

Respi Viruses & Children

How many, how much ?

Hans De Beenhouwer



Introduction

- Acute respiratory tract infection (RTI) are responsible for 75% of all consults for acute complaints in the world
- Every child of < 7 y has 3 to 8 times/year an acute upper respiratory infection (URTI).
- In USA children < 2 y: between 12 to 32 $\cdot 10^9$ episodes/y
- URTI are linked to asthmatic attacks, otitis media, lower respiratory infections (LRTI), ...



Clinical

Spectrum of clinical pictures going from common cold (rhinorrhea) to life threatening pneumonia

- ✓ Role environment, socio-economic ...
- ✓ Role individu (underlying diseases ...)
- ✓ Role of the infecting agent



Respi Viruses & Children

How many, how much ?



Etiology : how many ?

- number of 'agents' is growing fast
- Huge amount of viruses which have been implicated in RTI (in children) !
- "Old" chaps
- "New" chaps



Etiology : how many ?

"Old Chaps"

- Influenza A
- Influenza B
- RSV
- Parainfluenza 1-3
- Adenovirus
- Rhinovirus

"New Chaps"

- hMPV
- Parainfluenza 4
- CoronaV
- BocaV
- ParvoV 4 & 5
- Polyoma KI, WU
- ...



Detection, why ?

Better diagnostic tools are important:

→ patient:

- ✓ correct therapy (influenza)
- ✓ avoid unnecessary antibiotics, unnecessary examinations or hospitalizations
- ✓ correct prognosis ⇒ comforting patients
⇒ comforting parents



Why detection ?

→ doctors:

- Diagnostic reflexes in respiratory infections are underdeveloped, underused because results too slow, sampling problems, sensitivity problems, ...
 - ⇒ no interest in correct diagnosis
 - ⇒ hazy therapeutic policies
- Rapid diagnosis ⇒ Learning curve
 - ⇒ Better care



How to detect ?

A diagnostic test should be:

- ✓ rapid (same day - results)
- ✓ sensitive and specific
- ✓ simple sampling method and transport
- ✓ spectrum as large as possible / necessary
 - ⇒ Molecular testing
- ✓ easy
- ✓ cheap !



Large spectrum: Multi - detection

- Detection of Multiple pathogens :
 - ✓ Hexaplex +
 - ✓ Prodesse (7 viruses)
 - ✓ X-Tag (19 viruses)
 - ✓ MultiCode-Plx (17 viruses)
 - ✓ Respifinder (viruses + bacteria)
 - ✓ Res.plex I & II (viruses + bacteria)
 - ✓ Multiplex MassTag (viruses + bacteria)
 - ✓ Micro arrays
 - ✓ ...



Bilulu examined !

Performance of Respifinder Smart 22 for fast diagnosis of respiratory tract infections

EMMD
12-14/10/2011

RAYMAEKERS M.^{1*}, CARTUYVELS R.^{1*}, DUSON G.^{2*}, FRANS J.^{2*}, LAMBINI S.^{3*}, VAN DEN ABEELE A.^{3*}, VANKEERBERGHEN A.^{4*}, DIERICKX K.^{4*}, DE BEENHOUWER H.^{4*}

* on behalf of the Bilulu Study Group

¹ Jessa Ziekenhuis, campus Virga Jesse, Hasselt, ² Imelda Ziekenhuis, Bonheiden, ³ AZ Sint-Lucas, Gent, ⁴ Onze-Lieve-Vrouw Ziekenhuis, Aalst

Corresponding author: Stadsomvaart 11, 3500 Hasselt, BELGIUM Phone: +32-11 30 97 02 Fax: +32-11 30 97 50
Email: marijke.raymaekers@jessazh.be



OBJECTIVES

More than 20 viruses have been shown to cause respiratory tract infections. Identification of the etiological agent is useful to provide – or avoid – specific therapy to take the necessary hospital hygiene measures and for epidemic control. Nucleic acid amplification techniques allow broad spectrum detection of respiratory tract infections. (Smart), a multiplex analysis detecting 16 RNA viruses, 2 DNA viruses and 2

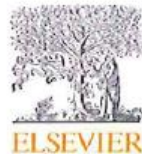
METHODS

Journal of Clinical Virology 52 (2011) 314–316

Contents lists available at SciVerse ScienceDirect

Journal of Clinical Virology

journal homepage: www.elsevier.com/locate/jcv



Timely diagnosis of respiratory tract infections: Evaluation of the performance of the Respifinder assay compared to the xTAG Respiratory Viral Panel assay

M. Raymaekers^{a,*}, B. de Rijke^b, I. Pauli^a, A.-M. Van den Abeele^b, R. Cartuyvels^a

^a Clinical Laboratory, Jessa Hospital, Site Virga Jesse, Stadsomvaart 11, B-3500 Hasselt, Belgium

^b Clinical Laboratory, AZ Sint-Lucas-Volkskliniek, Groenebriel 9000, Gent, Belgium



Why detection ?

Better diagnostic tools are important:

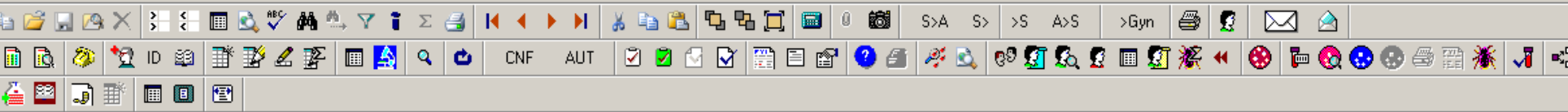
- ➔ Patient: optimal care
 - ✓ correct therapy (influenza)
 - ✓ avoid unnecessary antibiotics, unnecessary examinations or hospitalizations
 - ✓ correct prognosis ⇒ comforting patients/parents
- ➔ Doctors: “learning curve”
- ➔ Hospital: cohorting, infection control



Toolbar with various icons for file operations, navigation, and application settings.

A_Mollol_INF 07/12/10 11:25

		PCR HMPV	PCR HMPV CT	PCR RSV	PCR RSV CT	PCR INFA	PCR INFA CT	PCR INFB	PCR INFB CT
1)	RSV_POS-02031				29.90				
2)	HMPV_POS-0203		27.34						
3)	RSV_cDNA-1810				26.10				
4)	1631964 3697776 01					<NG	[]	<NG	[]
5)	2060034 3698068 01 *	<NG	[]	POS	19.34	<NG	[]	<NG	[]
6)	2058647 3698298 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
7)	1608982 3698350 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
8)	2047038 3698375 01	<NG	[]	POS	30.76	<NG	[]	<NG	[]
9)	2021298 3698482 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
10)	2028876 3698824 01	<NG	[]	POS	20.20	<NG	[]	<NG	[]
11)	2053335 3699037 01	<NG	[]	POS	21.02	<NG	[]	<NG	[]
12)	2055989 3699214 01	<NG	[]		38.91	<NG	[]	<NG	[]
13)	2060035 3699215 01	<NG	[]	POS	22.04	<NG	[]	<NG	[]
14)	1609461 3699543 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
15)	2059946 3699548 01	POS	25.12	<NG	[]	<NG	[]	<NG	[]
16)	2060146 3699584 01 *	<NG	[]	POS	28.31	<NG	[]	<NG	[]
17)	2060156 3699634 01 *	<NG	[]	POS	20.87	<NG	[]	<NG	[]
18)	2060152 3699636 01 *	<NG	[]	POS	22.32	<NG	[]	<NG	[]
19)	2060151 3699638 01 *	<NG	[]	POS	16.92	<NG	[]	<NG	[]
20)	1631965 3699658 01	<NG	[]	POS	20.99	<NG	[]	<NG	[]
21)	2060157 3699659 01 *	<NG	[]		40.46	<NG	[]	<NG	[]
22)	2060148 3699686 01 *	<NG	[]	POS	31.42	<NG	[]	<NG	[]
23)	2055253 3699690 01	<NG	[]	POS	28.90	<NG	[]	<NG	[]
24)	1633633 3699694 01	<NG	[]	POS	21.39	<NG	[]	<NG	[]
25)	2009880 3699696 01	<NG	[]	POS	19.70	<NG	[]	<NG	[]

