

## Kocisko 1994 first cell-free conversion

recPrPC incubated with purified, partially GdnHCl-denatured PrPSc

sonication 

 shaking

## Saborio 2001 **PMCA**

infected brain homogenate + uninfected brain homogenate + incubation/sonication cycles PK resistance readout

## Zou & Cashman 2002

acid, detergents and shaking can cause PrPC to spontaneously adopt PrPSc-like properties

## Castilla 2005

demonstration that PMCA can amplify infectivity indefinitely

Saa 2006  
refinement of PMCA protocol

Legname 2004  
first synthetic prions infectious to transgenic mice

## Rogers 1965

Thioflavin T can detect amyloid

Colby 2007  
amyloid seeding assay (ASA)  
ThT to detect prion conversion

## Deleault 2007

**minimal co-factors and synthetic prions**  
PMCA on PrPC + lipids + RNA yields first synthetic prion infectious to wild-type mice

## Atarashi 2007

**“rPrP-PMCA”**  
recPrPC + infected brain homogenate

## Atarashi 2008

### **QuIC**

whole infected brainhomogenate + purified recPrPC subjected to shaking, detected with thioflavin T

## Wang 2010

**synthetic recPrP prions**  
PMCA on recPrP + RNA + POPG first synthetic prion made from recPrP infectious to wild-type mice

## Atarashi 2011

### **RT-QuIC**

endpoint titration using QuIC on serial dilutions

## McGuire 2012

validation of RT-QuIC as diagnostic test for CSF

## Shi 2013

### **qRT-QuIC**

alternate readout without using serial dilutions



**clinical use**