EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

# Clinical Trial Information System (CTIS) Bitesize talk

## Notifications (part 1)

Presented by Laura Pioppo and Ana Rodriguez on 28 September 2022
European Medicines Agency

The European Medicines Agency developed this training material to enhance public access to information on the Clinical Trial Information System (CTIS). This material describes a preliminary version of CTIS and may therefore not entirely describe the system as it is at the time of use of this material. The Agency does not warrant or accept any liability in relation to the use (in part or in whole) or the interpretation of the information contained in this training material by third parties.

# CTIS Bitesize talk:

## Notifications: trial and recruitment period

| | |
|---|---|
| 14:30 - 14:35 | **Introduction** |
| 14:35 – 15:55 | **CTIS Demonstration Sessions followed by live Q&A Sessions** |
| 15:55 – 16:00 | **Closing remarks** |

*For questions, go to **www.sli.do** & use event code **#bt28sep***

# A few housekeeping rules

For your questions,
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Or scan
**slido**
**QR code:**

## Tips for optimal screen viewing

- Make use of the instructions under the embedded video in the event page and connect directly to the **IBM channel** *for the full-screen experience*
- Increase the *video quality* from the **HD button** on the right bottom of the screen setting it to 720p (or 1080p).
- Have a stable internet connection

# A few housekeeping rule

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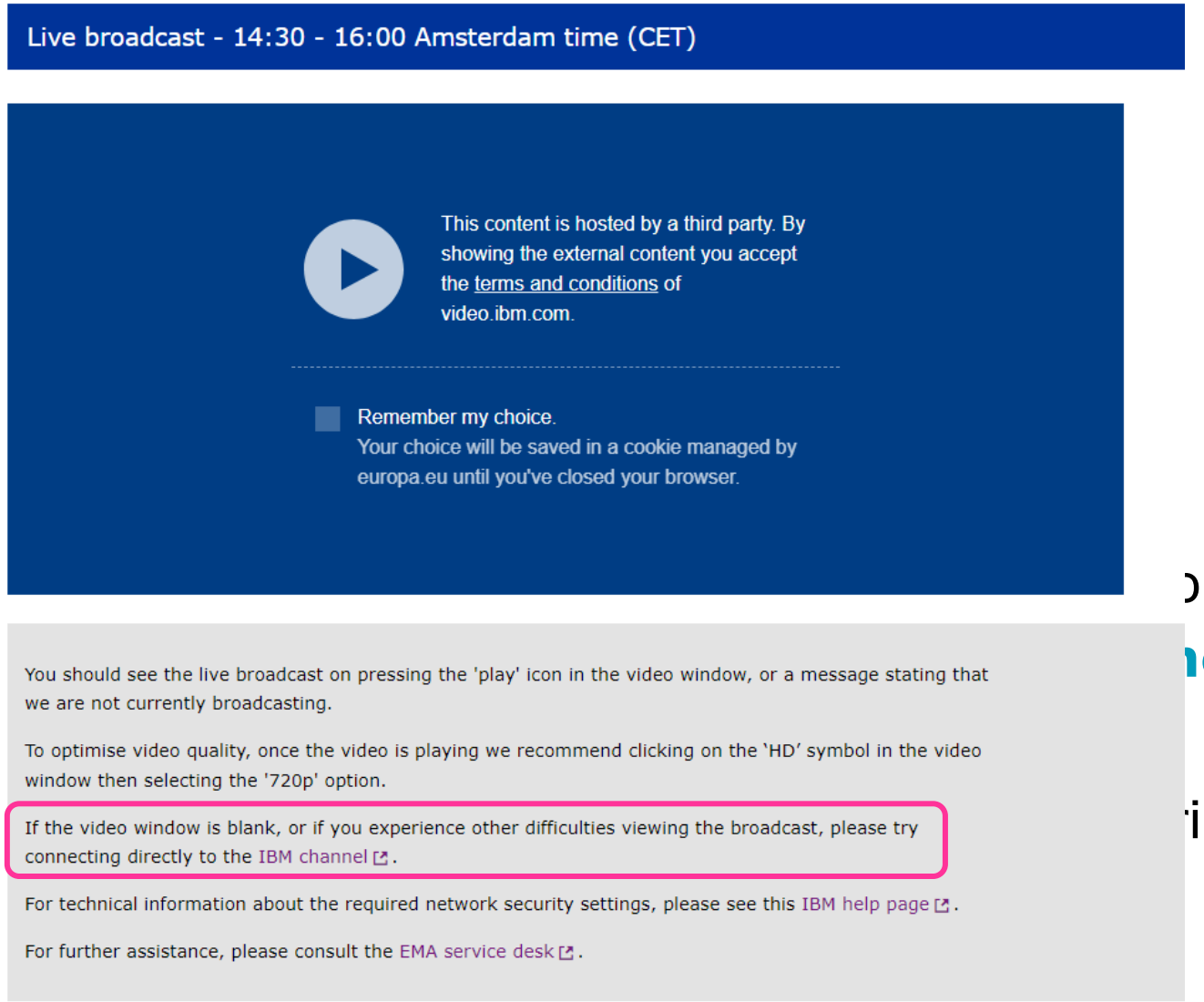
- Increase … botto … right



