

The logo for the European Bioanalysis Forum (EBF) is located in the top right corner of the slide. It consists of the letters 'EBF' in a white, sans-serif font. Below the letters is a white graphic element consisting of two curved lines that sweep upwards and to the right, resembling a stylized 'E' or a checkmark.

European
Bioanalysis
Forum

EBF Focus Workshop

Discussion points for Breakout

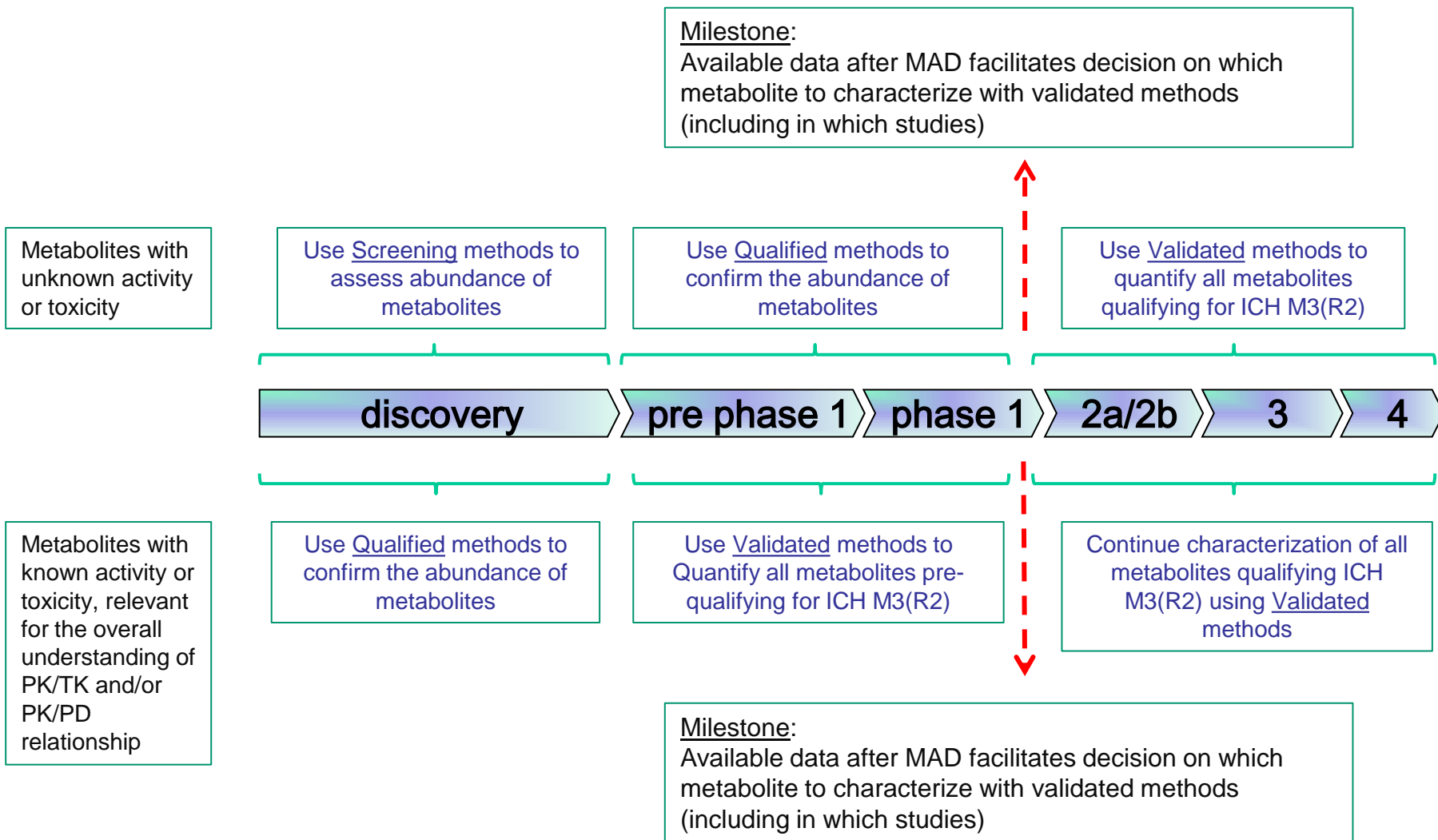
Philip Timmerman, on behalf of EBF

Introduction

The dual challenge:

