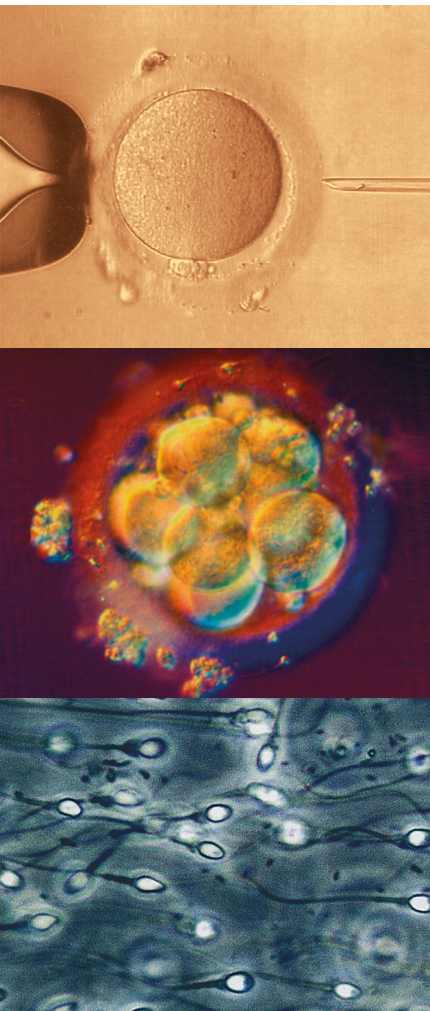


Journal of

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– Journal für Reproduktionsmedizin und Endokrinologie –

Andrology • Embryology & Molecular Biology • Endocrinology • Ethics & Law • Genetics
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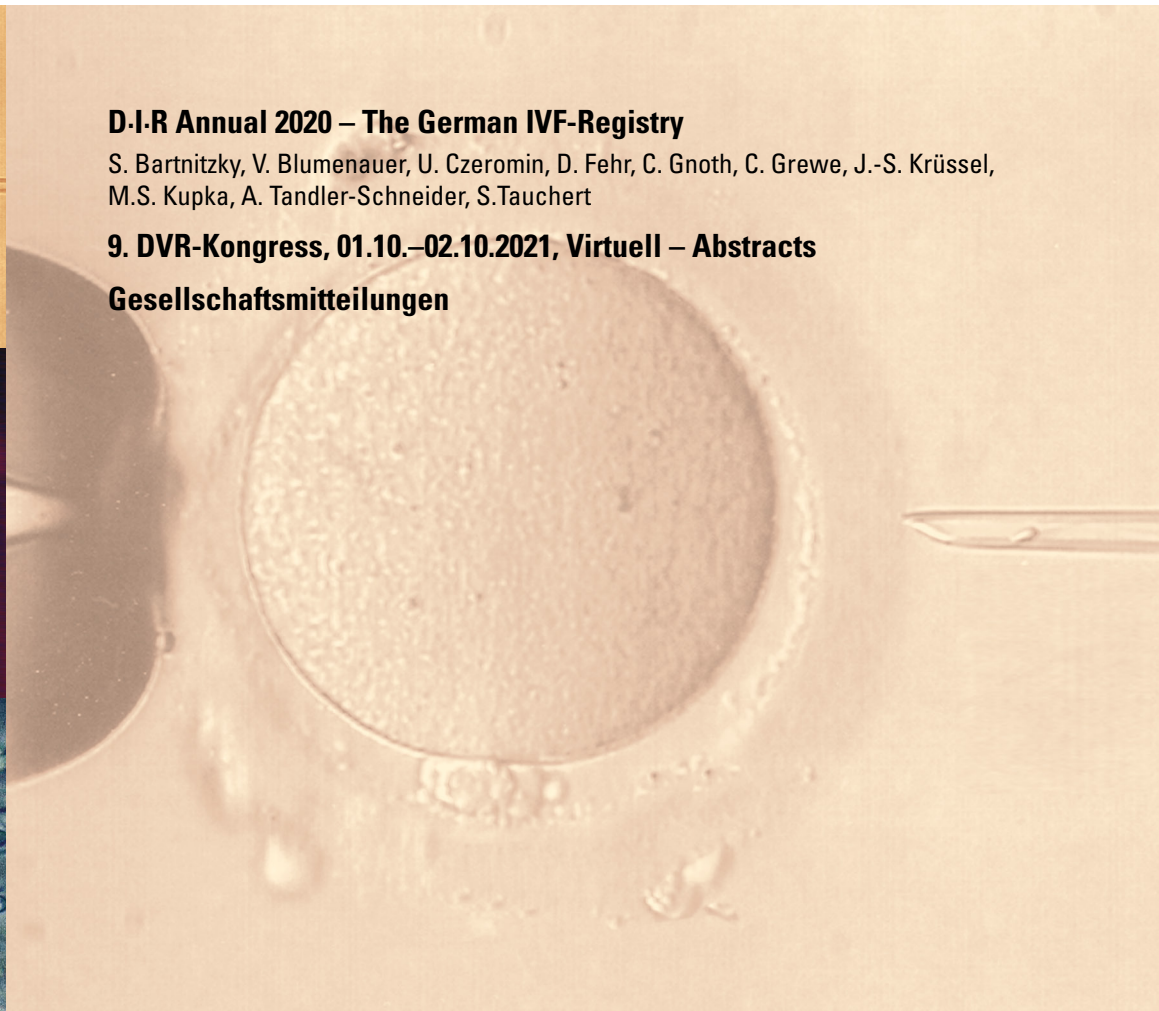


D-I-R Annual 2020 – The German IVF-Registry

S. Bartnitzky, V. Blumenauer, U. Czeromin, D. Fehr, C. Gnoth, C. Grewe, J.-S. Krüssel,
M.S. Kupka, A. Tandler-Schneider, S. Tauchert

9. DVR-Kongress, 01.10.–02.10.2021, Virtuell – Abstracts

Gesellschaftsmitteilungen



Offizielles Organ: AGRBM, BRZ, DVR, DGA, DGGEF, DGRM, D-I-R, OEGRM, SRBM/DGE



D·I·R
DEUTSCHES IVF REGISTER

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Preface

D-I-R Annual 2020 – The German IVF-Registry

S. Bartnitzky, V. Blumenauer, U. Czeromin, D. Fehr, C. Gnoth, C. Grewe, J.-S. Krüssel, M.S. Kupka, A. Tandler-Schneider, S. Tauchert

Once again it is our great pleasure to present the annual report of the data collected by the German IVF-Registry (D-I-R). The 2020 Registry looks back onto more than 30 years of diligent data collection, extraction, and evaluation.

Since 1996, the German IVF-Registry has been electronically collecting data for each initiated treatment cycle.

Meanwhile, nearly 2.0 million ART cycles (ART: Assisted Reproduction Technology) since 1982 and more than 340,000 children born since 1997 have been documented in the database. The prospective documentation as well as the cycle-by-cycle data collection are of particular value and make the German IVF-Registry unique world-wide.

This Yearbook 2020 is like the Yearbook 2019 a yearbook of change and of organizational efforts undertaken by all authors and contributors. After years of data control and reconciliation, administrative work and programming, the implementation of a new data system has been rolled out to the IVF-clinics. This is still a major change and required huge modifications within the existing, heterogeneous IT environments of those pioneering centers. We want to express our thanks to everyone contributing to this process: the software companies, the D-I-R data management group, the D-I-R office, and particularly the associates within those centers.

The 2020 yearbook is based on exploitable exports from 134 centers, for which

a number of 111,752 plausible cycles was documented. A total of 62,431 women were treated in 2020, resulting in an average of 1.9 cycles per woman. We are happy to note that the pregnancy rate per transfer in fresh cycles was 31.9%, whereas the pregnancy rate per transfer in cryo cycles was 29.4%. In the previous year, 2019, the documented birth rate per embryo transfer was 23.3% in fresh cycles and 20.4% in cryo cycles, with an acceptable general pregnancy outcome documentation of 93.0%.

Remarkably, 70.1% of the pregnancies led to a birth. A birth rate of 22.2% per embryo transfer was achieved. For comparison, the probability for a 25 year old woman to become pregnant WITHOUT any medical assistance averages at 23%. For a 35 year old woman this probability decreases to 16%, according to the Bundesgesundheitsblatt [BGBI 2013; 56: 1633–41].

In the last years particular attention was placed on the known problems of multiple pregnancies within ART cycles. In 2019, the rate of twin births was 17.8% in fresh cycles and 13.0% in cryo cycles. This rate is still too high, but we have been able to see a small decrease in multiple birth since two years!

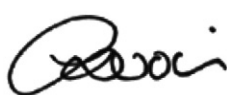
The data presented in this yearbook demonstrate the high quality of reproductive medicine in Germany, leading to impressive results even under the restrictive legislation: in accordance with the “Embryo Protection Act” of 1990, the elective single embryo transfer (eSET)

is still unlawful in Germany. Egg donation is forbidden as well. Cryopreservation is allowed for usually 2-PN stage oocytes only. The regulation that only a maximum of three embryos may be transferred has certainly led to positive effects. Due to the increasing quality of stimulation, improvements in oocyte and embryo treatment, and changes in transfer technology, the average number of transferred embryos has decreased by more than 25% in fresh cycles since 1997. Nevertheless on average 1.7 embryos after IVF and ICSI were transferred.

Good and diligent documentation is proof of a highly organized and efficient workflow. The prospective cycle-by-cycle data collection is another quality feature of the German IVF-Registry, a feature that sets the German IVF-Registry apart from any other registry in the world. The results of the comprehensive data collection in the German IVF-Registry and its reports have been cited in numerous scientific publications, nationally as well as internationally, lending support and impetus to scientific research.

Furthermore, the data have contributed to the continuous improvement of quality within the German centers for reproductive medicine. Physicians, biologists, and patients alike have benefited from the tedious work necessary to collect, export, and evaluate as well as publish the data. Therefore, we express our gratitude to all those involved.

Thank you and keep up the good work!



Dr. med. Ute Czeromin
President of the Board



Prof. Dr. med. Jan-Steffen Krüssel



Dr. med. Andreas Tandler-Schneider



Responsible for this Edition

Deutsches IVF-Register e.V. (D·I·R)[®] German IVF Registry

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Prof. Dr. med. Frank Nawroth
PD Dr. rer. nat. Verena Nordhoff
Prof. Dr. med. Nicole Sanger
Moritz Suerdieck
Prof. Dr. med. Michael von Wolff
Head Office: Weidornweg 17 · 35041 Marburg/Lahn
Tel.: +49 (0) 64 20 - 305 05 83
Email: info@fertiprotekt.com
www.fertiprotekt.com

D·I·R Office and D·I·R Data Management

Markus Kimmel · Kimmelnet
Lise-Meitner-Strae 14 · 40591 Dusseldorf
Tel. +49 (0)211 913 84 800
e-mail: geschaeftsstelle@deutsches-ivf-register.de
www.german-ivf-registry.com

Technology Interface ARTbox[®] and D·I·R-own acquisition program DIRproNOVA[®]

CRITEX GmbH
Stobaeusplatz 4 · 93047 Regensburg
Tel. +49 (0)941 569 98 770
e-mail: mail@critex.de

Evaluation Software

Qlik Technologies Inc., King of Prussia, PA 19406, USA
QlikTech GmbH, Dusseldorf, Germany
www.qlik.com/de-de
Transact - Gesellschaft fur Software & Analyse mbH
Hamburg, Germany
www.transact.de

Design and Layout

U&MI Visual Communication
Dipl.-Des. Soo-Hee Kim

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Number of Treatments in 2020

Centers for IVF-, ICSI-, and Cryo Transfer Treatments

Members of the German IVF-Registry 2020	n=139
Registry Participants 2020*	n=134
Data Received by Deadline May 6th 2021*	n=134
Documented Treatment Cycles	n=116,306
Number of Women Treated**	n=62,431
Mean Number of Treatment Cycles per Woman	1.9

Type of plausible treatment 2016 – 2020

IVF, ICSI, IVF/ICSI, Cryo Transfer – Prospective and Retrospective Data

	2016		2017		2018		2019		2020	
	n	%	n	%	n	%	n	%	n	%
IVF	15,586	15.5	15,586	15.2	17,254	16.7	18,159	16.9	18,724	16.8
ICSI	47,554	47.2	45,548	44.4	46,027	44.5	45,984	42.7	44,336	39.7
IVF/ICSI	1,201	1.2	1,135	1.1	1,424	1.4	1,338	1.2	1,471	1.3
Freeze all - MII			1,200	1.2	1,630	1.6	1,824	1.7	1,879	1.7
Freeze All - PNs and Embryos	3,319	3.3	4,089	4.0	4,750	4.6	5,100	4.7	5,476	4.9
Cryo	25,181	25.0	26,084	25.5	27,878	26.9	30,411	28.2	32,945	29.5
Mixed Fresh and Cryo Cycles	891	0.9	1,014	1.0	1,058	1.0	999	0.9	681	0.6
None (= Break-off before oocyte treatment or thawing)	7,022	7.0	7,831	7.6	3,524	3.4	3,875	3.6	6,240	5.6
Total Plausible Cycles	100,754		102,487		103,545		107,690		111,752	

*) Five centers could not be part of the nationale analyses

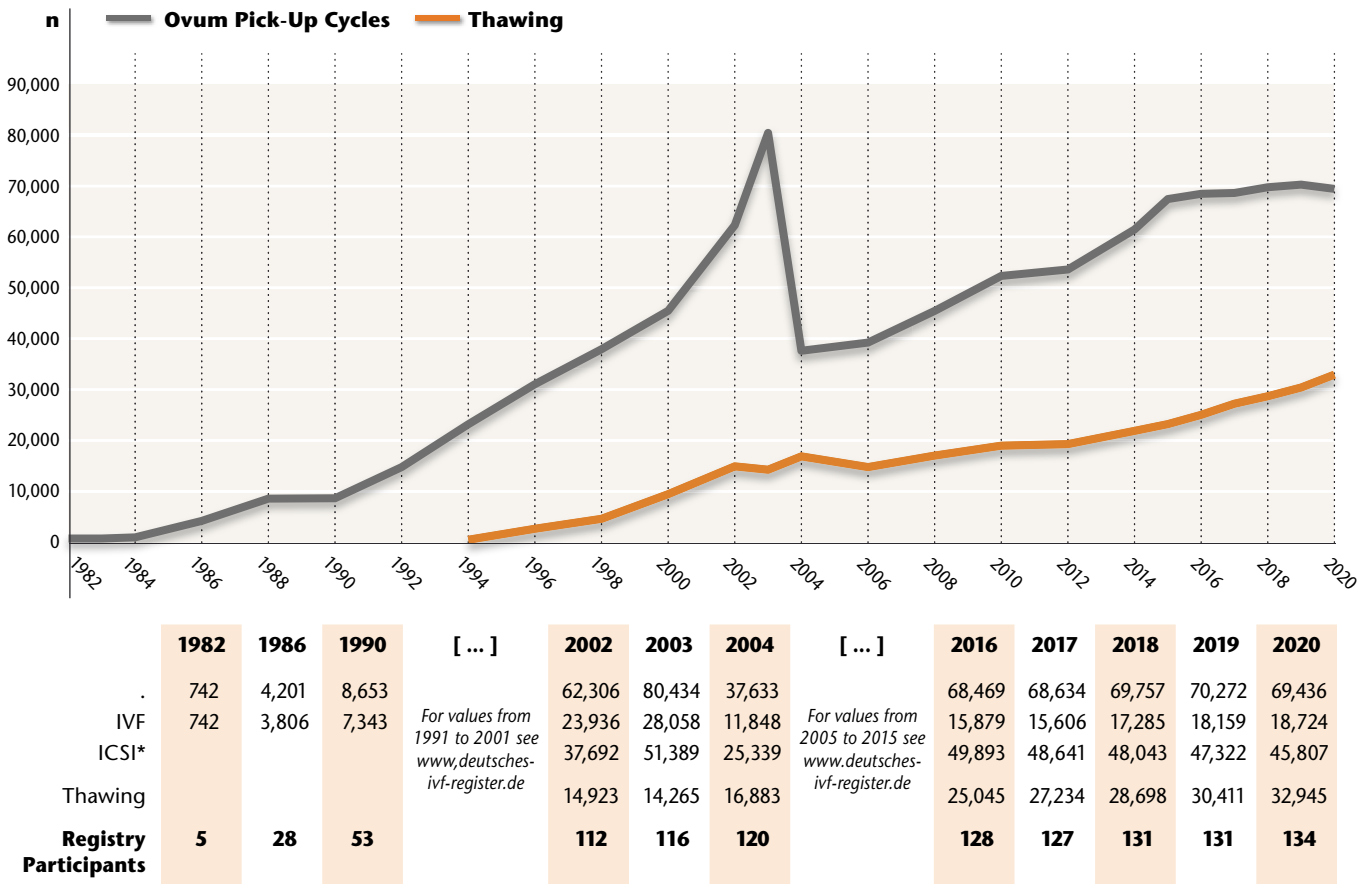
**) Base quantity: Total number of women, including implausible treatment cycles

Number of Oocyte Retrievals (Freshcycles) 1982 – 2020

Number of Thawing Cycles 1994 – 2020

Registry Participants 1982 – 2020

IVF, ICSI* – Prospective and Retrospective Data

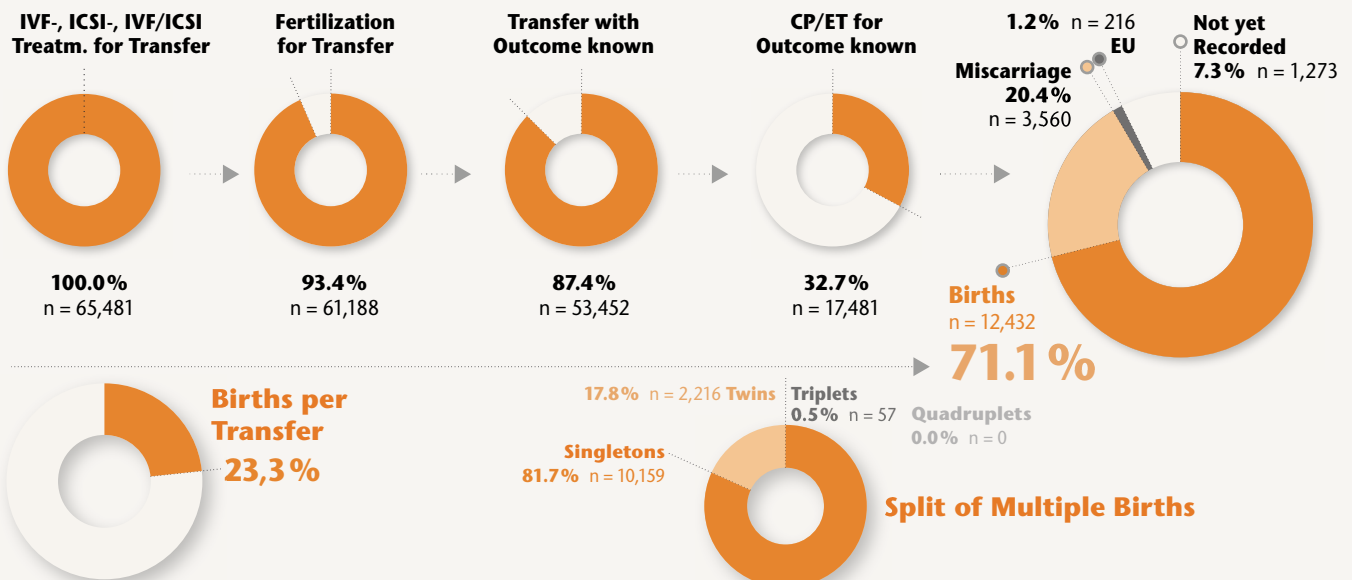


Data for 1982 to 2015 are published and available. Separate presentation of GIFT, ZIFT, IVF/ICSI was abstained from.

*) Where IVF/ICSI is not explicitly mentioned, the treatments were added to ICSI.

Summary of Statistics in Brief 2019 – CoD May 6th 2021

IVF, ICSI and IVF/ICSI – Prospective and Retrospective Data



Quality of Documentation 2019/2020

Plausible and Prospectively Documented Cycles, Cycle- and Pregnancy-Outcomes



Analyses results can only be as good as the underlying data.

The D-I-R-team is grateful to all participating centers and their meticulous work!

Plausible Cycles 2020

111,752 plausible cycles out of **116,363** documented cycles. **96.1%** of documented cycles are plausible. This does unfortunately not imply completeness of data required, thus impeding detailed analyses.

Prospectively Documented Cycles 2020

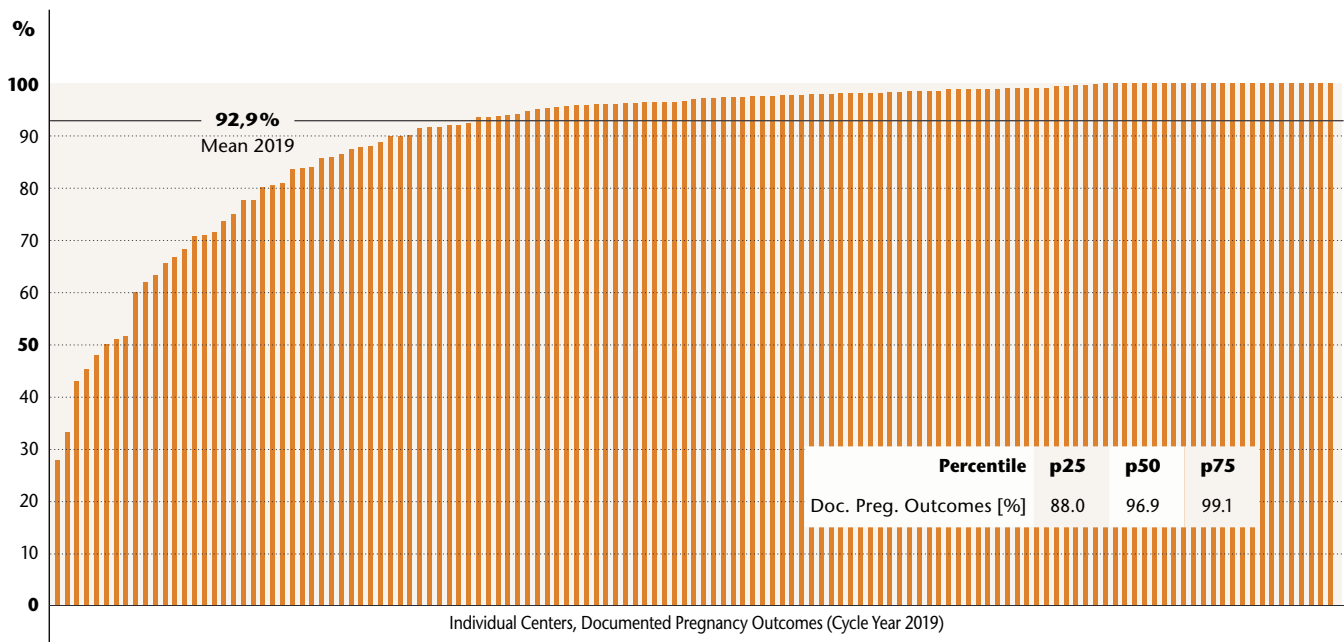
100,736 prospectively documented cycles out of **111,752** plausible cycles. **90.1%** of plausible cycles were prospectively documented. Worldwide, only the German IVF Registry separately shows the prospective cycles. This in itself is an indicator for quality!

Documented Cycle-Outcomes 2020

In **83,136** out of **83,889** embryo transfers, the cycle outcome was documented (**99.1%**).

Documented Pregnancy Outcomes 2019

In **24,400** out of **26,275** clinical pregnancies, the pregnancy outcome was documented (**92.9%**).



We are well aware that the meticulous documentation of these relevant data is troublesome, laborious and resource-binding.

