

Netherlands Brain Bank



Progress Report 2009-2010

Netherlands Brain Bank

Progress Report 2009 - 2010

Editors

Inge Huitinga
Michiel Kooreman
Marleen C. Rademaker
Wilma T.P. Verweij

Correspondence

Netherlands Brain Bank
Meibergdreef 47
1105 BA Amsterdam
The Netherlands

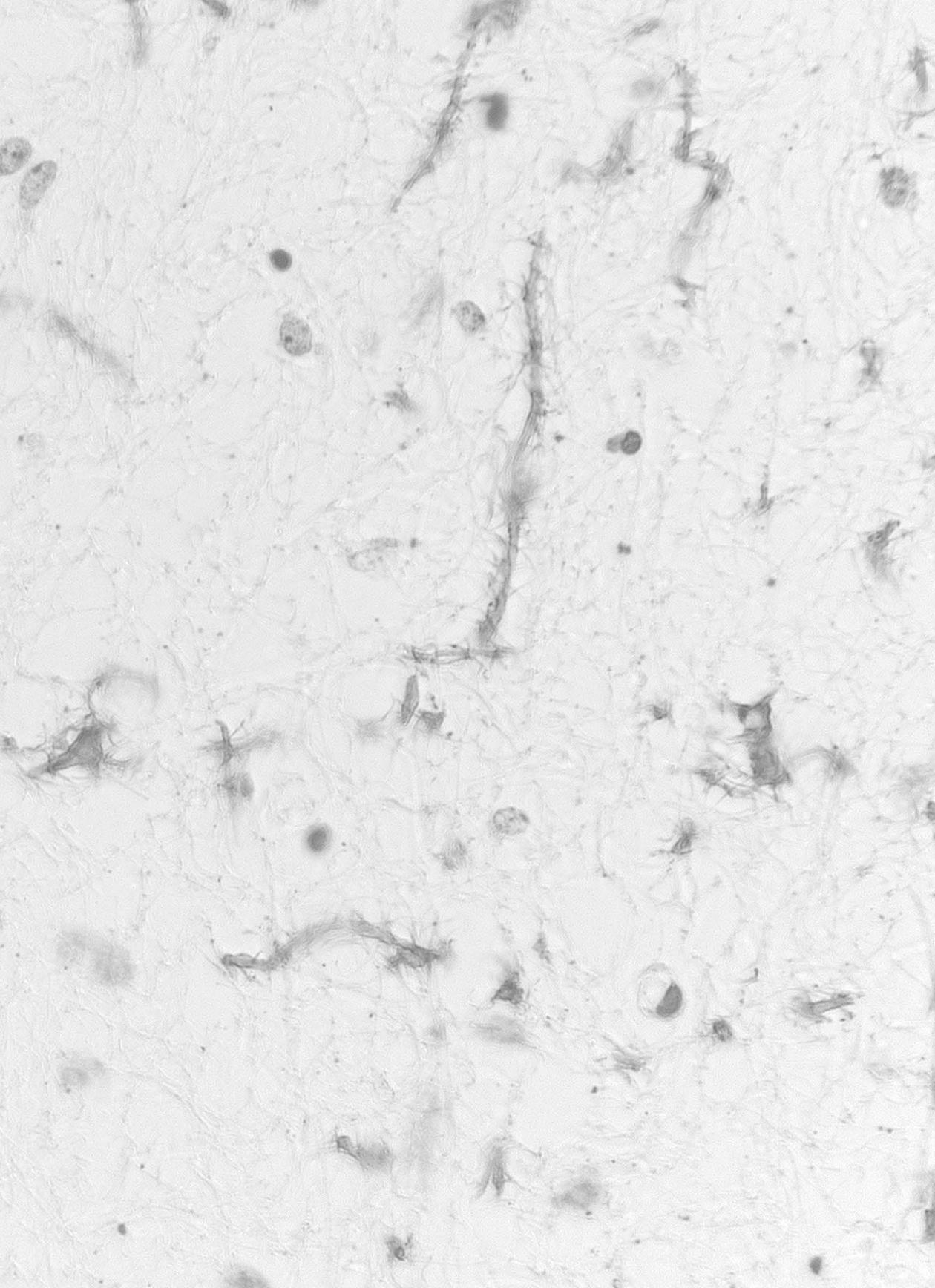
T (+31) 20 566 5499
F (+31) 20 691 8466
secretariaatnhb@nin.knaw.nl
www.brainbank.nl
www.nin.knaw.nl

Booklet design: Henk Stoffels

Print: Gildeprint BV, Enschede

Contents

Introduction	7
Objectives NBB	9
Donor registrations	11
Autopsies	17
Tissue supply	21
BrainNet Europe	25
Finances	29
Research projects 2009-2010	31
Publications 2006-2010	41
Staff and collaborations	56
Appendix	58
Abbreviations	60



Introduction

It is with great pleasure that I present the 2009/2010 progress report of the Netherlands Brain Bank (NBB). The primary activities of the NBB are to motivate people to register as brain donor, to perform autopsies and to disseminate tissue to researchers worldwide. In 2006 we professionalized all NBB procedures, from procurement and storage to dissemination to researchers. New informed consent forms, informational DVDs, improved tissue application procedures, advisory committees and increased public relation activities were all effective, resulting in increased numbers of donor registrations and autopsies. The number of annual registrations increased from 180 in 2006 up to a steady 290 in 2008-2010. Since the foundation of the NBB in 1985 the annual numbers of autopsies have kept climbing. In 2006, 82 autopsies were recorded and in 2010 the NBB reached the record autopsy number of 127!

After the successful reactivation of donor programs and the professionalization of brain banking procedures, we have now reached a point where our donor programs need more focus. To ensure a more effective dissemination of tissue we shall need to gear any new donor registrations to the most frequently requested types of tissues. In the coming years our focus will thus be on brain diseases for which we receive many tissue requests, and on patients who participate in clinical cohorts and are therefore well-documented. As a consequence, the registration of complex cases that are seldom requested by researchers will have a lower priority.

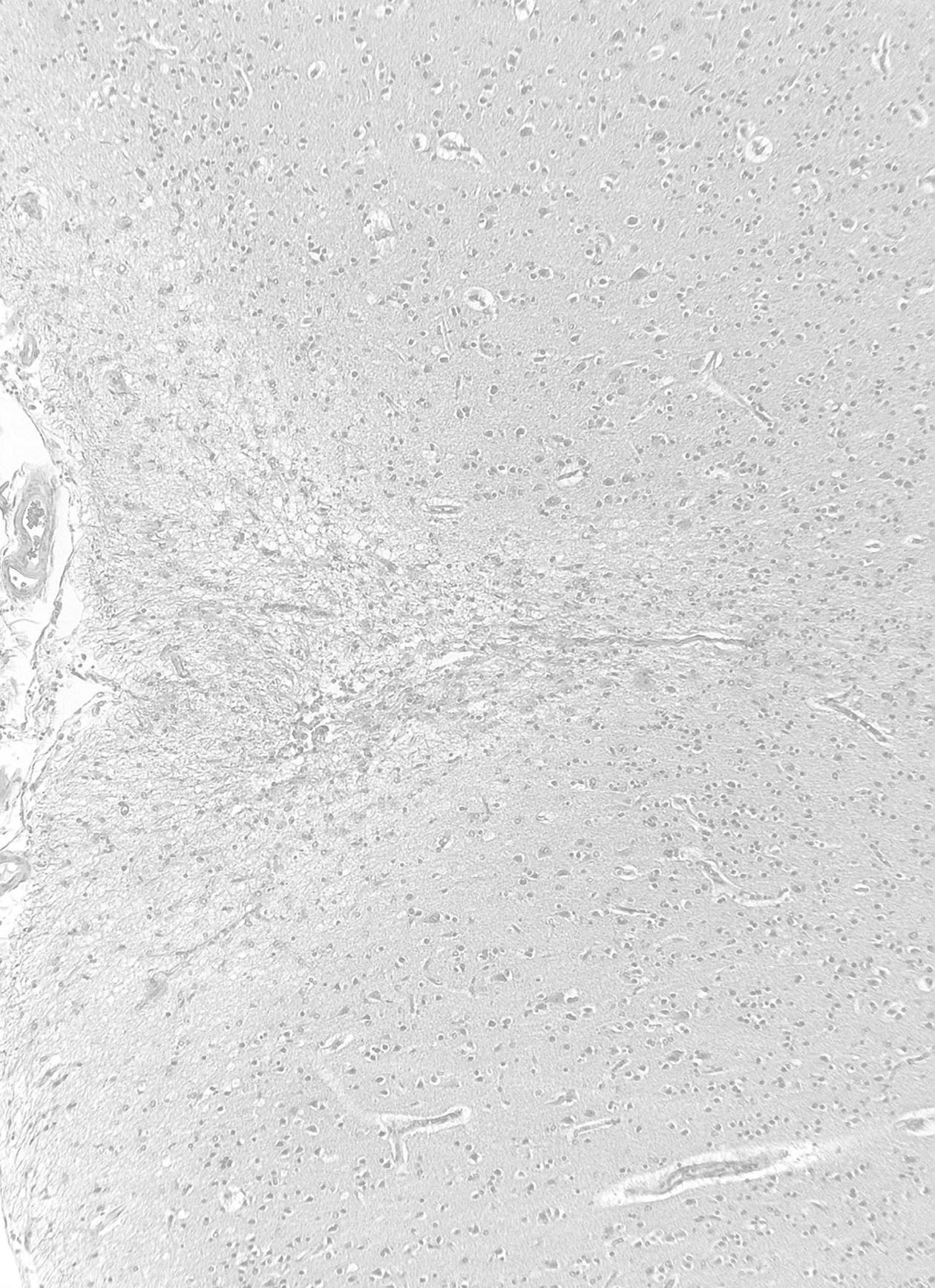
I would like to express my gratitude to the *NIN*, *KNAW*, *Stichting MS Research*, *Internationale Stichting Alzheimer Onderzoek*, *Internationaal Parkinson Fonds* and *Hersenstichting Nederland*, as well as to private backers, for their financial support, which is indispensable for the continuation of the NBB.

I also thank the members of the autopsy team for their guidance and help with the autopsies, day or night. Many of these are PhD students and technicians, who have volunteered to help us out despite their own busy programs and work commitments. Also indispensable are the autopsy assistants and pathologists at VUmc, to whom I would like to express my gratitude for their willingness to perform the autopsies.

Last but not least, I thank the donors, without whose willingness to donate their brain, worldwide scientific research of the brain and brain disease would not be possible.

Inge Huitinga

Director Netherlands Brain Bank

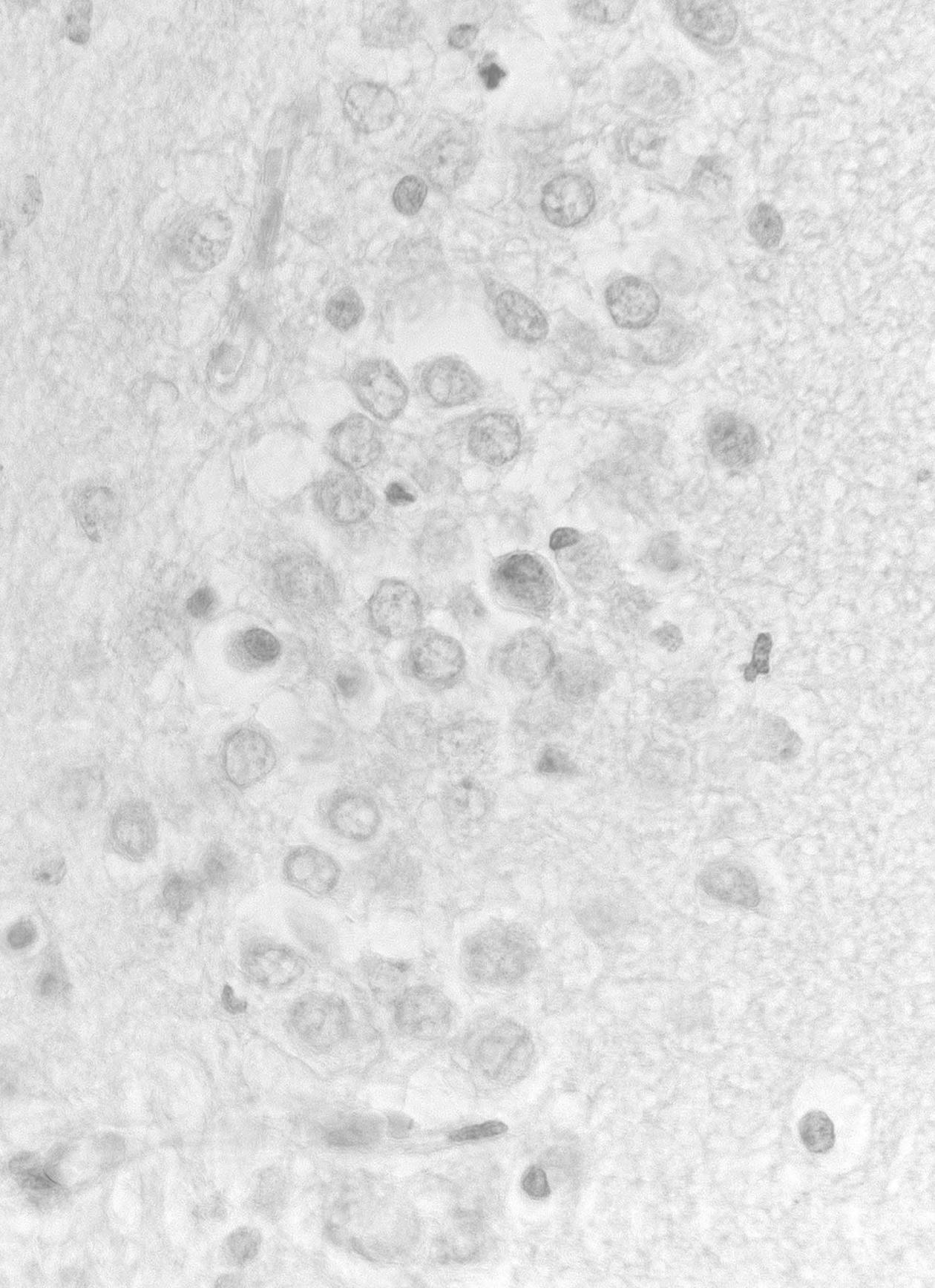


Objectives NBB

The Netherlands Brain bank was founded in 1985 by Professor Dick Swaab (1944) to obtain brain tissue for his Alzheimer research. While setting up an infrastructure to register Dutch people for a rapid brain autopsy for research purposes, he realized that such a facility would be of great value also for other researchers in neuroscience. The NBB has thus been open to tissue applications from researchers worldwide from the very start.

The primary objective of the NBB still is to collect, characterize and disseminate tissue of the human brain and spinal cord for scientific research worldwide. The ultimate goal is to increase knowledge of the human brain and to find cures for neurological and psychiatric brain diseases.

An overview of the current composition of the NBB can be found in the Appendix (Figure 9).



Donor Registrations

The NBB is one of the few brain banks in the world with an active donor program, which means that the NBB actively tries to motivate people with neurological, psychiatric and neuroendocrine disorders, as well as healthy individuals, to register as brain donor at the NBB. With this registration, donors give informed consent to the NBB to perform a rapid autopsy after death and to donate the brain tissue to reviewed research projects around the world. The donors also give permission to the NBB for the release of their medical information after they have passed away. Currently, 2374 living donors with a variety of disorders are registered at the NBB.

New Informed Consent Forms

In 2008, the NBB created new registration forms and accompanying informational brochures (informed consent forms), which are in line with regulations and guidelines issued by international key organizations, such as the Council of Europe, the European Commission, the World Medical Association and the World Health Organization. The informational brochures and registration forms were reviewed by the Medical Ethics Committee of VUmc and officially approved on October 30, 2009.

Biannual Newsletter

In order to inform our donors about the progress made within the NBB and about the scientific output achieved with material provided by the NBB, we initiated a biannual Newsletter for all our registered donors. The first edition (n = 2250) was issued in April 2009. The NBB received many positive reactions from the registered donors as well as from their relatives. An additional advantage of this large-scale mailing was that it cleaned up our database: 83 registrations (3.7%) could be removed. In some cases the newsletter elicited a notice from a family that the prospective donor had already passed away. The most frequently given reason why the NBB had not been contacted at time of death was that the spouse or children found the prospect of a brain autopsy too difficult.

Registrations

Figure 1 shows the number of registrations in 2009 and 2010, compared to the registrations in the period 2007-2008. The total number of registrations in 2009/2010 increased slightly in comparison with those in 2007/2008 (545 vs. 513), although the

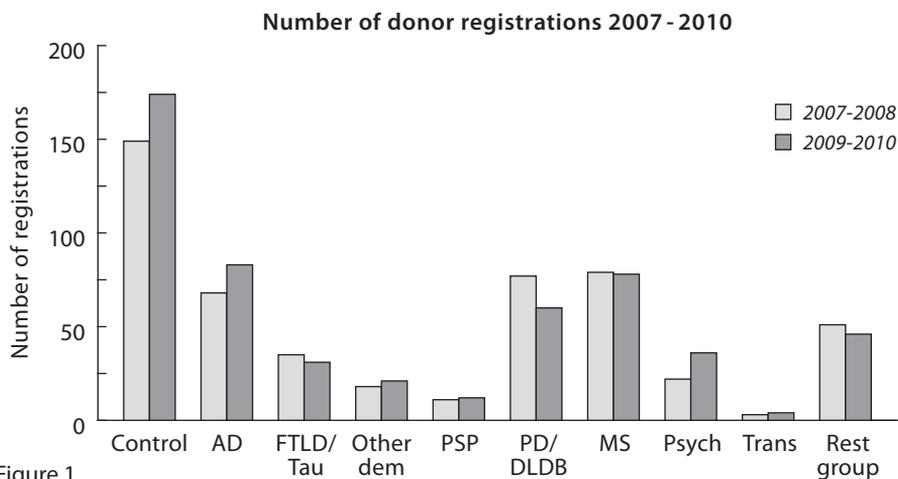


Figure 1

registrations did not increase for every diagnosis. The NBB is very pleased with the increase of donors with psychiatric disorders (depression, schizophrenia). The increase for multiple sclerosis (MS) and non-demented control donors is probably due to the promotional DVDs which can be ordered via our Dutch website. Especially the MS DVD 'De Oplossing zit in de Hersenen' is frequently ordered by (potential) donors: 130 times since its release in 2007. It gives an elaborate overview of the procedures of the NBB with regard to MS. In the last two years, the NBB sent more than 1000 information packs to individuals, neurologists and nursing home physicians (Table 1). The registration forms can also be downloaded from our website.

