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11th Annual Symposium of the European LeukemiaNet

Mannheim February 4 - 5, 2014

Coordinator: Prof. Dr. Rüdiger Hehlmann

Summary

The European LeukemiaNet is an EU - initiated research network of excellence with the aims to integrate leukemia research across Europe, to improve the management of leukemia, spread excellence globally and expand knowledge.

The 11th Annual Symposium of the European LeukemiaNet (ELN) was held on February 4 to 5, 2014 in Mannheim, Germany. The ELN Network management center (NMC) organized the scientific programme and provided the operational and organizational infrastructure of the symposium and workpackage (WP) meetings. This includes the scientific program, meeting facilities, catering, accommodations and reimbursement of travel costs. It was a major goal and challenge to get the members of all ELN workpackages face-to-face together. In total, more than 496 participants from 37 countries attended the Symposium including three invited keynote speakers from China and the US. The programme was available for download on the ELN homepage. Traditionally this event is combined with the Annual Symposium of the German Competence Network on Leukemia (KNL), which directly follows the ELN Symposium and in 2014 already took place for the 15th time.

The 11th Annual Symposium of the European LeukemiaNet was supported with grants from the European Science Foundation, the Deutsche Forschungsgemeinschaft, the ELN-Foundation and synergies with KNL. The ELN is grateful for the support.

Scientific content of and discussion at the Symposium

On the first day (**4.2.2014**), all ELN-Workpackages had their meetings. Challenges and new directions in leukemia and related disease entities were highlighted during the workpackage meeting in three



consecutive sessions. Each WP reported on the different projects and discussed future objectives of the group for the coming year.

Session I (11:30 – 13:30) WP4, WP7, WP8

WP4 (chronic myeloid leukemia (CML)) and **WP17** (registry) promoted the spread of statistical standards by forming an ELN statistics committee and providing teaching/workshops on statistics. Progress of the EURO-SKI study was presented. The Russian and Italian registries of pregnancies during CML summarized their current results. The application for HORIZON2020 was discussed and the EURO-SKI study was chosen to supply for a grant. The session closed with the discussion of a new pharmaco-economics study in CML. Background of this study is the loss of patent exclusivity of imatinib in 2017.

WP7 (chronic lymphocytic leukemia (CLL/ ERIC)) reported on their activities during 2013. The ERIC Novel Mutation project was presented as well as new approaches with MRD harmonisation by flow cytometry. The ERIC p53 group presented their ongoing accreditation in TP53 mutation analysis. **WP8** (myelodysplastic syndromes (MDS)) presented the progresses in the ELN EU-MDS registry project, for instance the implementation of IPSS-R, the impact of ESA and of iron chelation. The results of the 6th ELN workshop of flow cytometry in MDS were reported. The trial platform: EMSCO gave an overview and planned trial(s) were discussed. Progress of guidelines for allogeneic SCT in MDS was presented.

Session II (15:00 – 17:00)

WP5 (acute myeloid leukemia (AML)) together with **WP17** (registry) presented a new study on AML patients around 60 and dose response, with high-dose AraC given in younger and older patients. An update of the Volasertib AML study was given, a phase II trial on adult patients with previously untreated AML, ineligible for intensive therapy. A comparative analysis of different post-/remission treatment strategies in younger adults with AML exhibiting an intermediate-risk karyotype and a triple negative genotype were given by the ELN/EBMT group. A proposal of a position paper on biometrical methods addressing the time to treatment bias in analyses comparing conventional chemotherapy with blood stem cell transplantation was presented. Furthermore the group discussed the prevalence and the clinical characteristics of APL. Two additional reports dealt with molecular MRD and flow cytometric MRD in AML.

WP6 (European working group for ALL (EWALL)) discussed the prognostic impact of recurrent gene deletions in patients with Philadelphia positive ALL transplanted in complete first remission. The



preclinical results of the treatment with PF-114, a highly selective ABL-directed kinase inhibitor, were presented as well as an update of the IRON II study.

A lecture on CALR mutations was given in **WP9** (chronic myeloproliferative diseases, CMPD) to stimulate new clinical research questions. New ELN treatment recommendations for myelofibrosis based on bone marrow features in patients with myelofibrosis were presented and discussed. The routes to the development of new drug therapies in MPNs were described. An update on the clinical and register studies was given.

Session III (17:30 – 19:30)

WP10 (diagnostics) and **WP12** (minimal residual disease) had a joint meeting and reported on cell cycle, IDH1/2 mutations. **WP10** gave an update on the morphology faculty with 23 members from 18 countries started in 2013. A roundtable about accreditation in flow cytometry followed. **WP12** had two presentations dealing with the NPM1 mutants. **WP11** (cytogenetics) and **WP13** (Next generation sequencing) also organized a joint meeting, starting with a presentation on the evolution of monosomy 7 in myeloid malignancies and the question when the loss of the Y-chromosome is clonal and when not. Reports on the studies regarding next generation sequencing in MDS followed. In their separate session **WP11** discussed the karyotypic complexity in AML and the question whether in ALL the high TP53 mutation frequency is a specific clinical entity. Further topics were the balance condition of the karyotypes in CML and the determination of chromosome banding resolution in tumor cytogenetics. In the end a possible proposal for HORIZON2020 was discussed.

WP13 had a presentation dealing with the analysis of histone ChIP-Seq and new mutations in *JAK2* and *CRLF2*. Further discussions were about the usage of NGS regarding the diagnostic in CLL and MDS.

WP14 (stem cell transplantation) gave an overview on projects concerning the topic HCT vs CT randomized study in elderly patients with AML, and the conditioning regime before transplantation in MDS (RICMAC) study. Information on the Worldwide Network for Blood and Marrow Transplantation (**WBMT**) were given. The Auto/RIC-allo versus Auto in Myeloma NMAM2000 study was presented with information about the role of CR. A further topic was the prospective non-interventional study on allogeneic hematopoietic stem cell transplantation in T-prolymphocytic leukemia (T-PLL). In the end the recommendation for allogeneic and autologous stem cell transplantation in T-PLL, a EBMT/ERIC proposal, was discussed.

ECIL (European Conference on Infections in Leukemia) a joint effort by the ELN, EBMT, EORTC and ICHS gave an update on their work with the topics of management of viral hepatitis, pneumocystis jirovecii infections, antifungal prophylaxis including recommendations.



The joint **ELN/ESF-ELN Steering Committee Meeting (20:30 – 22:30)** followed in the evening on the first day (by invitation only, separate protocol).

On the second day (**5.2.2014**), each workpackage summarized results in a plenary session (8:00 – 10:00).

The scientific symposium started with the presentations of two ELN Merit Awards to PD Dr. Susanne Saußebe (Mannheim) and Dr. Nicola Gökbuget (Frankfurt).

Seven keynote lectures followed with invited speakers from Europe, USA and China. M. Baccarani from Bologna, Italy, reported on the new management recommendations for CML: the balance between practice and research. C. H. Pui from Memphis, USA, gave a lecture on recent advances in ALL. Z. Zheng and W. Chen from Peking, China, gave overviews on Leukemia in children and adults in China. D. Grimwade from London, UK, reported on standardization of molecular APL-monitoring. F. Lo-Coco from Rome, Italy, introduced new standards of cure of APL and D. Hoelzer from Frankfurt, Germany, gave a lecture in MRD-directed treatment of ALL.

The ELN **General Assembly** approved eleven new institutions, so that to date 204 institutions participate from 40 countries. There was one new country (Kyrgyzstan).

The tenth information letter was prepared for the symposium in 2014, highlighting the current progress on projects and listing studies and upcoming meetings. It fosters cooperation amongst network members and informs the public on current topics in leukemia.

Assessment of results and impact of the Symposium on the future direction of the field

The 11th Annual Symposium highlighted upfront European research in leukemia, also reaching out to other continents like the US and Asia. Leukemia is a rare disease and European clinical trials are important to gain a broad patient collective, to discuss and compare results and offer optimal treatment for patients. European harmonisation efforts are of major importance for progress in leukemias worldwide. Important activities include consensus decisions in clinical study endpoints, the set up of patient registries for all leukemias, common standardisation procedures and classification systems in



diagnosis and follow up (molecular monitoring, cytogenetics, minimal residual disease assessment), but also harmonisation in data evaluation and reporting. Cooperative research is the only way to cure leukemia.

The ELN offers innovative ideas and internationally recognized research. The ELN integrates a large portfolio of new disease markers, novel targets, drugs, drug-combinations and dose-optimisation studies, vaccination approaches and next generation high-throughput technologies developed in a harmonised setting of European collaborations. All WPs and projects are the basis for high quality research, essential for European excellence in the field of leukemia research and cure of leukemia. This impacts patients, health care economics and society and results in faster implementation of best treatment options for all patients across Europe and beyond.