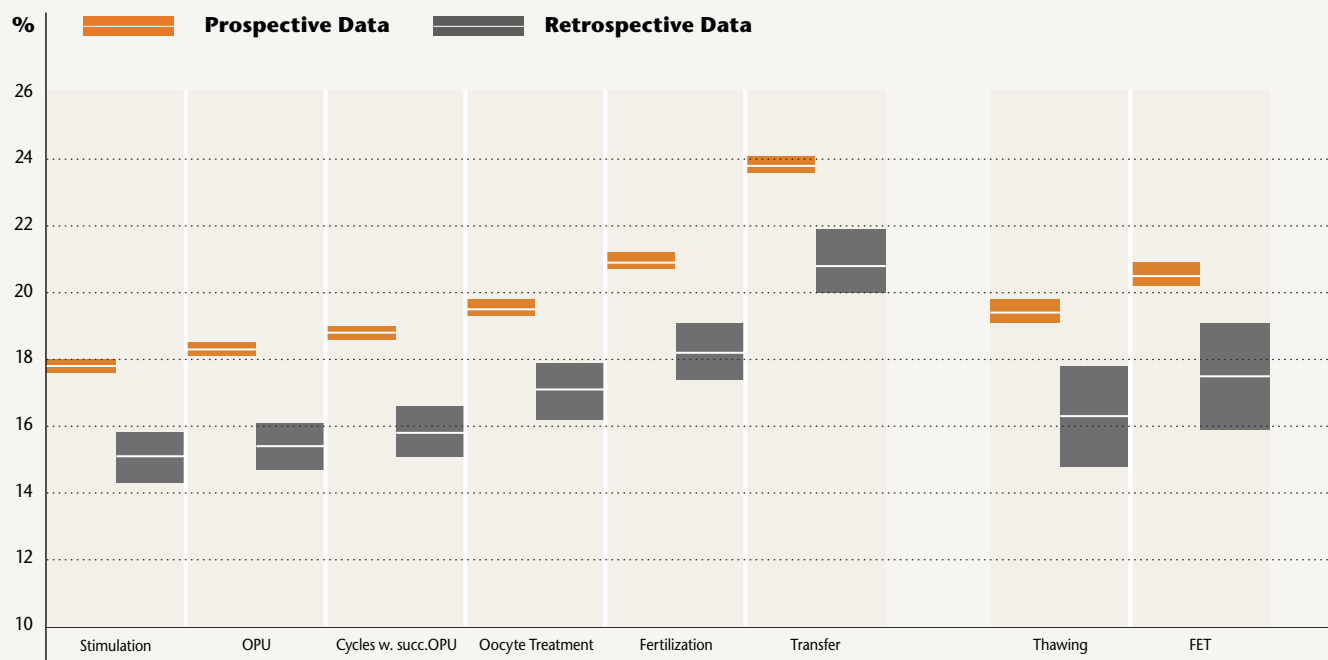
The D-I-R data analyses team is therefore very happy to state that for the third year the percentage of prospectively documented cycles is **above 90%**.

Unfortunately, the percentage of documented pregnancy outcomes has decreased (**95.0% -> 92.9%**). Only **62.6%** of individual centers have reached the goal of documenting more than **95%** of pregnancy outcomes (last annual: **72.1%**).

The dedicated goal of the D-I-R board of directors and board of trustees is to motivate the members of the D-I-R to furthermore keep their focus on the prospectivity and to achieve also for the next year a rate of documented pregnancy outcomes of **95 plus** percent.

Birth Rate per Treatment Level in Fresh and Cryo Treatment Cycles 2018 and 2019

Prospective and Retrospective Data



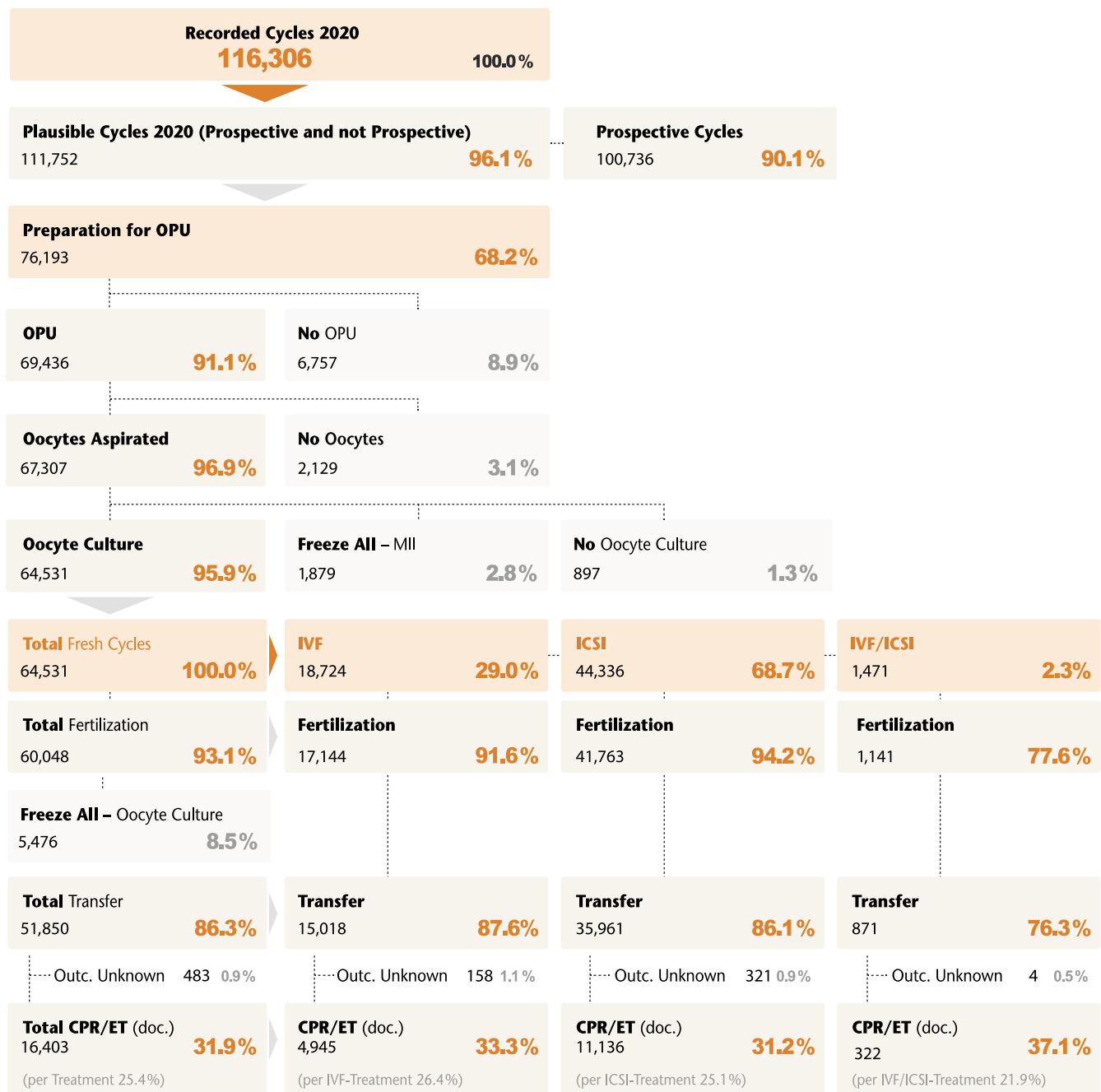
	Stimulation	OPU	Cycles with Successful OPU	Oocyte Treatment	Fertilization	Transfer	Thawing	FET
Prospective Numbers	133,925	130,576	126,897	122,112	113,842	100,289	55,871	52,885
Birth	23,846	23,846	23,846	23,846	23,846	23,846	10,862	10,862
Upper Confid. Limit*	18.0	18.5	19.0	19.8	21.2	24.1	19.8	20.9
Birth/Treatm. %	17.8	18.3	18.8	19.5	20.9	23.8	19.4	20.5
Lower Confid. Limit*	17.6	18.1	18.6	19.3	20.7	23.6	19.1	20.2
Retrospective Numbers	9,150	8,949	8,696	8,074	7,552	6,636	2,342	2,181
Birth	1,378	1,378	1,378	1,378	1,378	1,378	382	382
Upper Confid. Limit*	15.8	16.1	16.6	17.9	19.1	21.9	17.8	19.1
Birth/Treatm. %	15.1	15.4	15.8	17.1	18.2	20.8	16.3	17.5
Lower Confid. Limit*	14.3	14.7	15.1	16.2	17.4	20.0	14.8	15.9

* With a 95%-probability, the true mean lies within the defined confidence interval.

D-I-R Statistics in Brief – Fresh Cycles 2020 (CoD May 6th 2021)



German IVF Registry – Prospective and Retrospective Data



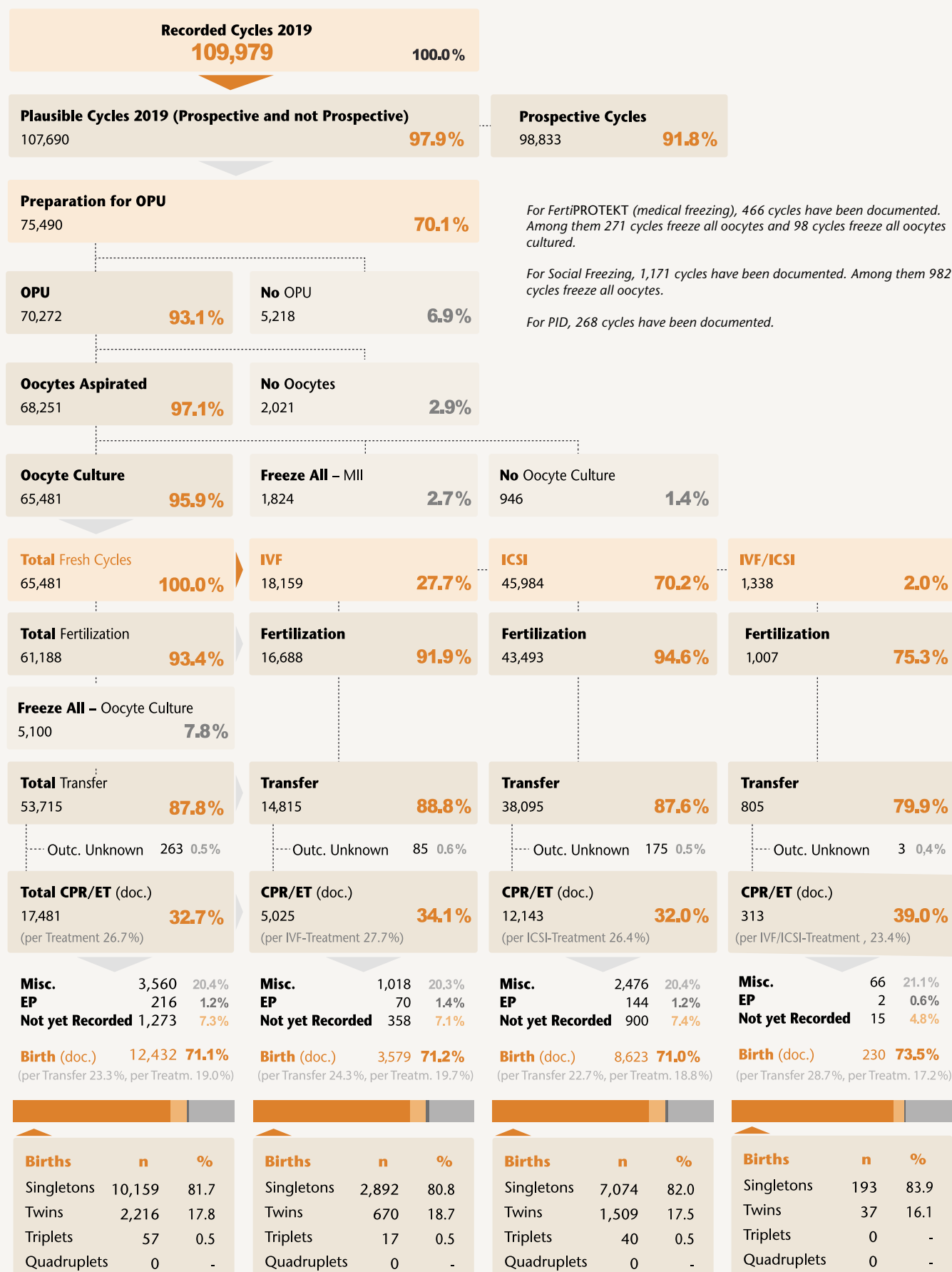
For FertiPROTEKT (medical freezing), 512 cycles have been documented. Among them 320 cycles freeze all oocytes and 82 cycles freeze all oocytes cultured.

For Social Freezing, 1,565 cycles have been documented. Among them 1,281 cycles freeze all oocytes.

For PID, 288 cycles have been documented.

D-I-R Statistics in Brief – Fresh Cycles 2019 (CoD May 6th 2021)

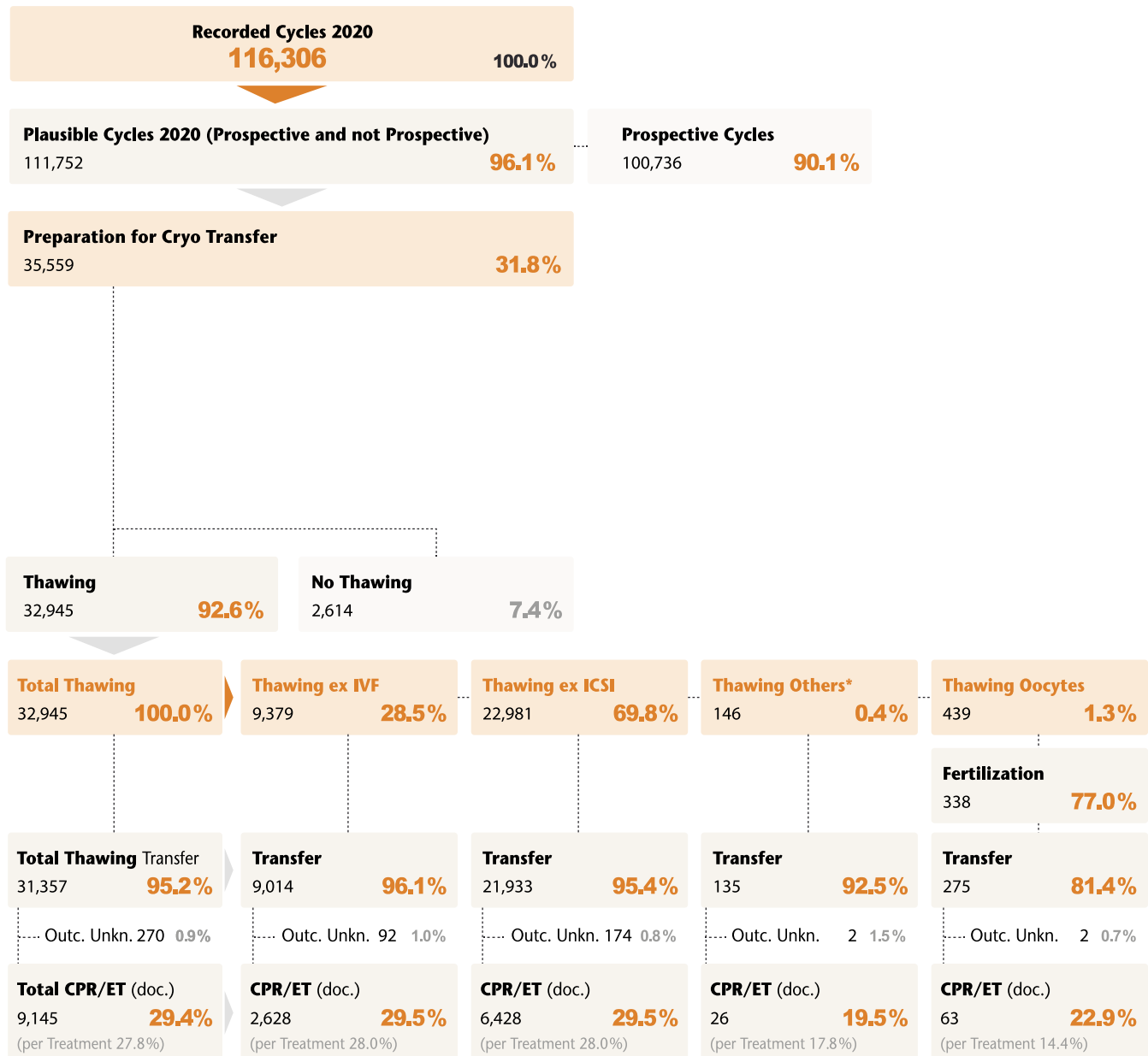
German IVF Registry – Prospective and Retrospective Data



D-I-R Statistics in Brief – Cryo Cycles 2020 (CoD May 6th 2021)



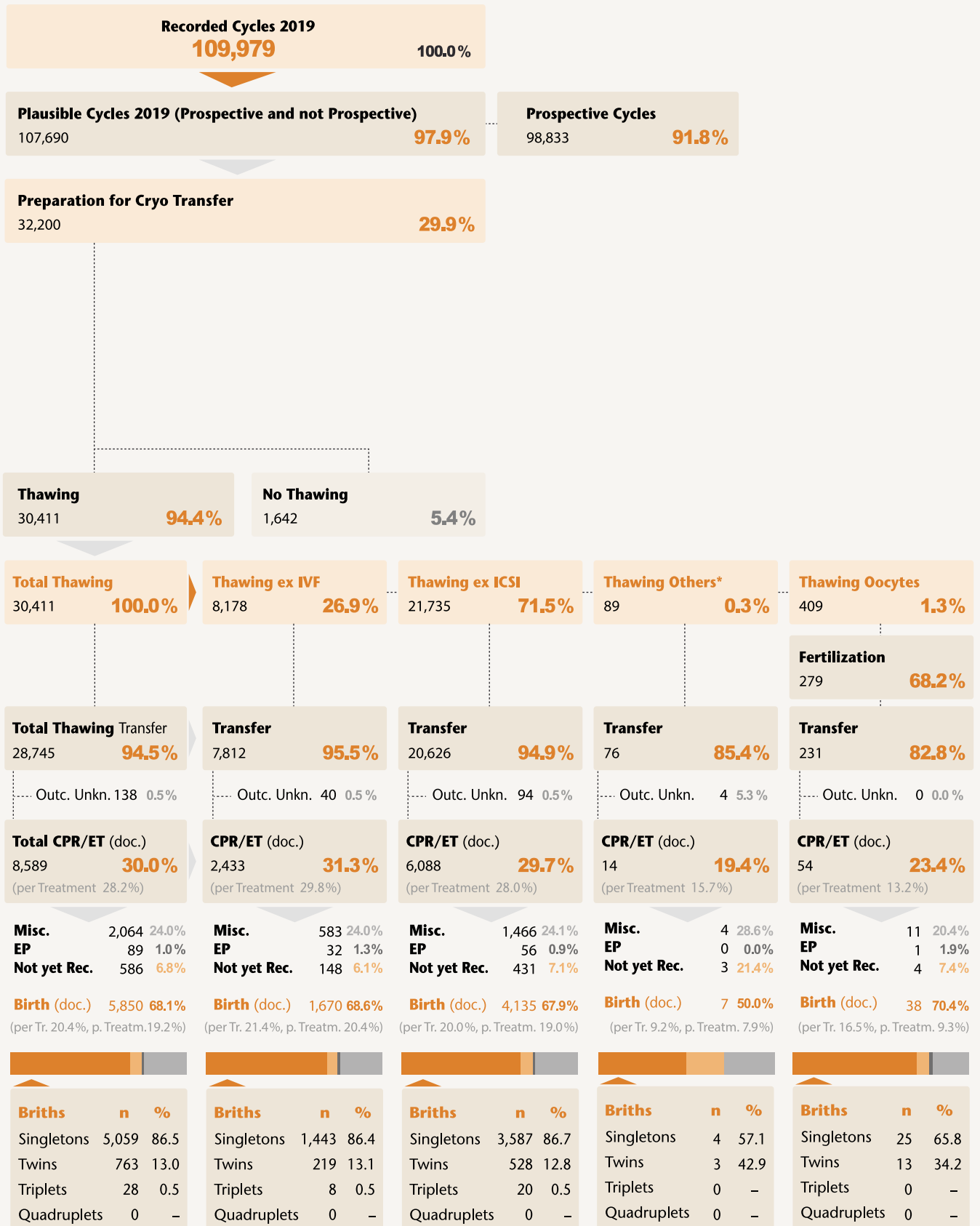
German IVF Registry – Prospective and Retrospective Data



*) Thawing others means cycles with unknown previous treatment, previous treatment not documented or previous treatment has partly been IVF and ICSI.

D-I-R Statistics in Brief – Cryo Cycles 2019 (CoD May 6th 2021)

German IVF Registry – Prospective and Retrospective Data



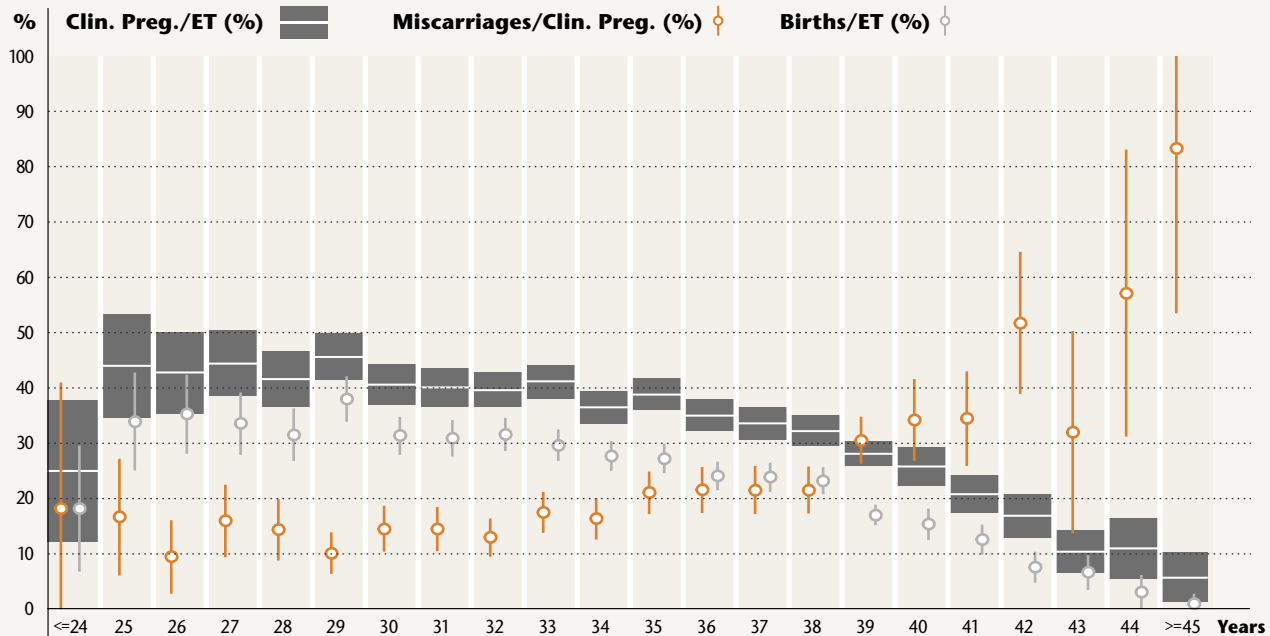
*) Thawing others means cycles with unknown previous treatment, previous treatment not documented or previous treatment has partly been IVF and ICSI.



