

Ard van Sighem<sup>1</sup>, Eline Op de Coul<sup>2</sup>, T. Sonia Boender<sup>1</sup>, Birgit van Benthem<sup>2</sup>, Jolande Bouwhuis<sup>3</sup>, Kees Brinkman<sup>4</sup>, Peter Reiss<sup>1,5,6</sup>, on behalf of the ATHENA national HIV cohort

<sup>1</sup>Stichting HIV Monitoring, Amsterdam, NL; <sup>2</sup>National Institute for Public Health and the Environment, Bilthoven, NL; <sup>3</sup>Isala, Zwolle, NL; <sup>4</sup>OLVG, Amsterdam, NL; <sup>5</sup>Academic Medical Centre, Amsterdam, NL; <sup>6</sup>Amsterdam Institute for Global Health and Development, Amsterdam, NL

The ATHENA database is maintained by Stichting HIV Monitoring and supported by a grant from the Dutch Ministry of Health, Welfare and Sport through the Centre for Infectious Disease Control of the National Institute for Public Health and the Environment.

**CLINICAL CENTRES** (\* denotes site coordinating physician)

**Academisch Medisch Centrum der Universiteit van Amsterdam (AMC-Uva):** HIV treating physicians: M. van der Valk\*, S.E. Geerlings, M.H. Godfriën, A. Goorhuis, J.W. Hovius, J.T.M. van der Meer, T.W. Kuijpers, F.J.B. Nellen, D. Pajkrt, T. van der Pol, J.M. Prins, P. Reiss, H.J. Scherpien, M. van Vugt, W.J. Wiersinga, F.W.M.N. Wit, HIV nurse consultants: M. van Duinen, J. van Eden, A.M.H. van Hes, M. Mutschelknauss, H.E. Nobel, F.J.J. Pijnappel, A.M. Weijenfeld, HIV clinical virologists/chemists: S. Jürrasits, N.K.T. Back, H.L. Zaaijer, B. Berkhouwt, M.T.E. Cornelissen, C.J. Schinkel, K.C. Wolthers. **Emma Kinderziekenhuis (AMC-Uva):** HIV nurse consultants: A. van der Plas, A.M. Weijenfeld. **Admiraal De Ruyter Ziekenhuis, Goes:** HIV treating physicians: M. van den Bergé, A. Stegeman, HIV nurse consultants: S. Baas, L. Hage de Looff. HIV clinical virologists/chemists: B. Wintermans, J. Veenemans. **Catharina Ziekenhuis, Eindhoven:** HIV treating physicians: M.J.H. Pronk\*, H.S.M. Ammerlaan, HIV nurse consultants: E.S. de Munck, HIV clinical virologists/chemists: A.R. Jansz, J. Tjolie, M.C.A. Wedgdam, B. Deiman, V. Scharnhorst. **DC AMC HIV Behandelcentrum:** HIV treating physicians: A. van Eeden\*, M. van der Valk, HIV nurse consultants: W. Broekking, M. Groot, L.J.M. Elsenburg, HIV clinical virologists/chemists: M. Damen, I.S. Kwa. **Elisabeth-Tweesteden Ziekenhuis, Tilburg:** HIV treating physicians: M.E.E. van Kasteren\*, A.E. Brouwer, HIV nurse consultants: R. van Erve, B.A.F.M. de Kruijff-van der Wiel, S. Keelan-Pfaf, B. van der Ven. Data collection: B.A.F.M. de Kruijff-van der Wiel, B. van der Ven. HIV clinical virologists/chemists: A.G.M. Buiting, P.J. Kabel, D. Versteeg. **Erasmus MC, Rotterdam:** HIV treating physicians: M.E. van der Ende\*, H.J. Bak, E.C.M. van Gorp, J.L. Nouwen, B.J.A. Rijnders, C.A.M. Schurink, A. Verbon, T.E.M.S. de Vries-Slijps, N.C. de Jong-Peltenburg, HIV nurse consultants: N. Bassant, J.E.A. van Beek, M. Vriesde, L.M. van