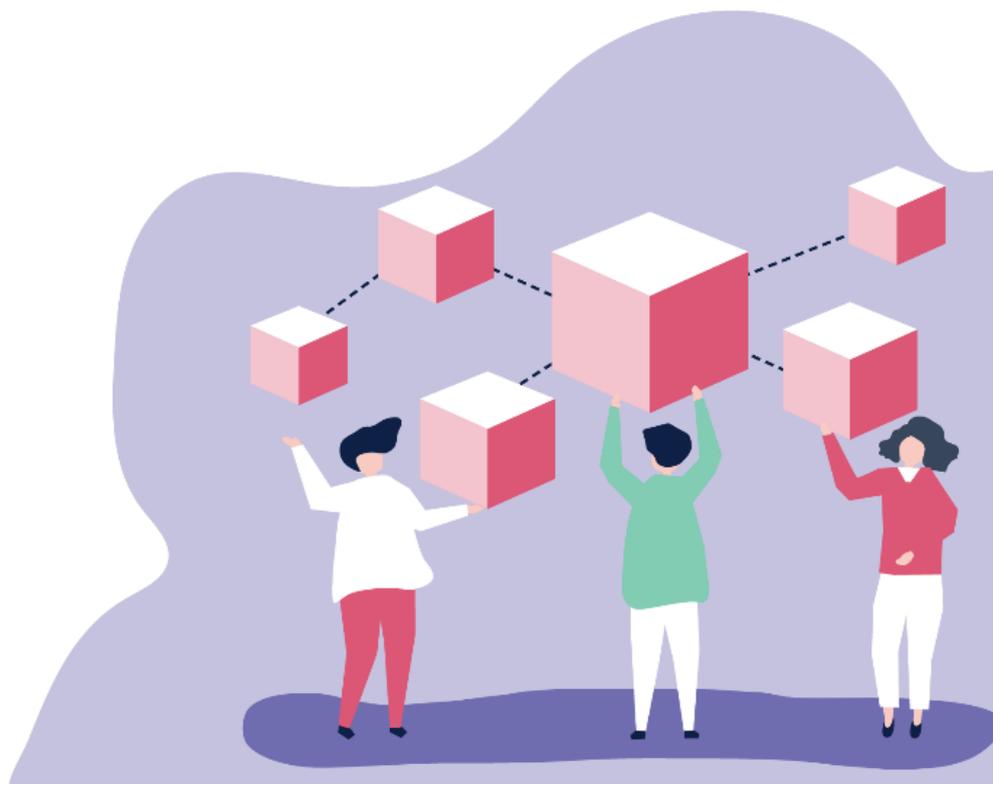




GATE KEEPER

User manual. GastricAITool

Deliverable No.		Due Date	
Description	User manual		
Type	Report	Dissemination Level	CO
Work Package No.	WP3	Work Package Title	
Version	1.0	Status	Final



Authors

Name and surname	Partner name	e-mail
Rocío Aznar	ITAINNOVA	raznar@itainnova.es
Rubén Muñoz	ITAINNOVA	rmunoz@itainnova.es
Asunción García	IIS Aragón	agarcia.iacs@aragon.es
Patricia Carrera	IIS Aragón	pcarreralasfuentes@gmail.com
Vega Rodríguez	ITAINNOVA	vrodriガルvarez@itainnova.es
Rafael del Hoyo	ITAINNOVA	rdelhoyo@itainnova.es

History

Date	Version	Change
13/04/2023	1.0	Final version

Key data

Keywords	User, patient, tool, diagnosis, prognosis
Lead Editor	
Internal Reviewer(s)	

Abstract

The objective of this document is to present the user manual of the GastricAITool (diagnostic and prognostic tool for gastric cancer).

Statement of originality

This deliverable contains original unpublished work except where clearly indicated otherwise. Acknowledgement of previously published material and of the work of others has been made through appropriate citation, quotation or both.

Table of contents

TABLE OF CONTENTS	3
LIST OF FIGURES	4
1 INTRODUCTION	5
1.1 PROJECT OBJECTIVE	5
1.2 PURPOSE OF THE DOCUMENT	5
2 USER MANUAL	6
2.1 SYSTEM LOGIN	6
2.2 USER MANAGEMENT	6
2.3 LIST OF PATIENTS	7
2.4 NEW PATIENT BY DIAGNOSIS/PROGNOSIS.....	8
2.5 RESULTS.....	10

List of figures

FIGURE 1. MAIN SCREEN. GASTRICAITool.....	6
FIGURE 2. USERS' MANAGEMENT. GASTRICAITool.....	6
FIGURE 3. CREATE NEW USER. GASTRICAITool.....	7
FIGURE 4. LIST OF PATIENTS.	8
FIGURE 5. NEW PATIENT BY DIAGNOSIS. GASTRICAITool.....	9
FIGURE 6. NEW PATIENT BY PROGNOSIS. GASTRICAITool.	9

1 Introduction

1.1 Project objective

The main objective of the proposal is to develop and deploy an innovative diagnostic and prognostic tool for gastric cancer (GC) built on big data technologies and personalised in GATEKEEPER infrastructure that helps the clinician to take critical decisions and detect risks about patients in a specific moment, based on the information available at this specific time and on an advanced health monitoring based on historical data. This tool will allow the development of more personalised follow-up strategies, early interventions, and therapeutic actions, improving the healthcare system.

