FBN1 mutation in our family.

- Personal history
- DNA discovery
- Distribution of the mutation.
- Reactions and Concerns.

Personal Medical History

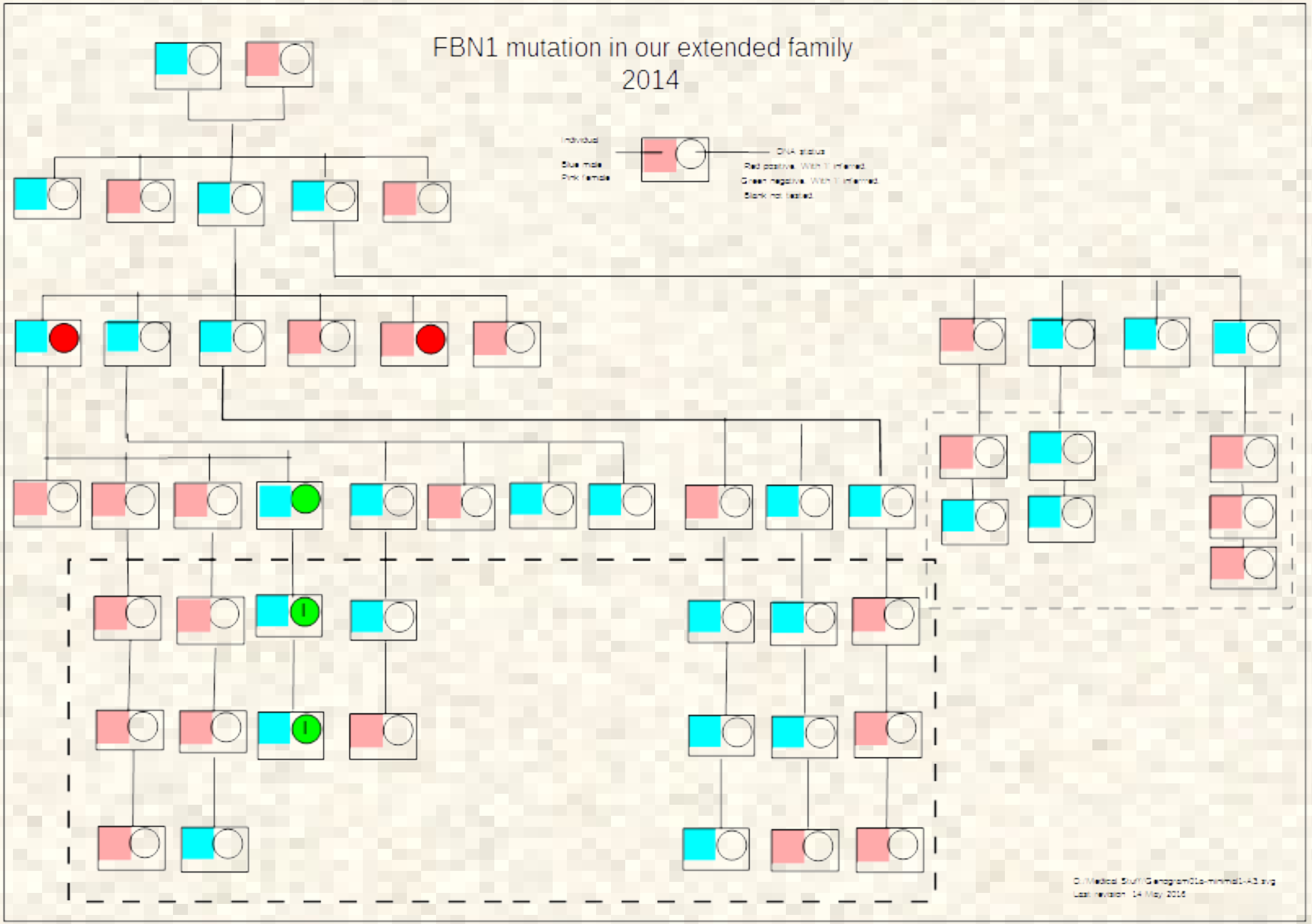
- 09/1937 born in Maastricht, Holland. (78 now)
- 1995 Abdominal Aortic Aneurysm repair.
- 2006 My sister suffers a dissection
- 2007 Left iliac aneurysm embolised.
- 04/2008 Right popliteal aneurysm repair
- 06/2008 Left popliteal anneurysm repair
- 01/2012 Right femoral aneurysm repair
- 05/2012 Became infected. Repair redone. Excision of infected seroma.
- 10/2013 Left femoral aneurysm repair.
- 01/2014 Right popliteal repaired with Viaban Stent.
- 03/2014 DNA test
- 01/2015 Aortic root and valve repaired Bental's procedure + Pacemaker..
- 10/2015 Acute thrombosis of Viaban Stent.
- On going surveillance of juxtarenal aneurysm