# Main Session: CTIS Demo with Q&A

## Notifications:

### trial and recruitment period

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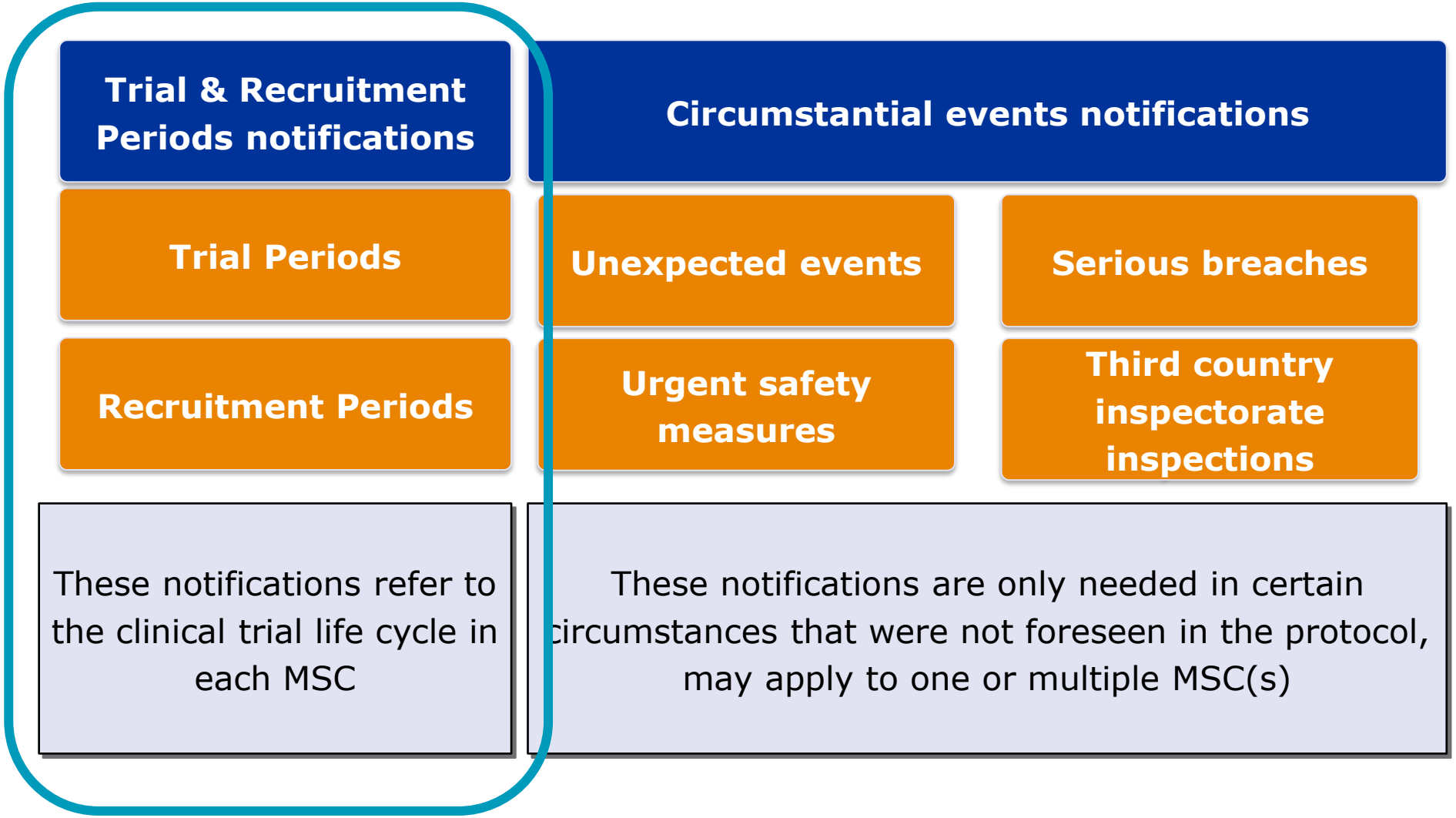
CTIS Delivery Workstream Lead