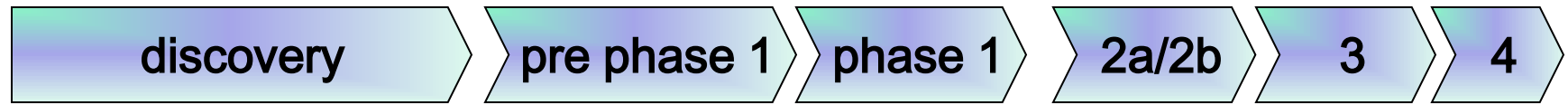
- Metabolite profiling
- Metabolite quantification

EBF position (2010) on Metabolite quantification



Ref: Best practices in a tiered approach to metabolite quantification: views and recommendations of the European Bioanalysis Forum. *Bioanalysis*. 2010 Jul;2(7):1185-94.

EBF position (2010) on regulations for BA



non GLP

Screening

Or

Qualified
Methods

Clinical, GLP and non GLP

Validated Methods

(non) rodent plasma (unchanged drug)
Other matrices - metabolites as appropriate

Human plasma
Other matrices - metabolites as appropriate

Additional preclinical species
Other matrices - metabolites as appropriate

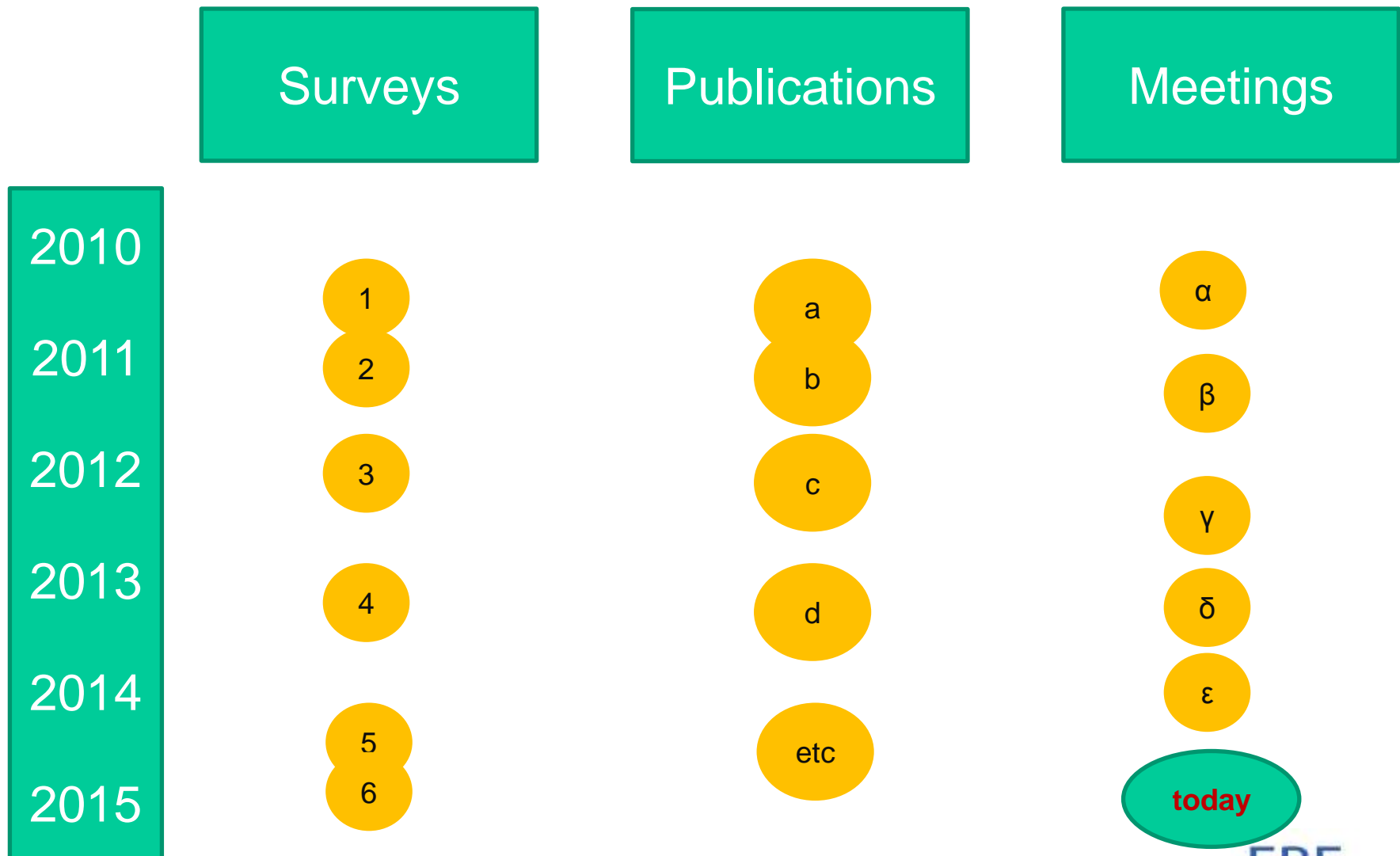
Qualified Methods

- Samples from:
 - *In vitro* (e.g. PPB),
 - some mechanistic PK, PK/PD
 - most non standard matrices (e.g. tissues)
- Quantification of metabolites

EBF position (2010) on Metabolite Profiling

Although we gathered some data in our first survey in 2009, this subject was out of scope