1.2 Purpose of the document

The purpose of this document is to present the user manual for the GastricAITool (the diagnostic and prognostic tool for gastric cancer). This tool provides diagnosis and/or prognosis of gastric cancer based on patient information. It is an intuitive and easy-to-use tool that has been designed in collaboration with project partners, always with clinical needs in mind. Collaborative work between partners (the Aragon Health Research Institute (IISA): the translational research in digestive pathology group, and ITAINNOVA from the Big Data and Cognitive systems side) has been crucial.

This document includes all the necessary information to use the tool, specifically related to:

- System login.
- User management.
- Creation and editing of patient data.
- Visualization of diagnostic and prognostic model results.

2 User manual

2.1 System login

Figure 1 displays the main screen of the tool. Here, the user must log in by entering the user name and password. The system will allow access to the tool only if the user name is on the list of users. Otherwise, the administrator must add it to the list.



Figure 1. Main screen. GastricAITool.

2.2 User management

Figure 2 shows the User Management System. Only the administrator has permissions to manage users (option located in the upper right corner: "Users management"). Therefore, the administrator is responsible for adding new users to the list, editing already created users, or removing them from the list.

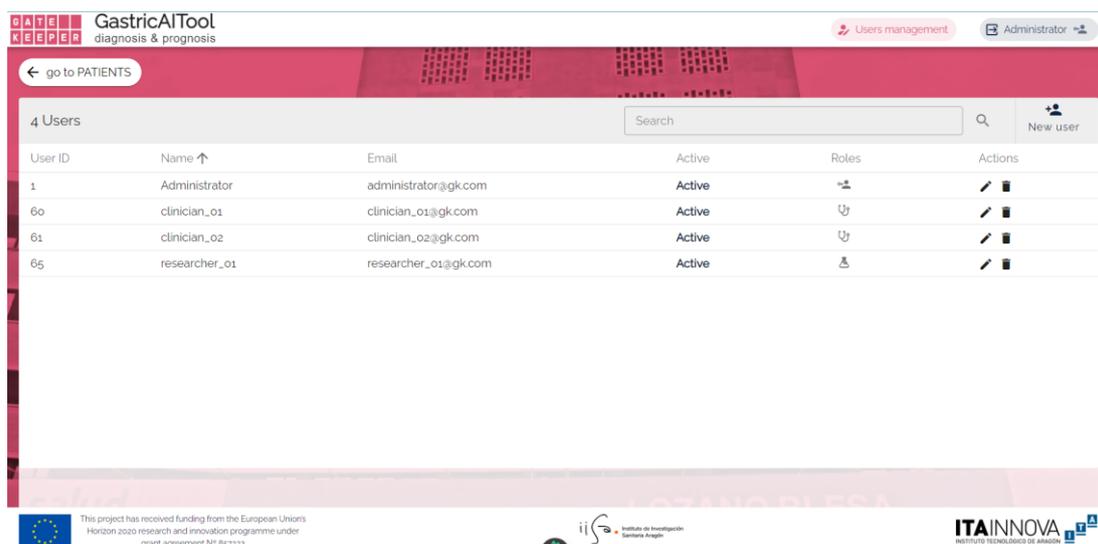


Figure 2. Users' management. GastricAITool.

To create a new user (option located in the upper right corner), the following screen appears as shown in Figure 3. Each created user is automatically assigned an identifier.

New User

username

password

email

active

Roles

Administrator

Clinician

Researcher

CANCEL

Figure 3. Create new user. GastricAITool.

The fields that identify each user are:

- Username and password: Required to access the tool.
- Email
- Status (active or inactive): Only active users and results of their patients will be displayed in the tool.
- Roles: They refer to the user's permissions to create new patients, delete patients, view patients' diagnostic and prognostic results, and manage users. The options are as follows:
 - Administrator: They have all permissions. Responsible for managing users. Can create and delete patients and view results of all patients.
 - Clinician: They can add new patients, view and delete the patients they have created. They cannot view patients of other clinicians.
 - Researcher: They have only read permissions. Can view the results of all patients.

2.3 List of patients

Once the user logs in, the first screen that appears is the list of patients as shown in Figure 4.