In total we received 268 new registrations in 2009 and 277 new registrations in 2010 (Figure 2). After the increase in annual registrations from 2007 onward, the number of registrations per year now seems to have stabilized. The number of new female registrations remains higher than that of new male registrations. This might

Table 1 PR Material NBB

Information pack	2009	% registered	2010	% registered
Informed consent	194	40%	132	38%
Authorization	23		11	
Informed consent via physician	221		155	
Authorization via physician	179		140	
<i>Total number of information packs sent</i>	617		438	

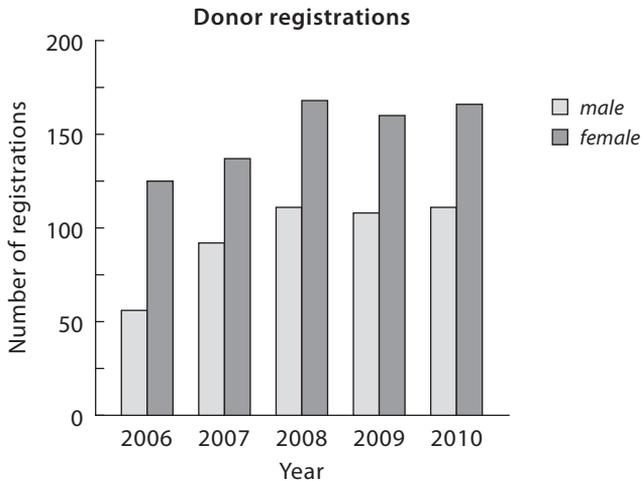


Figure 2

be caused by the increase of female MS and non-demented control registrations. The higher prevalence of MS (two-fold) in females probably explains this. As to the disproportionate increase of female control donors, we are as yet unable to offer an explanation for this development. This difference between the numbers of male and female registrations is not seen in organ donation for transplantation purposes (source: www.donorvoorlichting.nl).

The increase in the total numbers of annual registrations is reflected in the annual number of autopsies, which has increased from 82 in 2006 to 127 in 2010 (Table 3). Indeed, 46 % of the registrations occur in the year before autopsy (Table 4).

Increased focus of the NBB donor program

The NBB shifted its focus from general donor recruitment to recruitment from clinical cohorts. Many academic hospitals have clinical cohorts of patients with a specific neurological or psychiatric disorder to study disease course and the effect of experimental therapies. These patients are studied longitudinally and therefore many medical data are available in a standardized manner. This makes them a very interesting group for post mortem research. Moreover, these people - willing to participate in research during life - tend to be willing to donate tissue after their demise as well.

Presentations and articles

In the past two years the NBB has spent a great deal of time and effort into raising awareness of the importance of research with human brain tissue and the possibility of brain donation. We visited patient meetings to give presentations on the work

of the NBB and the possibility to become a donor. Being able to show the kind of research that is performed on tissue donated to the NBB - research that might help find a cure - evokes many positive reactions and has led to many new donor registrations. Table 2 gives an overview of the articles that were published about the work of the NBB in 2009 and 2010. We always make sure to mention that not only patients with neurological or psychiatric diseases, but also healthy control donors are crucial for good scientific research. In that way, we may also persuade many non-diseased family members to register as brain donor.

Websites of the NBB

Nowadays the internet is a very popular source for patients trying to learn more about their illness. By making sure that the NBB is mentioned on the websites of the various patient organizations, we try to enhance public awareness of the importance of brain donation. Also our donor website (www.hersenbank.nl) is updated regularly to inform our (potential) donors about the work of the NBB. The English website of the NBB (www.brainbank.nl) has been renewed for the sake of convenience for researchers and now provides more information on our procedures, diagnostics and the availability of tissue.

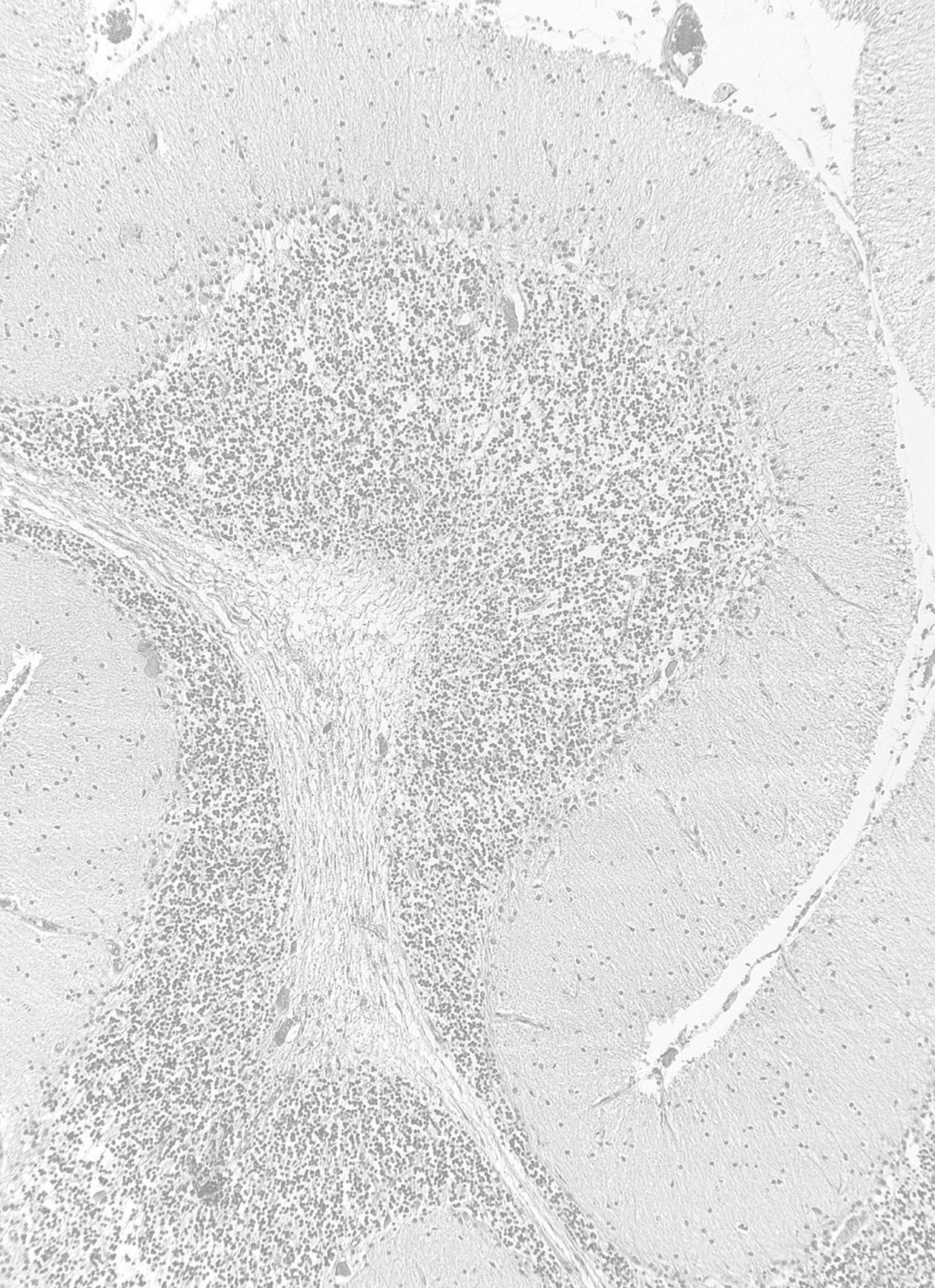
Table 2 Overview of PR activities in 2009 and 2010

Date	Article / presentation	Media
01-01-09	Interview technical coordinator NBB	Noorderlicht (radio show)
01-02-09	Interview director NBB	De Praktijk (radio show)
01-05-09	Radio commercial by director NBB	Radio commercial as part of World MS Day
01-09-09	'It is our mission to find solutions for brain diseases'	HersenMagazine (Magazine of patient organization)
04-10-09	Information stand	National Science Day in the Academic Medical Centre
01-10-09	Information stand	Publieksdag Hersenstichting Nederland
01-11-09	Radio commercial by director NBB	Radio commercial for MS Research is repeated
07-11-09	'MS research and the NBB'	Theme day of MSVN, Schagen
1-2-2010	'Looking the last will in the eye'	TV guide VPRO
1-6-2010	'A brain never gets normal'	Volkskrant (Daily newspaper)
01-06-10	'Brain researchers by accident'	Volkskrant (Daily newspaper)

Future plans

In the upcoming years the NBB will continue to pay special attention to people with psychiatric disorders such as autism, depression, schizophrenia and various addictions. Even though most psychiatric patients are able to give informed consent, they are often reluctant to register as donors at the NBB. The NBB will therefore continue to work together with physicians, psychiatrists and psychiatric nurses to inform patients in clinical cohorts about the importance of brain donation.

The NBB wishes to acknowledge and thank all donors and their families for their generosity and the invaluable gift they are giving to future generations.



Autopsies

Increase in annual number of autopsies

Since 1985 the NBB has performed over 3400 brain autopsies in all. The NBB performed 110 autopsies in 2009 and 127 in 2010. Table 3 shows the number of autopsies per year over the last 5 years, clearly showing that the total number of annual autopsies is increasing.

In the last five years, the number of autopsies per month ranged from 1 to 19. The fluctuations per month, however, show no significant differences of numbers of autopsies per month between the different months of the year ($p = 0.4$, Kruskal–Wallis one-way analysis of variance).

Relation between registration and autopsy

Based on the difference between year of registration and year of autopsy, the NBB calculated the average duration of the registration specified by the neuropathological diagnosis (Table 4). Interestingly, the number of registrations in the year of demise is generally double that of one year before demise. Moreover, 46 % of the registrations occur during the year of, or one year prior to, demise. However, this does not apply for the non-demented control group, where the duration of registration fluctuates more. The registration duration of MS donors shows a different pattern, probably caused by the fluctuating course of the disease. This suggests that the moment of

Table 3 Annual numbers of autopsies divided by disease

Diagnosis	2006	2007	2008	2009	2010
Contr	11	12	16	17	13
AD	30	28	30	29	16
FTLD/tau	2	8	15	15	7
Other dem	9	7	13	6	9
PSP	2	3	5	9	4
MS	8	14	10	11	8
PD/DLBD	6	7	12	16	12
Psych	4	2	4	2	2
Other	10	9	5	5	5
PANR					51
<i>Total</i>	<i>82</i>	<i>90</i>	<i>110</i>	<i>110</i>	<i>127</i>

Table 4 Duration of registration in months, specified by neuropathological diagnosis

	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	>15
Contr	23	17	7	8	15	18	9	9	12	5	8	9	9	8	4	6	7
AD	136	65	45	53	29	19	16	8	4	4	4	5	2	2	2		2
FTLD/tau	30	15	7	7	3	8	2	6	1		3	1					
Other dem	44	20	12	8	5	5	4	1	2	2	3	1	1	2	2	3	1
PSP	18	10	3	1	2	2	1	1	1								
PD/DLBD	43	24	9	6	3	1		4	1	2	5	2		2			1
MS	13	6	7	6	6	12	8	5	9	5	6	7	7	3	2	3	6
Psych	9	2	2	4	1				2		1	1	1	1	2		1
Rest group	27	24	8	8	5	4	1	1	1		1		2	1	1		1

registration is motivated by the amount of suffering caused by Alzheimer’s disease (AD), but also by understanding the necessity to donate your brain in order to find a cure for brain diseases (MS and controls). Importantly, the results in figure 6 indicate that the increasing number of annual autopsies can be attributed to donor recruitment efforts of the past couple of years.

Mean age of NBB donors at the time of death

The mean age at time of death from the autopsies performed in the period 2006-2010 is 70.8 years for males and 75.5 years for females. However, there are significant differences between the different diagnoses. Figure 3 shows the mean age at time of death (2006-2010) specified by diagnosis. The last column shows the Dutch life expectancy numbers (at birth) of 2009 (source: Centraal Bureau voor de Statistiek, www.cbs.nl), which are 78.8 and 82.6 for male and female respectively. Striking is the lower age at time of death for frontotemporal dementias and multiple sclerosis. These data are in line with the shorter life expectancy for those suffering from these neurological disorders (Hodges et al. 349-54; Sadovnick et al. 991-94; Sumelahti et al. 350-55).

Post mortem delay

Due to autolytic processes, tissue of the central nervous system quickly decays after death and there is thus only a small window of opportunity for brain autopsy. The post mortem delay (PMD: time elapsed from a person’s demise to removal of the brain) depends on several factors: time of notification of the donor’s death, distance

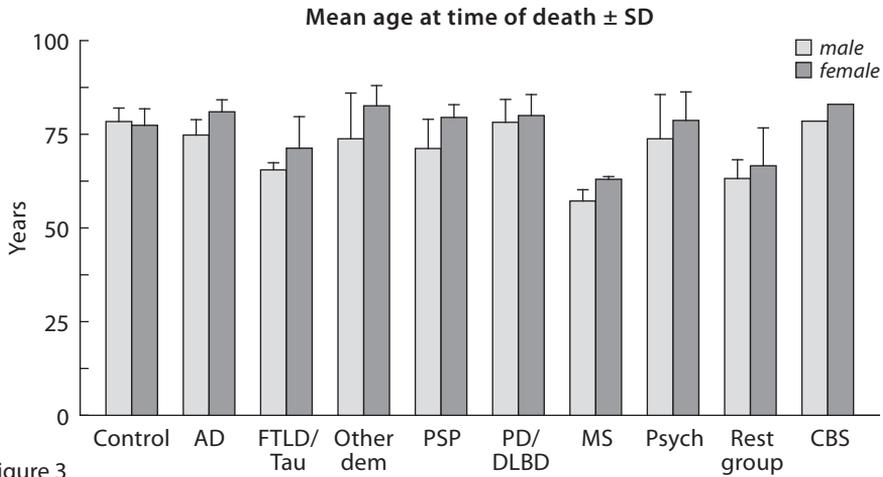


Figure 3

and time for transportation of the corpse and the availability of brain bank staff to perform the autopsy. Because PMD has a strong impact on the quality of the tissue (i.e. RNA, DNA and proteins; see references Chapter BrainNet Europe), several brain banks established rapid autopsy protocols relying on 24/7 availability of staff. The NBB achieves short PMDs, with 65 % of all autopsies having a PMD between 4 to 8 hours, whereas the average PMD of other European brain banks is more than 12 hours, even when they work with a 24/7 availability of staff (manuscript in prep.). Over the last 5 years the average PMD of the NBB autopsies has been extremely stable (Figure 4).

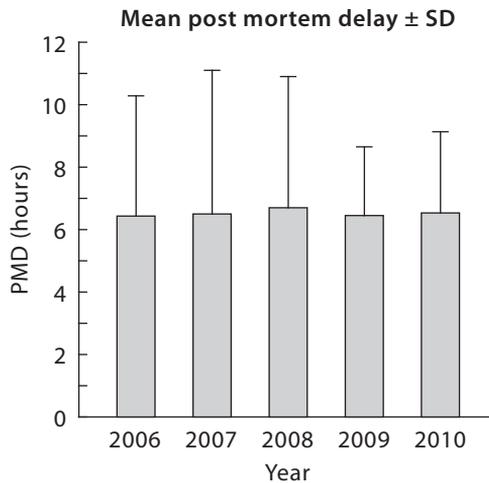


Figure 4

References

- Hodges, J.R., Davies, R., Xuereb, J., Kril, J., and Halliday, G. (2003). Survival in frontotemporal dementia. *Neurology* 61, 349-354.
- Sadovnick, A.D., Ebers, G.C., Wilson, R.W., and Paty, D.W. (1992). Life expectancy in patients attending multiple sclerosis clinics. *Neurology* 42, 991-994.
- Sumelahti, M.L., Tienari, P.J., Wikstrom, J., Salminen, T.M., and Hakama, M. (2002). Survival of multiple sclerosis in Finland between 1964 and 1993. *Mult. Scler.* 8, 350-355.

Tissue Supply

Number of research institutes that receive NBB tissue

In 2006 the NBB undertook to review all its procedures, which led to new informed consent forms and to professionalization of the application and tissue dissemination procedures. A Material Transfer Agreement (MTA) was drafted and put into use, to ensure the rights and obligations of the recipients of the tissue as well as those of the NBB. The first MTA was signed in June 2007. By now the NBB has entered into agreement with more than 70 universities/research institutes and 15 pharmaceutical companies worldwide. Once both parties have signed the MTA, which is valid for an indefinite period of time, any researcher within the institute can apply for tissue.

Number of tissue applications

The number of tissue applications has been on the increase since the introduction of the new procedures, but has stabilized now (Figure 5). Researchers have the possibility to inquire about the availability of samples, which in most cases leads to an application. When it concerns a new research project, the application is reviewed by the NBB's scientific committee. If approved, a new project number is assigned and the necessary paperwork is done, after which the tissue is supplied. The review process takes approximately four weeks. When the application concerns an existing, already reviewed, research project this is called a supplementary application. The option of filing a supplementary application was introduced in 2007, together with

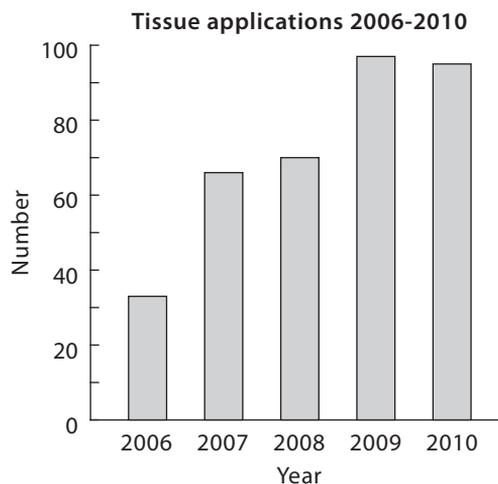


Figure 5

the MTA. With the original research project already approved, the requested tissue, if available, can be supplied even more quickly.

In 2009 and 2010 there were 24 cases (out of 192) where tissue inquiries did not lead to actual applications or where applications could not be approved. Inquiries can be for new applications as well as for supplementary applications. The main reasons why tissue inquiries or applications foundered are:

- an application form was sent to the researcher, but the researcher never actually applied for tissue;
- the researcher had to cancel the application due to financial problems (rejected grant applications);
- the NBB did not have the requested tissue.

