# **Linking the International Wheat Genome Sequencing Consortium**

# <sup>2</sup> bread wheat reference genome sequence to wheat genetic and

# 3 phenomic data

4

5	Michael Alaux <sup>1</sup> *, Jane Rogers <sup>2</sup> , Thomas Letellier <sup>1</sup> , Raphaël Flores <sup>1</sup> , Françoise Alfama <sup>1</sup> , Cyril
6	Pommier <sup>1</sup> , Nacer Mohellibi <sup>1</sup> , Sophie Durand <sup>1</sup> , Erik Kimmel <sup>1</sup> , Célia Michotey <sup>1</sup> , Claire Guerche <sup>1</sup> ,
7	Mikaël Loaec <sup>1</sup> , Mathilde Lainé <sup>1</sup> , Delphine Steinbach <sup>1†</sup> , Frédéric Choulet <sup>3</sup> , Hélène Rimbert <sup>3</sup> ,
8	Philippe Leroy <sup>3</sup> , Nicolas Guilhot <sup>3</sup> , Jérôme Salse <sup>3</sup> , Catherine Feuillet <sup>3</sup> ‡, International Wheat
9	Genome Sequencing Consortium <sup>2</sup> , Etienne Paux <sup>3</sup> , Kellye Eversole <sup>2</sup> , Anne-Françoise Adam-
10	Blondon <sup>1</sup> , Hadi Quesneville <sup>1</sup>
11	
12	<sup>1</sup> URGI, INRA, Université Paris-Saclay, 78026, Versailles, France.
13	<sup>2</sup> International Wheat Genome Sequencing Consortium, 2841 NE Marywood Ct, Lee's
14	Summit, MO 64086, USA.
15	<sup>3</sup> GDEC, INRA, Université Clermont Auvergne, 63000, Clermont-Ferrand, France.
16	† current address: GQE-Le Moulon UMR 320, INRA, Université Paris-Sud, Université Paris-
17	Saclay, CNRS, AgroParisTech, Ferme du Moulon, 91190, Gif-sur-Yvette, France.
18	‡ current address: Bayer CropScience, 3500 Paramount Pkwy., Morrisville, NC 27560, USA.
19	*Corresponding author. E-mail:michael.alaux@inra.fr
20	
21	

# 22 Keywords

23 data integration - information system - big data - wheat genomics, genetics and phenomics

24

25

## 26 Abstract

The Wheat@URGI portal (https://wheat-urgi.versailles.inra.fr) has been developed to provide 27 28 the international community of researchers and breeders with access to the bread wheat 29 reference genome sequence produced by the International Wheat Genome Sequencing Consortium. Genome browsers, BLAST, and InterMine tools have been established for in 30 depth exploration of the genome sequence together with additional linked datasets including 31 physical maps, sequence variations, gene expression, and genetic and phenomic data from 32 other international collaborative projects already stored in the GnpIS information system. The 33 34 portal provides enhanced search and browser features that will facilitate the deployment of the latest genomics resources in wheat improvement. 35

- 36
- 37

## 38 Background

The International Wheat Genome Sequencing Consortium (IWGSC) [1] is an international 39 40 collaborative group of growers, academic scientists, and public and private breeders that was established to generate a high quality reference genome sequence of the hexaploid bread 41 wheat, and to provide breeders with state-of-the-art tools for wheat improvement. The vision 42 43 of the consortium is that the high quality, annotated ordered genome sequence integrated with physical maps will serve as a foundation for the accelerated development of improved 44 varieties and will empower all aspects of basic and applied wheat science to address the 45 important challenge of food security. A first analysis of the reference sequence produced by 46 the consortium (IWGSC RefSeq v1.0) was recently published [2]. 47

To ensure that wheat breeding and research programs can make the most of this extensive 48 genomic resource, the IWGSC endorsed the establishment of a data repository at URGI (Unité 49 de Recherche Génomique Info / research unit in genomics and bioinformatics) from INRA 50 (Institut National de la Recherche Agronomique / French national institute for agricultural 51 research) to develop databases and browsers with relevant links to public data available 52 worldwide. The IWGSC data repository is thus hosted by URGI to support public and private 53 parties in data management as well as analysis and usage of the sequence data. Wheat 54 functional genomics (expression, methylation, etc.), genetic, and phenomic data has increased 55 concurrently, requiring the development of additional tools and resources to integrate 56 different data for biologists and breeders. To manage this escalation of data, URGI have built 57 this data repository for the wheat community with the following specific aims: (i) store 58 59 resources for which no public archive exists (e.g. physical maps, phenotype information); (ii) enable pre-publication access to specific datasets (e.g. sequence assemblies and annotations, 60 physical maps, markers); and (iii) rapid release of integrated resources upon publication. The 61 62 repository has been designed in accordance with the "FAIR" principles [3] to ensure that the data are Findable, Accessible, Interoperable and Reusable. To address the challenge of 63 64 integrating diverse data types from multiple sources, URGI employs solutions that provide enhanced features for data exploration, mining and visualisation using the GnpIS information 65 system [4] combined with a high level of data interoperability. 66

Here we describe the data and tools currently available through the Wheat@URGI portal [5], the primary resource for the reference sequence of the bread wheat genome (IWGSC RefSeq v1.0) and other IWGSC wheat genomic data. The links to functional genomics, genetic and phenomic data from many other large wheat projects are also described.

- 71
- 72

## 73 A large wealth of data is available throughout the Wheat@URGI

## 74 portal

The data hosted by the Wheat@URGI portal are available through flat files stored in the IWGSC data repository and through the GnpIS information system [4]. GnpIS encompasses a set of integrated databases to manage genomic data using well-known tools such as BLAST, JBrowse, GBrowse and InterMine, and an in-house database called GnpIS-coreDB developed by URGI to manage genetic and phenomic data.