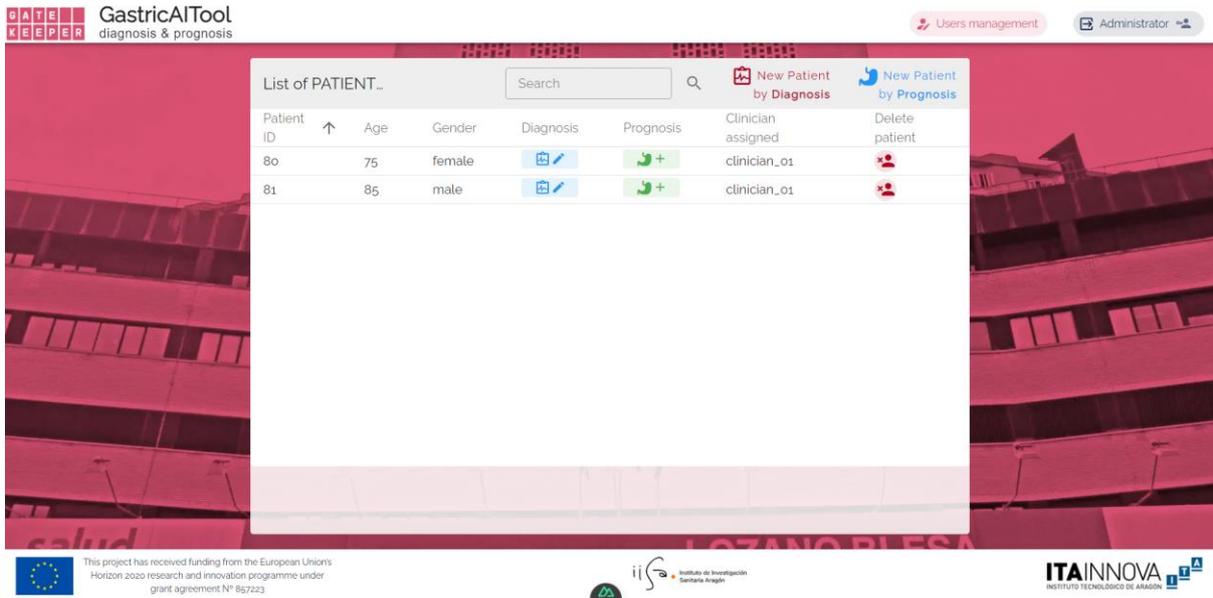


Figure 4. List of patients.

It displays the patients already included with the following information:

- Patient's tool identifier.
- Patient's age.
- Patient's gender.
- Their assigned clinician.
- Option to edit the patient's data related to diagnosis or prognosis or view the results (blue button).
- Option to add patient's data created related to diagnosis or prognosis (green button).
- Delete the patient.
- Include new patients to predict their diagnosis or prognosis ("New Patients by Diagnosis/Prognosis").

Also, a free text search for the patient by identification, age, or sex is offered.

2.4 New Patient by Diagnosis/Prognosis

Figure 5 and Figure 6 show the screens that appear when adding a new patient to predict their diagnosis ("New Patient by Diagnosis") or prognosis ("New Patient by Prognosis"), respectively.

GastricAITool
diagnosis & prognosis

Users management Administrator

New Patient

Age Gender Assigned to
Age is required Gender is required Clinician is required

Socio-Clinical Variables

Family History of Gastric Cancer H. pylori infection Smoking
 Yes No Yes No Current Never / Former

Genetic Variables

Carrier status
 Yes No

rs1060825 rs10759932 rs17655 rs20417
 rs2074522 rs2228000 rs2345060 rs4072037
 rs4150415 rs4986764 rs599443 rs5768
 rs6679677 rs909253 rs9894945

fill Clinician, Age, Gender fields — SAVE 24 fields empty — SAVE & EXECUTE

This project has received funding from the European Union's Horizon 2020 research and innovation programme under grant agreement N° 867223

ITAINNOVA

Figure 5. New Patient by Diagnosis. GastricAITool.

GastricAITool
diagnosis & prognosis

Users management Administrator

New Patient

Age Gender Assigned to
Age is required Gender is required Clinician is required

Clinical Variables

Radiotherapy Chemotherapy Surgery
 Yes No Yes No Yes No

Tumor location TNM Stage Metastasis
 Cecal Distal in situ I II III IV Yes No

Genetic Variables

Carrier status
 Yes No

rs1052133 rs1109956 rs1271521 rs13181
 rs144848 rs1799795 rs1800470 rs1896830
 rs2074522 rs207906 rs20779 rs2738120
 rs273859 rs293794 rs388874 rs4072037
 rs423459 rs4986764 rs4987876 rs595682
 rs7744 rs779265 rs7932765 rs8305
 rs941504

fill Clinician, Age, Gender fields — SAVE 24 fields empty — SAVE & EXECUTE

This project has received funding from the European Union's Horizon 2020 research and innovation programme under grant agreement N° 867223

ITAINNOVA

Figure 6. New Patient by Prognosis. GastricAITool.