Full list of speakers and participants



States*	Title**	Firstname	Surname	Nationality	Date of Birth	Gender	Department	Institute	University	Organization	Address	Postal Code	Town	Country	Telephone	Fax	Email
Participant	Ms	Maris	Abbigor	n.a.		female						37007	Salamanca	SPAIN			mymarv@usal.es
Participant	Dr.	Elisabetta	Abruzzese	n.a.		female					Piazza dell'UMANesimo 10	4133	Rome	ITALY			elisabetta.abruzzese@uniroma2.it
Participant	Dr.	Nicola	Adams	n.a.		male					Bunsenstr. 7a	82452	Martineried	GERMANY			nicola@labpmm.de
Participant	Dr.	Andreas	Agathangelidis	n.a.		male					Via Olgettina, 58	20132	Milan	ITALY			agathangelidis@gmail.com
Participant	Ms	Anni	Agarholm	n.a.		female					Tage-Hansens Gade 2	3000	Aarhus C	DENMARK			annia@agor.dk
Participant	Dr.	Mridul	Agarwal	n.a.		male							Mannheim	GERMANY			Mridul.Agarwal@medma.uni-heidelberg.de
Participant	Dr.	Sigrid	Aigner	n.a.		female					Neuhofstraße 6	6341	Basar	SWITZERLAND			sigrid.aigner@bms.com
Participant	Dr.	Suzanna	Akiki	n.a.		female					Metkley Park Road, Birmingham	B152TG	Birmingham	UNITED KINGDOM			suzanna.akiki@bwhct.nhs.uk
Participant	Dr.	Andreas	Albuquerque	n.a.		male					Max-Lebeche-Platz 31	81377	München	GERMANY			andreas.albuquerque@mll.com
Participant	Dr.	Emil	Aleksov	n.a.		male					44 Ivan Evtz. Geshov	1612	Sofia	BULGARIA			emil.aleksov@novartis.com
Speaker	Professor	Jane F.	Apperley	n.a.		female					Du Cone Road	W 12 0MN	London	UNITED KINGDOM			japperley@imperial.ac.uk
Participant	Mr	Samy	Arroum	n.a.		male					Rue Pierre Lhonnme		Courbevoie	FRANCE			samy.arroum@novartis.com
Participant	Ms	Natalia	Babkina	n.a.		female					Parenmky ave 19/121	63115	Kyiv	UKRAINE			natalia.babkina@gmail.com, leuk@ukr.net
Speaker	Professor	Michelle	Baccarani	n.a.		female					Via Casilina, 5	00182	Rome	ITALY			michela.baccarani@uniroma2.it
Participant	Professor	Gabriele	Basilocher	n.a.		male					Freiburgstr. 4	3010	Bern	SWITZERLAND			gabriele.basilocher@insel.ch
Participant	Mr	Georgii	Balatsenko	n.a.		male					6 Plovdivsko pole	1000	Sofia	BULGARIA			balatsenko@hotmail.com
Participant	Ms	Imma	Barber	n.a.		female					Avda. Diagonal 615, 3 floor	08028	Barcelona	SPAIN			ibarber@otcuka.es
Speaker	Professor	Tiziano	Barbui	n.a.		male					Largo Barozzi n. 1	24128	Bergamo	ITALY			tbarbui@ospedalsirriuniti.bergamo.it
Participant	Dr.	Katerina	Barnpour	n.a.		female					Nikandrov Papamias Ave. 15	22680	Lefkosia	CYPRUS			
Participant	Professor	Giovanni	Barosi	n.a.		male					Viale Golgi 9	21100	Pavia	ITALY			barosi@zmatteo.pv.it
Participant	Mr	Christoph	Bartenhagen	n.a.		male					Albert-Schweitzer-Campus 1	48144	Münster	GERMANY			Christoph.Bartenhagen@uni-muenster.de
Participant	Ms	Gabriele	Bartsch	n.a.		female							Mannheim	GERMANY			gabriele.bartsch@medma.uni-heidelberg.de
Speaker	Professor	Renato	Bazzan	n.a.		male					Via Pacequale 11	30114	Mestre-Venezia	ITALY			renato.bazzan@ulz.it, ve.it
Participant	Ms	Andrea	Bauer	n.a.		female					Roonstraße 25	30423	Nienberg	GERMANY			andreas.bauer@novartis.com
Speaker	Ms	Anja	Baumgartner	n.a.		female							München	GERMANY			Anja.Baumgartner@med.uni-muenchen.de
Participant	Professor	Ali	Bazarbachi	n.a.		male					P.O. Box 11-0236 Riad El-Solk, Ha	1107 2020	Beirut, Lebanon	LEBANON			bazarbachi@aub.edu.lb
Participant	Dr.	Alia	Bzabanova	n.a.		female					Lev Tolstov 1140	72007	Bishkek	KYRGYZSTAN			bzabanova312@gmail.com
Participant	Dr.	Heiko	Becker	n.a.		male					Hugstetterstr. 55	79106	Freiburg	GERMANY			heiko.becker@uniklinik-freiburg.de
Participant	Ms	Monika	Belickova	n.a.		female						12870	Prague	CZECH REPUBLIC			monika.belickova@uhkt.cz
Participant	Dr.	Amin	Ben Lazoued	n.a.		male					Hopital Nord, Chemin des Bourrel	13315	Marseille	FRANCE			amin.benlazoued@ap-hm.fr
Speaker	Professor	Marie C.	Béné	n.a.		female					3 Avenue de la Foret de Hays	54500	Yandoevres les Noy	FRANCE			mariecbene@gmail.com
Participant	Professor	Zwi	Berneman	n.a.		male					Wilrijkstraat 10	2650	Edegem	BELGIUM			zwi.berneman@uzb.be
Participant	Mr	Ralf	Bieber	n.a.		male							Mannheim	GERMANY			ralf.bieber@medma.uni-heidelberg.de
Participant	Mr	Christoph	Biernacki	n.a.		male					Joseph-Stelmann-Straße 3	50924	Köln	GERMANY			christoph.biernacki@uk-koeln.de
Participant	Dr.	Stephan	Bischoff	n.a.		male					Parkstr. 10 14 48	63004	Heidelberg	GERMANY			stephan.bischoff@t-online.de
Participant	Dr.	Marina	Bischoff	n.a.		female					Dr.-Ottmar-Koller-Strasse 2	55743	Idar-Oberstein	GERMANY			marina.bischoff@io.chg-kliniken.de
Participant	Dr.	Ole W.	Bjerrum	n.a.		male					Blegdamsvej no. 3	2100	Copenhagen	DENMARK			ole.w@bjierrum@rh.region.dk
Participant	Professor	Maquss	Björkholm	n.a.		male					Radiumkemet, pl. 2, Karolinska St	17176	Stockholm	SWEDEN			maquss.bjorkholm@karolinska.se
