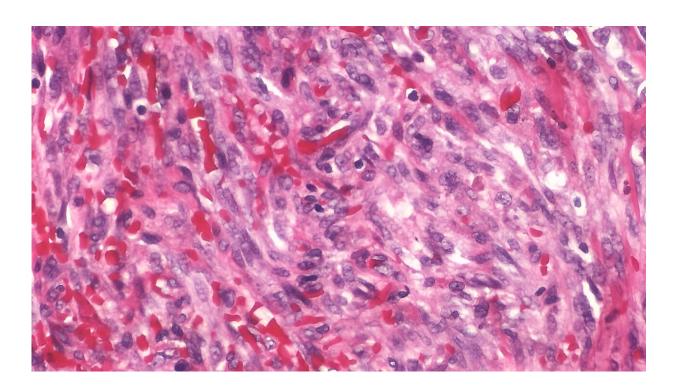


Scientists uncover the 'Rosetta stone' of sarcoma research to personalize treatment

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Kaposi sarcoma (cropped image). Credit: Yale Rosen, CC BY-SA 2.0

Scientists have unveiled the first and largest encyclopedia of protein alterations in soft tissue sarcomas—opening the door to a new era of understanding and treatment for this group of rare cancers, which account for around 4,000 cases in the U.K. each year.

The pioneering encyclopedia, or "Rosetta Stone," of sarcoma biology



compiles a wealth of information that could help untangle the <u>protein</u> <u>changes</u> driving sarcomas' growth, spread and survival.

The findings, published in *Nature Communications*, will allow researchers to tap into the potential of immunotherapy in this <u>cancer</u> type, which although rare in adults, makes up to 15% of cancers in kids, teenagers and young adults. It could also identify people at higher risk of relapse and provide new ways to personalize treatments.

Researchers hope better treatment in this group of patients could lead to more patients being cured of their disease, and also help them live longer cancer-free lives with fewer long term side effects from harsh chemotherapy.

The encyclopedia, developed by a team of researchers and clinicians led by The Institute of Cancer Research, London, comprises data from 321 people—including children, teenagers and adults—with 11 different types of sarcoma tumors. Some patients with sarcoma currently face limited treatment options.

Researchers looked at patient samples to uncover the "proteomic profile" of the sarcoma tumors. These proteomic profiles are snapshots of all the different proteins in different tumors, which help researchers better understand how the proteins are functioning and driving them.

Since most approved cancer drugs work by targeting proteins, understanding more about the proteins driving sarcoma could lead to new targeted drugs.

The new insights have also uncovered new immunotherapy targets, including proteins involved in the complement cascade—a key part of our innate immune system. The new targets could ultimately lead to new ways of treating "immune cold" sarcomas, which are not properly



recognized by the immune system and do not respond well to current immune checkpoint drugs.

The encyclopedia has also allowed researchers to identify proteins that can be measured to help predict which tumors are particularly high-risk and aggressive. This could, in the future, pave the way for more tailored treatment for individuals, for example by identifying patients who may benefit from more aggressive treatment.

The researchers were also able to categorize tumors into a series of interconnected biological networks called "proteomic modules." They identified 14 different modules, which can act as "signatures," or clues, that could help researchers and clinicians to better understand tumors, predict outcomes and assess clinical risk to improve patient care.

Study leader Dr. Paul Huang, Team Leader of the Molecular and Systems Oncology Team at the ICR, said, "Our 'encyclopedia' of <u>protein</u> alterations is a vital resource—a Rosetta Stone of sorts that will help us unravel the complex language of sarcoma tumors. This will allow us to explore new avenues of personalized treatment, offering renewed hope for people with sarcoma.

"Our findings hold promise for the development of new targeted treatments and immunotherapies that can disrupt the underlying processes driving sarcoma. I am hopeful that this study will lead to improved treatment outcomes and a better quality of life for <u>sarcoma</u> patients."

More information: Jessica Burns et al, The proteomic landscape of soft tissue sarcomas, *Nature Communications* (2023). <u>DOI:</u> <u>10.1038/s41467-023-39486-2</u>



Provided by Institute of Cancer Research

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