DATA NOTE

Squalomix: shark and ray genome analysis consortium and its data sharing platform [version 1; peer review: 2 approved]

Osamu Nishimura1*, John Rozewicki1*, Kazuaki Yamaguchi1, Kaori Tatsumi1, Yuta Ohishi1, Tazro Ohta2, Masaru Yagura3, Taiki Niwa3,4, Chiharu Tanegashima1, Akinori Teramura5, Shotaro Hirase5, Akane Kawaguchi3, Milton Tan6, Salvatore D’Aniello7, Filipe Castro8,9, André Machado8, Mitsumasa Koyanagi10, Akihisa Terakita10, Ryo Misawa11, Masayuki Horie12, Junna Kawasaki13, Takashi Asahida14, Atsuko Yamaguchi15, Kiyomi Murakumo16, Rui Matsumoto16, Iker Irisarri17, Norio Miyamoto18, Atsushi Toyoda19, Sho Tanaka20, Tatsuya Sakamoto21, Yasuko Semba22, Shinya Yamauchi23, Kazuyuki Yamada24, Kiyonori Nishida25, Itsuki Kiyatake25, Keiichi Sato16, Susumu Hyodo26, Mitsutaka Kodota1, Yoshinobu Uno27, Shigehiro Kuraku1,3,4

1Laboratory for Phyloinformatics, RIKEN Center for Biosystems Dynamics Research, Kobe, Hyogo, 657-0024, Japan
2Joint Support-Center for Data Science Research, Database Center for Life Science, Mishima, Shizuoka, 411-8540, Japan
3Molecular Life History Laboratory, National Institute of Genetics, Mishima, Shizuoka, 411-8540, Japan
4Department of Genetics, Sokendai (Graduate University for Advanced Studies), Mishima, Shizuoka, Japan
5Fisheries Laboratory, Graduate School of Agricultural and Life Sciences, University of Tokyo, Hamamatsu, Shizuoka, 431-0214, Japan
6Illinois Natural History Survey, Prairie Research Institute, University of Illinois at Urbana-Champaign, Champaign, Illinois, USA
7Biology and Evolution of Marine Organisms, Stazione Zoologica Anton Dohrn, Villa Comunale, Napoli, Italy
8Interdisciplinary Centre of Marine and Environmental Research, University of Porto, Porto, Portugal
9Faculty of Sciences, University of Porto, Porto, Portugal
10Department of Biology, Graduate School of Science, Osaka Metropolitan University, Osaka, Osaka, Japan
11Japan Fisheries Research and Education Agency, Hachinohe, Aomori, Japan
12Graduate School of Veterinary Science, Osaka Metropolitan University, Izumisano, Osaka, Japan
13Waseda Research Institute for Science and Engineering, Waseda University, Tokyo, Japan
14School of Marine Biosciences, Kitasato University, Sagamihara, Kanagawa, Japan
15Graduate School of Fisheries and Environmental Sciences, Nagasaki University, Nagasaki, Nagasaki, Japan
16Okinawa Churaumi Aquarium, Motobu, Okinawa, Japan
17Centre for Molecular Biodiversity Research, Leibniz Institute for the Analysis of Biodiversity Change (LIB), Museum of Nature-Zoology, Hamburg, 20146, Germany
18X-STAR, Japan Agency for Marine-Earth Science and Technology, Yokosuka, Kanagawa, Japan
19Comparative Genomics Laboratory, National Institute of Genetics, Mishima, Shizuoka, 411-8540, Japan
20School of Marine Science and Technology, Tokai University, Shizuoka, Shizuoka, Japan
21Ushimado Marine Institute, Graduate School of Natural Science and Technology, Okayama University, Setouchi, Japan., Okayama, Japan
22Highly Migratory Resources Division, Fisheries Resources Institute, Japan Fisheries Research and Education Agency, Shizuoka, Shizuoka, Japan
23Aquamarine Fukushima, Iwaki, Fukushima, Japan
24Marine Science Museum, Tokai University, Shizuoka, Shizuoka, Japan
25Osaka Aquarium Kaiyukan, Osaka, Osaka, Japan
26Laboratory of Physiology, Atmosphere and Ocean Research Institute, University of Tokyo., Kashiwa, Chiba, Japan
27Department of Life Sciences, Graduate School of Arts and Sciences, The University of Tokyo, Tokyo, Tokyo, Japan
Abstract
The taxon Elasmobranchii (sharks and rays) contains one of the long-established evolutionary lineages of vertebrates with a tantalizing collection of species occupying critical aquatic habitats. To overcome the current limitation in molecular resources, we launched the Squalomix Consortium in 2020 to promote a genome-wide array of molecular approaches, specifically targeting shark and ray species. Among the various bottlenecks in working with elasmobranchs are their elusiveness and low fecundity as well as the large and highly repetitive genomes. Their peculiar body fluid composition has also hindered the establishment of methods to perform routine cell culturing required for their karyotyping. In the Squalomix consortium, these obstacles are expected to be solved through a combination of in-house cytological techniques including karyotyping of cultured cells, chromatin preparation for Hi-C data acquisition, and high fidelity long-read sequencing. The resources and products obtained in this consortium, including genome and transcriptome sequences, a genome browser powered by JBrowse2 to visualize sequence alignments, and comprehensive matrices of gene expression profiles for selected species are accessible through https://github.com/Squalomix/info.