80

### 81 IWGSC data

Through its concerted efforts to achieve a high quality, functionally annotated reference wheat genome sequence, the IWGSC has developed a variety of resources for the bread wheat (*Triticum aestivum L.*) accession Chinese Spring. The IWGSC data hosted in the Wheat@URGI portal within the IWGSC data repository are shown in Table 1. They fall into four broad categories: (i) physical maps, (ii) sequence assemblies and annotations, (iii) gene expression, and (iv) variation data.

Physical maps: physical maps assembled by IWGSC scientists for the 21 bread wheat 88 chromosomes, based on high information content fluorescence fingerprinting (HICF) [6] or 89 whole genome profiling (WGP<sup>TM</sup>) [7] of flow-sorted chromosome or chromosome-arm specific 90 91 BAC libraries, are stored and displayed. The positions of individual BAC clones, markers, and deletion bins are mapped onto physical contigs. The database maintains all released versions 92 of each physical map with the software used to produce the BAC clone assemblies (FPC [8] or 93 LTC [9]), information from the group that produced the map and a link to order the BAC clones 94 from the French plant genomic resource centre [10]. 95

Sequence assemblies and annotations: the IWGSC wheat genome sequence assemblies 96 available for download, BLAST [11], and display in genome browsers include the draft survey 97 sequence assemblies released in 2014 (IWGSC Chromosome Survey Sequence (CSS) v1) and 98 two improved versions (CSS v2 and v3) [12], and the chromosome 3B reference sequence (the 99 first reference quality chromosome sequence obtained by the consortium) [13]. Associated 100 101 with these assemblies are the virtual gene order map generated for the CSS (Genome Zipper), 102 the POPSEQ data used to order sequence contigs on chromosomes [14] and mapped marker sets. The reference sequence of the bread wheat genome (IWGSC RefSeg v1.0, 14.5 Gb 103 assembly with super scaffold N50 of 22.8Mb) was obtained by integrating whole genome 104 shotgun Illumina short reads assembled with NRGene's DeNovoMAGIC<sup>™</sup> software with the 105 wealth of IWGSC map and sequence resources [2]. The IWGSC RefSeq v1.0 is available for 106 download, BLAST, and browser display. Users can access the whole genome, pseudomolecules 107 108 of individual chromosomes or chromosome arms, and scaffolds with the structural and functional annotation of genes, transposable elements, and non-coding RNAs generated by 109 110 the IWGSC. In addition, mapped markers as well as alignments of nucleic acid and protein evidence supporting the annotation are available. Updated versions of the annotation for 111 genes belonging to specific gene families or regions of specific chromosomes that have been 112 113 manually annotated (ca. 3685 genes) can be found in the IWGSC RefSeq v1.1 annotation. 114 In addition to the bread wheat sequence, the IWGSC also assembled seven diploid and tetraploid wheat related species: Triticum durum cv. Cappelli, Triticum durum cv. Strongfield, 115

116 Triticum durum cv Svevo, Triticum monococcum, Triticum urartu, Aegilops speltoides, Aegilops

*sharonensis* [12]. Download and BLAST is available for these data.

*Expression data*: RNA-Seq expression data are available as reads counts and transcripts per
 kilobase million (TPM) for the IWGSC RefSeq v1.1 annotation. It is a transcriptome atlas

- 120 developed from 850 RNA-Seq datasets representing a diverse range of tissues, development
- 121 stages and environmental condition [15].
- 122 Variation data: These data consists of downloadable VCF files from genotyping by sequencing
- and whole exome capture experiments of 62 diverse wheat lines [16] and of the IWGSC
- 124 **3,289,847** inter-varietal SNPs [17].
- 125 <u>Table 1</u>
- 126 IWGSC data summary in open access hosted in the IWGSC Data Repository of the
- 127 Wheat@URGI portal in March 2018.

Data	Details	Tools	Contacts	
IWGSC RefSeq v1.0 assembly	scaffolds, superscaffolfs, pseudomolecules	Download, BLAST and browser	IWGSC	
IWGSC RefSeq v1.0 annotation	genes, transposable elements, ncRNAs, markers, functional annotation, RNA-seq	Download and browser	IWGSC	
IWGSC WGA v0.4	scaffolds, superscaffolfs, pseudomolecules	Download and BLAST	IWGSC	
IWGSC Survey sequence v2 assembly	contigs, gene models, Genome Zipper, POPSEQ	Download and BLAST	IWGSC, Mihaela Martis, Manuel Spannagl, Klaus Mayer, Nils Stein	
IWGSC Survey sequence v2 annotation	genes, markers, physical contigs	Browser	IWGSC, Curtis Pozniak, Eduard Akhunov	
IWGSC Survey sequence v3 assembly	scaffolds	Dowload and browser	Andy Sharpe, David Konkin, Curtis Pozniak	
IWGSC SNPs	intervarietal SNPs	Download	Etienne Paux	
3B reference sequence assembly	contig, scaffolds, pseudomolecule	Download and BLAST	Frédéric Choulet, Etienne Paux	
3B reference sequence annotation	genes, transposable elements, RNAs, markers	Browser	Frédéric Choulet, Etienne Paux	
Other wheat species WGS assemblies	Triticum durum cv. Cappelli, Triticum durum cv. Strongfield, Triticum monococcum, Aegilops speltoides, Aegilops Sharonensis, Triticum urartu, Aegilops tauschii	Download and BLAST	Jon Wright, Mario Caccamo	
Transcriptome	Deep transcriptome sequencing	Download	Lise Pingault, Etienne Paux	
	Triticum urartu and Triticum turgidum (Graingenes)	Download	Jorge Dubcovsky	
Variations	GBS and WEC	Download	Eduard Akhunov	
Physical maps	1AS v1 and v2	Download and browser	James Breen, Thomas Wicker, Beat Keller	
	1AL v1 and v2	Download and browser	Stuart Lucas, Hikmet Budak	
	2AS	Download and browser	Kuldeep Singh	
	2AL	Download and browser	Kuldeep Singh	

