

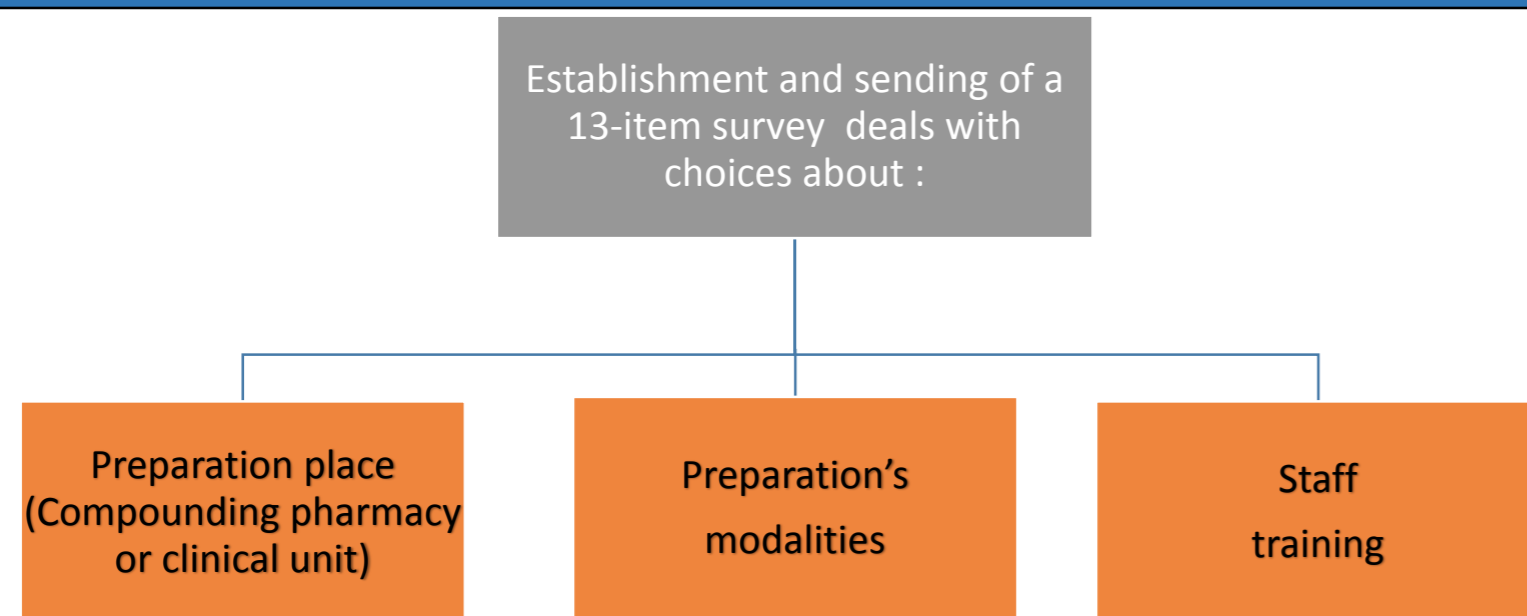
## 1- Context

On 08/27/2018 Patisiran got market access authorization for the treatment of hereditary transthyretin amyloidosis. This is the first small interfering double-stranded RNA drug. Thereby, Patisiran need to be reconstituted with specific steps, which are a 0.45µm filtration and a dilution for a total volume of 200mL. As a result, we can wonder if the product has to be reconstituted in the preparatory of the hospital or in clinical unit by nurses. Arguments, whether pharmaco-technicals, economics or practical are debated.

## 2 - Aim

- Compare professional reconstitution practices with the SPC in 19 centers.

