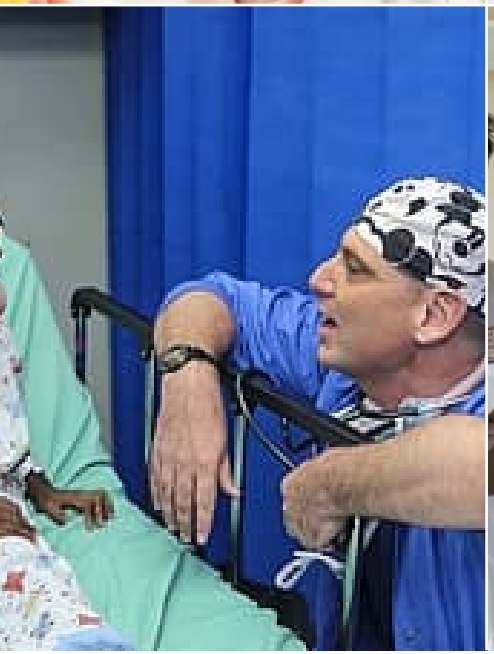


# Role of Principal Investigator

**The PI assumes the overall responsibility for all clinical research activities and ensures that the study is run in accordance with the requirements of the relevant regulatory agencies and the Guidelines for Good Clinical Practice.**