Pregnancy Rate and Ongoing Pregnancy as a Function of Female Age 2019

Prospective Data

IVF 2019



n	44	109	175	269	368	547	695	743	943	976	1,048	1,116	1,064	1,027	1,125	1,610	617	575	346	243	127	105	ET
%	25.0	44.0	42.8	44.4	41.6	45.6	40.6	40.1	39.6	41.2	36.5	38.8	35.0	33.6	32.2	28.1	25.8	20.8	16.9	10.4	11.0	5.7	CP/ET
n	8	37	61	90	116	206	218	229	295	288	287	303	255	244	261	272	94	72	26	16	4	1	Birth

Age in Years	<= 29	30 – 34	35 – 39	40	41	42	43	44	>=45	Total
OPU	1,898	5,377	7,217	746	698	432	307	163	135	16,973
Oocytes ¹	11.5	10.5	8.4	6.6	6.4	6.0	5.3	4.8	4.0	9.0
Insemination ¹	11.4	10.3	8.2	6.6	6.2	5.9	5.3	4.8	3.9	8.9
ET	1,512	4,405	5,942	617	575	346	243	127	105	13,872
ET/OPU %	79.7	81.9	82.3	82.7	82.4	80.1	79.2	77.9	77.8	81.6
Trans. Embr. ¹	1.67	1.63	1.65	1.65	1.65	1.75	1.66	1.75	1.71	1.65
CP	652	1,728	1,958	158	119	58	25	14	6	4,718
CP/OPU %	34.4	32.1	27.1	21.2	17.0	13.4	8.1	8.6	4.4	27.8
CP/ET Upper Conf. Limit*%	45.9	40.9	34.3	29.3	24.1	20.8	14.3	16.5	10.2	35.0
CP/ET %	43.4	39.5	33.1	25.8	20.8	16.9	10.4	11.0	5.7	34.2
CP/ET Lower Conf. Limit*%	40.8	38.0	31.9	22.3	17.4	12.9	6.6	5.6	1.3	33.4
CP/ET %: 2 Embr. Transf. + min. 2 2-PN Surplus	49.9	47.1	39.9	33.9	27.1	28.6	13.5	25.7	16.7	42.6
CP/ET %: 1 Embr. Transf. + min. 3 2-PN Surplus	44.3	36.6	35.7	27.8	29.0	12.5	10.5	0.0	0.0	36.1
Misc./CP Upper Conf. Limit*%	15.3	17.0	25.4	41.6	43.0	64.6	50.3	83.1	100.0	21.3
Misc./CP %	12.7	15.3	23.5	34.2	34.5	51.7	32.0	57.1	83.3	20.2
Misc./CP Lower Conf. Limit*%	10.2	13.6	21.6	26.8	25.9	38.9	13.7	31.2	53.5	19.1
Births/ET Upper Conf. Limit*%	36.8	31.4	23.6	18.2	15.3	10.4	9.8	6.2	2.8	25.2
Births/ET %	34.4	30.1	22.6	15.4	12.6	7.6	6.7	3.1	1.0	24.5
Births/ET Lower Conf. Limit*%	32.0	28.7	21.5	12.5	9.9	4.8	3.5	0.1	0.0	23.8

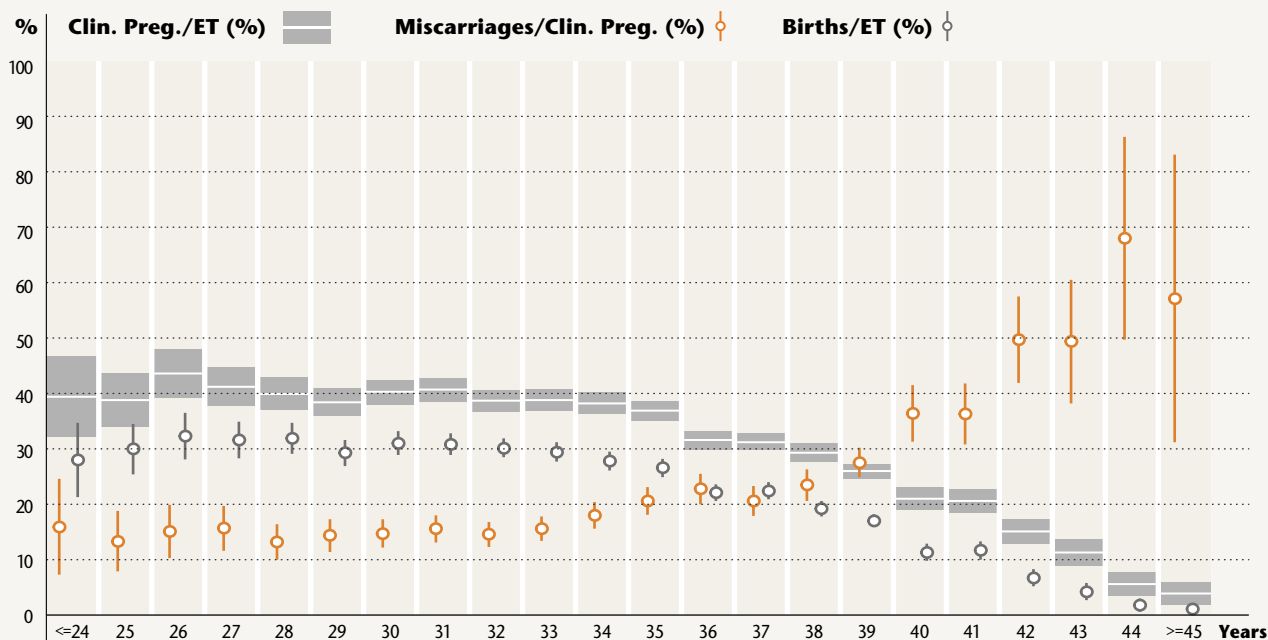
1) Mean

* With a 95%-probability, the true mean lies within the defined confidence interval.

Pregnancy Rate and Ongoing Pregnancy as a Function of Female Age 2019

Prospective Data

ICSI 2019



n	175	389	487	764	1,087	1,437	1,825	2,097	2,441	2,608	2,595	2,827	2,889	2,857	2,872	4,104	1,620	1,422	1,056	687	451	359	ET
%	39.4	38.8	43.6	41.2	39.9	38.4	40.3	40.7	38.7	38.8	38.2	36.9	31.6	31.2	29.3	26.0	21.0	20.6	15.1	11.3	5.6	3.9	CP/ET
n	49	116	157	240	344	419	565	642	733	764	718	748	635	639	549	696	182	166	71	29	8	4	Birth

Age in Years	<= 29	30 – 34	35 – 39	40	41	42	43	44	>=45	Total
OPU	5,230	13,753	18,476	1,987	1,754	1,345	880	583	476	44,484
Oocytes ¹	12.3	11.1	8.8	7.2	6.6	6.0	5.6	5.1	4.4	9.5
Injection ¹	9.6	8.7	7.0	5.7	5.3	4.9	4.5	4.1	3.6	7.5
ET	4,339	11,566	15,549	1,620	1,422	1,056	687	451	359	37,049
ET/OPU %	83.0	84.1	84.2	81.5	81.1	78.5	78.1	77.4	75.4	83.3
Trans. Embr. ¹	1.70	1.70	1.69	1.71	1.73	1.70	1.74	1.70	1.69	1.70
CP	1,725	4,515	4,739	338	292	159	77	25	14	11,884
CP/OPU %	33.0	32.8	25.6	17.0	16.6	11.8	8.8	4.3	2.9	26.7
CP/ET Upper Conf. Limit* %	41.4	40.1	31.3	23.0	22.7	17.3	13.6	7.7	5.9	32.7
CP/ET %	39.9	39.2	30.6	21.0	20.6	15.1	11.3	5.6	3.9	32.2
CP/ET Lower Conf. Limit* %	38.5	38.3	29.9	19.0	18.5	12.9	8.9	3.4	1.9	31.7
CP/ET %: 2 Embr. Transf. + min. 2 2-PN Surplus	45.4	45.3	38.9	27.0	28.4	20.8	18.6	10.1	12.8	40.7
CP/ET %: 1 Embr. Transf. + min. 3 2-PN Surplus	39.2	38.8	33.0	29.4	22.1	15.0	16.7	33.3	12.5	35.4
Misc./CP Upper Conf. Limit* %	16.0	16.8	24.3	41.5	41.8	57.5	60.5	86.3	83.1	21.1
Misc./CP %	14.4	15.8	23.1	36.4	36.3	49.7	49.4	68.0	57.1	20.4
Misc./CP Lower Conf. Limit* %	12.7	14.7	21.9	31.3	30.8	41.9	38.2	49.7	31.2	19.7
Births/ET Upper Conf. Limit* %	32.1	30.6	21.7	12.9	13.4	8.3	5.8	3.0	2.2	23.4
Births/ET %	30.7	29.7	21.1	11.3	11.7	6.7	4.2	1.8	1.1	23.0
Births/ET Lower Conf. Limit* %	29.3	28.9	20.4	9.8	10.0	5.2	2.7	0.6	0.0	2.5

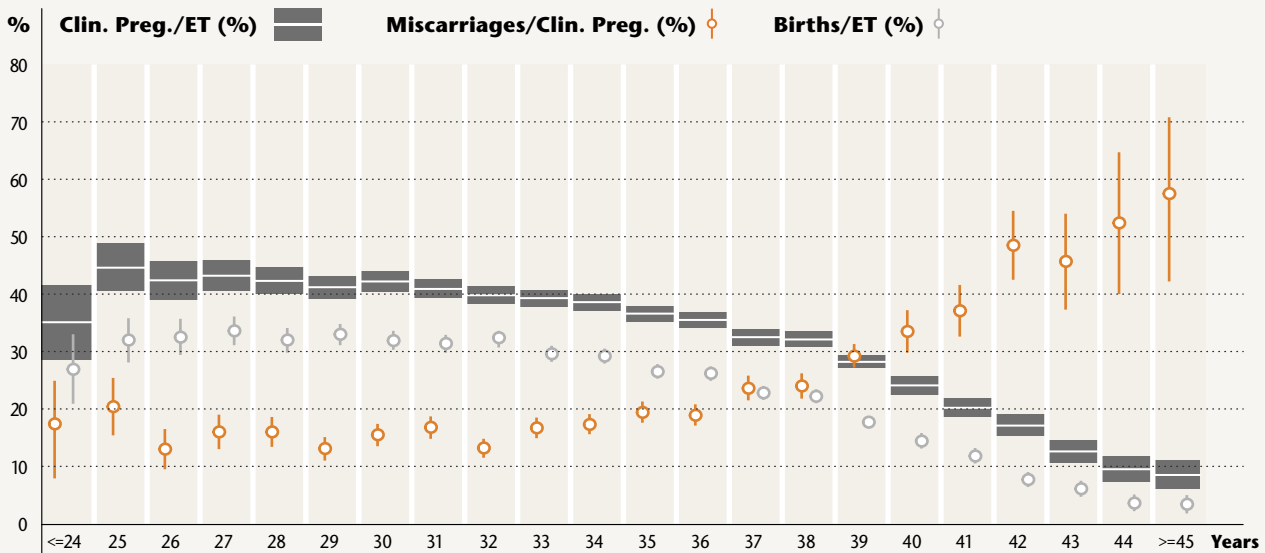
1) Mean
* With a 95%-probability, the true mean lies within the defined confidence interval.



Pregnancy Rate and Ongoing Pregnancy as a Function of Female Age 2015 – 2019

Prospective Data

IVF 2015 – 2019



n	208	560	839	1,360	1,824	2,467	3,057	3,421	3,967	4,325	4,591	4,861	4,765	4,518	4,695	6,665	2,624	2,210	1,566	1,100	662	472	ET
%	35.1	44.6	42.4	43.2	42.3	41.2	42.2	40.9	39.8	39.3	38.6	36.6	35.5	32.5	32.1	28.2	24.1	20.2	17.1	12.6	9.5	8.5	CP/ET
n	56	179	271	456	582	810	974	1,070	1,271	1,277	1,336	1,286	1,242	1,028	1,039	1,178	377	260	120	67	24	16	Birth

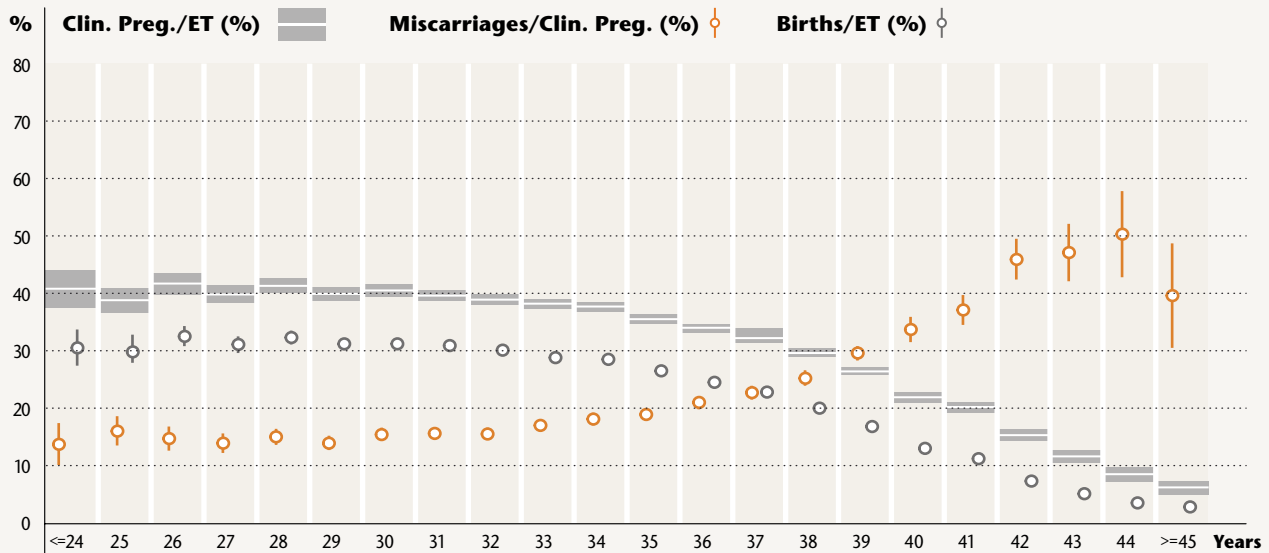
Alter in Jahren	<= 29	30 – 34	35 – 39	40	41	42	43	44	>=45	Gesamt
OPU	8,884	22,981	30,138	3,117	2,692	1,947	1,405	810	637	72,611
Oocytes ¹	11.7	10.5	8.5	6.9	6.6	6.1	5.4	5.1	4.2	9.2
Insemination ¹	11.3	10.1	8.1	6.6	6.3	5.8	5.3	5.0	4.2	8.8
Transfer	7,258	19,361	25,504	2,624	2,210	1,566	1,100	662	472	60,757
ET/OPU %	81.7	84.2	84.6	84.2	82.1	80.4	78.3	81.7	74.1	83.7
Trans. Embr. ¹	1.76	1.74	1.75	1.77	1.77	1.80	1.79	1.79	1.72	1.75
CP	3,044	7,716	8,301	630	445	268	138	63	40	20,645
CP/OPU %	34.3	33.6	27.5	20.2	16.5	13.8	9.8	7.8	6.3	28.4
CP/ET Upper Confidence Limit* %	43.2	40.7	33.2	25.8	21.9	19.0	14.5	11.8	11.0	34.5
CP/ET %	42.1	40.0	32.7	24.1	20.2	17.1	12.6	9.5	8.5	34.1
CP/ET Lower Confidence Limit* %	40.9	39.3	32.1	22.5	18.6	15.3	10.6	7.3	6.0	33.7
CP/ET %: 2 Embr. Transf. + min. 2 2-PN Surplus	47.5	46.6	39.8	32.8	29.4	25.4	13.2	10.0	8.9	39.4
CP/ET %: 1 Embr. Transf. + min. 3 2-PN Surplus	43.4	38.8	34.0	26.0	29.2	24.1	12.8	9.7	8.7	34.4
Misc./CP Upper Confidence Limit* %	16.3	16.7	24.0	37.2	41.6	54.5	54.0	64.7	72.8	21.0
Misc./CP %	15.0	15.9	23.1	33.5	37.1	48.5	45.7	52.4	57.5	20.5
Misc./CP Lower Confidence Limit* %	13.8	15.1	22.2	29.8	32.6	42.5	37.3	40.0	42.2	19.9
Births/ET Upper Confidence Limit* %	33.6	31.4	23.2	15.8	13.2	9.0	7.5	5.1	5.0	25.0
Births/ET %	32.5	30.7	22.7	14.4	11.8	7.7	6.1	3.6	3.4	24.6
Births/ET Lower Confidence Limit* %	31.5	30.1	22.2	13.1	10.5	6.4	4.7	2.2	1.8	24.3

1) Mean
* With a 95%-probability, the true mean lies within the defined confidence interval.

Pregnancy Rate and Ongoing Pregnancy as a Function of Female Age 2015 – 2019

Prospective Data

ICSI 2015 – 2019



n	838	2,106	2,746	4,083	5,798	7,463	9,105	10,430	11,628	12,491	13,398	14,289	14,027	13,744	13,927	19,011	7,949	6,475	4,898	3,296	2,004	1,807	ET
%	40.8	38.8	41.7	39.8	41.3	39.9	40.5	39.6	38.9	38.2	37.7	35.5	34.0	32.2	29.6	26.4	21.9	20.2	15.3	11.6	8.5	6.2	CP/ET
n	256	628	892	1,265	1,867	2,324	2,832	3,210	3,494	3,590	3,804	3,775	3,424	3,125	2,777	3,193	1,030	721	355	166	70	50	Birth

Alter in Jahren	<= 29	30 – 34	35 – 39	40	41	42	43	44	>=45	Gesamt
OPU	26,909	65,853	87,041	9,556	7,934	6,156	4,230	2,601	2,533	212,813
Oocytes ¹	12.4	11.2	8.9	7.3	6.8	6.3	5.9	5.4	4.8	9.7
Insemination ¹	9.5	8.7	7.0	5.7	5.4	5.0	4.6	4.3	3.7	7.5
Transfer	23,034	57,052	74,998	7,949	6,475	4,898	3,296	2,004	1,807	181,513
ET/OPU %	85.6	86.6	86.2	83.2	81.6	79.6	77.9	77.0	71.3	85.3
Trans. Embr. ¹	1.81	1.80	1.80	1.80	1.82	1.79	1.79	1.77	1.73	1.80
CP	9,287	22,111	23,354	1,733	1,304	749	380	169	111	59,198
CP/OPU %	34.5	33.6	26.8	18.1	16.4	12.2	9.0	6.5	4.4	27.8
CP/ET Upper Confidence Limit* %	41.0	39.2	31.6	22.8	21.2	16.3	12.7	9.7	7.3	32.9
CP/ET %	40.4	38.8	31.2	21.9	20.2	15.3	11.6	8.5	6.2	32.7
CP/ET Lower Confidence Limit* %	39.8	38.4	30.9	21.0	19.2	14.3	10.5	7.2	5.0	32.5
CP/ET %: 2 Embr. Transf. + min. 2 2-PN Surplus	44.6	44.0	38.5	29.0	28.3	22.6	16.0	16.9	15.7	40.4
CP/ET %: 1 Embr. Transf. + min. 3 2-PN Surplus	39.6	39.2	32.1	26.1	24.0	19.8	17.2	25.0	20.0	35.2
Misc./CP Upper Confidence Limit* %	15.2	16.9	24.0	35.9	39.7	49.5	52.1	57.8	48.7	20.9
Misc./CP %	14.5	16.4	23.4	33.7	37.1	45.9	47.1	50.3	39.6	20.6
Misc./CP Lower Confidence Limit* %	13.7	15.9	22.9	31.5	34.5	42.4	42.1	42.8	30.5	20.2
Births/ET Upper Confidence Limit* %	32.1	30.1	22.1	13.8	12.0	8.0	5.8	4.3	3.5	3.9
Births/ET %	31.5	29.7	21.8	13.0	11.2	7.3	5.1	3.5	2.8	23.7
Births/ET Lower Confidence Limit* %	30.9	29.4	21.5	12.3	10.4	6.5	4.3	2.7	2.0	23.5

1) Mean
* With a 95%-probability, the true mean lies within the defined confidence interval.

Results IVF, ICSI (COHS) and IVF and ICSI in Natural Cycles 2019

Prospective Data

IVF 2019

	n	%	Fertilization %	Embryo %	Transfer %	Clin. Preg. %
Performed ICSI Treatm.	16,989	100.0				
Successful Fertilization*	15,605	91.9	100.0			
Freeze All Oocyte Culture	1,212	7.1	7.8			
Minimum 1 Embryo**	14,338	90.9	99.6	100.0		
ET Performed	13,866	81.6	88.9	96.7	100.0	
Clin. Pregnancy	4,714	27.7	30.2	32.9	34.0	100.0
Birth	3,380	19.9	21.7	23.6	24.4	71.7
Miscarriage	951					20.2
Ectopic Pregnancy	65					1.4
Not Yet Recorded	318					6.7

ICSI 2019

	n	%	Fertilisation %	Embryo vorh.%	Transfer %	Klin. SS %
Performed ICSI Treatm.	44,506	100.0				
Successful Fertilization*	41,855	94.0	100.0			
Freeze All Oocyte Culture	3,363	7.6	8.0			
Minimum 1 Embryo**	38,320	93.1	99.6	100.0		
ET Performed	36,581	82.2	87.4	95.5	100.0	
Clin. Pregnancy	11,755	26.4	28.1	30.7	32.1	100.0
Birth	8,385	18.8	20.0	21.9	22.9	71.3
Miscarriage	2,397					20.4
Ectopic Pregnancy	141					1.2
Not Yet Recorded	832					7.1

IVF and ICSI in Natural Cycles 2019**

	n	%	Behandlung %	Fertilisation %	Embryo vorh. %	Transfer %	Klin. SS %
Started Cycles	3,812	100.0					
No Oocyte Treatment	1,041	27.3					
Treatm. in Natural Cycles***	2,771	72.7	100.0				
Successful Fertilization*	2,353	61.7	84.9	100.0			
Freeze All Oocyte Culture	343	9.0	12.4	14.6			
Minimum 1 Embryo**	1,988	57.3	81.9	98.9	100.0		
ET Performed	1,920	50.4	69.3	81.6	96.6	100.0	
Clin. Pregnancy	518	13.6	18.7	22.0	26.1	27.0	100.0
Birth	381	10.0	13.7	16.2	19.2	19.8	73.6
Miscarriage	112						21.6
Ectopic Pregnancy	7						1.4
Not Yet Recorded	18						3.5

*) Successful fertilization of at least one oocyte per cycle.

**) %-rates adjusted by freeze all oocytes culture.

***) Selection of cycles: without stimulation or stimulation only by Clomiphene or Letrozole.

Results of Thawing-Cycles, TESE, IVF and ICSI with Donor Semen 2019

Prospective Data

Cryo Transfer Cycles 2019

	n	%	PN/Embryo %	Transfer %	Clin. Preg. %
Cryo Transfer Cycles	28,961	100.0			
Thawed PN/Embryo	28,709	99.1	100.0		
ET Performed	27,554	95.1	96.0	100.0	
Clin. Pregnancy	8,262	28.5	28.8	30.0	100.0
Birth	5,647	19.5	19.7	20.5	68.3
Miscarriage	2,011				24.3
Ectopic Pregnancy	83				1.0
Not Yet Recorded	521				6.3

TESE 2019

	n	%	Fertilization %	Embryo %	Transfer %	Clin. Preg. %
Perf. ICSI/TESE Treatm.	2,159	100.0				
Successful Fertilization*	1,980	91.7	100.0			
Freeze All Oocyte Culture	169	7.8	8.5			
Minimum 1 Embryo**	1,777	89.3	98.1	100.0		
ET Performed	1,688	78.2	85.3	95.0	100.0	
Clin. Pregnancy	491	22.7	24.8	27.6	29.1	100.0
Birth	366	17.0	18.5	20.6	21.7	74.5
Miscarriage	94					19.1
Ectopic Pregnancy	5					1.0
Not Yet Recorded	26					5.3

IVF and ICSI with Donor Semen 2019

	n	%	Fertilization %	Embryo %	Transfer %	Clin. Preg. %
ART-Treatm. (donor sperm)***	1,404	100.0				
Successful Fertilization*	1,353	96.4	100.0			
Freeze All Oocyte Culture	87	6.2	6.4			
Minimum 1 Embryo**	1,251	95.0	98.8	100.0		
ET Performed	1,189	84.7	87.9	95.0	100.0	
Clin. Pregnancy	435	31.0	32.2	34.8	36.6	100.0
Birth	319	22.7	23.6	25.5	26.8	73.3
Miscarriage	99					22.8
Ectopic Pregnancy	4					0.9
Not Yet Recorded	13					3.0

*) Successful fertilization of at least one oocyte per cycle.