## 3 - Material and Method



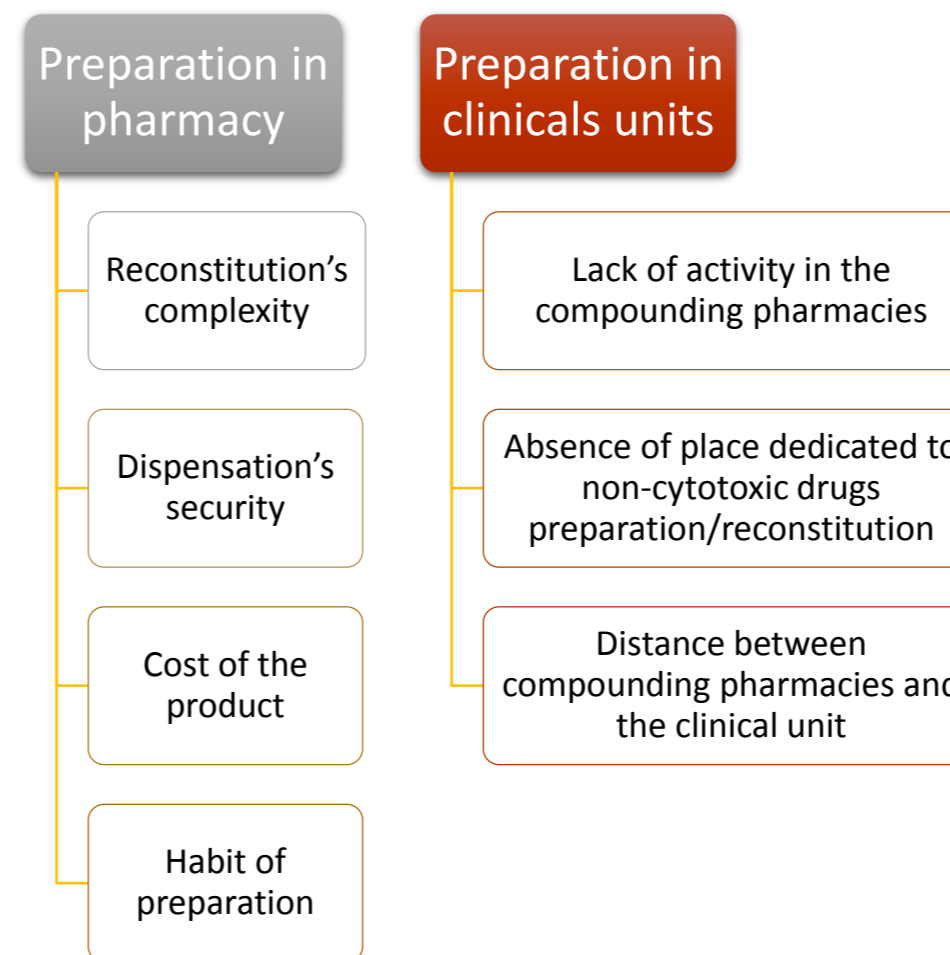
## 5 - Discussion

Because of the complexity of the reconstitution the majority of hospitals have opted to reconstitute the Patisiran in the hospital compounding pharmacies. However, 2 of them wish to delegate the reconstitution to the nurses in clinical unit after a training period.

In our center, clinical unit nurses reconstitute. They filter when injecting the Patisiran in the infusion bag with 0.45µm filter beforehand moisten. Thanks to that the retention volume is decreased from 1mL to 0.1ml, allowing to limit the losses. We don't respect the dilution in 200mL as it is written in the SCP because it's due to American practice and the fact that they have commercial 200mL infusion bags but no to pharmacological and stabilities arguments.