# Notifications



# Trial and recruitment period notifications

**Start trial**

*The first act of recruitment of a potential subject for a specific CT*

It could be the date of initiation of the clinical trial in the first site, or the date when the first trial specific advertisement is published

**Start recruitment**

*The first visit of the first subject.*

Recruitment shall begin within 2 years from trial authorisation

To extend start recruitment beyond 2 years, substantial modification requires to be authorized before initial timeline expires

**End recruitment**

*The act of not recruiting anymore subjects in a MSC*

**End trial**

*The last visit of the last subject, or a later point in time as defined in the protocol*

*Includes **also early terminations** and whether related to a matter of subject safety and/or benefit-risk balance, or not*

# Trial and recruitment period notifications

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**Temporary halt**

*An interruption not provided for in the protocol of the conduct of a trial by the sponsor with the intention to resume it*

There are two types depending if the temporary halt is related to subject safety and/or benefit-risk balance (B/R), or not

A halted trial is automatically ended if not restarted within 2 years

For an extension to restart the halted trial beyond 2 years, substantial modification requires to be authorised before initial timeline expires

**Restart trial**

*The act of restarting the trial after a temporary halt, or after suspension of the trial by an MSC*

If temporary halt is related to subject safety and/or benefit-risk balance, a **prior substantial modification must be submitted and authorised** to enable notification

**Restart recruitment**

*The act of restarting the recruitment of subjects*

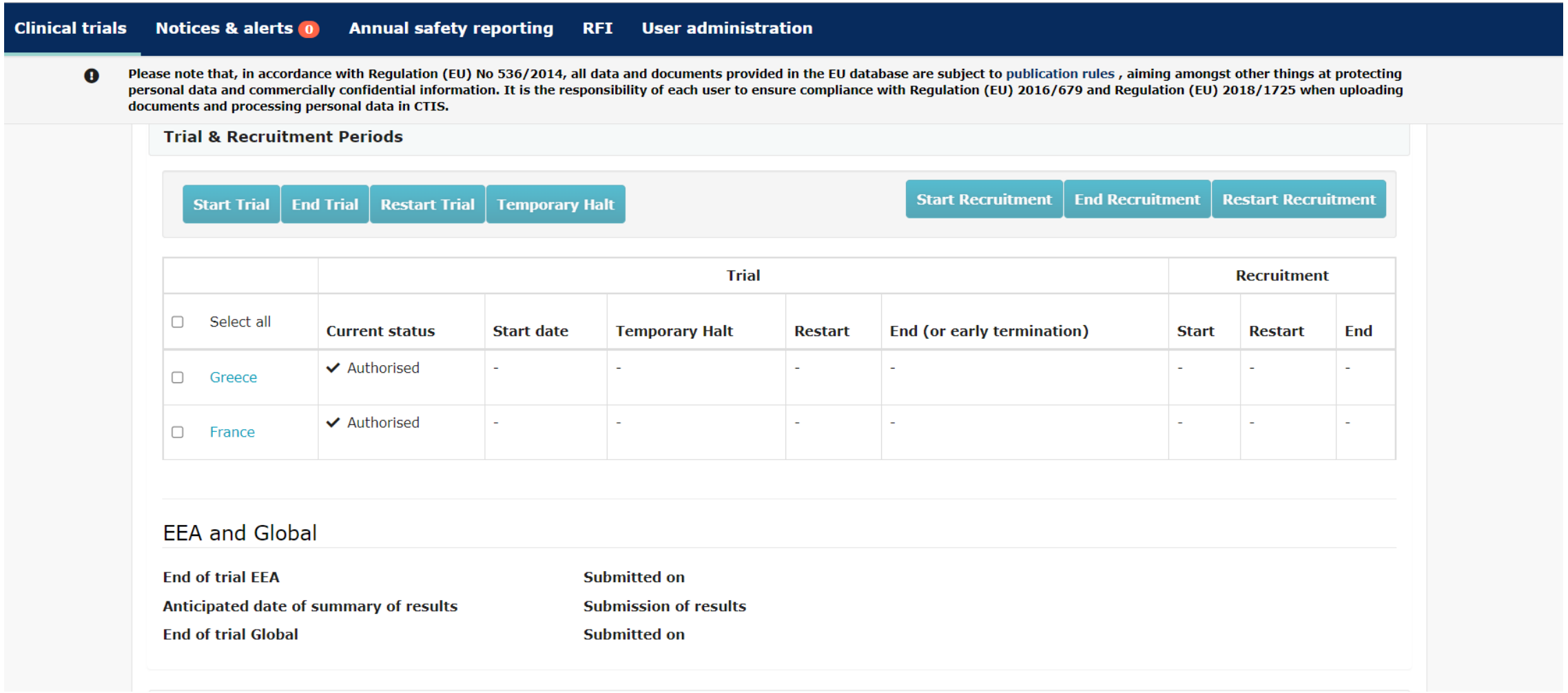
The trial must be restarted to enable restart the recruitment notification