Zonneveld. Data collection: H. van der Berg-Cameron, J. de Groot, M. de Zeeuw-de Man. HIV clinical virologists/chemists: C.A.B. Boucher, M.P.G. Koopmans, J.J.A. van Kampen, S.D. Pas. **Erasmus MC-Sophia, Rotterdam:** HIV treating physicians: P.L.A. Fraaij, A.M.C. van Rossum, C.L. Vermont, HIV nurse consultants: L.C. van der Knaap, E. Visser. **Flevoziekenhuis, Almere:** HIV treating physicians: J. Branger\*, A. Rijkeboer-Mes, HIV nurse consultant: C.J.H.M. Duijf-van de Ven. **HagaZiekenhuis, Den Haag:** HIV treating physicians: E.F. Schippers\*, C. van Nieuwkoop, HIV nurse consultants: J.M. van Dijperen, J. Gellings. Data collection: G. van der Hut. HIV clinical virologist/chemist: N.D. van Burgel. **HMC (Haaglanden Medisch Centrum), Den Haag:** HIV treating physicians: E.M.S. Leyten\*, L.B.S. Gelinck, HIV nurse consultants: A.Y. van Hartingsveld, C. Meerkirk, G.S. Wildenbeest. HIV clinical virologists/chemists: E. Heikens. **Izala, Zwolle:** HIV treating physicians: P.H.P. Groeneweld\*, J.W. Bouwhuis, A.J.J. Lammers. HIV nurse consultants: S. Kraan, A.G.W. van Hulzen. Data collection: G.L. van der Bleek, P.C.J. Bor. HIV clinical virologists/chemists: P. Bloembergen, M.J.H.M. Wolfhagen, G.J.H. Ruijs. **Leids Universitair Medisch Centrum, Leiden:** HIV treating physicians: F.P. Kroon\*, M.G.J. de Boen, H. Schepers, H. Jolink, A.M. Vollaard. HIV nurse consultants: W. Dorama, N. van Holten. HIV clinical virologists/chemists: E.C.J. Claas, E. Wessels. **Maaststad Ziekenhuis, Rotterdam:** HIV treating physicians: W.H.P. Groeneweld\*. Data collection: J.W. Bouwhuis, A.J.J. Lammers. HIV nurse consultants: R. Kastelijns, J.V. Smit, E. Smit, D. Struijk-Kalkman, C. Tearnio. Data collection: T. van Niekerk. HIV clinical virologists/chemists: O. Pontesilli. **Maastricht UMC+, Maastricht:** HIV treating physicians: S.H. Lowe\*, A.M.L. Oude Lashof, D. Posthouwer. HIV nurse consultants: R.P. Ackens, K. Burgers, J. Schippers. Data collection: B. Weijenberg-Maes. HIV clinical virologists/chemists: I.H.M. van Loo, T.R.A. Havenith. **MC Stotterhart, Amsterdam:** HIV treating physicians: J.W. Mulder\*, S.M.E. Vrouwerts, F.N. Lauw. HIV nurse consultants: M.C. van Broekhuizen, D.J. Vlastuin. HIV clinical virologists/chemists: P.H.M. Smits. **MC Zuiderzee, Leylestad:** HIV treating physicians: S. Weijer\*, R. El Moussaoui. HIV nurse consultant: A.S. Bosma. **Medisch Centrum Leeuwarden, Leeuwarden:** HIV treating physicians: M.G.A.van Vonderen\*, D.P.F. van Houte, L.M. Kamperschuur. HIV nurse consultants: K. Dijkstra, S. Faber. HIV clinical virologists/chemists: J. Weel. **Medisch Spectrum Twente, Enschede:** HIV treating physicians: G.J. Kootstra\*, C.E. Delsing. HIV nurse consultants: M. van der Burg-van de Plas, H. Heins. Data collection: E. Lucas. **Noordwest Ziekenhuisgroep, Alkmaar:** HIV treating physicians: W. Kortmann\*, G. van Twillert\*, R. Renckens. HIV