**) %-rates adjusted by freeze all oocytes culture.

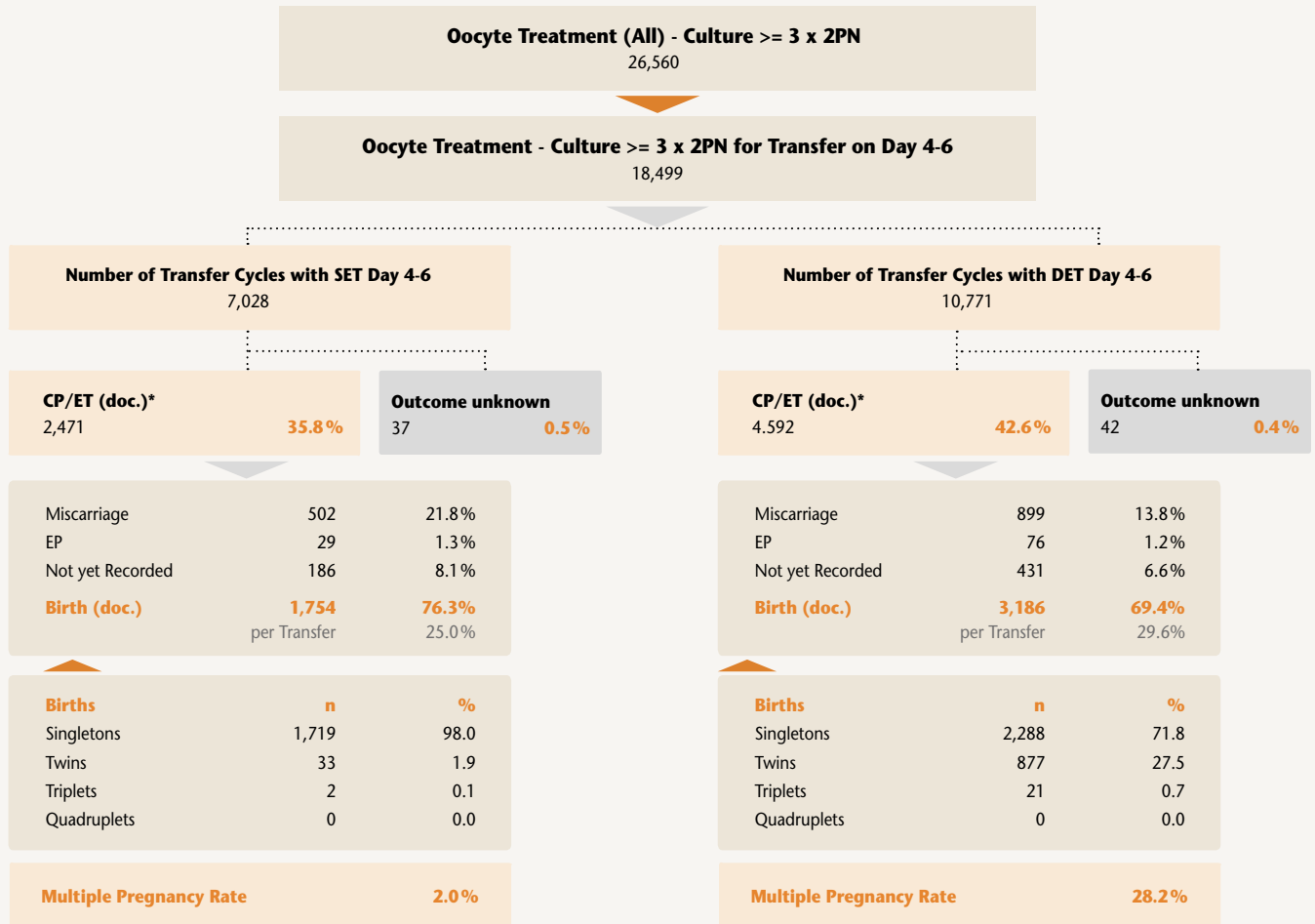
***) Currently, the data processing is not completely validated. Only cycles with definite data validation were included in this analysis.



Culture According to the "German Middle Way" and Impact on Therapy Outcome – Fresh Cycles 2020

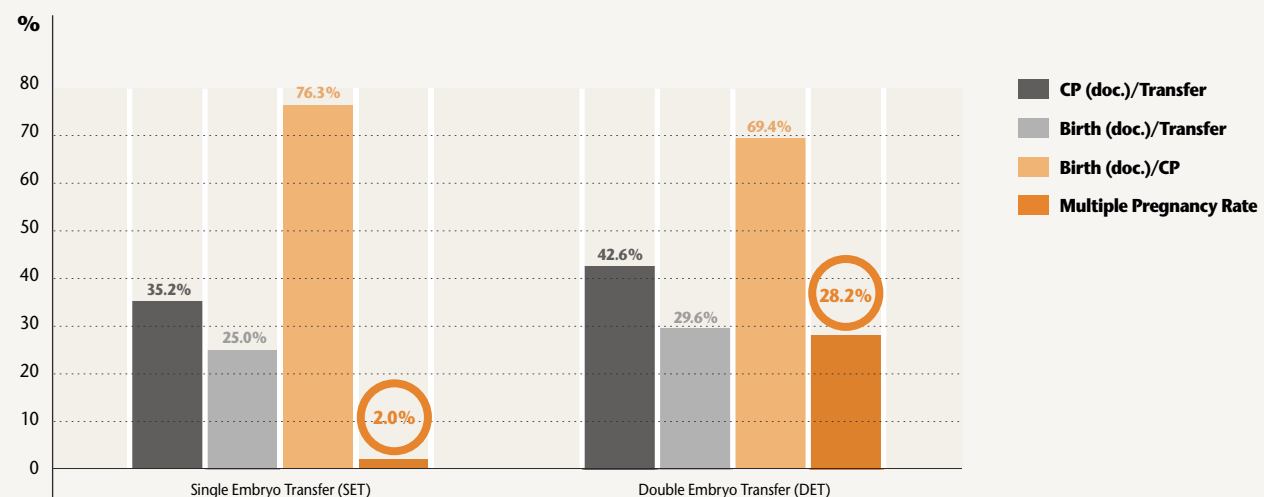
Prospective and Retrospective Data

Number of centers choosing $\geq 3 \times 2PN$ for extended culture: n=128



*) please note: documented clinical pregnancies (10,418) per transfer on other days than 4-6 (35,916): 29.0%

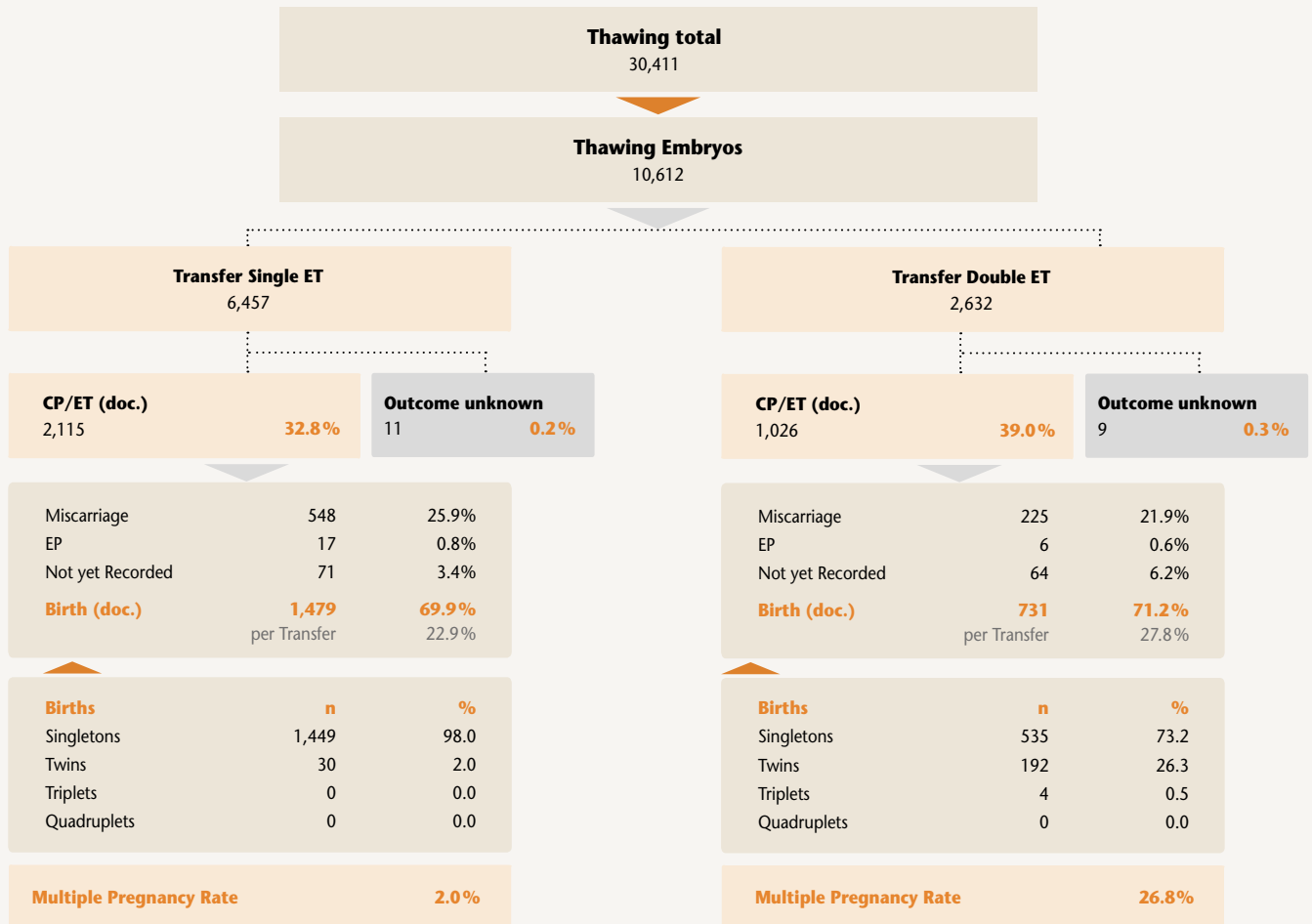
Comparison SET and DET "German Middle Way" Fresh Cycles 2019



Culture According to the "German Middle Way" and Impact on Therapy Outcome – Thawing Cycles Embryos 2019

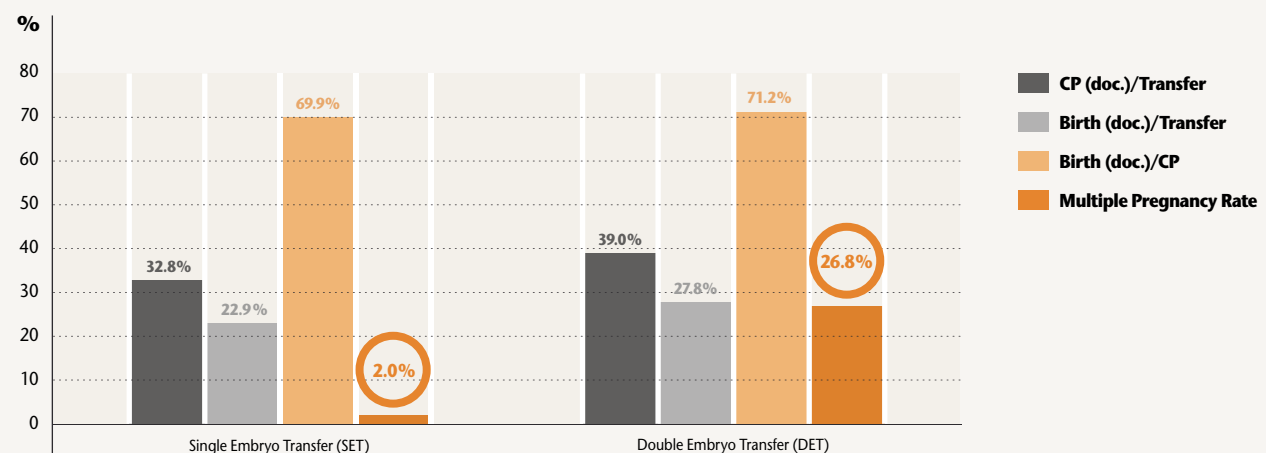
Prospective and Retrospective Data

Number of centers transferring previously cryopreserved embryos: n=121



Compared to thawing 2PNs:
 Documented clinical pregnancies (5,066) / transfers with outcome known(18,262): 27.7%
 - thereof day of transfer 2/3: documented clinical pregnancies (2,289) / transfers with outcome known (10,388): 22.0%
 - thereof day of transfer 5/6: documented clinical pregnancies (2,390) / transfers with outcome known (6,511): 36.70%
 - 1,363 transfers could not be allocated clearly.

Comparison SET and DET "German Middle Way" Thawing Cycles Embryos 2019



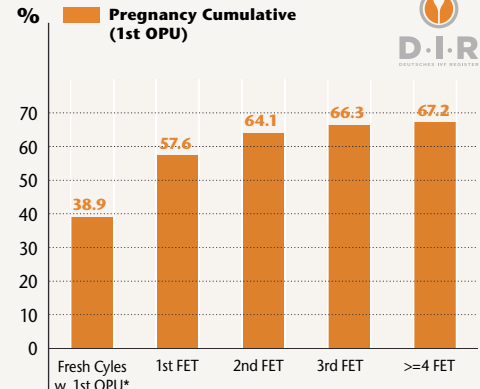
Pregnancies Cumulative 2017 – 2019 Based on First OPU



IVF, ICSI, Cryo Cycles – Prospective Data

The analysis deals with cumulative pregnancy rates after the first oocyte pickup (OPU) with fresh embryo transfer (ET) as well as the consecutive frozen embryo transfers (FET) including all cycles for which a cryopreservation of surplus fertilized oocytes had been available. Cycles without fresh transfer (freeze all) were excluded. Cryopreservation of surplus fertilized oocytes had been performed for 34% of the primary oocyte pickups.

For that subgroup the pregnancy rate (PR) after fresh embryo transfer was 39% of all patients of all ages. The pregnancy rates for up to four transfers added up to a cumulative rate of 67%.



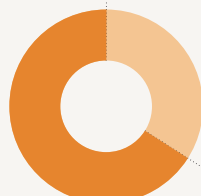
2017 – 2019 (Total)	1st OPU*	Cryo Cycles Based on 1st OPU	CP	CP/OPU / CP/FET (%)	CP Cumulative	CP/OPU Cumulative (%)
Fresh Cycles with 1st OPU*	32,725		12,724	38.9	12,724	38.9
1st Transfer (FET)		20,902	6,113	29.2	18,837	57.6
2nd Transfer (FET)		8,201	2,151	26.2	20,988	64.1
3rd Transfer (FET)		2,824	695	24.6	21,683	66.3
>= 4th Transfer (FET)		975	317	32.5	22,000	67.2
1st OPU 2017	1st OPU*	Cryo Cycles Based on 1st OPU	CP	CP/OPU / CP/FET (%)	CP Cumulative	CP/OPU Cumulative (%)
Fresh Cycles with 1st OPU*	11,819		4,644	39.3	4,644	39.3
1st Transfer (FET)		7,964	2,244	28.2	6,888	58.3
2nd Transfer (FET)		3,150	798	25.3	7,686	65.0
3rd Transfer (FET)		1,087	260	23.9	7,946	67.2
>= 4th Transfer (FET)		386	131	33.9	8,077	68.3
1st OPU 2018	1st OPU*	Cryo Cycles Based on 1st OPU	CP	CP/OPU / CP/FET (%)	CP Cumulative	CP/OPU Cumulative (%)
Fresh Cycles with 1st OPU*	10,336		3,969	38.4	3,969	38.4
1st Transfer (FET)		6,674	2,033	30.5	6,002	58.1
2nd Transfer (FET)		2,651	722	27.2	6,724	65.1
3rd Transfer (FET)		924	230	24.9	6,954	67.3
>= 4th Transfer (FET)		325	102	31.4	7,056	68.3
1st OPU 2019	1st OPU*	Cryo Cycles Based on 1st OPU	CP	CP/OPU / CP/FET (%)	CP Cumulative	CP/OPU Cumulative (%)
Fresh Cycles with 1st OPU*	10,570		4,111	38.9	4,111	38.9
1st Transfer (FET)		6,264	1,836	29.3	5,947	56.3
2nd Transfer (FET)		2,400	631	26.3	6,578	62.2
3rd Transfer (FET)		813	205	25.2	6,783	64.2
>= 4th Transfer (FET)		264	84	31.8	6,867	65.0

*) Cycles without fresh transfer (freeze all) and cycles without cryopreservation were excluded. Sample sizes: 1st OPU = 106,721; 1st OPU without freeze all = 96,354; Cycles without cryopreservation = 63,629; Cycles with Cryopreservation = 32,725.

1st OPU without Freeze All: Ratio Cycles with and without Cryopreservation



1st OPUs WITHOUT Cryopreservation
66%



1st OPUs WITH Cryopreservation
34%

The analysis reveals for the first time the increased cumulative probability of pregnancy when cryopreservation of surplus fertilized oocytes is possible and achievable.

Pregnancies Cumulative 2016 – 2019 as a Function of Female Age

IVF, ICSI, Cryo Cycles – Prospective Data

Age Groups ≤ 29	Retrievals	CP (Fresh Cycles)	CP (Fresh Cycles) in %	Cryo Cycle w. Transfer	CP (Cryo Cycles)	CP/ET (Cryo Cycles) in %	Cum. CP	Cumulative Pregnancy Rate in %
1. Transfer	12,358	5,198	42.1	2,473	903	36.5	6,101	41.1
2. Transfer	4,039	1,583	39.2	4,717	1,566	33.2	9,250	62.4
3. Transfer	1,852	700	37.8	2,618	799	30.5	10,749	72.5
4. Transfer	861	312	36.2	1,312	386	29.4	11,447	77.2
>4 Transfers	528	236	44.7	828	353	42.6	12,036	81.2
Age Groups 30-34	Retrievals	CP (Fresh Cycles)	CP (Fresh Cycles) in %	Cryo Cycle w. Transfer	CP (Cryo Cycles)	CP/ET (Cryo Cycles) in %	Cum. CP	Cumulative Pregnancy Rate in %
1. Transfer	28,646	11,638	40.6	6,190	2,259	36.5	13,897	39.9
2. Transfer	9,853	3,668	37.2	10,808	3,584	33.2	21,149	60.7
3. Transfer	5,048	1,791	35.5	6,213	1,839	29.6	24,779	71.1
4. Transfer	2,391	860	36.0	3,384	981	29.0	26,620	76.4
>4 Transfers	1,630	696	42.7	2,204	950	43.1	28,266	81.1
Age Groups 35-39	Retrievals	CP (Fresh Cycles)	CP (Fresh Cycles) in %	Cryo Cycle w. Transfer	CP (Cryo Cycles)	CP/ET (Cryo Cycles) in %	Cum. CP	Cumulative Pregnancy Rate in %
1. Transfer	33,461	10,839	32.4	6,305	2,041	32.4	12,880	32.4
2. Transfer	13,620	3,956	29.0	9,933	2,800	28.2	19,636	49.4
3. Transfer	7,204	2,024	28.1	5,718	1,528	26.7	23,188	58.3
4. Transfer	3,241	895	27.6	3,262	805	24.7	24,888	62.6
>4 Transfers	2,161	793	36.7	2,283	864	37.8	26,545	66.8
Age Groups ≥ 40	Retrievals	CP (Fresh Cycles)	CP (Fresh Cycles) in %	Cryo Cycle w. Transfer	CP (Cryo Cycles)	CP/ET (Cryo Cycles) in %	Cum. CP	Cumulative Pregnancy Rate in %
1. Transfer	9,782	1,708	17.5	2,209	508	23.0	2,216	18.5
2. Transfer	3,940	586	14.9	2,196	409	18.6	3,211	26.8
3. Transfer	1,980	323	16.3	1,173	177	15.1	3,711	31.0
4. Transfer	1,008	135	13.4	613	82	13.4	3,928	32.8
>4 Transfers	620	142	22.9	453	105	23.2	4,175	34.8
2017 – 2019 (total)	Retrievals	CP (Fresh Cycles)	CP (Fresh Cycles) in %	Cryo Cycle w. Transfer	CP (Cryo Cycles)	CP/ET (Cryo Cycles) in %	Cum. CP	Cumulative Pregnancy Rate in %
1. Transfer	84,740	29,497	34.8	17,247	5,727	33.2	35,224	34.5
2. Transfer	31,655	9,854	31.1	27,773	8,396	30.2	53,474	52.4
3. Transfer	16,191	4,869	30.1	15,789	4,357	27.6	62,700	61.5
4. Transfer	7,554	2,223	29.4	8,613	2,263	26.3	67,186	65.9
>4 Transfers	4,961	1,874	37.8	5,794	2,279	39.3	71,339	70.0

Further improvements of these cumulative analysis could be implemented in the annual. These analysis now reflect the patient's perspective and therefore the patient's journey as exact as possible.

Positive Pregnancy Outcomes 2019



IVF, ICSI – Prospective and Retrospective Data

	Fresh Cycles		Cryo Cycles	
	n	%	n	%
Clinical Pregnancies	17,481	100.0	8,589	100.0
Outcome documented	17,218	98.5	8,451	98.4
Transfer	53,715		28,745	
Births	12,432		5,850	
Life-Birth-Rate/Birth	12,405	99.8	5,828	99.6
Life-Birth-Rate/ET	12,405	23.1	5,828	20.3
SET Pat. <= 35 Y., >= 4 2PN				
Number of Transfers	5,070			
Life-Birth-Rate/ET	1,472	29.0		
Multiple Pregnancies	27	1.8		
DET/TET Pat. <= 35 Y., >= 4 2PN				
Number of Transfers	12,701			
Life-Birth-Rate/ET	4,367	34.4		
Number of Multiple Births	1,314	30.1		

Loss of Pregnancy 2019

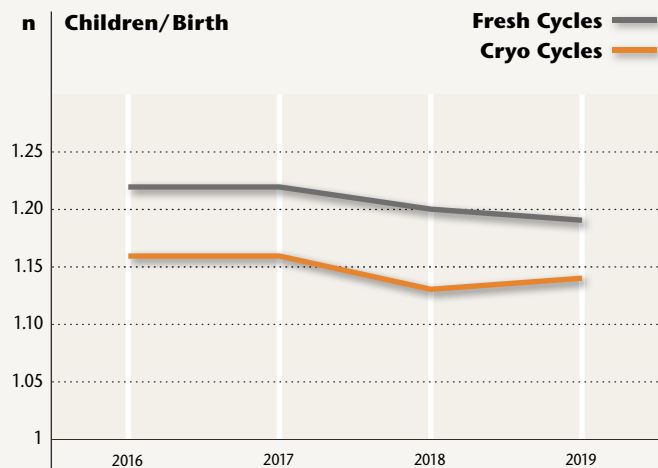
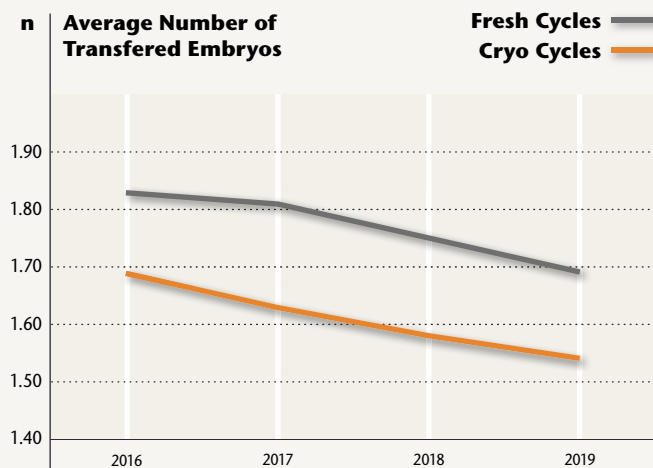


Prospective and Retrospective Data

	Fresh Cycles		Cryo Cycles	
	n	%	n	%
Clinical Pregnancies	17,481	100.0	8,589	100.0
Treatments with Known Cycle Outcome	17,218	98.5	8,451	98.4
Miscarriages	3,560	20.7	2,064	25.8
Among those: Induced Abortions	179	5.0	72	3.5
Stillbirths	27	0.2	22	0.3

Embryos per Transfer¹ and Children per Birth 1997 – 2019

IVF, ICSI – Prospective and Retrospective Data



	1997	[...]	2016	2017	2018	2019
Fresh Cycles	Average Number of Transf. Embryos	2.56	1.83	1.81	1.75	1.69
	Children/Birth	1.29	1.22	1.22	1.20	1.19
Cryo Cycles	Average Number of Transf. Embryos	2.34	1.69	1.63	1.58	1.54
	Children/Birth	1.14	1.16	1.16	1.13	1.14

For values from 1998 to 2015 see www.deutsches-ivf-register.de

The number of children originating from multiple pregnancies has almost remained stable within the last decades. But In 2018 and 2019, we are glad to notice a trend to a reduction.

In summary: the number of embryos per transfer is an important factor worth looking at. It is important to allow children resulting from an ART-treatment to come into life as a singleton. Siblings are important and desirable – one at a time!