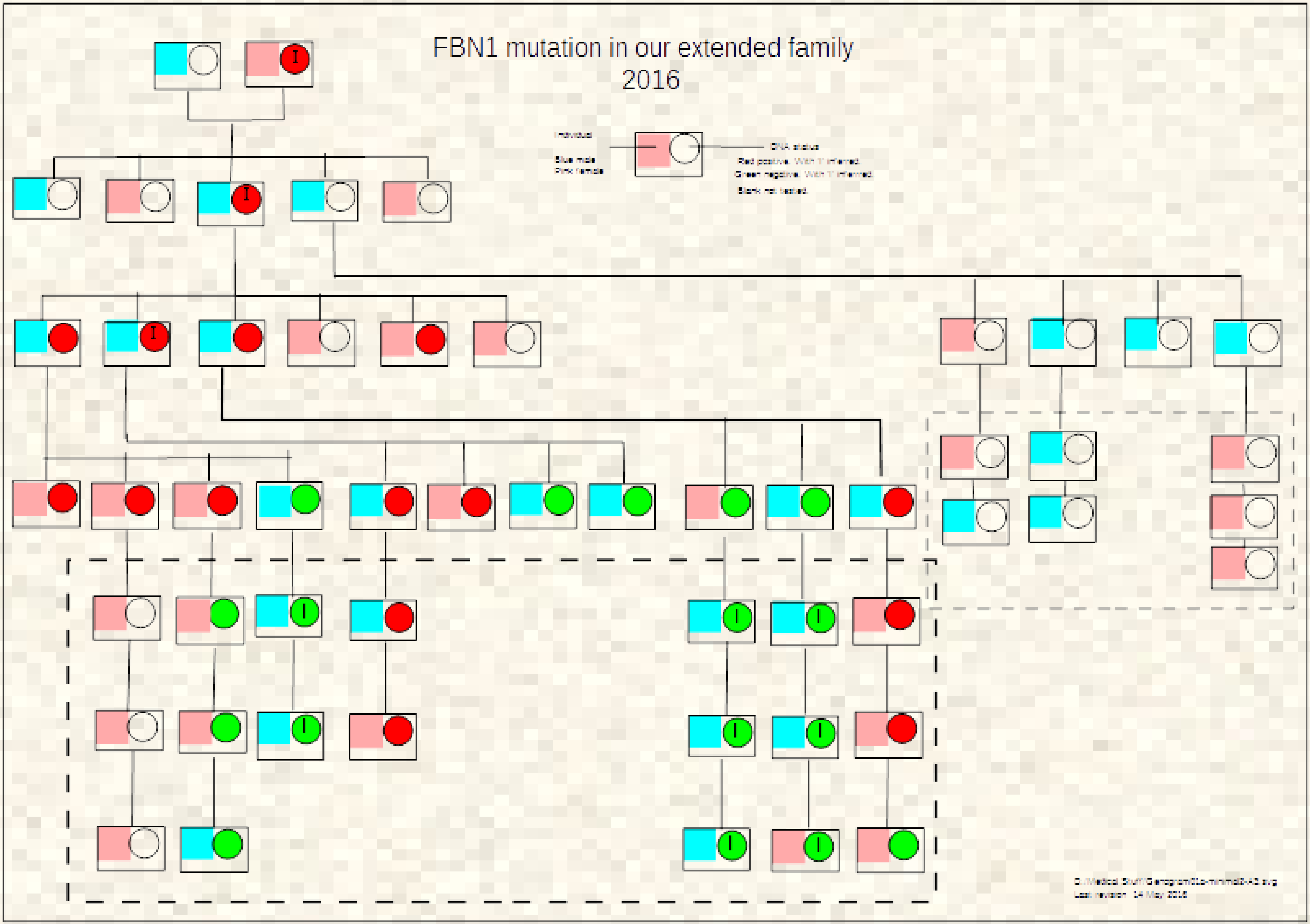
New FBN1 pathogenic variant.

From the report of the Collagen Diagnostic lab, University of Washington. (March 2014).

- No mutation in COL3A1, TGFBr1,TGFBR2, ACTA2, SMAD3, TGB2. (The usual suspects)
- Identification of a likely pathogenic variant in one allele of the FBN1 Gene, that encodes the protein fibrillin 1.
- The variant is predicted to result in substitution of arginine (Arg) by Cystine (Cys) at proteine position 2360.
- Not reported in UMD-FBN1 mutations database, dbSNP, ClinVar or Exome Variant Server.

Taken altogether, the available information supports that this is a likely pathogenic variant that explains the vascular phenotype in the patient.


