User manual

For the

Briganti Nomograms

Briganti 2012 Nomogram
Briganti 2017 Nomogram
Briganti 2019 Nomogram

Version 2, June 2024, in English

1. The Evidencio platform

The Evidencio platform facilitates the creation, use, validation and implementation of medical prediction models and clinical decision support tools. This user manual specifically relates to the Briganti Nomogram. The User Manual can also be referred to as the Instructions For Use (IFU).

Throughout this manual CE-marked content and the term medical device are used interchangeably.

2. Disclaimer

Evidencio provides information, models, calculators, equations, and algorithms (tools) intended for use by healthcare professionals. Some of these tools have been certified as CE-medical devices. For such CE-marked content the 'Official Legal Disclaimer for CE-marked content' applies. All other content and tools provided by Evidencio are explicitly only covered by the 'Official Legal Disclaimer for non CE-marked content' both are available here: https://www.evidencio.com/disclaimer

3. Warnings

1. Warnings for CE-marked content

Calculations alone should never dictate patient care, and are no substitute for professional judgement. This tool is only to be used by physicians in a clinical setting, and is not for patient use.

Always read the intended use before using this tool.

Before reading the result, double check the filled in values to prevent errors.

Results that concern risk percentages, do not guarantee certain outcomes. When there is a risk present, do not expect an event to not occur at all, even if the risk is very small.

This model is only intended for use in settings where the usage and result of a model are never immediately needed.

4. Model landing page

The medical device model on the Evidencio platform is shown in **Figure 1**. The model landing page contains the following sections, that are indicated in **Figure 1**.

