress** to overcome the challenges
  - We will use our work (and others) as examples
- Describe the **future** with “Smart ICUs”
  - We will show examples of what can be done

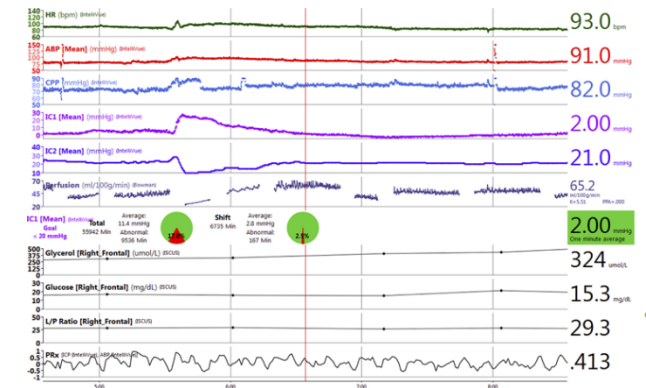
# Next steps





- Project for Precision Management of TBI
- Multi-center Trial: BOOST3

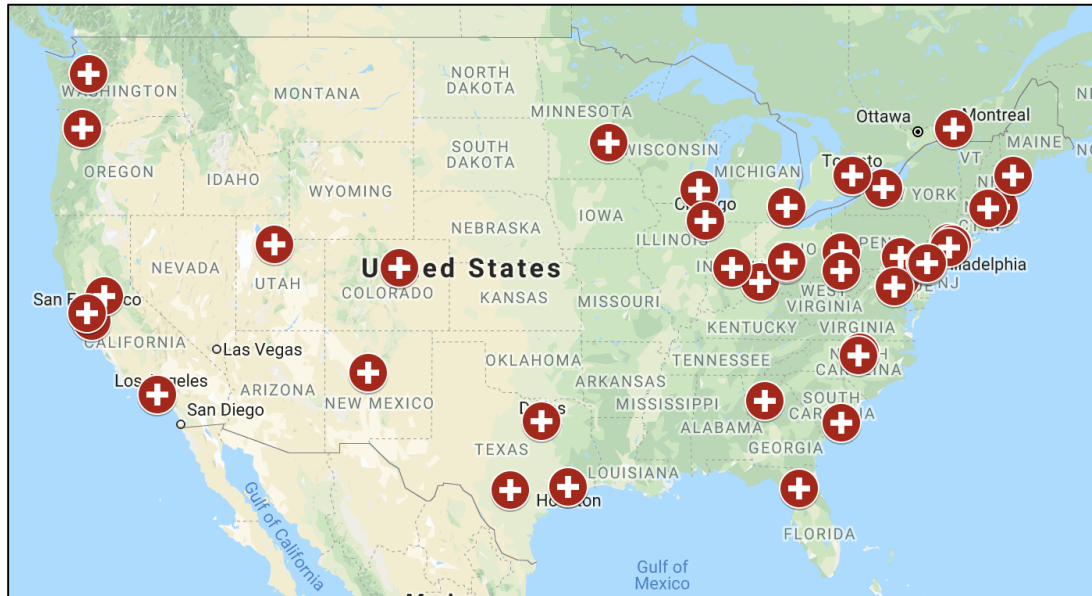
- Project
  - A More Meaningful Medical Record for the Brain to Enable Precision Management of TBI
  - Three year, DOD-funded, collaborative project
- Goals:
  - Standards: nomenclature, annotations
  - Architecture: Standardized data pathways, API
  - Define what should be in the medical record (and monitoring) that makes it useful for managing brain injury
- Organizations:
  - Moberg Research, MGH, U Cincinnati
  - Large group of collaborators and stakeholders



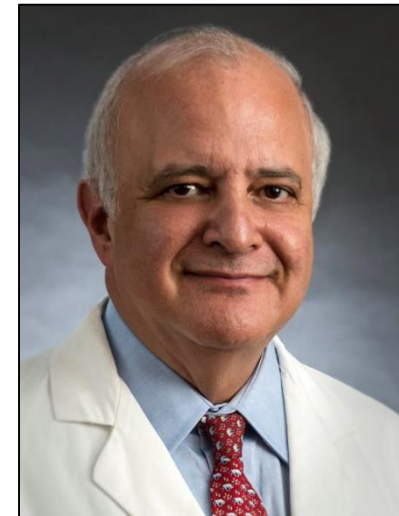
# Project: BOOST3 – A Multicenter Trial

## BOOST-3 Clinical Trial

- > 1000 Patients
- ~46 Participating Sites
- Compare TBI Treatment Strategies
  - ICP-only vs. ICP + PbtO2



- Baylor College of Medicine
- Harvard Medical School
- University of Maryland
- University of Cincinnati
- Columbia University Medical Center
- Duke University Medical Center
- Emory University
- University of North Carolina School of Medicine
- Henry Ford Health System
- Indiana University (IU Health Methodist Hospital)
- Medical College of Wisconsin
- University of Chicago
- Regions Hospital (Sub-Hub for U of Minnesota)
- University of Montreal
- North Shore University Hospital
- Oregon Health & Science University
- University of Rochester Medical Center
- Ohio State University Wexner Medical Center
- Penn State Hershey Milton S. Hershey Medical Center
- University of Pittsburgh
- Riverside Methodist Hospital - OhioHealth
- Stanford Medical
- Kings County
- Cooper University Hospital
- Washington Hospital Center/Georgetown University
- University of Texas Health Science Center at San Antonio
- University of Texas Health Sciences Center at Houston
- St. Michaels- University of Toronto
- Maine Medical Center
- UC Davis Medical Center
- University of California, Los Angeles
- Queen's Medical Center, HI
- University of California, San Francisco
- University of Massachusetts
- University of Pennsylvania
- University of Utah
- Parkland Hospital
- University of Washington
- Wayne State University
- University of New Mexico Hospital
- Medical University of South Carolina
- West Virginia University
- Thomas Jefferson University Hospital
- University of Florida
- University of Colorado School of Medicine



Ramon Diaz-Arrastia, MD, PhD



Penn Medicine

<https://siren.network/clinical-trials/boost-3>

# Two Projects



Develop a standardized platform for data used in precision management

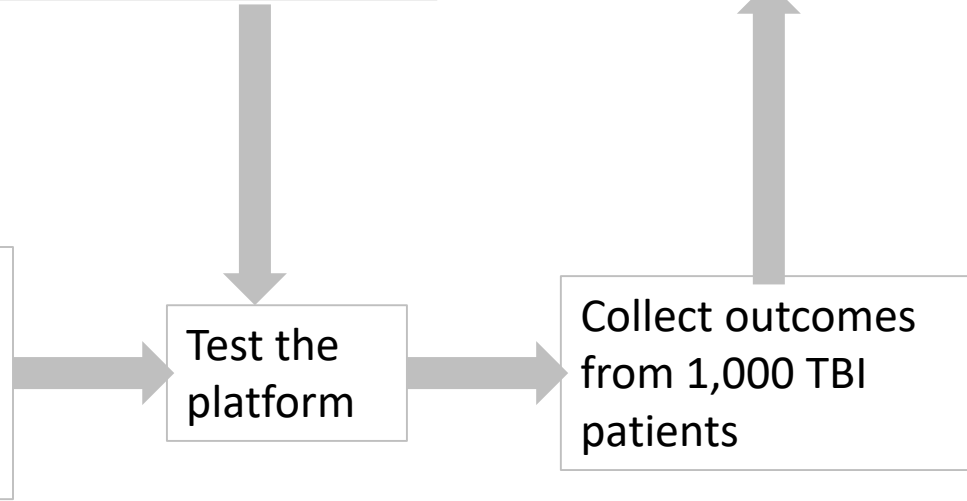
Develop a new medical record for the brain