nurse consultants and data collection: D. Ruiter-Pronk, F.A. van Truijen-Oud. HIV clinical virologists/chemists: J.W.T. Cohen Stuart, E.P. IJzerman, R. Jansen, W. Rozemeijer W. A. van der Reijden. **OLVG, Amsterdam:** HIV treating physicians: K. Brinkman\*, G.E.L. van den Berk, W.L. Blok, P.H.J. Frissen, K.D. Lettinga, W.E.M. Schouten, J. Veenstra. HIV nurse consultants: C.J. Brouwer, G.F. Geerlings, K. Hoeksma, M.J. Kleene, I.B. van der Meché, M. Spelbrink, A.J.M. Toonen, S. Wijnands. HIV clinical virologists: D. Kwa. Data collection: R. Regez (coordinator). **Radboudumc, Nijmegen:** HIV treating physicians: R. van Crevel\*, M. Keuter, A.J.A.M. van der Ven, H.J.M. ter Hofstede, A.S.M. Dofferhoff, S.S.V. Henriet, M. van de Flier, K. van Aerde, J. Hoogerwerf. HIV nurse consultants: M. Albers, J.T.J. Grintjes-Huisman, M. de Haan, M. Marneef, A. Hairwassers. HIV clinical virologists/chemists: J. Rahamat-Langendoen, F.F. Stelma. HIV clinical pharmacology consultant: D. Burger. **Rijnstate, Arnhem:** HIV treating physicians: E.H. Gisolf\*, R.J. Hassing, M. Claassen. HIV nurse consultants: G. ter Beest, P.H.M. van Bentum, N. Langebeek. HIV clinical virologists/chemists: R. Tiemessen, C.M.A. Swanink. **Sparreine Gasthuis, Haarlem:** HIV treating physicians: S.F.L. van Lelyveld\*, R. Soetekouw. HIV nurse consultants: L.M.M. van der Pijt, J. van der Swaluw. Data collection: N. Bermon. HIV clinical virologists/chemists: W.A. van der Reijden, R. Jansen, B.L. Hépers, D. Veenendaal. **Medisch Centrum Jan Goyen, Amsterdam:** HIV treating physicians: D.W.M. Verhaegen. HIV nurse consultants: M. van Wijk. **Universitair Medisch Centrum Groningen, Groningen:** HIV treating physicians: W.F.W. Bierman\*, M. Bakker, J. Kleinijnenjans, E. Kloese, E.H. Scholvinck, Y. Stienstra, C.L. Vermont, K.R. Wilting, M. Wouthuyzen-Bakker. HIV nurse consultants: A. Boomstra, H. de Groot-de Jonge, P.A. van der Meulen, D.A. de Weerd. HIV clinical virologists/chemists: H.G.M. Niesters, C.C. van Leer-Buter, M. Knoester. **Universitair Medisch Centrum Utrecht, Utrecht:** HIV treating physicians: A.I.M. Hoepelman\*, J.E. Arends, R.E. Barth, A.H.W. Bruns, P.M. Ellerbroek, T. Mudrikova, J.J. Oosterheert, E.M. Schadd, M.W.M. Wassenberg, M.A.D. van Zoelen. HIV nurse consultants: K. Aarsman, D.H.M. van Elst-Laurijssen, I. de Kroon, C.S.A.M. van Rooten. Data collection: M. van Berkel, C.S.A.M. van Rooten. HIV clinical virologists/chemists: R. Schuurman, F. Verduyn-Lunel, A.M.J. Wensing. **VUmc, Amsterdam:** HIV treating physicians: E.J.G. Peters\*, M.A. van Agtmael, M. Bomers. HIV nurse consultants: M. Heitmueller, L.M. Laan. HIV clinical virologists/chemists: C.W. Ang, R. van Houdt, A.M. Pettersson, C.M.J.E. Vandernoucke-Grauls. **Wilhelmina Kinderziekenhuis, UMC Utrecht, Utrecht:** HIV treating physicians: L.J. Bont, S.P.M. Geelen, T.F.W. Wolfs. HIV nurse consultants: N. Nauta.