The following information appears:

- **Patient:** First, the patient's information (age and gender) and their assigned clinician appear. In this example (Figure 5 and Figure 6), since the role is that of an administrator, it allows assigning from the list of clinicians. If the user were a clinician, they would be automatically assigned to themselves.
- **Clinical variables:** Second, the clinical variables information appears. Help is provided to detail each information.
- **Genetic variables:** Finally, the genetic variables to be completed are shown, providing possible options.

- **Autofill:** In this section, the option to autofill genetic variables is offered if this information is not available, provided that at least 80% of the genetic variables have been completed.

Options:

- **Save:** This button must be clicked first. It allows saving the completed information of the new patient in the database. To do so, it is required to have completed at least the age, gender, and assigned clinician.

After saving the information initially, an identifier is automatically created in the tool for the new patient, without the possibility of identifying them with the real patient. It will be the responsibility of the assigned clinician to establish this identification in a way that complies with GDPR.

- **Save & Execute:** It allows applying the diagnosis or prognosis model given the patient's information. All information must be completed. As a result, the predicted probabilities by the model are provided, as well as global and local explainability graphs that allow the clinician to interpret the results and understand how the model arrived at that outcome.

Likewise, the tool provides help functions that offer detailed explanations in the points where they are needed.

The same screen shown in Figure 5 and Figure 6 also appears with the option to edit or add patient information from the patient list (blue and green buttons, Figure 4). If the edit option is selected (blue button), the information will appear already completed, along with the results provided by the model, below.

2.5 Results

Once all the patient information has been completed, the "Save & Execute" button applies the diagnostic or prognostic model and provides the following information:

- **Diagnosis**
 - Probability of gastric cancer risk.
 - Global Explanation. Features contribution: A graph that provides an explanation of the general behaviour of the model.
 - Local Explanation. Features contribution: A graph that provides an explanation of how the model made the prediction for the particular individual.
 - Local vs. Global Explanation. Features contribution: A graph that incorporates absolute local and global contribution.
 - Genetic Risk Score (GRS): The GRS, normalized between 0 and 10, resulting from the patient's genetic information.
- **Prognosis**
 - Survival probability at 1,5, 3, 5 and 10 years.
 - Predicted survival curve.
 - Global Explanation. Features contribution.
 - Local Explanation. Features contribution.

- Genetic Risk Score (GRS): The GRS, normalized between 0 and 10, resulting from the patient's genetic information.

All graphs are accompanied by explanatory text.

These results can be saved as a PDF report. The following pages show examples of diagnosis and prognosis, respectively.

Diagnosis Prediction

Patient

Patient ID 	Age	Gender
80	75	female ▼

Assigned to

Clinician
clinician_01 ▼

Socio-Clinical Variables

Family History of Gastric Car	H. pylori Infection	Smoking
<input type="radio"/> Yes <input checked="" type="radio"/> No	<input checked="" type="radio"/> Yes <input type="radio"/> No	<input checked="" type="radio"/> Current
		<input type="radio"/> Never / Former

Genetic Variables

Carrier Ilra2

Yes No

rs1060826 AG ▼	rs10759932 CT ▼	rs17655 CG ▼	rs20417 GG ▼
rs2074522 AA ▼	rs2228000 CT ▼	rs2345060 GG ▼	rs4072037 TT ▼
rs4150416 TT ▼	rs4986764 CT ▼	rs569143 CG ▼	rs5788 CC ▼
rs6679677 CC ▼	rs909253 AA ▼	rs9894946 GG ▼	