Keywords
Shark, ray, chimaera, biodiversity genomics, whole genome sequencing, karyotype

This article is included in the Genomics and Genetics gateway.
Corresponding author: Shigehiro Kuraku (skuraku@nig.ac.jp)

Author roles: Nishimura O: Data Curation, Methodology, Resources; Rozewicki J: Formal Analysis, Resources, Software; Yamaguchi K: Formal Analysis; Tatsumi K: Methodology, Resources; Ohishi Y: Formal Analysis; Ohta T: Formal Analysis; Yagura M: Formal Analysis; Niwa T: Formal Analysis; Tanegashima C: Methodology, Resources; Teramura A: Formal Analysis; Hirase S: Formal Analysis; Kawaguchi A: Formal Analysis; Tan M: Formal Analysis; D’Aniello S: Formal Analysis; Castro F: Formal Analysis; Machado A: Formal Analysis; Koyanagi M: Formal Analysis; Terakita A: Formal Analysis; Misawa R: Resources; Horie M: Resources; Kawasaki J: Resources; Asahida T: Supervision; Yamaguchi A: Supervision; Murakumo K: Resources; Matsumoto R: Resources; Irisarri I: Supervision; Miyamoto N: Resources; Toyoda A: Data Curation, Formal Analysis; Tanaka S: Supervision; Sakamoto T: Resources; Semba Y: Resources; Yamauchi S: Resources; Yamada K: Resources; Nishida K: Resources, Supervision; Kiyatake I: Resources; Sato K: Resources, Supervision; Hyodo S: Supervision; Kadota M: Data Curation, Methodology, Resources; Uno Y: Formal Analysis; Kuraku S: Conceptualization, Data Curation, Funding Acquisition, Project Administration, Supervision, Writing – Original Draft Preparation, Writing – Review & Editing

Competing interests: No competing interests were disclosed.

Grant information: The consortium is funded by intramural budgets granted by RIKEN and the National Institute of Genetics, Japan, as well as JSPS KAKENHI Grant Numbers 20H03269 and 16H06279 (PAGS).

The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

Copyright: © 2022 Nishimura O et al. This is an open access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

How to cite this article: Nishimura O, Rozewicki J, Yamaguchi K et al. Squalomix: shark and ray genome analysis consortium and its data sharing platform [version 1; peer review: 2 approved] F1000Research 2022, 11:1077 https://doi.org/10.12688/f1000research.123591.1

First published: 21 Sep 2022, 11:1077 https://doi.org/10.12688/f1000research.123591.1
Introduction
Although usually recognized as a kind of ‘fish’ like actinopterygian fishes, cartilaginous fishes (chondrichthyans) form a distinct class of vertebrates with more than 1,200 species, known mostly as sharks and rays (Figure 1; Nelson et al., 2016). This taxonomic class has the longest evolutionary history among vertebrates of about 400 million years, in terms of the divergence of extant members (Naylor et al., 2012). Whereas its diversity might not be widely recognized, species in this taxon are characterized by several unique traits including electromagnetic sensing (all cartilaginous fishes), electricity generation (electric rays), diverse morphology sometimes with a flattened body (angelsarks and most rays) and/or a toothed rostrum (sawsharks and sawfishes). The highlight of their biological enigmas is in their reproductive modes with high plasticity between oviparity and viviparity, and occasionally parthenogenesis and intersexuality (Penfold and Wyffels, 2019). Mainly because of overfishing, many cartilaginous fish populations are declining (Pacoureau et al., 2021), and evidence-based resource management would greatly benefit from the establishment of genomic platforms.