1) Mean

Oocyte Maturity Depending on Stimulation Protocol 2020

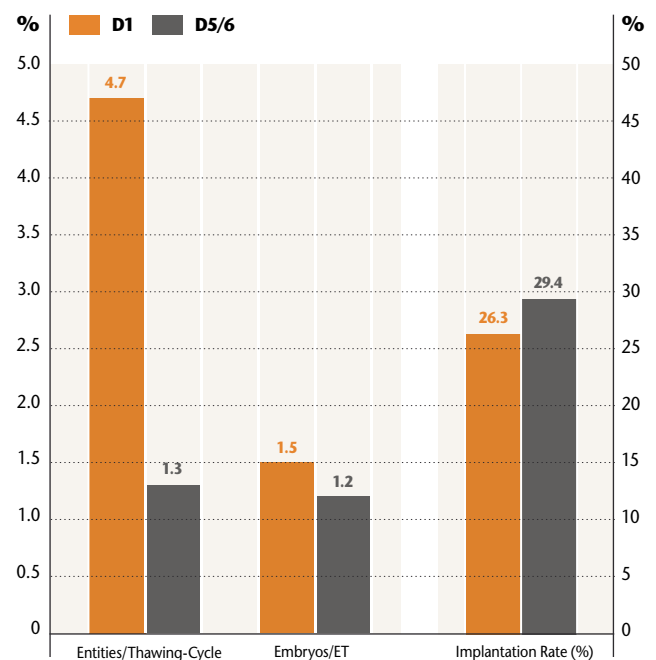
ICSI – Prospective and Retrospective Data

	OPU with Oocytes Maturity known	Mean Age	Oocytes Retrieved	Ooc. per Retrieval	Mature Oocytes	Mature Ooc. per Retrieval	2PN	2PN/Oocyte Retrieval	Embryos for ET	Gestational Sacs	Implant. Rate (%)
Agonist	7,024	35.7	64,689	9.2	47,235	6.7	31,469	4.5	9,622	2,108	21.9
Antagonist	29,654	35.1	288,855	9.7	214,085	7.2	145,040	4.9	38,405	8,471	22.1

Cycles and Implantation-Rates with Transfer of Day 5/6 Embryos after Cryopreservation either on Culture Day 1 or Day 5/6 2020

Thawing – Prospective and Retrospective Data

	Freezing on Culture D1	Freezing on Culture D5/6
Thawing-Cycles resulting in ET	7,585	10,022
PN/Embryos Thawed	35,610	13,397
Entities/Thawing-Cycle	4.7	1.3
Re-Cryo of Embryos	2,227	28
Embryos Available for Transfer	11,502	11,904
Re-Cryo/Cycle	0.3	0.0
Transfer	7,585	10,022
Gestational Sacs (Intrauterine)	3,022	3,494
Embryos/ET	1.5	1.2
Implantation Rate (%)	26.3	29.4

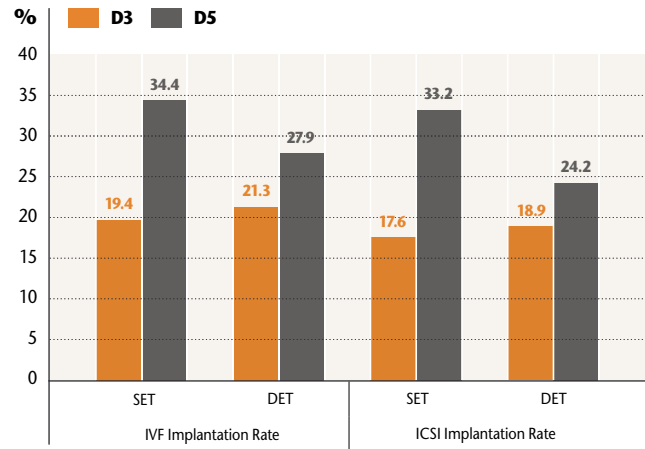




Implantation Rates of D3- and D5-Embryos and Number of Embryos per Transfer 2020

IVF, ICSI – Prospective and Retrospective Data

		D3 ImplRate %	D5 ImplRate %
IVF Implantation Rate	SET	19.7	34.4
	DET	21.3	27.9
ICSI Implantation Rate	SET	17.6	33.2
	DET	18.9	24.2

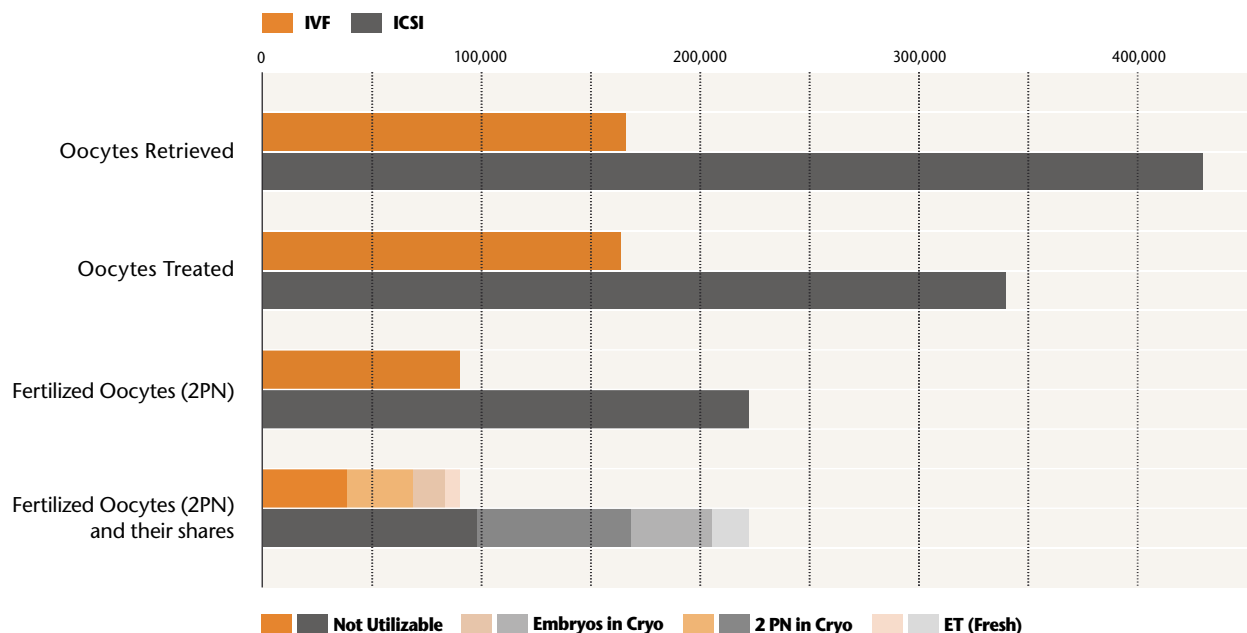


Evolution of Retrieved Oocytes (IVF or ICSI) 2020



IVF, ICSI – Prospective and Retrospective Data

	IVF		%		ICSI		%	
Oocytes Retrieved	165,637	100.0			429,862	100.0		
Oocytes Treated	163,399	98.6	100.0		339,517	79.0	100.0	
Fertilized Oocytes (2PN)	90,123		55.2	100.0	221,995		65.4	100.0
2PN Cryopreserved	30,071			33.4	70,450			31.7
ET (Fresh)	15,018			16.7	36,832			16.6
Embryos Cryopreserved	6,463			7.2	16,655			7.5



Clinical Pregnancies (CP)/Fresh Transfer as a Function of Embryo Quality 2020

IVF, ICSI, IVF/ICSI – Prospective Data

Quality		<= 29 Years		30 – 34 Years		35 – 39 Years		>= 40 Years		Total*	
Ideal	Not Ideal	ET	CP/ET %	ET	CP/ET %	ET	CP/ET %	ET	CP/ET %	ET	CP/ET %
0	1	240	17.6	806	14.5	1,197	12.9	537	2.6	2,780	11.8
0	2	278	22.3	747	27.5	1,288	21.1	434	11.0	2,474	21.4
0	3	6	16.7	18	11.8	21	15.0	29	3.4	74	9.7
1	0	1,637	38.6	5,235	35.5	6,982	28.0	2,717	14.1	16,571	29.2
1	1	289	34.9	1,017	30.6	1,491	28.5	564	18.0	3,361	27.9
1	2	25	40.0	61	40.0	100	26.8	43	20.0	229	30.6
2	0	2,427	49.1	7,066	45.6	8,737	37.8	2,861	23.3	21,092	39.8
2	1	6	33.3	22	28.6	46	37.8	64	26.6	138	30.9
3	0	18	41.2	113	27.1	30	32.5	267	19.7	628	26.3
Total*		4,978	41.5	15,204	38.2	20,244	30.9	7,574	17.1	48,001	32.1

*) 654 transfers could not be allocated.

Clinical Pregnancies (CP)/Frozen Transfer as a Function of Embryo Quality 2020

Cryo Transfer – Prospective Data

Quality		IVF		ICSI	
ideal	Not Ideal	ET	CP/ET %	ET	CP/ET %
0	1	432	15.5	1,224	14.8
0	2	401	16.3	1,241	19.0
0	3	5	40.0	48	12.5
1	0	3,955	30.2	9,241	31.5
1	1	666	28.3	1,481	27.9
1	2	11	27.3	54	22.2
2	0	2,829	33.4	6,866	33.6
2	1	25	44.0	73	20.5
3	0	71	28.2	234	28.8
Total**		8,680	32.5	21,191	29.8

***) 1,014 transfers could not be allocated.

Children as a Function of Week of Gestation (WoG) and Birth Weight (BW) 2019

Prospective and Retrospective Data

IVF, ICSI, IVF/ICSI

Current WoG	20 - 26		27 - 31		32 - 37		38 - 41		≥ 42		Total	% of total
Singletons (n and %)	36	0.4	93	1.0	853	9.5	6,954	77.1	1,087	12.0	9,023	69.4
Average Birth Weight (g)	630		1,063		2,463		3,332		3,578		3,245	
Twins (n and %)	76	2.0	282	7.3	1,860	48.3	1,606	41.7	18	0.5	3,848	29.6
Average Birth Weight (g)	609		1,196		2,222		2,714		2,930		2,327	
Triplets (n and %)	12	9.5	39	31.0	72	57.1	3	2.4	-	-	126	1.0
Average Birth Weight (g)	638		1,039		1,856		3,293		-		1,528	

Percentage of preterm deliveries in singleton pregnancies is 10.9%.

Percentage of preterm deliveries in twin pregnancies is 57.6%.

Percentage of preterm deliveries in triplets pregnancies is 97.6%.

Cryo Transfer

Current WoG	20 - 26		27 - 31		32 - 37		38 - 41		≥ 42		Total	% of total
Singletons (n and %)	15	0.3	38	0.8	321	7.0	3,354	73.4	844	18.5	4,572	75.9
Average Birth Weight (g)	676		1,079		2,530		3,463		3,660		3,405	
Twins (n and %)	14	1.0	48	3.5	638	46.2	666	48.3	14	1.0	1,380	22.9
Average Birth Weight (g)	587		1,220		2,311		2,774		2,854		2,483	
Triplets (n and %)	6	8.3	24	33.3	42	58.3	-	-	-	-	72	1.2
Average Birth Weight (g)	690		1,033		1,736		-		-		1,433	

Percentage of preterm deliveries in singleton pregnancies is 8.2%.

Percentage of preterm deliveries in twin pregnancies is 50.7%.

Percentage of preterm deliveries in triplets pregnancies is 100.0%.

Children Born 1997 – 2019

Prospective and Retrospective Data

Total (IVF, ICSI, IVF/ICSI, Cryo Transfer)

	Singletons		Twins		Triplets		Quadruplets		Total
	n	%	n	%	n	%	n	%	n
1997	4,175	58.7	1,902	32.8	492	8.4	8	0.1	6,577
1998	5,357	58.2	3,152	34.2	702	7.6	0	-	9,211
1999	6,116	60.5	3,396	33.6	600	5.9	4	0.0	10,116
2000	6,143	60.5	3,504	34.5	507	5.0	4	0.0	10,158
2001	7,726	62.2	4,252	34.3	435	3.5	0	-	12,413
2002	8,930	63.8	4,662	33.3	387	2.8	8	0.1	13,987
2003	11,922	63.1	6,334	33.6	597	3.2	24	0.1	18,877
2004	6,891	65.6	3,336	31.8	273	2.6	0	-	10,500
2005	7,038	65.8	3,440	32.1	213	2.0	12	0.1	10,703
2006	7,419	66.9	3,450	31.1	222	2.0	4	0.0	11,095
2007	8,407	66.4	4,076	32.2	183	1.4	4	0.0	12,670
2008	8,444	65.7	4,142	32.3	240	1.9	8	0.1	12,834
2009	9,016	67.3	4,152	31.0	216	1.6	8	0.1	13,392
2010	8,619	66.2	4,156	31.9	249	1.9	0	-	13,024
2011	9,388	63.3	5,131	34.7	300	2.0	0	-	14,819
2012	10,188	66.4	4,906	32.0	249	1.6	0	-	15,343
2013	11,713	64.9	6,003	33.3	327	1.8	8	0.0	18,051
2014	13,092	65.5	6,566	32.9	309	1.5	12	0.1	19,979
2015	13,702	65.4	6,942	33.2	297	1.4	8	0.0	20,949
2016	13,692	66.0	6,800	32.8	258	1.2	4	0.0	20,754
2017	14,353	67.2	6,686	31.3	324	1.5	8	0.0	21,371
2018	15,088	69.7	6,332	29.3	222	1.0	0	-	21,642
2019	15,325	71.0	6,002	27.8	261	1.2	0	-	21,588
Total	222,744	65.5	109,322	32.1	7,863	2.3	124	0.0	340,053

WUPPERTAL
Population of 355,004

BIELEFELD
Population of 333,509

340,053

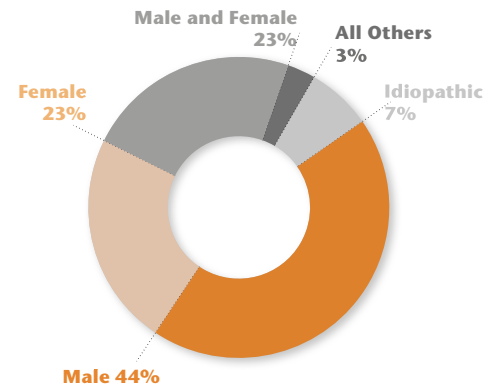
Children born 1997 – 2019

Distribution of Indications 2020

IVF and ICSI – Prospective Data

	Treatments		Transfers		Clin. Preg.	
	n		n	%	n	%
No Information	743		653	87.9	227	34.8
Idiopathic	4,163		3,246	78.0	1,069	32.9
Male	26,265		21,542	82.0	7,108	33.0
Female	13,622		10,751	78.9	3,392	31.6
Male and Female	13,936		10,959	78.6	3,231	29.5
Missing Male Partner	645		537	83.3	145	27.0
Homosexuality	354		313	88.4	121	38.7
Total	59,728		48,001	80.4	15,293	31.9

Shares of Indications (Cycles) 2020



IVF

Male Factor	Normal		Red. Semen Quality		Unknown		Other**		Total***	
	n	%	n	%	n	%	n	%	n	%
Female Factor										
Normal	3,132	14.7	639	3.0	91	0.4	473	2.2	4,335	20.3
Tubal Pathology	2,614	12.3	524	2.5	48	0.2	467	2.2	3,653	17.1
Endometriosis	2,068	9.7	440	2.1	45	0.2	469	2.2	3,022	14.2
Hyperandrog./PCO	654	3.1	131	0.6	16	0.1	170	0.8	971	4.6
Ovulatory Dysf.	966	4.5	378	1.8	42	0.2	285	1.3	1,671	7.8
Psychogen. Factors	25	0.1	6	0.0	2	0.0	8	0.0	41	0.2
Age	867	4.1	281	1.3	31	0.1	231	1.1	1,410	6.6
Other*	3,779	17.7	1,075	5.0	94	0.4	1,233	5.8	6,181	29.0
No Information	28	0.1	3	0.0	9	0.0	10	0.0	50	0.2
Total***	14,133	66.2	3,477	16.3	378	1.8	3,346	15.7	21,334	100.0

ICSI

Male Factor	Normal		Red. Semen Quality		Azoospermia		Unknown		Other**		Total***	
	n	%	n	%	n	%	n	%	n	%	n	%
Female Factor												
Normal	5,165	10.0	8,839	17.1	1,193	2.3	200	0.4	2,211	4.3	17,608	34.1
Tubal Pathology	901	1.7	1,837	3.6	70	0.1	43	0.1	664	1.3	3,515	6.8
Endometriosis	1,122	2.2	2,273	4.4	117	0.2	79	0.2	929	1.8	4,520	8.7
Hyperandrog./PCO	403	0.8	1,045	2.0	94	0.2	36	0.1	356	0.7	1,934	3.7
Ovulatory Dysf.	804	1.6	2,123	4.1	177	0.3	74	0.1	848	1.6	4,026	7.8
Psychogen. Factors	20	0.0	44	0.1	8	0.0	0	0.0	13	0.0	85	0.2
Age	930	1.8	1,706	3.3	229	0.4	73	0.1	692	1.3	3,630	7.0
Other*	5,056	9.8	6,944	13.4	730	1.4	215	0.4	3,276	6.3	16,221	31.4
No Information	73	0.1	29	0.1	5	0.0	6	0.0	16	0.0	129	0.2
Total***	14,474	28.0	24,840	48.1	2,623	5.1	726	1.4	9,005	17.4	51,668	100.0

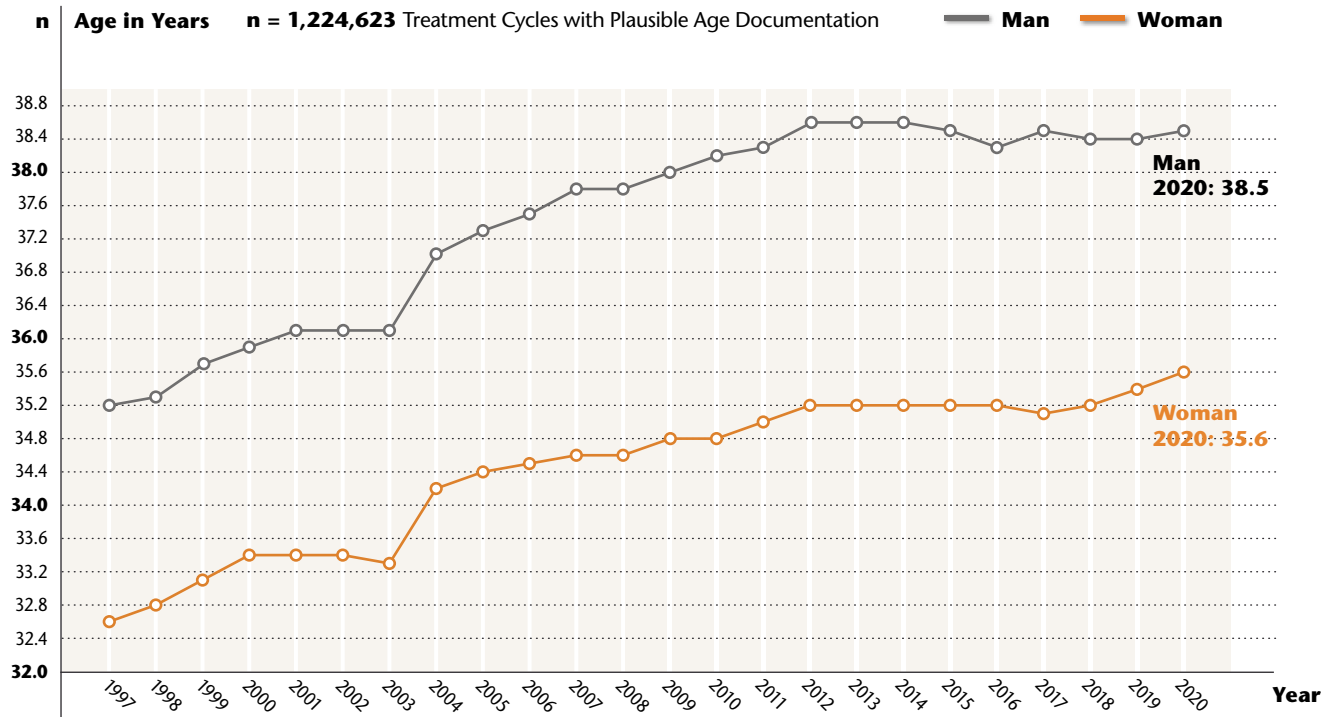
*) This includes the following indications: medical freezing, genetic testing (polar body analysis, trophoctoderm biopsy for PGT-A, PGT-SR, PGT-M), homosexuality, diminished ovarian reserve, social freezing, uterine or cervical factor, single women, others.

**) This includes following indications: anejaculation, azoospermia, medical freezing, preimplantation genetic testing, psychogenic disturbances, failed or bad fertilization rate in conventional ivf-procedure, CBAVD, retrograde ejaculation, urogenital malformation, condition after genital cancer, condition after severe genital infection, condition after vasectomy, others.

***) Multiple answers per cycle permitted.

Mean Age for Women and Men 1997 – 2020

IVF, ICSI, IVF/ICSI – Prospective and Retrospective Data



Social Freezing 2017 – 2020

Indication Social Freezing (Therapy or Patient)

Fresh Cycles – Prospective and Retrospective Data

	2020	2019	2018	2017*
No. of Centers	98	89	84	75
Recorded Cycles	1,565	1,171	969	667
Plausible Cycles	1,448	1,159	963	659
Plausible Cycles %	92.5	99.0	99.4	98.8
Ø-Age of Patients	35.7	35.6	35.5	35.5
OPU	1,324	1,075	899	616
Oocytes Aspirated	1,281	1,043	875	598
Avg. Oocytes Aspirated / Cycle	10.5	10.6	10.3	10.0
Freeze All MII	1,180	975	795	488

Follow up social freezings: up to now, only a few pregnancies and births out of former social freezing cycles are recorded.

* Possibility of documentation of social freezing only started in 2017.

Clinical Pregnancy Rate as a Function of Stimulation 2020



Prospective Data

Total	recFSH	hMG	recFSH a. recLH	recFSH a. hMG	Long-Acting recFSH	hrFSH	Antiestrogen +/- Gonadotropin	Other*	No Inform.	Total
Stimulations (n)	25,285	7,453	16,042	6,003	2,528	767	3,290	3,134	1,848	66,350
Transfers (n)	18,699	4,960	11,638	4,165	1,793	502	1,663	2,458	1,054	46,932
Transfer (%)	74.0	66.6	72.5	69.4	70.9	65.4	50.5	78.4	57.0	70.7
CP (n)	6,587	1,307	3,799	1,121	508	187	332	811	371	15,023
CP/ET (%)	35.2	26.4	32.6	26.9	28.3	37.3	20.0	33.0	35.2	32.0
CP/Stim. (%)	26.1	17.5	23.7	18.7	20.1	24.4	10.1	25.9	20.1	22.6
Ø-Age of Patients	33.9	37.6	35.6	36.5	36.9	33.8	38.5	35.4	35.9	35.4

Short GnRH _a	recFSH	hMG	recFSH a. recLH	recFSH a. hMG	Long-Acting recFSH	hrFSH	Antiestrogen +/- Gonadotropin	Other*	No Inform.	Total	Share (%) from total
Stimulations (n)	438	425	551	411	32	50	122	27	44	2,100	3.2
Transfers (n)	297	273	386	208	17	40	32	13	35	1,301	2.8
Transfer (%)	67.8	64.2	70.1	50.6	53.1	80.0	26.2	48.1	79.5	62.0	
CP (n)	73	69	119	50	7	14	6	5	11	354	2.4
CP/ET (%)	24.6	25.3	30.8	24.0	41.2	35.0	18.8	38.5	31.4	27.2	
CP/Stim. (%)	16.7	16.2	21.6	12.2	21.9	28.0	4.9	18.5	25.0	16.9	
Ø-Age of Patients	35.4	38.0	36.9	37.2	37.9	34.0	39.3	37.9	35.9	36.9	

Long GnRH _a	recFSH	hMG	recFSH a. recLH	recFSH a. hMG	Long-Acting recFSH	hrFSH	Antiestrogen +/- Gonadotropin	Other*	No Inform.	Total	Share (%) from total
Stimulations (n)	2,738	1,562	2,738	1,479	164	36	22	464	291	9,494	14.3
Transfers (n)	2,137	1,181	2,096	1,132	125	22	9	336	190	7,228	15.4
Transfer (%)	78.0	75.6	76.6	76.5	76.2	61.1	40.9	72.4	65.3	76.1	
CP (n)	770	339	681	323	34	6	1	109	69	2,332	15.5
CP/ET (%)	36.0	28.7	32.5	28.5	27.2	27.3	11.1	32.4	36.3	32.3	
CP/Stim. (%)	28.1	21.7	24.9	21.8	20.7	16.7	4.5	23.5	23.7	24.6	
Ø-Age of Patients	34.0	36.8	35.6	36.1	37.6	33.2	38.4	36.3	36.7	35.5	

GnRH-Antagonists	recFSH	hMG	recFSH a. recLH	recFSH a. hMG	Long-Acting recFSH	hrFSH	Antiestrogen +/- Gonadotropin	Other*	No Inform.	Total	Share (%) from total
Stimulations (n)	20,472	4,682	11,805	3,761	2,131	530	2,155	2,285	1,210	49,031	73.9
Transfers (n)	15,364	3,087	8,675	2,646	1,538	332	1,250	1,819	670	35,381	75.4
Transfer (%)	75.0	65.9	73.5	70.4	72.2	62.6	58.0	79.6	55.4	72.2	
CP (n)	5,472	804	2,859	705	435	121	252	603	241	11,492	76.5
CP/ET (%)	35.6	26.0	33.0	26.6	28.3	36.4	20.2	33.2	36.0	32.5	
CP/Stim. (%)	26.7	17.2	24.2	18.7	20.4	22.8	11.7	26.4	19.9	23.4	
Ø-Age of Patients	33.8	37.8	35.5	36.5	36.8	33.9	38.6	35.5	35.7	35.3	

4,724 stimulations (7.1 %) have been realized without Analoga or Antagonists, resulting in 2,226 transfers and 660 clinical pregnancies (29.1 %).