**COORDINATING CENTRE**  
Director: P. Reiss. Data analysis: D.O. Bezemer, A.I. van Sighem, C. Smit, F.W.M.N. Wit, T.S. Boender. Data management and quality control: S. Zaheri, M. Hillebrecht, A. de Jong. Data monitoring: D. Bergsma, S. Grivell, A. Jansen, M. Raethke, R. Meijering, T. Rutkens. Data collection: L. de Groot, M. van den Akker, Y. Bakker, M. Bezemer, E. Claessen, A. El Berkouit, J. Geerlings, J. Koops, E. Krijnje, C. Lodewijk, R. van der Meer, L. Munijnish, F. Paling, B. Peek, C. Ree, R. Regtop, Y. Ruijs, M. Schoor, A. Timmerman, E. Tuijn, L. Veenenberg, S. van der Vliet, A. Wisse, E.C. de Witte, T. Woudstra. Patient registration: B. Tuk.

## Contact

Ard van Sighem  
Stichting HIV Monitoring  
E: a.i.vansighem@amc.uva.nl  
T: +31205666781  
www.hiv-monitoring.nl

## Background

- The HIV care continuum summarises engagement in various stages of HIV care but does not provide information on the time to reach these stages (Figure 1).
- We aimed to estimate changes in the time to reach successive stages of the HIV care continuum in the Netherlands since 2000.

## Methods

### Patient population

- All 23,376 HIV-1-positive individuals in the ATHENA national observational HIV cohort aged ≥18 years at the time of diagnosis.

### Time to diagnosis

- ECDC HIV Modelling Tool used to estimate time between HIV infection and diagnosis.
- Estimation mainly based on changes over calendar time in the distribution of CD4 counts at diagnosis.

### Time to reach other stages

- Survival methods were used to estimate time between successive stages from HIV diagnosis onwards.
- Patients who did not start combination antiretroviral treatment (cART) or reach viral suppression were censored at their last follow-up visit.
- No censoring for reaching HIV care because the ATHENA cohort only includes individuals ever enrolled in care.