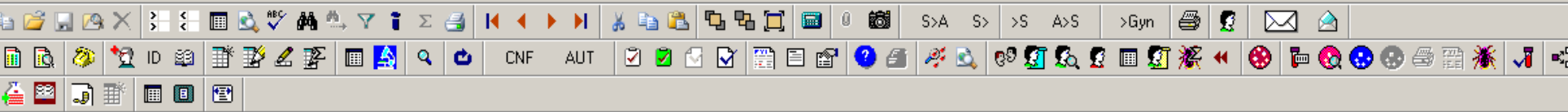


A. M... 07/12/10 11:25

		PCR HMPV	PCR HMPV CT	PCR RSV	PCR RSV CT	PCR INFA	PCR INFA CT	PCR INFB	PCR INFB CT
1)	RSV_POS-02031				29.90				
2)	HMPV_POS-0203		27.34						
3)	RSV_cDNA-1810				26.10				
4)	1631964 3697776 01					{<NG}	[]	{<NG}	[]
5)	2060034 3698068 01 *	{<NG}	[]	POS	19.34	{<NG}	[]	{<NG}	[]
6)	2058647 3698298 01	{<NG}	[]	{<NG}	[]	{<NG}	[]	{<NG}	[]
7)	1608982 3698350 01	{<NG}	[]	{<NG}	[]				
8)	2047038 3698375 01	{<NG}	[]	POS	30.76				
9)	2021298 3698482 01	{<NG}	[]	{<NG}	[]				
10)	2028876 3698824 01	{<NG}	[]	POS	20.20				
11)	2053335 3699037 01	{<NG}	[]	POS	21.02				
12)	2055989 3699214 01	{<NG}	[]		38.91				
13)	2060035 3699215 01	{<NG}	[]	POS	22.04				
14)	1609461 3699543 01	{<NG}	[]	{<NG}	[]	{<NG}	[]	{<NG}	[]
15)	2059946 3699548 01	POS	25.12	{<NG}	[]				
16)	2060146 3699584 01 *	{<NG}	[]	POS	28.91				
17)	2060156 3699634 01 *	{<NG}	[]	POS	20.97				
18)	2060152 3699636 01 *	{<NG}	[]	POS	22.32				
19)	2060151 3699638 01 *	{<NG}	[]	POS	16.92				
20)	1631965 3699658 01	{<NG}	[]	POS	20.99				
21)	2060157 3699659 01 *	{<NG}	[]		40.46				
22)	2060148 3699686 01 *	{<NG}	[]	POS	31.42				
23)	2055253 3699690 01	{<NG}	[]	POS	28.90				
24)	1633633 3699694 01	{<NG}	[]	POS	21.39				
25)	2009880 3699696 01	{<NG}	[]	POS	19.70				

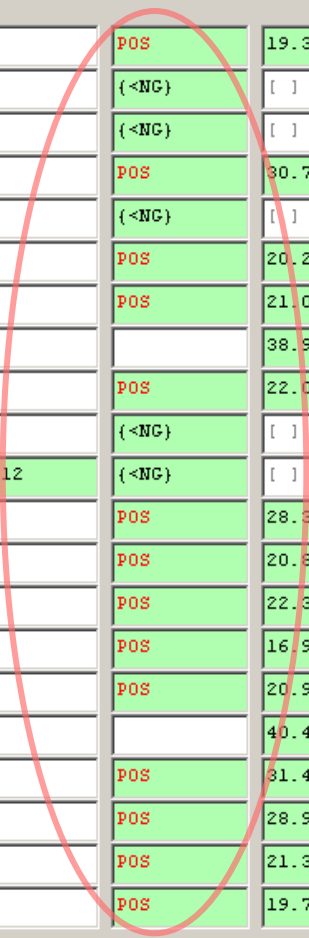
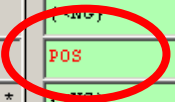
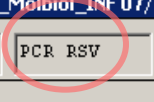
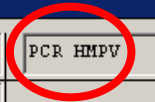
{<NG}	[]	{<NG}	[]
{<NG}	[]	{<NG}	[]

{<NG}	[]	{<NG}	[]
-------	-----	-------	-----



A_Molbiol_INF 07/12/10 11:25

		PCR HMPV	PCR HMPV CT	PCR RSV	PCR RSV CT	PCR INFA	PCR INFA CT	PCR INFB	PCR INFB CT
1)	RSV_POS-02031				29.90				
2)	HMPV_POS-0203		27.34						
3)	RSV_cDNA-1810				26.10				
4)	1631964 3697776 01					<NG	[]	<NG	[]
5)	2060034 3698068 01 *	<NG	[]	POS	19.34	<NG	[]	<NG	[]
6)	2058647 3698298 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
7)	1608982 3698350 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
8)	2047038 3698375 01	<NG	[]	POS	30.76	<NG	[]	<NG	[]
9)	2021298 3698482 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
10)	2028876 3698824 01	<NG	[]	POS	20.20	<NG	[]	<NG	[]
11)	2053335 3699037 01	<NG	[]	POS	21.02	<NG	[]	<NG	[]
12)	2055989 3699214 01	<NG	[]		38.91	<NG	[]	<NG	[]
13)	2060035 3699215 01	<NG	[]	POS	22.04	<NG	[]	<NG	[]
14)	1609461 3699543 01	<NG	[]	<NG	[]	<NG	[]	<NG	[]
15)	2059946 3699548 01	POS	25.12	<NG	[]	<NG	[]	<NG	[]
16)	2060146 3699584 01 *	<NG	[]	POS	28.31	<NG	[]	<NG	[]
17)	2060156 3699634 01 *	<NG	[]	POS	20.37	<NG	[]	<NG	[]
18)	2060152 3699636 01 *	<NG	[]	POS	22.32	<NG	[]	<NG	[]
19)	2060151 3699638 01 *	<NG	[]	POS	16.92	<NG	[]	<NG	[]
20)	1631965 3699658 01	<NG	[]	POS	20.99	<NG	[]	<NG	[]
21)	2060157 3699659 01 *	<NG	[]		40.46	<NG	[]	<NG	[]
22)	2060148 3699686 01 *	<NG	[]	POS	31.42	<NG	[]	<NG	[]
23)	2055253 3699690 01	<NG	[]	POS	28.90	<NG	[]	<NG	[]
24)	1633633 3699694 01	<NG	[]	POS	21.39	<NG	[]	<NG	[]
25)	2009880 3699696 01	<NG	[]	POS	19.70	<NG	[]	<NG	[]



<NG	[]	<NG	[]
<NG	[]	<NG	[]

<NG	[]	<NG	[]
-----	-----	-----	-----

Why detection ?

Better diagnostic tools are important:

- Patient: optimal care
- Doctors: "learning curve"
- Hospital: cohorting, infection control
- Government: cost-effectif



JOURNAL OF CLINICAL MICROBIOLOGY, Aug. 2000, p. 2824-2828
 0095-1137/00/\$04.00+0
 Copyright © 2000, American Society for Microbiology. All Rights Reserved.

Vol. 38, No. 8

Clinical and Financial Benefits of Rapid Detection of Respiratory Viruses: an Outcomes Study

JOAN BARENFANGER,^{1*} CHERYL DRAKE,¹ NIDIA LEON,² TINA MUELLER,¹
 AND TAMMY TROUTT¹

Laboratory Medicine, Memorial Medical Center,¹ and Internal Medicine Department, Southern Illinois University School of Medicine,² Springfield, Illinois 62781

Cost-Effectiveness of Rapid Diagnosis of Viral Respiratory Tract Infections in Pediatric Patients

PATRICK C. Y. WOO,¹ SUSAN S. CHIU,² WING-HONG SETO,¹ AND MALIK PEIRIS^{1*}

Departments of Microbiology¹ and Paediatrics,² Queen Mary Hospital, University of Hong Kong, Hong Kong

Received 12 December 1996/Returned for modification 7 February 1997/Accepted 5 March 1997

Rapid diagnosis of respiratory viral infections in children resulted in significantly reduced hospital stays, antibiotic use, and laboratory utilization compared with those of a matched group of patients from the previous year who were diagnosed by virus culture. We demonstrate that rapid diagnosis of respiratory infections in children is a cost-effective procedure.

The Cost of Community-managed Viral Respiratory Illnesses in a Cohort of Healthy Preschool-aged Children
 Stephen B. Lambert; Kelly M. Allen; Robert C. Carter; Terence M. Nolan



Why detection ?