1,003 stimulations (1.5 %) could not be allocated to a specific protocol, resulting in 756 transfers and 185 clinical pregnancies (24.7 %).

*) e.g. uFSH, uFSH and hMG etc.

Ovarian Hyperstimulation Syndrome (OHSS) as a Function of Stimulation Protocol and Age Cohort 2020

IVF, ICSI, IVF/ICSI – Prospective Data

	Stimulations Started	%	Oocytes Retrieved	OHSS III (WHO)	OHSS III/Cycles %
Short GnRHa	2,100	3.2	7.1	1	0.0
<= 29 Years	130		10.0	0	0.0
30 – 34 Years	449		9.0	0	0.0
35 – 39 Years	930		6.6	0	0.0
>= 40 Years	591		5.5	1	0.2
Long GnRHa	9,494	14.5	9.3	50	0.5
<= 29 Years	840		11.8	15	1.8
30 – 34 Years	2,905		10.5	18	0.6
35 – 39 Years	4,271		8.7	17	0.4
>= 40 Years	1,478		7.0	0	0.0
GnRHa-Antagonists	49,031	75.0	9.5	133	0.3
<= 29 Years	5,483		12.2	34	0.6
30 – 34 Years	15,389		11.1	62	0.4
35 – 39 Years	20,165		8.9	33	0.2
>= 40 Years	7,994		6.2	4	0.1
No Analoga / no Antagonists	4,722	7.2	6.6	3	0.1
<= 29 Years	395		9.8	0	0.0
30 – 34 Years	1,222		9.2	2	0.2
35 – 39 Years	2,010		6.3	1	0.0
>= 40 Years	1,095		3.8	0	0.0
Total*	65,347	100	9.2	187	0.3

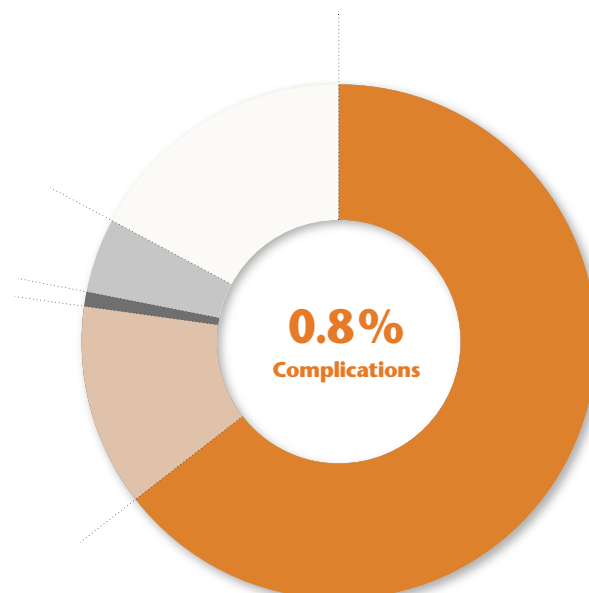
*) in 1,003 cycles, the protocol could not be reliably determined

Complications as a Function of Ovum Pick-up (OPU) 2020

IVF, ICSI, IVF/ICSI, Prospective Data

Total OPU's	62,199	100.0%
No Information	539	0.9%
No Complications	61,212	98.4%
Complications	448	0.8%

Complications	n	%
Vaginal Bleeding	289	64.5
Intraabdom. Bleeding	57	12.7
Intestinal Tract Injury	4	0.9
Peritonitis	21	4.7
Other	77	17.2
Total	448	100.0

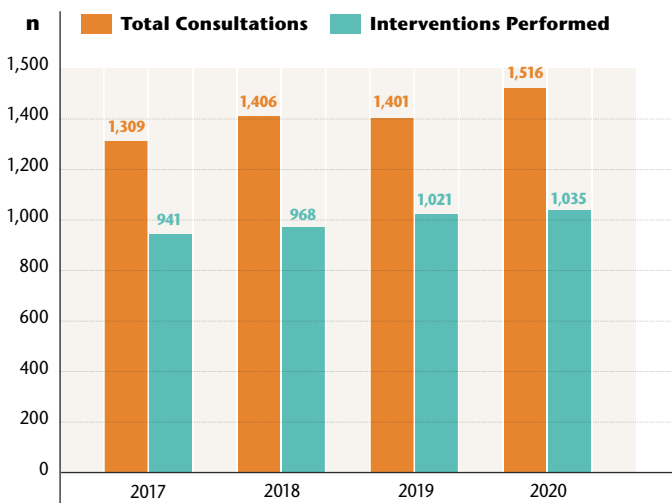




The *FertiPROTEKT* Network e.V. is pleased to present again the data from all documented consultations and therapies in the scope of medically indicated fertility preservation from the recent and the previous years. To give an impression of the excellent activities of the network the following tables and figures list important individual entities and therapies for fertility preservation.

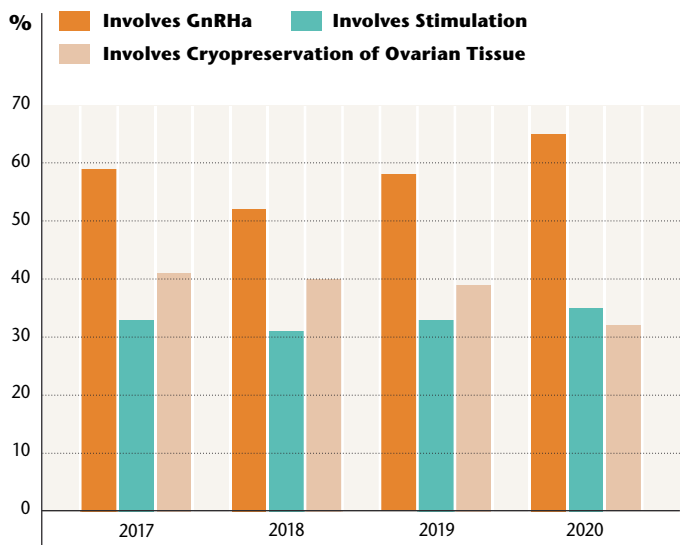
In the coming years, it remains to be seen whether the implementation of the new law regarding the assumption of costs for fertility preservation will have an effect on the amount and type of therapies carried out; this taking also into account that GnRH agonists are not and cryopreservation of germ cells/tissue is somewhat limited covered. Hope remains that many women will get the opportunity for counselling.

Consultations and Interventions – *FertiPROTEKT* Network e.V.



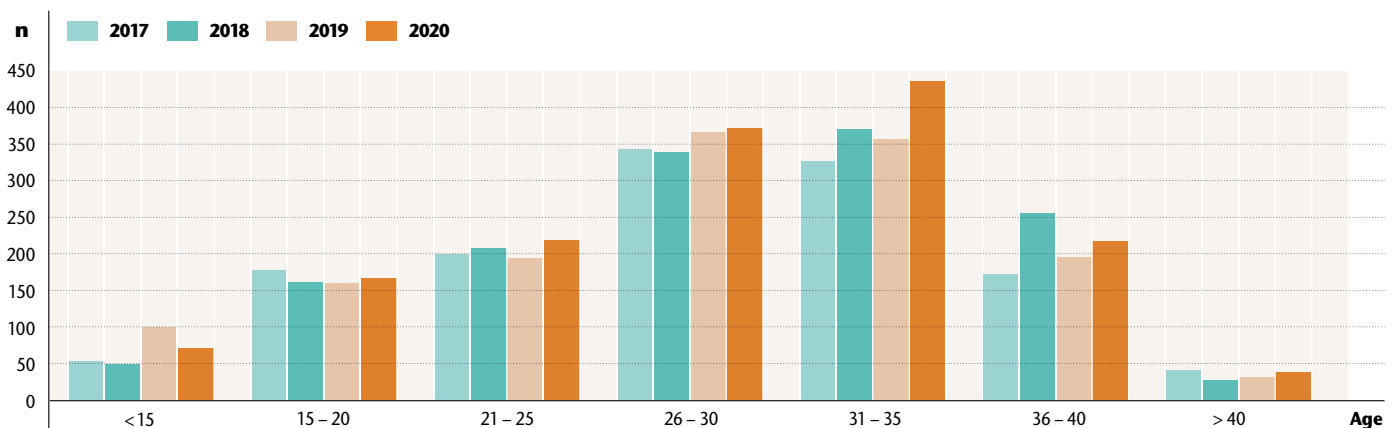
The number of consultations documented in the *FertiPROTEKT* Network e.V. rose again in 2020 compared to previous years. Interventions have also increased compared to previous years, so that a very positive upward trend can be seen here. While the ovarian stimulations performed remained stable,

Frequency Distribution of Fertility-Protective Interventions



GnRH agonists in particular were used significantly more frequently. However, a downward trend in the cryopreservation of ovarian tissue has been observed in recent years.

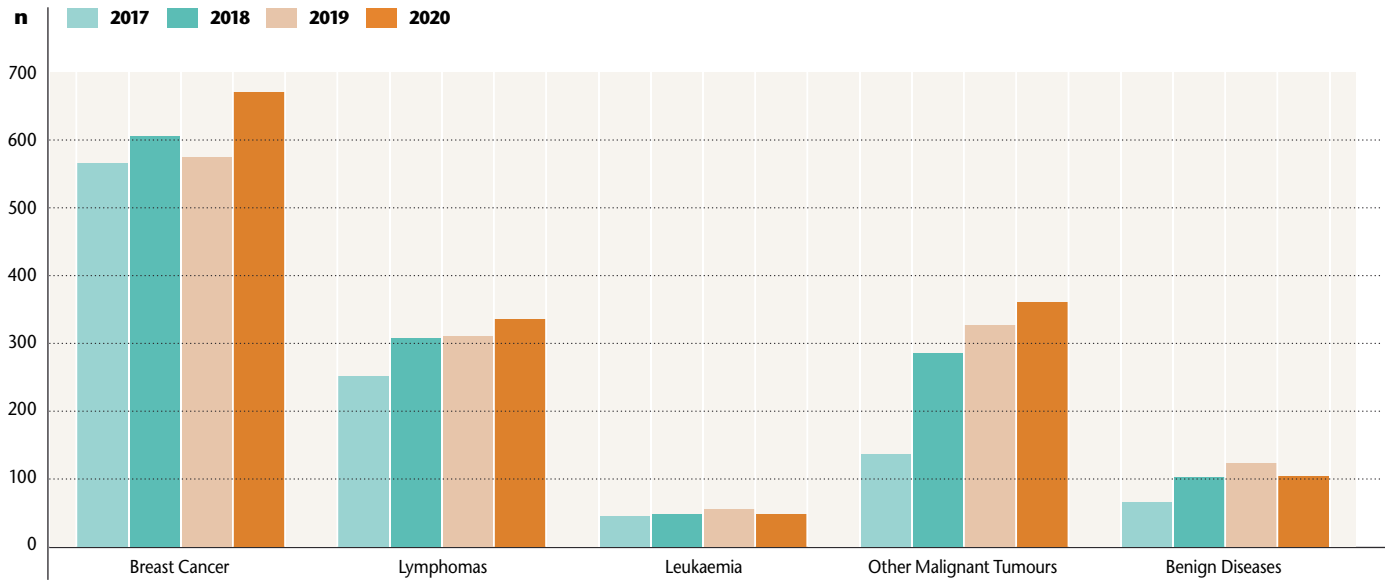
Age-Distribution during Consultation – *FertiPROTEKT* Network e.V.



This year, we see a significant increase of consultations especially in women aged 31-35 years. Presumably, the increasing number of initial consultations of women with breast cancer could explain this observation. The remaining age groups are

roughly at the level of last year. The total number of consultations of "children" under the age of 15 has evened off at around 70.

Diagnosis at Initial Consultation

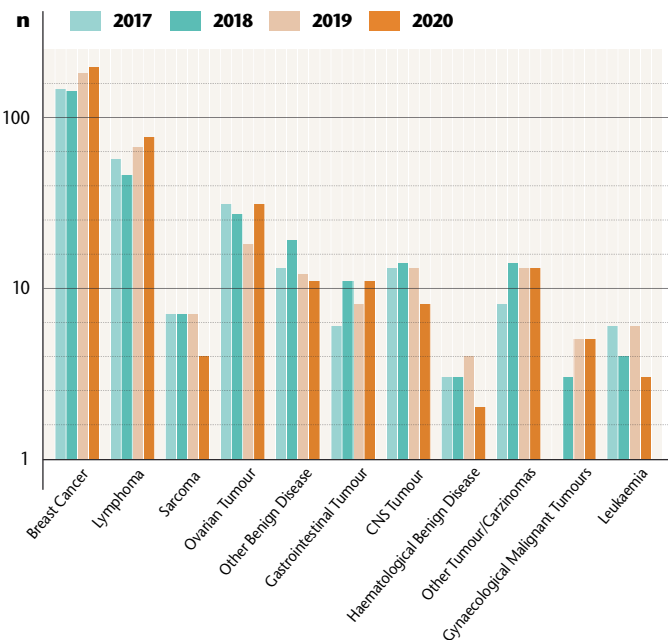


In line with the increased number of initial presentations, all groups of oncological diseases show slightly higher patient numbers in the year 2020. Only the group of breast cancer

patients shows a disproportionate increase compared to the previous years. Patients with benign diseases, on the other hand, were less frequently counselled in that year.

Underlying Disease at Time of Ovarian Stimulation

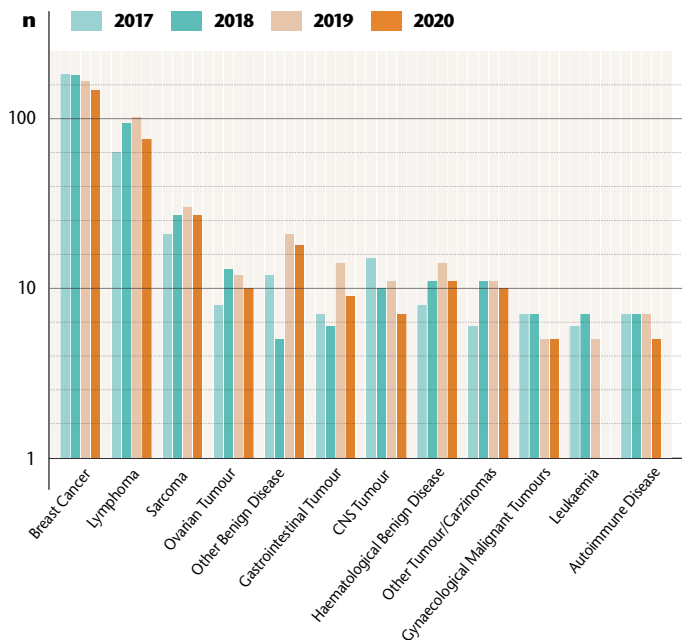
Logarithmic Scale



Considering performed therapies in relation to the underlying disease, the overall rate of ovarian tissue cryopreservation is lower, with only one case of tissue cryopreservation in case of leukaemia. Additionally, fewer ovarian stimulations have been done in

Underlying Disease at Time of Ovarian Tissue Cryopreservation

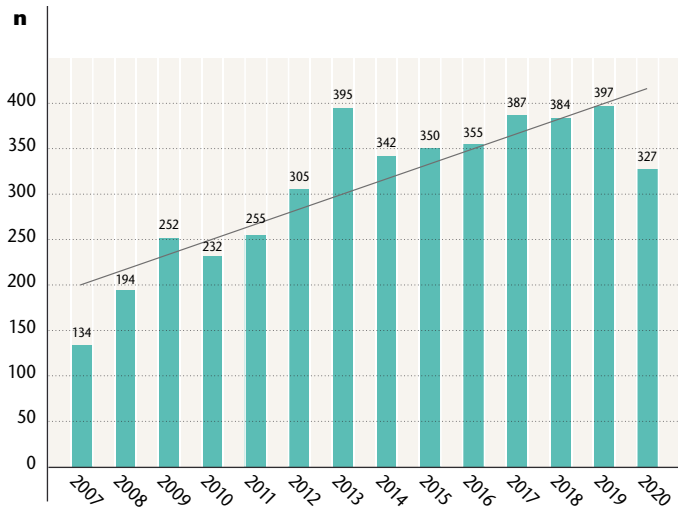
Logarithmic Scale



leukaemia patients while counselling a constant number of these patients; so the total number of interventions in this entity decreased markedly.

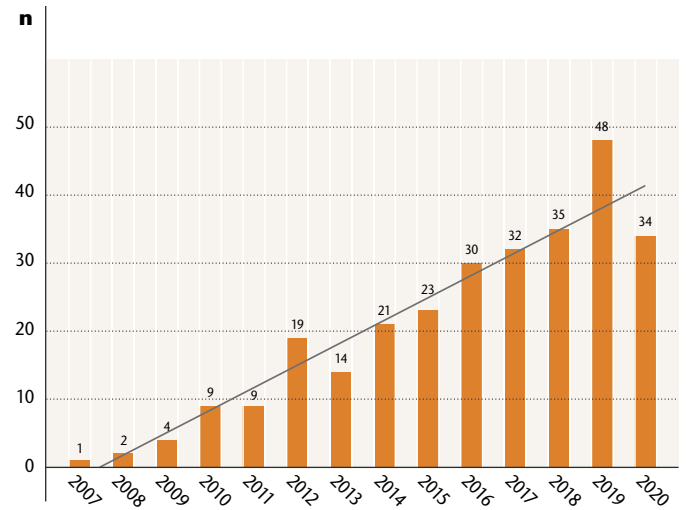
Number of Ovarian Tissue Cryopreservations for Later Transplantation

per Year from 2007–2020, n=4,309



Number of Ovarian Tissue Transplantations

per Year from 2007–2020, n=281

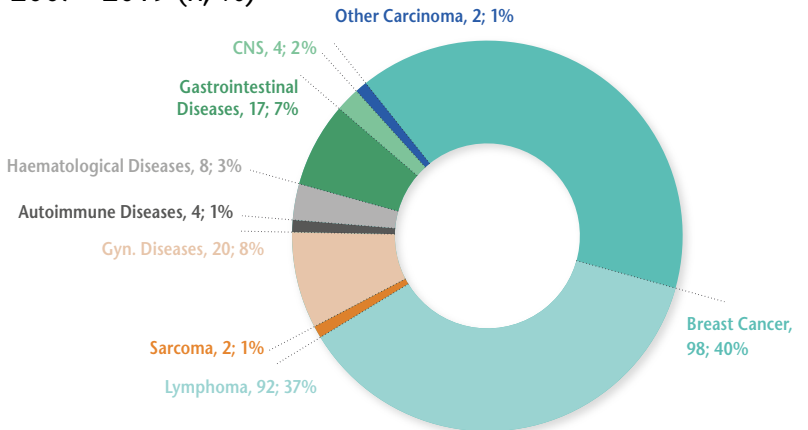


Overall Outcome of Ovarian Tissue Transplantations from 2007–2019

The data represent the results of 27 university and non-university FertiPROTEKT centres (Germany: 20, Switzerland: 4, Austria: 3) since 2007.

In the year 2019, 247 ovarian tissue transplantations have been performed in 200 patients. The current usage rate is 6.5 %.

Underlying Diseases of Patients who have Undergone a Ovarian Tissue Transplantation, 2007–2019 (n; %)



Characteristics of Patients

	Total Number of Transplantations Performed 2007-2019 (n)	Total Number of Patients Undergoing at least 1 Transplantation (n)	Mean Age at Time of Cryopreservation in Years (min-max)	Mean Age at Time of Transplantation in Years (min-max)	Mean Storage Length of Cryopreserved Ovarian Tissue in Years (min-max)
Total Transplantations	247	200	31.1 (17-44)	35.9 (23-47)	4.9 (1-16)
Transplantations with Ovarian Tissue after Overnight Transportation Before Cryopreservation	127	101	31.4 (20-44)	35.8 (25-47)	4.5 (0-11)

Restitution of Endocrine Activity after Transplantation of Cryopreserved Ovarian Tissue

	Endocrine Activity after Transplantation in Relation to Total Number of Transplantations (%)	Endocrine Activity after Transplantation in Relation to Total Number of Patients Transplanted (%)	Patients Without POI* Before Transplantation in Relation to Total Number of Patients Transplanted (%)
Total Transplantations	181/247 (73.3%)	143/200 (71.5%)	26/200 (13.0%)
Transplantations with Ovarian Tissue after Overnight Transportation Before Cryopreservation	107/127 (84.3%)	83/101 (82.2%)	6/101 (5.9%)

*) POI – Premature Ovarian Failure

Pregnancies after Transplantation of Cryopreserved Ovarian Tissue

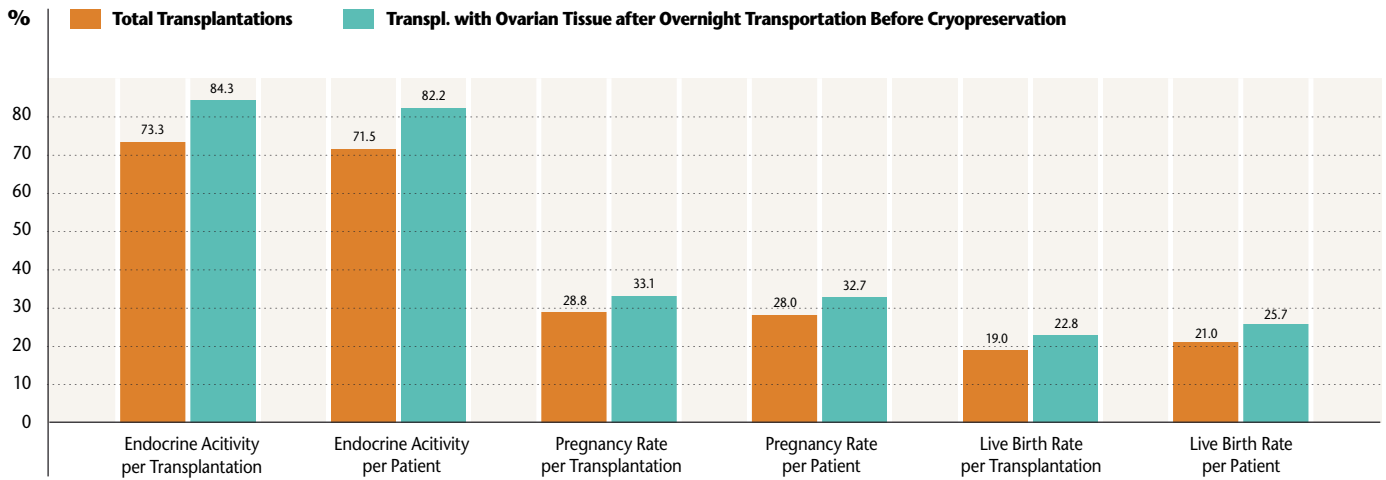
	Pregnancies in Relation to Total Number of Transplantations (%)	Pregnancies in Relation to Total Number of Patients Transplanted (%)	Spontaneous Conceptions in Relation to Total Number of Pregnancies (%)	Conceptions Using ART** in Relation to Total Number of Pregnancies (%)
Total Transplantations	71/247 (28.8%)	56/200 (28.0%)	46/71 (64.8%)	25/71 (35.2%)
Transplantations with Ovarian Tissue after Overnight Transportation Before Cryopreservation	39/127 (33.1%)	33/101 (32.7%)	24/39 (61.5%)	15/39 (38.5%)

** ART – Assisted Reproductive Techniques

Live Births after Transplantation of Cryopreserved Ovarian Tissue

	Live Births in Relation to Total Number of Transplantations (%)	Live Births in Relation to Total Number of Patients Transplanted (%)	Ongoing Pregnancies in Relation to Total Number of Pregnancies (%)
Total Transplantations	47/247 (19.0%)	42/200 (21.0%)	5/56 (8.9%)
Transplantations with Ovarian Tissue after Overnight Transportation Before Cryopreservation	29/127 (22.8%)	26/101 (25.7%)	3/39 (7.7%)

Graphical Presentation of Success Rates ***

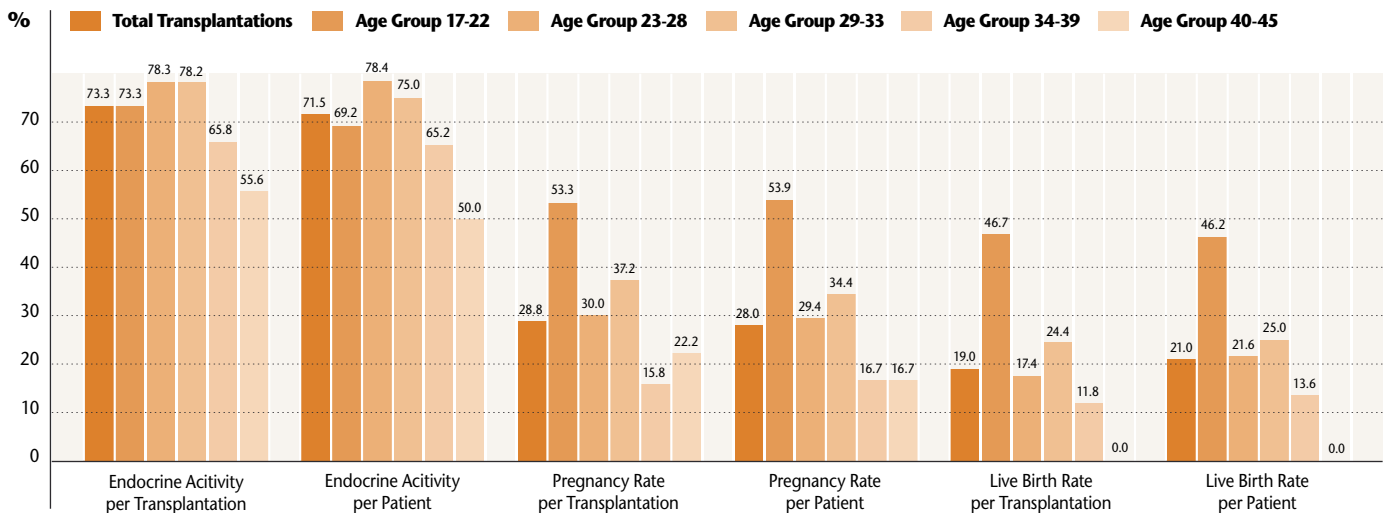


*** The success rates (in %) were calculated without taking into account the age, the site of transplantation, the number of transplants, the underlying disease, the ovarian reserve (the latter determined by serum AMH level before cryopreservation).