# Trial and recruitment period notifications

- The legal basis for notifications submission is in Article 36 and Article 37 of the Regulation (EU) No 536/2014
- The relevant training material on the topic is in Module 5 published on EMA website - online training modules: https://www.ema.europa.eu/en/human-regulatory/research-development/clinical-trials/clinical-trials-information-system-ctis-online-modular-training-programme
- Relevant clarifications can be found also in section 10 of the COM Q&A document: https://health.ec.europa.eu/system/files/2022-09/regulation5362014_qa_en.pdf

  Latest version 6.2 published in Sept 2022

# Trial and recruitment period notifications



# Trial and recruitment period notifications

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- Notifications to a MSC can be submitted as soon as the trial is authorised by that MSC
- The sponsor does not need to wait to have the trial authorised in all MSC before a notification can be submitted
- As soon as a trial is authorised the sponsor can submit Start of trial (SoT) or early termination notifications
- Submission of these notifications is sequential and follows predefined logic
- Notifications can be submitted to one or more MSC simultaneously with the multi select functionality
- CTR defines specific timelines for notifications submission however the system is flexible in terms of business rules to allow submission beyond due date (MS can monitor compliance though)
- Documents can be attached to the notifications – but this is optional

# Demo on SoT/SoR/EoR and 1st Q&A session

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# Trial and recruitment period notifications

Demo includes:

- Entry point for notifications
- How to populate and submit a notification
- Document upload
- Sequential flow
- Update functionality
- Withdrawal
- Publication aspects

# Temporary halt and restart of trial

# Trial and recruitment period notifications

## Temporary halt and restart of trial

Article 2(2)(28)

*Temporary halt of a clinical trial' means an interruption not provided in the protocol of the conduct of a clinical trial by the sponsor with the intention of the sponsor to resume it;*