Despite these outstanding evolutionary and biological importance, modern genomic approaches have only recently been applied to cartilaginous fishes (reviewed in Kuraku, 2021). The only exception is the effort commenced before 2010 on the elephant fish Callorhinchus milii (Venkatesh et al., 2014), a member of the Holocephali (chimaeras and ratfishes), the more species-poor chondrichthyan lineage, with a relatively small genome size of about 1.9 giga basepairs (Gbp). In contrast, most elasmobranchs have genomes of more than 3 Gbp plagued with abundant repetitive elements.

Figure 1. Chondrichthyan phylogeny and taxon sampling in the Squalomix Consortium. This figure includes some chondrichthyan species selected to represent the individual taxonomic orders that reflect the local fauna of Japan and are/will be analyzed by the consortium by genome or transcriptome sequencing (as of April 10, 2022). The full list of species and current status can be found in https://github.com/Squalomix/info.
Squalomix: consortium scope and organization

The Squalomix Consortium (Figure 2A) was launched in 2020 aiming to provide the genome sequence and other genome-wide data for chondrichthyan species including transcriptomes and epigenomes. Sample processing and data production is conducted by the Molecular Life History Laboratory at the National Institute of Genetics, Mishima, Japan, and the Laboratory for Phyloinformatics in RIKEN Kobe, Japan, which harbors a DNA Analysis Facility. The consortium is funded by academic agencies as of May 2022 and is seeking additional funding sources, especially from industrial groups oriented toward the conservation of biodiversity and marine environments. In November 2020, the Squalomix Consortium became affiliated with Earth BioGenome Project (EBP), the global initiative to promote biodiversity genomics (Lewin et al., 2022). The collaborative network at the Squalomix Consortium includes an extensive range of expertise and worldwide distribution.

Versatile sample collection featuring the local fauna

In Squalomix, sample collection is performed cautiously to minimize the sacrifice of wildlife—especially those with an endangered status. The collection focuses mainly on the rich marine fauna in Japan’s neighboring temperate waters, with occasional sources from death stranding for elusive species. The project collaborates closely with local aquariums oriented toward academic science. Their contributions play indispensable roles in relaying offshore sampling and enable sustainable sampling of embryos and blood from live individuals, although the latter approach is limited to species that can be bred in captivity and are amenable to husbandry.

Another strength of the Squalomix Consortium is its expertise in laboratory solutions that are not confined to DNA sequencing, but additionally explore post-genome approaches to decipher the molecular basis of chondrichthyan phenotypic evolution. Access to fresh tissues from local aquaria facilitates embryological analysis, genome size quantification with flow cytometry, and karyotyping from cell cultures (Figure 3). Remarkably, cell culture in cartilaginous fishes, which was long thought difficult because of their high body fluid osmolarity, was enabled by modifying the culture medium with balancing osmolytes (Uno et al., 2020). Our cytological expertise also allowed various epigenomic analyses that benefit from whole genome sequencing, on transcription factor binding with ChIP-seq (Hara et al., 2018) and chromatin openness with ATAC-seq, in addition to long-range DNA interactions with Hi-C (Kadota et al., 2020; Onimaru et al., 2021). These techniques contributed to biological analyses based on the draft genome sequences of three shark species (Hara et al., 2018), which launched the Squalomix Consortium.