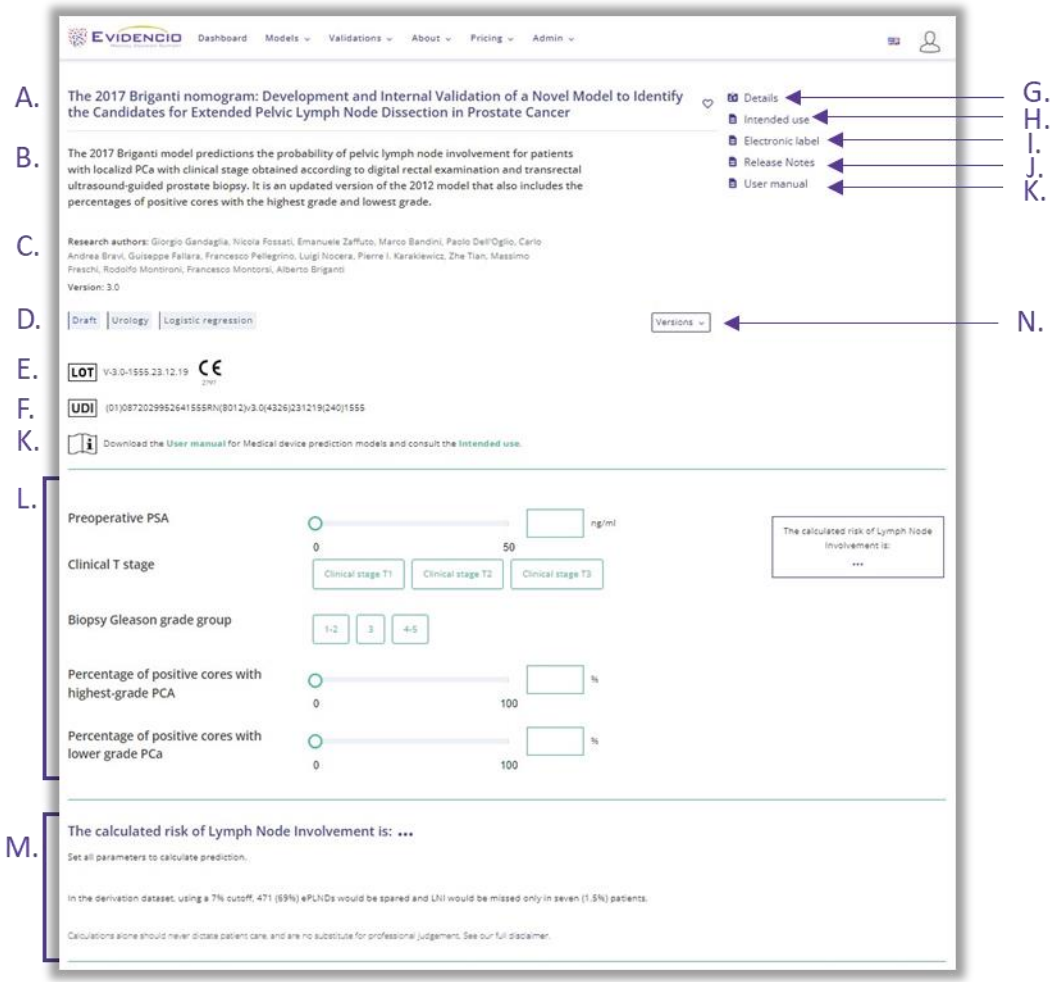

***Figure 1**. An example of a model landing page.*

A. Model title

This is the title and name of the model.

B. Model description

This is a short description of the model.

C. Research authors

These are the research authors of the paper that originally published the model.

D. Model tags

These are the tags that are assigned to the model. Evidencio has the following status tags: “Draft”, “Public”, “Private”, “Under review”. Evidencio has the following model type tags: “Composite model”, “Sequential model”, “API model”. Evidencio has the following calculation method tags: “Linear model”, “Logistic regression”, “Cox regression”, “RScript” and “Custom model”. Next to this, there are tags that indicate the specialty e.g. “Cardiology”.

E. LOT number

The LOT number indicated the model version, the model identifier, and the model publication date. Publication date is indicated as YY.MM.DD.

Additionally, the CE mark is displayed next to the LOT number. This way, medical devices can be easily recognized.

F. UDI number

The UDI number is an international tool that helps users identify and find information on products. UDI stands for Unique Device Identifier. Evidencio's UDIs have the following format:

(01)[UDI-DI number](8012)[versionnumber](4326)[releasedate](240)[identificationnumber]

The UDI-DI number is a unique numeric code. For each medical device of Evidencio, a unique UDI-DI is ascribed. This UDI-DI is used as an "access key" for information stored in a unique device identification database (UDID). Information on Evidencio's medical devices can be found by searching for the UDI-DI number in the following data base: https://gepir.gs1.org/index.php/search-by-gtin

G. Details button

On the top right of the model page, several clickable buttons are displayed that show a pop-up when clicked. The first button opens a pop-up concerning additional information about the model. This pop-up has three sections: Details, Study characteristics and Supporting publications & related files.

Details

The first part of the additional information concerns the details of the model as shown in **Figure 2**.

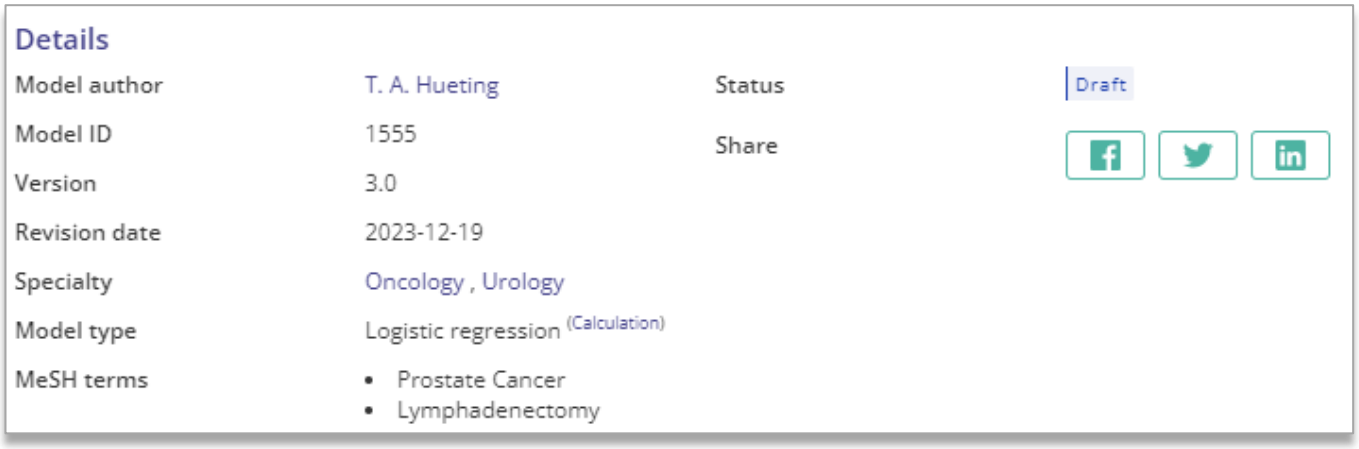

***Figure 2**. The model details.*

Study characteristics

Below the 'Details section' the section labelled 'Study characteristics' provides information on the characteristics of the patient data used to derive and validate the model. Additional information is provided on the methods used to develop and/or validate the model.