Better diagnostic tools are important:

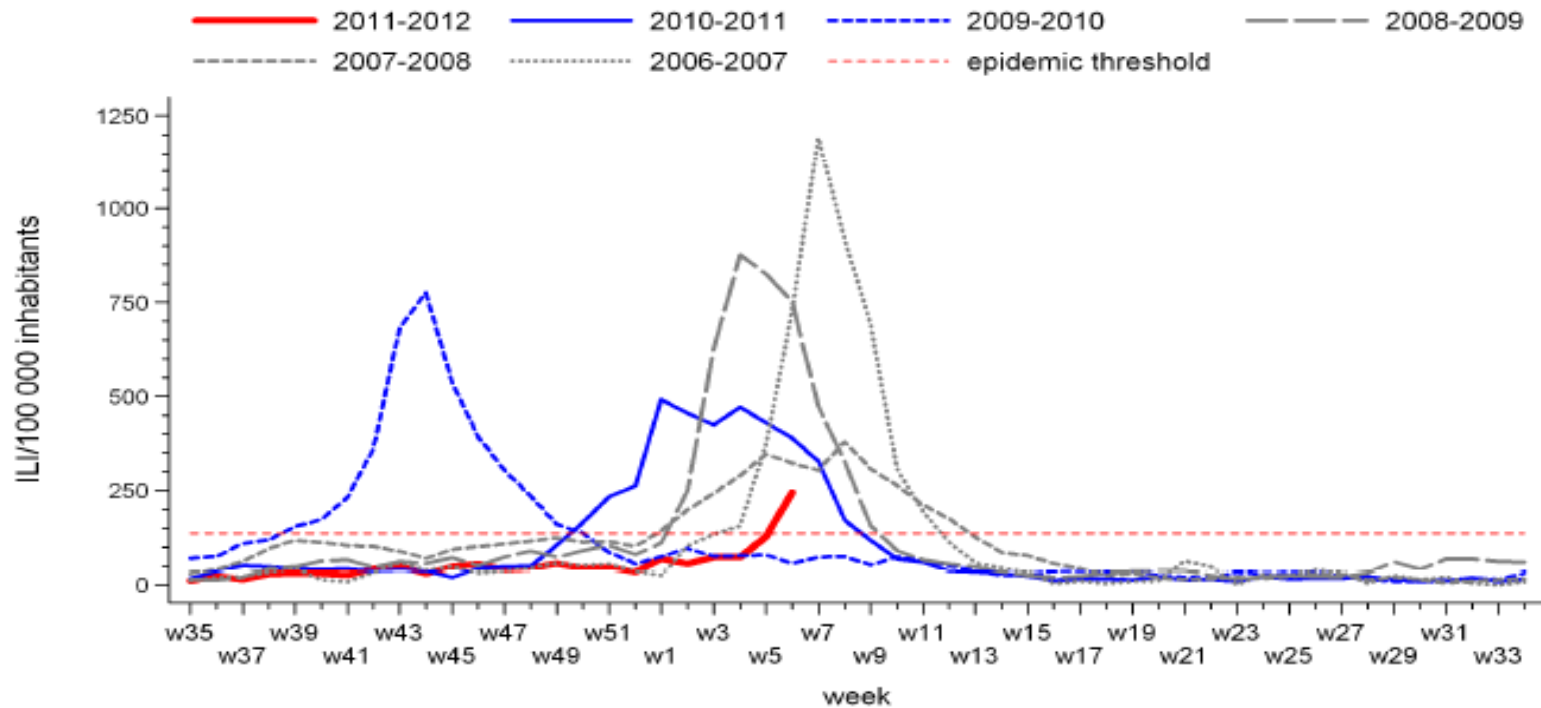
- Patient: optimal care
- Doctors: "learning curve"
- Hospital: cohorting, infection control
- Government: cost-effectif
- Public Health: more correct epidemiology
(epidemics, patterns, new viruses, variants, ...)



Epidemio WIV



GRIEP en acute luchtweginfecties (ALI) in België



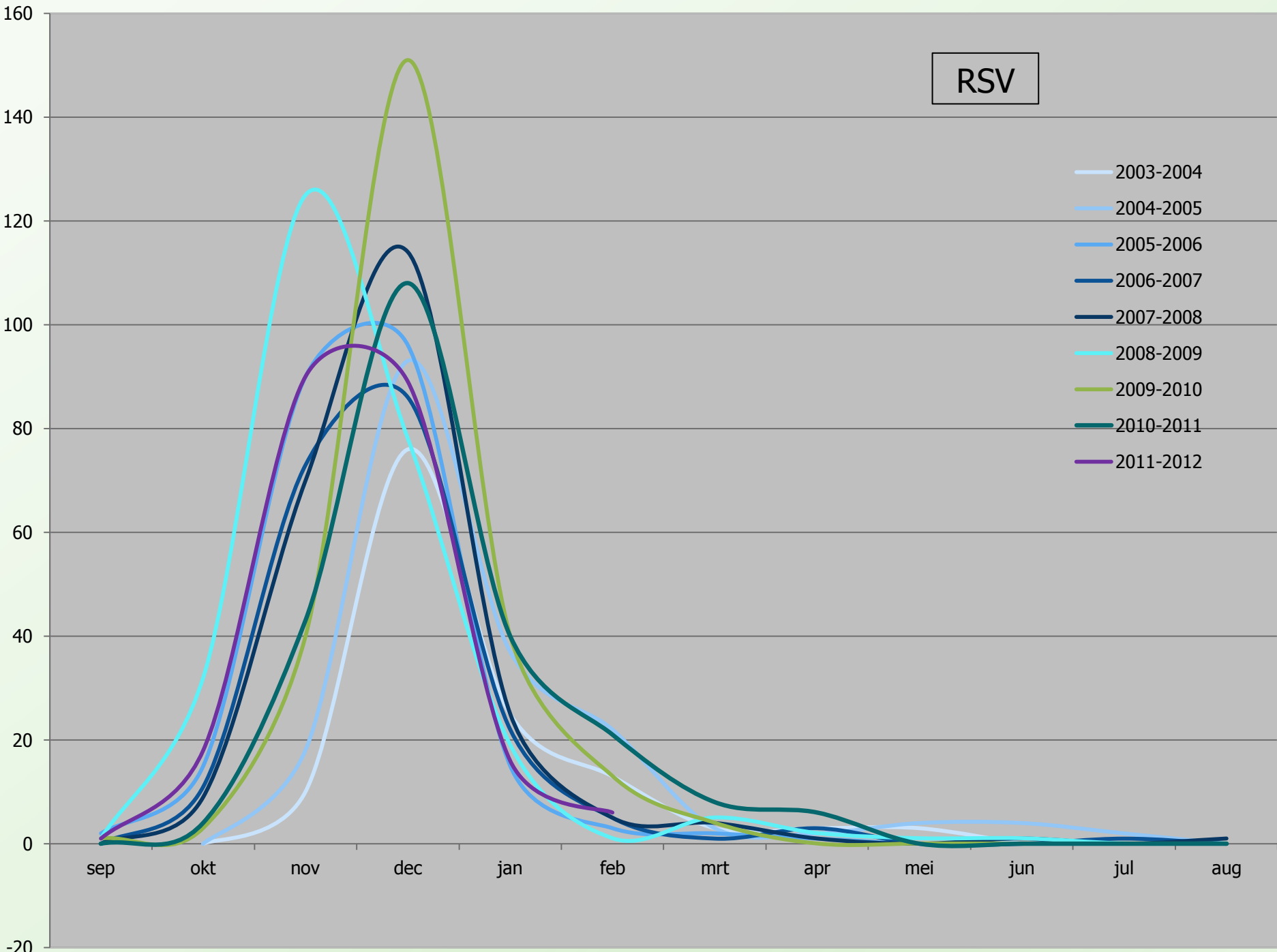
Epidemio Aalst

- Respiratory pathogen detection started in 2003 : 2 viruses (RSV & hMPV)
- Actually up to 15 viruses/bacteria