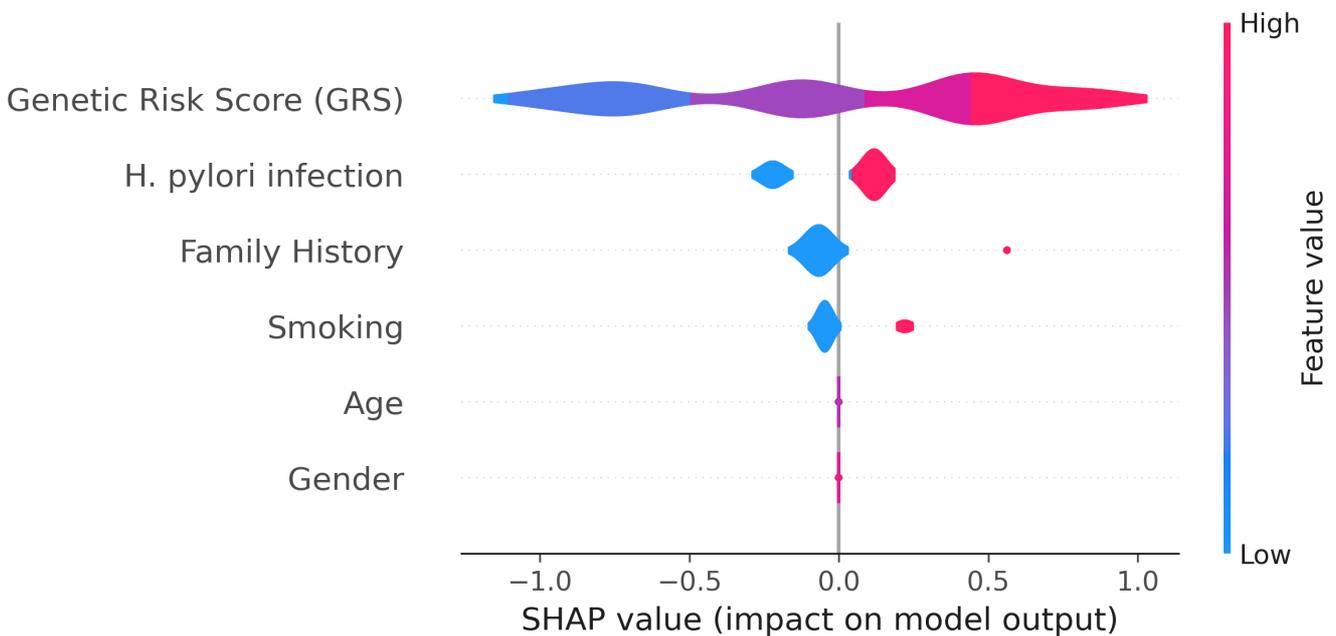
RESULTS



Probability of gastric cancer risk

Global Explanation. Features contribution

i This graph represents the global explainability of the model through the contribution that each variable has had in the constructed model. The position of the x-axis is determined by the Shapley value, and the position of the y-axis is determined by the variable, ordered from most to least important. Positive shap values indicate risk of gastric cancer, and negative values indicate the opposite. The color indicates the value of the variable, represented by blue for the lowest values and red for the highest values.



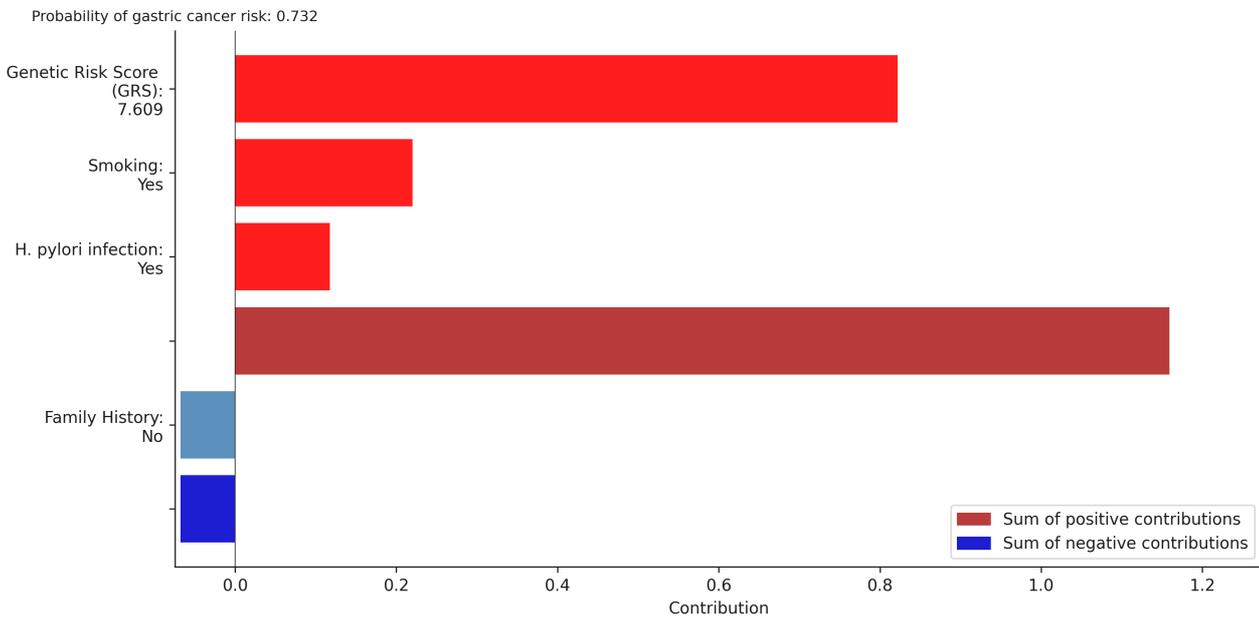
The variables of H. pylori infection, family history or smoking are binary variables where 0 indicates absence and 1, the opposite. The graph indicates that genetics, H. pylori infection and family history are the factors that have most globally influenced the model's decision. High values of the genetic risk score and having a family history of gastric cancer have an increased risk of gastric cancer. Conversely, low values of the genetic risk score or not being infected by H. pylori bacteria are protective factors.

Local Explanation. Features contribution

i This graph represents the local explainability, i.e. the contribution of each variable (>0) that influenced the model's decision for the particular patient. Positive contributions indicate gastric

cancer risk and negative contributions the opposite.