The latter shows the need to increase the number of donors with a specific neurological or psychiatric disorder and was one of the reasons to start donor recruitment efforts among clinical cohorts referred to in the section on Donor Registrations.

Tissues disseminated for research projects

Figure 6 shows the specification of supplied samples by diagnosis in 2009 and 2010, compared to the tissue supply in 2007 and 2008. Especially for research on multiple sclerosis and Parkinson’s disease more samples were supplied. Also the number of samples supplied from non-demented control donors increased considerably. Thus, in spite of the fact that the number of applications remained stable, numbers of tissue units per application increased.

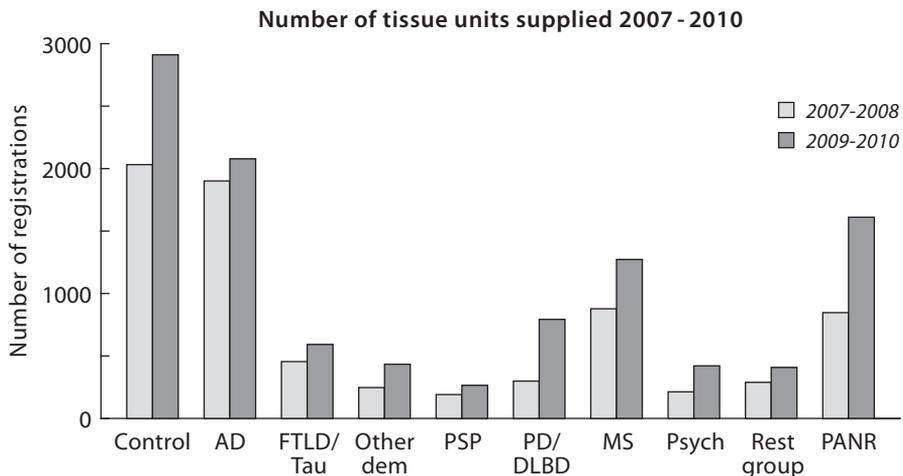


Figure 6

Figure 7 displays the specification of the samples by type of storage. The NBB not only provides frozen or formalin fixed paraffin embedded samples, but also fresh tissue and formalin fixed tissue. The different treatments of the tissue allow the possibility of different kinds of research approaches.

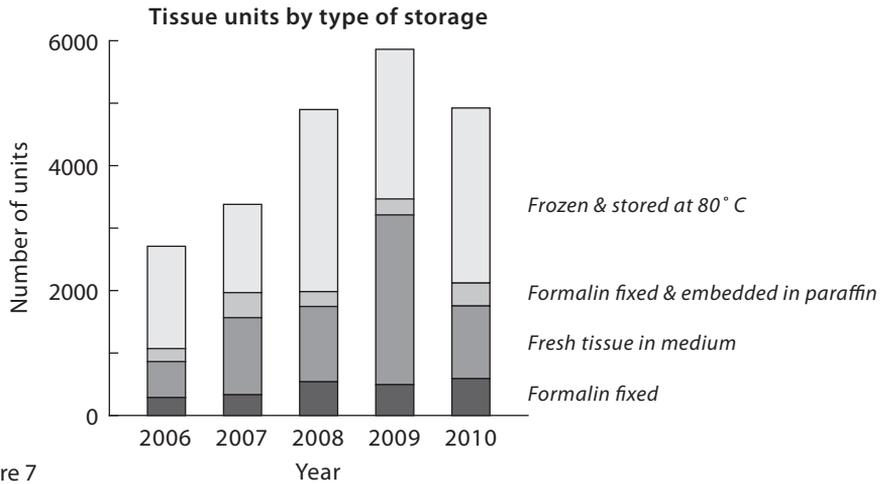
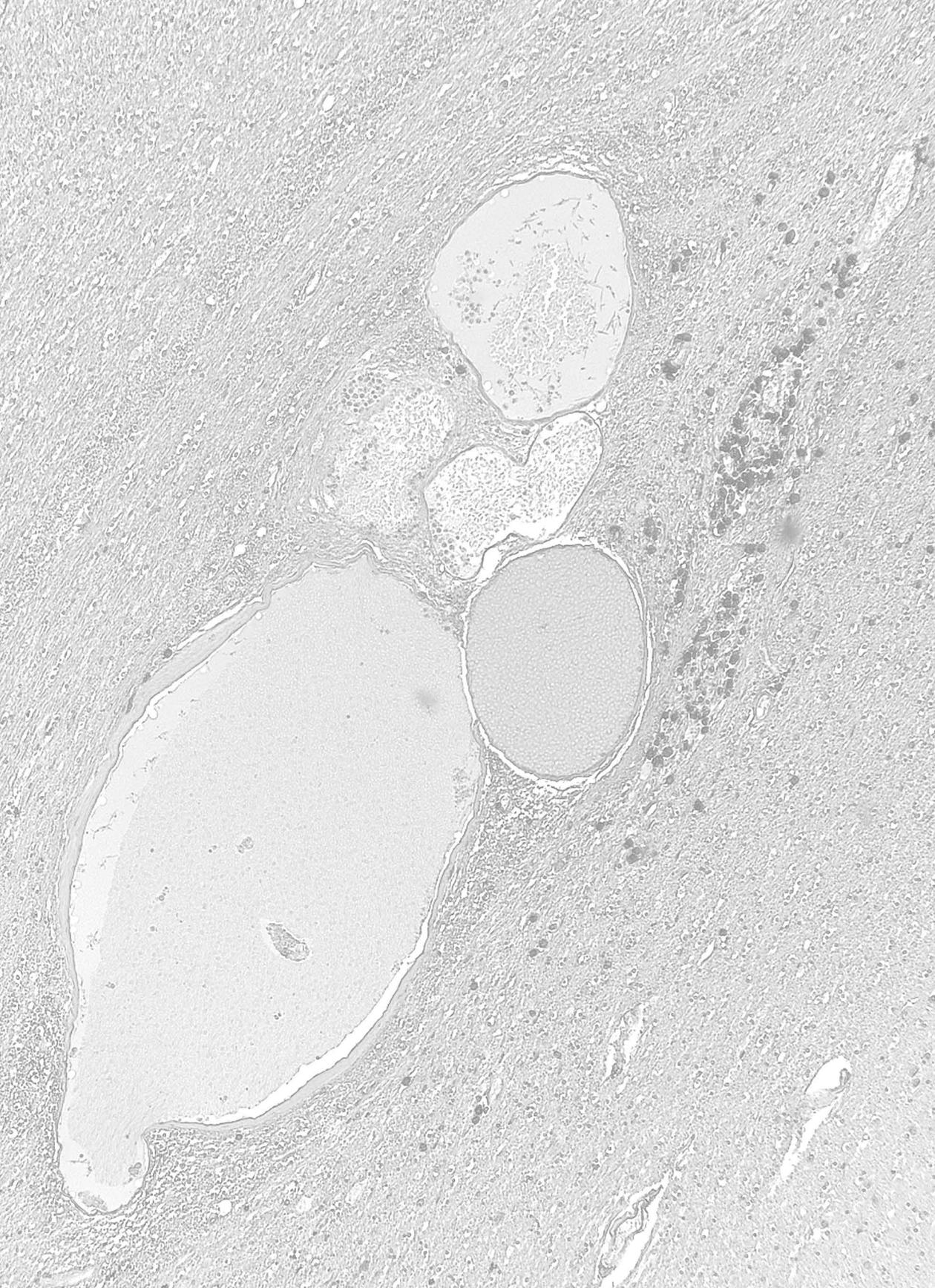


Figure 7



BrainNet Europe

BrainNet Europe II (BNE II) is a 'Network of Excellence', established in the 6th Framework Programme of Life Sciences of the European Commission (FP6) (www.brainnet-europe.org). The Consortium consists of 19 brain banks across Europe. Until 2009, BNE II was funded by the European Commission (EC), in order to carry out work with regard to its objectives, which are, among other things:

- Harmonization of neuropathological diagnostic criteria in Europe;
- Development of gold standards for quality, safety and ethics for obtaining and handling of human tissue;
- Sharing of knowledge and dissemination of the information to neuroscientists and the general public.

Now that the funding by the EC has come to an end, BNE is in transition: from an FP6 funded network it has to transform into a registered BNE Society. All necessary preparations are being made to continue BNE in an independent manner, starting with the BNE Charter which is currently being written.

The NBB has been a longstanding member of the BNE Consortium and an active participant designated to carry out work with regard to the ethical and legal issues in brain banking and recruitment of donors (donor programs).

As the leader of the work package on legal and ethical issues, the NBB has developed a series of documents that should provide a general ethical framework (on Consortium level) and could function as a guideline on the level of the individual organization (on brain bank level). The NBB used a structure that focuses on globally accepted bioethical principles and international doctrine. For this purpose the NBB formulated a BNE Code of Conduct, which covers basic legal rules and bioethical principles involved in brain banking and is based on various sources available in the field of bioethics. Such sources include laws, regulations and guidelines issued by international governmental and non-governmental key organizations, such as the Council of Europe, the European Commission, the World Medical Association and the World Health Organization. In June 2008, all BNE II partners signed the Code of Conduct. The NBB observes all rules and regulations of the Code of Conduct.

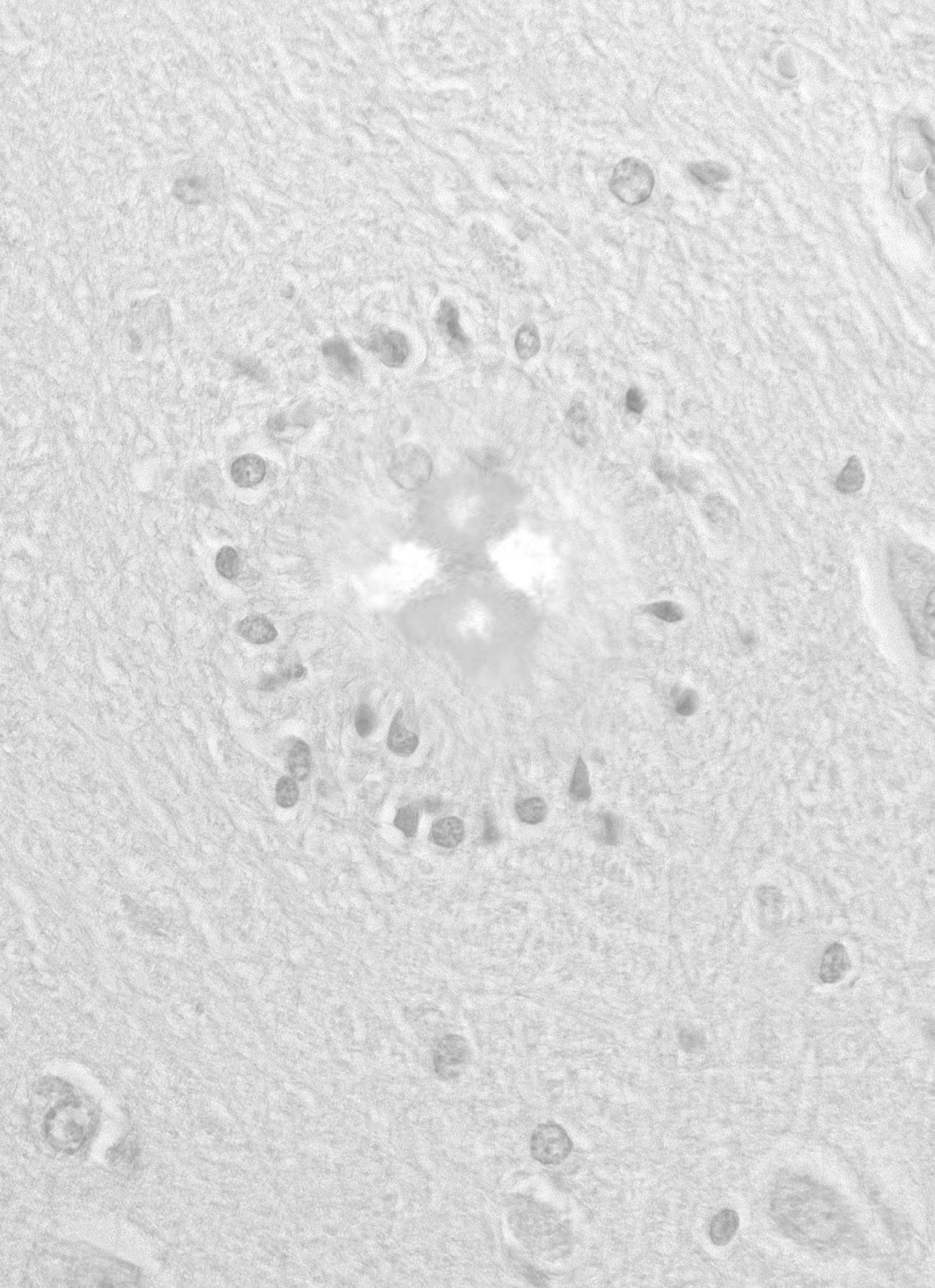
The Code of Conduct addresses fundamental topics, such as the rights of the persons donating their tissue, the obligations of the brain bank with regard to respect and observance of such rights, informed consent, confidentiality, the protection of personal data, the collection and management of human biological material, and transparency and accountability within the organization of a brain bank. As the

Code of Conduct only sets a framework of ground rules and general principles, more concrete guidelines are included in another document called the Brain Bank Regulations. To support the daily practice and ensure compliance with the above-mentioned documents, the NBB has also developed a set of model forms and contracts - indispensable for the daily practice of any well-established brain bank. These forms and contracts include Informed Consent forms, Material Transfer Agreements and Confidentiality Agreements and have been made available to all members of the BNE Consortium. Currently the NBB is preparing a publication on the Code of Conduct.

Publications BNE

- Alafuzoff, I., et al. "Staging of neurofibrillary pathology in Alzheimer's disease: a study of the BrainNet Europe Consortium." *Brain Pathol.* 18.4 (2008): 484-96.
- Alafuzoff, I., et al. "Staging/typing of Lewy body related alpha-synuclein pathology: a study of the BrainNet Europe Consortium." *Acta Neuropathol.* 117.6 (2009): 635-52.
- Alafuzoff, I., et al. "Assessment of alpha-synuclein pathology: a study of the BrainNet Europe Consortium." *J.Neuropathol.Exp.Neurol.* 67.2 (2008): 125-43.
- Alafuzoff, I., et al. "Interlaboratory comparison of assessments of Alzheimer disease-related lesions: a study of the BrainNet Europe Consortium." *J.Neuropathol.Exp.Neurol.* 65.8 (2006): 740-57.
- Alafuzoff, I., et al. "Inter-laboratory comparison of neuropathological assessments of beta-amyloid protein: a study of the BrainNet Europe consortium." *Acta Neuropathol.* 115.5 (2008): 533-46.
- Alafuzoff, I., et al. "Assessment of beta-amyloid deposits in human brain: a study of the BrainNet Europe Consortium." *Acta Neuropathol.* 117.3 (2009): 309-20.
- Bell, J. E., et al. "Management of a twenty-first century brain bank: experience in the BrainNet Europe consortium." *Acta Neuropathol.* 115.5 (2008): 497-507.
- Durrenberger, P. F., et al. "Effects of antemortem and postmortem variables on human brain mRNA quality: a BrainNet Europe study." *J.Neuropathol.Exp.Neurol.* 69.1 (2010): 70-81.
- Ferrer, I., et al. "Effects of Formalin Fixation, Paraffin Embedding, and Time of Storage on DNA Preservation in Brain Tissue: A BrainNet Europe Study." *Brain Pathol.* (2007).
- Grunblatt, E., et al. "Comparison analysis of gene expression patterns between sporadic Alzheimer's and Parkinson's disease." *J.Alzheimers.Dis.* 12.4 (2007): 291-311.
- Huitinga, I., Rademaker, M. and Klioueva, N.. The art of brain banking in Europe: ethical, legal and practical guidelines for donor recruitment, tissue handling and tissue distribution. Abstract in *J. Neural Transm.* 115 (2008), 1715.
- Jacob, C. P., et al. "Alterations in expression of glutamatergic transporters and receptors in sporadic Alzheimer's disease." *J.Alzheimers.Dis.* 11.1 (2007): 97-116.
- Kretschmar, H. "Brain banking: opportunities, challenges and meaning for the future." *Nat. Rev.Neurosci.* 10.1 (2009): 70-78.
- Mackenzie, I. R., et al. "Nomenclature for neuropathologic subtypes of frontotemporal lobar

- degeneration: consensus recommendations." *Acta Neuropathol.* 117.1 (2009): 15-18.
- Mackenzie, I. R., et al. "Nomenclature and nosology for neuropathologic subtypes of fronto-temporal lobar degeneration: an update." *Acta Neuropathol.* 119.1 (2010): 1-4.
- Monoranu, C. M., et al. "pH measurement as quality control on human post mortem brain tissue: a study of the BrainNet Europe consortium." *Neuropathol.Appl.Neurobiol.* 35.3 (2009): 329-37.
- Schmitt, A., et al. "How a neuropsychiatric brain bank should be run: a consensus paper of Brainnet Europe II." *J.Neural Transm.* 114.5 (2007): 527-37.



Finances

The NBB receives structural financial support from the KNAW and the NIN, but apart from that it is almost completely dependent upon grants, donations and the financial contributions from researchers who use NBB material.

The “Stichting tot Ondersteuning van de Hersenbank” (Foundation for the Support of the NBB) was founded in 1986 and helps realize the objectives of the NBB by giving financial support. Since January 2008, the foundation has been deemed an ‘Algemeen Nut Beogende Instelling’ (Institution for Public Advancement) by the Dutch Tax Authority. The assets of this Foundation are formed by donations, testamentary dispositions and legacies (Trade Register Amsterdam, S205869).

The work of the NBB would not be possible without the support of numerous foundations, patient organizations, and the enthusiastic dedication of private individuals.

Grants	2009	2010
Structural contribution of the KNAW	€ 224,321	€ 224,144
Structural contribution of the NIN	€ 100,000	€ 100,000
Stichting MS Research	€ 109,036	€ 106,253
Internationale Stichting Alzheimer Onderzoek	€ 30,000	€ 17,407
Internationaal Parkinson Fonds	€ 25,000	€ 25,000
Hersenstichting Nederland	€ 10,000	€ 10,000

The necessity of grants

Due to the received funding, the NBB is able to continue brain banking. The costs to make tissue available for research are enormous and continue to grow annually. Without the help of patient organizations the NBB would not be able to maintain its high standards.