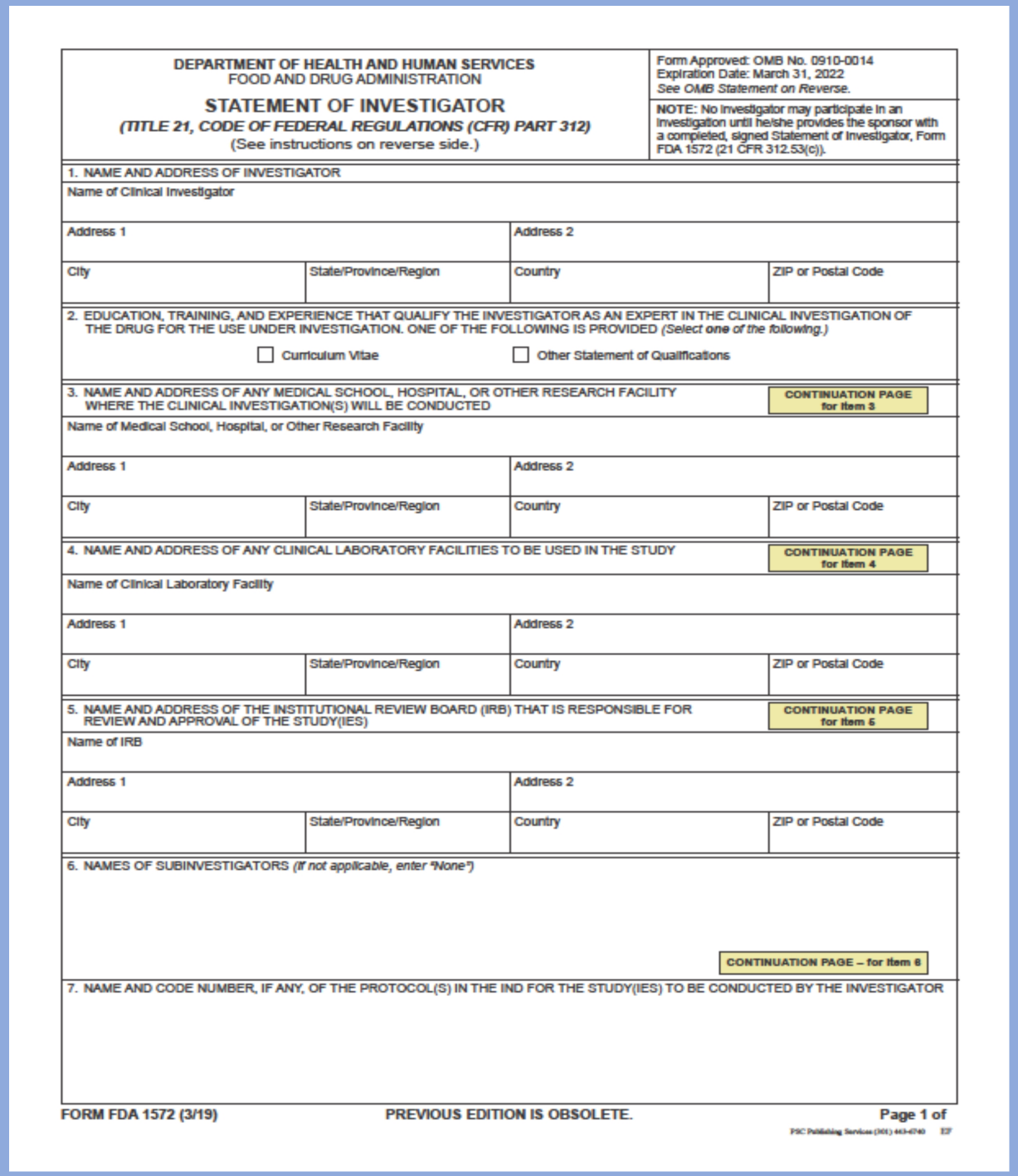
**These responsibilities include:**

1) **Conducting and supervising the research activities, e.g. ensuring that the protocol is scientifically sound and of value and the facility is equipped (personnel, equipment, space, funding) to perform the research.**
2) **Obtaining all necessary approval, specifically Institutional Review Board approval .**
3) **Select personnel that is qualifies and certified to perform the tasks they are assigned to during the trial.**
4) **Protect the rights, safety and welfare of the study participants e.g. by ensuring that the study is performed in accordance with all applicable regulatory requirements.**

# FDA 1572

**As physicians we all sign contracts.**

**As Principal Investigator you will sign a contract with the Federal Government.**



DEPARTMENT OF HEALTH AND HUMAN SERVICES
FOOD AND DRUG ADMINISTRATION

**STATEMENT OF INVESTIGATOR**

***(TITLE 21, CODE OF FEDERAL REGULATIONS (CFR) PART 312)***

(See instructions on reverse side.)

Form Approved: OMB No. 0910-0014
Expiration Date: March 31, 2022
*See OMB Statement on Reverse.*

**NOTE:** No investigator may participate in an investigation until he/she provides the sponsor with a completed, signed Statement of Investigator, Form FDA 1572 (21 CFR 312.53(c)).

1. NAME AND ADDRESS OF INVESTIGATOR

Name of Clinical Investigator

| Address 1 | | Address 2 | |
|---|---|---|---|
| City | State/Province/Region | Country | ZIP or Postal Code |

2. EDUCATION, TRAINING, AND EXPERIENCE THAT QUALIFY THE INVESTIGATOR AS AN EXPERT IN THE CLINICAL INVESTIGATION OF THE DRUG FOR THE USE UNDER INVESTIGATION. ONE OF THE FOLLOWING IS PROVIDED *(Select **one** of the following.)*

☐ Curriculum Vitae ☐ Other Statement of Qualifications

3. NAME AND ADDRESS OF ANY MEDICAL SCHOOL, HOSPITAL, OR OTHER RESEARCH FACILITY WHERE THE CLINICAL INVESTIGATION(S) WILL BE CONDUCTED — CONTINUATION PAGE for Item 3

Name of Medical School, Hospital, or Other Research Facility

| Address 1 | | Address 2 | |
|---|---|---|---|
| City | State/Province/Region | Country | ZIP or Postal Code |

4. NAME AND ADDRESS OF ANY CLINICAL LABORATORY FACILITIES TO BE USED IN THE STUDY — CONTINUATION PAGE for Item 4

Name of Clinical Laboratory Facility

| Address 1 | | Address 2 | |
|---|---|---|---|
| City | State/Province/Region | Country | ZIP or Postal Code |