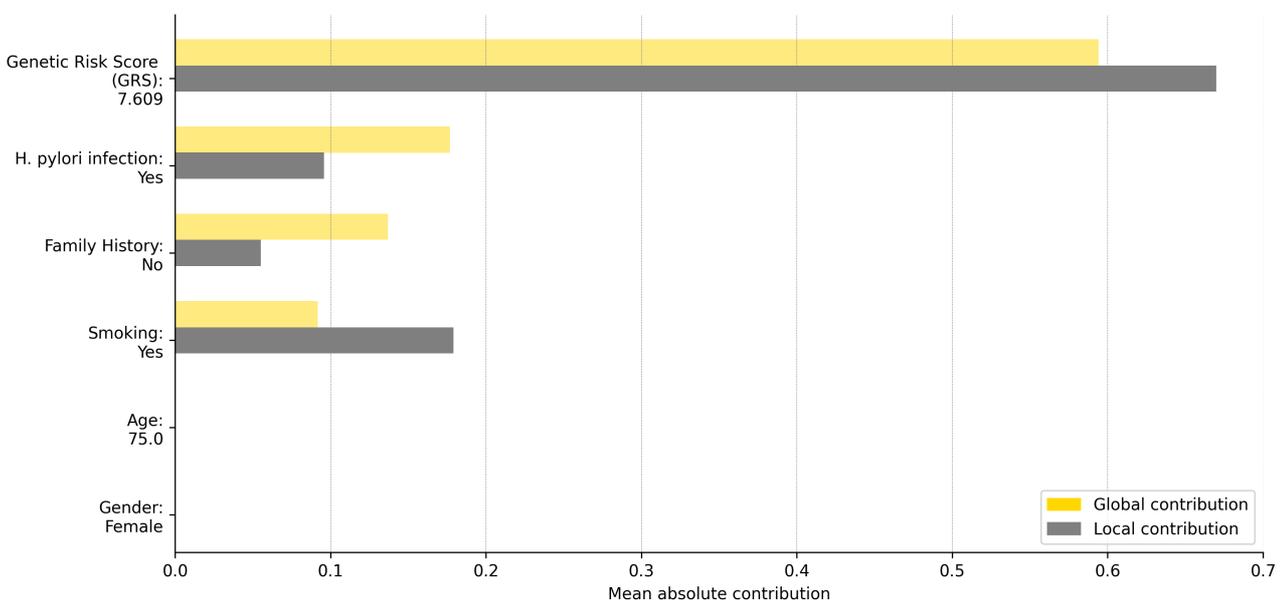
Local Explanation. Features contribution



Local vs. Global Explanation. Features contribution

i This graph represents both the local and the global explainability of the model. The mean absolute contribution of each variable to the model prediction is presented. The sum of the contributions of all variables is 1. This graph allows to compare the importance of the variables globally and in the prediction of the individual.

Global vs. Local Contribution



Genetic Risk Score



Value from 0 to 10

Prognosis Estimation

Patient

Patient ID 	Age	Gender
82	80	male 

Assigned to

Clinician
clinician_01 

Clinical Variables

Radiotherapy 

Yes No

Chemotherapy 

Yes No

Surgery 

Yes No

Tumor location

Cardia Distal

TNM Stage

In situ I II III
 IV

Metastasis

Yes No

Genetic Variables

Carrier Ilra2

Yes No

rs1052133 CG 	rs11086565 AG 	rs12711521 AA 	rs13181 GT 
rs144848 GT 	rs1799796 GG 	rs1800470 TT 	rs1898830 AG 
rs2074522 GG 	rs207906 AG 	rs26779 AG 	rs2738120 GG 
rs2738169 AA 	rs293794 CT 	rs3088074 GG 	rs4072037 TT 