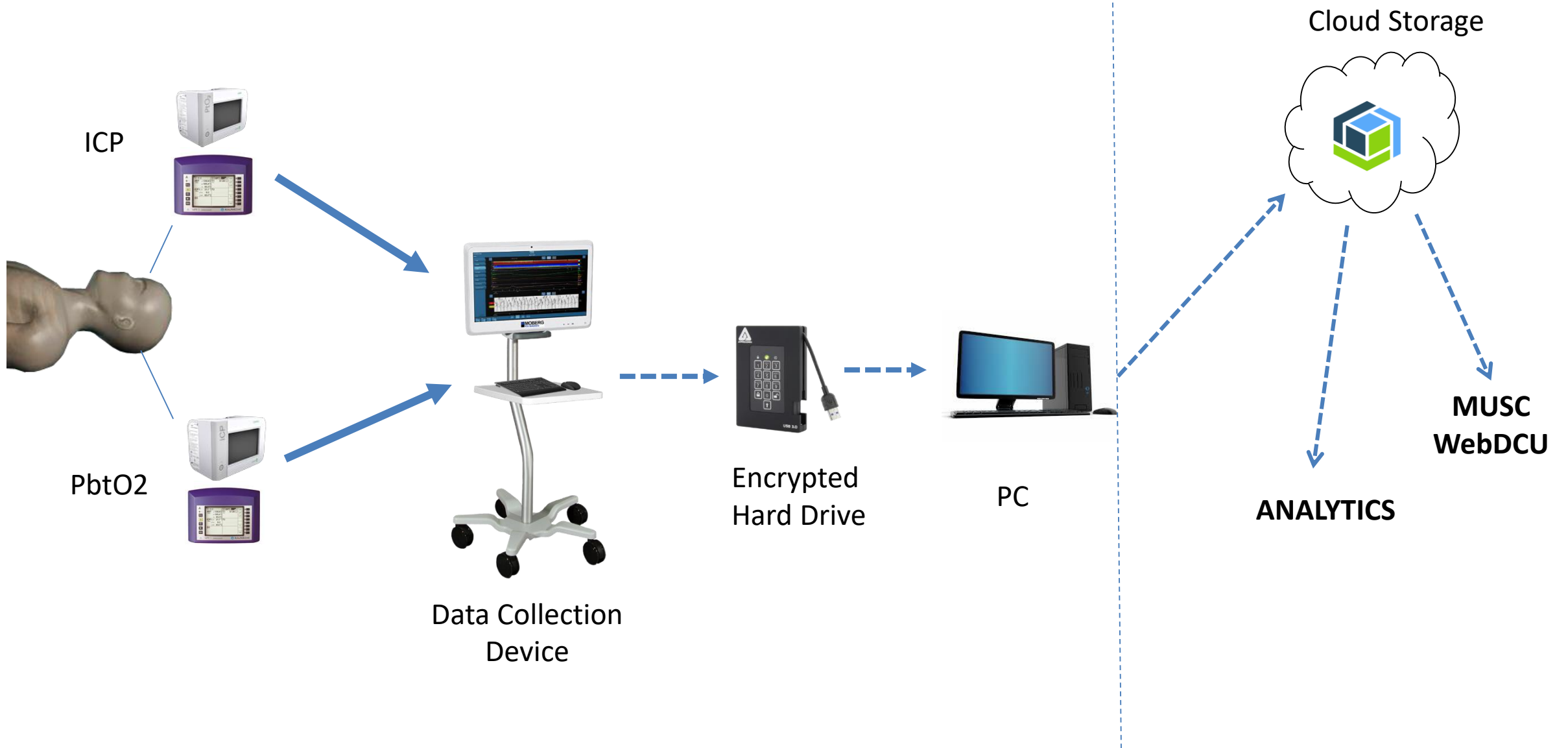
Collect comprehensive data from 1,000 TBI patients

Test the platform

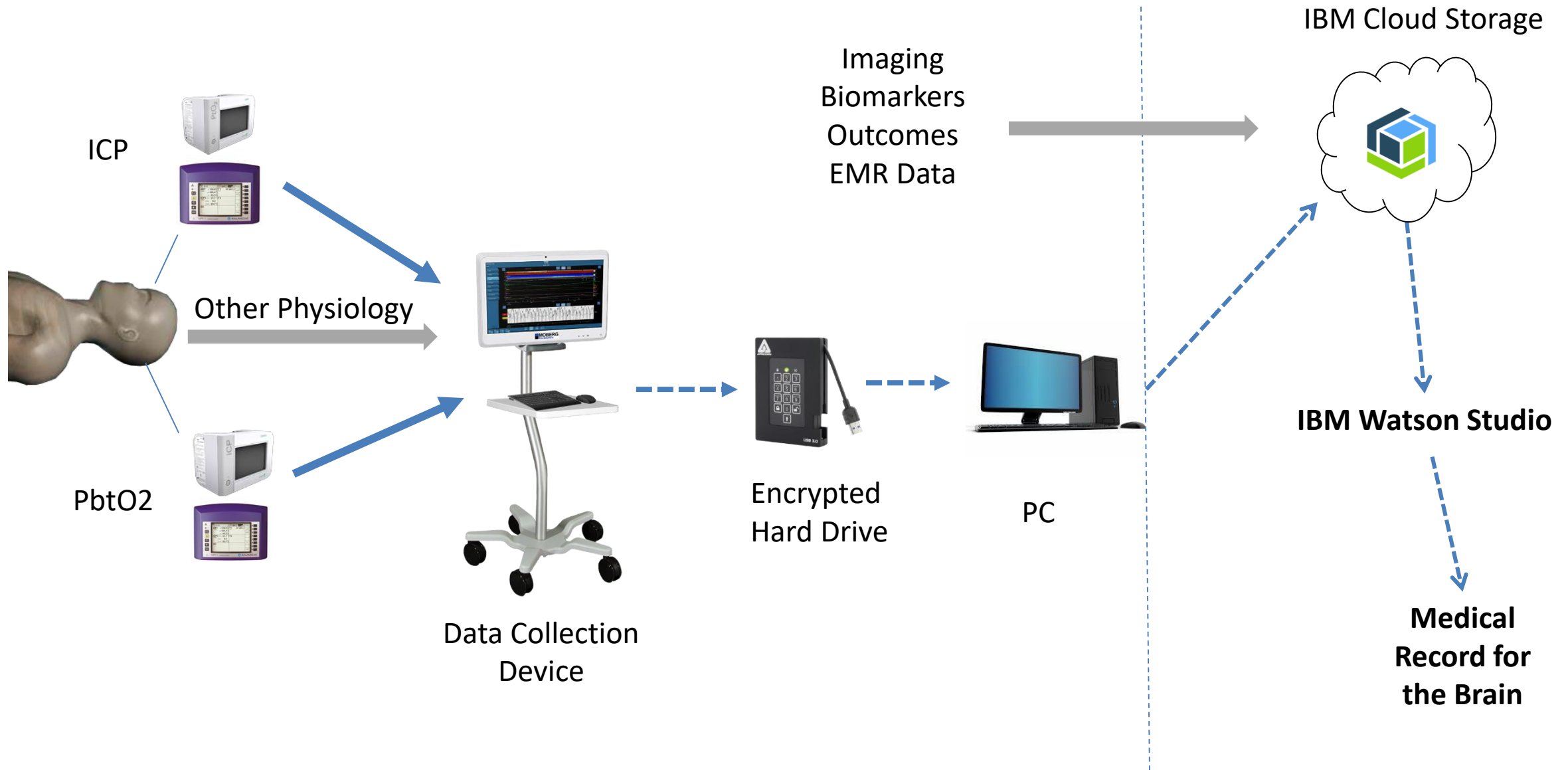
Collect outcomes from 1,000 TBI patients