The Stichting MS Research (www.msresearch.nl) has funded the NBB for many years, resulting in an increase of the number of MS donors and availability of MS tissue. Due to the special MRI-guided dissection protocol, the autopsy costs for MS are higher than for other autopsies. Moreover, the clinical files of people with MS are often more extensive and the summarization of their medical information requires a greater effort. Finally, in-depth neuropathological diagnostics of the MS plaques is time-consuming, but indispensable for good tissue dissemination. MS Research covers the costs of all MS - and some control - autopsies.

The funding of the Internationale Stichting Alzheimer Onderzoek (www.alzheimer.nl) has made it possible for the NBB to produce a new informative DVD, with the objective to raise awareness on the possibility of brain donation for research purposes, and to start up a DNA bank to keep up with the latest developments in research, where genotyping is becoming the important bridge between clinical and neuropathological characteristics.

The grants of the Internationaal Parkinson Fonds (www.parkinsonfonds.nl) cover the costs of a part of the Parkinson autopsies and some donor recruitment activities, which would not be possible without this extra funding.

Funding of the Hersenstichting Nederland (www.hersenstichting.nl) is used to cover donor recruitment, autopsy and administration costs.

Research Projects 2009-2010

The abstracts can be downloaded from our website (www.brainbank.nl)

National

- Aronica, E. et al. Department of Pathology, Academic Medical Centre, Amsterdam. Switch in the expression of mGlu1 and mGlu5 metabotropic glutamate receptors in the cerebellum of mice developing experimental autoimmune encephalomyelitis and in autoptic cerebellar samples from patients with multiple sclerosis.
- Bergen, A. et al. Department of Molecular Ophthalmogenetics, Netherlands Institute for Neuroscience, Amsterdam. The molecular machinery of the choroid plexus in health and disease.
- Bonifati, V. Department of Clinical Genetics, Erasmus MC Rotterdam. Characterization of the FBXO7 (PARK15) protein.
- Borgers, A.J., Alkemade, A. et al. Department of Endocrinology and Metabolism, Academic Medical Centre, Amsterdam. Distribution of serotonin transporters in the human hypothalamus.
- Borgers, A.J., Alkemade, A. et al. Department of Endocrinology and Metabolism, Academic Medical Centre, Amsterdam. Arginine vasopressin immunoreactivity in the suprachiasmatic nucleus is decreased in patients treated for a suprasellar tumor leading to visual field defects.
- Bossers, K. et al. Netherlands Institute for Neuroscience, Amsterdam. Transcriptional alterations during the progression of Alzheimer's disease.
- Dijkstra, A.A., Van de Berg, W.D.J. et al. Department of Anatomy and Neurosciences, Neuroscience Campus Amsterdam, VU University Medical Center, Amsterdam. Canonical pathways involved in the prodromal and motor phase of the Parkinson's disease.
- Garcia-Falgueras, A., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Galanin neurons in the Intermediate nucleus (van Noort et al. 694-703) of the human hypothalamus in relation to sex, age and gender identity.
- García Vallejo, J.J. and Van Kooyk, Y. Departments of Molecular Cell Biology & Immunology, VU University Medical Center, Amsterdam. Glycosylation controls immune homeostasis in the human brain.
- Ishunina, T.A. and Swaab, D.F. Netherlands Institute for Neuroscience, Amsterdam. Estrogen receptor- α splice variants in the Alzheimer's disease brain.
- Kan, A.A., De Graan, P.N.E. et al. Department of Neuroscience & Pharmacology, Rudolf Magnus institute of Neuroscience, Utrecht. Towards unraveling the activated immune system in refractory temporal lobe epilepsy patients.
- Kipp, M., Amor, S. et al. Department of Pathology, VU University Medical Center, Amsterdam. Astrocyte-derived signals to promote myelin formation and repair.

- Klok, M.D., DeRijk, R.H. et al. Division of Medical Pharmacology, Leiden/Amsterdam Center for Drug Research, Leiden University, Leiden. Differential corticosteroid receptor mRNA expression in postmortem brain regions of patients with major depressive disorder.
- Kondova, I. Division of Pathology and Microbiology, Department of Animal Science, Biomedical Primate Research Centre, Rijswijk. Age-related neurological disorders: comparison of brain tissues from humans, chimpanzees and rhesus macaques and exploring the role of miRNAs and small non-coding RNAs (ncRNAs) in the pathogenesis of neurodegeneration.
- Kooi, E., Geurts, J.J.G. et al. Department of Anatomy and Neuroscience, Division of Clinical Neuroscience and Department of Pathology, VU University Medical Center, Amsterdam. Cholinergic imbalance in the multiple sclerosis hippocampus.
- Kreft, K.L., Hintzen, R.Q. et al. Department of Neurology and MS Centre ErasMS, Erasmus University Medical Center, Rotterdam. Genetic determinants of kinesin expression in multiple sclerosis patients.
- Laman, J.D. et al. Department of Immunology, Erasmus medical centre, Rotterdam. Pathogenic mechanisms during multiple sclerosis in the central nervous system and the draining cervical lymph nodes.
- Lucassen, P.J. et al. Center for Neuroscience, Swammerdam Institute of Life Sciences, University of Amsterdam, Amsterdam. Neurogenesis and cellular proliferation in the Alzheimer hippocampus.
- Lucassen, P.J. et al. Center for Neuroscience, Swammerdam Institute of Life Sciences, University of Amsterdam, Amsterdam. Hippocampal neurogenesis during major depression.
- Lucassen, P.J. et al. Center for Neuroscience, Swammerdam Institute of Life Sciences, University of Amsterdam, Amsterdam. Glucocorticoid receptor (GR) in the hippocampus during aging, Alzheimer's disease (AD) and in depression.
- Luchetti, S., Huitinga, I. et al. Department of Neuroimmunology, Netherlands Institute for Neuroscience, Amsterdam. Neurosteroids and multiple sclerosis: lack of synthesis of molecules that mediate neuroprotection and remyelination?
- Luchetti, S., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Neurosteroid biosynthetic pathways changes in prefrontal cortex in Alzheimer's disease.
- Luchetti, S., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Neurosteroid Biosynthetic Pathway Changes in Substantia Nigra and Caudate Nucleus in Parkinson's disease.
- Luchetti, S., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Neurosteroid synthetic pathways in the human brain: differences in relation to age and brain area but not to sex.
- Melief, J., Huitinga, I. et al. Department of Neuroimmunology, Netherlands Institute for Neuroscience, Amsterdam. Isolation and characterization of human post-mortem microglia.
- Middeldorp, J. and Hol, E. Astrocyte Biology & Neurodegeneration, Netherlands Institute for Neuroscience, Amsterdam. Astrocyte subtypes in Alzheimer's disease.

- Nabuurs, R. et al. Departments of Radiology, Pathology and Anatomy, Leiden University Medical Center, Leiden. Histological basis of MRI visualization of AD/CAA in ex vivo human brain tissue.
- Ophoff, R. et al. Department of Psychiatry, Rudolf Magnus Institute, UMC, Utrecht. Genetic studies of schizophrenia: molecular analysis of human brain tissue.
- Prins, M., Van Dam, A. et al. VU University Medical Center, Neuroscience Campus Amsterdam, Amsterdam. Do glia-derived factors determine hippocampal neuronal fate in Parkinson's disease?
- Qi, X., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. The Neurobiology of Depression.
- Reijerkerk, A., De Vries, E. et al. Blood-brain barrier Research Group, Molecular Cell Biology and Immunology, VU University Medical Center, Amsterdam. MicroRNAs in the brain vasculature.
- Riese, H., Niezen-Koning, K. et al. University Medical Center Groningen, Groningen. Comparison of methylation and expression of the serotonin reuptake transporter gene in amygdala tissue, cerebrospinal fluid and peripheral blood.
- Rizzu, P., Heutink, P. et al. Medical Genomics, department of Clinical Genetics, VU Medical Center, Amsterdam. A collection of region specific brain cDNA libraries for verification of the importance of identified SNPs for neurological traits.
- Rozemuller, J.M. et al. Department of Pathology, VU Medical Center Amsterdam. Disease mechanisms in the pathology of Alzheimer's disease and related disorders.
- Scheper, W. et al. Department of Genome Analysis and Department of Neurology, Academic Medical Center and Department of Neuropathology VU Medical Center, Amsterdam. Activation of the unfolded protein response in neurodegenerative tauopathies.
- Schuurman, K. Huitinga, I. et al. Department of Neuroimmunology, Netherlands Institute for Neuroscience, Amsterdam. Analysis of gene expression in MS lesions and normal appearing white matter.
- Shan, L., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. The Histaminergic system neuropsychiatry disorders: a postmortem study.
- Siljee-Wong, J., Alkemade, A. et al. Netherlands Institute for Neuroscience, Amsterdam. Melanocortin-4 Receptor expression in the hypothalamus.
- Temel, Y. et al. Departments of Neuroscience and Neurosurgery, Maastricht University, Maastricht. Serotonin expression in the dorsal raphe nucleus and prefrontal cortex after deep brain stimulation for Parkinson's disease.
- Van den Berge, S. and Hol, E. Astrocyte Biology & Neurodegeneration, Netherlands Institute for Neuroscience, Amsterdam. Neural stem cells in the Parkinson's disease brain.
- Van den Berge, S., Van Strien, M. and Hol, E. Astrocyte Biology & Neurodegeneration, Netherlands Institute for Neuroscience, Amsterdam. Neural stem cells in the adult human brain.
- Van der Star, B., Van der Valk, P. and Amor, S. Department of Pathology, VU University Medical Center, Amsterdam. Axonal damage in MS: engulfment of axons and phagocytosis of neuronal antigens in MS.

- Van Kuppeveld, F. et al. Department of Medical Microbiology, Radboud University Nijmegen Medical Centre, Nijmegen. Role of Saffold virus, a recently identified human cardiovirus, in multiple sclerosis?
- Van Leeuwen, F.W. and Gentier R. Department of Neuroscience, Maastricht University, Maastricht. Proteasomal dysfunction: a way to classify FTD subjects?
- Van Noort, J.M., Amor, S. et al. Department of Pathology, VU University Medical Center, Amsterdam. Alpha B-crystallin is a target for adaptive immune responses, and a trigger of innate responses in preactive multiple sclerosis lesions.
- Van Swieten, J.C. et al. Department of Neurology, Erasmus MC, Rotterdam. Immunohistochemical and biochemical characterization of frontotemporal dementia and progressive supranuclear palsy.
- Van Velzen, M, Verjans, G.M.G.M. et al. Department of Virology, Erasmus MC, Rotterdam. Latent herpes simplex virus and varicella zoster virus infections of sensory neurons of the peripheral nervous system.
- Van Wamelen, D.J., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Hypothalamic changes in Huntington's disease.
- Veerhuis, R. and Hoozemans, J. Department of Clinical chemistry and Alzheimer center and department of Pathology, VU university medical center, Amsterdam. Analysis of mediators of inflammation in Alzheimer's disease.
- Verwer, R.W.H. and Swaab, D.F. Netherlands Institute for Neuroscience, Amsterdam. Reactivation and functional activity of neurons in cultured postmortem brain tissue slices.
- Wilhelmus, M.M.M. et al. Department of Anatomy and Neurosciences, Neuroscience Campus Amsterdam, VU University Medical Center, Amsterdam. Transglutaminases and transglutaminase-catalyzed cross-links colocalize with hyperphosphorylated tau aggregates in tauopathies.
- Wijte, D. et al. Leids Universiteit Medisch Centrum, Leiden. Detection of breakdown products in CSF during the progression of Alzheimer's disease.
- Zhao, J., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Alterations in GABA and glutamate pathways in the prefrontal cortex in major depressive disorder, bipolar disorder and depression in Alzheimer's disease.
- Zhao, T., Swaab, D.F. et al. Netherlands Institute for Neuroscience, Amsterdam. Dendritic cell nuclear protein-1, a novel depression-related protein, upregulates corticotropin-releasing hormone expression.

International

- Andreyeva, A. and Gottmann, K. Institute for Neuro- and Sensory Physiology, University Düsseldorf, Düsseldorf, Germany. Role of N-cadherin in the aging human brain and in Alzheimer's disease.
- Avila, J. et al. Centro de Biología Molecular Severo Ochoa, Madrid, Spain. Expression of somatostatin, dopamine, ghrelin and neurotensin systems is altered in the temporal lobe of Alzheimer disease patients.
- Avila, J. et al. Centro de Biología Molecular Severo Ochoa, Madrid, Spain. Tissue non-specific alkaline phosphatase and muscarinic receptors in the temporal lobe of Alzheimer disease patients.
- Bayer, T. et al. Department of Psychiatry, University Medicine Goettingen, Germany. Intraneuronal Abeta accumulation in Alzheimer's disease.
- Berrocal, M. et al. 'Departamento de Bioquímica y Biología Molecular y Genética, Facultad de Ciencias, University of Extremadura, Badajoz, Spain. Study of tau pathology, amyloid β pathology and calcium pumps in the progression of Alzheimer disease.
- Berson, A., Soreq, H. et al. Department of Biological Chemistry and the Edmond and Lily Safra Center of Neuroscience, The Hebrew University of Jerusalem, Jerusalem, Israel. Global gene expression analysis identifies novel molecular players in neurodegeneration.
- Borea, P.A. and Varani, K. Institute of Pharmacology, University of Ferrara, Italy. A_{2A} adenosine receptor overexpression and functionality, as well as TNF- α , correlate with motor symptoms in Parkinson's disease patients.
- Brust, P. and Deuther-Conrad, W. Helmholtz-Zentrum Dresden-Rossendorf e.V., Leipzig, Germany. Use of human pineal organ for affinity determination of novel ligands for $\alpha_3\beta_4$ nicotinic acetylcholine receptors.
- Chen, X-N. et al. Hefei National Laboratory for Physical Sciences at Microscale, Hefei, China. The Involvement of Retinoic Acid Receptor- α in Corticotropin-Releasing Hormone Gene Expression and Affective Disorders.
- Csiba, L. and Farkas, S. Department of Neurology, University of Debrecen, Debrecen, Hungary. A comparative analysis of expressed and functionally active dopamine receptors in the human brain obtained from Parkinson's disease patients and age matched controls.
- Curtis, M.A. and Graham, S. Centre for Brain Research, Auckland University. Cannabinoid receptor expression in MS lesions.
- Delalle, I. et al. Boston University School of Medicine and Harvard NeuroDiscovery Center, USA. Exosomal and cell-class specific miRNA-profiles in bipolar disorder.
- Farkas, S., Csiba, L. et al. Department of Neurology, University of Debrecen, Debrecen, Hungary. A comparative analysis of expressed and functionally active dopamine receptors in the human brain obtained from Parkinson's disease patients and age matched controls.
- Galter, D. Karolinska Institutet, Department of Neuroscience, Stockholm, Sweden. Study of the mitochondrial serine- protease OMI/HTRA2 in Alzheimer's disease.

- Gao, S., Bao, A. et al. Department of Neurobiology; Key Laboratory of Medical Neurobiology of Ministry of Health of China; Zhejiang Province Key Laboratory of Neurobiology, Zhejiang University School of Medicine, Hangzhou, Zhejiang, P.R. China. Dysregulation of GABAergic neurotransmission in mood disorder: a postmortem study in the hypothalamic paraventricular nucleus and cerebrospinal fluid.
- Giordana, M.T. et al. University of Turin, Turin, Italy. Expression of tumour necrosis factor- α , its receptors (tnfr 1/2), and ask1 in the spinal cord of amyotrophic lateral sclerosis patients.
- Giordana, M.T. et al. University of Turin, Turin, Italy. Characterisation of detergent-insoluble proteins in amyotrophic lateral sclerosis.
- Gulyás B. and Halldin, C. Department of Clinical Neuroscience, Karolinska Institute, Stockholm, Sweden. Autoradiographic examination of mGluR5 in the human brain in Huntington's disease with ^3H and ^{14}C labeled ABP688.
- Hampton, D. et al. The Euan Macdonald Centre for Motor Neurone Disease Research, Centre for Neuroregeneration, Edinburgh, UK. Glutamate re-uptake and recycling and verification of novel transgenic models of FTD.
- Hellings, N. et al. Biomedical Research Institute, Hasselt University and School of Life Sciences, Diepenbeek, Belgium. CX₃CR1 drives cytotoxic CD4+CD28- T cells into the brain of multiple sclerosis patients.
- Hellings, N. et al. Biomedical Research Institute, Hasselt University and School of Life Sciences, Diepenbeek, Belgium. HLA-restricted regulatory CD8 T cells are immunologically altered in multiple sclerosis.
- Ikemoto, K. et al. Department of Neuropsychiatry, Fukushima Medical University School of Medicine, Fukushima, Japan. DNA methylation status of MAOA and MAOB genes in post-mortem brains of patients with schizophrenia.
- Johnston, J. University Belfast, School of Medicine, Dentistry and Biomedical Sciences, Centre for Public Health, RVH ICSB, N. Ireland. Processing of precursor proteins implicated in Alzheimer's disease, Parkinson's disease, and Dementia with Lewy bodies.
- Jones, E. et al. Wolfson Centre for Age-Related Diseases, King's College London, London, UK. Functional Genetic Analysis: Mechanisms of Dementia in People with Down syndrome.
- Kellenberger, S. et al. Department of Pharmacology and Toxicology, University of Lausanne, Lausanne, Switzerland. Analysis of ASIC subunit RNA expression by quantitative RT-PCR.
- Kaiser, C. and Meinl, E. Max-Planck Institute for Neurobiology, Martinsried, Germany. Targets of autoantibodies in MS.
- Klafki, H. et al. Department of Psychiatry and Psychotherapy, University of Duisburg-Essen. Identification of post-translational modifications of the β -amyloid peptides in amyloid plaques.
- Kosmidis, M. Neuroimmunology Laboratory, Athens University Medical School, Greece. JC virus infection in primary and secondary demyelinating diseases of the central nervous system.