Participant	Professor	Clara	Bloomfield	n.a.		female					1216 James Cancer Hospital, 300 V		Columbus OH 432	USA			clara.bloomfield@osumc.edu
Participant	Professor	Monica	Bocchia	n.a.		female					Viale Bracci 16	53100	Siena	ITALY			bocchia@unisi.it
Participant	Ms	Daniela	Bodschel	n.a.		female					Arnulfstraße 23	80636	München	GERMANY			daniela.bodschel@bms.com
Participant	Dr.	Andrija	Boqdanovic	n.a.		male					2 Dr. Koste Todorovic str	11000	Belgrade	SERBIA			aboqdan@eunet.rs
Participant	Mr	Uwe	Böhm	n.a.		male					Theodor-Kutzer-Ufer 1-3	68167	Mannheim	GERMANY			uwe.boehm@medma.uni-heidelberg.de
Participant	Dr.	Nicolaz	Bonadies	n.a.		male					Peterzgraben 4	4562	Basel	SWITZERLAND			nicolaz.bonadies@usb.ch
Participant	Dr.	Stefan	Bosbach	n.a.		male					Kreuzmatte 1f	6260	Raiden	SWITZERLAND			stefan.bosbach@sanet.ch
Participant	Dr.	Colin	Bradley	n.a.		male					West Glen Road	PA6 7GU	Houston, Refrews	UNITED KINGDOM			colin.bradley@cmofoundation.org
Participant	Ms	Jutta	Bradtke	n.a.		female					Langhansstr. 10	35332	Gießen	GERMANY			Jutta.Bradtke@paediat.med.uni-giessen.de
Participant	Dr.	Andreas	Brünniger	n.a.		male					Langhansstr. 10	35332	Gießen	GERMANY			andreas.brueunniger@patho.med.uni-giessen.de
Participant	Mr	Jörg	Brenn	n.a.		male					Kastor & Pollux - Platz der Einheit	60327	Frankfurt am Main	GERMANY			joerg.brenn@arid.com
Participant	Dr.	Jana	Brczinová	n.a.		female						128 20	Prague	CZECH REPUBLIC			brca@uhkt.cz
Participant	Professor	Tim Henrik	Brümmendorf	n.a.		male					Pawelstr. 30	52074	Aachen	GERMANY			tbrueummendorf@ukaachen.de
Participant	Professor	Mats	Bruno	n.a.		male					Sahlgrenska University Hospital	41345	Göteborg	SWEDEN			bruno@medfak.gu.se
Participant	Professor	Dieter	Buchheidt	n.a.		male					Theodor-Kutzer-Ufer 1-3	68167	Mannheim	GERMANY			dieter.buchheidt@umm.de
Speaker	Professor	Thomas	Buechner	n.a.		male					Albert-Schweitzer-Campus 1, Geb.	48123	Münster	GERMANY			buechner@uni-muenster.de
Participant	Dr.	Lutz	Bullinger	n.a.		male					Albert-Einstein-Allee 23	63081	München	GERMANY			lutz.bullinger@uniklinik-wlm.de
Participant	Dr.	Horst	Bumbas	n.a.		male					163 Spahid Independenței, sector 5	050098	Bucharest	ROMANIA			horstbumbas@gmail.com
Participant	Professor	Andreas	Burchert	n.a.		male					Baldingerstraße	35033	Münster	GERMANY			burchert@staff.uni-marburg.de
Participant	Dr.	Yolanda	Cabrero	n.a.		female					Gran Via de les Cortes Catalanes, 78	8013	Barcelona	SPAIN			yolanda.cabrero@novartis.com
Participant	Mr	Luz Felipe	Casado	n.a.		male							Virgen de la Salud	SPAIN			
Participant	Dr.	Jaroslav	Cermak	n.a.		male					U nemocnice 1	128 20	Prague	CZECH REPUBLIC			cermak@uhkt.cz
Participant	PD Dr.	Yves	Chalandon	n.a.		male					4 rue Gabrielle-Perrot-Gentil	1211	Geneva	SWITZERLAND			yves.chalandon@hcuge.ch
Participant	Dr.	Ekaterina	Chelysheva	n.a.		female					Novozykovsk pr. 4a	125167	Moscow	RUSSIA			denvs@bk.ru
Speaker	Professor	Wenming	Chen	n.a.		male					Capital Medical University	No 8, Gongti	Beijing	CHINA			wzxw@ccmu.edu.cn
Participant	Ms	Sibina	Chiarotti	n.a.		female							Rome	ITALY			
Participant	Professor	Luisi	Chicco-Bianchi	n.a.		male					Via Gattamelata 64	05128	Perugia	ITALY			Luisi.Chicco-Bianchi@uni-perugia.it
Participant	Dr.	Hans-Heinrich	Christoffers	n.a.		male					Frankfurter Ring 133a	68087	München	GERMANY			h.h.christoff@t-online.de
Participant	Dr.	Laura	Ciccioni	n.a.		female					V.le Oxford, 81	00133	Rome	ITALY			claudia.ciccioni@eptvonline.it
Participant	PD Dr.	Johannes	Clausen	n.a.		male					Müller Hauptstr. 48	5020	Salzburg	AUSTRIA			jo.clausen@zalk.at
Participant	Ms	Celine	Cleary-Goffette	n.a.		female					3 rue Joseph Monier	32500	Rueil Malmaison	FRANCE			celine.cleary-goffette@bms.com
Participant	Ms	Rena	Conti	n.a.		female							Chicago	USA			rconti@chicago.edu
Participant	Dr.	Lucia	Copáková	n.a.		female					Klenova 1	83310	Bratislava	SLOVAKIA			lucia.copakova@nos.sk
Participant	Dr.	Alfredo	Covelli	n.a.		male					Largo Umberto Boccioni 1	21040	Origaio (VA)	ITALY			alfredo.covelli@novartis.com
Participant	Dr.	Ana Manuela	Crisan	n.a.		female					no 258, Fundeni street, sector 2	72437	Bucharest	ROMANIA			cristianamanuela@yahoo.com
Participant	Professor	Florence	Cymbalista	n.a.		female					125 rue de Stalingrad	93003	Bobigny	FRANCE			florence.cymbalista@svc.spp.fr
Participant	Dr.	Samvel	Danielyan	n.a.		male					Muratsan 114	00715	Yerevan	ARMENIA			danielcymel@yahoo.com
Participant	Dr.	Svenja	Daskalov	n.a.		female					Bunsenstr. 7a	82152	Martineried	GERMANY			sdschick@medma.de
Participant	Dr.	Michael	Daskalovic	n.a.		male					Freiburgstrasse	3010	Bern	SWITZERLAND			michael.daskalovic@insel.ch