A lot happened since → new insights increased experience

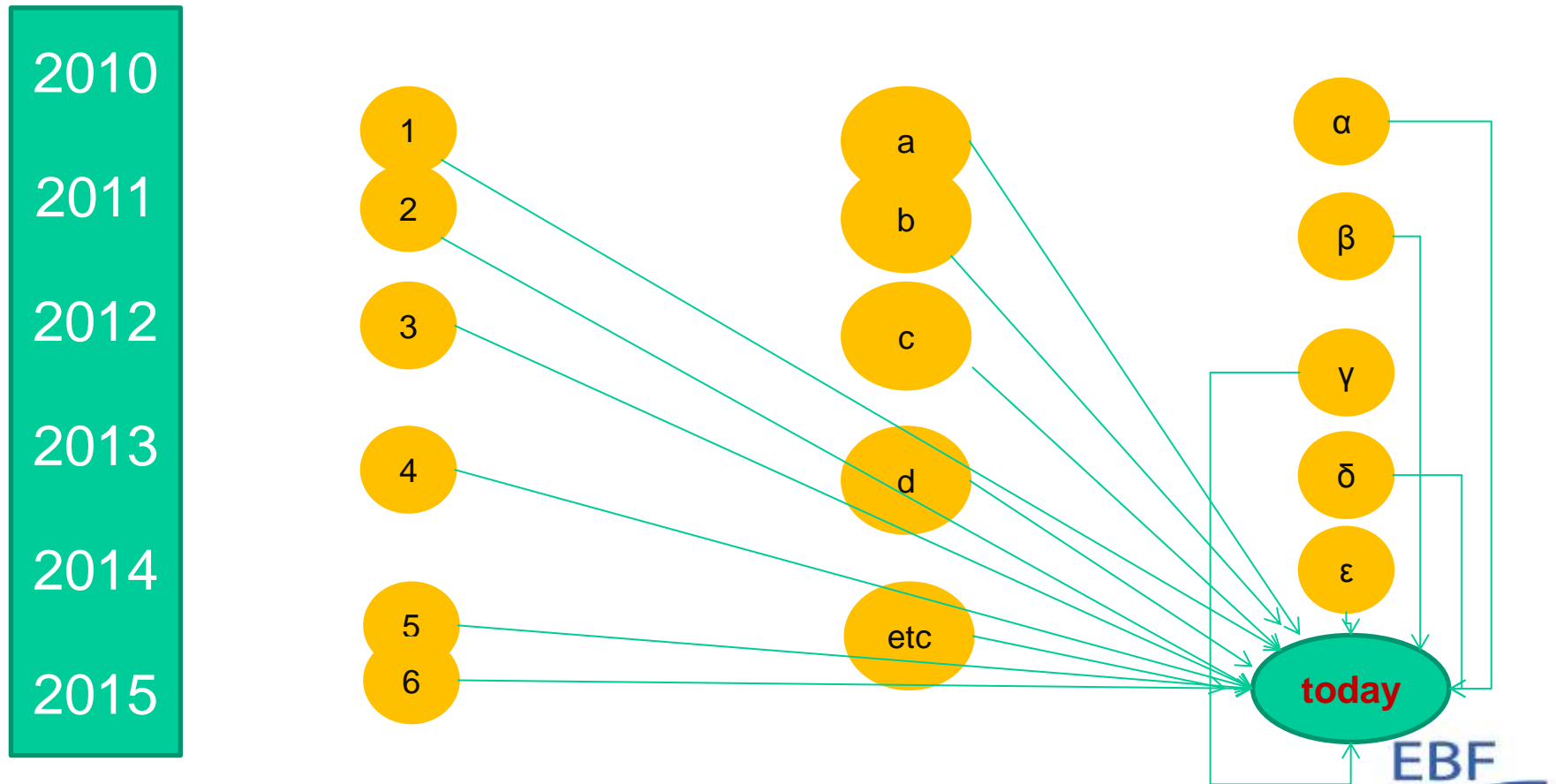


And we want to use them.....