3AS v1	Download and browser	Sunish Sehgal, Bikram Gill
3AS v2	Download and browser	Sunish Sehgal, Bikram Gill
3AL	Download and browser	Vijay Kumar Tiwari
4AS	Download and browser	Miroslav Valarik, Jaroslav Dolezel
4AL v1 and v2	Download and browser	Miroslav Valarik, Jaroslav Dolezel
5AS	Download and browser	Simone Scalabrin
5AL	Download and browser	Simone Scalabrin
6AS	Download and browser	Naser Poursarebani
6AL	Download and browser	Naser Poursarebani
7AS	Download and browser	Gabriel Keeble-Gagnere
7AL	Download and browser	Gabriel Keeble-Gagnere
1BS v1, v2, v3 and v5	Download and browser	Dina Raats, Zeev Frenkel, Abraham Korol
1BL v1 and v2	Download and browser	Etienne Paux
2BS	Download and browser	John Jacobs
2BL	Download and browser	John Jacobs
3B	Download and browser	Etienne Paux
4BS	Download and browser	John Jacobs
4BL	Download and browser	John Jacobs
5BS	Download and browser	Elena Salina
5BL	Download and browser	John Jacobs
6BS v1 and v2	Download and browser	Fuminori Kobayashi, Hirokazu Handa
6BL v1 and v2	Download and browser	Fuminori Kobayashi, Hirokazu Handa
7BS	Download and browser	Tatiana Belova, Odd-Arne Olsen
7BL	Download and browser	Tatiana Belova, Odd-Arne Olsen
1D	Download and browser	Bikram Gill, Sunish Sehgal, Vijay Kumar Tiwari
2DS	Download and browser	John Jacobs
2DL	Download and browser	John Jacobs
3DS v1 and v2	Download and browser	Jan Bartos, Jaroslav Doleze
3DL	Download and browser	Jon Wright, Mario Caccam Mike Bevan
4D	Download and browser	Bikram Gill, Sunish Sehgal, Vijay Kumar Tiwari
		Hikmet Budak. Bala Ani
5DS	Download and browser	Akpinar
5DS 5DL	Download and browser Download and browser	Akpinar John Jacobs
	Download and browser Download and browser	Akpinar John Jacobs
5DL	Download and browser	Akpinar John Jacobs Bikram Gill, Sunish Sehgal,

128 Enquiries about these data should be addressed to <u>communications@wheatgenome.org</u> and

129 urgi-contact@inra.fr.

130

#### 132 Wheat gene pool

As well as IWGSC resources, URGI also hosts other open access wheat sequence data to facilitate research into the wheat gene pool. Sequence assemblies available for download and BLAST include the bread wheat whole genome sequence assembly *Triticum aestivum* TGACv1 [18] and the diploid progenitor of *Aegilops tauschii* [19].

137

### 138 Genetic and phenomic data

In addition to sequence data, the Wheat@URGI portal hosts, within GnpIS-coreDB, several sets of genetic and phenomic wheat data [20] that have been produced from French, European, and international projects since 2000 [21]. A significant amount of these data is available without restriction. However, access to restricted data can be obtained through a material transfer or intellectual property agreement. Table 2 presents the types and number of genetic and phenomic data hosted in the GnpIS-coreDB database.

Genetic information corresponds to genetically mapped markers, quantitative trait loci (QTLs), genetic resources (germplasms), and genetic studies (genome wide association studies -GWAS). Genomic information consists of variation from SNP discovery experiments, genotyping, comparative genomics (synteny) and expression data (microarray, RNA-Seq). Phenomic data are available as whole trials including phenotypic and environmental observations recorded using ontologies controlled variables with MIAPPE [22] compliant metadata.

Germplasm data were mainly provided by the French small grain cereals genebank maintained by INRA at Clermont-Ferrand [23] but also by partners of several EU projects. They were linked together with related genotyping or phenotyping characterization data. Generally, genetic and phenomic data have been produced by INRA and its partners in large collaborative projects.

#### 157 <u>Table 2</u>

- 158 Genetic and phenomic wheat data summary hosted in the GnpIS-coreDB database of the
- 159 Wheat@URGI portal in March 2018.

Data type	Object	#Total	#Open access	#Restricted access to projects
Genetic Resources	Taxon	56	56	0
Genetic Resources	Accession	12839	10016	2823
	Мар	30	29	1
Genetic Maps	Marker	704822	34164	670658
	QTL	749	465	284
	Sequence Variation	4189312581	90	4189312491
SNP discovery	SNP, indel	724132	95	724037
	Experiment	23	2	21
Genotyping (high throughput)	Sample	8885	47	8872
	Marker	668540	0	668540
	Trial	850	821	29
Phonotyning	Plot	3660	2985	901
Phenotyping	Variable	282	89	195
	Observation	1171172	527981	643191
	Analysis	1555	43	1512
	Sample	2365	1839	526
GWAS	Variable	359	37	322
	Marker	123866	4109	119757
	Association	824217	48596	775621

160 Questions about these data can be addressed to <u>urgi-contact@inra.fr</u>.

161

162

# 163 Browsing and searching a large variety of integrated data

Data can be easily accessed through the Wheat@URGI portal [5] using (i) tabs at the top of the pages allowing access in one click to the data, tools, and projects descriptions as well as the IWGSC data repository, (ii) direct links on the home page to the different data types (e.g. clicking on "Physical maps" opens the physical maps browser), and (iii) data discovery and InterMine [24] tools on the home page.

169 The IWGSC data repository [25] allows accessing consortium data by (i) clicking on a

170 chromosome to open a pop-up menu with all related data (*e.g.* 3A, 3B, etc.), (ii) using the tabs

171	on the left to access the data by type (e.g. Assemblies, Annotations, etc.), or useful links to the
172	news, the BLAST tool, the FAQ, the access status of the data (e.g. open access), etc.
173	

#### 174 **Physical maps browser**

A GBrowse [26] displays the physical maps generated by the IWGSC members [27]. A clickable image on the top of the browser gives access to all versions of the physical map for each chromosome. The browser displays physical contigs, BACs, deletions bins, and markers. From the BACs track, it is possible to order BAC clones directly at the INRA French plant genomic resource centre [10]. From the BACs and markers tracks, one can go directly to the corresponding region in the IWGSC RefSeq v1.0 browser.