# The Data Pathway for BOOST3

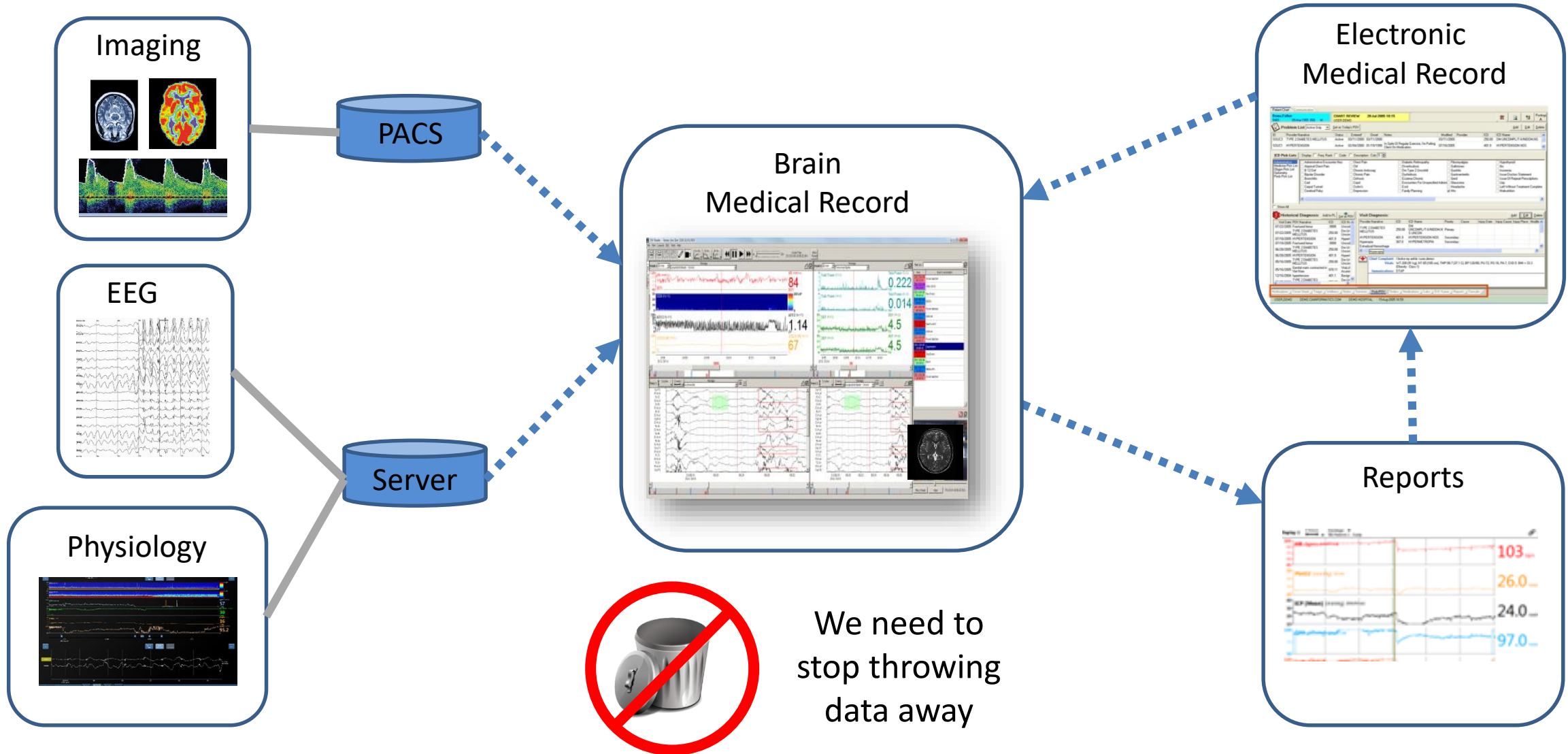


# The Data Pathway for BOOST3 – Ancillary Projects

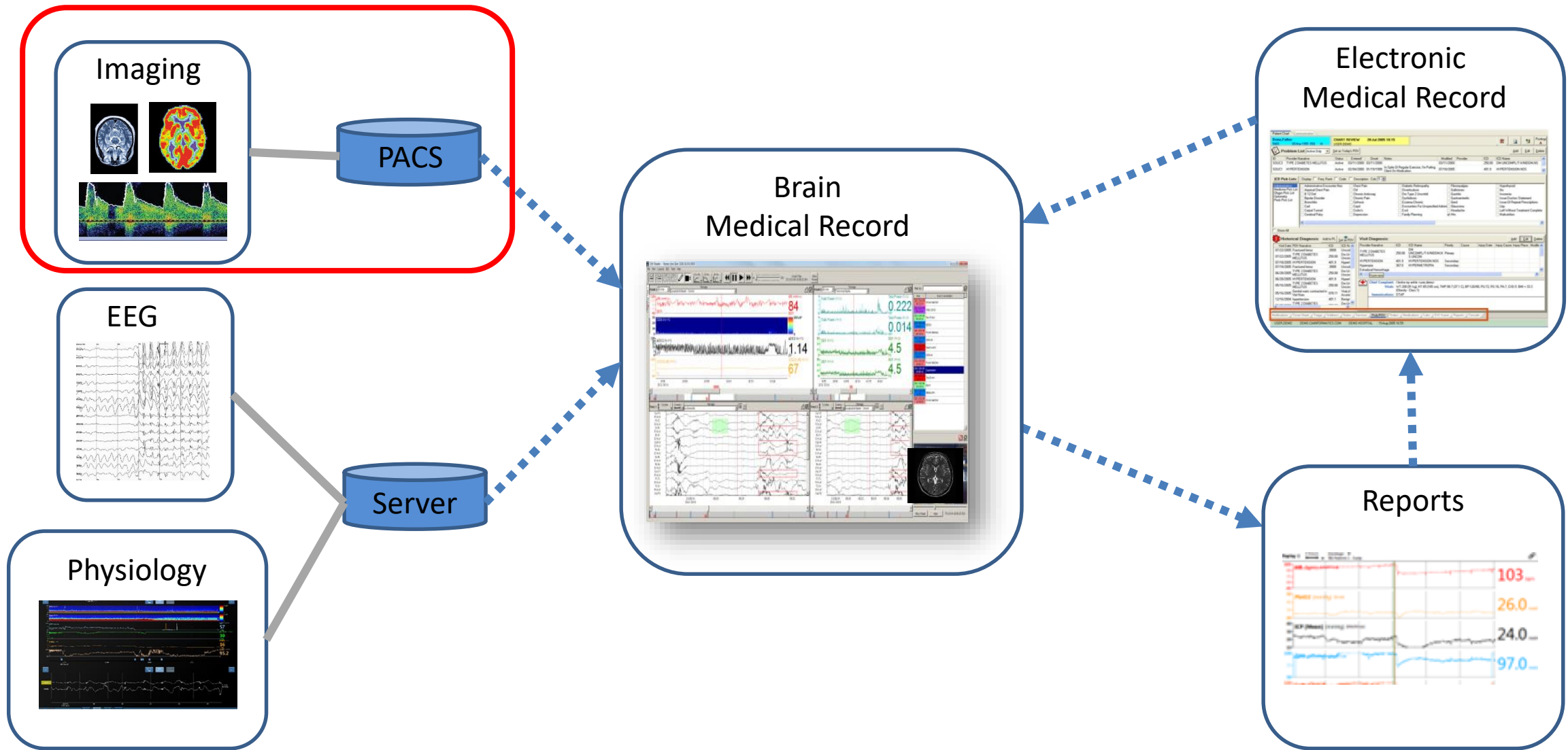


# A More Meaningful Medical Record for the Injured Brain

We need to collect data in a way that enables further use

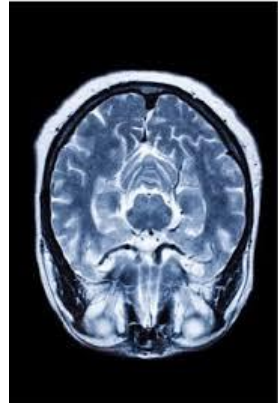


# A New Medical Record for the Brain – Data Collection





# Transform Image Data