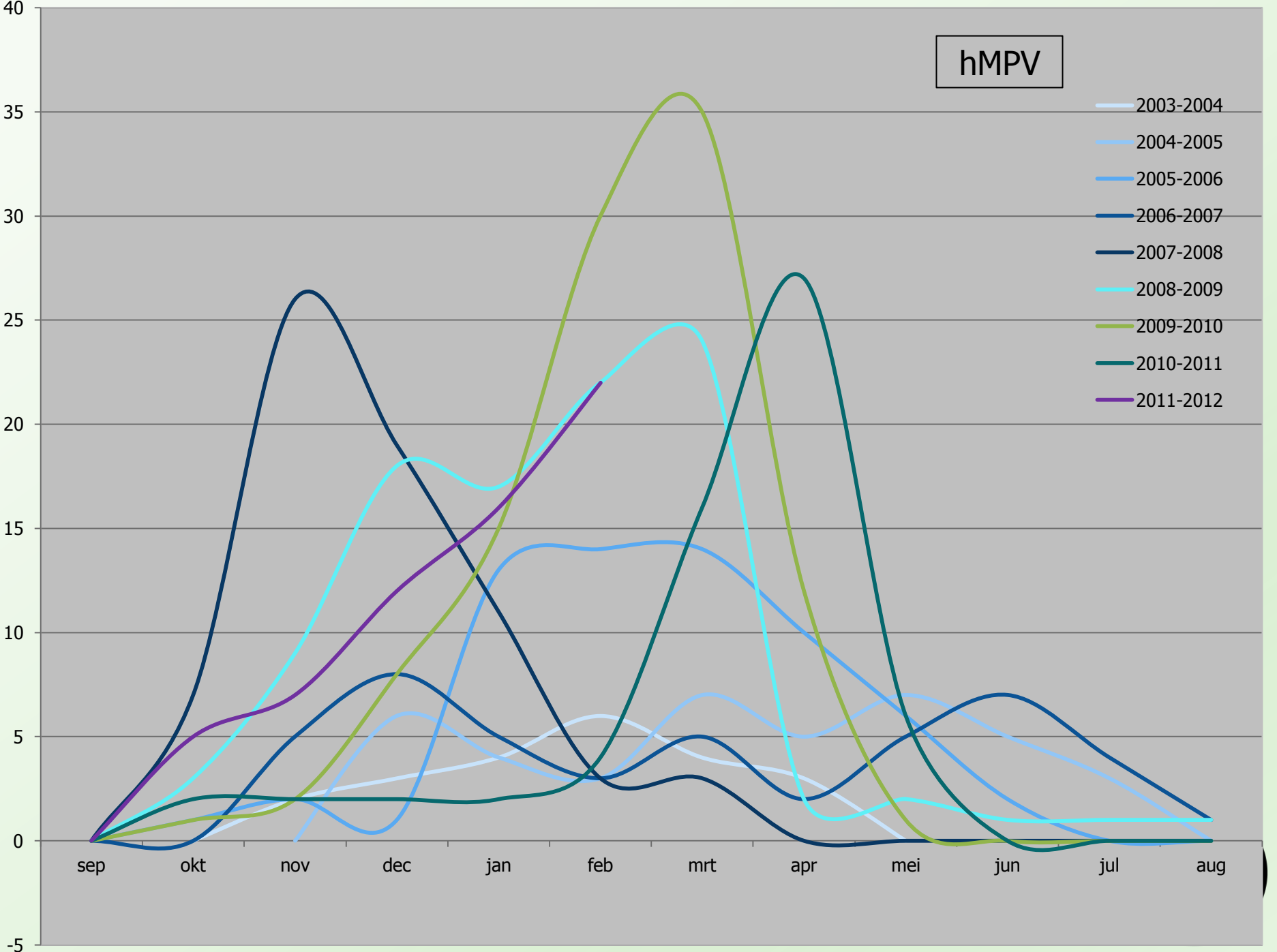
RSV

- 2003-2004
- 2004-2005
- 2005-2006
- 2006-2007
- 2007-2008
- 2008-2009
- 2009-2010
- 2010-2011
- 2011-2012



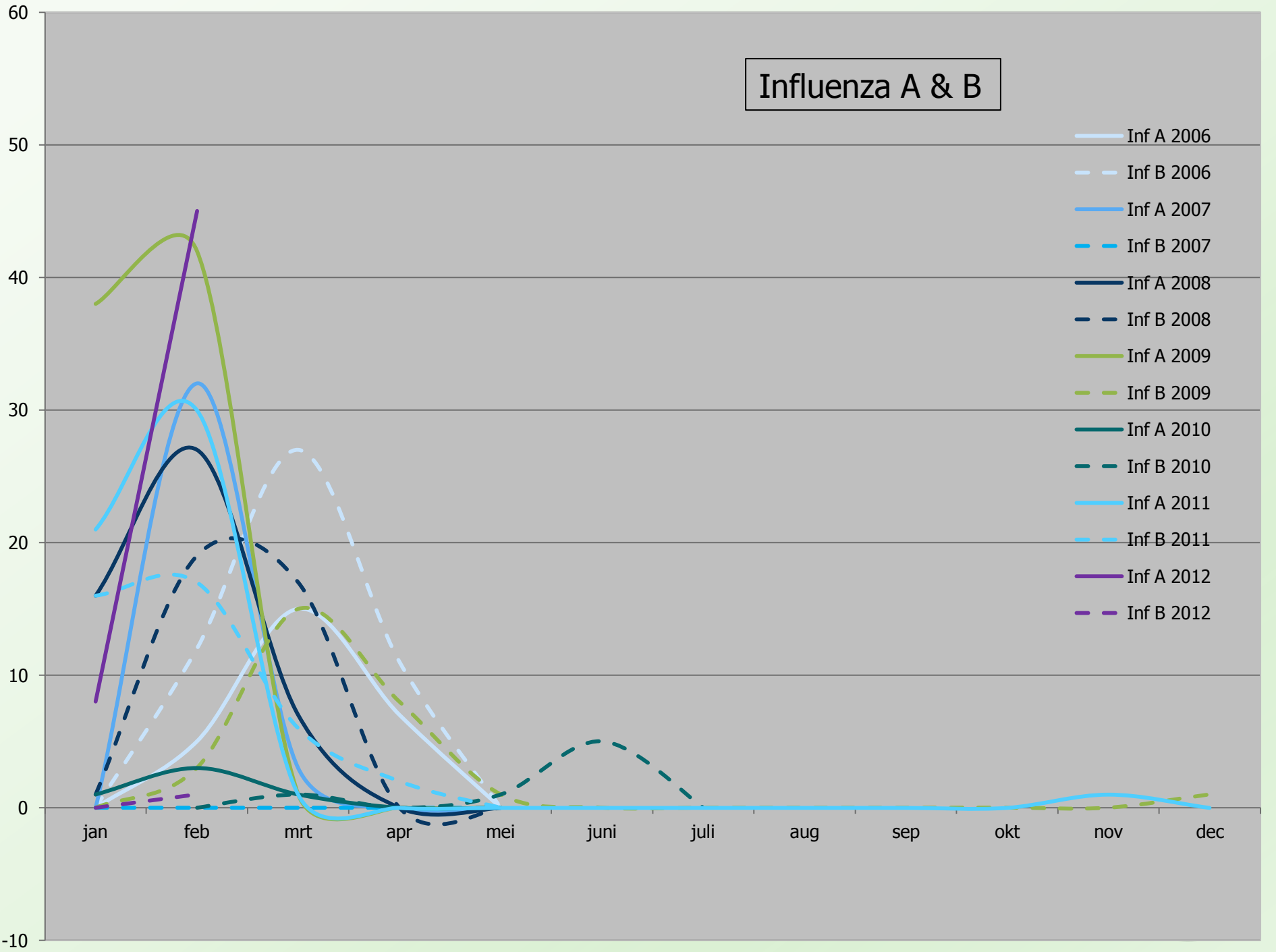
hMPV

- 2003-2004
- 2004-2005
- 2005-2006
- 2006-2007
- 2007-2008
- 2008-2009
- 2009-2010
- 2010-2011
- 2011-2012