Sequencing strategy and recent progress

The sequencing strategy in the Squalomix Consortium is designed to accommodate genomic characteristics of cartilaginous fishes, mostly with large, repetitive genomes. In the standard protocol formulated in January 2021 (Figure 3), we start by estimating genome size using flow cytometry and karyotyping as well as by ‘survey’ sequencing of transcriptomes, which serves for species identity verification with an assembled mitochondrial DNA sequence. These initial steps ensure sample authenticity and quality. We then proceed to genome sequencing, which employs both short-read and long-read high-fidelity (‘HiFi’) sequencing platforms, together with Hi-C data production for chromosome-scale scaffolding based on three-dimensional DNA interactions. The long-read data are obtained using the Sequel II or IIe platforms (Pacific Biosciences, Inc.) with a minimum sequencing depth of 20x. The assembly outputs are evaluated with reference to their coverage of protein-coding gene space, as well as transcriptome data, genome size, and karyotypic organization obtained separately. These validations allow us to scrutinize the inclusion of those genomic regions that are difficult to sequence and assemble, such as the Hox C genes that were previously thought to be missing in elasmobranchs but were retrieved by elaborate annotation (Hara et al., 2018; reviewed in Kuraku, 2021). Complete genome assemblies are critical to validate gene loss and variations in gene repertoires via syntenic/phylogeny comparisons, previously suggested for.
visual opsins and conventional olfactory receptors (Hara et al., 2018). The standard procedure outlined above (Figure 3) has been applied to several study species, including the red stingray Hemitrygon akajei (Figure 2B) for which a draft genome assembly has been made available for BLAST searches at the Squalomix sequence archive (Figure 4A; https://transcriptome.riken.jp/squalomix/).

Cooperation toward the global goals
The Squalomix Consortium aims not only to sequence and analyze the genomes but also to tightly interact with other research groups whose target species list contains cartilaginous fishes including other EBP-affiliated projects (see below). To maximize mutual benefit among those projects, some animal samples from our collection could be provided for genome sequencing at other sites. The Squalomix Consortium offers laboratory experiments for genome size quantification or karyotype analysis for species listed by other consortia, provided that fresh cells are available. The sample transfer will be processed in accordance with the Nagoya Protocol and other relevant regulations. Inclusive cooperation respecting complementary expertise is expected to overcome the long-standing difficulty in studying elasmobranchs sustainably and contribute to disentangling the marine ecosystems for effective conservation.

Data sharing platforms
Once produced, genome assemblies pass rigid quality controls and are deposited in the NCBI Genome under the NCBI BioProject ID PRJNA707598 and made available as database for BLAST searches at our Squalomix sequence archive (https://transcriptome.riken.jp/squalomix/). This archive also has a link to the up-to-date listing of the species for which genome sequences are available, filed by the GenomesSync database (http://genomesync.org/). The archive website also hosts a gateway to genome browsers powered by JBrowse2 that allow users to visualize specific genomic regions and load additional tracks including base composition, gene models, repetitive elements, and aligned RNA-seq reads.
We also provide comprehensive matrices of expression profiles for predicted genes of the brownbanded bamboo shark *Chiloscyllium punctatum* and the cloudy catshark *Scyliorhinus torazame* that were already quantified and normalized based on RNA-seq data of various tissues for our past publication (Hara et al., 2018).

Other pioneering efforts tackling elasmobranch genomes

Some elasmobranch genomes have already been sequenced by other pioneering working groups (https://www.ncbi.nlm.nih.gov/data-hub/genome/?taxon=7777&reference_only=true). This includes the Vertebrate Genomes Project (VGP), whose data production format employs a suite of modern promising solutions including optical mapping and Hi-C scaffolding as well as long-read and short-read sequencing, to cover all vertebrate species (Rhie et al., 2021). The initial VGP progress report released the genome sequences of the thorny skate *Amblyraja radiata* (NCBI Genome ID, GCA_010909765.2). The Darwin Tree of Life (DToL) Project partly links with VGP and aims to sequence all eukaryotic species in Britain and Ireland. DToL’s first chondrichthyan genome is that of the small-spotted catshark *Scyliorhinus canicula*, the egg-laying species most widely studied in developmental biology and endocrinology (NCBI Genome ID, GCA_902713615.1). The recently launched European Reference Genome Atlas (ERGA) also plans to produce reference chromosome anchored genomes of multiple species from this geography including cartilaginous fish aiming to empower conservation efforts (Formenti et al., 2022). Researchers in China launched the Fish10K project that partially targets cartilaginous fishes (Fan et al., 2020). In addition, the DNA Zoo project puts special emphasis on Hi-C scaffolding (Rao et al., 2014), often using available genome assemblies already released by other groups as input and performing chromosome-scale genome scaffolding using Hi-C data even in the presence of intra-specific genomic variations. So far, the DNA Zoo effort produced the chromosome-scale genome assemblies of the brownbanded bamboo shark *C. punctatum* and the whale shark *Rhincodon typus*, each of which was produced using samples from multiple individuals (Hoencamp et al., 2021). All the above efforts are expected to be coordinated under the overarching EBP initiative, in order to play complementary roles towards the global aim of generating high-quality genomic resources.
Data availability
Products from this consortium are deposited in NCBI under the BioProject ID PRJNA707598 and are available at our Squalomix sequence archive (https://transcriptome.riken.jp/squalomix/).