rs4234259
AG ▼

rs4986790
AG ▼

rs4987876
GT ▼

rs6151662
GG ▼

rs7744
AA ▼

rs7797466
GG ▼

rs7932766
CC ▼

rs8305
AA ▼

rs9841504
CC ▼

RESULTS



Survival probability at 1.5 years



Survival probability at 3 years



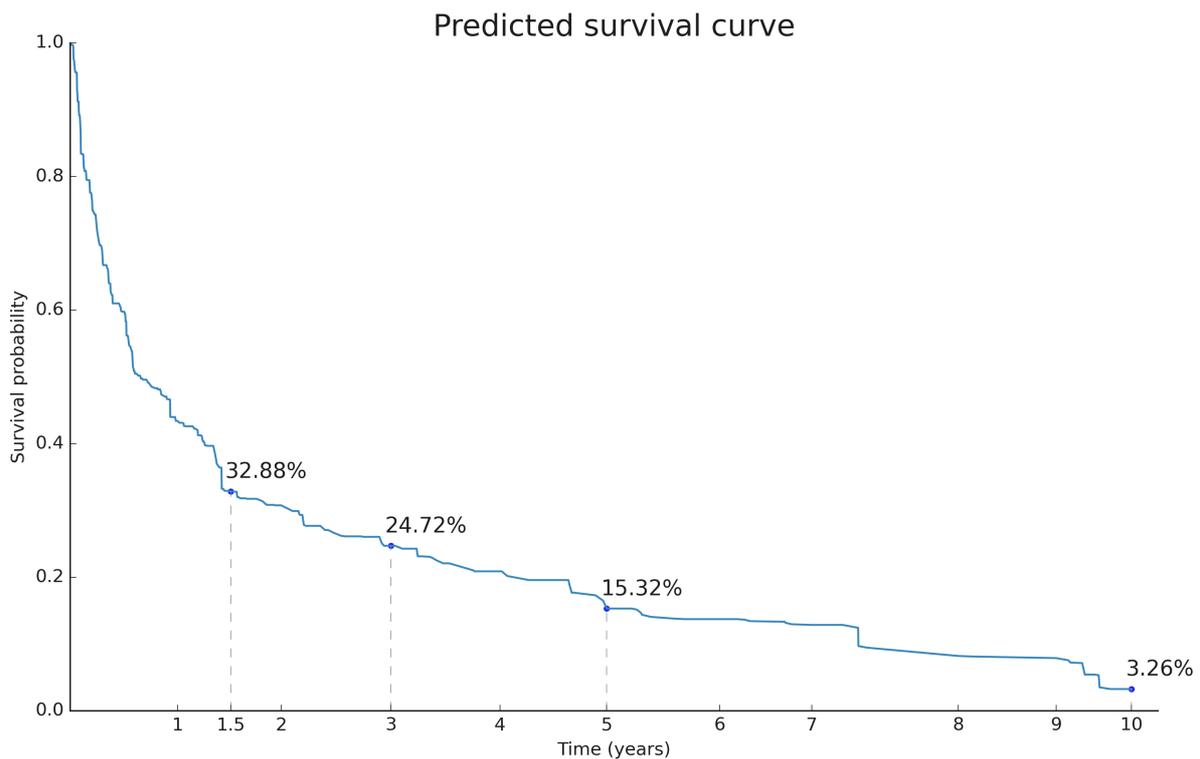
Survival probability at 5 years



Survival probability at 10 years

Survival curve

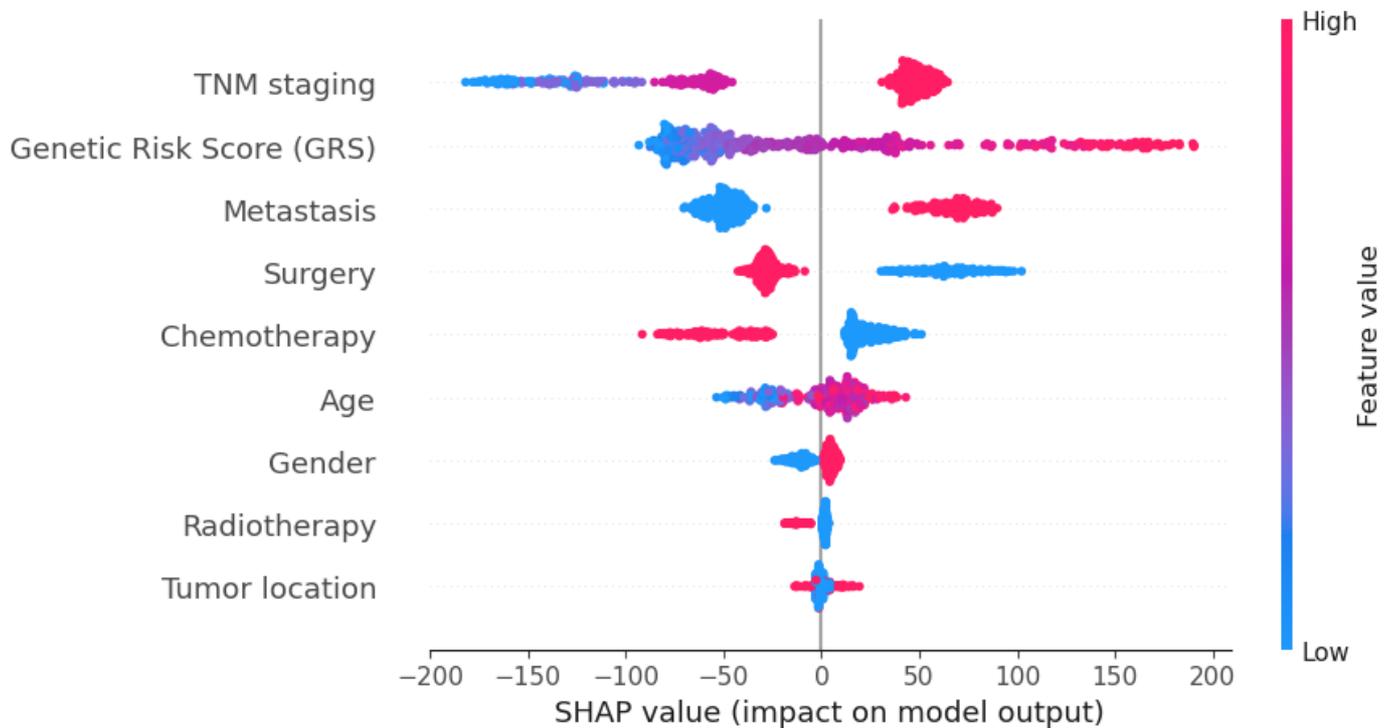
- i** This graph represents the survival curve predicted by the model with a time period up to 10 years. The survival probabilities at 1.5, 3, 5 and 10 years are shown.