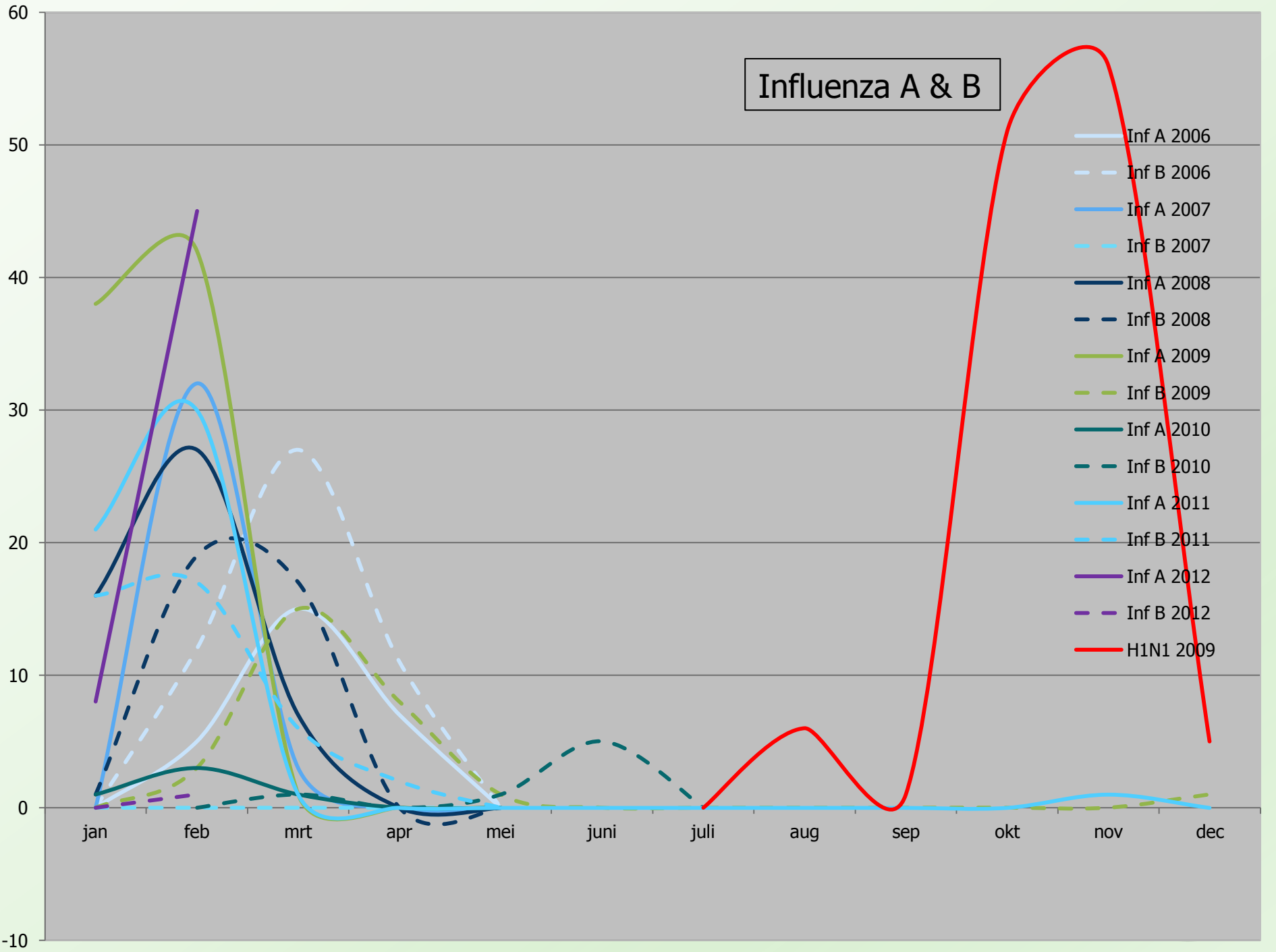


Influenza A & B

- Inf A 2006
- - Inf B 2006
- Inf A 2007
- - Inf B 2007
- Inf A 2008
- - Inf B 2008
- Inf A 2009
- - Inf B 2009
- Inf A 2010
- - Inf B 2010
- Inf A 2011
- - Inf B 2011
- Inf A 2012
- - Inf B 2012