5. NAME AND ADDRESS OF THE INSTITUTIONAL REVIEW BOARD (IRB) THAT IS RESPONSIBLE FOR REVIEW AND APPROVAL OF THE STUDY(IES) — CONTINUATION PAGE for Item 5

Name of IRB

| Address 1 | | Address 2 | |
|---|---|---|---|
| City | State/Province/Region | Country | ZIP or Postal Code |

6. NAMES OF SUBINVESTIGATORS *(If not applicable, enter "None")*

CONTINUATION PAGE – for Item 6

7. NAME AND CODE NUMBER, IF ANY, OF THE PROTOCOL(S) IN THE IND FOR THE STUDY(IES) TO BE CONDUCTED BY THE INVESTIGATOR

FORM FDA 1572 (3/19) PREVIOUS EDITION IS OBSOLETE. Page 1 of

PSC Publishing Services (301) 443-6740 EF

# Pharmaceutical Sponsored vs Investigator Initiated

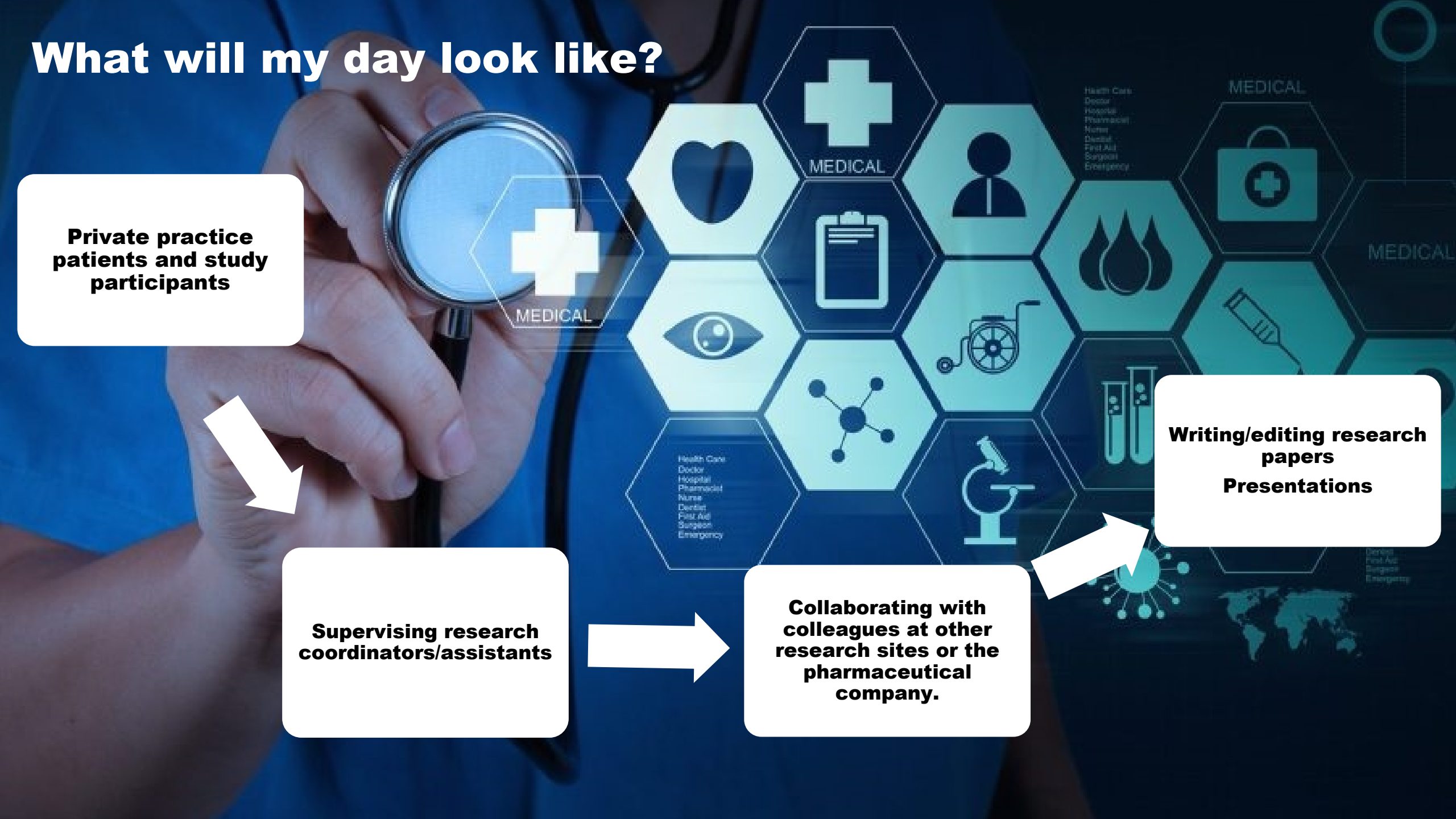
## Multisite Clinical Trials – Pharmaceutical Grants