to a Usable Format



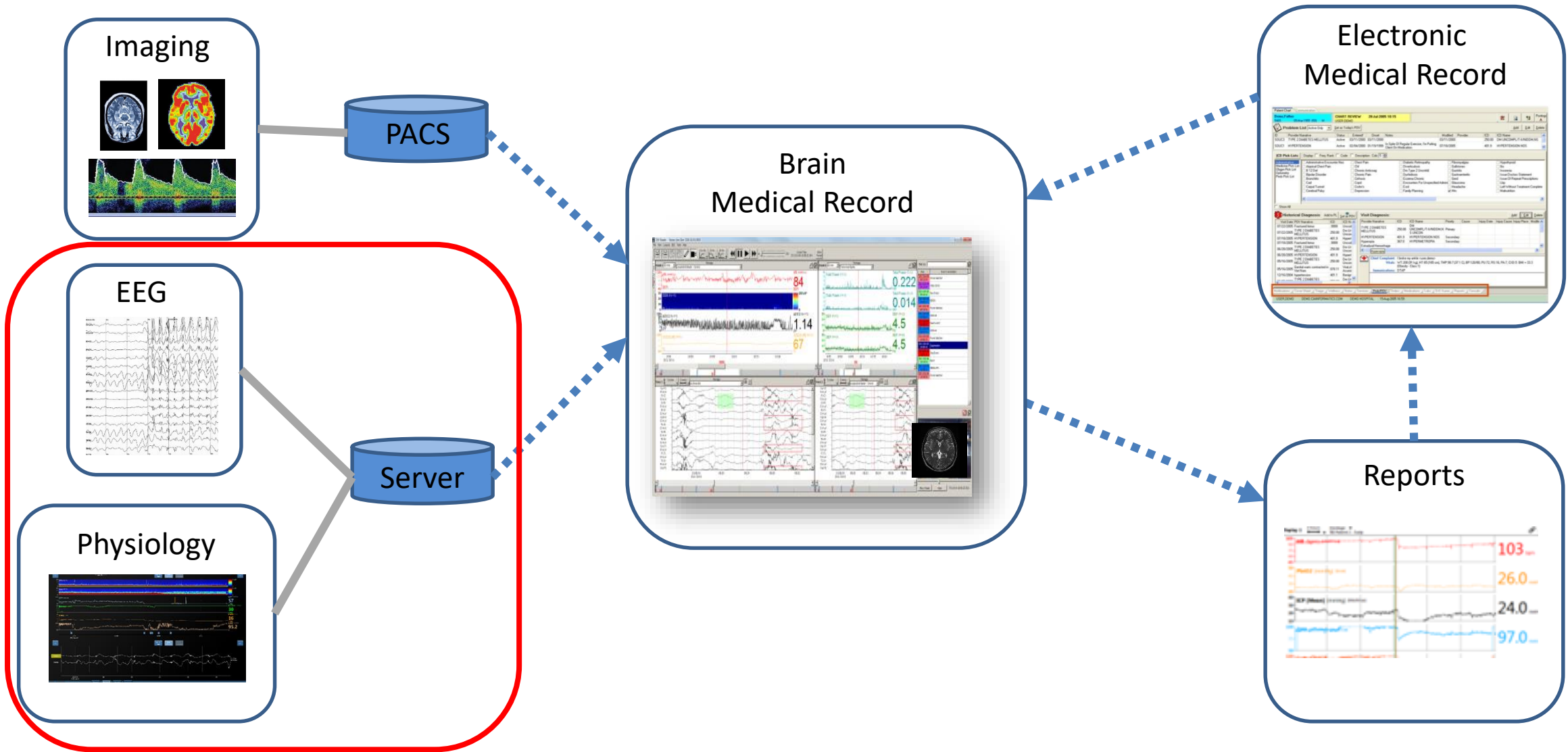
Automated Medical  
Image Description

Max Wintermark  
Stanford



Argonne's IBM Blue  
Gene/Q Supercomputer  
U.S. Dept of Energy

# A New Medical Record for the Brain – Data Collection



## EEG



Automatic  
Detection of  
Significant  
Descriptors



Ancillary Study



## Database

## ECG

**MIMIC/PhysioNet**

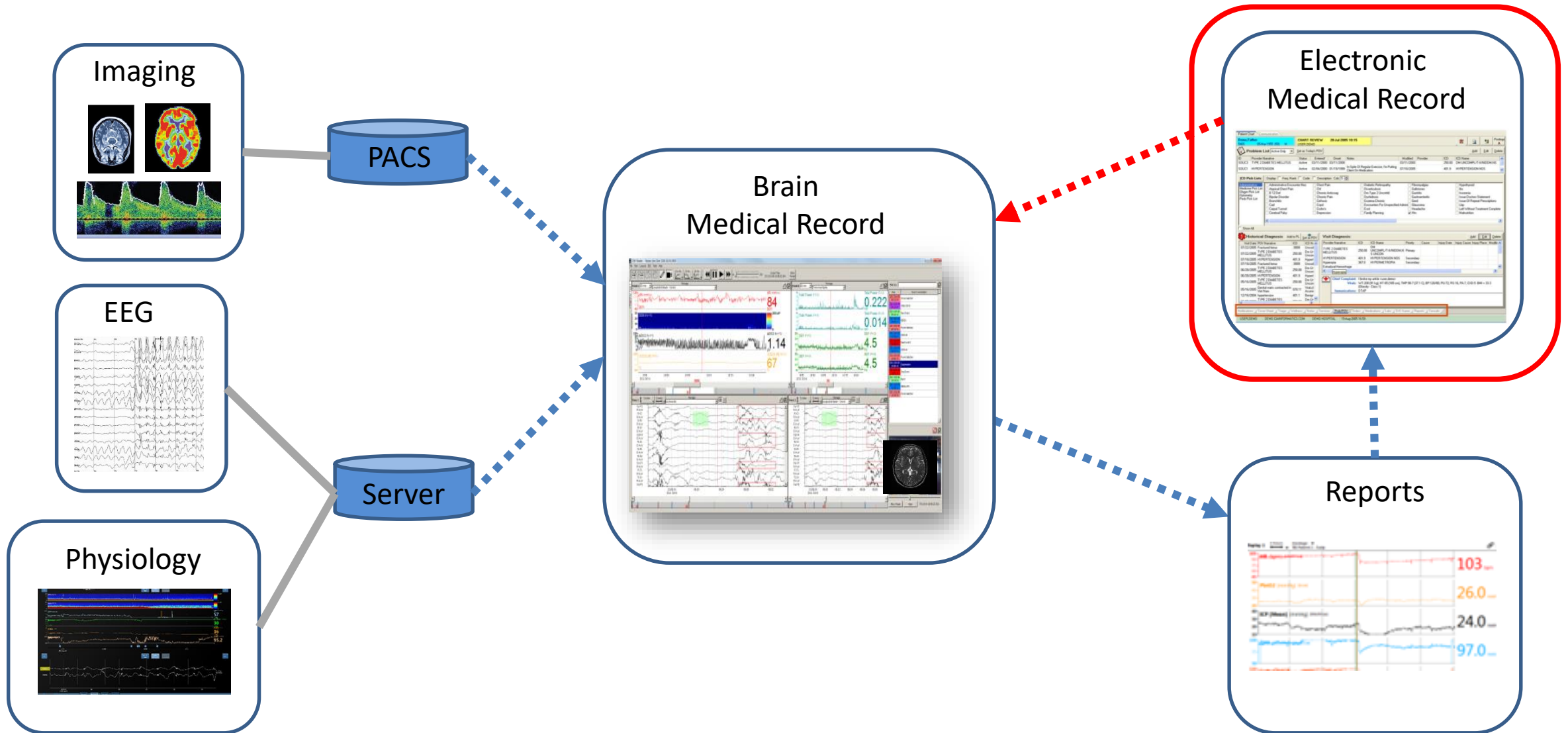