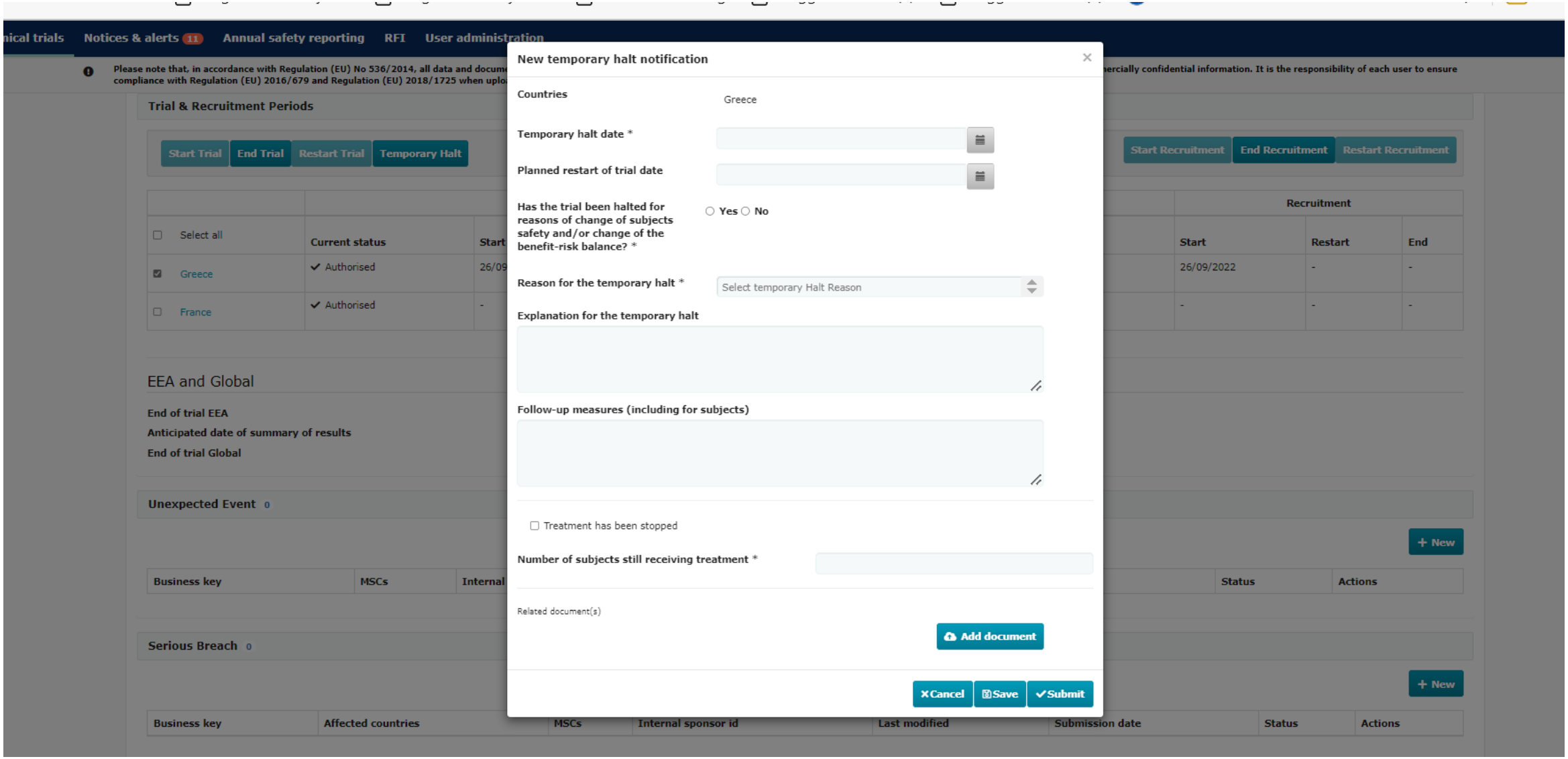
- Sponsor can stop trial activity described in the protocol (i.e., recruitment only or recruitment and treatment), due to unexpected circumstances that could (or not) affect the benefit/risk ratio
- In case of a change of the B/R the sponsor can only restart the trial if a substantial modification (part I only, part II only or part I and II) has been authorised by the MSC
- Substantial modification to restart should be submitted with 2 years since the halt started

# Trial and recruitment period notifications

From COM Q&A:

- In case of temporary halt that have no potential effect on the benefit/risk balance (e.g., lack of supply of IMP/shortages), the sponsor should notify in CTIS the restart of trial within 15 days from its occurrence
- If a temporarily halted CT is not resumed within **two years**, the expiry date of this period or the date of the decision of the sponsor not to resume the clinical trial, whichever is earlier, shall be deemed to be the **date of the end of the CT**.
- A sponsor can submit within the two-year period following a temporary halt a SM requesting a restart date **after** the 2-year period. This SM can only be submitted **before the expiry** of the 2-year period and applies to temporary halts for reasons of subject safety or not.