181

#### 182 Genome browser and BLAST

183 The IWGSC RefSeq v1.0 is displayed in a dedicated JBrowse [28], [29]. The "markers track" provides links to additional genetic information stored in GnpIS-coreDB which includes access 184 185 to the position of the marker in cM on genetic maps and to the overlapping QTLs. The most popular tool of the IWGSC data repository is the BLAST search tool (476,000 BLAST searches 186 187 launched in 2017). All of the wheat sequences available on the Wheat@URGI portal are 188 indexed for BLAST search (see [30] for the complete list). A set of databanks can be selected: 189 e.g. IWGSC RefSeq v1.0 and IWGSC CSS v3 for a given chromosome. The result is presented in a classical tabular format with (i) links to download the data (matching contigs and high scoring 190 191 pairs - HSP), (ii) links on the genome browsers directly zooming in on the matching region and (iii) external links to EnsemblPlants [31]. 192

193

#### 194 Genetic and phenomic data in GnplS-coreDB

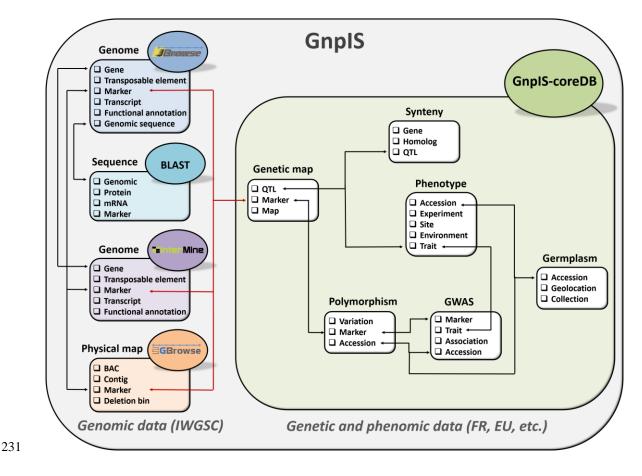
The IWGSC sequence data are linked to genetic and phenomic data within the GnpIS 195 information system [4]. This integration is organized around key data, also called "pivot data" 196 197 as they are pivotal objects which allow integration between data types. The key objects used to link genomic resources to genetic data are markers and QTLs. Markers are mapped on the 198 genome sequences and provides information on neighbour genes and their function. They 199 also have links to GnpIS-coreDB genetic maps, QTLs, genotyping and GWAS data. Additional 200 information on the marker itself can be found regarding the marker type (e.g. SSR, DArT), the 201 primers sequence for PCR amplification, and SNP details (including the flanking sequences) 202 when relevant. QTLs link the genetic data to the phenomic data in GnpIS-coreDB and to 203

synteny data displayed by the PlantSyntenyViewer tool [32], [33].

The accession (i.e. germplasm) and the variables (i.e. observed trait) described with dedicated 205 206 ontologies are another important key data for genetic studies as they allow linking phenotype 207 data to genetic associations or QTLs through traits and to genotype diversity data. The genetic resources stored in GnpIS-coreDB displays the unambiguous identification of the accession 208 209 used (with digital object identifier - DOI) and a rich set of associated data following the MCPD (multi-crop passport descriptors, [34]) standard: a picture, synonyms, descriptors, geolocation 210 211 of the sites (origin, collecting and evaluation), the collections or panels it belongs to, the 212 holding stock centre with a link to order the accession when possible. The phenotype data 213 includes traceability on trials with timing, like year and temporal series, location, and 214 environment including soil and cultural practices. The phenotype and environment variables follow the crop ontologies format [35] that includes unique identifiers for each variable which 215 are composed of a trait description (e.g. grain yield, plant height top, spike per area), a unit 216 and a method. All these data are displayed in the GnpIS-coreDB web interface and can be 217 218 downloaded in different file formats, all compliant with the MIAPPE standard [22].

### 220 Mining and data discovery tools

- To complete this already rich integrated datasets, a gene centric data warehouse, the 221 WheatMine, has been set-up using the well-established InterMine tool [24]. The gene card 222 displays gene function, gene ontology terms, and overlapping genomic features. WheatMine 223 [36] provides access to IWGSC RefSeq v1.0 annotation data (genes, mRNA, polypeptides, 224 transposable elements), polymorphisms (markers) and, through pivotal objects, to genetic 225 data (QTL, metaQTL). It is also possible to navigate from a gene card to its position on the 226 227 wheat genome browser or to relevant marker details in GnpIS-coreDB. Figure 1 presents the concept and the tools to navigate through the key data in GnpIS. 228
- 229 Figure 1
- 230 Conceptual view of wheat data links in GnpIS



233	Finally, to facilitate data search and access to this wealth of data, we developed a data
234	discovery tool, which, similar to a google search, allows the user to enter keywords or terms
235	to find all the matching information in the various data warehouses. The results are presented
236	in a table with details on the matches (database source, type, species, description) and a direct
237	link to the feature (e.g. a gene in a browser, a marker page in GnpIS-coreDB, etc.).
238	A practical use case describing how to use the Wheat@URGI portal to go from a gene
239	sequence to find the genetic studies related is detailed in the Supplementary data.
240	

241

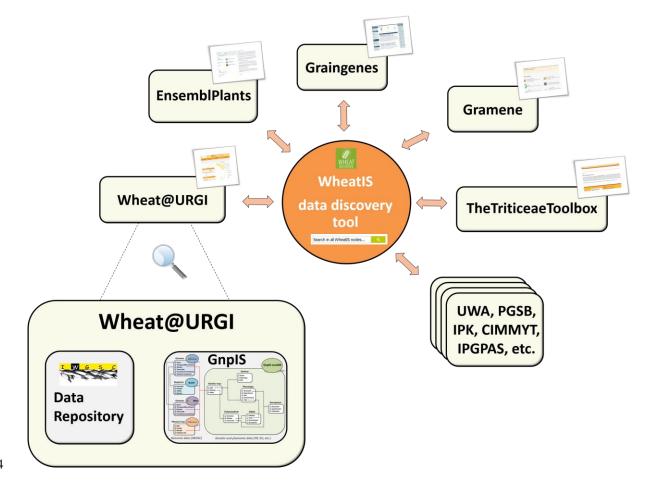
## 242 Conclusion and future directions

The Wheat@URGI portal hosts and gives access to essential, high quality wheat data from the IWGSC, European, and international projects. Furthermore, its added value is that it integrates different data type altogether (genomics, genetics and phenomics) and provides dedicated tools to explore them.