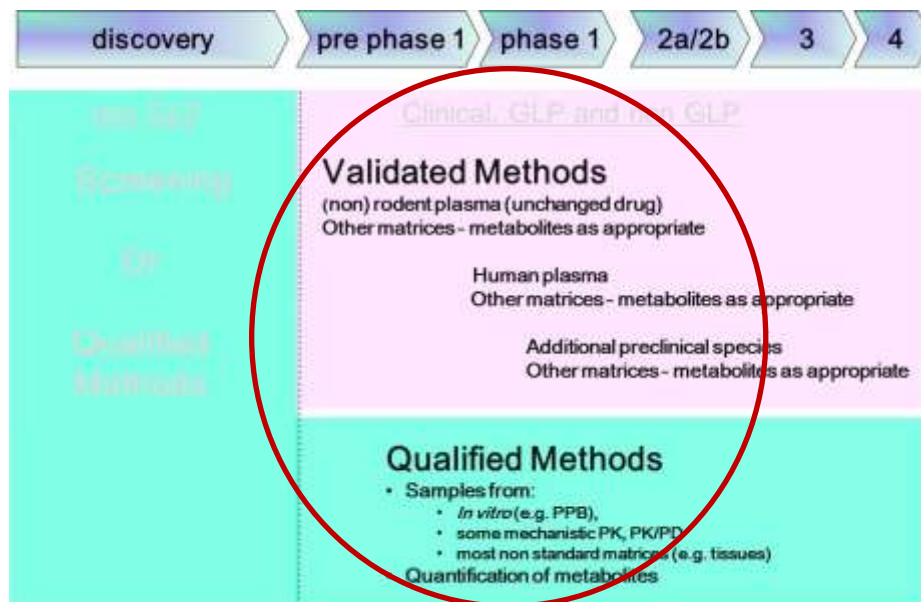
Surveys

Publications

Meetings



Updated EBF position on regulations for BA



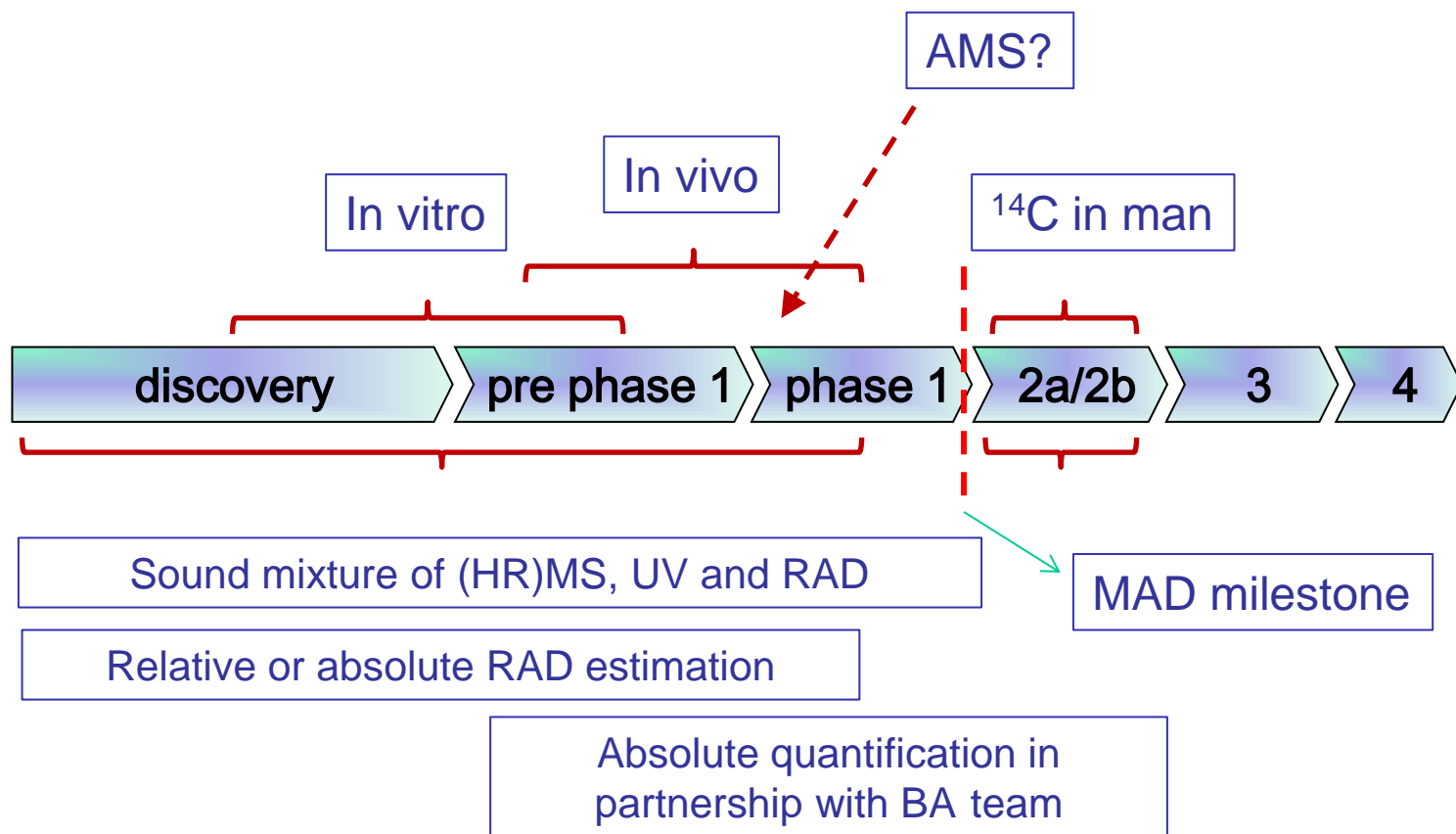
Refined definition of validated methods and proposed timing/usage into

- Regulatory validation:
 - Late stage studies
- Scientific validation:
 - 5 areas: dosed/unchanged drug/metabolites in tissue homogenates, urine, early clinical (e.g. SAD/MAD), early preclinical and 'pre-ICH(M3)' metabolites in plasma
 - Provided guidance on pre and in study validation criteria, study acceptance and reporting for regulatory validation