An important part of the Study characteristics is the information on Supporting publications and related files. These sections can be found at the bottom of the Details-pop-up as shown in **Figure 3**.

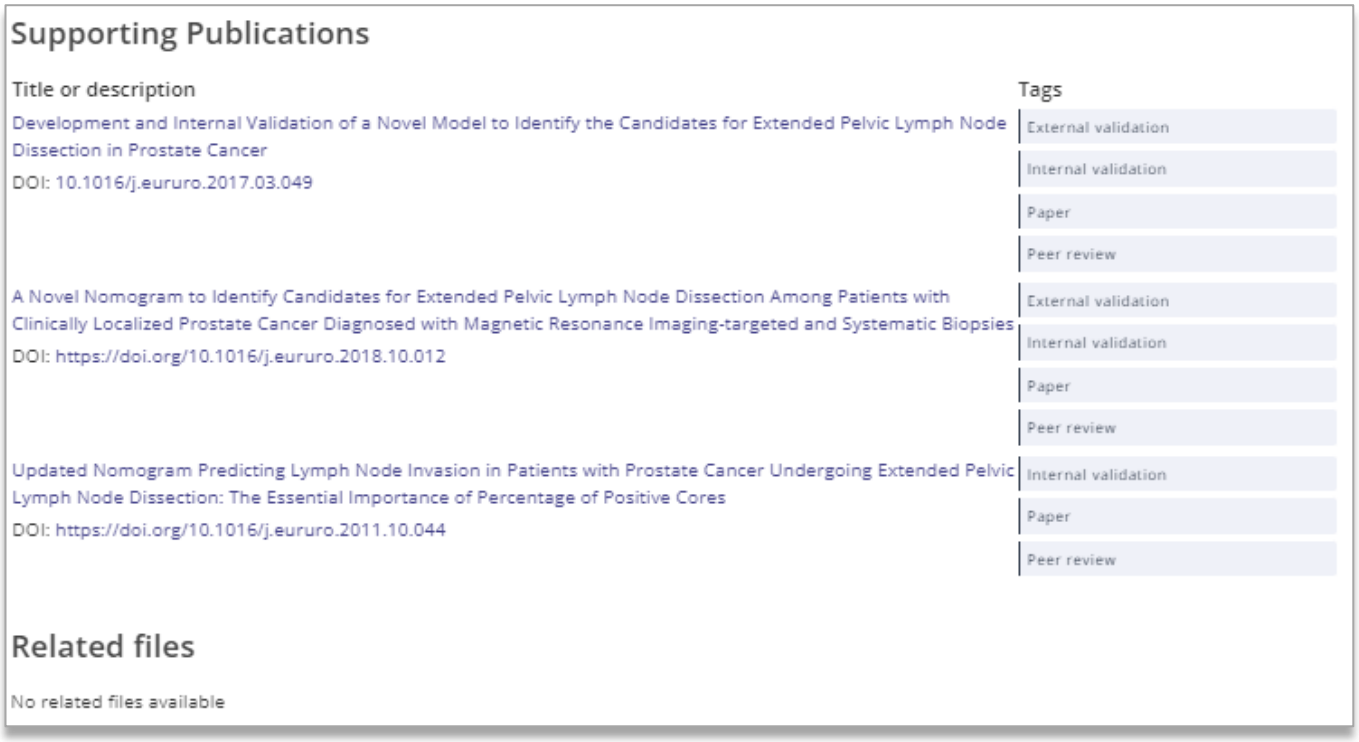

***Figure 3**. An example of Supporting publications & related files.*

H. Intended use button

Intended medical use

The device is intended to be used by professional users who are capable of operating the device and interpreting its results. It can be used to estimate the probability of lymph node involvement in patients with clinically localized Prostate Cancer (PCa). The medical device software (MDSW) includes three algorithms, available, the 2012, 2017 and 2019 algorithms.