# Trial and recruitment period notifications

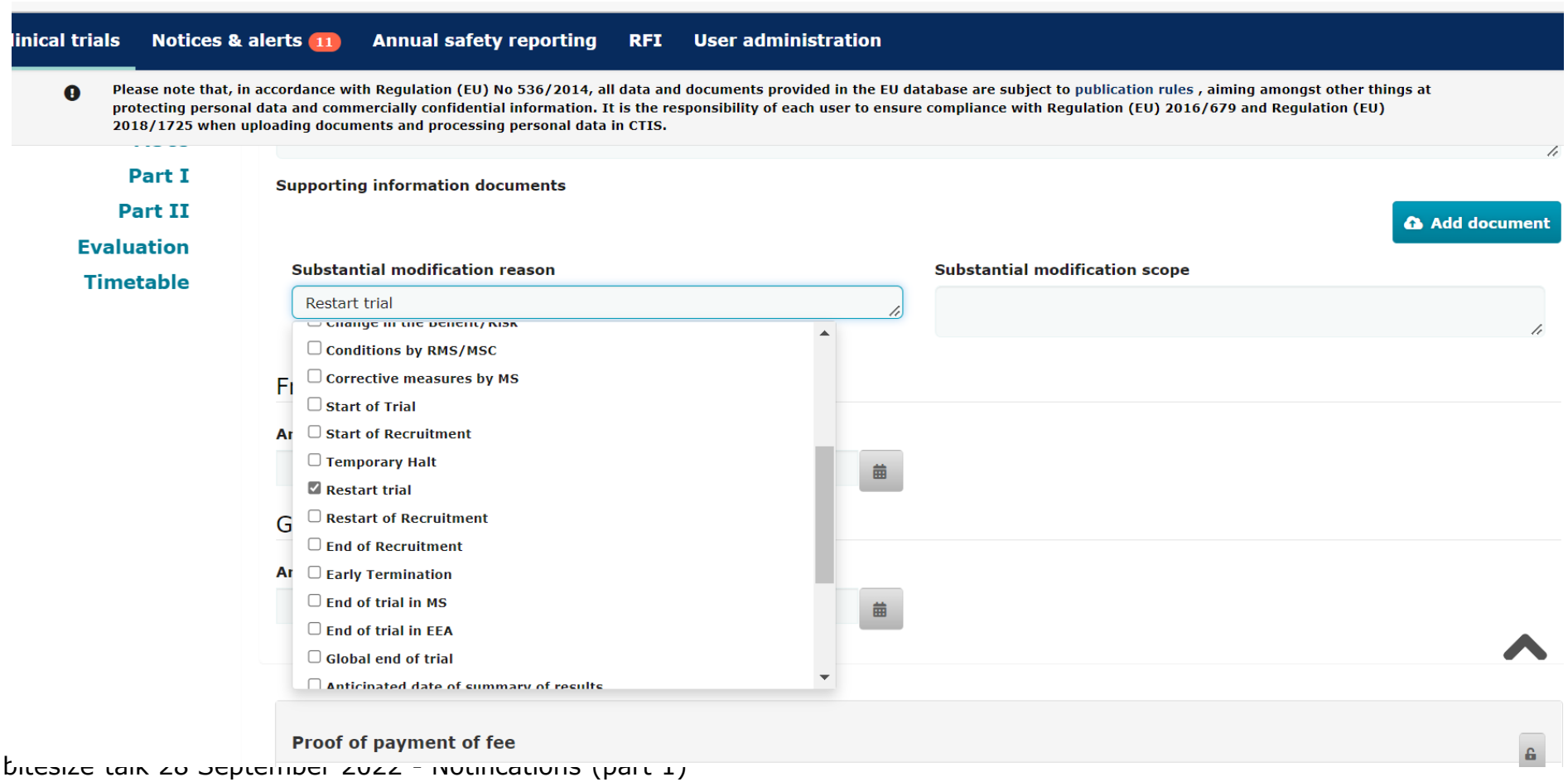


# Trial and recruitment period notifications

- A temporary halt (TH) not B/R related is published as soon as they are submitted
- A temporary halt B/R related is also published as soon as it is submitted with the exception of the field on reason/explanation for TH
- After an assessment is performed by the MSC the reason/explanation of the TH form is published
- Restart of trial for temporary halt not B/R related can be submitted, as applicable, when issue solved
- Restart of trial for temporary halt B/R related can only be submitted after an SM has been authorised

# Trial and recruitment period notifications

- It is very important that when a substantial modification is submitted to restart a trial following a temporary halt impacting B/R (benefit/risk) the sponsor select the correct value ‘’ restart trial” from the SM form