- Koutsilieri, E. et al. Institute of Virology and Immunobiology, Wuerzburg, Germany. Role of NMDA receptor subunits in Alzheimer's disease.
- Kravitz, E. and Biegon, A. The Joseph Sagol Neuroscience Center, Sheba Medical Center, Tel Hashomer, Israel. Measuring neuroinflammation by TSPO autoradiography in Alzheimer's disease brains post-mortem.
- Kretzschmar, H. et al. Institute of Neuropathology and Prion Research, Munich, Germany. Genetics and Neuropathology of cases with the diagnosis 'Neurodegeneration with brain iron accumulation (NBIA, formerly Hallervorden-Spatz syndrome)'.
 Li, A. et al. Key Lab for Organ Failure Research, Ministry of Education of China, Nanfang Hospital, Southern Medical University, Guangzhou, Guangdong, China. A descriptive study on the expression of advanced oxidation protein products in different brain areas of Alzheimer disease and vascular dementia.
- Longone, P. et al. Molecular Neurobiology Unit, Fondazione Santa Lucia, Rome, Italy. Molecular and cellular mechanisms of the motor system neurodegenerative pathologies.
- Marcello, E. et al. Department of Pharmacological Sciences and Centre of Excellence on Neurodegenerative Diseases, University of Milan, Italy. SAP97-mediated local trafficking is altered in hippocampus of Alzheimer Disease patients.
- Mohan, H. and Meinel, E. Max-Planck Institute for Neurobiology, Martinsried, Germany. What determines remyelination? Expression profiles of shadow plaques.
- Nielsen, H.M., Wennström, M. et al. Lund University, Dept of Clinical Sciences, Molecular Memory Research Unit, Skåne University Hospital, Malmö, Sweden. Analyses of distribution and activation of NG2-cells in the Alzheimer's brain.
- O'Callaghan, P. et al. Institute for Public Health and Caring Sciences, Uppsala University, Uppsala, Sweden. The fractal dimensions of Ab plaque complexity; an insight to the dynamics of Ab pathology.
- O'Neill, C. et al. Alzheimer's Disease Research Lab, Department of Biochemistry, BioSciences Institute, University College Cork, Cork, Ireland. Pathological signal transduction in Alzheimer's disease: focus of the IGF-I / insulin receptor–Akt pathway, an emerging mechanism central to neurodegeneration in Alzheimer's disease.
- Pavlakis P.P. et al. Department of Pathophysiology, Medical School, University of Athens, Athens, Greece. Autoimmune peripheral neuropathy and Sjögren syndrome: Clinical and Immunological approach.
- Pévet, P. Institut des neurosciences Cellulaire et Intégratives, CNRS et Université de Strasbourg, Strasbourg, France. Structure-function determination of a human brain large non-coding RNA.
- Riederer, P. et al. Clinic and Policlinic of Psychiatry, Psychosomatics and Psychotherapy, University of Wuerzburg, Germany. Neuromelanin of substantia nigra-pathological aspects of Parkinson's disease.

- Rodríguez-Cueto, C. et al. Instituto Universitario de Investigación en Neuroquímica, Department of Biochemistry and Molecular Biology, Faculty of Medicine, Complutense University, Madrid, Spain. Changes in different elements (CB₁ and CB₂ receptors, and FAAH enzyme) of the cannabinoid signalling in post-mortem tissues of patients with spinocerebellar ataxias.
- Somers, V. et al. Biomedical Research Institute, Hasselt University and School of Life Sciences, Diepenbeek, Belgium. Expression profiles of novel MS biomarkers identified via cDNA phage display technology.
- Tofighi, R., Ceccatelli, S. et al. Department of Neuroscience, Karolinska Institutet and Karolinska Hospital, Stockholm, Sweden. Galanin and its receptors in human pituitary adenoma: Evidence for association of GalR₃ with tumor relapse.
- Tsamis, K.I. et al. Laboratory of Neuropathology, 1st Department of Neurology, Aristotle University, Thessaloniki, Greece. Synaptic alterations of human caudate nucleus in Alzheimer's disease.
- Van de Nes, J.A.P. et al. Institute of Pathology and Neuropathology, University Hospital Essen, Essen Germany. Methylation of somatostatin and IST receptor genes in Alzheimer's disease.
- Wang, Y. et al. Institute Of Neuroscience, Shanghai Institutes for Biological Sciences, Shanghai, China. Possible roles of Trpc6 channels in Alzheimer's disease.
- Webb, S. et al. Department of Neurology, Institute of Neurosciences, Southern General Hospital, Glasgow, Scotland. A comparison of viral infections in lymph nodes of patients with multiple sclerosis and normal controls.
- Wennström, M., Nielsen, H.M. et al. Lund University, Dept of Clinical Sciences, Molecular Memory Research Unit, Skåne University Hospital, Malmö, Sweden. Determination of amyloid-associated proteins present in amyloid plaques found in the brains of Dementia with Lewy bodies patients compared to Alzheimer's disease patients.
- Willnow, T.E. et al. Max-Delbrück Center for Molecular Medicine, Berlin, Germany. Identification of Alzheimer's disease risk haplotype that predicts efficiency of *SORL1/SORLA* expression in the brain.
- Yoon, S.-Y. and Kim, D.-H. Department of Anatomy and Cell Biology, University of Ulsan College of Medicine, Seoul, Korea. Search for the key pathogenic molecules in Alzheimer's disease brain.

Pharmaceutical companies

Asterand UK Ltd.

Expression of potential Alzheimer's Disease (AD)-related proteins in FFPE sections of cerebellum from donors with late stage AD.

Expression of therapeutic candidate G-protein coupled receptors in the Dorsal Root Ganglia from male and female donors.

Expression of therapeutic candidate gene in cerebellum from Type 2 Diabetic and control donors.

Bayer Schering Pharma AG

Interrelationship of activated microglia & reactive astrocytes in AD.

Characterization of alpha-synuclein binding molecules.

Characterization of small molecules binding to dementia related pathological targets.

Biopta Ltd.

Studies on Isolated Tissues to Investigate Drug Actions.

GlaxoSmithKline

Identification of potential therapeutic targets for amyotrophic lateral sclerosis and Huntington's disease.

Identification and validation of potential therapeutic targets for multiple sclerosis.

Innogenetics

Isolation of native Tau and PHF Tau from brain material.

Loke Diagnostics ApS

Imaging MS on brain tissue samples from Alzheimer's disease (AD) patients.

Neurimmune Therapeutics AG

Development of novel human antibody therapeutics for the treatment of neurodegenerative disorders such as Alzheimer's disease, Parkinson's disease and amyotrophic lateral sclerosis.

NeuroNova AB

Identification and validation of target protein receptors in brain tissue from Parkinson's patients.

Novartis AG

Localization of S1P receptors in MS brain tissue Examination of MS lesions for macrophage activation and inhibitory markers'

Pfizer Ltd.

Pharmacological profiling of allergy related targets known to be expressed in brain frontal cortex.

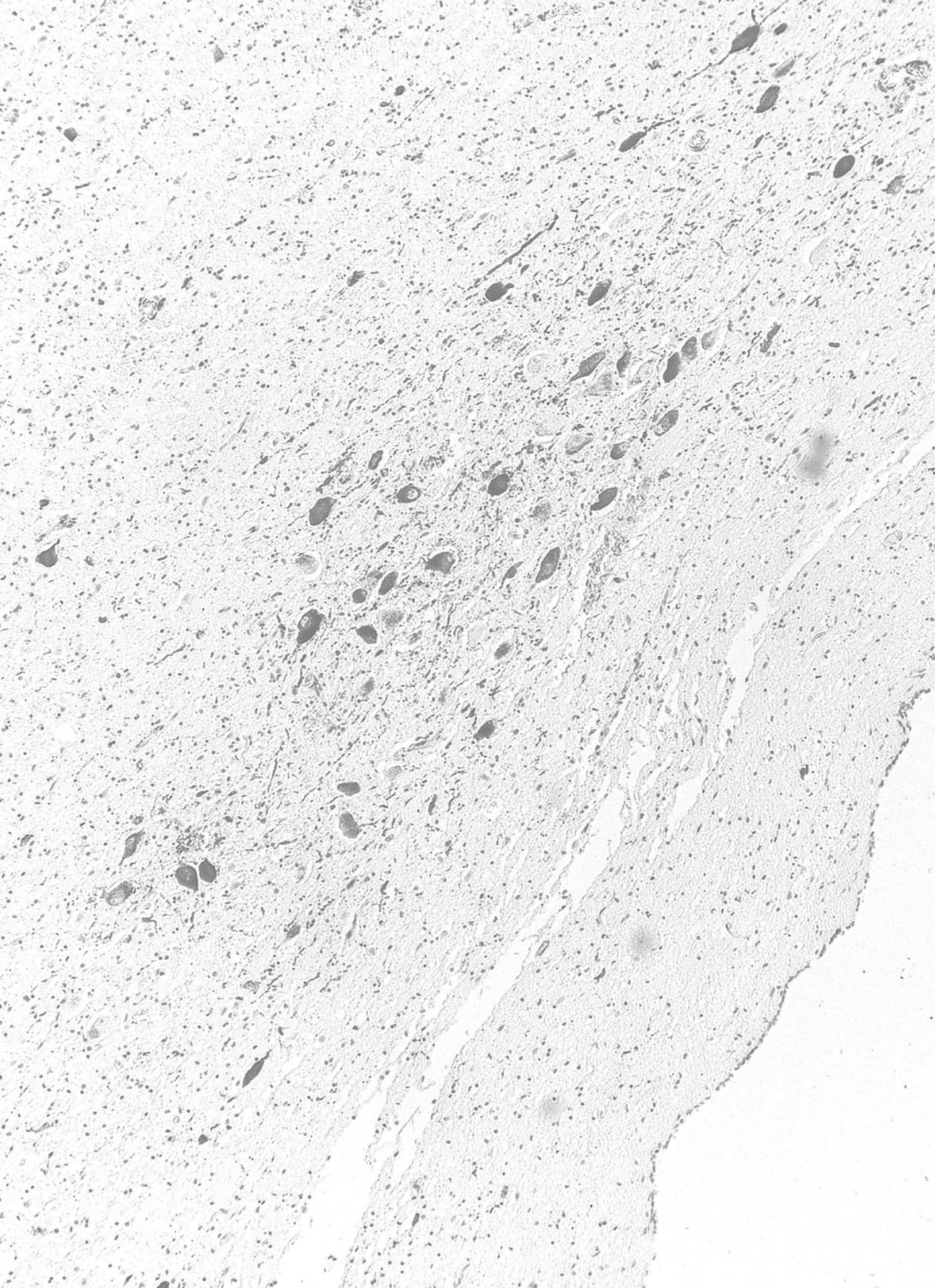
Investigation of the presence and tissue distribution of potential drug targets in human tissue.

Target Characterisation

Assessment of potential urological dysfunction target using human cerebral arteries.

Schering-Plough Biopharma

Analysis of novel proteins and RNA expression in multiple sclerosis.



Publications 2006-2010

The following publications were realized through the use of NBB tissue

- Abildayeva, K. et al. Human apolipoprotein C-I expression in mice impairs learning and memory functions. *J.Lipid Res.* 49.4 (2008): 856-69.
- Alkemade, A. et al. Novel neuroanatomical pathways for thyroid hormone action in the human anterior pituitary. *Eur.J.Endocrinol.* 154.3 (2006): 491-500.
- Alt, S.R. et al. Differential expression of glucocorticoid receptor transcripts in major depressive disorder is not epigenetically programmed. *Psychoneuroendocrinology* 35.4 (2010): 544-56.
- Amor, S. et al. Inflammation in neurodegenerative diseases. *Immunology* 129.2 (2010): 154-69.
- Anand, U. et al. Cannabinoid receptor CB₂ localisation and agonist-mediated inhibition of capsaicin responses in human sensory neurons. *Pain* 138.3 (2008): 667-80.
- Anand, U. et al. The effect of neurotrophic factors on morphology, TRPV₁ expression and capsaicin responses of cultured human DRG sensory neurons. *Neurosci.Lett.* 399.1-2 (2006): 51-56.
- Anand, U. et al. TRPA₁ receptor localisation in the human peripheral nervous system and functional studies in cultured human and rat sensory neurons. *Neurosci.Lett.* 438.2 (2008): 221-27.
- Aziz, A. et al. Hypocretin and melanin-concentrating hormone in patients with Huntington disease. *Brain Pathol.* 18.4 (2008): 474-83.
- Bao, A.M. and Swaab, D.F.. Gender difference in age-related number of corticotropin-releasing hormone-expressing neurons in the human hypothalamic paraventricular nucleus and the role of sex hormones. *Neuroendocrinology* 85.1 (2007): 27-36.
- Bao, A.M. et al. A direct androgenic involvement in the expression of human corticotropin-releasing hormone. *Mol Psychiatry* 2006; 11: 567-76.
- Basso, M. et al. Characterization of detergent-insoluble proteins in ALS indicates a causal link between oxidative stress and aggregation in pathogenesis. *PLoS.One.* 4.12 (2009): e8130.
- Berrocal, M. et al. Altered Ca²⁺ dependence of synaptosomal plasma membrane Ca²⁺-ATPase in human brain affected by Alzheimer's disease. *FASEB J.* 23.6 (2009): 1826-34.
- Berson, A. et al. Changes in readthrough acetylcholinesterase expression modulate amyloid-beta pathology. *Brain* 131 (2008), 109-119.
- Beyer, N. et al. ZnT3 mRNA levels are reduced in Alzheimer's disease post-mortem brain. *Mol. Neurodegener.* 4 (2009): 53.
- Bharathi and Rao, K.S. Molecular understanding of copper and iron interaction with alpha-synuclein by fluorescence analysis. *J.Mol.Neurosci.* 35.3 (2008): 273-81.
- Bharathi et al. A new insight on Al-maltolate-treated aged rabbit as Alzheimer's animal model. *Brain Res.Rev.* 52.2 (2006): 275-92.

- Bharathi, Ravid, R. and Rao, J.K.S. Role of metal in neuronal apoptosis: Challenges associated with neurodegeneration. *Journal of Current Alzheimer Research* 2006; 3.
- Bhardwaj, R.D. et al. Neocortical neurogenesis in humans is restricted to development. *Proc Natl Acad Sci USA* 2006; 103: 12564-8.
- Bo, L. et al. Grey matter pathology in multiple sclerosis. *Acta Neurol Scand Suppl* 2006; 183: 48-50.
- Bo, L. et al. Lack of correlation between cortical demyelination and white matter pathologic changes in multiple sclerosis. *Arch. Neurol.* 64 (2007), 76-80.
- Boekhoorn, K., Joels, M., and Lucassen, P.J. Increased proliferation reflects glial and vascular-associated changes, but not neurogenesis in the presenile Alzheimer hippocampus. *Neurobiol Dis* 2006; 24: 1-14.
- Booij, J.C. et al. The dynamic nature of Bruch's membrane. *Prog. Retin. Eye Res.* 29.1 (2010): 1-18.
- Boor, I. et al. MLC1 is associated with the dystrophin-glycoprotein complex at astrocytic endfeet. *Acta Neuropathol.* 114.4 (2007): 403-10.
- Bossers, K. et al. Analysis of gene expression in Parkinson's disease: possible involvement of neurotrophic support and axon guidance in dopaminergic cell death. *Brain Pathol.* 19.1 (2009): 91-107.
- Bossers, K. et al. Concerted changes in transcripts in the prefrontal cortex precede neuropathology in Alzheimer's disease. *Brain* 133.Pt 12 (2010): 3699-723.
- Bossers, K. et al. Intensity-based analysis of dual-color gene expression data as an alternative to ratio-based analysis to enhance reproducibility. *BMC Genomics* 11 (2010): 112.
- Boven, L.A. et al. Myelin-laden macrophages are anti-inflammatory, consistent with foam cells in multiple sclerosis. *Brain* 2006; 129: 517-26.
- Breij, E.C. et al. Homogeneity of active demyelinating lesions in established multiple sclerosis. *Ann. Neurol.* 63 (2008), 16-25.
- Bronner, I.F. et al. Comprehensive mRNA expression profiling distinguishes tauopathies and identifies shared molecular pathways. *PLoS. One.* 4.8 (2009): e6826.
- Brunner, P. et al. Pineal and cortical melatonin receptors MT1 and MT2 are decreased in Alzheimer's disease. *Eur J Histochem* 2006; 50: 311-6.
- Bruno, M.A. and Cuello, A.C. Activity-dependent release of precursor nerve growth factor, conversion to mature nerve growth factor, and its degradation by a protease cascade. *Proc Natl Acad Sci USA* 2006; 103: 6735-40.
- Bsibsi, M. et al. Identification of soluble CD14 as an endogenous agonist for Toll-like receptor 2 on human astrocytes by genome-scale functional screening of glial cell derived proteins. *Glia* 55 (2007), 473-482.
- Bsibsi, M. et al. The microtubule regulator stathmin is an endogenous protein agonist for TLR3. *J. Immunol.* 184.12 (2010): 6929-37.
- Bsibsi, M. et al. Toll-like receptor 3 on adult human astrocytes triggers production of neuroprotective mediators. *Glia* 2006; 53: 688-95.
- Cao, Y. et al. Changed clathrin regulatory proteins in the brains of Alzheimer's disease patients and animal models. *J. Alzheimers. Dis.* 22.1 (2010): 329-42.