The device combines preoperative Prostate Specific Antigen (PSA), Clinical T-stage, primary Gleason Grade, Secondary Gleason Grade and Percentage of positive cores for the 2012 algorithm, preoperative PSA, Clinical T-stage, Biopsy Gleason Grade Group, Percentage of positive cores with highest-grade PCa and percentage of positive cores with lower grade PCA for the 2017 algorithm and Preoperative PSA, clinical stage at mpMRI, maximum lesion diameter at mpMRI, biopsy Gleason grade group at MRI-targeted biopsy and percentage of cores with clinically significant PCa at systematic biopsy for the 2019 algorithm, to predict the risk of pelvic lymph node involvement. Using a specific version depends on the available patient data.

The device is intended to be used for patients with clinically localized PCa. The result of the device is intended to be reviewed and interpreted by qualified medical specialists only. The device is not intended for use by patients on their own.

The device is not intended to replace clinical decision-making. It can only provide information to the user on the estimation of pelvic lymph node involvement. The user can use this information to support clinical decision-making regarding optimal treatment options. In practice, this typically entails the decision to perform an extended pelvic lymph node dissection.

Clinical Benefit

Correct functioning of the Briganti Nomogram can result in these clinical benefits:

- The Briganti Nomogram can assist in risk stratification of patients.
- Risk stratification can reduce the burden of (invasive and intensive) medical procedures in patients with low risks, reducing, shortening or avoiding adverse events caused by the procedures.
- Risk stratification can reduce the unnecessary consumption of (scarce) medical resources, decreasing costs and increasing their availability for high risk patients.

Intended target population and exclusion
The Briganti nomogram is intended to be used only for a specific group of patients, corresponding to the below indications and contra-indications.

Clinical inclusion criteria
The Briganti nomogram is intended for:
- Patients with clinically localized PCa

Clinical exclusion criteria
The Briganti nomogram should not be used for patients that do not fulfil the inclusion criteria, i.e. it is not intended for:
- Patients with incomplete pathologic or biopsy data required for calculation of (one of) the Briganti Nomogram(s)

User profile
The Briganti nomogram is intended to be used by Healthcare Professionals or automatically calculated through Evidencio's API. Results shall always be reviewed and interpreted by qualified medical specialists only, in the context of the patient's clinical history and other diagnostic test results. Healthcare professionals do not require additional training prior to the use of the medical device. The device is not intended for use by patients on their own.

Intended use environment
The MDSW can be used as made available on the Evidencio platform in any actively supported web-browser on personal computers, mobile devices, or tablet PCs, and on the mobile app provided by Evidencio. The MDSW can also be used through Evidencio's iFrame representation as an embedded view, provided that the specific Evidencio guidelines for iFrame implementations of this MDSW are adhered to. Automated calculation of the device is enabled through Evidencio's API. The device is only intended for use in healthcare settings where the immediate application and outcomes of the device are not required.

Physical interaction
The MDSW is stand-alone software and does not come into contact with any bodily or other material of the patient, user or otherwise.

Versions of the MDSW
The versions of the Briganti Nomogram concerned in this document were developed by Briganti et al. in 2012. Gandaglia et al. in 2017, and Gandaglia et al. in 2019 [R1, R3, R4], resulting in the Briganti 2012 Nomogram, Briganti 2017 Nomogram, and Briganti 2019 Nomogram respectively. The versions mainly differ in the variables required to calculate the risk of Lymph Node Invasion. The 2017 model adds a precise assessment of cancer involvement within the biopsy core and intraprostatic heterogeneity to the Briganti 2012 Monogram model, while the Briganti 2019 Monogram model is specifically based on multiparametric magnetic resonance imaging data instead of ultrasound-based data.

Functioning, physical principle
The MDSW's underlying mathematical formula concerns a logistic regression based statistical model. The acquisition and processing of the data, the analyses to assemble the relevant criteria for the MDSW as well as the setup and refinement are provided in the instructions for use. Entering the details for an individual in the MDSW initiates the estimation of pelvic lymph node involvement risk.