Acknowledgments
The authors representing the Squalomix Consortium thank the animal caretakers and administrative staff at the aquaria and the DNA sequencing facilities that are assisting the consortium. Computations were partially performed on the NIG supercomputer at ROIS National Institute of Genetics.

References

Bueks R, Eric Y, Desch CM, et al.: JBrowse: a dynamic web platform for genome visualization and analysis. Gen. Biol. 2016; 17: 66.
PubMed Abstract | Publisher Full Text
Fan G, Song Y, Yang J, et al.: Initial data release and announcement of the 10,000 Fish Genomes Project (Fish10K). GigaScience. 2020; 9: giaa080.
PubMed Abstract | Publisher Full Text
Formenti G, et al.: The era of reference genomes in conservation genomics. Trends Genet. 2022; 38: 197–202.
PubMed Abstract | Publisher Full Text
Hara Y, Yamaguchi K, Onimaru K, et al.: Shark genomes provide insights into elasmobranch evolution and the origin of vertebrates. Nat. Ecol. Evol. 2018; 2: 1761–1771.
PubMed Abstract | Publisher Full Text
Hoencamp, et al.: 3D genomics across the tree of life reveals condensin II as a determinant of architecture type. Science. 2021; 372: 984–989.
PubMed Abstract | Publisher Full Text
Kadota M, Nishimura O, Miura H, et al.: Multifaceted Hi-C benchmarking: what makes a difference in chromosome-scale genome scaffolding? Gigascience. 2020; 9: giz158.
PubMed Abstract | Publisher Full Text
Kuraku S: Shark and ray genomics for disentangling their morphological diversity and vertebrate evolution. Dev. Biol. 2021; 477: 262–272.
PubMed Abstract | Publisher Full Text
Kuraku S, Zmasek CM, Nishimura O, et al.: aLeaves facilitates on-demand exploration of metazoan gene family trees on MAFFT sequence alignment server with enhanced interactivity. Nuc. Acids Res. 2013; 41: W22–W28.
Publisher Full Text | Reference Source
Lee, et al.: The Earth BioGenome Project 2020: Starting the clock. Proc. Natl. Acad. Sci. USA. 2022; 119: e2115635118.
PubMed Abstract | Publisher Full Text
Naylor GJP, Cairn JN, Jensen K, et al.: Elasmobranch Phylogeny: A mitochondrial estimate based on 595 species. Carrier JC, Musick JA, Heithaus MR, editors. The Biology of Sharks and Their Relatives. Boca Raton: CRC Press, Taylor & Francis Group; 2012; pp. 31–56.
Publisher Full Text
Nelson JS, Grande T, Wilson MVH: Fishes of the world. Fifth ed. Hoboken, New Jersey: John Wiley & Sons; 2016; p. 1 online resource.
Onimaru K, Tatsunami K, Tanegashima C, et al.: Developmental hourglass and heterochronic shifts in fin and limb development. eLife. 2021; 10: e62865.
PubMed Abstract | Publisher Full Text
Pacoureau N, Rigby CL, Kyne PM, et al.: Half a century of global decline in oceanic sharks and rays. Nature. 2021; 589: 567–571.
PubMed Abstract | Publisher Full Text
Penfold LM, Wyffels JT: Reproductive Science in Sharks and Rays. Adv. Exp. Med. Biol. 2019; 1200: 465–488.
PubMed Abstract | Publisher Full Text
Rao SS, Huntley MH, Durand NC, et al.: A 3D map of the human genome at kilobase resolution reveals principles of chromatin looping. Cell. 2014; 159: 1665–1680.
PubMed Abstract | Publisher Full Text
Rhee A, McCarthy SA, Fedrigo O, et al.: Towards complete and error-free genome assemblies of all vertebrate species. Nature. 2021; 592: 737–746.
PubMed Abstract | Publisher Full Text
 Uno Y, Nozu R, Kiyatake I, et al.: Cell culture-based karyotyping of orectolobiform sharks for chromosome-scale genome analysis. Commun. Biol. 2020; 3: 652.
PubMed Abstract | Publisher Full Text
Venkatesh B, Lee AP, Ravi V, et al.: Elephant shark genome provides unique insights into gnathostome evolution. Nature. 2014; 565: 174–179.
PubMed Abstract | Publisher Full Text
Yamaguchi K, Koyanagi M, Kuraku S: Visual and nonvisual opsin genes of sharks and other nonosteichthyan vertebrates: genomic exploration of underwater photoreception. J. Evol. Biol. 2020; 34: 968–976.
Publisher Full Text
I greet this initiative with great enthusiasm. The description is well written, clear and easy to follow. I have just a few comments that I hope the authors will consider.