Participant	Mr	Stephane	de Botton	n.s.		male			114 Rue Edward Vaillant		Villjuif	FRANCE		Stephane.DEBOTTON@gustavroussy.fr
Participant	Professor	Cirmino	de Sousa	n.s.		male			Rua Carlos Kiesel 74, Int 12	130025-020	Campinas	BRAZIL		csrmino@unicamp.br
Speaker	Professor	Theo J.M.	de Witte	n.s.		male			Geert Grooteplein 10	6500 HB	Nijmegen	THE NETHERLANDS		T.deWitte@nmls.ru.nl
Participant	Ms	Cassandra	Dean	n.s.		female			Pettenkoferstr. 22	68169	Mannheim	GERMANY		cassandra.dean@medma.uni-heidelberg.de
Participant	Ms	Sabine	Dean	n.s.		female			Pettenkoferstraße 22	68169	Mannheim	GERMANY		sabine.dean@med3.ma.uni-heidelberg.de
Participant	Ms	Yasna	Deserno	n.s.		female			Liebfrauenstr. 47	52249	Erdweiler	GERMANY		vdcseno@uksachen.de
Participant	Dr.	Irina	Dmytrenko	n.s.		female			av. Pobeda 119/121	03115	Kiev	UKRAINE		louk@ukr.net
Speaker	Professor	Hartmut	Döhner	n.s.		male			Albert-Einstein-Allee 23	83081	Ulm	GERMANY		hartmut.doehner@uniklinik-ulm.de
Participant	PD Dr.	Konstanze	Döhner	n.s.		female			Albert-Einstein-Allee 23	83081	Ulm	GERMANY		konstanze.doehner@uniklinik-ulm.de
Speaker	Professor	Hervé	Dombret	n.s.		male			1 Avenue Claude Vellefaux	75475	Paris Cedex 05	FRANCE		herve.dombret@slz.sp-hop-paris.fr
Participant	Dr.	Michael	Doubek	n.s.		male			Jihlavská 20	63300	Brno	CZECH REPUBLIC		mdoubek@fnbrno.cz
Participant	Ms	Katarina	Dubavova	n.s.		female			Pražska 4	04011	Kozice	SLOVENIA		sombid.vckum@gmail.com
Speaker	Professor	Martin	Dugas	n.s.		male			Albert-Schweitzer-Campus 1, Geb. 1	48143	Münster	GERMANY		dugas@uni-muenster.de
Participant	Ms	Melanie	Dujda	n.s.		female			Route 141 & Henry Clay Road	03115	Wilmington, DE 19	UNITED KINGDOM		mdujda@incvts.com
Participant	Dr.	Irina	Dvornik	n.s.		female			119/121 Porcelan	03115	Kyiv	UKRAINE		lovik@ukr.net
Speaker	Dr.	Hans	Ebert	n.s.		male			Theodor-Stern Kai 7	60530	Frankfurt am Main	GERMANY		h.berth@med.uni-frankfurt.de
Participant	Dr.	Eckhard	Eigendorff	n.s.		male			Erlanger-Alle 101	07740	Jena	GERMANY		eckhard.eigendorff@med.uni-jena.de
Participant	Mr	Philipp	Eißmann	n.s.		male					Gröfeling	GERMANY		
Participant	Dr.	Hakim	El Housni	n.s.		male			808 Route de Lenik	4070	Bruxelles	BELGIUM		helhousn@ulb.ac.be
Participant	Ms	Ibrahim	El Housni	n.s.		male				4056	Basel	SWITZERLAND		ibrahim.elhousni@novartis.com
Participant	Dr.	Ahmet Emre	Eksazan	n.s.		male			Kocamustafapasa	34303	Istanbul	TURKEY		emreeksazan@hotmail.com
Participant	Dr.	Monika E.	Eszink	n.s.		female			Theaterstr. 6	22880	Wedel	GERMANY		m.eszink@meduc.de
Participant	Dr. rer. nat.	Alice	Fabarius	n.s.		female			Pettenkoferstraße 22	68169	Mannheim	GERMANY		alice.fabarius@medma.uni-heidelberg.de
Participant	Dr.	Edgar	Faber	n.s.		male			I.P. Pavlova 6	175 20	Dlomouc	CZECH REPUBLIC		edgar.faber@fnol.cz
Participant	Dr.	Christian	Fabrizi	n.s.		male			Salvador-Allende-Platz 29	07147	Jena	GERMANY		christian.fabrizi@med.uni-jena.de
Participant	Dr.	Paul	Fabry	n.s.		male			Siemenstraße 5B	63263	Berlin	GERMANY		paul.fabry@genzyme.com
Participant	Ms	Sonja	Farber	n.s.		female					Berlin	GERMANY		
Participant	Dr.	Larisa	Fechina	n.s.		female			Serafima Dryubina, 32	620149	Yekaterinburg	RUSSIA		laridrens_oncolog@mail.ru
Speaker	Professor	Pierre	Fenaux	n.s.		male			125 Rue de Stalingrad	93009	Bobigny	FRANCE		pietre.fenaux@svc.aphp.fr
Participant	Ms	Kalitra	Filiovski	n.s.		female			Largo Umberto Boccioni 1	21040	Orioglio / VA	ITALY		kalitra.filiovski@novartis.com
Participant	Dr.	Katerina	Filonenko	n.s.		female			Lomonosova str. 33/43	03022	Kiev	UKRAINE		kfilonenko@yahoo.com
Participant	Mr	Markus	Fischer	n.s.		male			Roonstraße 25	90429	Nürnberg	GERMANY		markus-f.fischer@novartis.com
Participant	Ms	Klaudia	Filmelova	n.s.		female			Galvaniho 17C	82016	Bratislava	SLOVAKIA		kludia.filmelova@medirex.sk
Participant	Dr.	Juliane	Folkerts	n.s.		female			Im Neuenheimer Feld 280	63120	Heidelberg	GERMANY		J.Folkerts@dkfs.de
Participant	Mr	Mikhail	Fominikh	n.s.		male			2-a Sovetskaya 16	191024	St. Petersburg	RUSSIA		m.s.fominikh@gmail.com
Participant	Professor	Michaels	Fontenay	n.s.		female			27, rue du Faubourg Saint-Jacques	75014	Paris	FRANCE		michaels.fontenay@inscm.fr
Participant	Dr.	Frank	Fox	n.s.		male			Milkenstr. 5	40225	Düsseldorf	GERMANY		frankfox@uni-due.de
Participant	Mr	Martin	Fox	n.s.		male			Plaça Europa 17-19	080	Barcelona	SPAIN		martin.fox@barcelonaquasos.com
Participant	PD Dr.	Olivier	Frank	n.s.		male			Roonstraße 25	90429	Nürnberg	GERMANY		olivier.frank@novartis.com