Graphical Presentation of Success Rates in Different Age Groups (%)



Age Grouping of Patients was done at the Time of Cryopreservation



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List of D-I-R Members

Sorted by postal code in ascending order – as at August 2021

If calling from outside Germany, dial +49 and omit 0 in the area code.

Kinderwunschzentrum Dresden

Dr. med. univ. Birgit Leuchten

Dr. med. univ. Birgit Leuchten, Dr. med. Hans-Jürgen Held, FÄ Sophia Müller, FÄ Nuria Simó, Dr. med. Stefanie Reißner, M.Sc. Lisa Baldauf, Dr. med. Christina Weiner, Dr. rer. nat. Bernd Junkersdorf
 Prager Straße 8a
 01069 Dresden
 T: 0351 5014000, F: 0351 50140028
 buero@ivf-dresden.de
 www.ivf-dresden.de

Gynäkologische Endokrinologie und Reproduktionsmedizin der Universitätsfrauenklinik Dresden

Universitäres Kinderwunschzentrum

Dr. med. Maren Goeckenjan-Festag, Dr. med. Berit Thieme, Ina Trinkaus
 Fetscherstraße 74
 01307 Dresden
 T: 0351 458 3491, F: 0351 458 5351
 gyn-ivf@uniklinikum-dresden.de
 www.uniklinikumdresden.de/gyn

Kinderwunschzentrum Leipzig-Chemnitz Standort Leipzig

Dipl. med. Jens-Peter Reiher, Dr. med. Petra Jogschies, Dörte Geistert, Laila Shugair, Dr. med. Beate Decker, Dipl.-Biol. Verona Blumenauer
 Goldschmidtstraße 30
 04103 Leipzig
 T: 0341 141200, F: 0341 1412081
 info@ivf-leipzig.de
 www.ivf-leipzig.de

Kinderwunschzentrum Praxisklinik City Leipzig

Standort Leipzig

Dr. med. Astrid Gabert, Dr. med. Katharina Bauer, Dr. med. Isabel Schwandt, FÄ Jana Sonneck, Dr. med. Doreen Marx, Prof. Dr. med. Henry Alexander, Dr. rer. nat. Stefanie Breuer
 Petersstraße 1
 04109 Leipzig
 T: 0341 2158550, F: 0341 21585517
 info@ivf-city-leipzig.de
 www.praxisklinik-city-leipzig.de

Universitätsklinikum Halle (Saale)

Martin-Luther-Universität Halle-Wittenberg, Zentrum für Reproduktionsmedizin und Andrologie

Univ.-Prof. Dr. med. Hermann M. Behre, OA PD Dr. med. Gregor Seliger, Dr. rer. nat. Thomas Greither
 Ernst-Grube-Straße 40
 06120 Halle (Saale)
 T: 0345 557 4782, F: 0345 557 4788
 ZRA@uk-halle.de
 www.kinderwunsch-halle.de

Zentrum für Reproduktionsmedizin Jena & Erfurt

Gemeinschaftspraxis Dres. Fritzsche

PD Dr. med. habil. Heidi Fritzsche, Dr. med. Andreas Fritzsche, Msc.
 Markt 4
 07743 Jena
 T: 03641 474440, F: 03641 4744442
 information@kinderwunsch-thueringen.de
 www.kinderwunsch-thueringen.de

Klinik und Poliklinik für Frauenheilkunde und Fortpflanzungsmedizin, Universitätsklinikum Jena der Friedrich-Schiller-Universität

Uni-Kinderwunsch- und Hormonzentrum

Univ.-Prof. Dr. med. Ingo B. Runnebaum, Dr. rer. nat. Ines Hoppe, Dr. Kristin Nicolaus, Dr. med. Gabriele Pretzsch, Dr. med. Dominik Bräuer, Dr. med. Robert Sczesny, Abdunasser Shtian
 Am Klinikum 1
 07747 Jena
 T: 03641 9329 116, F: 03641 9329 129
 kinderwunsch@med.uni-jena.de
 https://www.uniklinikum-jena.de/frauenheilkunde/KinderwunschHormonzentrum.html

Kinderwunschzentrum Leipzig-Chemnitz Standort Chemnitz

Dipl. med. Jens-Peter Reiher, Dr. med. Petra Jogschies, Dörte Geistert, Laila Shugair, Dr. med. Beate Decker
 Jakobikirchplatz 4
 09111 Chemnitz
 T: 0371 5034980, F: 0371 50349881
 info@ivf-chemnitz.de
 www.ivf-chemnitz.de

Kinderwunschzentrum Praxisklinik City Leipzig

Standort Chemnitz

Dr. med. Astrid Gabert, Dr. med. Katharina Bauer, Dr. med. Isabel Schwandt, FÄ Jana Sonneck, Dr. med. Doreen Marx, Dr. rer. nat. Stefanie Breuer
 Flemmingstraße 2a
 09116 Chemnitz
 T: 0371 4331300, F: 0371 43313017
 info@kinderwunschzentrum-chemnitz.de
 www.kinderwunschzentrum-chemnitz.de

Kinderwunschzentrum am Potsdamer Platz

Dr. med. Hanadi Awwadeh, Dr. med. Anja Mutz
 Leipziger Platz 7
 10117 Berlin
 T: 030 20089500, F: 030 200895099
 info@kinderwunsch-potsdamerplatz.de
 www.kinderwunsch-potsdamerplatz.de

MVZ Kinderwunschteam Berlin GmbH

Dr. med. Gülden Halis, Prof. Dr. med. Jürgen Weiss, Dr. med. Elisabeth Weiss, Dr. sc. hum. Karen Rosenberg, Dr. biol. Jana Krüger
 Friedrichstraße 79
 10117 Berlin
 T: 030 2065805 0, F: 030 2065805 20
 info@kinderwunschteam.berlin
 www.kinderwunschteam.berlin

Praxis für Fertilität

Gynäkologische Endokrinologie und Reproduktionsmedizin

Dr. med. David J. Peet, Dr. med. Wibke Wilkening, Constanze Glaser, Dr. med. Annette Nickel, Dr. med. Rolf Metzger, Dr. med. Susann Kreuz
 Friedrichstraße 150
 10117 Berlin
 T: 030 2630231 0, F: 030 2630231 19
 info@fertilitaet.de
 www.fertilitaet.de

MVZ TFP Berlin GmbH

Dr. med. Christine F. Nöldechen, Dr. med. Dagmar V. Geiß, Dr. med. Ulrike Bergmann-Hensel, Dr. med. Birgit Bestvater, Astrid Kim, Dr. med. Maurus J. Asen
 Kronenstraße 55-58
 10117 Berlin
 T: 030 2062672 0, F: 030 2062672 18
 berlin@tfp-fertility.com
 www.tfp-fertility.com/de-de/tfp-kinderwunsch-berlin

Wunschkind Berlin

Dr. med. Björn Horstkamp, PD Dr. med. Bernd Krause, Dr. med. Petra Rudolph, Dr. med. Katja Schwenn, Dr. med. vet. Sophia Herzfeld, Dipl.-Biol. Werner Hoppenstedt
 Uhlandstraße 20-25
 10623 Berlin
 T: 030 880349056, F: 030 880349035
 info@wunschkind-berlin.de
 www.wunschkind-berlin.de

Ceres – Kinderwunschzentrum Dr. Hannen und Dr. Stoll

Dr. med. Reinhard Hannen, Dr. med. Christian Friedrich Stoll
 Landgrafenstraße 14
 10787 Berlin
 T: 030 2639830, F: 030 26398399
 info@kinderwunschzentrum.de
 www.kinderwunschzentrum.de

Kinderwunschzentrum an der Gedächtniskirche

Dr. med. Matthias Bloechle, Dr. med. Silke Marr, Astrid Kim
 Rankestraße 34
 10789 Berlin
 T: 030 2190920, F: 030 21909299
 info@kinderwunsch-berlin.de
 www.kinderwunsch-berlin.de

Kinderwunschzentrum am Innsbrucker Platz Berlin

Babette Remberg, Dr. med. Susanne Tewordt-
Thyselius, Dr. med. Jutta Sidor, Dr. rer. nat.
Thomas Jeziorowski
Hauptstraße 65
12159 Berlin
T: 030 85757930, F: 030 85757935
praxis@kinderwunschpraxis-berlin.de
www.kinderwunschpraxis-berlin.de

**Kinderwunschzentrum Dres. Hoffmann
Praxis-Klinik Dres. Hoffmann**

Dr. med. Svetlana Hoffmann, Dr. med. Ivan
Hoffmann, Dr. Vladimir Fait, M. Sc.
Bölschestraße 62
12587 Berlin
T: 030 84599450
info@praxisklinik-dres-hoffmann.de
www.praxisklinik-dres-hoffmann.de

MVZ für Gynäkologie Helle-Mitte

Dr. med. Muna Zaghoul-Abu Dakah, Dr. med.
Gothild Matheus
Stendaler Straße 24
12627 Berlin
T: 030 9927790, F: 030 99277922
info@berliner-kinderwunsch.de
www.berliner-kinderwunsch.de

MVZ Fertility Center Berlin

Auf dem Gelände der DRK Kliniken Westend

Dr. med. Andreas Tandler-Schneider, Dr. med.
Gabriele Stief, Dr. med. Anette Siemann, Prof.
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Dr. rer. nat. Ulrike Montag, Dr. rer. nat. Lorena
Fernández González, Caroline Hirschfeld-Ihlow
Spandauer Damm 130
14050 Berlin
T: 030 233208110, F: 030 233208119
info@fertilitycenterberlin.de
www.fertilitycenterberlin.de

Kinderwunschartze Berlin GbR

**Zentrum für Kinderwunschbehandlung und
Fertilitätsprotektion**

Dr. med. Andreas Jantke, Dr. med. Anna Stegel-
mann, Frau Bettina Jantke, Dr. rer. nat. Yves
Charron, Dipl.-Biol. Nicole Klauke
Clayallee 225a
14195 Berlin
T: 030 814576565, F: 030 814576566
info@kinderwunschaerzte-berlin.de
www.kinderwunschaerzte-berlin.de

Kinderwunschzentrum Potsdam MVZ GmbH

Dr. med. Kay-Thomas Moeller, Dr. med.
Kathleen Linca, Dr. med. Anja Bretschnei-
der-Schwarz, Dr. med. Hendrikje Lukoschus,
Elaine Hempel, PD Dr. Dr. Heide Reil
Babelsberger Straße 8
14473 Potsdam
T: 0331 23189292, F: 0331 23189293
info@kinderwunsch-potsdam.de
www.kinderwunschzentrum-potsdam.de

**Praxis für Fertilität
MVZ GmbH**

PD Dr. med. Heiner Müller, Annette Busecke,
Anja Bossow, Dr. med. Anne Koenen
Südring 81
18059 Rostock
T: 0381 44012030, F: 0381 44012031
info@ivf-rostock.de
www.ivf-rostock.de

**Medizinisches Versorgungszentrum
Fertility Center Hamburg GmbH**

Dr. med. (IL) Robert Fischer, Dr. med. Kay
Christian Löbbecke, Dr. med. Heike Boppert,
Thomas Meyer, Prof. Dr. med. Wolfgang
Schulze
Speersort 4
20095 Hamburg
T: 040 30804400, F: 040 30804900
fch-service@amedes-group.com
www.fertility-center-hh.de

amedes experts

**Facharzt-Zentrum für Kinderwunsch,
Pränatale Medizin, Endokrinologie und
Osteologie Hamburg**

Prof. Dr. med. Frank Nawroth, Prof. Dr. med.
Barbara Sonntag, Dr. med. Astrid Dangel, Prof.
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Meibes, Dr. med. Raquel Pozo-Ugarte, Prof.
Dr. med. Sabine Segerer, Dr. rer. nat. Beatrice
Maxrath
Mönckebergstraße 10 (Barkhoffpassage)
20095 Hamburg
T: 0800 5891688, F: 040 380708310
kinderwunsch-hamburg@amedes-group.com
www.amedes-experts-hamburg.de

Kinderwunsch Valentinshof

Dr. med. Anja Dawson, Dr. med. Nuray Aytakin,
PD Dr. med. Ulrich A. Knuth
Caffamacherreihe 8
20355 Hamburg
T: 040 709751 10, F: 040 709751 39
empfang@kinderwunsch-valentinshof.de
www.kinderwunsch-valentinshof.de

Kinderwunsch Hamburg Mitte

Dr. med. Anja Dawson, Prof. Dr. (Univ. Bs. As.)
Miguel Hinrichsen, Dr. med. Nuray Aytakin
Caffamacherreihe 8
20355 Hamburg
T: 040 6963244 60, F: 040 6963244 79
empfang@kinderwunsch-hh-mitte.de
www.kinderwunsch-hh-mitte.de

**Kinderwunschzentrum Altonaer Straße
(MVZ) im Gynaecologikum Hamburg (GbR)**

Dr. med. Annick Horn, Dr. med. Tim Cordes,
Prof. Dr. med. Markus S. Kupka
Altonaer Straße 59
20357 Hamburg
T: 040 306836 0, F: 040 306836 69
info@ivf-hamburg.de
www.ivf-hamburg.de

KinderwunschZentrum HAFENCITY Hamburg

Dr. med. Ekbert Göhmann, Dr. med. Thomas
Krämer
Sumatrankontor, Überseeallee 1
20457 Hamburg
T: 040 30088 100, F: 040 30088 1010
kontakt@kinderwunsch-hafencity.de
www.kinderwunsch-hafencity.de

Kinderwunsch Praxisklinik Fleetinsel Hamburg

Dr. univ. Ist. Semsettin E. Koçak, Dr. med. Peter
List, PD Dr. med. Kay Neumann, Dr. med.
Sonja Scheuß, Dr. rer. nat. Uwe Weidner
Admiralitätstraße 4
20459 Hamburg
T: 040 38605550, F: 040 38605551
info@kinderwunschfleetinsel.de
www.kinderwunschfleetinsel.de

**Praxis für Kinderwunsch & Hormone –
Hamburger Straße**

Dr. med. Urte Reinhardt, Dr. med. Tina Oster-
holz-Zaleski, Jennifer Hajek MSC.
Klinikweg 23
22081 Hamburg
T: 040 6000 379 0, F: 040 6000 379 29
welcome@ivf-hh.de
www.ivf-hh.de

**Universitäres Kinderwunschzentrum Lübeck
und Manhagen**

Zentrum für Gynäkologische Endokrinologie
und Reproduktionsmedizin am Universitäts-
klinikum Schleswig-Holstein, Universitäre
Kinderwunschzentren GmbH
Prof. Dr. med. Georg Griesinger, M. Sc., PD Dr.
med. Askan Schultze-Mosgau, Dr. med. Marion
Depenbusch
Ratzeburger Allee 111-125
23562 Lübeck
T: 0451 505778 10, F: 0451 505778 299
ParkKlinik Manhagen, Zufahrt: Hansdorfer
Straße 9
22927 Großhansdorf
T: 04102 777 686 0, F: 04102 777 686 309
kinderwunsch@uksh.de
www.uksh.de/Kinderwunsch_Luebeck/

Kinderwunsch Holstein

Dr. med. Peter Kunstmann, Dr. biol. hom. Claas
Mehnert
Lübecker Straße 68
23611 Bad Schwartau
T: 0451 498955 22, F: 0451 498955 25
info@ivf-badschwartau.de
www.ivf-badschwartau.de

fertilitycenter Schleswig-Holstein

fertilitycenterkiel / fertilitycenterflensburg

Dr. med. Antonia Wenners, Dr. med. Martin
Völckers, Dr. med. Nevin Inan
Prüner Gang 15
24103 Kiel
T: 0431 97413 33, F: 0431 97413 89
Bahnstraße 23c
24937 Flensburg
T: 0461 50506 20, F: 0461 50506 22
info@fertilitycenter.de
www.fertilitycenter.de

Universitäres Kinderwunschzentrum Kiel
 PD Dr. med. Sören von Otte, Prof. Dr. med. Liselotte Mettler, Dr. med. Wiebke Junkers, Dr. med. Veronika Günther
 Arnold-Heller-Straße 3, Haus C
 24105 Kiel
 T: 0431 500 922 20, F: 0431 500 922 24
 kinderwunsch-kiel@uksh.de
 www.uksh.de/Kinderwunsch_Kiel/

KinderwunschKiel
 Dr. med. Angela Carstensen, Dr. med. Kirsten Schem, Dr. sc. agr. Raphael Schütt
 Im Brauereiviertel 5
 24118 Kiel
 T: 0431 553433, F: 0431 5192745
 info@kinderwunschkiel.de
 www.kinderwunschkiel.de

Team Kinderwunsch Oldenburg GbR MVZ
 Dr. med. Saif Jibril, Nina Wezel, Anja Scheffer
 Leo-Trepp-Straße 5
 26121 Oldenburg
 T: 0441 2171570, F: 0441 21715798
 info@teamkinderwunsch.de
 www.teamkinderwunsch.de

Tagesklinik Oldenburg
 Dr. med. Jörg Hennefründ, Dr. Firas Alhalabieh
 Achternstraße 21
 26122 Oldenburg
 T: 0441 922700, F: 0441 9227028
 info@tagesklinik-oldenburg.de
 www.tagesklinik-oldenburg.de

Kinderwunschzentrum Ostfriesland
 Dr. med. Grita Hasselbach, Dr. med. Alice Rachidi, Dr. rer. nat. Isabell Motsch
 Hafestraße 6d
 26789 Leer (Ostfriesland)
 T: 0491 454250, F: 0491 4542510
 info@kinderwunschleer.de
 www.kinderwunschostfriesland.de

Kinderwunsch Bremen
 Dr. med. Christoph Grewe, Tanja Finger, Sennur Erikli-Koc
 Emmastraße 220
 28213 Bremen
 T: 0421 224910, F: 0421 2249122
 info@kinderwunschbremen.de
 www.kinderwunschbremen.de

Bremer Zentrum für Fortpflanzungsmedizin (BZF)
 Dr. med. Sebastian Grewe, Dr. med. Olaf Drost
 Gröpelinger Heerstraße 406-408
 28239 Bremen
 T: 0421 61021212, F: 0421 61021213
 kontakt@icsi.de
 www.icsi.de

Team Kinderwunsch Hannover
 Dr. med. Nabil Saymé, Dipl.-Biol. T. Krebs
 Aegidientorplatz 2b
 30159 Hannover
 T: 0511 450 34410, F: 0511 450 34419
 info@team-kinderwunschhannover.de
 www.team-kinderwunsch-hannover.de

Medizinische Hochschule Hannover (MHH) Klinik für Frauenheilkunde und Geburtshilfe, Abteilung Reproduktionsmedizin und gynäkologische Endokrinologie
 Prof. Dr. med. Cordula Schippert, Prof. Dr. med. Frauke von Versen-Höyneck, Dr. med. Guillermo-José Garcia-Rocha, Dr. med. Jill-Caren Philippeit, Dr. rer. nat. Dagmar Töpfer
 Carl-Neuberg-Straße 1, Gebäude K11, Ebene SO
 30625 Hannover
 T: 0511 5326099, F: 0511 5326088
 Frauenklinik-Kinderwunsch@mh-hannover.de, Schippert.Cordula@mh-hannover.de
 www.mh-hannover.de/kinderwunsch.html

Kinderwunschzentrum Langenhagen & Wolfsburg MVZ
 Dr. med. Thilo Schill, Dr. med. Martina Mueseler-Albers, Dr. med. Natascha Peper, Dr. med. Christina Nardmann, Herr Christian Bell, Dr. med. Kristin Remke, Dr. med. Mareike Albers
 Ostpassage 9
 30853 Langenhagen
 T: 0511 972300, F: 0511 9723018
 praxis@kinderwunsch-langenhagen.de
 www.kinderwunsch-langenhagen.de

Zentrum für Reproduktionsmedizin & Humangenetik MVZ
 Dr. med. Georg Wilke, Dr. med. Jan-Simon Lanowski, Dr. med. Notker Graf, Anja Scheffer, Natalia Kellermann, Dr. med. Gabriele Lanowski, Dr. med. Janina Bartels, Dr. rer. hum. biol. Susanne Gärtner-Hübsch, Tom Seeling M. Sc., Ksenia Evseeva B. Sc.
 Gartenstraße 18-20
 31141 Hildesheim
 T: 05121 206790, F: 05121 2067911
 praxis@kinderwunsch-hildesheim.de
 www.kinderwunsch-hildesheim.de

Deutsche Klinik Bad Münder – Hannover Zentrum für IVF und Reproduktionsmedizin MVZ wagnerstibbe für Gynäkologie, Reproduktionsmedizin, Zytologie, Pathologie und Innere Medizin
 Dr. med. Christina Baßler, Dr. med. Elmar Breitbach, Dr. med. Arvind Chandra, Dr. med. Frauke Kramer, Dr. med. Nadine Kundu, Dr. med. Sabine Leßmann, Dr. med. Franziska Wegener, Iris Krause, Ulrike Hasenjäger
 Hannoversche Straße 24
 31848 Bad Münder
 T: 05042 940 360, F: 05042 940 308
 info@kinderwunsch.com
 www.kinderwunsch.com