I. Electronic label button

The electronic label button opens a pop-up with the location and address of Evidencio, the LOT number, the UDI number, the CE-mark, the medical device logo and a download link for the declaration of conformity of the medical device. The example of the electronic label is shown in **Figure 4**.

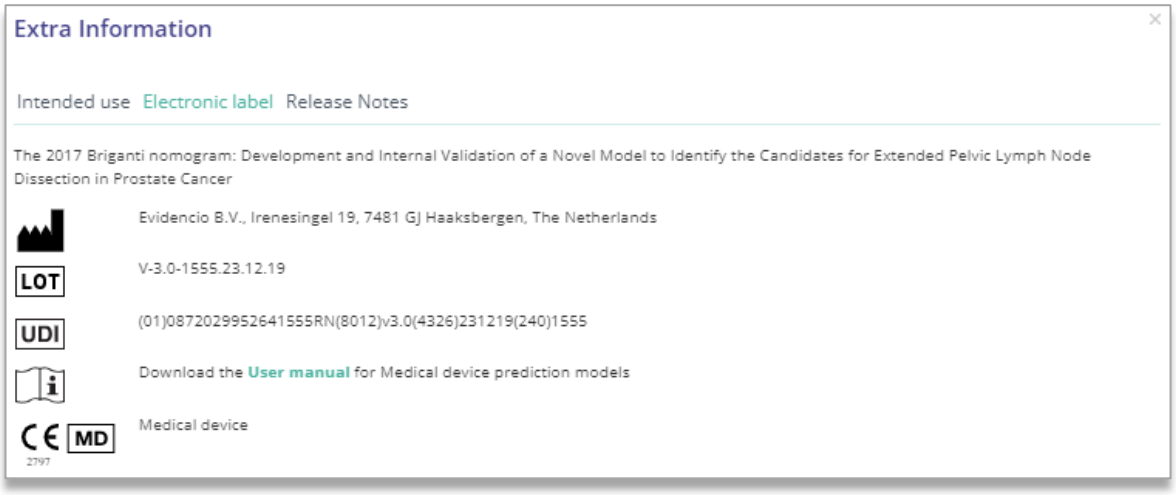

Figure 4. Example of the electronic label

J. Release notes

The 'Release Notes' button opens a pop-up with the latest release notes of the model. Here you can find what has changed over the last versions of the model. Additionally, if there are any known residual anomalies the user should be aware of, they are listed here.

K. User Manual

This user manual can be found in three places: 1) under the short description, 2) on the right of the model page, and 3) in the electronic label. Additionally, all versions of the user manual can be found in the general page for all user manuals for medical devices. The page can be found under the 'About' drop-down menu button as shown in **Figure 5**. The user manual page is shown in **Figure 6**.

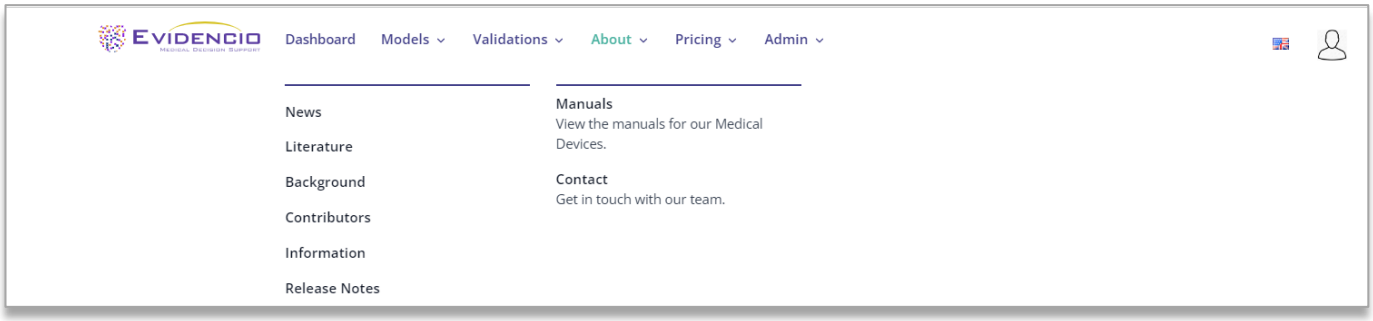

Figure 5. The drop-down menu where the user manual page can be found.