- Protocols are provided to the site
- Some or all of the clinical supplies are provided
- Number of participants may vary from a 100 to several thousand
- Pharmaceutical company oversees the conduct of the trial
- Central or Local Institutional Review Board

## Investigator Initiated Clinical Trials – Federal, Patient Advocacy, Private Foundations, University

- Principal Investigator develops and writes the protocol
- Clinical supplies are provided by the investigator
- Small number of participants
- Local Institutional Review Board

# What will my day look like?




**Private practice patients and study participants**

**Supervising research coordinators/assistants**

**Collaborating with colleagues at other research sites or the pharmaceutical company.**

**Writing/editing research papers**

**Presentations**

# Clinical Research vs Clinical Practice

| Clinical Practice | Clinical Research |
| --- | --- |
| Safety and well being | Safety and well being |
| Expectation of treatment | No expectation of treatment |
| FDA approved medication | Investigational Product |
| Off Label prescriptions as agreed between the physician and patient | FDA approved medication that is being tested in a particular patient population or new disease |
| Patient appointments are determined by patient and/or insurance coverage | Participant visits are set by the protocol and ***must*** be followed.<br><br>The PI may have additional appointment based on their discretion for safety. |
| Fees are paid by the patient or third party. | The sponsor pharmaceutical company pays the PI for the entire study. |
| Patients do not receive compensation for their time and efforts. | Participants frequently receive a stipend for their time and efforts. |
| Both the patient and physician now what is being prescribed. | In double blind trials neither the physician nor the participant knows. |

# Consent: Mission Critical in research


Consent for procedures & release of information are obtained in practice.

In Clinical Research, consent/assent is obtained **BEFORE** any aspect of the trial may be executed.

- Informed Consent Forms will be scrutinized by the IRB, FDA, Sponsor.
- They must be signed/dated.
- The participate must receive a sign/dated copy.
- The process must be documented, and signed off.

- Check In – Warm
- Determined by need of patient and decision of MD
  - Vital Signs
  - Examinations
  - Laboratory
  - Scans/Procedures
  - Follow Up
- Conversation – Warm, caring, concerned, congenial

- Check In – Neutral
- Protocol Determines
  - Vital Signs
  - Examinations
  - Laboratory
  - Scans/Procedures
  - Follow up
- MD may override any and all to protect participant safety
- Conversation – Neutral, caring

# Why the neutrality?
# Placebo Response the growing conundrum in clinical research

**Maintaining the blind is paramount in clinical research. It is a medical myth that only CNS trails have "placebo response."**

**Which of the following studies have not had placebo responses in clinical trials?**

1. **Hypertension**
2. **Diabetes Mellitus**
3. **Rheumatoid Arthritis**
4. **Seizure Disorders**
5. **Knee Replacement**