As new wheat resources such as GWAS, genomic selection, and pan-genome data are generated in the frame of ongoing projects, GnpIS will allow their management and integration with other data already available in the information system, linking new upcoming data to this central IWGSC genomic resource.

At a wider scale, an expert working group (EWG) of the international Wheat Initiative build an international wheat information system, called WheatIS, with the aim of providing a singleaccess web-based system to all available wheat data resources and bioinformatics tools [37]. The Wheat@URGI portal is a major node of the WheatIS federation that expose genomic, genetic and phenomic integrated data to the community. The WheatIS data discovery tool allows a one-stop search in GnpIS [4] (including IWGSC browsers, InterMine and GnpIS-

- coreDB; URGI), EnsemblPlants (EMBL-EBI) [31], CrowsNest [38] (PGSB), CR-EST [39], GBIS [40]
- and MetaCrop [41] (IPK), The Triticeae Toolbox (Triticeae CAP), CIMMYT DSpace and
- 259 Dataverse (CIMMYT), Gramene [42] (CSH, OSU, EMBL-EBI), Cropnet (IPGPAS), WheatPan [43]
- 260 (UWA) and GrainGenes [44] (USDA).
- 261 The Figure 2 presents the WheatIS ecosystem.
- 262 Figure 2
- 263 The Wheat@URGI portal node in the WheatIS ecosystem



264

265

Data integration is fundamental for researchers and breeders that want to use genomic information to improve wheat varieties. However, the diversity of data type and the concomitant lack of data harmonisation and standards hamper cross-referencing and metaanalysis. A joint action between the WheatIS EWG and a group of linked data scientists created the Wheat Data Interoperability Working Group under the Research Data Alliance (RDA) 14

- umbrella [45] to help tackle this difficult issue [46]. The Wheat@URGI portal continuously
- evolves its repository to follow the standard recommendations [47].
- 273
- 274

## 275 Abbreviations

- 276 IWGSC: international wheat genome sequencing consortium
- 277 INRA: institut national de la recherche agronomique / French national institute for agricultural
- 278 research
- 279 URGI: unité de recherche génomique info / research unit in genomics and bioinformatics
- 280 **FAIR:** findable, accessible, interoperable, reusable
- 281 **BLAST:** basic local alignment search tool
- 282 HICF: high-information-content fingerprinting
- 283 WGP<sup>TM</sup>: whole genome profiling
- 284 BAC: bacterial artificial chromosome
- 285 **FPC:** fingerprinted contig
- 286 LTC: linear topological contig
- 287 **CSS:** chromosome survey sequence
- 288 **POPSEQ:** population sequencing
- 289 **RNA:** ribonucleic acid
- 290 TPM: transcripts per kilobase million
- 291 VCF: variant call format
- 292 SSR: simple sequence repeats
- 293 SNP: single nucleotide polymorphism
- 294 **DArT:** diversity arrays technology

- **QTL:** quantitative trait loci
- **GWAS:** genome-wide association study
- **cM:** centimorgan
- HSP: high scoring pairs
- 299 PCR: polymerase chain reaction
- 300 DOI: digital object identifier
- 301 MCPD: multi-crop passport descriptors
- **MIAPPE:** minimum information about a plant phenotyping experiment
- **EWG:** expert working group
- **EMBL-EBI:** European bioinformatics institute
- **PGSB:** plant genome and systems biology group
- **IPK:** Leibniz institute of plant genetics and crop plant research
- **CIMMYT:** international maize and wheat improvement center
- **CSH:** Cold Spring Harbor laboratory
- **OSU:** Ohio State University
- **IPGPAS:** institute of plant genetics of the Polish academy of science
- 311 UWA: University of Western Australia
- 312 USDA: U.S. department of agriculture
- **EWG:** expert working group
- **RDA:** research data alliance

## **Declarations**

318 Ethics approval and consent to participate

- 319 Not applicable.
- 320
- 321 **Consent for publication**
- 322 Not applicable.
- 323

#### 324 Availability of data and materials

- 325 The open access data (including all the IWGSC data) are available through the Wheat@URGI
- 326 portal: <u>https://wheat-urgi.versailles.inra.fr</u>.
- 327

#### 328 Competiting interest

- 329 The authors declare that they have no competing interests.
- 330

#### 331 Funding

- 332 The development of the information system and the integration of wheat data was supported
- 333 by INRA and several projects: BreedWheat (ANR-10-BTBR-03, France Agrimer, FSOV), Whealbi
- 334 (EU FP7-613556), TriticeaeGenome (EU FP7-KBBE-212019), 3BSEQ (ANR-09-GENM-025,
- 335 FranceAgrimer), TransPLANT (EU FP7-283496).
- 336

### 337 Authors' contributions

- 338 MA, JR, TL, FA, KE designed, developed and filled the IWGSC data repository.
- 339 MA, TL, RF, FA, CP, NM, SD, EK, CM, CG, MLo, MLa, DS, AFAB, HQ designed, developed and
- 340 filled the GnpIS information system.
- FC, HR, PL, NG, JS, CF, IWGSC, EP generated, submitted the data and give feedback on the
   tools.
- 343 MA, JR, EP, KE, AFAB, HQ draft the manuscript.
  - 17