National treasure  
30 years of funding



## Multimodal Neuro Database

Start to develop annotated  
multimodal neurotrauma  
database with BOOST data

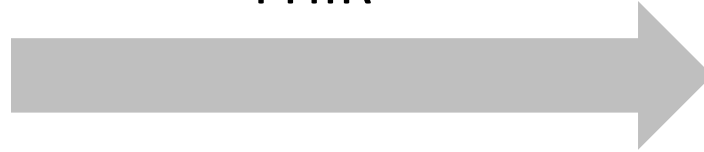
# A New Medical Record for the Brain





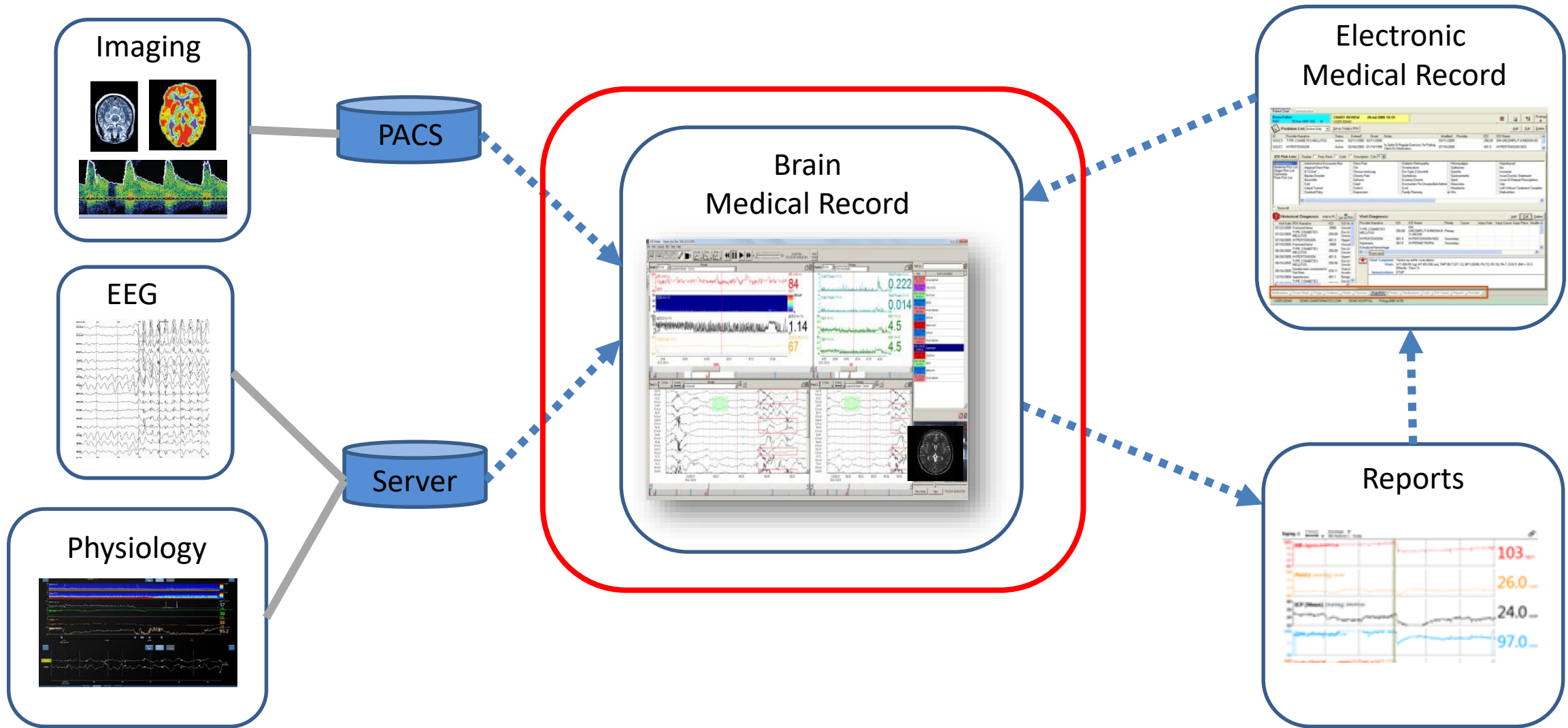
**Epic**

Redox  
FHIR



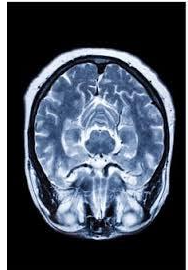
ADT  
Medications  
Lab Data  
Nursing Interventions