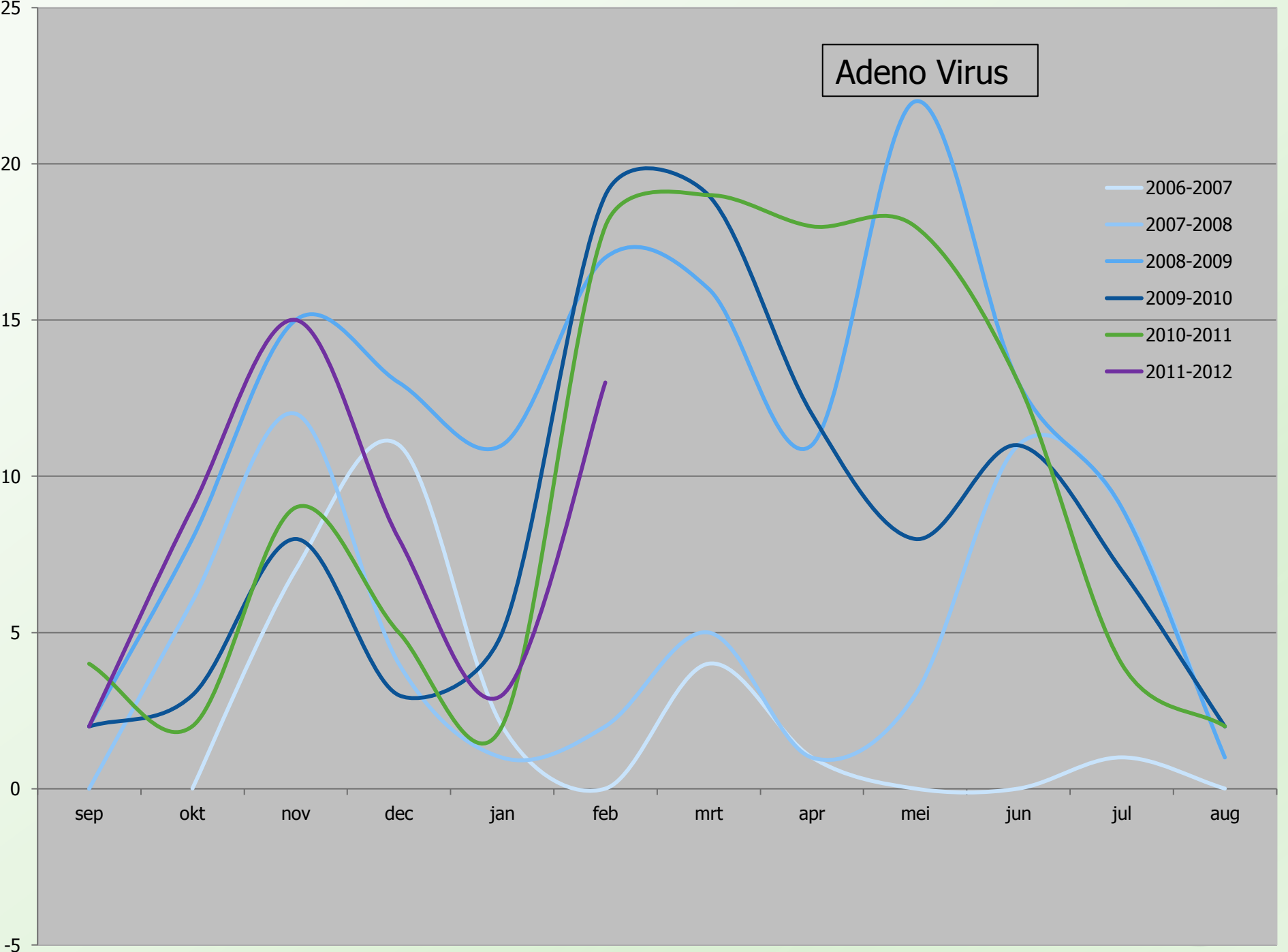


Influenza A & B



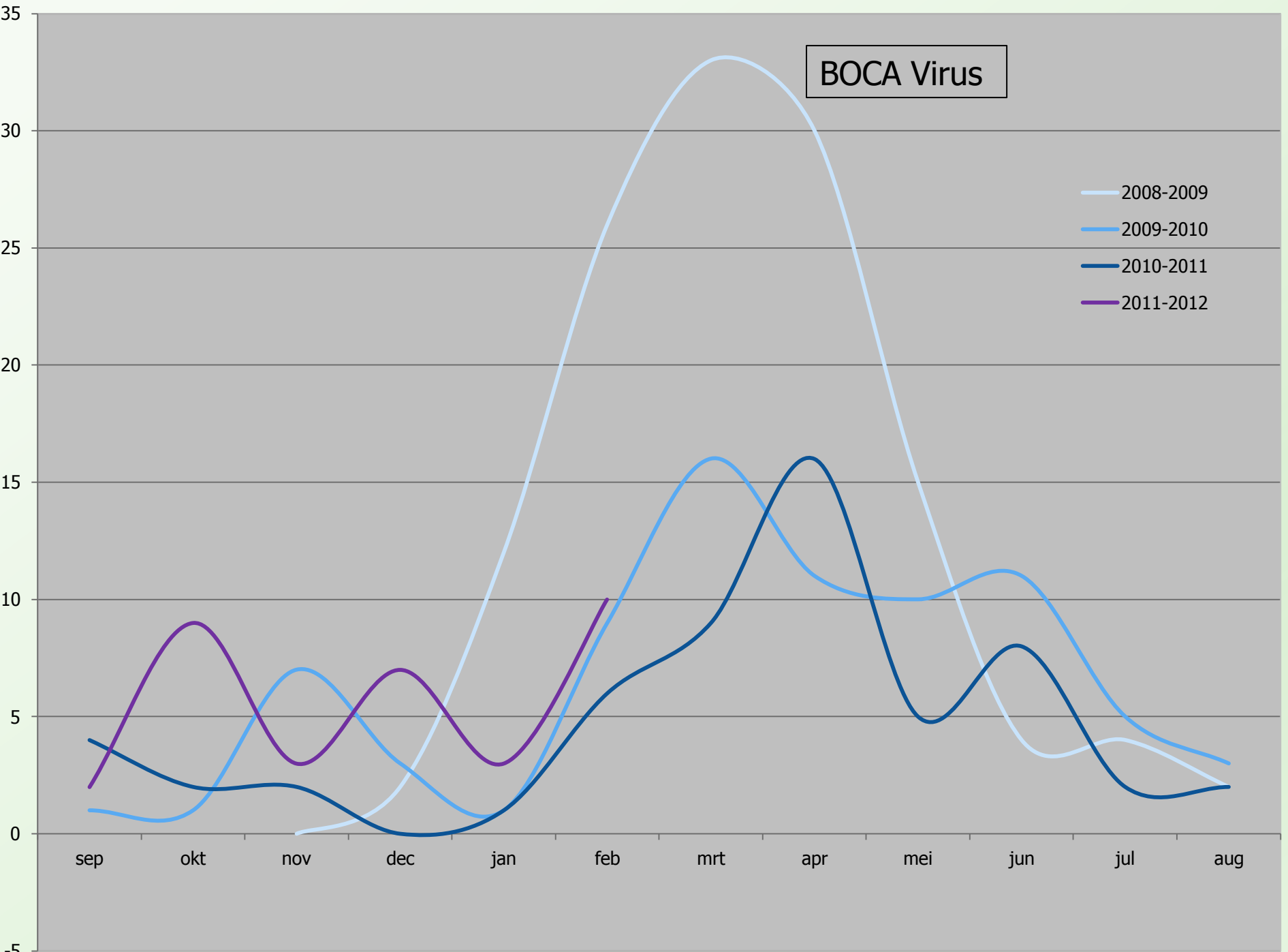
Adeno Virus

- 2006-2007
- 2007-2008
- 2008-2009
- 2009-2010
- 2010-2011
- 2011-2012



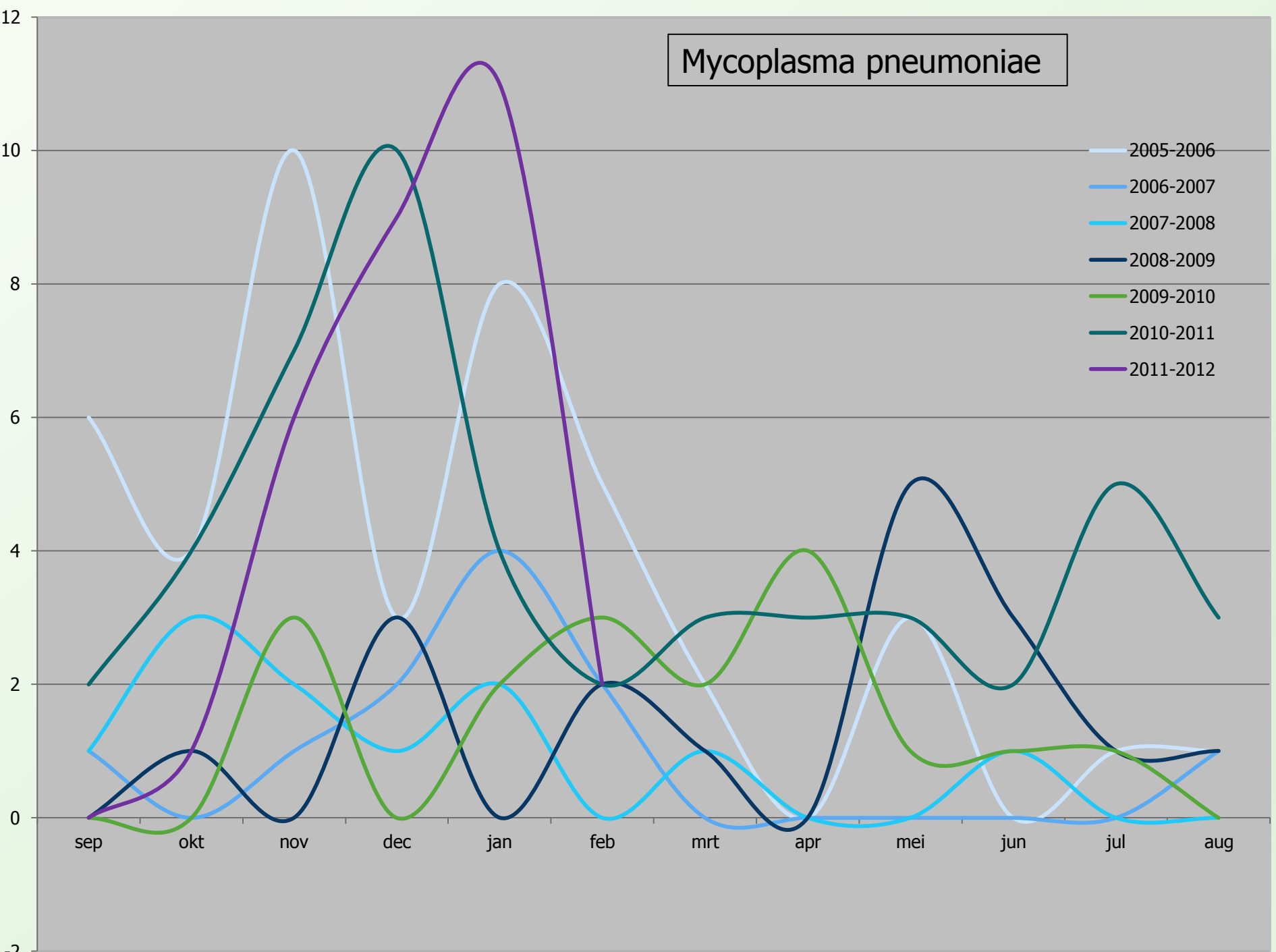
BOCA Virus

- 2008-2009
- 2009-2010
- 2010-2011
- 2011-2012



Mycoplasma pneumoniae

- 2005-2006
- 2006-2007
- 2007-2008
- 2008-2009
- 2009-2010
- 2010-2011
- 2011-2012



Etiology : how much

detection = infection ?



Detection = infection ?

- ✓ is detection of an agent = etiological agent of the present disease ?
- ✓ does viral colonisation exist in the respiratory tract?
- ✓ is persistency important? detectable ?

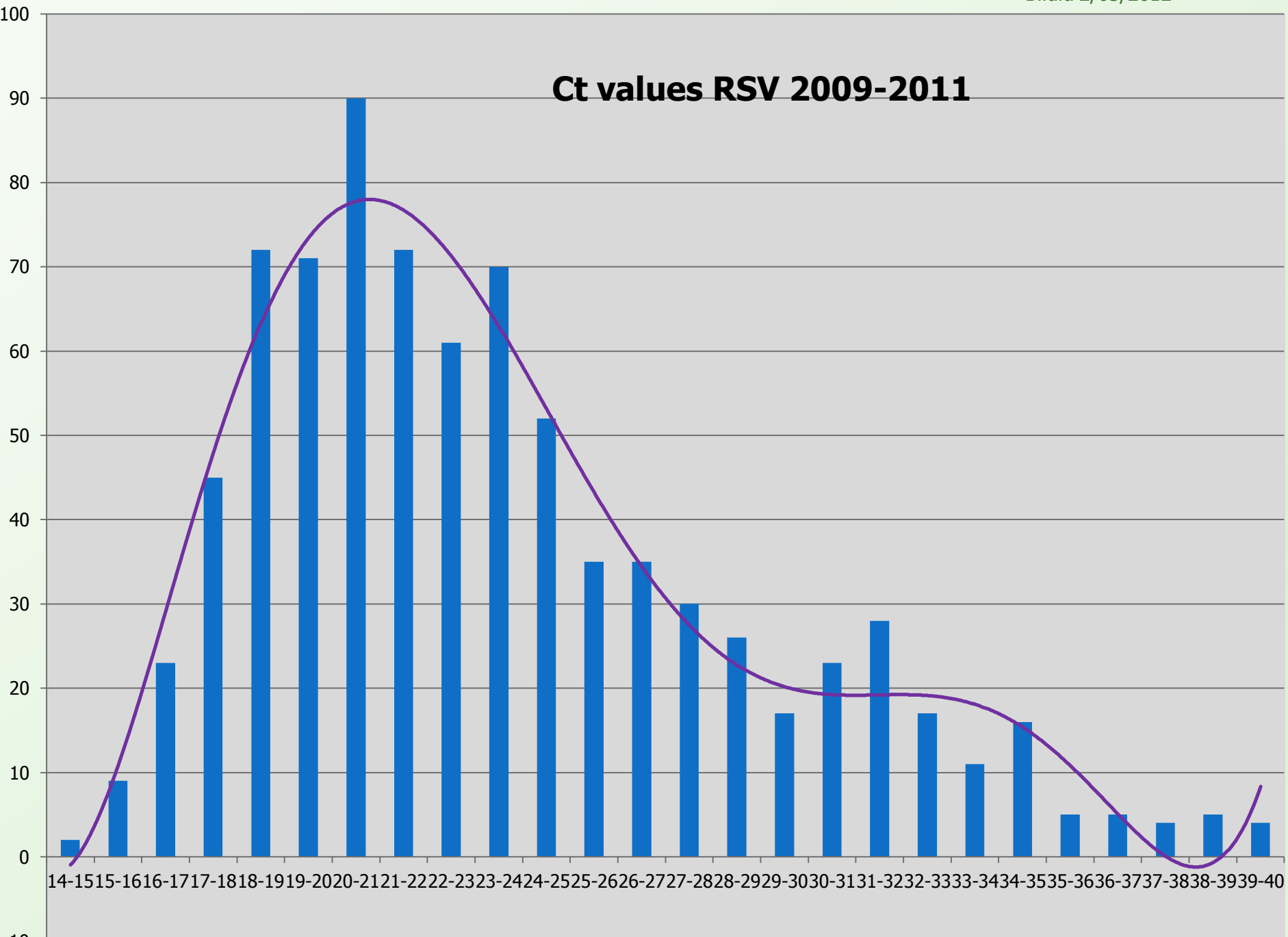
if yes (in analogy to bacterial infection):