#### All authors read and approved the final manuscript.

345

### 346 Acknowledgements

- 347 The authors would like to thank for their help or advices at various stages of the project, the
- following people from INRA-URGI: Véronique Jamilloux, Joëlle Amselem, Dorothée Charruaud,
- 349 Guillaume Cornut, Laura Burlot, Florian Philippe, Nicolas Francillonne, Loïc Couderc, Daphné
- 350 Verdelet, Baptiste Brault, Kirsley Chennen; from INRA-GDEC: Jacques Le Gouis, Gilles Charmet,
- 351 François Balfourier, Pierre Sourdille, Catherine Ravel, François-Xavier Oury, Audrey Didier;
- 352 from INRA-DIST: Esther Dzale, Sophie Aubin, Odile Hologne; and from INRA-Agronomie:
- 353 Arnaud Gauffreteau.
- 354 Thanks to Isabelle Caugant (IWGSC), Hélène Lucas (Wheat Initiative), the International Wheat
- 355 Genome Sequencing Consortium and its sponsors, the WheatIS expert working group, the
- 356 URGI platform, and all the data submitters.
- 357
- 358

### 359 **References**

- 1. IWGSC website. http://www.wheatgenome.org/. Accessed 10 April 2018.
- 361 2. IWGSC, 2018, under review.

362 3. Wilkinson MD, Dumontier M, Aalbersberg IJJ, Appleton G, Axton M, Baak A, et al. The FAIR Guiding
 363 Principles for scientific data management and stewardship. Sci Data. 2016;3:160018.

4. Steinbach D, Alaux M, Amselem J, Choisne N, Durand S, Flores R, et al. GnpIS: an information system to
integrate genetic and genomic data from plants and fungi. Database J Biol Databases Curation.
2013;2013:bat058.

- 367 5. Wheat@URGI portal. https://wheat-urgi.versailles.inra.fr. Accessed 10 April 2018.
- 6. Nelson WM, Bharti AK, Butler E, Wei F, Fuks G, Kim H, et al. Whole-Genome Validation of HighInformation-Content Fingerprinting. Plant Physiol. 2005;139:27–38.
- 370 7. Philippe R, Choulet F, Paux E, van Oeveren J, Tang J, Wittenberg AH, et al. Whole Genome Profiling

provides a robust framework for physical mapping and sequencing in the highly complex and repetitive wheat
 genome. BMC Genomics. 2012;13:47.

- 8. Soderlund C, Humphray S, Dunham A, French L. Contigs built with fingerprints, markers, and FPC V4.7.
- 374 Genome Res. 2000;10:1772–87.
  - 18

- 9. Frenkel Z, Paux E, Mester D, Feuillet C, Korol A. LTC: a novel algorithm to improve the efficiency of contig
  assembly for physical mapping in complex genomes. BMC Bioinformatics. 2010;11:584.
- 10. French plant genomic resource centre. https://cnrgv.toulouse.inra.fr/en. Accessed 10 April 2018.
- 11. Altschul SF, Gish W, Miller W, Myers EW, Lipman DJ. Basic local alignment search tool. J Mol Biol.
  1990;215:403–10.
- 12. International Wheat Genome Sequencing Consortium (IWGSC). A chromosome-based draft sequence of the
   hexaploid bread wheat (Triticum aestivum) genome. Science. 2014;345:1251788.
- 13. Choulet F, Alberti A, Theil S, Glover N, Barbe V, Daron J, et al. Structural and functional partitioning of
   bread wheat chromosome 3B. Science. 2014;345:1249721.
- 14. Mascher M, Muehlbauer GJ, Rokhsar DS, Chapman J, Schmutz J, Barry K, et al. Anchoring and ordering
   NGS contig assemblies by population sequencing (POPSEQ). Plant J Cell Mol Biol. 2013;76:718–27.
- 386 15. Ramirez-Gonzalez et al., 2018, submitted.
- 16. Jordan KW, Wang S, Lun Y, Gardiner L-J, MacLachlan R, Hucl P, et al. A haplotype map of allohexaploid
   wheat reveals distinct patterns of selection on homoeologous genomes. Genome Biol. 2015;16:48.
- 17. Rimbert H, Darrier B, Navarro J, Kitt J, Choulet F, Leveugle M, et al. High throughput SNP discovery and
   genotyping in hexaploid wheat. PloS One. 2018;13:e0186329.

18. Clavijo BJ, Venturini L, Schudoma C, Accinelli GG, Kaithakottil G, Wright J, et al. An improved assembly
 and annotation of the allohexaploid wheat genome identifies complete families of agronomic genes and provides
 genomic evidence for chromosomal translocations. Genome Res. 2017;

19. Luo M-C, Gu YQ, You FM, Deal KR, Ma Y, Hu Y, et al. A 4-gigabase physical map unlocks the structure
and evolution of the complex genome of Aegilops tauschii, the wheat D-genome progenitor. Proc Natl Acad Sci
U S A. 2013;110:7940–5.

- 397 20. GnpIS wheat data. https://wheat-urgi.versailles.inra.fr/Data. Accessed 10 April 2018.
- Samson D, Legeai F, Karsenty E, Reboux S, Veyrieras J-B, Just J, et al. GénoPlante-Info (GPI): a collection
   of databases and bioinformatics resources for plant genomics. Nucleic Acids Res. 2003;31:179–82.
- 400 22. Ćwiek-Kupczyńska H, Altmann T, Arend D, Arnaud E, Chen D, Cornut G, et al. Measures for
- interoperability of phenotypic data: minimum information requirements and formatting. Plant Methods.2016;12:44.
- 403 23. French small grain cereals genebank.
- 404 https://www6.clermont.inra.fr/umr1095\_eng/Teams/Research/Biological-Resources-Centre. Accessed 10 April
   405 2018.
- 406 24. Kalderimis A, Lyne R, Butano D, Contrino S, Lyne M, Heimbach J, et al. InterMine: extensive web services
   407 for modern biology. Nucleic Acids Res. 2014;42:W468-472.
- 408 25. IWGSC data repository. https://wheat-urgi.versailles.inra.fr/Seq-Repository. Accessed 10 April 2018.
- 409 26. Stein LD, Mungall C, Shu S, Caudy M, Mangone M, Day A, et al. The generic genome browser: a building
  410 block for a model organism system database. Genome Res. 2002;12:1599–610.
- 411 27. GnpIS: Physical map browser. https://urgi.versailles.inra.fr/gb2/gbrowse/wheat\_phys\_pub. Accessed 10
   412 April 2018.
- 28. Skinner ME, Uzilov AV, Stein LD, Mungall CJ, Holmes IH. JBrowse: a next-generation genome browser.
  Genome Res. 2009;19:1630–8.