Participant	Dr.	Georg-Nikolaus	Frank	n.s.		male			Johannallee	04103	Leipzig	GERMANY		georg-nikolaus.frank@medizin.uni-leipzig.de
Participant	Dr.	Petra	Fritsche	n.s.		female			Roonstraße 25	90429	Nürnberg	GERMANY		petra.fritsche@novartis.com
Participant	Ms	Sabine	Fröhlich	n.s.		female			Arnulfstraße 29	80636	München	GERMANY		sabine.froehlich@bms.com
Participant	Mr	Michael	Fuchs	n.s.		male			Kerpenerstr. 62	50324	Köln	GERMANY		michael.fuchs@uk-koeln.de
Participant	Ms	Ota	Fuchs	n.s.		female			U Nemocnice 2	12820	Prague	CZECH REPUBLIC		ota.fuchs@uhkt.cz
Participant	Professor	Roland	Fuchs	n.s.		male			Pauwelsstr. 30	52074	Aachen	GERMANY		rofuchs@uksachen.de
Participant	Dr.	Christian	Gahrman	n.s.		male			Süther Str. 3	50396	Köln	GERMANY		gahrman@philanthropy-consulting.eu
Participant	Professor	Gösta	Gahrton	n.s.		male			Huddinge University Hospital	14186	Stockholm	SWEDEN		gosta.gahrton@ki.se
Participant	Ms	Yasna	Gaidzik	n.s.		female					Ulm	GERMANY		
Participant	Professor	Robert	Gale	n.s.		male			11633 San Vicente Blvd. #335	90049	Los Angeles	USA		robertpetraqlo@gmail.com; rqa@celgene.com
Participant	Dr.	Tatyana	Ganeva	n.s.		female			44 Ivan Evst. Gechev	1612	Sofia	BULGARIA		tatyana.ganeva@novartis.com
Speaker	Professor	Arnold	Ganser	n.s.		male			Carl-Neuberg-Str. 1	30625	Hannover	GERMANY		ganser.arnold@mh-hannover.de
Speaker	Dr. nat. techn.	Christina	Ganster	n.s.		female			Robert-Koch-Straße 40	37075	Göttingen	GERMANY		christina.ganster@med.uni-goettingen.de
Participant	Ms	Hege	Garelius	n.s.		female					Göteborg	SWEDEN		
Participant	Professor	Winfried	Gaszmann	n.s.		male			Kampenstr. 51	57072	Siegen	GERMANY		w.gaszmann@marienkrankenhaus.com
Participant	Professor	Stefan	Gattenloehner	n.s.		male			Langhansstr. 10	35392	Gießen	GERMANY		stefan.gattenloehner@patho.med.uni-giessen.de
Participant	Ms	Dagmar	Gehenio	n.s.		female			Beneststraße 1	61352	Bad Homburg	GERMANY		dagmar.gehenio@medpharma.de
Participant	Dr.	Otilia	Georgescu	n.s.		female			258, Fundeni Flood	22328	Bucharest	ROMANIA		georgescu.otilia@yahoo.com
Participant	Professor	Ulrich	Germinq	n.s.		male			Milkenstr. 5	40225	Düsseldorf	GERMANY		germinq@med.uni-duesseldorf.de
Speaker	Dr.	Paolo	Ghile	n.s.		male			Via Casettina 56	20132	Milano	ITALY		paolo.ghile@unimi.it
Speaker	PD Dr.	Aristoteles	Gliganidis	n.s.		male			Rochusstr. 2	40473	Düsseldorf	GERMANY		aristoteles.gliganidis@vhkd-kliniken.de
Participant	Mr	Arthur	Gil	n.s.		male			Marchionniestr. 15	81337	München	GERMANY		gil@fbc.med.uni-muenchen.de
Participant	Dr.	Larisa	Girshova	n.s.		female			Akkuratova str. 2	107341	St. Petersburg	RUSSIA		sqirchov@mail.ru
Participant	Ms	Silke	Gloaguen	n.s.		female			Fetscherstr. 74	01307	Dresden	GERMANY		silke.gloaguen@uniklinikum-dresden.de
Participant	Dr.	Michal	Gniot	n.s.		male			Szamarzewskiego 82/84	60-569	Poznan	POLAND		michal.gniot@zppp.edu.pl
Participant	Mr	Rainer	Göbel	n.s.		male			Eiswaldstraße 27 D	12249	Berlin	GERMANY		rainer.goebel@leukemie-in-berlin.org
Speaker	Dr.	Nicola	Gölkbuget	n.s.		female			Theodor-Stern-Kai 7	60530	Frankfurt am Main	GERMANY		goelkbuget@cm.uni-frankfurt.de
Participant	Mr	Michael	Gorray	n.s.		male			Linkstr. 10	10785	Berlin	GERMANY		michael.gorray@pfizer.com
Participant	Dr.	Dariko	Gozencs	n.s.		male					Mannheim	GERMANY		dariko.gozencs@medma.uni-heidelberg.de
Speaker	Professor	Martin	Grzeschhammer	n.s.		male			Hans-Noltz-Str. 1	32439	Minden	GERMANY		martin.grzeschhammer@muehlenkreiskliniken.de
Participant	Ms	David James	Grimsdale	n.s.		female			8th Floor, Giv's Tower, St. Thomas	SE1 3RT	London	UNITED KINGDOM		david.grimsdale@genetics.kcl.ac.uk
Participant	Professor	Margarita	Ganeva	n.s.		female			6 Plovdivsko pole	1000	Sofia	BULGARIA		margareta@mail.bg
Speaker	Professor	Francois	Guilhot	n.s.		male			2, Rue de la Militrie	86021	Poitiers	FRANCE		f.guilhot@chu-poitiers.fr; fr.guilhot@wanadoo.fr
Participant	Dr.	Joelle	Guilhot	n.s.		female			2, Rue de la Militrie	86021	Poitiers	FRANCE		j.guilhot@chu-poitiers.fr
Participant	Mr	Josef	Gulden	n.s.		male			Kirchstraße 29	82284	Grafsthr	GERMANY		josgul@sol.com
Participant	Mr	Joscha	Guszmán	n.s.		male			Linkstr. 10	10785	Berlin	GERMANY		Joscha.guszmán@pfizer.com
Speaker	Professor	Dietlef	Hasse	n.s.		male			Robert-Koch-Straße 40	37075	Göttingen	GERMANY		hasse.onkologie@med.uni-goettingen.de
Participant	Ms	Sabine	Hasse	n.s.		female			Rochusstr. 2	40473	Düsseldorf	GERMANY		sabine.hasse@vhkd-kliniken.de
Participant	Dr.	Björn	Hackanson	n.s.		male			Hugstetter Str. 55	73106	Freiburg	GERMANY		bjorn.hackanson@uniklinik-freiburg.de