# Demo on Temporary halt and restart 2nd Q&A session

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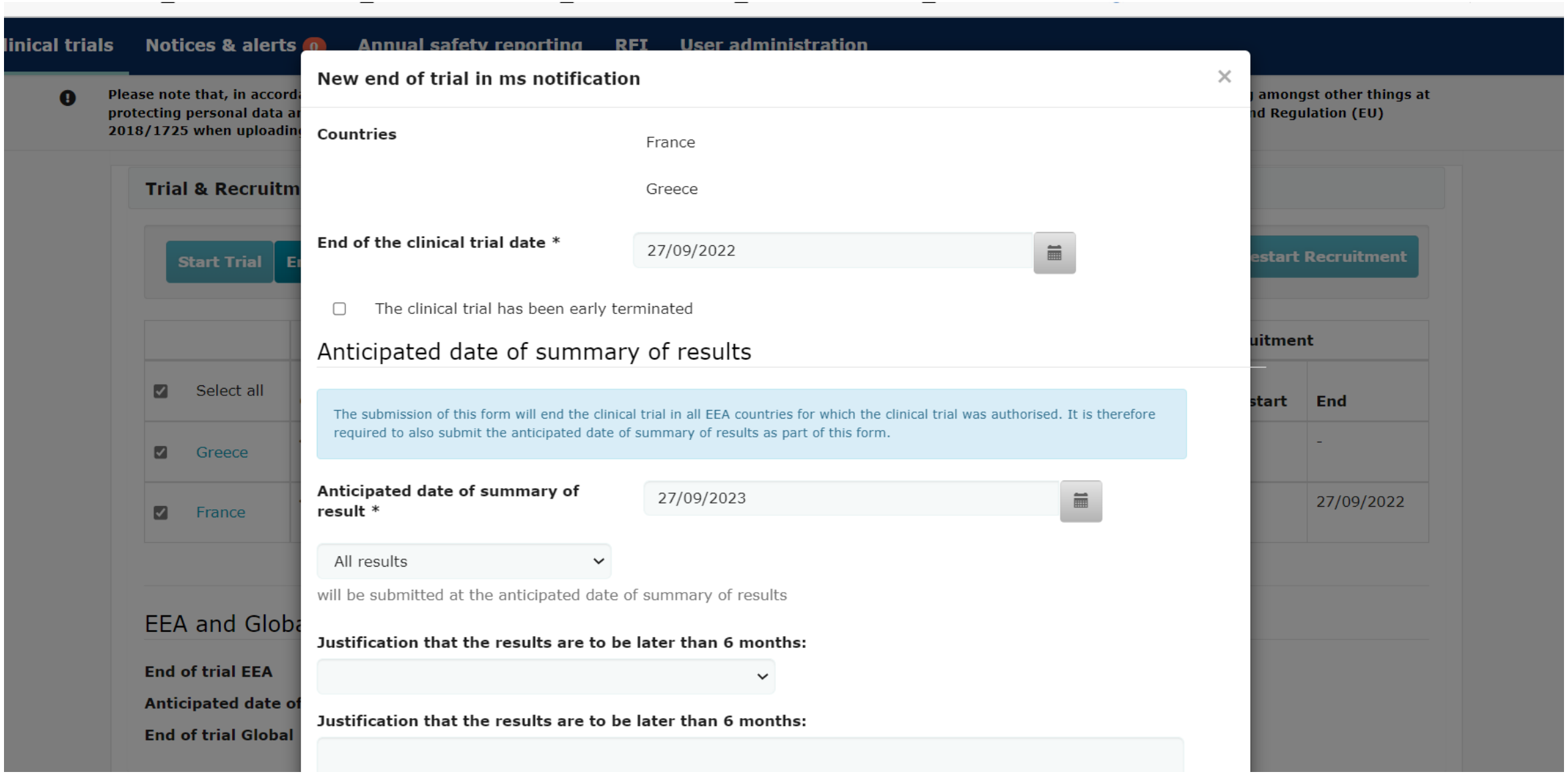
# End of trial and summary of results