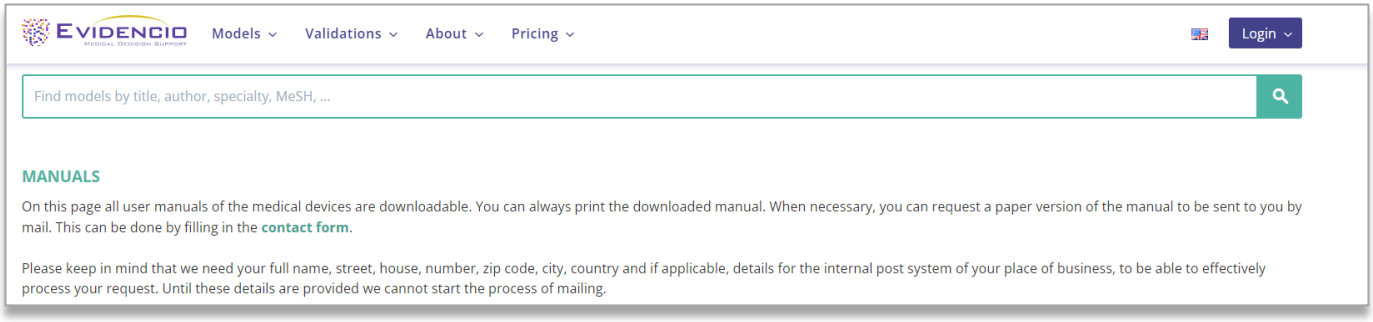

Figure 6. The user manual page for all user manuals.

You (The user) can always print this downloaded manual. When necessary, you can request a paper version of the manual to be sent to you by mail. Evidencio's contact details are listed in Chapter 6 of this user manual.

L. Input section

The Evidencio platform allows two separate input variables; categorical, and continuous variables.

Categorical variables

In the example shown in Figures 7 and 8, the **Biopsy Gleason grade group** variable concerns a categorical variable. The patient status can be entered by clicking on either button. The selected button changes to green, as seen in Figure 8.

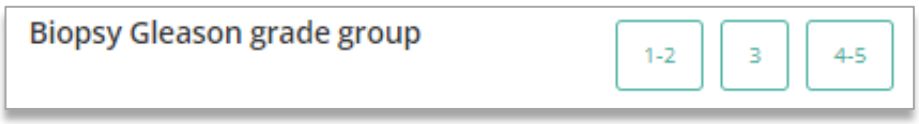

Figure 7. The variable for Biopsy Gleason grade group, where no button has been clicked, and thus no input has been provided by the user.

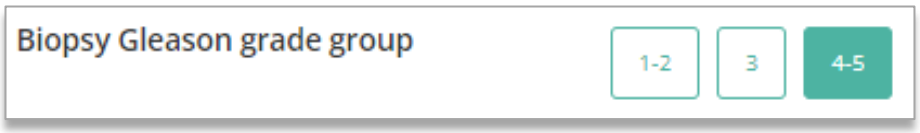

Figure 8. The variable for Biopsy Gleason grade group, where the "4-5" button has been clicked.

Continuous variables

In the example shown in Figure 9, the **Preoperative PSA** variable, exemplifies a continuous variable. The plausible ranges for the variables are used for the model.

The details for a patient can be entered by sliding the button to the correct value, or by entering the correct value in the box on the right-hand side (i.e., where the 20 is entered for **Preoperative PSA**).

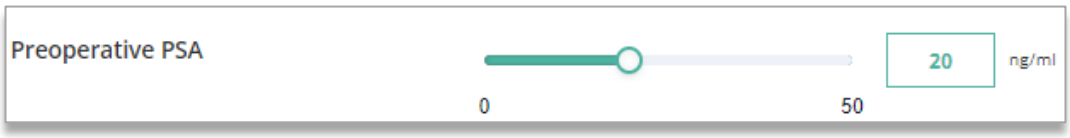

Figure 9. The variable for Preoperative PSA, where "20" has been entered

M. Result section

At the bottom of the page, the results of the model are shown.