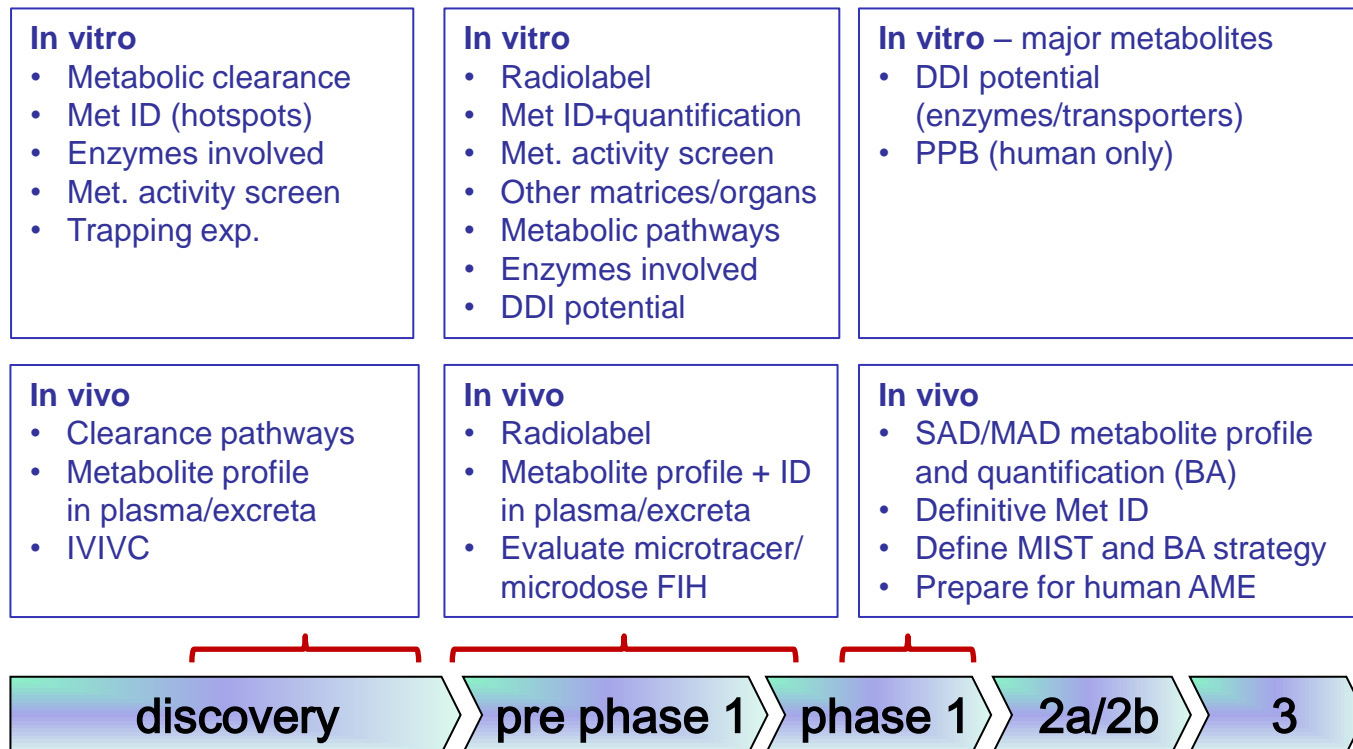
Ref:

- *Tiered approach into practice - scientific validation for chromatography-based assays: a recommendation from the European Bioanalysis Forum. Bioanalysis, in press*
- *Feedback from the European Bioanalysis Forum Workshop: taking tiered approach to the next level. Bioanalysis 6:19, 2593-2598.*