- Chen, X.N. et al. The involvement of retinoic acid receptor-alpha in corticotropin-releasing hormone gene expression and affective disorders. *Biol. Psychiatry* 66.9 (2009): 832-39.
- Chesik, D. et al. Insulin-like growth factor binding proteins: regulation in chronic active plaques in multiple sclerosis and functional analysis of glial cells. *Eur.J.Neurosci.* 24.6 (2006): 1645-52.
- Choi, Y. et al. Minocycline attenuates neuronal cell death and improves cognitive impairment in Alzheimer's disease models. *Neuropsychopharmacology* 32 (2007), 2393-2404.
- Christensen, D.Z. et al. Accumulation of intraneuronal Aβ correlates with ApoE4 genotype. *Acta Neuropathol.* 119.5 (2010): 555-66.
- Cogswell, J.P. et al. Identification of miRNA changes in Alzheimer's disease brain and CSF yields putative biomarkers and insights into disease pathways. *J.Alzheimers.Dis.* 14.1 (2008): 27-41.
- Colsch, B. et al. Sulfogalactosylceramides in motor and psycho-cognitive adult metachromatic leukodystrophy: relations between clinical, biochemical analysis and molecular aspects. *Biochim. Biophys. Acta* 1780 (2008), 434-440.
- Coon, K.D. et al. A high-density whole-genome association study reveals that APOE is the major susceptibility gene for sporadic late-onset Alzheimer's disease. *J.Clin.Psychiatry* 68.4 (2007): 613-18.
- Copani, A. et al. DNA polymerase-beta is expressed early in neurons of Alzheimer's disease brain and is loaded into DNA replication forks in neurons challenged with beta-amyloid. *J.Neurosci.* 26.43 (2006): 10949-57.
- Corneveaux, J.J. et al. Association of CR1, CLU and PICALM with Alzheimer's disease in a cohort of clinically characterized and neuropathologically verified individuals. *Hum. Mol. Genet.* 19.16 (2010): 3295-301.
- Coulson, D.T. et al. BACE1 mRNA expression in Alzheimer's disease post-mortem brain tissue. *J. Alzheimers. Dis.* 22.4 (2010): 1111-22.
- Coulson, D.T. et al. Identification of valid reference genes for the normalization of RT qPCR gene expression data in human brain tissue. *BMC. Mol. Biol.* 9 (2008), 46.
- Couturier, N. et al. Mast cell transcripts are increased within and outside multiple sclerosis lesions. *J. Neuroimmunol.* 195 (2008), 176-185.
- Cuello, A.C. and Bruno, M.A. The Failure in NGF Maturation and its Increased Degradation as the Probable Cause for the Vulnerability of Cholinergic Neurons in Alzheimer's Disease. *Neurochem. Res.* (2007).
- Davidson, Y. et al. Ubiquitinated pathological lesions in frontotemporal lobar degeneration contain the TAR DNA-binding protein, TDP-43. *Acta Neuropathol.* 113.5 (2007): 521-33.
- De Pril, R., Fischer, D.F. and Van Leeuwen, F.W. Conformational diseases: an umbrella for various neurological disorders with an impaired ubiquitin-proteasome system. *Neurobiol.Aging* 27.4 (2006): 515-23.
- De Silva, R. et al. An immunohistochemical study of cases of sporadic and inherited frontotemporal lobar degeneration using 3R- and 4R-specific tau monoclonal antibodies. *Acta Neuropathol (Berl)* 2006; 111: 329-40.

- Diaz-Hernandez, M. et al. Tissue-nonspecific alkaline phosphatase promotes the neurotoxicity effect of extracellular tau. *J. Biol. Chem.* 285.42 (2010): 32539-48.
- Duan, X.H. et al. Novel anilinophthalimide derivatives as potential probes for beta-amyloid plaque in the brain. *Bioorg. Med. Chem.* 18.3 (2010): 1337-43.
- Durrenberger, P.F. et al. Prostanoid receptor EP1 and Cox-2 in injured human nerves and a rat model of nerve injury: a time-course study. *BMC.Neurol.* 6 (2006): 1.
- Eikelenboom, P. et al. Neuroinflammation in plaque and vascular beta-amyloid disorders: clinical and therapeutic implications. *Neurodegener. Dis.* 5 (2008), 190-193.
- Elliott, E., Tsvetkov, P. and Ginzburg, I. BAG-1 associates with Hsc70.Tau complex and regulates the proteasomal degradation of Tau protein. *J.Biol.Chem.* 282.51 (2007): 37276-84.
- Eriksson, M. et al. The NMDAR subunit NR3A interacts with microtubule-associated protein 1S in the brain. *Biochem.Biophys.Res.Comm.* 361.1 (2007): 127-32.
- Facer, P. et al. Differential expression of the capsaicin receptor TRPV1 and related novel receptors TRPV3, TRPV4 and TRPM8 in normal human tissues and changes in traumatic and diabetic neuropathy. *BMC.Neurol.* 7 (2007): 11.
- Familian, A. et al. Inhibitory effect of minocycline on amyloid beta fibril formation and human microglial activation. *Glia* 2006; 53: 233-40.
- Familian, A., Eikelenboom, P. and Veerhuis, R. Minocycline does not affect amyloid beta phagocytosis by human microglial cells. *Neurosci.Lett.* 416.1 (2007): 87-91.
- Farkas S. et al. A comparative analysis of expressed and functionally active dopamine receptors in the human brain obtained from Parkinson's disease patients and age matched controls, *European Journal of Neurology* 2010, 17 (Suppl. 3), 267.
- Farkas S. et al. Functional autoradiography of dopamine D2 receptors in Parkinson's disease on human brain samples. *EME Orvos- és Gyógyszerésztudományi Szakosztálya; XX. Tudományos Ülésszak; Orvostudományi értesítő* 2010;83,1:12-13 (Hungarian).
- Fasano, M., Bergamasco, B. and Lopiano, L. Is neuromelanin changed in Parkinson's disease? Investigations by magnetic spectroscopies. *J.Neural Transm.* 113.6 (2006): 769-74.
- Fasano, M., Bergamasco, B. and Lopiano, L. Modifications of the iron-neuromelanin system in Parkinson's disease. *J.Neurochem.* 96.4 (2006): 909-16.
- Fazio, F. et al. Switch in the expression of mGlu1 and mGlu5 metabotropic glutamate receptors in the cerebellum of mice developing experimental autoimmune encephalomyelitis and in autoptic cerebellar samples from patients with multiple sclerosis. *Neuropharmacology* 55 (2008), 491-499.
- Fliers, E. et al. Hypothalamic thyroid hormone feedback in health and disease. *Prog.Brain Res.* 153 (2006): 189-207.
- Fliers, E., Unmehopa, U.A. and Alkemade, A. Functional neuroanatomy of thyroid hormone feedback in the human hypothalamus and pituitary gland. *Mol.Cell Endocrinol.* 251.1-2 (2006): 1-8.
- Fonfria, E. et al. Tissue distribution profiles of the human TRPM cation channel family. *J.Recept.Signal.Transduct.Res.* 26.3 (2006): 159-78.

- Fonfria, E. et al. TRPM2 is elevated in the tMCAO stroke model, transcriptionally regulated, and functionally expressed in C13 microglia. *J.Recept.Signal.Transduct.Res.* 26.3 (2006): 179-98.
- Gahete, M.D. et al. Expression of Somatostatin, cortistatin, and their receptors, as well as dopamine receptors, but not of neprilysin, are reduced in the temporal lobe of Alzheimer's disease patients. *J. Alzheimers. Dis.* 20.2 (2010): 465-75.
- Gahete, M.D. et al. Expression of the ghrelin and neurotensin systems is altered in the temporal lobe of Alzheimer's disease patients. *J. Alzheimers. Dis.* 22.3 (2010): 819-28.
- Garcia-Falgueras, A. and Swaab, D.F. A sex difference in the hypothalamic uncinate nucleus: relationship to gender identity. *Brain* 131.Pt 12 (2008): 3132-46.
- Geurts, J. J. et al. Does high-field MR imaging improve cortical lesion detection in multiple sclerosis? *J.Neurol.* 255.2 (2008): 183-91.
- Geurts, J.J. and Barkhof, F. (2008). Grey matter pathology in multiple sclerosis. *Lancet Neurol.* 7, 841-851.
- Geurts, J.J. et al. Extensive hippocampal demyelination in multiple sclerosis. *J. Neuropathol. Exp. Neurol.* 66 (2007), 819-827.
- Geurts, J.J. et al. Multiple sclerosis as an "inside-out" disease. *Ann. Neurol.* 68.5 (2010): 767-68.
- Gosso, F.M. et al. Exploring the functional role of the CHRM2 gene in human cognition: results from a dense genotyping and brain expression study. *BMC.Med.Genet.* 8 (2007): 66.
- Goto-Inoue, N. et al. A new lipidomics approach by thin-layer chromatography-blot-matrix-assisted laser desorption/ionization imaging mass spectrometry for analyzing detailed patterns of phospholipid molecular species. *J. Chromatogr. A* 1216.42 (2009): 7096-101.
- Gouw, A.A. et al. Heterogeneity of white matter hyperintensities in Alzheimer's disease: post-mortem quantitative MRI and neuropathology. *Brain* 131 (2008), 3286-3298.
- Govarts, C. et al. Analysis of antibody reactivity in paired cerebrospinal fluid and serum of a relapsing remitting multiple sclerosis patient. *Autoimmunity* 42.8 (2009): 699-704.
- Grunblatt, E. et al. Comparison analysis of gene expression patterns between sporadic Alzheimer's and Parkinson's disease. *J. Alzheimers. Dis.* 12 (2007), 291-311.
- Gupta, V.B., Indi, S.S. and Rao, K.S. Studies on the role of amino acid stereospecificity in amyloid beta aggregation. *J.Mol.Neurosci.* 34.1 (2008): 35-43.
- Ha, T.Y. et al. S100a9 knockdown decreases the memory impairment and the neuropathology in Tg2576 mice, AD animal model. *PLoS.One.* 5.1 (2010): e8840.
- Heesen, C. et al. Stress and hypothalamic-pituitary-adrenal axis function in experimental autoimmune encephalomyelitis and multiple sclerosis - a review. *Psychoneuroendocrinology* 32 (2007a), 604-618.
- Heesen, C. et al. Stress regulation in multiple sclerosis – current issues and concepts. *Multiple sclerosis* 12 (2006) 1-6.
- Heesen, C. et al. Stress regulation in multiple sclerosis: current issues and concepts. *Mult. Scler.* 13 (2007b), 143-148.
- Heidbrink, C. et al. Reduced cortisol levels in cerebrospinal fluid and differential distribution of 11beta-hydroxysteroid dehydrogenases in multiple sclerosis: implications for lesion pathogenesis. *Brain Behav. Immun.* 24.6 (2010): 975-84.

- Hellstrom-Lindahl, E., Viitanen, M. and Marutle, A. Comparison of Abeta levels in the brain of familial and sporadic Alzheimer's disease. *Neurochem. Int.* 55.4 (2009): 243-52.
- Hellstrom-Lindahl, E., Ravid, R. and Nordberg, A. Age-dependent decline of neprilysin in Alzheimer's disease and normal brain: inverse correlation with A beta levels. *Neurobiol. Aging* 29.2 (2008): 210-21.
- Hellstrom-Lindahl, E., Ravid, R., and Nordberg, A. Age-dependent decline of neprilysin in Alzheimer's disease and normal brain: Inverse correlation with Abeta levels. *Neurobiol Aging* 2006.
- Hofman, M.A., and Swaab, D.F. Living by the clock: the circadian pacemaker in older people. *Ageing Res Rev* 2006; 5: 33-51.
- Hoogendijk, W.J. et al. Increased cerebrospinal fluid cortisol level in Alzheimer's disease is not related to depression. *Neurobiol Aging* 2006; 27: 780 e1-780 e2.
- Hoozemans, J.J. et al. Cyclooxygenase-1 and -2 in the different stages of Alzheimer's disease pathology. *Curr. Pharm. Des* 14 (2008), 1419-1427.
- Hoozemans, J.J. et al. The unfolded protein response affects neuronal cell cycle protein expression: implications for Alzheimer's disease pathogenesis. *Exp Gerontol* 2006; 41: 380-6.
- Hoozemans, J.J. et al. The unfolded protein response is activated in pretangle neurons in Alzheimer's disease hippocampus. *Am. J. Pathol.* 174.4 (2009): 1241-51.
- Hou, H.L. et al. Alterations of hHrd1 expression are related to hyperphosphorylated tau in the hippocampus in Alzheimer's disease. *J.Neurosci.Res.* 84.8 (2006): 1862-70.
- Huisman, E., Uylings, H.B. and Hoogland, P.V. Gender-related changes in increase of dopaminergic neurons in the olfactory bulb of Parkinson's disease patients. *Mov Disord.* 23.10 (2008): 1407-13.
- Huitinga I. et al. Characterization of human post mortem microglia isolated during density gradient and FACS sorting. Abstract in: *Glia*, volume 57, issue S13, page S134, October 2009.
- Hull, M. et al. Amyloid beta peptide (25-35) activates protein kinase C leading to cyclooxygenase-2 induction and prostaglandin E2 release in primary midbrain astrocytes. *Neurochem.Int.* 48.8 (2006): 663-72.
- Ikemoto, K. Striatal D-neurons: in new viewpoints for neuropsychiatric research using post-mortem brains. *Fukushima J. Med. Sci.* 54 (2008), 1-3.
- Ikemoto, K. Ventral Tegmental Area. *Japanese Journal of Molecular Psychiatry* 10: 214-218 (2010).
- Ishunina, T.A. and Swaab, D.F. Decreased alternative splicing of estrogen receptor-alpha mRNA in the Alzheimer's disease brain. *Neurobiol.Aging* (2010).
- Ishunina, T.A. and Swaab, D.F. Hippocampal estrogen receptor-alpha splice variant TADDI in the human brain in aging and Alzheimer's disease. *Neuroendocrinology* 89.2 (2009): 187-99.
- Ishunina, T.A. and Swaab, D.F. Age-dependent ERalpha MB1 splice variant expression in discrete areas of the human brain. *Neurobiol. Aging* 29 (2008a), 1177-1189.
- Ishunina, T.A. and Swaab, D.F. Estrogen receptor-alpha splice variants in the human brain. *Gynecol. Endocrinol.* 24 (2008b), 93-98.

- Ishunina, T.A. and Swaab, D.F. Hippocampal Estrogen Receptor-Alpha Splice Variant TADDI in the Human Brain in Aging and Alzheimer's Disease. *Neuroendocrinology* (2008c).
- Ishunina, T.A., Fischer, D.F., and Swaab, D.F. Estrogen receptor alpha and its splice variants in the hippocampus in aging and Alzheimer's disease. *Neurobiol. Aging* 28 (2007), 1670- 1681.
- Ishunina, T.A., Fischer, D.F., and Swaab, D.F. Estrogen receptor alpha and its splice variants in the hippocampus in aging and Alzheimer's disease. *Neurobiol Aging* 2006.
- Jacob, C.P. et al. Alterations in expression of glutamatergic transporters and receptors in sporadic Alzheimer's disease. *J. Alzheimers. Dis.* 11 (2007), 97-116.
- Jamali, S. et al. Large-scale expression study of human mesial temporal lobe epilepsy: evidence for dysregulation of the neurotransmission and complement systems in the entorhinal cortex. *Brain* 2006; 129: 625-41.
- Jansen, C. et al. Prion protein amyloidosis with divergent phenotype associated with two novel nonsense mutations in PRNP. *Acta Neuropathol.* 119.2 (2010): 189-97.
- Junker, A. et al. MicroRNA profiling of multiple sclerosis lesions identifies modulators of the regulatory protein CD47. *Brain* 132.Pt 12 (2009): 3342-52.
- Kaat, L.D. et al. Frontal presentation in progressive supranuclear palsy. *Neurology* 69 (2007), 723-729.
- Kalsbeek, A. et al. Vasopressin and the output of the hypothalamic biological clock. *J. Neuroendocrinol.* (2010).
- Kim, H. S. et al. Swedish amyloid precursor protein mutation increases phosphorylation of eIF2alpha in vitro and in vivo. *J.Neurosci.Res.* 85.7 (2007): 1528-37.
- Klok, M.D. et al. Decreased expression of mineralocorticoid receptor mRNA and its splice variants in post-mortem brain regions of patients with major depressive disorder. *J. Psychiatr. Res.* (2010).
- Koning, N. et al. Distribution of the immune inhibitory molecules CD200 and CD200R in the normal central nervous system and multiple sclerosis lesions suggests neuron-glia and glia-glia interactions. *J. Neuropathol. Exp. Neurol.* 68.2 (2009): 159-67.
- Koning, N. et al. Downregulation of macrophage inhibitory molecules in multiple sclerosis lesions. *Ann. Neurol.* 62 (2007), 504-514.
- Koning, N. et al. Expression of the inhibitory CD200 receptor is associated with alternative macrophage activation. *J. Innate. Immun.* 2.2 (2010): 195-200.
- Koning, N. et al. Restoring immune suppression in the multiple sclerosis brain. *Prog. Neurobiol.* 89.4 (2009): 359-68.
- Kontostavlaki, D.P. et al. Co-expression of tyrosine hydroxylase and GTP cyclohydrolase I in arginine vasopressin-synthesizing neurons of the human supraoptic nucleus demonstrated by laser microdissection and real-time PCR. *Neuroendocrinology* 2006; 84: 386-95.
- Kooi, E.J. et al. Abundant extracellular myelin in the meninges of patients with multiple sclerosis. *Neuropathol. Appl. Neurobiol.* 35.3 (2009): 283-95.
- Kooi, E.J. et al. Meningeal inflammation is not associated with cortical demyelination in chronic multiple sclerosis. *J. Neuropathol. Exp. Neurol.* 68.9 (2009): 1021-28.

- Kooij, G. et al. T lymphocytes impair P-glycoprotein function during neuroinflammation. *J. Autoimmun.* (2009).
- Kovacech, B. et al. A novel monoclonal antibody DC63 reveals that inhibitor 1 of protein phosphatase 2A is preferentially nuclearly localised in human brain. *FEBS Lett.* 581 (2007), 617-622.
- Kravitz, E. and Biegon, A. Measuring neuroinflammation by TSPO autoradiography in Alzheimer's disease brains post-mortem. *Israeli Society for Neuroscience abstracts 2010*, p. 56.
- Kravitz, E. and Biegon, A. Sex and region dependent NMDA receptor (NMDAR) loss and neuroinflammation in Alzheimer's disease brains post-mortem. Program No. 543.26. 2008 Neuroscience Meeting Planner. Washington, DC: Society for Neuroscience, 2008.
- Kravitz, E., Gaisler-Salomon, I. and Biegon, A. Overexpression of gonadal hormone biosynthesis and receptor genes in Alzheimer's disease brains post-mortem Program No. 43.23 Abstract Viewer/Itinerary Planner. Chicago: Society for Neuroscience, 2009.
- Kravitz, E., Gaisler-Salomon, I. and Biegon, A. Overexpression of gonadal hormone biosynthesis and receptor genes in Alzheimer's disease brains post-mortem. *Israeli Society for Neuroscience abstracts 2008*, p. 42.
- Krumbholz, M. et al. CCL19 is constitutively expressed in the CNS, up-regulated in neuroinflammation, active and also inactive multiple sclerosis lesions. *J. Neuroimmunol.* 190 (2007), 72-79.
- Kuhn, H.G. et al. Changes in neurogenesis in dementia and Alzheimer mouse models: are they functionally relevant? *Eur. Arch. Psychiatry Clin. Neurosci.* 257 (2007), 281-289.
- Kuipers, H.F. et al. Simvastatin affects cell motility and actin cytoskeleton distribution of microglia. *Glia* 53.2 (2006): 115-23.
- Kuipers, H.F. et al. CC chemokine receptor 5 gene promoter activation by the cyclic AMP response element binding transcription factor. *Blood* 112 (2008), 1610-1619.
- Kumaran, R. et al. DJ-1 (PARK7) is associated with 3R and 4R tau neuronal and glial inclusions in neurodegenerative disorders. *Neurobiol.Dis.* 28.1 (2007): 122-32.
- Kunii, Y. et al. Fukushima Brain Bank and our results. *Japanese Journal of Biological Psychiatry* 21: 105-112, 2010.
- Liu, C.Q. et al. A quantitative in situ hybridization protocol for formalin-fixed paraffin-embedded archival post-mortem human brain tissue. *Methods* 52.4 (2010): 359-66.
- Liu, R.Y. et al. Glucocorticoids suppress vasopressin gene expression in human suprachiasmatic nucleus. *J. Steroid Biochem. Mol. Biol.* 98 (2006), 248-253.
- Lopez-Aranda, M.F. et al. Localization of the GoLoco motif carrier regulator of G-protein signalling 12 and 14 proteins in monkey and rat brain. *Eur J Neurosci* 2006; 23: 2971-82.
- Lucassen, P.J. et al. Decreased numbers of progenitor cells but no response to antidepressant drugs in the hippocampus of elderly depressed patients. *Neuropharmacology* 58.6 (2010): 940-49.
- Lucassen, P.J. et al. Stress, depression and hippocampal apoptosis. *CNS Neurol Disord Drug Targets* 2006; 5: 531-46.
- Luchetti, S. et al. Neurosteroid biosynthetic pathway changes in substantia nigra and caudate nucleus in Parkinson's disease. *Brain Pathol.* 20.5 (2010): 945-51.