# A New Medical Record for the Brain

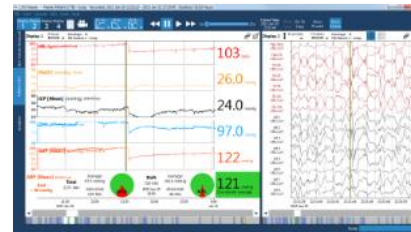
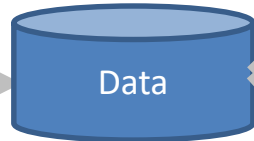
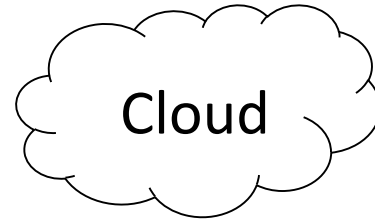


# Our Approach

## Contextual Data



## Patient State Determination



CNS Envision

## Guidance

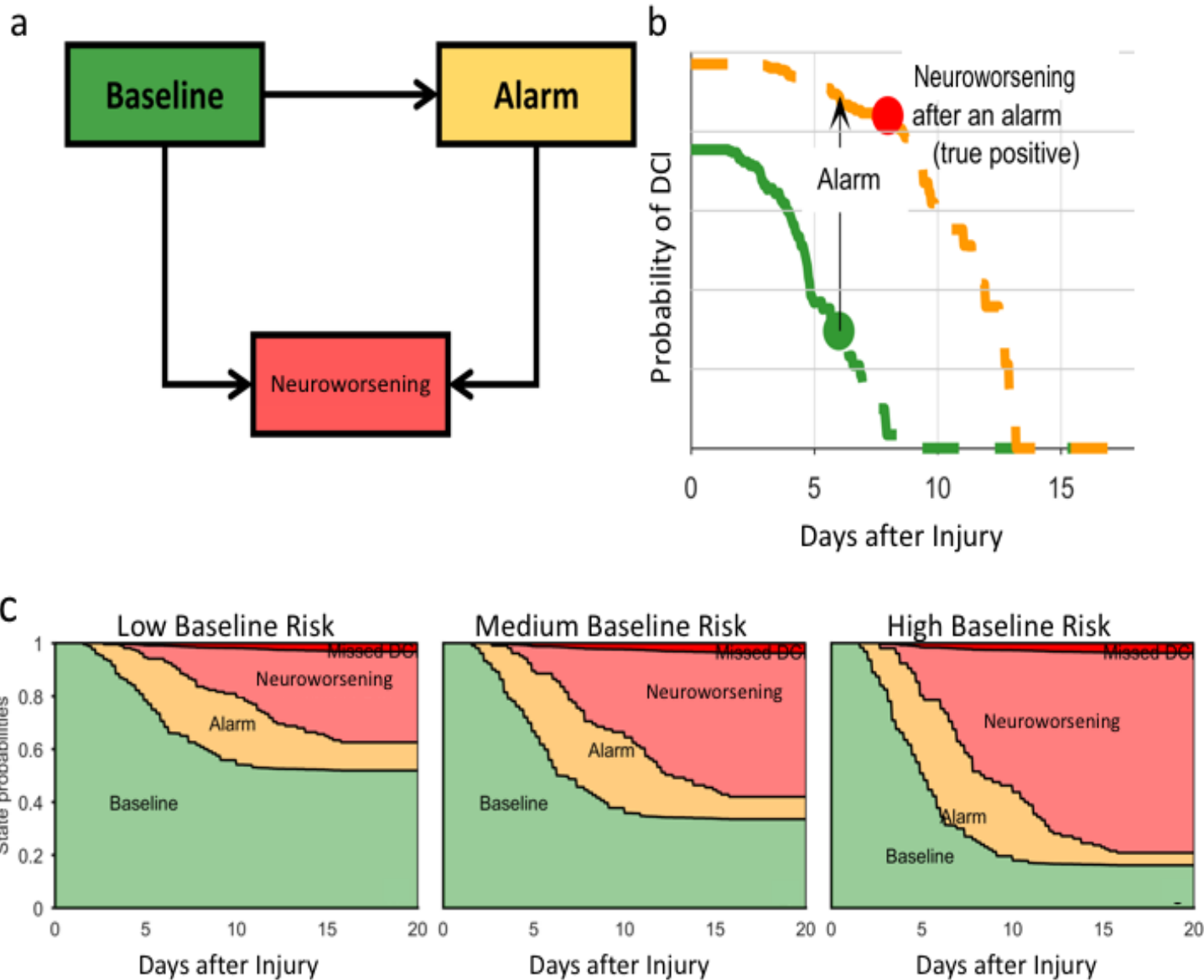
Apps

- Data cleaning
- Event detection
- Prediction
- Classification
- Machine learning
- Data mining