⇒ it is diagnostically important to make the difference !

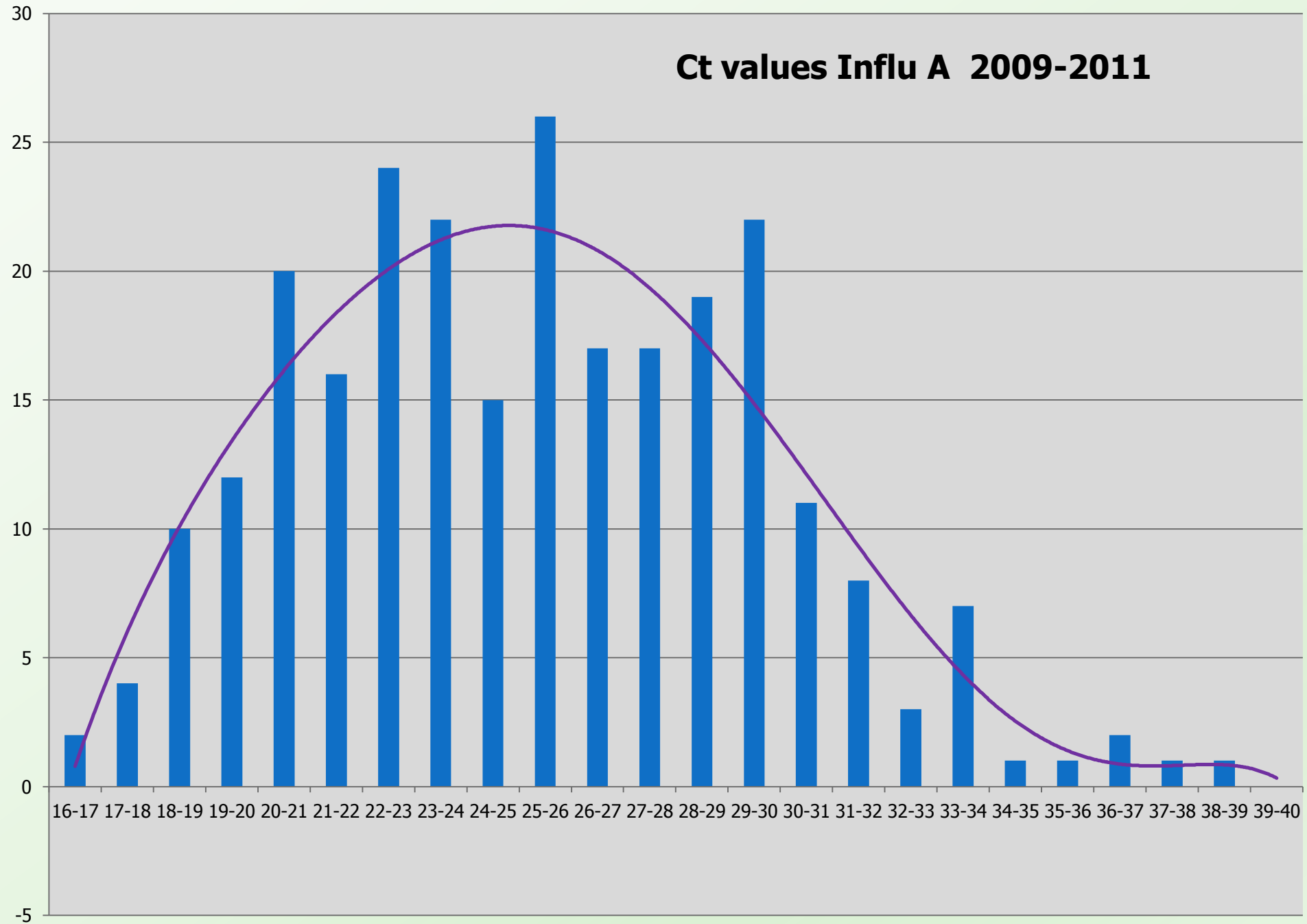
⇒ how to make the difference ?