In the introduction, the authors describe Chondrichthyes as the oldest vertebrate class ("longest evolutionary history"). However, this is due to the imprecise use of the term "class" in vertebrate taxonomy where both Chondrichthyes and Mammalia are designated as classes. Thus, classes are not of equal temporal rank. Furthermore, the authors' statement is not quite true, because even Agnatha has the taxonomic rank as a vertebrate class and would thereby be even earlier than Chondrichthyes. I would recommend the authors to describe Chondrichthyes instead as one of the two lineages resulting from the first bifurcation or divergence in (the infraphylum of) Gnathostomata (jawed vertebrates). This, by the way, means that Osteichthyes is as old as Chondrichthyes!

Please correct the grammar of the expression "Despite these outstanding evolutionary and biological importance...". It probably needs to be rephrased, perhaps like this: "Despite the outstanding evolutionary and biological importance of chondrichthynes..." (or elasmobranchs if you prefer to focus on these).

Shouldn't "giga basepairs" be one word as one would surely write megabasepairs and kilobasepairs.

Figure 1: The expression "Other osteichthynes" is imprecies. I assume the authors want to avoid the term "Sarcopterygians" for this group pterygi means fins, and tetrapods of course don't have them. Maybe it's better to just delete "Other osteichthynes" and let this branch be called "Tetrapods, lungfishes and coelacanths" (without capital initial letters).
Is the rationale for creating the dataset(s) clearly described?
Yes

Are the protocols appropriate and is the work technically sound?
Yes

Are sufficient details of methods and materials provided to allow replication by others?
Yes

Are the datasets clearly presented in a useable and accessible format?
Yes

Competing Interests: No competing interests were disclosed.

Reviewer Expertise: Gene family evolution, pharmacology of G protein-coupled receptors, mechanism of long-term memory

We confirm that we have read this submission and believe that we have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

Reviewer Report 26 September 2022
https://doi.org/10.5256/f1000research.135712.r151220

© 2022 Debiais-Thibaud M. This is an open access peer review report distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Mélanie Debiais-Thibaud
Institut des Sciences de l’Evolution de Montpellier, ISEM, Université Montpellier, Montpellier, France

In this Data note, the authors describe and wrap-up all available material generated through their consortium named Squalomix, in which a set of biological material and sequence data obtained in elasmobranch organisms are made available to the research community. The rationale, protocol, material and data availability are clearly described in this Note.

Is the rationale for creating the dataset(s) clearly described?
Yes

Are the protocols appropriate and is the work technically sound?
Yes

Are sufficient details of methods and materials provided to allow replication by others?
Yes
Are the datasets clearly presented in a useable and accessible format? 
Yes

**Competing Interests:** No competing interests were disclosed.

**Reviewer Expertise:** Developmental genetics, EvoDevo

I confirm that I have read this submission and believe that I have an appropriate level of expertise to confirm that it is of an acceptable scientific standard.

The benefits of publishing with F1000Research:

- Your article is published within days, with no editorial bias
- You can publish traditional articles, null/negative results, case reports, data notes and more
- The peer review process is transparent and collaborative
- Your article is indexed in PubMed after passing peer review
- Dedicated customer support at every stage

For pre-submission enquiries, contact research@f1000.com