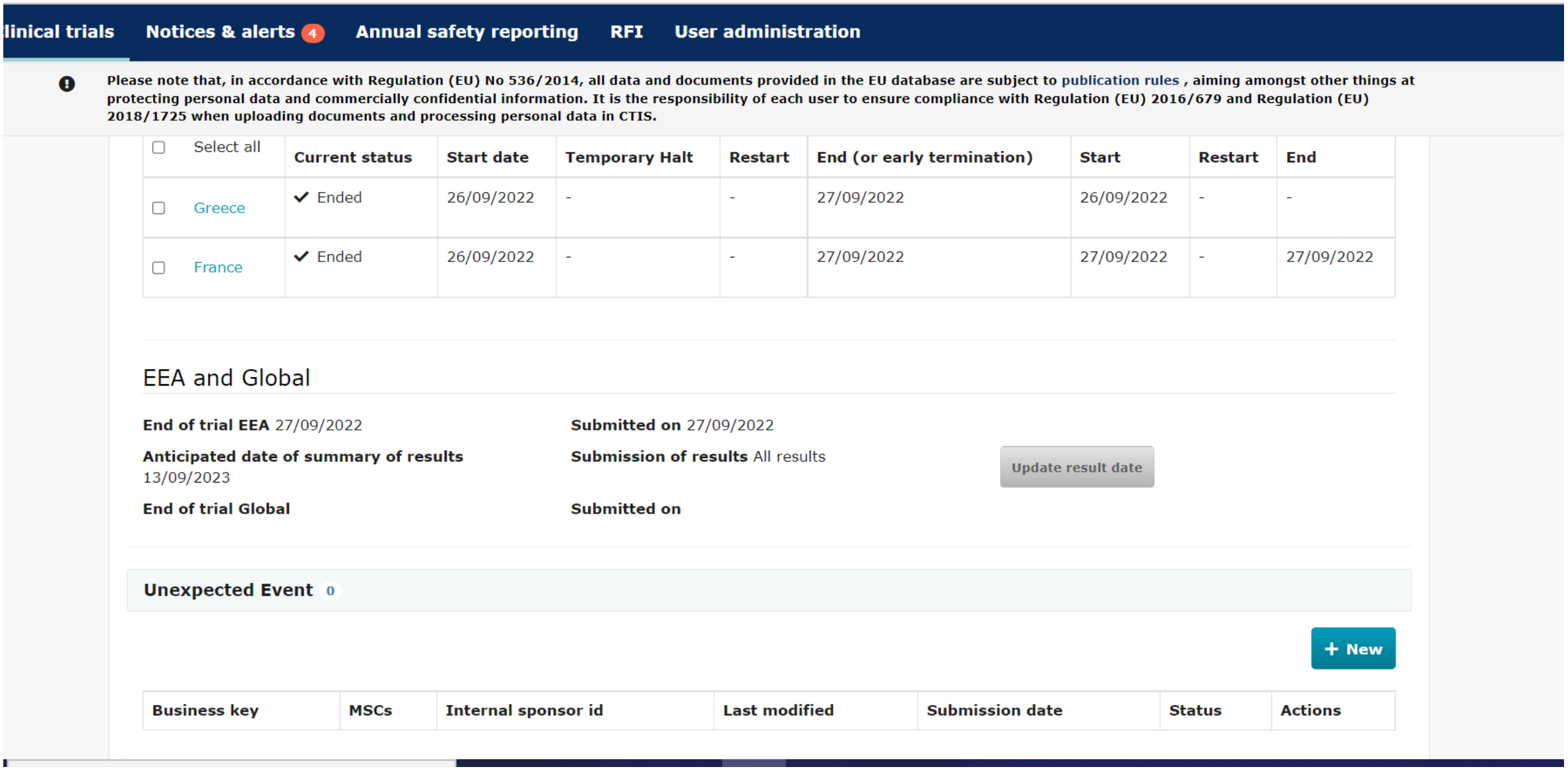
# Trial and recruitment period notifications

- End of trial notifications (like the other period notifications) can be submitted for individual MSC
- Last end of trial notification in a MSC is equal to end of the trial in the EU/EEA
- When completing the end of trial in the EU/EEA the sponsor is prompted to specify the anticipated date for submission of summary of results

# Trial and recruitment period notifications



# Trial and recruitment period notifications



# Demo on end of trial and last Q&A session

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# We ask for *your feedback* on this event

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# Info on upcoming events

# Upcoming CTIS events 2022 – Save the date

***Submit your questions in advance!***

| Date | CTIS event |
|---|---|
| 19 Oct | OMS TroubleShooting Session for CTIS users |
| 5 Oct<br>20 Nov | CTIS Walk-in clinics |
| 16 Nov | CTIS Webinar - 9 months on and going forward |
| 23 Nov | CTIS bitesize talk: Notifications - Part 2 |

*Clinical Trials Information System: training and support | European Medicines Agency*

*For satisfaction feedback, Please go to* ***www.sli.do*** *& use event code* ***#bt28sep***

# Thank you for attending today's event

*Next CTIS bitesize talk on 23 November*

## Further information

For CTIS communication, training & change management queries, e-mail CT.Communication@ema.europa.eu

For the Clinical Trials Newsletter sign up at CT.NewsletterSubscriptions@ema.europa.eu.

For upcoming CTIS events, visit the EMA event page.

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