FBN1 mutation in our extended family
2014
Individual
Blue male
Pink female
DNA status
Red positive. With TT inferred.
Green negative. With TT inferred.
Blank not tested.

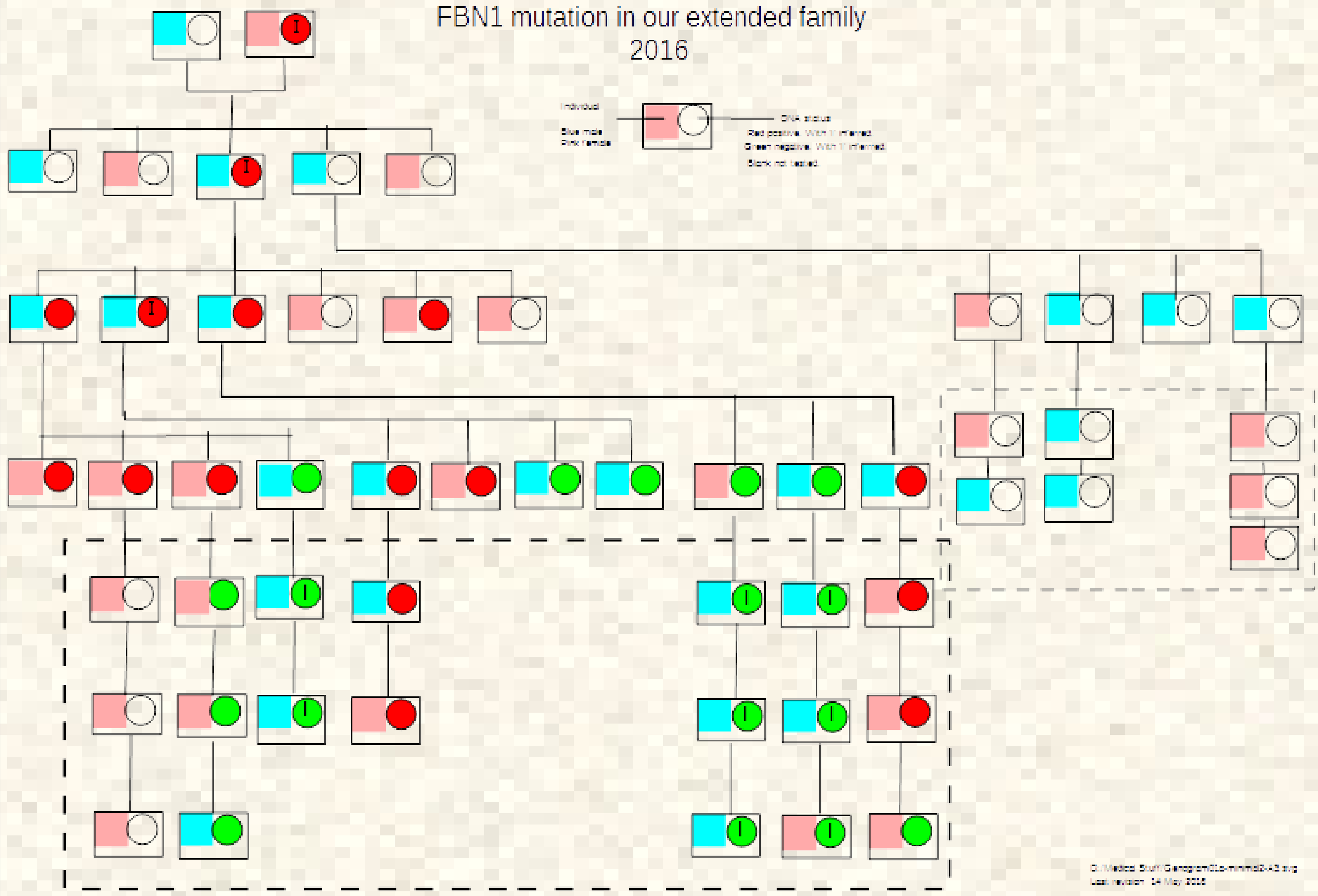

FBN1 mutation in our extended family
2016
Individual
Blue male
Pink female
DNA status
Red positive. With 'I' inferred.
Green negative. With 'I' inferred.
Blank not tested.
Last revision 14 May 2016

Summary

(excluding descendants of Ae4)

- Of my father's 36 descendants, 15 have been identified as carrying the mutation. (42%)
- If one caries the mutation one is highly likely to be tall, but one can be tall without the mutation. (4 cases)
- Already 6 members have been operated for enlarged aortas: M (77), R (71), C(61), X(52), B (44), R (24).
- Enlarged aortic root is emerging as a common feature.
- Although there is a small sprinkling of Marfan features there is no consistency or pattern.

FBN1 mutation in our extended family
2016




GrandFather
GrandMother
Wim
Francois