- 415 29. GnpIS: IWGSC RefSeq v1.0 browser.
- https://urgi.versailles.inra.fr/jbrowseiwgsc/gmod\_jbrowse/?data=myData%2FIWGSC\_RefSeq\_v1.0. Accessed
   10 April 2018.
- 418 30. GnpIS: IWGSC BLAST tool. https://urgi.versailles.inra.fr/blast\_iwgsc/blast.php. Accessed 10 April 2018.
- 31. Bolser DM, Staines DM, Perry E, Kersey PJ. Ensembl Plants: Integrating Tools for Visualizing, Mining, and
   Analyzing Plant Genomic Data. Methods Mol Biol Clifton NJ. 2017;1533:1–31.
- 421 32. GnpIS PlantSyntenyViewer. https://urgi.versailles.inra.fr/synteny/synteny/viewer.do#form/datasetId=6.
   422 Accessed 10 April 2018.
- 423 33. Pont C, Murat F, Guizard S, Flores R, Foucrier S, Bidet Y, et al. Wheat syntenome unveils new evidences of
- 424 contrasted evolutionary plasticity between paleo- and neoduplicated subgenomes. Plant J Cell Mol Biol.
- 425 2013;76:1030–44.
- 426 34. Multi-Crop Passport Descriptors V.2.1. https://www.bioversityinternational.org/e-
- library/publications/detail/faobioversity-multi-crop-passport-descriptors-v21-mcpd-v21/. Accessed 10 April
   2018.
- 429 35. Shrestha R, Matteis L, Skofic M, Portugal A, McLaren G, Hyman G, et al. Bridging the phenotypic and
- 430 genetic data useful for integrated breeding through a data annotation using the Crop Ontology developed by the 431 crop communities of practice. Front Physiol. 2012;3:326.
- 432 36. GnpIS: WheatMine tool. https://urgi.versailles.inra.fr/WheatMine. Accessed 10 April 2018.
- 433 37. WheaIS. http://www.wheatis.org/. Accessed 10 April 2018.
- 434 38. Spannagl M, Nussbaumer T, Bader KC, Martis MM, Seidel M, Kugler KG, et al. PGSB PlantsDB: updates
   435 to the database framework for comparative plant genome research. Nucleic Acids Res. 2016;44:D1141–7.
- 436 39. Künne C, Lange M, Funke T, Miehe H, Thiel T, Grosse I, et al. CR-EST: a resource for crop ESTs. Nucleic
  437 Acids Res. 2005;33:D619–21.
- 438 40. Oppermann M, Weise S, Dittmann C, Knüpffer H. GBIS: the information system of the German Genebank.
- 439 Database J Biol Databases Curation [Internet]. 2015 [cited 2017 Sep 18];2015. Available from:
- 440 http://www.ncbi.nlm.nih.gov/pmc/articles/PMC4423411/
- 41. Schreiber F, Colmsee C, Czauderna T, Grafahrend-Belau E, Hartmann A, Junker A, et al. MetaCrop 2.0:
  42 managing and exploring information about crop plant metabolism. Nucleic Acids Res. 2012;40:D1173–7.
- 443 42. Tello-Ruiz MK, Stein J, Wei S, Preece J, Olson A, Naithani S, et al. Gramene 2016: comparative plant 444 genomics and pathway resources. Nucleic Acids Res. 2016;44:D1133-1140.
- 445 43. Montenegro JD, Golicz AA, Bayer PE, Hurgobin B, Lee H, Chan C-KK, et al. The pangenome of hexaploid
  bread wheat. Plant J Cell Mol Biol. 2017;90:1007–13.
- 447 44. Carollo V, Matthews DE, Lazo GR, Blake TK, Hummel DD, Lui N, et al. GrainGenes 2.0. An Improved
  448 Resource for the Small-Grains Community. Plant Physiol. 2005;139:643–51.
- 449 45. Wheat Data Interoperability Working Group of the Research Data Alliance. https://rd-
- 450 alliance.org/groups/wheat-data-interoperability-wg.html. Accessed 10 April 2018.
- 46. Dzale Yeumo E, Alaux M, Arnaud E, Aubin S, Baumann U, Buche P, et al. Developing data interoperability
  using standards: A wheat community use case. F1000Research. 2017;6:1843.
- 453 47. Wheat Data Interoperability Working Group guidelines. https://ist.blogs.inra.fr/wdi/. Accessed 10 April
  454 2018.
- 455 48. Deng W, Nickle DC, Learn GH, Maust B, Mullins JI. ViroBLAST: a stand-alone BLAST web server for 456 flexible queries of multiple databases and user's datasets. Bioinforma Oxf Engl. 2007;23:2334–6.