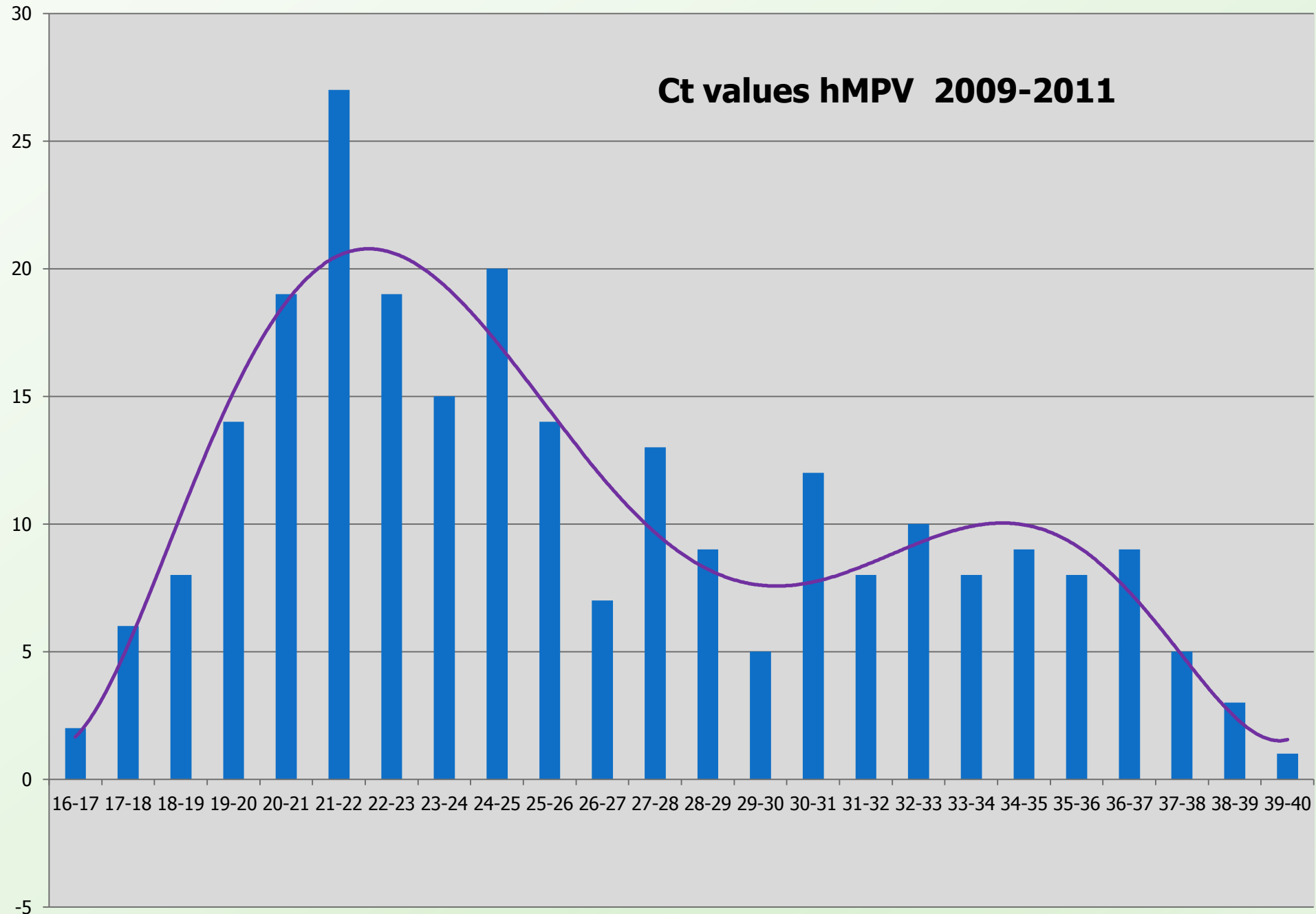
Ct values RSV 2009-2011



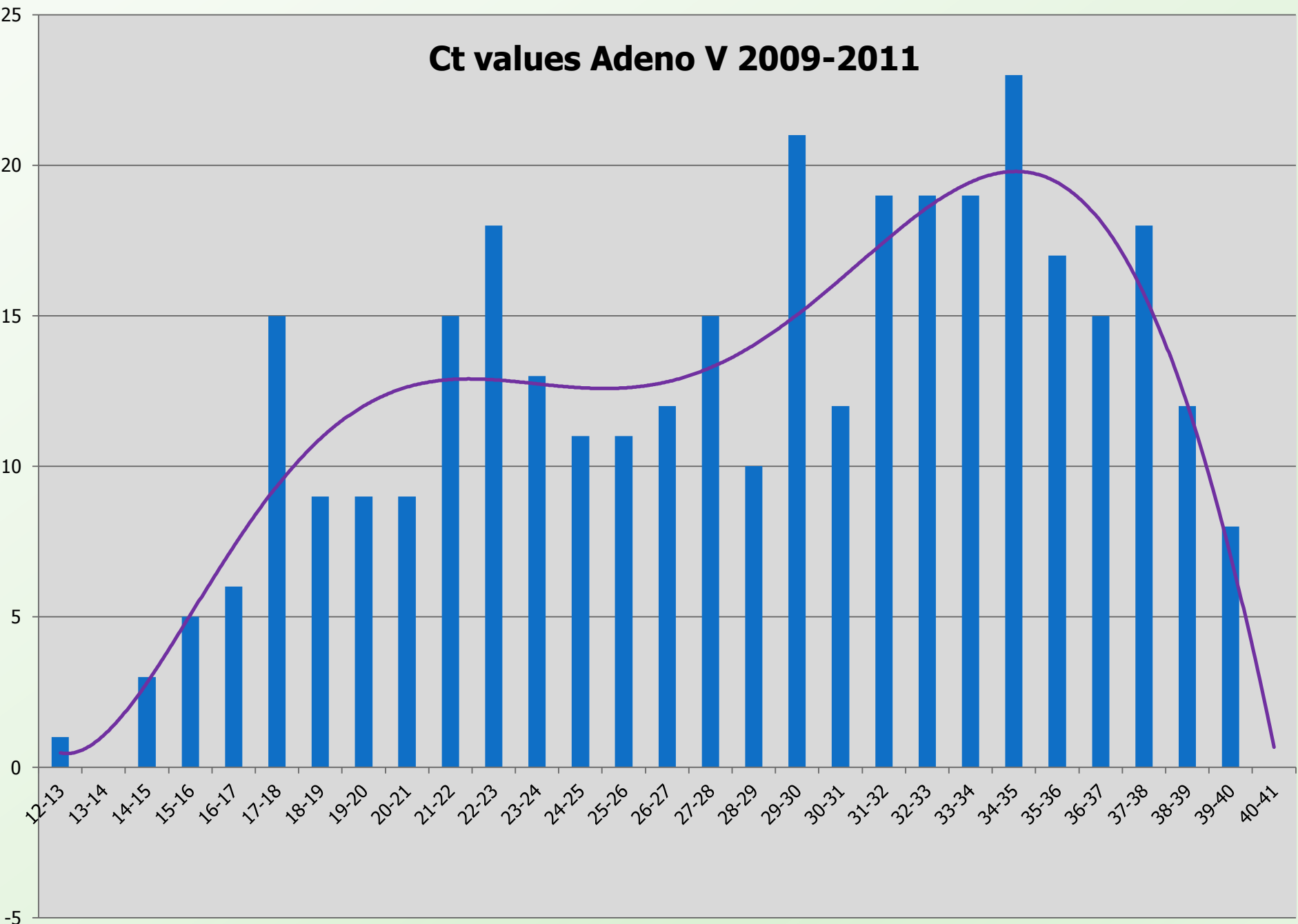
Ct values Influ A 2009-2011



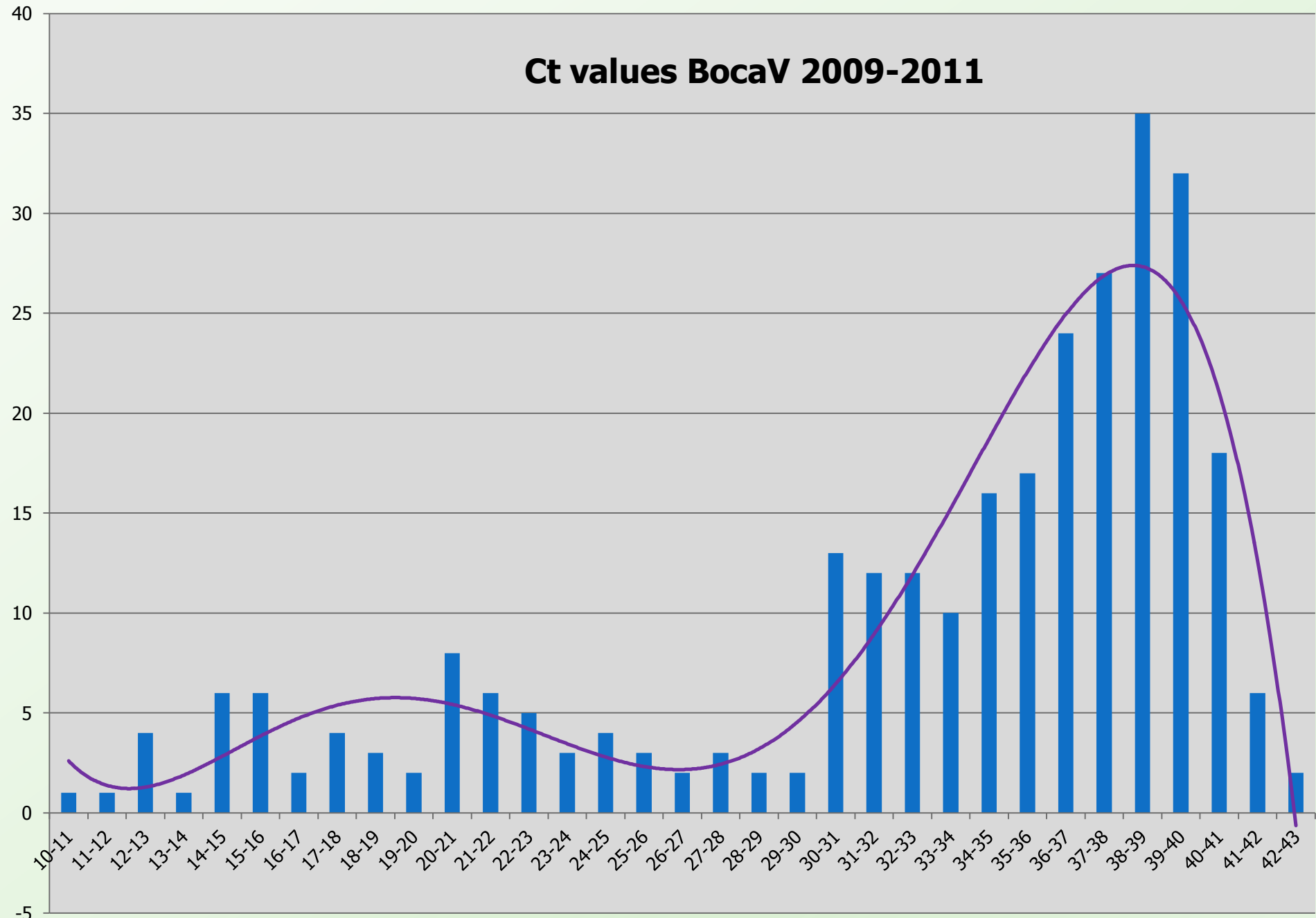
Ct values hMPV 2009-2011



Ct values Adeno V 2009-2011



Ct values BocaV 2009-2011



Etiology : how much

detection \neq infection ?

To be continued !
Come and see next time !



Thanks to

- ✓ Molbiol team (Magda, Elfi, Inge, Astrid, Freya, Karen)
- ✓ Anne Vankeerberghen (data !)
- ✓ An Boel, Kristien Van Vaerenbergh