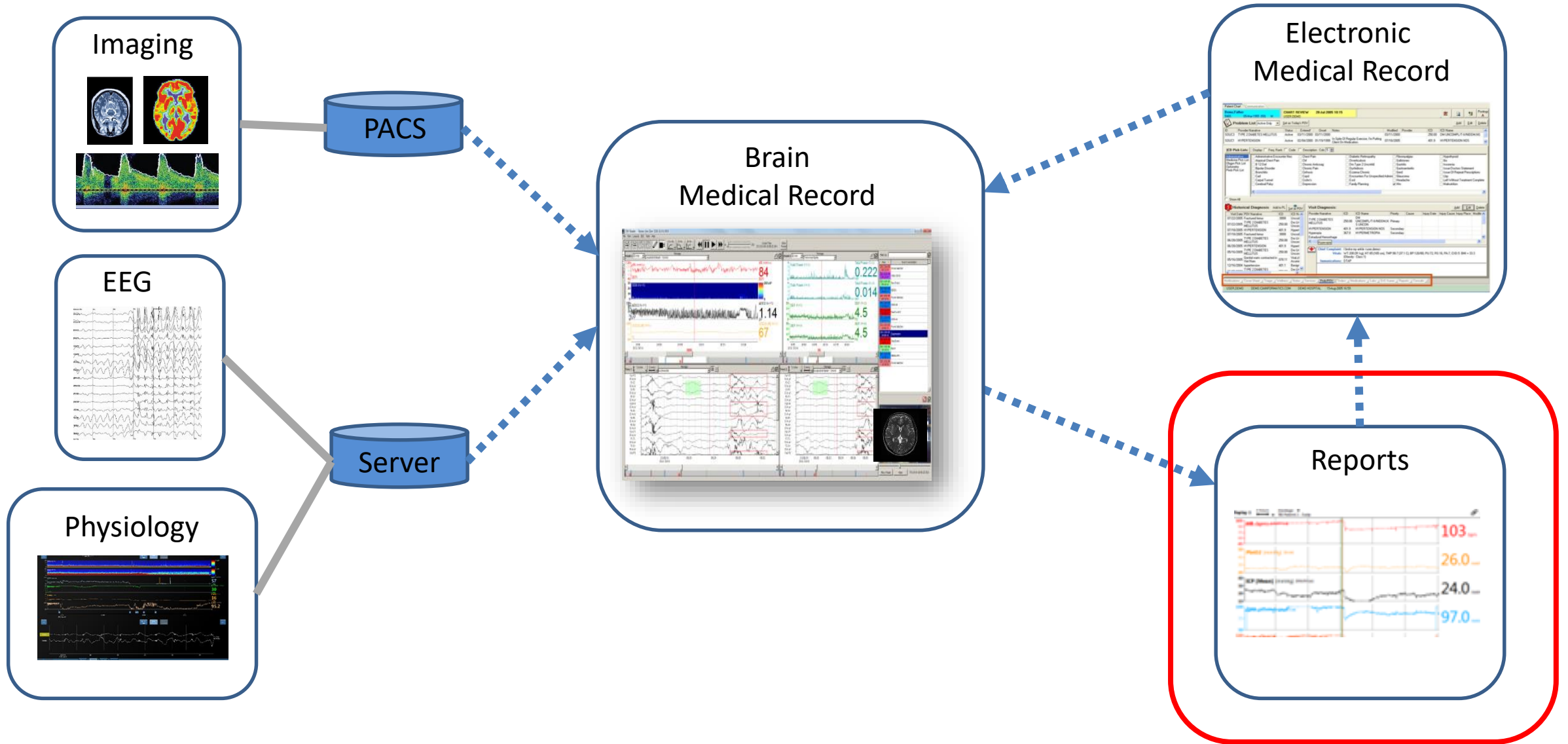
Apps

Visualization  
Guidance

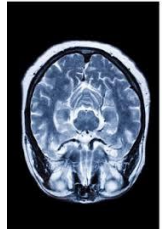
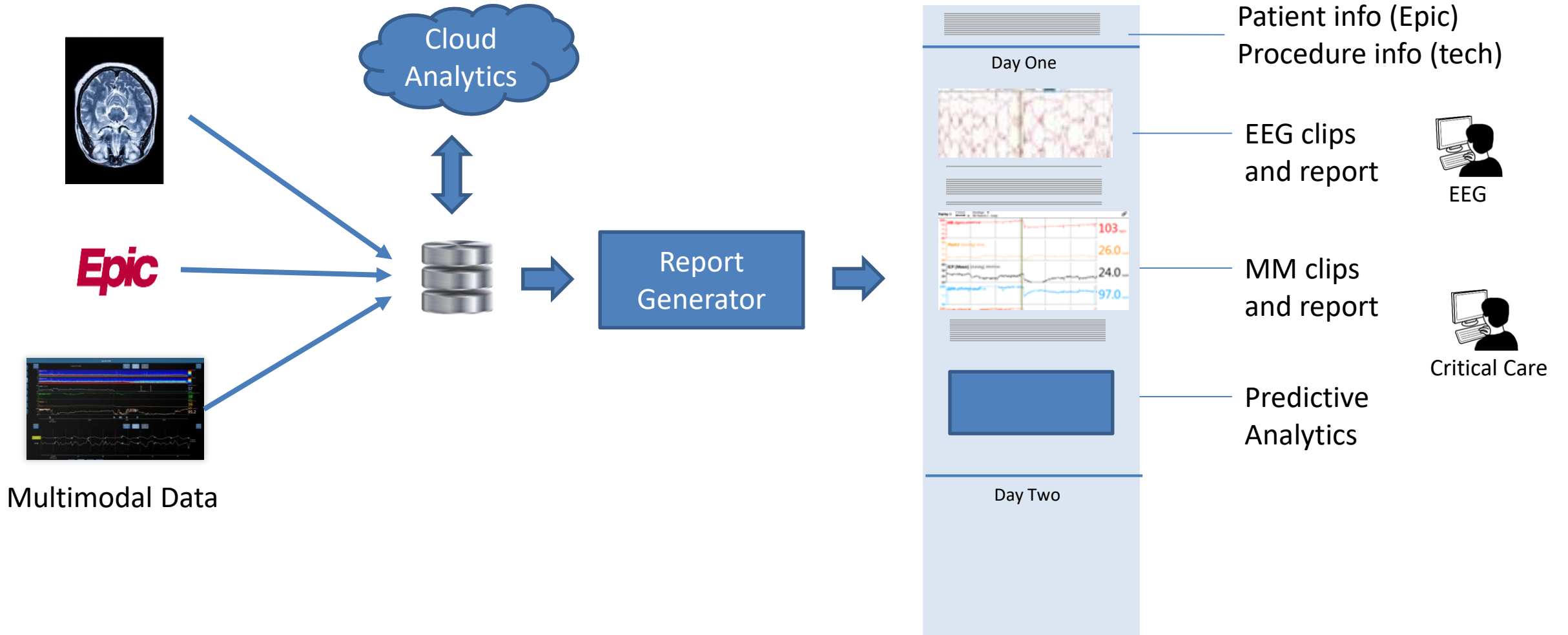
# Multi-state Predictive Models of Neuroworsening



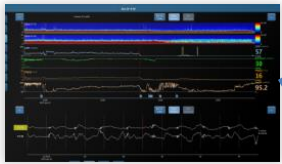




# Automated Comprehensive Reports



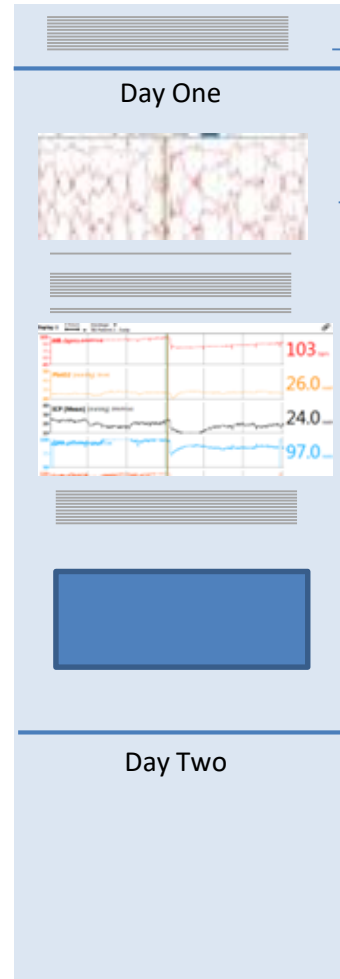
**Epic**



Multimodal Data



Report Generator



Patient info (Epic)  
Procedure info (tech)

EEG clips  
and report



MM clips  
and report



Critical Care

Predictive  
Analytics

Day Two

**BACKGROUND:**

1. Normal features: None.
2. Abnormal features: The background activity is poorly organized, with no discernable posterior dominant rhythm. Eye blinks are absent. The EEG is not reactive.