# 457 Supplementary data

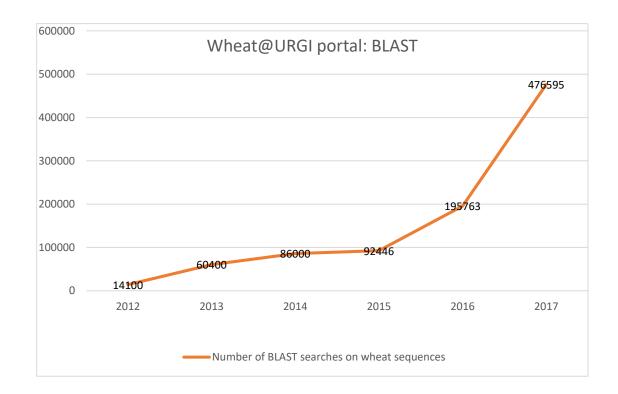
### 458 Software technologies

- The Wheat@URGI portal website is based on eZ Publish v4 open source content management system (<u>https://ez.no/</u>) using the PHP language and a MySQL database (<u>https://www.mysql.com/</u>).
- 462 The genome browsers are based on the GMOD (<u>http://gmod.org/wiki/Main\_Page</u>) GBrowse
- 463 v2.33 [26] and JBrowse v1.11.5 [28] built with JavaScript and HTML5. We customized GBrowse
- to display the physical map data. The gff3 file is generated from the .fpc file obtained by the
- data producer using the FPC [8] or the LTC [9] tools.
- 466 The stand-alone BLAST web interface implemented at URGI is based on ViroBLAST [48],
- 467 customized to obtain a user-friendly grouping of searched databanks and visualization of the
- results. A robust file download system was also developed using a home-made php script to
   handle big data volume.
- GnpIS-coreDB is a URGI development using state of the art technologies: Java EE framework 470 (http://www.oracle.com/technetwork/java/javaee/overview/index.html), GWT (Google Web 471 472 Toolkit, http://www.gwtproject.org/), Spring boot v1.4 (https://projects.spring.io/springboot/), PostgreSQL relational database v9.6 (https://www.postgresql.org/) and Elasticsearch 473 NoSQL database v2.3.3 (https://www.elastic.co/). To set-up a GnpIS-coreDB dedicated to the 474 wheat community, a filter allowing to display only the wheat data (Triticum, Aegilops) and 475 barley data (Hordeum) was developed. This filter relies on a variable-length multidimensional 476 arrays field in the PostgreSQL database. It is completely transparent to the user and allows 477 478 him to navigate in GnpIS-coreDB through wheat data only. New versions of the GnpIS-coreDB 479 software are deposited in the APP, the European body for protecting authors and publishers of digital works (http://www.app.asso.fr/en/welcome.html). 480

481	WheatMine uses InterMine [24] v1.8.3 which provides a fast, flexible and user friendly access
482	to integrated data by multiple ways: a browser, a query builder and a region search tool. Users
483	can filter their favorite features, save their own queries, and export results in many different
484	formats (GFF3, BED or XML). An On-line documentation and pre-computed queries are also
485	available.
486	The data discovery tool relies on the Solr full-text indexing technology v6.6.2
487	(http://lucene.apache.org/solr/). We used a restriction on the wheat and barley species to

- 488 search only the corresponding data in the indexed databases. The tool was packaged and is
- 489 downloadable (<u>https://wheat-urgi.versailles.inra.fr/Projects/Wheat-Information-</u>
- 490 <u>System/SolR-tool-package</u>).
- 491
- 492 Usage Statistics
- 493 Table S1. Usage statistics of genomics data in the Wheat@URGI portal (all numbers exclude
- 494 web-robots and internal IP).

Number of	2012	2013	2014	2015	2016	2017
Visits on the IWGSC Sequence						
Repository website	N/A	11440	20754	27070	20841	28151
Downloads of wheat sequence						
data	2253	4413	17783	19307	18724	22935
Visits on the wheat browsers	5869	9370	9130	22989	22373	18262
Number of BLAST searches on						
wheat sequences	14100	60400	86000	92446	195763	476595
Number of WheatIS data					13010	
discovery tool searches	N/A	N/A	N/A	N/A	13010	26480



495

#### 496 Use case example

497 A researcher in genomics works on his wheat favorite gene. He wants to explore all the 498 genomic data in the vicinity of this gene and find out if there are genetic studies pointing to 499 the genomic regions where the gene is located. He searches the gene name (e.g. 500 TraesCS5A01G033100) in the data discovery tool (<u>https://wheat-urgi.versailles.inra.fr</u>, Fig 501 S1.1A) or BLAST the sequence of the gene against the IWGSC RefSeq v1.0 502 (<u>https://urgi.versailles.inra.fr/blast\_iwgsc/</u>, Fig S1.1B).

503 The results are displayed in a table (Fig S1.2A, Fig S1.2B) with links to the JBrowse directly

- 504 zooming on the gene
- 505 (https://urgi.versailles.inra.fr/jbrowseiwgsc/gmod\_jbrowse/?data=myData%2FIWGSC\_RefSe
- 506 <u>q v1.0&loc=chr5A%3A30211546..30218715&tracks=DNA%2CHighConfidenceGenesv1.0%2C</u>
- 507 LowConfidenceGenesv1.0%2CrepeatRegion%2CrepeatMasker%2CDART\_PUBLIC\_SUMMARY
- 508 <u>&highlight=chr5A%3A30214481..30215784%20(-%20strand)%20(TraesCS5A01G033100)</u>). He

509	explores the region a	round the gene and	l finds a marker	r (e.g. wPt-4131	_PURr, Fig S1.3). By
510	clicking on the ma	rker, he obtains	additional info	ormation stored	d in GnpIS-coreDB
511	(https://urgi.versailles	s.inra.fr/GnpMap/m	apping/id.do?ad	ction=MARKER&	. <u>id=40393</u> , Fig S1.4)
512	showing that	the marker	is used	in a GW	AS experiments
513	( <u>https://urgi.versailles</u>	s.inra.fr/association/	association/vie/	wer.do?results/	markerIds=40393,
514	Fig S1.5A) an	d is linked	to a	QTL (e.g.	GY_ml.8.Np_5A,
515	https://urgi.versailles.	.inra.fr/GnpMap/ma	apping/id.do?ac	tion=QTL&id=59	9 <u>588</u> , Fig S1.5B).
516	From the	Trait	description	of	this QTL
517	( <u>https://urgi.versailles</u>	a.inra.fr/GnpMap/m	apping/id.do?ac	ction=TRAIT&id=	<u>255</u> , Fig S1.6), he
518	displays all the	phenotyping	experiment	performed	on this trait
519	( <u>https://urgi.versailles</u>	s.inra.fr/ephesis/eph	nesis/viewer.doa	#dataResults/tra	aitCode=SCNO:0000
520	0002 Eig (1.7)				

- 520 <u>0002</u>, Fig S1.7).
- 521 Figure S1. Printscreens of the web interfaces.