- Luchetti, S. et al. Neurosteroid biosynthetic pathways changes in prefrontal cortex in Alzheimer's disease. *Neurobiol. Aging* (2009).
- Mahad, D. et al. Modulating CCR2 and CCL2 at the blood-brain barrier: relevance for multiple sclerosis pathogenesis. *Brain* 2006; 129: 212-23.
- Maier, O., Baron, W. and Hoekstra, D. Reduced raft-association of NF155 in active MS-lesions is accompanied by the disruption of the paranodal junction. *Glia* 55 (2007a), 885-895.
- Maier, O., Baron, W. and Hoekstra, D. Reduced raft-association of NF155 in active MS-lesions is accompanied by the disruption of the paranodal junction. *Glia* 55 (2007b), 885-895.
- Mali, Y. and Zisapels, N. Gain of interaction of ALS-linked G93A superoxide dismutase with cytosolic malate dehydrogenase. *Neurobiol. Dis.* 32 (2008), 133-141.
- Marcello, E. et al. SAP97-mediated local trafficking is altered in Alzheimer disease patients' hippocampus. *Neurobiol. Aging* (2010).
- Matute, C. P2X7 receptors in oligodendrocytes: a novel target for neuroprotection. *Mol. Neurobiol.* 38 (2008), 123-128.
- Matute, C. et al. Excitotoxic damage to white matter. *J. Anat.* 210 (2007a), 693-702.
- Matute, C. et al. P2X(7) receptor blockade prevents ATP excitotoxicity in oligodendrocytes and ameliorates experimental autoimmune encephalomyelitis. *J. Neurosci.* 27 (2007b), 9525-9533.
- Maubach, K.A. et al. BGC20-1531, a novel, potent and selective prostanoid EP receptor antagonist: a putative new treatment for migraine headache. *Br. J. Pharmacol.* 156.2 (2009): 316-27.
- Medhurst, A.D. et al. GSK189254, a novel H3 receptor antagonist that binds to histamine H3 receptors in Alzheimer's disease brain and improves cognitive performance in preclinical models. *J.Pharmacol.Exp.Ther.* 321.3 (2007): 1032-45.
- Meli, G. et al. Direct in vivo intracellular selection of conformation-sensitive antibody domains targeting Alzheimer's amyloid-beta oligomers. *J. Mol. Biol.* 387.3 (2009): 584-606.
- Melief J. et al. Consequences of stress-axis activity for the severity of multiple sclerosis lesions. Abstract in: *Journal of Neuroimmunology*, volume 203, issue 2, pages 216-217, October 2008.
- Melief J. et al. Severe multiple sclerosis is associated with low stress-axis activity. Abstract in: *Journal of Neural Transmission*, volume 115, issue 12, pages 1718-1718, December 2008.
- Melief, J. et al. Cortisol affects microglia activation status in multiple sclerosis. Abstract in: *Glia*, volume 57, issue S13, page S134, October 2009.
- Meynen, G. et al. Hypothalamic oxytocin mRNA expression and melancholic depression. *Mol.Psychiatry* 12.2 (2007): 118-19.
- Meynen, G. et al. Increased arginine vasopressin mRNA expression in the human hypothalamus in depression: A preliminary report. *Biol Psychiatry* 2006; 60: 892-5.
- Meynen, G. et al. Relation between corticotropin-releasing hormone neuron number in the hypothalamic paraventricular nucleus and depressive state in Alzheimer's disease. *Neuroendocrinology* 85.1 (2007): 37-44.
- Middeldorp, J. Astrocytes in development, aging and disease - Starring GFAP. Thesis, supervisor: Dick Swaab, co-supervisor: Elly Hol. University of Amsterdam (10-06-2010).

- Middeldorp, J. et al. Specific human astrocyte subtype revealed by affinity purified GFAP antibody; unpurified serum cross-reacts with neurofilament-L in Alzheimer. *PLoS.One.* 4.11 (2009): e7663.
- Mohan, H. et al. Extracellular matrix in multiple sclerosis lesions: Fibrillar collagens, biglycan and decorin are upregulated and associated with infiltrating immune cells. *Brain Pathol.* 20.5 (2010): 966-75.
- Moloney, A.M. et al. Defects in IGF-1 receptor, insulin receptor and IRS-1/2 in Alzheimer's disease indicate possible resistance to IGF-1 and insulin signalling. *Neurobiol. Aging* 31.2 (2010): 224-43.
- Moloney, A.M. et al. Defects in IGF-1 receptor, insulin receptor and IRS-1/2 in Alzheimer's disease indicate possible resistance to IGF-1 and insulin signalling. *Neurobiol. Aging* (2008).
- Mulder, S.D. et al. CSF levels of PSA and PSA-ACT complexes in Alzheimer's disease. *Ann. Clin. Biochem.* 46.Pt 6 (2009): 477-83.
- Nabuurs, R.J. et al. High-field MRI of single histological slices using an inductively coupled, self-resonant microcoil: application to ex vivo samples of patients with Alzheimer's disease. *NMR Biomed.* (2010).
- Nielsen, H.M. et al. Astrocytic A beta 1-42 uptake is determined by A beta-aggregation state and the presence of amyloid-associated proteins. *Glia* 58.10 (2010): 1235-46.
- Nielsen, H.M. et al. Binding and uptake of A beta1-42 by primary human astrocytes in vitro. *Glia* 57.9 (2009): 978-88.
- Nilsson, A. et al. Analysis of NR3A receptor subunits in human native NMDA receptors. *Brain Res.* 1186 (2007): 102-12.
- Niwa, S. et al. Post-mortem Brain Studies in Schizophrenia. *Japanese Journal of Clinical Pharmacology* 12: 148-168 (2009).
- O'Callaghan, P. et al. Heparan sulfate accumulation with Abeta deposits in Alzheimer's disease and Tg2576 mice is contributed by glial cells. *Brain Pathol.* 18.4 (2008): 548-61.
- Omari, K.M. et al. Role for CXCR2 and CXCL1 on glia in multiple sclerosis. *Glia* 2006; 53: 24-31.
- Overeem, S. et al. Immunohistochemical screening for autoantibodies against lateral hypothalamic neurons in human narcolepsy. *J.Neuroimmunol.* 174.1-2 (2006): 187-91.
- Pavakis, P.P. et al. Peripheral neuropathies in Sjogren syndrome: a new reappraisal. *J. Neurol. Neurosurg. Psychiatry* (2010).
- Peferoen, L.A. et al. Epstein Barr virus is not a characteristic feature in the central nervous system in established multiple sclerosis. *Brain* 133. Pt 5 (2010): e137.
- Pereira, S. et al. Nuclear localization of a novel human syntaxin 1B isoform. *Gene* 423.2 (2008): 160-71.
- Perez, M. et al. The role of the VQIVYK peptide in tau protein phosphorylation. *J.Neurochem.* 103.4 (2007): 1447-60.
- Perng, M.D. et al. Glial fibrillary acidic protein filaments can tolerate the incorporation of assembly compromised GFAP-delta, but with consequences for filament organization and alphaBcrystallin association. *Mol. Biol. Cell* 19 (2008), 4521-4533.

- Persengiev, S. et al. Genome-wide analysis of miRNA expression reveals a potential role for miR-144 in brain aging and spinocerebellar ataxia pathogenesis. *Neurobiol. Aging* (2010).
- Petanjek, Z. et al. Lifespan alterations of basal dendritic trees of pyramidal neurons in the human prefrontal cortex: a layer-specific pattern. *Cereb.Cortex* 18.4 (2008): 915-29.
- Plumb, J. et al. Upregulation of ADAM-17 expression in active lesions in multiple sclerosis. *Mult.Scler.* 12.4 (2006): 375-85.
- Pollio, G. et al. Increased expression of the oligopeptidase THOP1 is a neuroprotective response to Abeta toxicity. *Neurobiol. Dis.* 31 (2008), 145-158.
- Reiman, E.M. et al. GAB2 alleles modify Alzheimer's risk in APOE epsilon4 carriers. *Neuron* 54.5 (2007): 713-20.
- Reppe, S. et al. Abnormal muscle and hematopoietic gene expression may be important for clinical morbidity in primary hyperparathyroidism. *Am.J.Physiol Endocrinol.Metab* 292.5 (2007): E1465-E1473.
- Robbins, M.J. et al. Evaluation of the mGlu8 receptor as a putative therapeutic target in schizophrenia. *Brain Res.* 1152 (2007): 215-27.
- Roberts, J.C. et al. Autoradiographical imaging of PPARgamma agonist effects on PBR/TSPO binding in TASTPM mice. *Exp. Neurol.* (2009).
- Rogaeva, E. et al. The neuronal sortilin-related receptor SORL1 is genetically associated with Alzheimer disease. *Nat. Genet.* 39 (2007), 168-177.
- Roll, P. et al. SRPX2 mutations in disorders of language cortex and cognition. *Hum Mol Genet* 2006; 15: 1195-207.
- Roos, R.A. and Aziz, N.A. Hypocretin-1 and secondary signs in Huntington's disease. *Parkinsonism.Relat Disord.* 13 Suppl 3 (2007): S387-S390.
- Roses, A.D. et al. A TOMM40 variable-length polymorphism predicts the age of late-onset Alzheimer's disease. *Pharmacogenomics. J.* (2009).
- Royer-Zemmour, B. et al. Epileptic and developmental disorders of the speech cortex: ligand/receptor interaction of wild-type and mutant SRPX2 with the plasminogen activator receptor uPAR. *Hum.Mol.Genet.* 17.23 (2008): 3617-30.
- Rubio, A., Avila, J. and Perez, M. Effect of acetylcholine on tau phosphorylation in human neuroblastoma cells. *J.Mol.Neurosci.* 30.1-2 (2006): 185-88.
- Santa-Mara, I. et al. Coenzyme q induces tau aggregation, tau filaments, and Hirano bodies. *J.Neuropathol.Exp.Neurol.* 67.5 (2008): 428-34.
- Schreibelt, G. et al. Protective effects of peroxiredoxin-1 at the injured blood-brain barrier. *Free Radic.Biol.Med.* 45.3 (2008): 256-64.
- Seabrook, T.J. et al. Angiogenesis is present in experimental autoimmune encephalomyelitis and pro-angiogenic factors are increased in multiple sclerosis lesions. *J. Neuroinflammation.* 7 (2010): 95.
- Seelaar, H. et al. Distinct genetic forms of frontotemporal dementia. *Neurology* 71 (2008), 1220-1226.
- Seelaar, H. et al. TDP-43 pathology in familial frontotemporal dementia and motor neuron disease without Progranulin mutations. *Brain* 130.Pt 5 (2007): 1375-85.

- Seewann, A. et al. Diffusely abnormal white matter in chronic multiple sclerosis: imaging and histopathologic analysis. *Arch. Neurol.* 66.5 (2009): 601-09.
- Seewann, A. et al. Translating pathology in multiple sclerosis: the combination of post-mortem imaging, histopathology and clinical findings. *Acta Neurol. Scand.* 119.6 (2009): 349-55.
- Shen, C. et al. Hydrogen peroxide promotes Abeta production through JNK-dependent activation of gamma-secretase. *J.Biol.Chem.* 283.25 (2008): 17721-30.
- Shiarli, A.M. et al. Comparison of extent of tau pathology in patients with frontotemporal dementia with Parkinsonism linked to chromosome 17 (FTDP-17), frontotemporal lobar degeneration with Pick bodies and early onset Alzheimer's disease. *Neuropathol Appl Neurobiol* 2006; 32: 374-87.
- Stockley, J.H. and O'Neill, C. Understanding BACE1: essential protease for amyloid-beta production in Alzheimer's disease. *Cell Mol. Life Sci.* 65 (2008), 3265-3289.
- Stockley, J.H., Ravid, R. and O'Neill, C. Altered beta-secretase enzyme kinetics and levels of both BACE1 and BACE2 in the Alzheimer's disease brain. *FEBS Lett* 2006; 580: 6550-60.
- Svedberg, M.M. et al. [(11)C]PIB-amyloid binding and levels of Abeta40 and Abeta42 in post-mortem brain tissue from Alzheimer patients. *Neurochem. Int.* 54.5-6 (2009): 347-57.
- Swahn, B.M. et al. Synthesis and evaluation of 2-pyridylbenzothiazole, 2-pyridylbenzoxazole and 2-pyridylbenzofuran derivatives as ¹¹C-PET imaging agents for beta-amyloid plaques. *Bioorg. Med. Chem. Lett.* 20.6 (2010): 1976-80.
- † Hart, B.A., Hintzen, R.Q. and Laman, J.D. Multiple sclerosis - a response-to-damage model. *Trends Mol. Med.* 15.6 (2009): 235-44.
- Teunissen, C.E. et al. Growth-associated protein 43 in lesions and cerebrospinal fluid in multiple sclerosis. *Neuropathol Appl Neurobiol* 2006; 32: 318-31.
- Timmons, S. et al. Akt signal transduction dysfunction in Parkinson's disease. *Neurosci. Lett.* 467.1 (2009): 30-35.
- Toiber, D. et al. N-acetylcholinesterase-induced apoptosis in Alzheimer's disease. *PLoS. ONE.* 3 (2008), e3108.
- Tong, Z. et al. Urine formaldehyde level is inversely correlated to mini mental state examination scores in senile dementia. *Neurobiol. Aging* (2009).
- Torkildsen, O. et al. Upregulation of immunoglobulin-related genes in cortical sections from multiple sclerosis patients. *Brain Pathol.* 20.4 (2010): 720-29.
- Trouw, L.A. et al. C4b-binding protein in Alzheimer's disease: binding to Abeta1-42 and to dead cells. *Mol.Immunol.* 45.13 (2008): 3649-60.
- Ulfman, L.H. et al. Homeostatic intracellular-free Ca²⁺ is permissive for Rap1-mediated constitutive activation of alpha4 integrins on eosinophils. *J.Immunol.* 180.8 (2008): 5512-19.
- Vallejo-Illarramendi, A. et al. Increased expression and function of glutamate transporters in multiple sclerosis. *Neurobiol Dis* 2006; 21:154-64.
- Van de Nes, J.A. et al. Beta-protein/A4 deposits are not associated with hyperphosphorylated tau in somatostatin neurons in the hypothalamus of Alzheimer's disease patients. *Acta Neuropathol.* 111.2 (2006): 126-38.
- Van den Berge, S.A. et al. Longterm quiescent cells in the aged human subventricular neurogenic system specifically express GFAP-delta. *Aging Cell* 9.3 (2010): 313-26.

- Van der Valk, P. and Amor, S. Preactive lesions in multiple sclerosis. *Curr. Opin. Neurol.* 22.3 (2009): 207-13.
- Van Dijk M. et al. The pre-eclampsia gene STOX1 controls a conserved pathway in placenta and brain upregulated in late-onset Alzheimer's disease. *J. Alzheimers. Dis.* 19.2 (2010): 673-79.
- Van Doorn R. et al. Sphingosine 1-phosphate receptor 1 and 3 are upregulated in multiple sclerosis lesions. *Glia* 58.12 (2010): 1465-76.
- Van Eijk M. et al. Differential expression of the EGF-TM7 family members CD97 and EMR2 in lipid-laden macrophages in atherosclerosis, multiple sclerosis and Gaucher disease. *Immunol. Lett.* 129.2 (2010): 64-71.
- Van Horssen J. et al. Nrf2 and DJ1 are consistently upregulated in inflammatory multiple sclerosis lesions. *Free Radic. Biol. Med.* 49.8 (2010): 1283-89.
- Van Horssen, J. et al. Extensive extracellular matrix depositions in active multiple sclerosis lesions. *Neurobiol.Dis.* 24.3 (2006): 484-91.
- Van Horssen, J. et al. Matrix metalloproteinase-19 is highly expressed in active multiple sclerosis lesions. *Neuropathol.Appl.Neurobiol.* 32.6 (2006): 585-93.
- Van Horssen, J. et al. NAD(P)H:quinone oxidoreductase 1 expression in multiple sclerosis lesions. *Free Radic.Biol.Med.* 41.2 (2006): 311-17.
- Van Horssen, J. et al. Severe oxidative damage in multiple sclerosis lesions coincides with enhanced antioxidant enzyme expression. *Free Radic. Biol. Med.* 45 (2008a), 1729-1737.
- Van Horssen, J. et al. The blood-brain barrier in cortical multiple sclerosis lesions. *J.Neuropathol.Exp.Neurol.* 66.4 (2007): 321-28.
- Van Leeuwen, F.W. et al. Frameshift proteins in autosomal dominant forms of Alzheimer disease and other tauopathies. *Neurology* 66.2 Suppl 1 (2006): S86-S92.
- Van Leeuwen, F.W. et al. Molecular misreading: the occurrence of frameshift proteins in different diseases. *Biochem.Soc.Trans.* 34.Pt 5 (2006): 738-42.
- Van Noort, J.M. and Bsibsi, M. Toll-like receptors in the CNS: implications for neurodegeneration and repair. *Prog. Brain Res.* 175 (2009): 139-48.
- Van Noort, J.M. Human glial cell culture models of inflammation in the central nervous system. *Drug Discov. Today* 11 (2006), 74-80.
- Van Noort, J.M. Stress proteins in CNS inflammation. *J. Pathol.* 214 (2008), 267-275.
- Van Noort, J.M. et al. Alfab-crystallin is a target for adaptive immune responses and a trigger of innate responses in preactive multiple sclerosis lesions. *J. Neuropathol. Exp. Neurol.* 69.7 (2010): 694-703.
- Van Noort, J.M. et al. Autoantibodies against alpha B-crystallin, a candidate autoantigen in multiple sclerosis, are part of a normal human immune repertoire. *Mult Scler* 2006; 12: 287-93.
- Van Swieten, J.C. et al. The DeltaK280 mutation in MAP tau favors exon 10 skipping in vivo. *J.Neuropathol.Exp.Neurol.* 66.1 (2007): 17-25.
- Van Swieten, J.C. and Heutink, P. Mutations in progranulin (GRN) within the spectrum of clinical and pathological phenotypes of frontotemporal dementia. *Lancet Neurol.* 7 (2008), 965-974.