Participant	Dr.	Perttu	Koskenvesa	n.a.	male					Hartmaninkatu 4	00029 HUS	Helsinki	FINLAND			perttu.koskenvesa@helsinki.fi
Participant	Dr. rer. nat.	Ute	Kossak-Roth	n.a.	female					Pettenkoferstraße 22	68163	Mannheim	GERMANY			ute.kossak@medma.uni-heidelberg.de
Participant	Dr.	Kostyantyn	Kotlyarchuk	n.a.	male					45, General Chupryvka str.	73044	Lviv	UKRAINE			hematology_k@yahoo.com
Participant	Dr.	Ute	Kotulla	n.a.	female					Alemannenstraße 14	44263	Dortmund	GERMANY			ute.kotulla@novartis.com
Participant	PD Dr.	Alla	Kovrigina	n.a.	female					Novosilovskii prosad 4	125167	Moscow	RUSSIA			kovrigina.alla@gmail.com
Participant	Dr. rer. nat.	Zhanna	Kosich	n.a.	female					tschisti schosze str 64-34	246012	gomel	BELARUS			janakosi@rambler.ru
Participant	Dr.	Sebastian	Kreil	n.a.	male					Theodor-Kutzer-Ufer 1-3	68167	Mannheim	GERMANY			sebastian.kreil@um.de
Participant	Professor	Hans Heinrich	Kreipe	n.a.	male					Carl-Neuberg-Str. 1	30625	Hannover	GERMANY			kreipe.hans@mh-hannover.de
Participant	Dr.	Dieta	Krisponeit	n.a.	male					Hoyer Str.2	28205	Bremen	GERMANY			dkrisponeit@t-online.de
Participant	Professor	Nicolas	Króger	n.a.	male					Martinistraße 52	20246	Hamburg	GERMANY			nkroeger@uke.uni-hamburg.de
Participant	Mr	Michael	Kühnlein	n.a.	male							Gröfeling	GERMANY			
Participant	Dr.	Sergai	Kulikov	n.a.	male					Novy Zykovskiy pr. 4a	125167	Moscow	RUSSIA			kulikov@blood.ru
Participant	Dipl. Stat.	Marijana	Kurzar-Bocklet	n.a.	female					Farbenstraße 68	65931	Frankfurt am Main	GERMANY			kurzar-bocklet@hotmail.de
Participant	PD Dr.	Hans Michael	Kvasnicka	n.a.	male					Joseph-Stolzmann-Str. 9	50931	Köln	GERMANY			hans-michael.kvasnicka@kqu.de
Participant	Dr.	Edward	Laane	n.a.	male					J. Sütiste tee 19	13419	Tallinn	ESTONIA			edward.laane@rejonaaishaiqls.ee
Participant	Ms	Juliette	Lambert	n.a.	female					177, Rue de Versailles	78157	Le Chesnay	FRANCE			jlambert@ch-versailles.fr
Participant	PD Dr.	Thoralf	Lange	n.a.	male					Numburger Str. 76	06637	Weißenfels	GERMANY			th.lange@asklepios.com
Participant	Professor	William	LaRoche	n.a.	male					15 Commercial Street	06443	Brandford	USA			bill.larochelle@proche.com
Participant	Dr.	Richard	Larson	n.a.	male					5841, S. Maryland Avenue, MC-211	60640 IL 604	Chicago	USA			rlarson@medicine.bsd.uchicago.edu
Participant	Dipl. Stat.	Michael	Lauseker	n.a.	male					Marchionistr. 15	61377	München	GERMANY			lauseker@ibe.med.uni-muenchen.de
Participant	Ms	Natalia	Lazorko	n.a.	female							St. Petersburg	RUSSIA			
Participant	Dr.	Oliver	Leizmann	n.a.	male					Roonstr. 25	90429	Nürnberg	GERMANY			oliver.leizmann@novartis.com
Participant	Ms	Michaels	Leitnerova	n.a.	female					Klekovs 1, Odd. Genetikt	83310	Bratislava	SLOVAKIA			michaels.leitnerova@nou.sk
Speaker	PD Dr.	Eva	Lenfelder	n.a.	female					Theodor-Kutzer-Ufer 1-3	68167	Mannheim	GERMANY			eva.lenfelder@um.de
Participant	Ms	Susanne	Leopold	n.a.	female							Mannheim	GERMANY			susanne.leopold@medma.uni-heidelberg.de
Participant	Dr.	Ke	Lin	n.a.	male					2nd Floor, Duncan Building, Daub	163 3GA	Liverpool	UNITED KINGDOM			k.lin@liverpool.ac.uk
Participant	Dr.	Markus	Lindsuer	n.a.	male					Am Gesundbrunnen 20-26	74078	Heilbronn	GERMANY			markus.lindsuer@stk-kliniken.de
Participant	Dipl.-Inf.	Doris	Lindörfer	n.a.	female					Marchionistr. 15	81377	München	GERMANY			lindorf@ibe.med.uni-muenchen.de
Speaker	Professor	Per	Ljungman	n.a.	male						14186	Stockholm	SWEDEN			per.ljungman@ki.se
Speaker	Professor	Francesco	Lo-Coco	n.a.	male					Viale Oxford 81	103	Rome	ITALY			francesco.lo.coco@uniroma2.it
Participant	Ms	Vera	Lohrbacher	n.a.	female					Rochuszstr. 2	40478	Duisburg	GERMANY			vera.lohrbacher@vmd-kliniken.de
Participant	Dr.	Elena	Lomala	n.a.	female					Akkuratova Str. 2	131341	St. Petersburg	RUSSIA			elomala@rambler.ru
Participant	Professor	Michael	Lübbert	n.a.	male					Hugstetter Straße 55	73106	Freiburg	GERMANY			michael.luebbert@uniklinik-freiburg.de
Participant	Ms	Tina	Lupberger	n.a.	female							Basel	SWITZERLAND			tina.lupberger@novartis.com
Participant	Ms	Insa	Lütge	n.a.	female							Mannheim	GERMANY			insa.lutge@medma.uni-heidelberg.de
Participant	Dr.	Katerina	Machova	n.a.	female					U Nemocnice 1	12820	Prague 2	CZECH REPUBLIC			katerina.machova@uhkt.cz
Participant	Dr.	Rolf	Maier	n.a.	male					Dechant-Decker-Str. 8	52849	Eschweiler	GERMANY			Rdmaier@yahoo.de
Participant	Professor	Francois-Xavier	Mahon	n.a.	male					146, rue Léon Saignat	33076	Bordeaux	FRANCE			francois-xavier.mahon@umr5540.u-bordeaux2.fr
Participant	Dr.	Jitka	Malcikova	n.a.	female					Jihlavská 20	62500	Brno	CZECH REPUBLIC			imalcikova@fnbrno.cz
Participant	Ms	Kalliopi	Manola	n.a.	female					Terma Patriarhou Oresqorion	15310	Athens	GREECE			pmanola@ipta.domokritos.gr
Participant	Dr.	Yuri	Marinov	n.a.	male					U nemocnice 1	12800	Prague	CZECH REPUBLIC			marinov@uhkt.cz; luri.marinov@uhkt.cz
Participant	Mr	Nicolas	Maroc	n.a.	male					163, Avenue de Luminy	13288	Marseille	FRANCE			nicolas.maroc@qisgen.com
Participant	PD Dr.	Hans	Martin	n.a.	male					Theodor-Stern-Kai 7	60530	Frankfurt	GERMANY			h.martin@em.uni-frankfurt.de
Participant	PD	Zoya	Martina	n.a.	female					119/121 Peremsky	03115	Kyiv	UKRAINE			zozymed@gmail.ru
Participant	Professor	Giovanni	Martinielli	n.a.	male					Via Mazzarati 9	40138	Bologna	ITALY			giovanni.martinielli@unibo.it; martg@tin.it
Participant	Dr.	Josefin	Martinez-Lopez	n.a.	female					Avd de Cordoba 2/n	28041	Madrid	SPAIN			martinezlopez@hematologiafoctubre.com
Participant	Dr.	Irina	Martynkevich	n.a.	female					2-ya Sovetskaya 16	191024	St. Petersburg	RUSSIA			iris2307@mail.ru
Participant	Professor	Zvenyolva	Matlyak	n.a.	female					45, General Chupryvka str.	73044	Lviv	UKRAINE			hematology_k@yahoo.com
Participant	Elke	Matzat	n.a.	female						Pettenkoferstraße 22	68163	Mannheim	GERMANY			elke.matzat@medma.uni-heidelberg.de
Participant	Professor	Jiri	Mayer	n.a.	male					Jihlavská 20	625 00	Brno	CZECH REPUBLIC			imayer@fnbrno.cz
Participant	Dr.	Mary Frances	McMullin	n.a.	female					Lisburn Road	BT3 TAB	Belfast	UNITED KINGDOM			m.mcmullin@qub.ac.uk
Participant	Professor	Cristina	Mecucci	n.a.	female					Via Brunamonti Policlinico Montel	6122	Perugia	ITALY			crimscuzzi@unipq.it; cristina.mecucci@unipq.it
Participant	Dr.	Hind	Medyouf	n.a.	female					Im Neuenheimer Feld	6122	Heidelberg	GERMANY			h.medyouf@dkfs.de
Participant	Dr.	Annerose	Meier	n.a.	female					Auguststraße 23	16303	Uckermark	GERMANY			a.meier@asklepios.com
Participant	Ms	Manuela	Meincke	n.a.	female					Roonstraße 25	90429	Nürnberg	GERMANY			manuela.meincke@novartis.com
Participant	Dr.	Drorit	Merkel	n.a.	female					Weizmann Str. 6	54239	Tel Aviv	ISRAEL			Drorit.merkel@sheba.health.gov
Participant	Dr.	Ralf Georg	Meyer	n.a.	male					Lanachbeckstrasse 1	55131	Mainz	GERMANY			qmeyer@uni-mainz.de
Participant	Professor	Jan Jacques	Mihail	n.a.	male					Wjilililstrat 10	62130	Edgemon, Antwerp	GERMANY			qodhsartcenter@upcmail.nl
Participant	Professor	Sergai	Mikhailov	n.a.	male					6 Perviyvok Pole Str.	1756	Sofia	BULGARIA			smikhailov@bio.com; qasombulg@gmail.com
Participant	Dr.	Dana	Mikulekova	n.a.	female					U nemocnice 1	12820	Prague	CZECH REPUBLIC			dana.mikulekova@uhkt.cz
Participant	Ms	Julia	Miroslubova	n.a.	female					str. Botshava Zolmina 23-6	191110	St. Petersburg	RUSSIA			julia_3702@yandex.ru
Participant	Professor	Maehe	Mittelman	n.a.	male					Weizmann Str. 6	64239	Tel Aviv	ISRAEL			moeht@tasmc.health.gov.il
Participant	Dr.	Briqitte	Mohr	n.a.	female					Fetscherstr. 74	01307	Dresden	GERMANY			briqitte.mohr@uniklinikum-dresden.de
Participant	Mr	Pau	Montezinos	n.a.	male							Valencia	SPAIN			
Participant	Mr	Marco	Montillo	n.a.	male							Milano	ITALY			
Speaker	Dr.	Carolina	Moreno	n.a.	female					Avenida de Sant Antoni Maria Clar	8025	Barcelona	SPAIN			cmoreno@zantpau.cat
Participant	Dr.	Elena	Morozeva	n.a.	female					L. Tolstov Str. 6/8		St. Petersburg	RUSSIA			dr_morozeva@mail.ru
Participant	Mr	Maximilian	Mozner	n.a.	male							Mannheim	GERMANY			
Participant	PD Dr.	Dmitry	Motorin	n.a.	male					Akkuratova Str. 2	131341	St. Petersburg	RUSSIA			dmotorin@mail.ru
Participant	Professor	Tariq	Muqbal	n.a.	male					23655 Currant Drive	CO 80401	Boulder-Denver	USA			tmughal311@hotmail.com
Participant	Ms	Barbara	Müller	n.a.	female					Pettenkoferstraße 22	68163	Mannheim	GERMANY			barbara.mueller@medma.uni-heidelberg.de
Participant	Professor	Marin C.	Müller	n.a.	male					Pettenkoferstraße 22	68163	Mannheim	GERMANY			marin.mueller@medma.uni-heidelberg.de
Participant	Dr.	Vasilje	Murceta	n.a.	male					Tectembanj str. 30	2025	Chicago	MOLDAVIA			vmurceta@bio.com; qasombulg@gmail.com
Participant	Dr.	Carl Thomas	Nebc	n.a.	male					Collinistr. 11	68161	Mannheim	GERMANY			thomas.nbc@hscms-labor.de
Participant	Ms	Line	Nederby	n.a.	female					Tage-Hansen Gade 2	8000	Aarhus	DENMARK			lnederby@hi.au.dk