**Over the past 25 years there has been a logarithmic increase in placebo response across all disease states.**

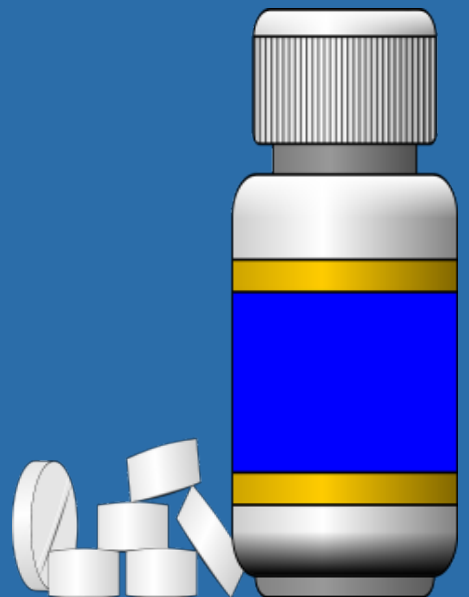
**We are now seeing clinical trial effects in long term oncology trials. Where the participants general health and sense of well being declines when the trial concludes, even when they remain on the same treatment regimen, they were on during the trial**

**Trust and honest communication between the PI and their participants is imperative.**

**Dampening the hype and hyperbole surrounding a new compound**

**Understanding that we are in this experiment together**

In practice you write a prescription and in clinical research …



## Medications

- FDA approved
  - Off label used
- May be written at physician's discretion, within state laws.
- Patient picks up at pharmacy
- No accountability
- You are not responsible for the chain of custody
- You are not responsible for how the patient takes or doesn't comply with treatment
- You are not responsible for monitoring on site storage requirements

## Investigational Product

- NOT FDA approved
  - FDA approved may be in trials
- Protocol directs how IP is to be administered
- Site dispenses/administers IP
- Accountability is documented at every visit
- You are responsible for chain of custody from the time the IP arrives onsite to when it is shipped back to the depot
- Storage conditions are determined by the protocol
- NO PRIOR authorizations!!!!!

# Clinical Research Staff

The ultimate responsibility for the conduct of the trail is the Principal Investigator.

- Site Director
- Front Desk
- Clinical Coordinators/Research Assistants
- Laboratory Technicians
- Recruitment Specialists
- Data Entry
- Regulatory Specialists
- Sub-Investigators

Integration of the pieces of the puzzle occurs with regular communication and collaboration.  As research is a highly regulated and strictly monitored environment all the pieces must fit precisely.

# Collegiality

**Principal Investigator**

**Principal Investigator**
**At other sites**

**Medical Monitors**

**Protocol Design**

**Recruitment**

**Operational Aspects**

**Results**

**Investigator Meetings**

**Ad hoc Meetings**

**Emails**

**Telephone**

# Academic Research

## MD/PhD, MD/MS, MD/MPH

- The PhD, MS, MPH may be obtained prior to medical school, in combination, or after residency