## 4 - Results

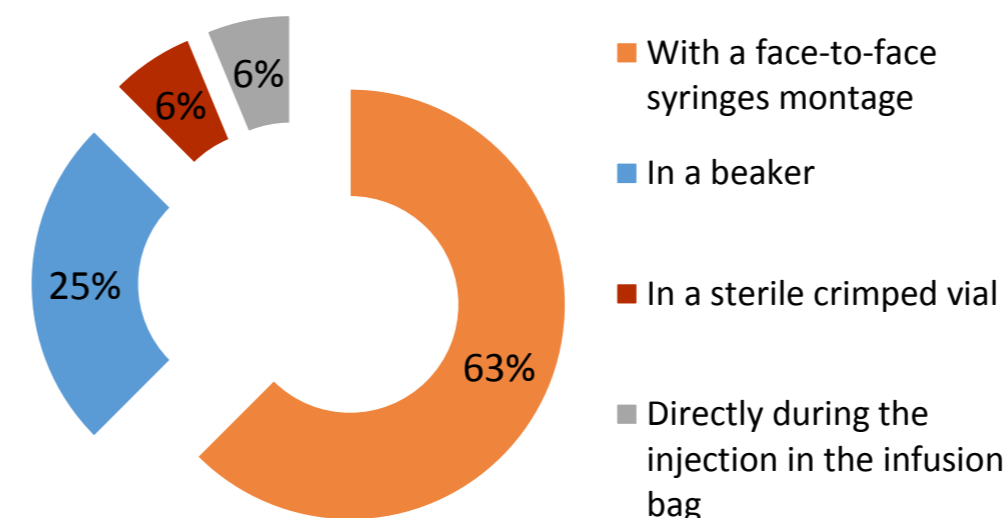
### Arguments in favor of :



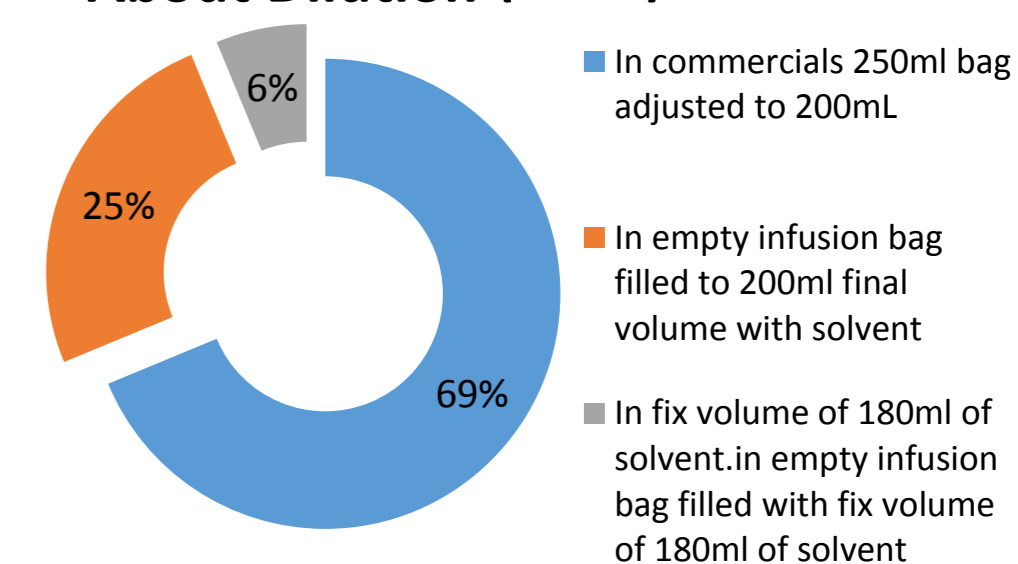
| Number of pharmacies which prepare | Number of clinicals units which prepare |
|------------------------------------|---|
| 11                                 | 5                                       |

- 16 centers out of 19 have answered
- 69% reconstitute in compounding pharmacy
- 91% of them have reported as arguments : the reconstitution's complexity (n=10)
- Mains arguments in favor of a preparation in clinicals units are :
  - The lack of activity in the compounding pharmacy (n=2)
  - Preparation improper if performed in cytotoxic drug reconstitution isolator (n=3)
- In this context, nurses have been trained on this act and a procedure of reconstitution has been joined with vials

### About Filtration (n=16)



### About Dilution (n=16)



- Main filtration methods are these with a face-to-face syringes montage or in a beaker
- 2 centers do not have an aseptic process : nurses filter patisiran in beaker on working area in clinicals units .
- 15 centers follow SPC instructions and dilute in 200mL
- In our center, we filter directly during the injection in the infusion bag filled with 180mL of solvent.**