Participant	Ms	Michaela	Neusser	n.a.							München	GERMANY		
Participant	Dr.	Suzanne	Neumann	n.a.							München	GERMANY		zneumann@celgene.com
Participant	Dr.	Michaela	Neusser	n.a.							München	GERMANY		michaela.neusser@med.uni-muenchen.de
Participant	Dr.	Emilia	Niculescu-Mitil	n.a.							Fundeni Road No 256	ROMANIA		nmemilia@gmail.com
Speaker	Professor	Dietger	Niederwieser	n.a.							Johannisallee 32A	GERMANY		dietger@medizin.uni-leipzig.de
Participant	Dr.	Dirk	Niemann	n.a.							Johannes-Müller-Str.	Koblenz	GERMANY	dirk.niemann@stfrunzjakob.de
Participant	Ms	Orlane	Noir	n.a.							5 rue Joseph Mozier	FRANCE		orlane.noir@bms.com
Participant	Mr	Florian	Nolte	n.a.							Pettenkoferstr. 22	Mannheim	GERMANY	florian.nolte@medma.uni-heidelberg.de
Participant	Dr.	Josap F.	Nomdedeu	n.a.							Avenida de Sant Antoni Maria Claret	Barcelona	SPAIN	inomed@eu@hsp.zanpana.es
Participant	Dr.	Daniel	Nowak	n.a.							Pettenkoferstr. 22	Mannheim	GERMANY	daniel.nowak@medma.uni-heidelberg.de
Participant	Dr.	Alexander	Nuszbaum	n.a.							Schulstr. 30A	München	GERMANY	alexander.nuszbaum@riad.com
Participant	Professor	Charlotte	Nyvold	n.a.							Tage-Hansens Gade 2	Aarhus	DENMARK	charlotn@biobase.dk; nyvold@hi.au.dk
Participant	Ms	Claudia	Oancea	n.a.								Frankfurt am Main	GERMANY	
Participant	PD Dr.	Fuat	Oduncu	n.a.							Ziemssenstr. 1	München	GERMANY	Fuat.Oduncu@med.uni-muenchen.de
Speaker	Dr.	Uta	Oelschlaegel	n.a.							Fetscherstr. 74	Dresden	GERMANY	uta.oelschlaegel@uniklinikum-dresden.de
Participant	Dr. rer. nat.	Elizabeth	Oppliger Leibundqu	n.a.							Freiburgstrasse	Bern	SWITZERLAND	elizabeth.oppliger@insel.ch
Participant	Mr	Arturo	Orazi	n.a.								New York	USA	
Participant	Dr.	Kerem	Organ	n.a.							Via V	Origgio / VA	ITALY	kerem.organ@novartis.com
Speaker	Professor	G.J.	Ossenkoppels	n.a.							De Boelelaan 1117, Postbus 7057	1007 MB Amsterdam	THE NETHERLANDS	g.ossenkoppels@vumc.nl
Speaker	Professor	Oliver G.	Ottmann	n.a.							Theodor-Stern-Kai 1	60530 Frankfurt am Main	GERMANY	ottmann@em.uni-frankfurt.de
Participant	Dr.	Pose Ann	Padua	n.a.							11 Ave Claude Vellefaux, Batiment 10	75010 Paris	FRANCE	pose-ann.padua@inserm.fr
Participant	Dr.	Tadej	Pajic	n.a.							Nicoesova 4	1000 Ljubljana	SLOVENIA	tadej.pajic@kclj.si
Participant	Dr.	Niels	Pallisaard	n.a.							Ksbbelroft 25	1100 Visele	DENMARK	Niels.Pallisaard@rsyd.dk
Participant	Dr.	Nikolaos	Papakonstantinou	n.a.							Exochi	Thessaloniki	GREECE	n.papakonstantinou83@gmail.com
Participant	Dr.	Elena	Parovchnikova	n.a.							Novozykovskiy pr.4	125167 Moscow	RUSSIA	pozharokaya@dmcc.ru
Participant	Dr.	Peter	Paschka	n.a.							Albert-Einstein-Allee 23	63081 Ulm	GERMANY	peter.paschka@uniklinik-ulm.de
Participant	Mr	Tomas	Pavlik	n.a.							Kamenice 3	62500 Brno	CZECH REPUBLIC	pvavlik@iba.muni.cz
Participant	Dr.	Andras	Pellagatti	n.a.							Adademic Block, John Radcliff Hos	OX39DU Oxford	UNITED KINGDOM	andrapellagatti@yahoo.co.uk
Participant	Dr.	Julian	Peres Ronco	n.a.							Postfach	4002 Basel	SWITZERLAND	julian.peres_roncio@novartis.com
Participant	Mr	Stefan	Peters	n.a.							Pettenkoferstr. 22	68163 Mannheim	GERMANY	stefanpeters1383@web.de
Speaker	Dr.	Markus	Pfirrmann	n.a.							Marchionistr. 15	81377 München	GERMANY	pfi@ibe.med.uni-muenchen.de
Participant	Dr.	Eckard	Picht	n.a.							Kennedyallee 40	53175 Bonn	GERMANY	Eckard.Picht@dfg.de
Participant	Ms	Viktoria	Plamerova	n.a.							Fetscherstr. 74	01307 Sofia	BULGARIA	
Speaker	Professor	Uwe	Platzbecker	n.a.							Pettenkoferstraße 22	68163 Mannheim	GERMANY	uwe.platzbecker@uniklinikum-dresden.de
Participant	Ms	Regina	Platitsch	n.a.							Pettenkoferstraße 22	68163 Mannheim	GERMANY	regina.platitsch@medma.uni-heidelberg.de
Participant	Ms	Karla	Plevoša	n.a.								Brno	CZECH REPUBLIC	
Participant	Ms	Andrea	Poetsch	n.a.							Pettenkoferstr. 22	68163 Mannheim	GERMANY	Sekretariat.Hellmann@medma.uni-heidelberg.de
Participant	Mr	Jaroslav	Polák	n.a.							U nemocnice 1	12820 Prague 2	CZECH REPUBLIC	jaroslav.polak@uhkt.cz
Participant	Dr.	Liubol	Polushkina	n.a.							2nd Sovetskaya str., 16	191024 St. Petersburg	RUSSIA	polushkina.lb@gmail.com
Participant	Dr.	Marina	Popova	n.a.							Lev Tolstoy 6/8	193022 St. Petersburg	RUSSIA	marina.popova.spb@gmail.com
Participant	Ms	Viola	Pöschel	n.a.								Hamburg	GERMANY	
Participant	Dr.	Sarka	Pospisilova	n.a.							Cernopolni 3	62500 Brno	CZECH REPUBLIC	sarka.pospisilova@fnbrno.cz
Participant	Dr.	Engelbert	Precht	n.a.							Dr. Molewaterplein 50, Suite Ee 151	3015GE Rotterdam	THE NETHERLANDS	E.precht@spr-bio.de
Participant	Dr.	Witold	Przejner	n.a.							ul. Dobinski 7	60211 Gdansk	POLAND	wpre@qumed.edu.pl
Participant	Dr.	Irena	Preložnik Zupan	n.a.							Zaloska C, 7	1000 Ljubljana	SLOVENIA	irena.zupan@kclj.si
Participant	Dr.	Ulrike	Proetel	n.a.							Pettenkoferstraße 22	68163 Mannheim	GERMANY	Ulrike.Proetel@medma.uni-heidelberg.de
Speaker	Dr.	China-Hon	Pui	n.a.							262 Danny Thomas Place	TH 38105-36 Memphis	USA	China-Hon.Pui@STJUDE.ORG
Participant	Dr.	Mari	Punab	n.a.							8 Puvsepa Str.	51014 Tartu	ESTONIA	mari.punab@unilium.ee
Participant	Dr.	Sarka	Rasodorfova	n.a.								128 20 Prague	CZECH REPUBLIC	
Participant	Dr.	Andri	Rawstron	n.a.							Beckett Street	Le3 TF Leeds	UNITED KINGDOM	andru.rawstron@sqw.work.ac.uk
Participant	Professor	Sophie	Ramaud	n.a.							30 Roire Momaine	06002 Nice Cedex 2	FRANCE	ramaud@unice.fr
Participant	Ms	Jana	Reichardt	n.a.								Heidelberg	GERMANY	J.Reichardt@stud.uni-heidelberg.de
Participant	Dr.	Anja	Reichert	n.a.							Edisonstraße 4	85716 Unterschleißheim	GERMANY	anja_reichert@baxter.com
Speaker	Professor	Andreas	Reiter	n.a.							Theodor-Kutzer-Ufer 1-3	68167 Mannheim	GERMANY	andreas.reiter@medma.uni-heidelberg.de
Speaker	Dr.	Jose-Maria	Ribera	n.a.							C/Canyet S/N	6816 Badalona	SPAIN	iribera@iconologia.net
Participant	Dr.	Johan	Richter	n.a.								221 85 Lund	SWEDEN	Johan.Richter@med.lu.se
Participant	Ms	Marion	Richter	n.a.								Nürnberg	GERMANY	marion.richter@novartis.com
Speaker	Professor	Harald	Rieder	n.a.							Universitätsstraße 1	40225 Düsseldorf	GERMANY	harald.rieder@uni-duesseldorf.de
Participant	Ms	Eva	Rietkötter	n.a.							Robert-Koch-Str. 40	37075 Göttingen	GERMANY	e.rietkoetter@med.uni-goettingen.de
Participant	Dr.	Julia	Ringel	n.a.							Schadenstr. 43	40212 Düsseldorf	GERMANY	julia.ringel@ad.com
Participant	Dr.	Richard	Rosenquist	n.a.							Diaga Hammarstiölds väg 20, Rudbeck	75185 Uppsala	SWEDEN	richard.rosenquist@igp.uu.se
Participant	PD Dr.	Alicia	Rowe	n.a.							Petersstraben 4	40131 Basel	SWITZERLAND	Alicia.Rowe@uhkt.ch
Participant	Professor	Jacob M.	Rowe	n.a.							Est Galim	31036 Haifa	ISRAEL	rowe@rambam.health.gov.il
Participant	PD Dr.	Martin	Rothardt	n.a.							Theodor-Stern-Kai 7	60530 Frankfurt am Main	GERMANY	rothardt@em.uni-frankfurt.de
Participant	Professor	Tapani	Rutu	n.a.							Haartmaninkatu 4	00029 HUS Helsinki	FINLAND	tsapani.rutu@helsinki.fi
Participant	Dr.	Tomasz	Sacho	n.a.							Ul. Kopernika 17	31-501 Krakow	POLAND	tschatom@gmail.com
Participant	Professor	Giuseppe	Scaglio	n.a.							A.O. San Luigi, Regione Gonzola, 11	10043 Orbassano	ITALY	giuseppe.scaglio@unito.it
Participant	Mr	Frank	Salomon	n.a.							Am Haag 6-7	62166 Gröfelling	GERMANY	frank.salomon@fresenius-biotech.com
Participant	Dr.	Hans-Jürgen	Salwender	n.a.							Paul-Ehrlich-Straße 1	22763 Hamburg	GERMANY	h.salwender@asklepios.com
Participant	Dr.	Constantina	Sambani	n.a.							Aghia Paraskevi	11510 Athens	GREECE	csambani@ipts.demokritos.gr
Participant	Professor	Valeria	Santini	n.a.							Viale Gaetano Pieraccini, 24	50139 Firenze	ITALY	vsantini@unifi.it
Participant	Dr.	Lilit	Sargayan	n.a.							Muratsen 114	00075 Yerevan	ARMENIA	lsalitsa@rambler.ru
Participant	Dr.	Nicholas J.	Sarlis	n.a.							Routes 141 & Henry Clay Road	Williamington, DE 19	UNITED KINGDOM	nsarlis@incyte.com
Speaker	PD Dr.	Suzanne	Savabele	n.a.							Pettenkofer Straße 22	68163 Mannheim	GERMANY	suzanne.savabele@medma.uni-heidelberg.de
Participant	Professor	Alexandar	Savic	n.a.							Hejduk Velićeva	21000 Novi Sad	SERBIA	asavic@uniosk.com
Participant	Dr.	Nilgün	Savinalp	n.a.							Hacettepe FM	06100 Sıhhiye - Ankara	TURKEY	nsavinal@gmail.com
Participant	Dr.	Lydia	Scarfo	n.a.							Via Diagetta 58	20132 Milano	ITALY	scarfo.lydia@hsr.it