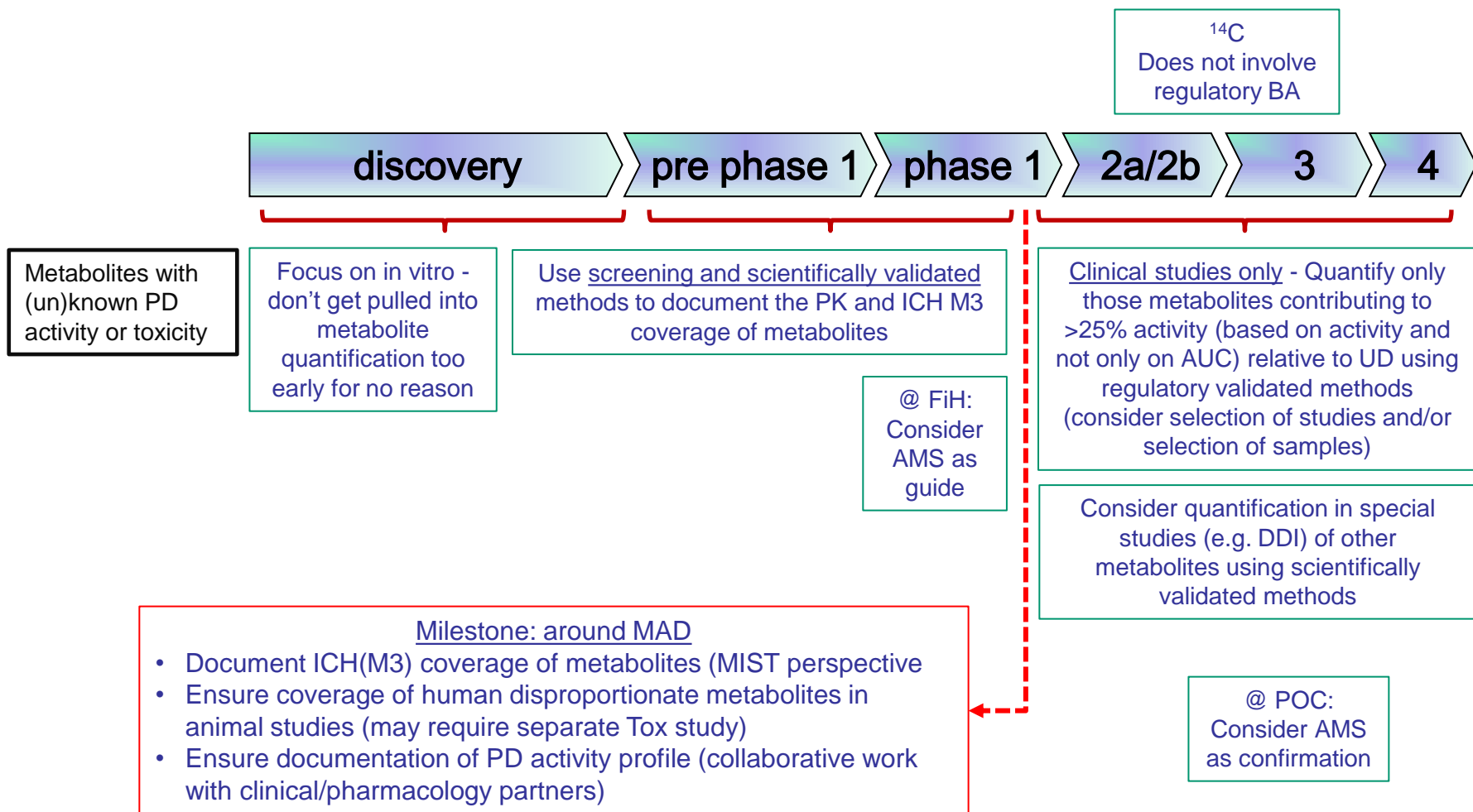
Starting proposal Metabolite profiling – when and how?



Starting proposal Metabolite profiling – what and when?



Starting proposal Metabolite quantification – what when and how (in vivo)?



Afternoon breakout

➤ Metabolite profiling subteam

- Move quickly in back of room
- Discuss starting proposal, consider early morning presentation, peer reviewed literature and your experience
- 5 moderators (Geert, Graeme, Martha, Stefan and Claude) will capture thoughts and plan forward and present @ 15.00

➤ Metabolite quantification subteam

- Stay or come a bit closer
- Discuss starting proposal, consider early morning presentation, peer reviewed literature and your experience
- 4 moderators (John, Steve, Stuart and Philip) will capture thoughts and plan forward and present @ 15.30

- Plan forward will become the basis of updated EBF recommendation on Metabolite Profiling and Quantification to be published in 2016