**SLEEP FEATURES:** Rudimentary sleep architecture is present.

**ATTENUATION:** None

**SLOWING:** Mild generalized slowing is present.

**HYPERVENTILATION:** None

**PHOTIC STIMULATION:** None

**TECHNOLOGIST'S IMPRESSION:** None

**EPILEPTIFORM DISCHARGES, RHYTHMIC OR PERIODIC PATTERNS:**

1. Abundant spikes, most prominent in the left fronto-temporal regions.

**SEIZURES OR CLINICAL EVENTS:**

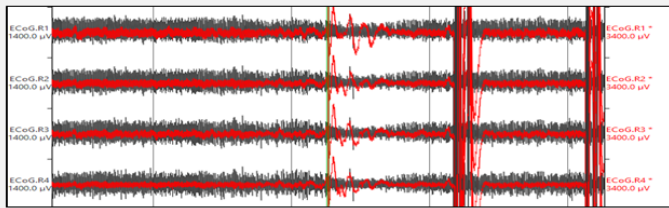
1. Electrographic seizures: Seizures occurring about 10 times per hour. The majority are subclinical. Clinical: Patient lying in bed. After a minute or two there is right facial twitching followed by head turn and gaze deviation to the right. There is head twitching to the right and sometimes the entire face right > left appears to be twitching. Toward the end of the seizure, she has a head turn to the left.

During some of the seizures she will have only facial pulling to the right. No head or eye deviation.

Electrographic: There is rhythmic delta over the left hemisphere. This evolves into left LRDA+S at 2-2.5z over the left fronto-centro-temporal region. At times there is left hemisphere fast activity during the seizure. The right hemisphere shows polymorphic slowing during this time. During the clinical seizures there is muscle artifact that obscures the right > left sided electrodes. Typically, during the clinical seizures after the muscle artifact ends the electrographic seizure continues for several seconds consisting of LRDA+FS throughout the left hemisphere that slows from 3Hz to 2Hz.

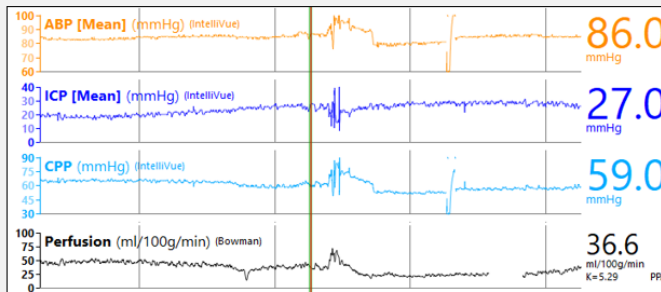
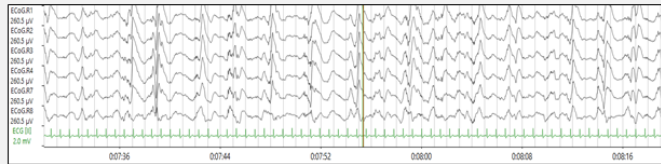
2. The patient is having frequent left foot jerking which is not suppressible. This has no EEG correlate.

3. Patient exhibits characteristics of spreading depolarization, which indicates a dangerous level of brain inactivity. Depolarization was observed on ECoG and was found to impact relevant multimodal measurements at the time of the spike / compression.



**MULTIMODAL DATA:**

1. **ABP:** ABP was also affected by the spreading depolarization, as levels can be seen to rise shortly after the event. ABP was found to be above 85 mmHg for 73.1% of the shift, while it remained increased for 71.3% of the total measurement time.
2. **ICP:** ICP averaged 5.9+/-3.8 mmHg. The ICP waveform demonstrated P2>P1. ICP was monotonous with very brief (<15 min) spikes to 30 mmHg at 16:00, 17:00, 20:15. There were scattered very low-amplitude b-waves with no plateau waves. ICP measurements were averaged per minute. Each average measurement was compared to the goal range of <20 mmHg. In total, 86% of measurements were above 20 mmHg, while the current period had 85.8% of the ICP measurements above 20 mmHg, indicating that the patient's status is critical.
3. **CPP:** The CPP averaged 78.2+/-8.5 mmHg. There were no sustained periods of hypoperfusion (CPP<60) throughout the recording. There was reciprocal decrease in CPP with ICP spikes described above transiently as low as 50 mmHg. However, ICP remained stable across a range of CPP. However, PbtO2 and CBF correlated with CPP.
4. **rCBF:** The PPA remained <2 throughout the recording. During brain fever (>39C) there were no CBF readings 2/2 technical limitations of the probe. CBF initially related to sedation and was as low as 6 ml/100g/min; after temperature was controlled overnight (after 22:00), the CBF ranged from 15-40 ml/100g/min and was correlated with CPP and linked by sedation. K value gradually increased from 4.99 at insertion to 5.10 at the end of the recording, consistent with <75% brain water content.



Combined multimodal physiology and cEEG.

Neuroworsening metrics

How well did nursing meet the target?  
 What improvements have we achieved?

**BACKGROUND:**

1. Normal features: None.
2. Abnormal features: The background activity is poorly organized, with no discernable posterior dominant rhythm. Eye blinks are absent. The EEG is not reactive.

**SLEEP FEATURES:** Rudimentary sleep architecture is present.

**ATTENUATION:** None

**SLOWING:** Mild generalized slowing is present.

**HYPERVENTILATION:** None

**PHOTIC STIMULATION:** None

**TECHNOLOGIST'S IMPRESSION:** None

**EPILEPTIFORM DISCHARGES, RHYTHMIC OR PERIODIC PATTERNS:**

1. Abundant spikes, most prominent in the left fronto-temporal regions.

**SEIZURES OR CLINICAL EVENTS:**

1. Electrographic seizures: Seizures occurring about 10 times per hour. The majority are subclinical. Clinical: Patient lying in bed. After a minute or two there is right facial twitching followed by head turn and gaze deviation to the right. There is head twitching to the right and sometimes the entire face right > left appears to be twitching. Toward the end of the seizure, she has a head turn to the left.  
  
During some of the seizures she will have only facial pulling to the right. No head or eye deviation.  
  
Electrographic: There is rhythmic delta over the left hemisphere. This evolves into left LRDA+S at 2-2.5z over the left fronto-centro-temporal region. At times there is left hemisphere fast activity during the seizure. The right hemisphere shows polymorphic slowing during this time. During the clinical seizures there is muscle artifact that obscures the right > left sided electrodes. Typically, during the clinical seizures after the muscle artifact ends the electrographic seizure continues for several seconds consisting of LRDA+FS throughout the left hemisphere that slows from 3Hz to 2Hz.
2. The patient is having frequent left foot jerking which is not suppressible. This has no EEG correlate.
3. Patient exhibits characteristics of spreading depolarization, which indicates a dangerous level of brain inactivity. Depolarization was observed on ECoG and was found to impact relevant multimodal measurements at the time of the spike / compression.