- Van Tijn, P. et al. The neuronal ubiquitinproteasome system: murine models and their neurological phenotype. *Prog. Neurobiol.* 85 (2008b), 176-193.
- Van Veen T. et al. CCL5 and CCR5 genotypes modify clinical, radiological and pathological features of multiple sclerosis. *J. Neuroimmunol.* 190.1-2 (2007): 157-64.
- Van Velzen, M. et al. Neuron-interacting satellite glial cells in human trigeminal ganglia have an APC phenotype. *J. Immunol.* 183.4 (2009): 2456-61.
- Van Wamelen, D.J. et al. Functional Increase of Brain Histaminergic Signaling in Huntington's Disease. *Brain Pathol.* (2010).
- Van Zwam M. et al. Myelin ingestion by macrophages promotes their motility and capacity to recruit myeloid cells. *J. Neuroimmunol.* 225.1-2 (2010): 112-17.
- Van Zwam, M. et al. Brain antigens in functionally distinct antigen-presenting cell populations in cervical lymph nodes in MS and EAE. *J. Mol. Med.* 87.3 (2009): 273-86.
- Van Zwam, M. et al. Surgical excision of CNS-draining lymph nodes reduces relapse severity in chronic-relapsing experimental autoimmune encephalomyelitis. *J. Pathol.* 217.4 (2009): 543-51.
- Vanderlocht, J. et al. Leukemia inhibitory factor is produced by myelin-reactive T cells from multiple sclerosis patients and protects against tumor necrosis factor-alpha-induced oligodendrocyte apoptosis. *J.Neurosci.Res.* 83.5 (2006): 763-74.
- Varani, K. et al. A2A adenosine receptor overexpression and functionality, as well as TNF-alpha levels, correlate with motor symptoms in Parkinson's disease. *FASEB J.* 24.2 (2010): 587-98.
- Verheijen, J. H. et al. Detection of a soluble form of BACE-1 in human cerebrospinal fluid by a sensitive activity assay. *Clin.Chem.* 52.6 (2006): 1168-74.
- Verjans, G.M. et al. Selective retention of herpes simplex virus-specific T cells in latently infected human trigeminal ganglia. *Proc. Natl. Acad. Sci. USA* 104.9 (2007): 3496-501.
- Verwer, R.W. et al. Mature astrocytes in the adult human neocortex express the early neuronal marker doublecortin. *Brain* 130.Pt 12 (2007): 3321-35.
- Verwey, N.A. et al. Quantification of amyloid-beta 40 in cerebrospinal fluid. *J. Immunol. Methods* 348.1-2 (2009): 57-66.
- Visser, L. et al. Phagocytes containing a disease-promoting Toll-like receptor/Nod ligand are present in the brain during demyelinating disease in primates. *Am J Pathol* 2006; 169: 1671-85.
- Wang, S.S. et al. Gene expression analysis in the human hypothalamus in depression by laser micro-dissection and real-time PCR: the presence of multiple receptor imbalances. *Mol. Psychiatry* 13 (2008), 786-99, 741.
- Waschbisch, A. et al. Interleukin-1 beta-induced expression of the prostaglandin E-receptor subtype EP3 in U373 astrocytoma cells depends on protein kinase C and nuclear factor-kappaB. *J.Neurochem.* 96.3 (2006): 680-93.
- Webster, J. A. et al. Sor11 as an Alzheimer's disease predisposition gene? *Neurodegener.Dis.* 5.2 (2008): 60-64.
- Westerlund, M. et al. Altered enzymatic activity and allele frequency of OMI/HTRA2 in Alzheimer's disease. *FASEB J.* (2010).

- White, J.H. et al. Identification of a novel asthma susceptibility gene on chromosome 1qter and its functional evaluation. *Hum.Mol.Genet.* 17.13 (2008): 1890-903.
- Wilczak, N. et al. IGF binding protein alterations on periplaque oligodendrocytes in multiple sclerosis: implications for remyelination. *Neurochem.Int.* 52.8 (2008): 1431-35.
- Wilhelmus, M.M. et al. Association of Parkinson disease-related protein PINK1 with Alzheimer disease and multiple sclerosis brain lesions. *Free Radic. Biol. Med.* (2010).
- Wilhelmus, M.M. et al. Novel role of transglutaminase 1 in corpora amylacea formation? *Neurobiol. Aging* (2009).
- Wilhelmus, M.M. et al. Presence of Tissue Transglutaminase in Granular Endoplasmic Reticulum is Characteristic of Melanized Neurons in Parkinson's Disease Brain. *Brain Pathol.* (2010).
- Wilhelmus, M.M. et al. Transglutaminases and Transglutaminase-Catalyzed Cross-Links Colocalize with the Pathological Lesions in Alzheimer's Disease Brain. *Brain Pathol.* (2008).
- Wirhth, O. et al. Identification of low molecular weight pyroglutamate abeta oligomers in Alzheimer disease: a novel tool for therapy and diagnosis. *J. Biol. Chem.* (2010).
- Wirhth, O. et al. Pyroglutamate Abeta pathology in APP/PS1KI mice, sporadic and familial Alzheimer's disease cases. *J. Neural Transm.* 117.1 (2010): 85-96.
- Witte, M.E. et al. Enhanced number and activity of mitochondria in multiple sclerosis lesions. *J. Pathol.* 219.2 (2009): 193-204.
- Witte, M.E. et al. Mitochondrial dysfunction: a potential link between neuroinflammation and neurodegeneration? *Mitochondrion.* 10.5 (2010): 411-18.
- Witte, M.E. et al. Parkinson's disease-associated parkin colocalizes with Alzheimer's disease and multiple sclerosis brain lesions. *Neurobiol. Dis.* 36.3 (2009): 445-52.
- Wu, L. et al. Neural stem cells improve neuronal survival in cultured post mortem brain tissue from aged and Alzheimer patients. *J. Cell Mol. Med.* 12 (2008), 1611-1621.
- Wu, Y.H., Fischer, D.F. and Swaab, D.F. A promoter polymorphism in the monoamine oxidase A gene is associated with the pineal MAOA activity in Alzheimer's disease patients. *Brain Res.* 1167 (2007): 13-19.
- Wu, Y.H. et al. Decreased MT1 melatonin receptor expression in the suprachiasmatic nucleus in aging and Alzheimer's disease. *Neurobiol.Aging* 28.8 (2007): 1239-47.
- Wu, Y.H. et al. Pineal clock gene oscillation is disturbed in Alzheimer's disease, due to functional disconnection from the master clock. *FASEB J.* 20.11 (2006): 1874-76.
- Yiangou, Y. et al. COX-2, CB2 and P2X7-immunoreactivities are increased in activated microglial cells/macrophages of multiple sclerosis and amyotrophic lateral sclerosis spinal cord. *BMC.Neurol.* 6 (2006): 12.
- Zhou, T. et al. Dendritic cell nuclear protein-1, a novel depression-related protein, upregulates corticotropin-releasing hormone expression. *Brain* 133.10 (2010): 3069-79.
- Zhu, H.Y. et al. Increased expression of the Nogo receptor in the hippocampus and its relation to the neuropathology in Alzheimer's disease. *Hum.Pathol.* 38.3 (2007): 426-34.
- Zouambia, M. et al. Proteasome subunit proteins and neuropathology in tauopathies and synucleinopathies: Consequences for proteomic analyses. *Proteomics.* 8.6 (2008): 1221-36.

Staff and Collaborations

Managing director NIN

R. van der Neut

r.van.der.neut@nin.knaw.nl

Director

I. Huitinga

i.huitinga@nin.knaw.nl

Technical coordinator

M. Kooreman

m.kooreman@nin.knaw.nl

Management assistant

M.C. Rademaker

m.rademaker@nin.knaw.nl

Secretariat

P. Brom

p.brom@nin.knaw.nl

Lab technicians

A. van den Berg

a.van.den.berg@nin.nl

P. Evers

p.evers@nin.knaw.nl

Medical writer

C. van Eden

c.van.eden@nin.knaw.nl

Legal advisor

N.M. Klioueva

n.klioueva@nin.knaw.nl

Neuropathologists

J.M. Rozemuller

jm.rozemuller@vumc.nl

Department of Pathology, VUmc

P. van der Valk

p.vandervalk@vumc.nl

Department of Pathology, VUmc

F. van de Goot (until 01-08-2009)

F.van.der.goot@nfi.minjus.nl

Netherlands Forensic Institute, Den Haag

M. Bugiani (starting 01-08-2009)

m.bugiani@vumc.nl

Department of Pathology, VUmc

P. van der Voorn

jp.vandervoorn@vumc.nl

Department of Pathology, VUmc

W. Kamphorst

w.kamphorst@vumc.nl

Department of Pathology, VUmc

Neurologist

C.H. Polman

ch.polman@vumc.nl

(For evaluation of clinical files of MS donors),
Department of Neurology, VUmc

Autopsy team

J. Anink, A. van den Berg, P. Evers, R. Fronczek (until 31-12-2009), S. Hoyng (started 01-08-2009), E. Klerkx, N. Koning (until 01-08-2009), M. Kooreman, J. Korecka, C. Mamber (started 01-10-2010), E. Møst (until 01-09-2010), K. Roet, K. Schuurman, L. Shan (started 01-11-2010), U. Unmehopa, Y. van der Werf.

We owe special thanks to the autopsy assistants of the Department of Pathology of VUmc, Amsterdam, A. van Berkomp, P. Kraaijeveld, T. Oldert and R. Vos, and to John, Frank and Jack of Rouwservice Nederland and the undertakers of Uitvaartcentrum Zuid for their dedication to the Netherlands Brain Bank.

Scientific Committee

I. Huitinga (NBB)

J. Verhaagen (Netherlands Institute for Neuroscience)

J.M. Rozemuller (Department of Pathology, VUmc)

M. Kooreman (NBB)

Advisory Board

Prof. dr. D.A.J.P. Denys (Psychiatry)

Prof. mr. dr. J.K.M. Gevers (Health Law)

Prof. dr. P. Heutink (Genetics)

Drs. A.A. Keizer (Geriatrics)

Prof. dr. H.P.H. Kremer (Neurology)

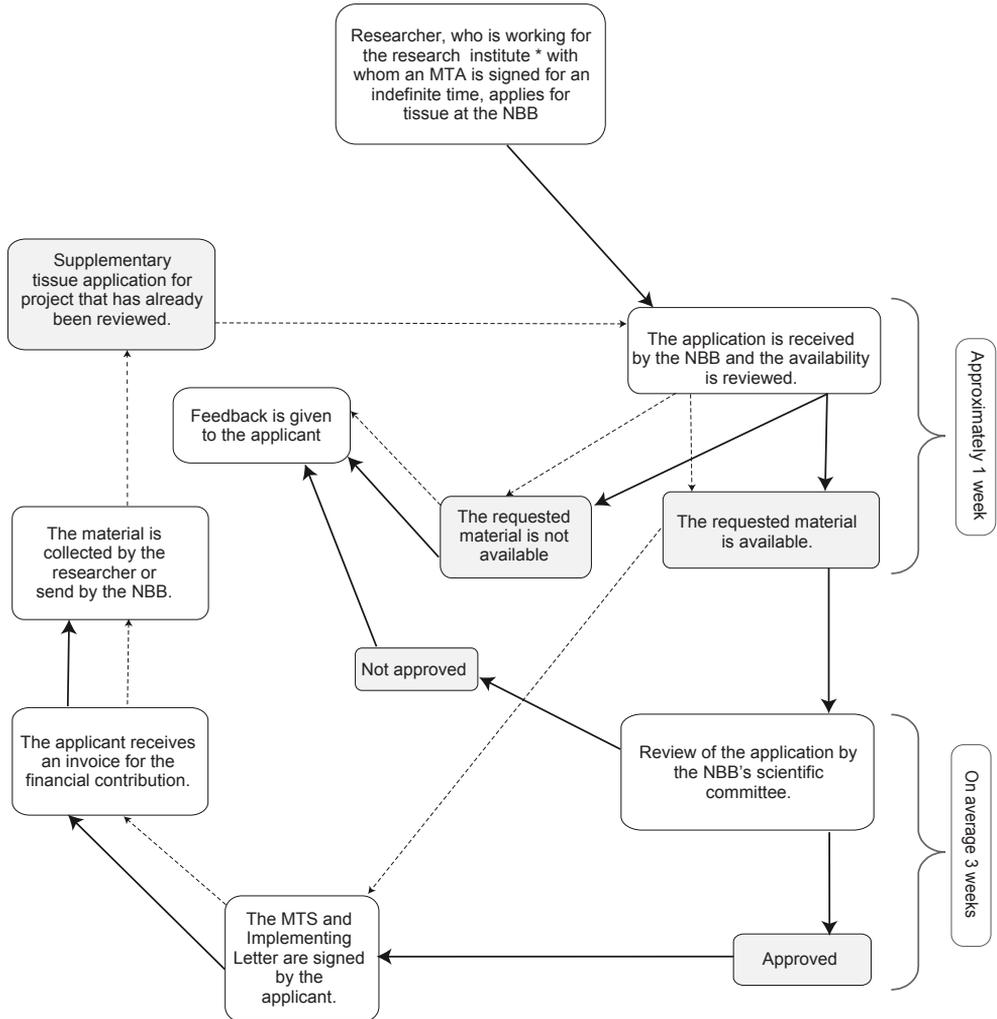
Dr. E. Marchiori (Informatics)

Prof. dr. C.H. Polman (Neurology)

Prof. dr. P. van der Valk (Pathology)

Appendix

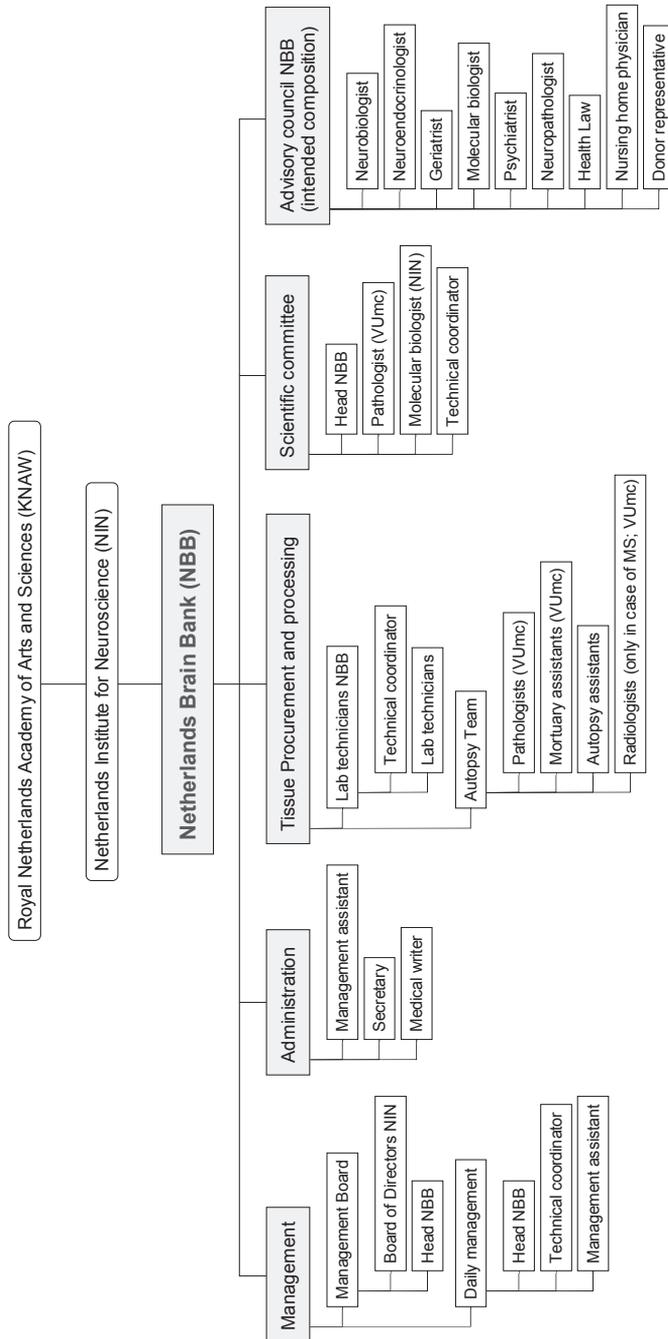
Figure 8 NBB's Procedure of Material Transfer



* The research institute is a legal entity with whom the MTA is signed. Legally, the research institute is thus a party of the agreement. The research institute is thus called "Recipient" of the Material in the MTA and *not* the researcher. In case no MTA for indefinite time has been signed at the institute/organisation where the researcher is working, the NBB will not supply any tissue. First, the authorized person (head manager or managing coordinator) needs to sign the MTA.

———— Application new project:
 - - - - - Supplementary application within reviewed project:

Figure 9 Non-hierarchic scheme of the organization of the Netherlands Brain Bank (NBB)



Abbreviations

AD	Alzheimer's disease
Contr	Non-demented controls
FTLD/tau	Frontotemporal lobar degeneration/Tauopathy
MS	Multiple sclerosis
Other dem	Other dementia
PANR	Pathological report not ready
PD/DLBD	Parkinson's disease/Diffuse Lewy body dementia
PSP	Progressive supranuclear palsy
Psych	Psychiatric disorders
Rest group	Other diagnoses
Trans	Transsexuality