Global Explanation. Features contribution

- i** This graph represents the global explainability of the model through the contribution that each variable has had in the constructed model. The position of the x-axis is determined by the Shapley value, and the position of the y-axis is determined by the variable, ordered from most to least important. Positive shap values indicate risk (poor prognosis), and negative values indicate the opposite. The color indicates the value of the variable, represented by blue for the

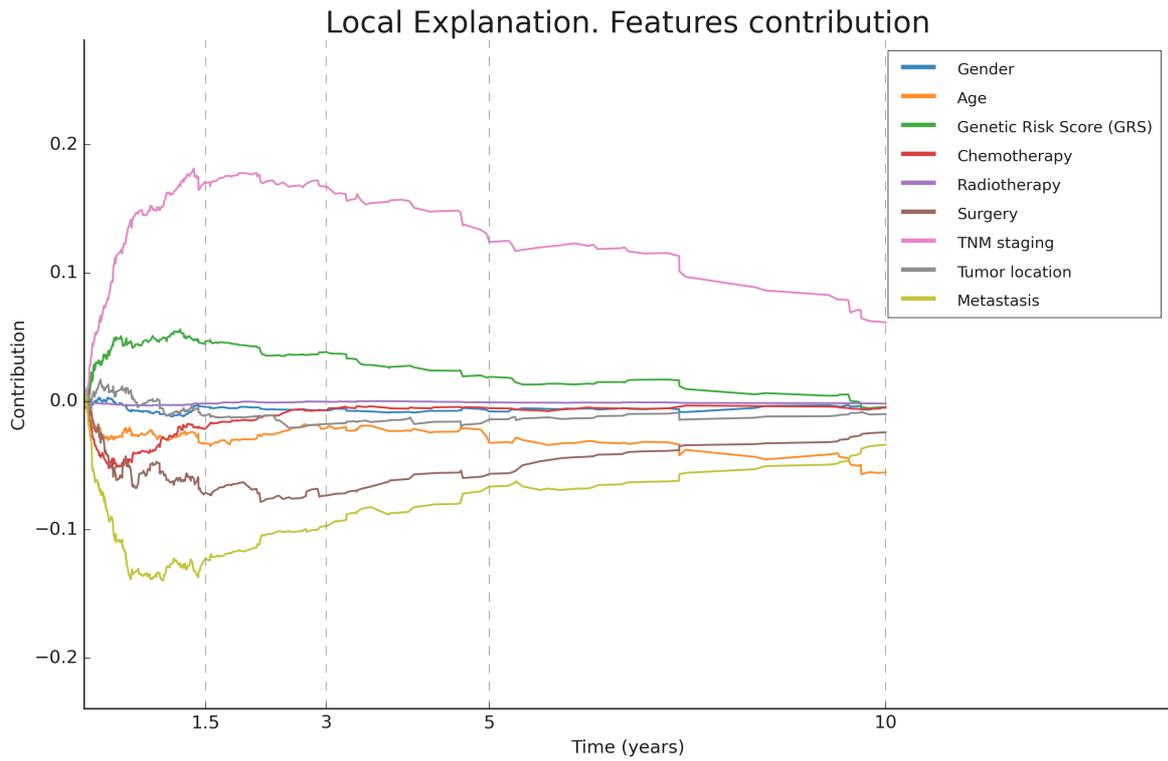
lowest values and red for the highest values. The variables of surgery, chemotherapy, and radiation therapy are binary variables where a value of 1 indicates that the patient has been treated, and 0 indicates the opposite. A value of 0 in the gender variable indicates female, and 1 indicates male.



The graph indicates that the TNM stage and genetics are the factors that have most globally influenced the model's decision, and, conversely, tumor location has had the least influence. Advanced TNM stages, high values of the genetic risk score, having metastasis at diagnosis, not having been treated with surgery or not having been treated with chemotherapy are factors with a negative impact on survival (poorer prognosis).

Local Explanation. Features contribution

- i** This graph represents the local explainability, i.e. the contribution of each variable that influenced the model's decision for the particular patient. Contributions are represented over time. Negative contributions indicate a higher risk (worse prognosis) and positive contributions the opposite.



Genetic Risk Score



Value from 0 to 10