

Research Updates

Tirisham Gyang, MD December 9th 2021



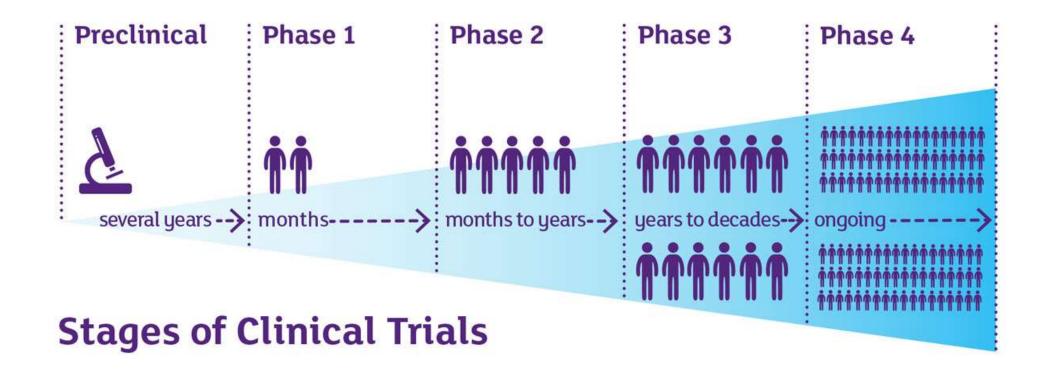
Poll question



Current research at OSU

- Clinical trials phase 3
 - RRMS study of BTK inhibitor
 - PPMS study of BTK inhibitor
- Bench and translational research
 - Neuroscience Research Institute Brain Bank & Biorepository (NRI-BBB)
 - Repair pathways in MS
 - Immune response to DMTs
 - Aging and MS
 - Neuromuscular function and MS



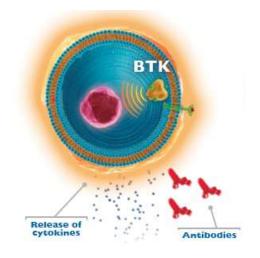


http://stemcellsaustralia.tac-web04.accsysit.com.au/About-Stem-Cells/What-are-clinical-trials-.aspx



BTK inhibitors in RRMS and PPMS

- Bruton's tyrosine kinase (BTK) inhibitor BTK-i
 - BTK is an enzyme found inside certain immune cells
 - B cells
 - Myeloid cells macrophage and granulocytes
 - Microglial cells in the central nervous system
 - Blocking BTK may have therapeutic benefits in
 - Certain types of malignancies leukemia, lymphoma
 - Graft vs host disease transplant patients
 - Autoimmune diseases





BTK-i in RRMS and PPMS

Potential advantages of BTK-i in MS

- Effects on both adaptive and innate immune cells
- Ability to penetrate the blood brain barrier
- Direct effect on microglia cells in the CNS
- May have neuroprotective effects
- Potential benefit in both relapsing and progressive MS

Preliminary studies

EAE – BTK-i effectively treats experimental mouse model of MS

Crespo O. J Clin Immunol. 2011;31(6):1010-1020

- RRMS BTK-i vs. placebo 12 weeks
 - 85% relative reduction in new gadolinium-enhancing lesions
 - 89% relative reduction in new or enlarging T2 lesions (secondary outcome)

Reich DS. Eur J Neurol. 2020;27(Suppl. 1):1–102.



Phase 3 BTK inhibitor (BTK-i) in MS

- Relapsing remitting MS
 - BTK-i 60mg daily vs. Teriflunomide 14mg daily
 - Primary end point annualized relapse rate
- Primary progressive MS
 - BTK-i 60mg daily vs. placebo
 - Primary end point time to onset of 6-month confirmed disability progression
- Secondary progressive MS (starting very soon)
 - BTK-i 60mg daily vs. placebo



Phase 3 BTK inhibitor (BTK-i) in MS

- Contact information
 - Kasturi Ganesh Barki
 - 614-293-6123 Office
 - Email: Kasturi.ganesh@osumc.edu
- Discuss with your MS doctor and let them know you are interested



Neuroscience Research Institute Brain Bank & Biorepository (NRI-BBB)

- The aim of this study is to collect and store biospecimens like blood, spinal fluid and tissue from patients undergoing routine testing for clinical care.
- If you are undergoing routine blood work or a procedure, you can be enrolled to provide an extra sample of blood, fluid or tissue to the biorepository
- These bio-samples will be used for research to increase our understanding of the disease process in MS and other neurological disorders



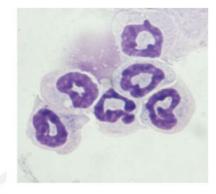
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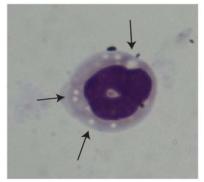
MSResearch@osumc.edu



Dr. Segal - Lab research

- Interrogation of <u>repair pathways in MS</u>
 - Investigating the potential of <u>novel white blood</u> <u>subsets</u>
 - suppress destructive inflammation
 - induce remyelination/ nerve fiber regeneration in mouse models of MS
- In depth analysis of immune responses in relapsing and progressive MS pre- and post-initiation of DMT
 - Goal: To <u>discover biomarkers</u> predictive of responsiveness to individual DMTs, or that reflect disease activity, and to elucidate new therapeutic targets







Aging and MS

- Dr. Zhang is recruiting for a study of biological age in MS
 - To see if people with MS age differently than those without MS
 - To understand the impact of aging on the disease course in MS
- You may be eligible if you are over 18 years of age and have relapsing-remitting MS or secondary-progressive MS.
- Participants will undergo a blood draw and participate in neurological exams and neuropsychological assessments.
- For more information, please email <u>MSResearch@osumc.edu</u>



Neuromuscular function in MS

- Dr. Zhang is planning a study of neuromuscular function in MS
 - Progressive weakness is common in older adults with MS
 - Age and MS both contribute to weakness
- Goals:
 - Assess feasibility of neuromuscular testing in older adults with MS
 - Measure neural and muscular factors involved leg strength
- The study will take place at Ohio University in Athens
- Recruitment anticipated to start in summer 2022



Thank You