Participant	Ms	Susanne	Schadlinger	n.a.	female				Roonstraße 25	90423	Nürnberg	GERMANY		susanne.schadlinger@novartis.com
Participant	Ms	Susanne	Schaffert	n.a.	female				Largo Umberto Boccioni 1	21040	Origgio	ITALY		susanne.schaffert@novartis.com
Participant	Dr.	Marcus	Schittenhelm	n.a.	male				Olfrid-Müller-Straße 10	72070	Tübingen	GERMANY		marcus.schittenhelm@med.uni-tuebingen.de
Participant	Professor	Richard	Schlenk	n.a.	male				Albert Einstein Allee 23	93081	Ulm	GERMANY		richard.schlenk@uniklinik-ulm.de
Participant	Dr.	Stefan	Schmidt	n.a.	male				Anichstr. 35 A	6020	Innsbruck	AUSTRIA		stefan.schmidt@i-med.ac.at
Participant	Professor	Annette	Schmitt-Gräff	n.a.	female				Breisacher Str. 114a	73104	Freiburg	GERMANY		annette.schmitt-graef@uniklinik-freiburg.de
Participant	Dr.	Stephanie	Schneider	n.a.	female				Marchioninstr. 15	81377	München	GERMANY		stephanie.schneider@med.lmu.de
Participant	Professor	Sebastian	Scholl	n.a.	male				Erlanger Allee 101	07740	Jena	GERMANY		sebastian.scholl@med.uni-jena.de
Participant	Ms	Nicole	Schomber	n.a.	female				Pettenkofer Straße 22	68169	Mannheim	GERMANY		nicole.schomber@medma.uni-heidelberg.de
Participant	Dr.	Anje	Schulze	n.a.	female				Nordhäuser Straße 74	93089	Erfurt	GERMANY		anje.schulze@helios-kliniken.de
Participant	Ms	Christiane	Schumann	n.a.	female				Pettenkoferstrasse 22	63169	Mannheim	GERMANY		christiane.schumann@medma.uni-heidelberg.de
Participant	Dr.	Gerrit Jan	Schuurhuis	n.a.	male				De Boelelaan 117, Postbus 7057	1007 MB	Amsterdam	THE NETHERLANDS		G.J.Schuurhuis@vumc.nl
Speaker	Dr.	Sebastian	Schwind	n.a.	male				Johannisallee 32A	04103	Leipzig	GERMANY		Sebastian.Schwind@uniklinik-leipzig.de
Participant	Ms	Barbara	Seipel	n.a.	female				Konrad-Adenauer Strasse 17	65218	Ingelheim	GERMANY		barbara.seipel@bioscientia.de
Participant	Dr.	Andrea	Sendler	n.a.	female				Leimenstr. 20	63450	Hanau	GERMANY		andrea_sendler@klinikum-hanau.de
Participant	Professor	Hubert	Serve	n.a.	male				Theodor-Stern-Kai 7	60590	Frankfurt am Main	GERMANY		serve@em.uni-frankfurt.de
Participant	Dr.	Katjooon	Shirneshan	n.a.	female				Robert-Koch-Straße 40	37075	Göttingen	GERMANY		shirneshan@med.uni-goettingen.de
Participant	Ms	Valentyna	Sholjko	n.a.	female				av. Pobeda 119/121	03115	Kiev	UKRAINE		val-sholjko@yandex.ru
Participant	Dr.	Vasily	Shuvaev	n.a.	male				2nd Sovetskaya str., 16	191024	St. Petersburg	RUSSIA		shuvaev77@mail.ru
Participant	Professor	Richard T.	Silver	n.a.	male				525 East 68th Street, Box 581, Pkwy	10065	New York	USA		rtsilver@med.cornell.edu
Speaker	Professor	Bengt	Simonsson	n.a.	male				Sankt Olofsgatan 10B	75185	Uppsala	SWEDEN		bengt.simonsson@medsci.uu.se
Participant	Dr.	Lukas	Smolej	n.a.	male				Sokolská 581	50005	Hradec Králové	CZECH REPUBLIC		smolej@seznam.cz; lukas.smolej@fnhk.cz
Participant	Dr. rer. nat.	Catherine	Sodan-Boyer	n.a.	female				Pettenkoferstr. 22	68169	Mannheim	GERMANY		catherine.sodan-boyer@medma.uni-heidelberg.de
Participant	Ms	Simona	Soverini	n.a.	female				Via Massarenti 3	40138	Bologna	ITALY		simona.soverini@tin.it
Participant	Ms	Petra	Spengler	n.a.	female				Roonstraße 25	90429	Nürnberg	GERMANY		petra.spengler@novartis.com
Participant	PD Dr.	Karsten	Spiekermann	n.a.	male				Marchioninstr. 15	81377	München	GERMANY		karsten.spiekermann@med.uni-muenchen.de
Participant	Dr.	Andreas	Springauf	n.a.	male				Friedrichstr. 2-6	60323	Frankfurt am Main	GERMANY		aspringauf@otsuka.de
Participant	Dr.	Elena	Stadnik	n.a.	female				Lev Tolstoy 6/8	197022	St. Petersburg	RUSSIA		elena_stadnik@mail.ru
Participant	Dr.	Viktoria	Stalbovsckaja	n.a.	female					4058	Basel	SWITZERLAND		viktorija.stalbovsckaja@novartis.com
Participant	Ms	Evangelia	Stalika	n.a.	female				Exochi	57010	Thessaloniki	GREECE		evangelia.stalika@gmail.com
Participant	Ms	Inge	Stallann	n.a.	female				Pettenkofer Straße 22	68169	Mannheim	GERMANY		inge.stallann@medma.uni-heidelberg.de
Participant	Dr.	Kostas	Stamatopoulos	n.a.	male					67010	Thessaloniki	GREECE		kostas.stamatopoulos@gmail.com
Speaker	Professor	Reinhard	Stauder	n.a.	male				Anichstraße 35	6020	Innsbruck	AUSTRIA		reinhard.stauder@i-med.ac.at
Participant	Professor	Juan Luis	Stegemann	n.a.	male				Diego de León, 52	28006	Madrid	SPAIN		ljustegemann@gmail.com
Participant	Dr.	Frank	Stegelmann	n.a.	male				Albert-Einstein-Allee 23	69081	Ulm	GERMANY		frank.stegelmann@uniklinik-ulm.de
Participant	Dr.	Gerold	Stellmacher	n.a.	male				Schildkrötstraße 17-19	68199	Mannheim	GERMANY		gstellmacher@cti-lifesciences.com
Participant	Dr.	Leif	Stenke	n.a.	male					17176	Stockholm	SWEDEN		leif.stenke@ki.se
Participant	Professor	Stephan	Stilgenbauer	n.a.	male				Albert-Einstein-Allee 23	69081	Ulm	GERMANY		stephan.stilgenbauer@uniklinik-ulm.de
Participant	Dr.	Susanne	Stoelben	n.a.	female				Schadowstraße 44	40212	Düsseldorf	GERMANY		Susanne.Stoelben@sigma-tau.de
Participant	Dr.	Heribert	Strotkotter	n.a.	male				Heusnerstr. 40	42283	Vuppertal	GERMANY		dr.strotkotter@t-online.de
Participant	Dr.	Milagna	Subar	n.a.	female				Route 206 & Provinceline Rd.	08543	Princeton, NJ	USA		milagna.subar@bms.com
Participant	PD	Irina	Subortseva	n.a.	female				Novy Zykovsky pr. 4a	125167	Moscow	RUSSIA		soubortseva@yandex.ru
Participant	Dr.	Andrey	Sudarikov	n.a.	male				Novy Zykovsky pr. 4	125167	Moscow	RUSSIA		dusha@blood.ru
Participant	Dr.	Lesley Ann	Sutton	n.a.	female				Rudbecklaboratoriet, Dag Hammarskjöldsgatan 1	75185	Uppsala	SWEDEN		lesley.sutton@IGP.uu.se
Speaker	Ms	Argiris	Symeonidis	n.a.	male						Patras	GREECE		
Participant	Dr.	Rodica	Talmaci	n.a.	female				258 Fundeni str, sector 2	022328	Bucharest	ROMANIA		rodica.talmaci@gmail.com
Participant	Dr.	Gevorg	Tamamyan	n.a.	male				Muratsan 114	00075	Yerevan	ARMENIA		gevorgtamamyan@gmail.com
Participant	Dr.	Christoph	Tapprich	n.a.	male				Preußenstr. 34	41464	Neuss	GERMANY		tapprich@hotmail.com
Participant	Dr.	Ambra	Tarquini	n.a.	female				P. le Umanesimo 10	00144	Rome	ITALY		elisabetta.abruzzese@uniroma2.it
Participant	Dr.	Aurelia	Tatic	n.a.	female				258, Fundeni Pload	022328	Bucharest	ROMANIA		auratic@yahoo.com
Participant	Mr	Jörg	Teubner	n.a.	male				Ulagenstr. 1	40724	Hilden	GERMANY		joerg.teubner@qiagen.com
Participant	Ms	Frauke	Theis	n.a.	female						Ulm	GERMANY		frauke.theis@uniklinik-ulm.de
Participant	Professor	Christian	Thiede	n.a.	male				Fetscherstr. 74	01307	Dresden	GERMANY		christian.thiede@uniklinikum-dresden.de
Participant	Mr	Jürgen	Thiele	n.a.	male						Köln	GERMANY		
Participant	Professor	Jürgen	Thiele	n.a.	male				An der Bismarckschule 18	30173	Hannover	GERMANY		jthiele@uni-koeln.de
Participant	Dr.	Noortje	Thielen	n.a.	female				De Boelelaan 117	2 PK 018 1091H	Amsterdam	Niederlande		n.thielen@vumc.nl
Participant	Dr.	Felicitas	Thol	n.a.	female				Carl-Neuberg-Str. 1	30625	Hannover	GERMANY		thol.felicitas@mh-hannover.de
Participant	Mr	Eustace	Thomas	n.a.	male				Roonstraße 25	90429	Nürnberg	GERMANY		eustace-john.thomas@novartis.com
Participant	Dr.	Gianni	Tognoni	n.a.	male						Santa Maria Imbari	ITALY		tognoni@negrisud.it
Participant	Ms	Natasa	Tozic	n.a.	female				Vojvode Stepe 444A	11010	Belgrade	SERBIA		nmtosic@sezampro.rs
Participant	Professor	Marek	Trněný	n.a.	male				U nemocnice 1	128 20	Praha	CZECH REPUBLIC		trneny@cesnet.cz
Participant	Dr.	Grigory	Tsaur	n.a.	male				Serafima Derqabina, 32	620149	Yekaterinburg	RUSSIA		283850@gmail.com