## Research Project Grant [RO1]

- Original and oldest mechanism by NIH
- Supports health-related research

## Pharmaceutical Grants

- University and Industry collaboration

## Patient Care

## Teaching

## Scholarly Journal Articles/Texts

## Presentations

# Broad Range of Therapeutic Areas

## Inpatient & Outpatient

## Early to Late Phase

**Pain**
- Post Operative
- Chronic

**Endocrine**
- Diabetes Mellitus
- Thyroid

**Cardiovascular**

**Hepatic**

**Renal**

**Central Nervous System**
- Alzheimer's
- MDD
- Schizophrenia
- GAD/OCD/PTSD
- Autism

**Vaccines**
- COVID
- Influenza
- Lyme's Disease

**Rare Diseases**

**Women's Health**

Evolution Research Group

# Evolution Research Group

# What it means to be part of ERG

**Professional Opportunities**

- Clinical Trials
- Professional Meetings
- Collaboration

**Administrative Support**

- Staffing
- Human Resources

**Regulatory Support**

**Technologic Support**

## Can I publish?

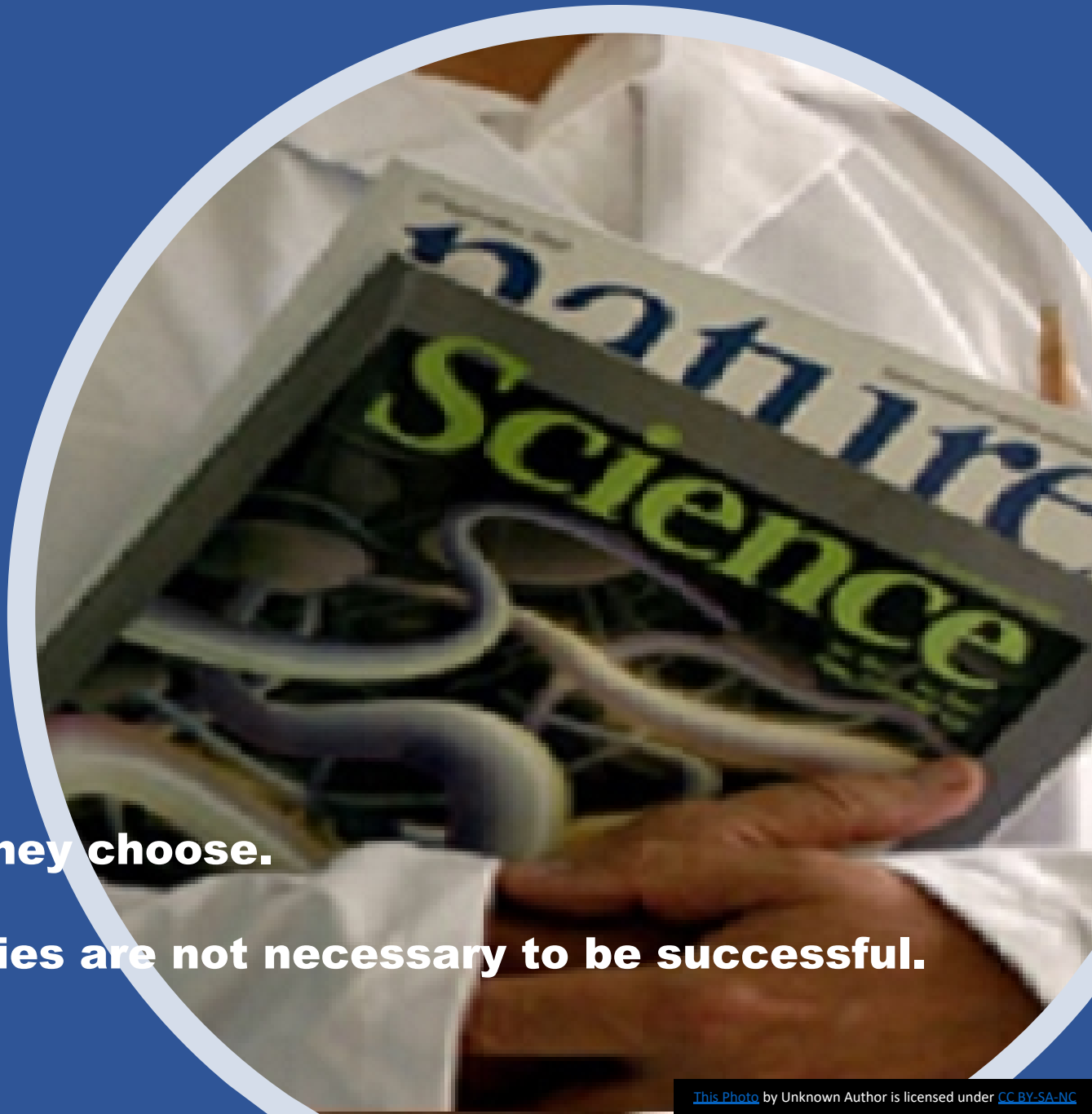
## Must I publish?

Collaboration on scientific papers.

Presentations at national and international conferences.

Participation on panels.

High level discussions with regulatory agencies.






All are part of a PI's professional life if they choose.

In private clinical research these activities are not necessary to be successful.


As a Physician Investigator you are the cutting edge of medicine.