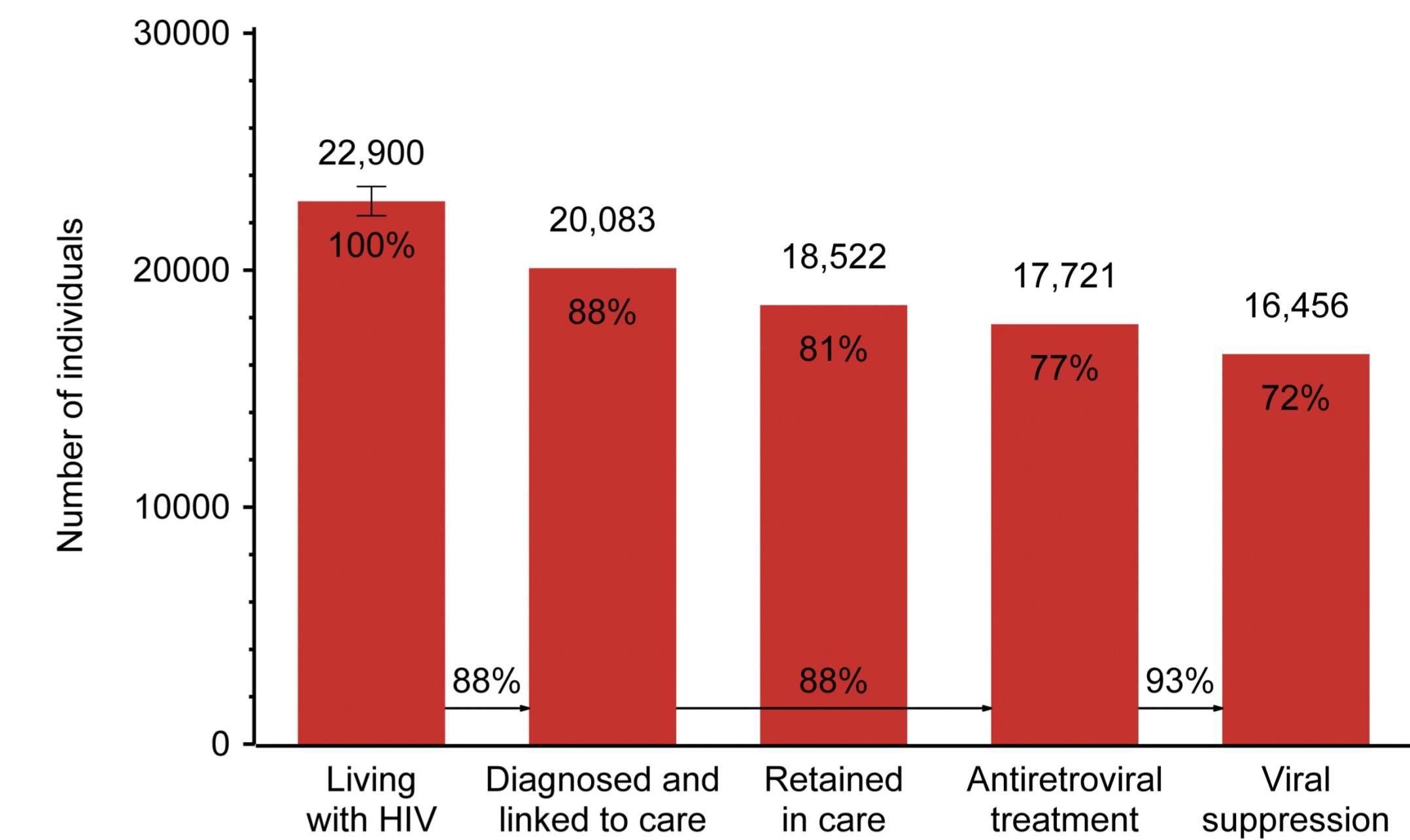
## Conclusions

- Time from acquiring HIV to reaching viral suppression has declined, but continues to be dominated by the duration of undiagnosed HIV infection.
- To further improve access to care and treatment, ensuring earlier diagnosis by expanding testing for HIV will be key.

**Figure 2: Estimated median time to reach key stages in the HIV care continuum for HIV-1-positive patients, including time from infection to diagnosis, from diagnosis to entry into care, from entry into care to starting combination antiretroviral treatment (cART), from starting cART to reaching viral suppression, and from diagnosis to viral suppression.**

1. HIV infection → diagnosis
  2. Diagnosis → care
  3. Care → cART
  4. cART → viral suppression
- Diagnosis → viral suppression

**Figure 1: Continuum of HIV care for the total HIV-positive population living in the Netherlands by the end of 2015. Percentages at the top of the bars are calculated relative to the estimated number living with HIV, while percentages at the bottom correspond to UNAIDS' 90-90-90 targets.**



## Results

- 17,053 (73%) individuals were diagnosed in 2000 or later.
- CD4 counts at the time of diagnosis increased from 260 (interquartile range, 78-460) cells/mm<sup>3</sup> in 2000 to 370 (150-560) cells/mm<sup>3</sup> in 2015.
- Estimated time from infection to diagnosis decreased from 4.7 years in 2000 to 2.8 years in 2015 (Table 1; Figure 2).
- Changes in time between HIV diagnosis and viral suppression mainly reflected changes in time to start cART.

**Table 1: Estimated median time and interquartile range between key stages in the HIV care continuum; viral suppression: first HIV RNA measurement <100 copies/ml.**

From	to	2000	2015
Infection	Diagnosis	4.7	2.8
		2.3-8.4	1.3-5.1
Diagnosis	Care	0.07	0.02
		0.02-0.24	0.01-0.04
Care	cART	0.27	0.07
		0.09-2.45	0.04-0.13
cART	Viral suppression	0.28	0.18
		0.18-0.57	0.08-0.32
Diagnosis	Viral suppression	0.85	0.37
		0.41-3.66	0.25-0.60