Result calculation

When all variables are filled in, a result will be calculated. No risk is displayed until all variables are filled in. The result section indicates *"Set all parameters to calculate prediction."*

Result interpretation

In the result interpretation, a risk stratification is given based on the risk score. An example of the information is shown In **Figure 10**.

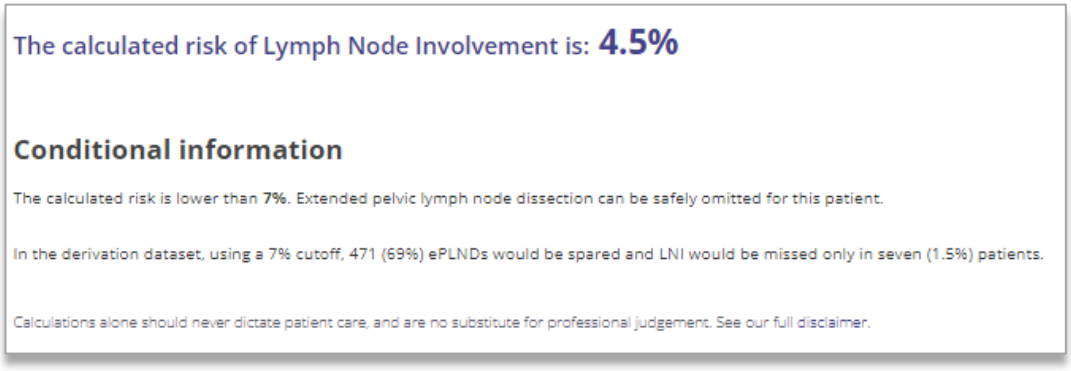

Figure 10. The result information

Relevant information for correct use of the model

At the bottom of the page, there is a link to Evidencio's terms and conditions of use, the privacy policy, and the disclaimer.

N. Versions

With this button, different versions, corresponding to different publications of the Briganti Nomogram, can be selected. Version **2.0** is the Briganti 2012 Nomogram, Version **3.0** the Briganti 2017 Nomogram, and Version **4.0** the Briganti 2019 Nomogram, originally published in **2012**, **2017** and **2019** respectively.

***Figure 10**. The Version selection window.*

5. Use of Medical devices

In general, and unless explicitly stated otherwise, CE-marked tools on Evidencio are only to be used by professionals, and are not for patient use.

To use the tool, Evidencio requires a stable internet connection and runs on the following devices:

- Personal computers or laptops using the following browsers:
  - Safari (the latest three versions)
  - Chrome (the latest three versions)
  - Firefox (the latest three versions)
  - Edge (the latest three versions)
- Tablets or smartphones running on the next operating systems:
  - IOS (the latest three versions)
  - Android (the latest three versions)

The medical device cannot be used in combination with Internet Explorer. The personal computers, laptops, tablets or smartphones used should at least be able to have an internet connection and use the browsers mentioned above. The minimal screen resolution should be 800x600.
Furthermore, the model may be used through the Evidencio iFrame representation of the calculator, as an embedded view, provided that the specific Evidencio guidelines for iFrame implementations of that model are adhered to.

The Evidencio MDSW models can be used with any browser settings that don't distort the regular display of websites, with a 50% to 500% zoom rate, and at a display resolution starting from 800x600. However, factory recommended browser settings, 100% zoom rate and regular display resolution are recommended.

This model is only intended for use in settings where the usage and result of a model are never immediately needed.

The Briganti nomogram can also be integrated in third party applications over the API. To use the API tools provided by Evidencio, authorization is required with an API key and access to the device must be granted. API keys and algorithm access requests must be submitted to info@evidencio.com. Specific requirements apply for the use of the device over the API. Use specifications are available upon request, and will be shared upon receiving the API key.

6. Manufacturer details

Any serious incident that has occurred in relation to the device should be reported to the manufacturer and the competent authority of the country in which you, the reader, are established. A competent authority is the institute that governs all issues related to medical devices in a country.
Contact details of your competent authority can be found here: https://www.ema.europa.eu/en/partners-networks/eu-partners/eu-member-states/national-competent-authorities-human

Please contact Evidencio when you suspect any malfunction or changes in the performance of a medical device. Do not use the device, until Evidencio replies to your message that it is safe to start using it again.

Contact details of Evidencio:

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