Zentrum für Kinderwunschbehandlung und pränatale Medizin GMP
 Dr. med. Michael Dumschat, Dr. med. Ralf Menkhous, Dr. med. Stefanie Strunk
 Simeonsplatz 17
 32423 Minden
 T: 0571 972600, F: 0571 9726099
 info@kinderwunsch-minden.net
 www.kinderwunsch.net

Praxisklinik Prof. Volz – FROG
 Prof. Dr. med. Joachim Volz, PD Dr. med. Stefanie Volz-Köster
 Adenauerplatz 7
 33602 Bielefeld
 T: 0521 9883060, F: 0521 98830622
 kinderwunsch@frog.de.com
 www.frog.de.com

Bielefeld Fertility-Center Zentrum für Reproduktionsmedizin und Gynäkologische Endokrinologie
 Dr. med. Karl Völklein, Beata Szypajlo, Dr. med. Wiebke Rübberdt, Paul A. Ebert, Dr. med. Kathrin Brandhorst, Christopher Eber
 Wertherstraße 266-268
 33619 Bielefeld
 T: 0521 101005, F: 0521 101079
 praxis@kinderwunsch-bielefeld.de
 www.kinderwunsch-bielefeld.de

MVZ für Reproduktionsmedizin am Klinikum Kassel
 Dr. med. Marc Janos Willi, Dr. med. Oswald Schmidt, Prof. Dr. (UBsAs) Miguel J. Hinrichsen
 Haus F, Mönchebergstraße 41-43
 34125 Kassel
 T: 0561 9802980, F: 0561 9802981
 info@ivf-kassel.de
 www.kinderwunsch-kassel.de

Reproduktionsmedizinisches Kompetenzzentrum Marburg und Gießen
 PD Dr. med. Volker Ziller, Prof. Dr. med. Uwe Wagner, FA Marcel Schuett
 Baldingerstraße
 35043 Marburg
 T: 06421 5861330, F: 06421 5867070
 Klinikstraße 33
 35392 Gießen
 T: 0641 98545207, F: 0641 98557099
 kinderwunsch@med.uni-marburg.de
 www.repko-ukgm.de

Kinderwunschzentrum Mittelhessen
 Dr. med. Amir Hajimohammad, Berthold Oels
 Sportparkstraße 9
 35578 Wetzlar
 T: 06441 2002020, F: 06441 20020299
 info@ivf-mh.de
 www.ivf-mh.de

Zentrum für Kinderwunsch und Reproduktionsmedizin gyn-medicum Göttingen
 Dr. med. Monica Tobler, PD Dr. med. Andreas Schmutzler, Dr. sc. agr. Manuela Ropeter-Scharfenstein und KollegInnen
 Waldweg 5
 37073 Göttingen
 T: 0551 41337, F: 0551 41722
 info@kinderwunsch-praxis-goettingen.de
 www.gyn-medicum.de

MVZ Kinderwunschzentrum Göttingen
 Dr. med. Rüdiger Moltrecht, Dr. med. Thomas Welcker, Dr. med. Filiz Sakin-Kaindl, Dr. rer. nat. Heike Eckel
 Kasseler Landstraße 25a
 37081 Göttingen
 T: 0551 998880, F: 0551 9988899
 info@kinderwunsch-zentrum-goettingen.de
 www.kinderwunsch-zentrum-goettingen.de

Kinderwunschzentrum Magdeburg

Dr. med. Evelyn Richter, Dr. med. Mohamed Gamal Ibrahim
Michael-Lotter-Straße 7
39108 Magdeburg
T: 0391 6624890, F: 0391 66248929
info@kiwumed.de
www.kinderwunschzentrum-magdeburg.de

**Otto-von-Guericke-Universität Magdeburg
Universitätsfrauenklinik, Bereich Reproduktionsmedizin und Gynäkologische Endokrinologie**

Dr. med. Julia Bartley, FÄ Dr. med. Claudia Matthes, Dr. rer. nat. Katja Seidel
Gerhart-Hauptmann-Straße 35
39108 Magdeburg
T: 0391 6717390, F: 0391 6717389
susan.eggert@med.ovgu.de
www.krep.ovgu.de

Kinderwunsch-Kö

Dr. med. Martina Behler, Tanja Emde, Dr. rer. nat. Suna Cukurcam
Königsallee 63-65
40215 Düsseldorf
T: 0211 3113550, F: 0211 31135522
info@kinderwunsch-koe.de
www.kinderwunsch-koe.de

MVZ TFP Düsseldorf GmbH

Dr. med. Kathrin Fleischer, Dipl. med. Kersten Marx, Katja Neldner, PD Dr. med. Andrea Schanz, Dr. med. Nilofar Roshandel, Drs. H.V.H. Mous, P.R. Beizermann, Dr. med. Peter Schröder, Dr. med. W. Fabry, Dr. med. Zübeyda Akya-zi-Oberhoffer, Dr. med. Jochen Tigges, Dr. med. Maria Quasdorf, Dr. rer. nat. Dipl.-Biol. David Jussen; Dr. rer. nat. Dipl.-Biol. Sarah Funke
Völklinger Straße 4
40219 Düsseldorf
T: 0211 901970, F: 0211 9019750
duesseldorf@tfp-fertility.com
www.tfp-fertility.com/de-de/TFP-kinderwunsch-duesseldorf

UniKiD

Universitäres interdisziplinäres Kinderwunschzentrum Düsseldorf

Prof. Dr. med. Jan-Steffen Krüssel, Prof. Dr. med. Alexandra Bielfeld, Dr. med. Sylvia Bartnitzky, Dr. med. Inke Bruns, Dr. med. Julia Bugler, Dr. med. Cornelius Döhmen, Dr. med. Philipp Edimiris, Dr. med. Tanja Freund-Schütt, Dr. med. Eva Klemm, Dr. med. Barbara Mikat-Drozdzynski, Chrysostomos Papadopoulos, Dr. rer. nat. Jens Hirchenhain, Dr. rer. nat. Dunja M. Baston-Büst, Dr. rer. nat. Sebastian Büst, Dr. rer. nat. Nele Weber, Dr. rer. nat. Jana Liebenthron
Moorenstraße 5
40225 Düsseldorf
T: 0211 8104060, F: 0211 8116787
info@unikid.de
www.unikid.de

Ki.Nd

Kinderwunschzentrum Niederrhein

Dr. med. Georg M. Döhmen, Dr. med. Thomas Schalk, Dr. med. Nina Röner, Dr. rer. nat. Ezzaldin Alazzeh, Melanie Süßmilch, Wiebke Grotherath, Claudia Kalus
Madriker Str. 6
41069 Mönchengladbach (Nordpark)
T: 02161 496860, F: 02161 4968619
47805 Krefeld
Zweigstelle: Melanchthonstraße 36
T: 02151 150 2310, F: 02151 150 2311
info@ki-nd.de
www.ki-nd.de

green-ivf

Grevenbroicher Endokrinologie- und IVF-Zentrum

Dr. med. Daniel Fehr, M.Sc., Prof. Dr. med. Christian Gnoth, Dr. med. Nina Bachmann, Dr. med. Mareen ten Busch, Dr. med. Richard Leblanc, Dr. med. Sandra Schott, Dr. med. Elina Schwake, Dr. med. Luisa Steiff, Dr. med. Nadine Sutter, Dr. med. Uta von Westernhagen, Dr. med. Carola Ziegler, Dr. rer. nat. Caroline Merino León, Ilka Haase, M.Sc., Viktoria Zent, M.Sc.
Rheydter Straße 143
41515 Grevenbroich
T: 02181 491513, F: 02181 491534
info@green-ivf.de
www.green-ivf.de

Bergisches Kinderwunschzentrum Remscheid

Dr. med. Johannes Luckhaus, Dr. med. Anke Beerkotte
Elberfelder Straße 49
42853 Remscheid
T: 02191 791920, F: 02191 7919239
mail@kinderwunsch-remscheid.de
www.kinderwunsch-remscheid.de

Überörtliche Berufsausübungsgemeinschaft Kinderwunsch Dortmund, Siegen, Dorsten, Wuppertal GbR

Prof. Dr. med. Stefan Dieterle, Dr. med. Andreas Neuer, Prof. Dr. med. Robert Greb, Dr. med. Katharina Möller-Morlang, Dr. med. Thomas von Ostrowski, Dr. med. Saskia Möckel, Petra Wilbrink, Dr. med. Karoline Hohenstein, Dr. med. Bernhard Mohr
Olpe 19
44135 Dortmund
T: 0231 5575450, F: 0231 55754599
Hermelsbacher Weg 41
57072 Siegen
T: 0271 7701810, F: 0271 77018129
Südwall 15
46282 Dorsten
T: 02362 27001, F: 02362 27002
Hofaue 93
42103 Wuppertal
T: 0202 4789930, F: 0202 47899329
info@kinderwunschzentrum.org
www.kinderwunschzentrum.org

novum - Zentrum für Reproduktionsmedizin Überörtliche Gemeinschaftspraxis

Prof. Dr. med. Thomas Katzorke, Dr. med. Susanne Wohlers, Najib N. R. Nassar, Prof. Dr. med. Peter Bielfeld, Dr. med. Nora Holtmann, Dr. med. Ruth Pankoke
Hauptbetriebsstätte: Akazienallee 8-12
45127 Essen
T: 0201 294290, F: 0201 2942914
Nebenbetriebsstätte: Friedrich-Wilhelm-Straße 71
47051 Duisburg
T: 0203 7139580, F: 0203 71395815
info@ivfzentrum.de
www.ivfzentrum.de

REProVita

Kinderwunschzentrum Recklinghausen

Dr. med. Cordula Pitone, Dr. med. Agnieszka Wendt
Hertener Straße 29
45657 Recklinghausen
T: 02361 904188 0, F: 02361 904188 41
info@reprovita.de
www.reprovita.de

Kinderwunschpraxis Gelsenkirchen

Wissenschaftspark Pav. 8, 1.OG
Dr. med. Ute Czeromin, Dr. med. Ina Walter-Göbel
Munscheidstraße 14
45886 Gelsenkirchen
T: 0209 167 1470, F: 0209 167 1471
info@kinderwunsch-gelsenkirchen.de
www.kinderwunsch-gelsenkirchen.de

Kinderwunschpraxis an der Promenade GMP Mempel & Stratmann

Dr. med. Andrea Mempel, Susanne Stratmann, Dr. rer. nat. Melanie Rickert-Föhring
Von-Vincke-Straße 14
48143 Münster
T: 0251 414312 0, F: 0251 414312 20
willkommen@kinderwunsch-promenade.de
www.kinderwunsch-promenade.de

MVZ Kinderwunsch- und Hormonzentrum Münster GmbH

Kinderwunschzentrum Münster

Dr. med. Caroline Niehoff, Prof. Dr. med. Axel Kamischke
Hötteweg 5-7
48143 Münster
T: 0251 482670, F: 0251 4826777
info@ivf-muenster.de
www.ivf-muenster.de

UKM Kinderwunschzentrum Universitätsklinikum Münster

Prof. Dr. med. Hermann M. Behre, Prof. Dr. med. Sabine Kliesch, Tanja Sperlbaum, PD Dr. rer. nat. Verena Nordhoff
Albert-Schweitzer-Campus 1, Gebäude D 11
48149 Münster
T: 0251 8358280, F: 0251 8348267
info-kinderwunsch@ukmuenster.de
www.ukm-kinderwunschzentrum.de

**Zentrum für Kinderwunschbehandlung
Osnabrück / Nordhorn**

Irene Coordes, Dr. med. Manfred Schneider
Friedrich-Janssen-Straße 1
49076 Osnabrück
T: 0541 404500, F: 0541 4045040
Osnabrücker Straße 1
48529 Nordhorn
info@kinderwunsch123.de
www.kinderwunsch123.de

Kinderwunschzentrum Königsdorf

Dr. med. Dieter Struller, Dr. med. Christof Etien
Aachener Straße 545
50226 Frechen-Königsdorf
T: 02234 6060600
info@kinderwunsch-erft.de
www.kinderwunsch-erft.de

MVZ PAN Institut GmbH**Interdisziplinäres Kinderwunschzentrum**

Dr. med. Stefan Palm, Dr. med. Mirko Dannhof,
Dr. med. Irene Pütz, PD Dr. med. Dolores Foth,
Dr. med. Julia Holtschmidt, Dr. med. Bartlomiej
Berger, Dr. rer. nat. Bastian Schäferhoff, Dipl.
biol. Manuel Kernbach, Dr. med. Dipl. bio.
Martina Kreiß, Dr. med. Susanne Stemmler,
Dr. med. Norbert Schöngen, Dr. med. Georg
Mansmann, PD Dr. med. Torsten Schmidt, Dr.
med. Diana Delle (MIC II), Dipl.-Psych. Andrea
Langness, Dipl.-Psych. (NL) Johanna Morris,
Prof. Dr. med. Martina Breidenbach
Zeppelinstraße 1 (Neumarkt Galerie)
50667 Köln
T: 0221 2776200, F: 0221 2776201
repro@pan-klinik.de
www.mvz-pan-institut.de

**Klinik und Poliklinik für Frauenheilkunde
und Geburtshilfe der Universität zu Köln
Gynäkologische Endokrinologie und Reproduktionsmedizin**

Prof. Dr. med. Gohar Rahimi, Dr. med. Ramona
Das, Dr. rer. nat. Evgenia Isachenko, Dr. rer.
nat. Vladimir Isachenko
Kerpener Straße 34
50931 Köln
T: 0221 478 87550, F: 0221 478 86201
kinderwunsch-info@uk-koeln.de
frauenklinik.uk-koeln.de/schwerpunkte/
kinderwunsch/

**MVZ amedes für IVF- und Pränatalmedizin
in Köln GmbH**

Dr. med. Markus Merzenich, Dr. Birgit Kruse-
notto, Eva Schwahn
Schönhauser Straße 3
50968 Köln
T: 0221 3403070, F: 0221 34030777
info@kinderwunschzentrum-koeln.de
www.kinderwunschzentrum-koeln.de

Kinderwunschzentrum Aachen

Dr. med. Bilge Kwiatkowski, Dr. med. Klaus
Grunwald, Verena Schroeder, Frau Verena Alt
Kasernenstraße 25
52064 Aachen
T: 0241 99774140, F: 0241 99774144
kinderwunschzentrum@t-online.de
www.kinderwunsch-aachen.de

**Klinik für Gynäkologische Endokrinologie und
Reproduktionsmedizin**

Uniklinik - RWTH Aachen
Prof. Dr. med. Joseph Neulen, Dr. rer. nat. Ute
Weißenborn
Pauwelsstraße 30
52074 Aachen
T: 0241 8088953, F: 0241 8082511
uweissenborn@ukaachen.de
www.ukaachen.de

**Kinderwunschzentrum Heinsberger Höfe GbR
Medizinische Kooperationsgemeinschaft**

Drs. Willem-Jan S. S. Cuypers, Drs. Nerissa B. E.
Cuypers, Dr. rer. nat. Ralf Böhm
Hochstraße 154
52525 Heinsberg
T: 02452 9966 900, F: 02452 9966 910
info@cuypers-cuypers.com
www.cuypers-cuypers.com

Universitätsklinikum Bonn**Gynäkologische Endokrinologie und Reproduktionsmedizin – VenusKIND am UKB**

Prof. Dr. med. Nicole Sängler, Dr. (univ Aleppo)
Fadi Bakjaji, Dr. med. Angela Meiser, Dr. med.
Eva Vokuhl, Dr. med. Julia John, Dr. Lena
Reichenbach, Dr. med. Nora Emrich, Dr. med.
Elena Leinweber, Dr. rer. nat. Andreas Schall-
moser, Cara Färber, Dr. Vanessa Hüren
Venusberg Campus 1, Gebäude 35
53127 Bonn
T: 0228 287 15779, F: 0228 28715795
officerepro@ukb.uni-bonn.de
www.kinderwunsch-uni-bonn.de

**MVZ für Frauenheilkunde und IvF-Medizin
Bonn GbR**

Prof. Dr. med. Katrin van der Ven, Dr. med.
Marietta Kühr, Dr. med. Martina Gördes,
Dr. med. Simone Sauter, Dipl.-Biol. Claudia
Grewenig
Godesberger Allee 64
53175 Bonn
T: 0228 9090440, F: 0228 90904411
info@praxis-godesberger-allee.de
www.praxis-godesberger-allee.de

kiwup®**Kinderwunschpraxis in Bonn**

Dr. med. Julia Ittstein, PD Dr. Dr. med. Gernot
Paul Prietl, Dr. med. Monika Prietl, Dr. med.
Peter Heuschen
Theaterplatz 18
53177 Bonn
T: 0228 3503910, F: 0228 364892
info@kiwup.de
www.kiwup.de

Kinderwunschzentrum Bonner Bogen

Dr. med. Ulrike Bohlen, Dr. med. Eva-Maria
Boogen, Dr. med. vet. Maria Köster
Joseph-Schumpeter-Allee 1
53227 Bonn
T: 0228 3388200, F: 0228 33882099
info@kinderwunschzentrum-bonnerbogen.de
www.kinderwunschzentrum-bonnerbogen.de

**Kinderwunsch Praxisklinik Trier
Wissenschaftspark (WIP)**

Dr. med. Mohsen Satari, Prof. Dr. med. Jürgen
P. Hanker
Max-Planck-Straße 15
54296 Trier
T: 0651 979060, F: 0651 9790620
info@kinderwunsch-trier.de
www.kinderwunsch-trier.de

Kinderwunsch Zentrum Mainz

Dr. med. Robert Emig, Dr. med. Christine
Molitor, Prof. Dr. med. Thomas Steck, Dr. med.
Marie-Theres Swayze
Rheinstraße 4
55116 Mainz
T: 06131 603020, F: 06131 6030210
info@kinderwunschzentrum-mainz.de
www.kinderwunschzentrum-mainz.de

**Kinderwunschzentrum der Universitäts-
medizin Mainz**

Dr. med. Ruth Gomez, PD Dr. med. Christine
Skala, Prof. Dr. med. Annette Hasenburg
Langenbeckstraße 1
55131 Mainz
T: 06131 172764, F: 06131 173415
kinderwunsch@unimedizin-mainz.de
www.unimedizin-mainz.de/kinderwunschzentrum/

Kinderwunschzentrum Mittelrhein

Dr. med. Josef Beran, Dr. med. Sebastian
Hagelauer, Ester Baumbach
Marktstraße 83
56564 Neuwied
Viktoriastraße 15
56068 Koblenz
T: 02631 39680, F: 02631 396829
info@kinderwunsch-mittelrhein.de
www.kinderwunsch-mittelrhein.de

Freyja IVF Hagen**Kinderwunsch- & Hormonzentrum**

Dr. med. Birgit Lühr
Bahnhofstraße 1
58095 Hagen
T: 02331 7390421, F:
info@kinderwunsch-hagen.de
www.kinderwunsch-hagen.de

**Kinderwunsch- und Hormonzentrum
Frankfurt am Main
Am Palmengarten**

Prof. Dr. med. Stefan Kissler, FÄ Ines Voß, Prof.
Dr. med. Inka Wiegratz
Gräfstraße 97
60487 Frankfurt am Main
T: 069 5060 68650, F: 069 5060 68651
info@kinderwunschzentrum-frankfurt.de
www.kinderwunschzentrum-frankfurt.de

**reproedicum Kinderwunschzentrum
Reproduktionsmedizin - Gyn. Endokrinologie**

Prof. Dr. med. Dr. med. habil. Ernst Siebzehn-
rübl, Dr. med. Anja Weidner
Hanauer Landstraße 328-330
60314 Frankfurt am Main
T: 069 4260770, F: 069 42607710
mail@reproedicum.de
www.reproedicum.de

Gynäkologische Endokrinologie und Reproduktionsmedizin Re-Pro-Gyn

Universitätsklinikum Frankfurt am Main
Dr. med. Annette Bachmann, Dr. med. Kristin Hawig, Dr. med. Liza Koch, Dr. med. Daniela Rebhan
Theodor-Stern-Kai 7
60590 Frankfurt am Main
T: 069 63015708, F: 069 63017120
info.ivf@kgu.de
www.kgu.de

Kinderwunsch- und Endometriosezentrum am Büsing Park

Dr. med. Konstantin Manolopoulos, Dr. med. Dr. rer. nat. Lutz Belkien, Dr. med. Elena Hartschuh, Dr. (tip) Nurgül Basogul, Dr. med. Ismini Psyraiki, Nadia El Messaoudi, Dr. med. Anna Klauz, Lina Tsiflika
Herrnstraße 51
63065 Offenbach
T: 069 80907571, F: 069 80907573
info@offenbach-kinderwunsch.de
www.offenbach-kinderwunsch.de

Kinderwunschzentrum Darmstadt

Dr. med. Aysen Bilgicyildirim, Dr. med. Matthias Inacker, Dr. med. H. Engelskirchen-Amran, Dr. med. J. Bratengeier, A. Weber-Lohrum
Bratustraße 9
64293 Darmstadt
T: 06151 500980, F: 06151 50098500
info@ivf-da.de
www.kinderwunschzentrum-da.de

MVZ TFP Rhein-Main GmbH

Univ.-Prof. Dr. med. Rudolf Seufert M.Sc., Dr. med. Michael Amrani, Dr. med. Kristina Bockmeyer, Dr. med. Birgit Borzager, Dr. med. Thomas Hahn, Dr. med. Kristin Hawig, Nicole Hunenbart, Dr. med. Julia Limberg, Dr. med. Wolfram Lorei, Dr. med. Eva Rau, Karin Schilberz, Dr. med. Martin Schorsch, Dr. med. Dieter Schrappner, Dr. med. Antje Kühne-Golombek, Dr. med. Dirk Wallmeier, Dr. med. Caroline Faulhaber, Sarah Warschauer
Mainzer Straße 98-102
65189 Wiesbaden
T: 0611 976320, F: 0611 9763210
wiesbaden@tfp-fertility.com
www.tfp-fertility.com/de-de/tfp-kinderwunsch-wiesbaden

IVF-SAAR Saarbrücken-Kaiserslautern

Dr. med. Lars Happel, Dr. med. Sascha Tauchert, Michaela von Blohn, Kathrin Alt, Dr. med. Anette Russu, Dr. rer. medic. Martin Greuner
Europaallee 15
66113 Saarbrücken
T: 06819 36320, F: 06819 363210
Maxstraße 13
67659 Kaiserslautern
T: 0631 70431, F: 0631 78568
zentrum@ivf-saar.de
www.ivf-saar.de

Klinik für Frauenheilkunde, Geburtshilfe und Reproduktionsmedizin

Universitätsklinikum des Saarlandes
Prof. Dr. med. E.-F. Solomayer, Dr. med. Simona Baus, Dr. rer. nat. Jasmin Ney
Kirrbergerstraße 1
66421 Homburg
T: 06841 1628101, F: 06841 1628110
frauenklinik.ivf@uks.eu
www.uks.eu/frauenklinik/kinderwunsch

Kinderwunschzentrum Ludwigshafen

Dr. med. Tobias Schmidt, Dr. med. Claudia Schmidt, Dr. rer. nat. Nicole Hirschmann
Ludwigstraße 54 b
67059 Ludwigshafen
T: 06215 9298688, F: 06215 9298690
kontakt@kinderwunsch-lu.de
www.kinderwunschzentrum-ludwigshafen.de

Kinderwunschzentrum der Universitätsmedizin Mannheim

Prof. Dr. med. Marc Suetterlin, Dr. med. Regine Schaffelder, Dr. med. Amelie Trebin, Dr. med. Thomas Große Steffen
Theodor-Kutzer-Ufer 1-3
68167 Mannheim
T: 0621 3833638, F: 0621 3832705
ivflabor@umm.de
www.umm.de

Viernheimer Institut für Fertilität

PD Dr. med. Stephanie Volz-Köster, Dr. med. Christina Nell
Walter-Gropius-Allee 2
68519 Viernheim
T: 06204 918290, F: 06204 9182910
info@vif-kinderwunsch.de
www.vif-kinderwunsch.de

Praxisgemeinschaft Kinderwunschzentrum Heidelberg

Dr. med. Daniela Seehaus, Dr. sc. hum. Suat Parta, Dr. med. Christina Thöne, Dr. med. Maria-Theresia Schröder
Römerstraße 3
69115 Heidelberg
T: 06221 893000, F: 06221 8930020
info@kwz-hd.de
www.kwz-hd.de

Universitäts-Frauenklinik Heidelberg Abt. Gynäkologische Endokrinologie und Fertilitätsstörungen

Prof. Dr. med. Thomas Strowitzki, Prof. Dr. med. Ariane Germeyer, Dr. med. Sabine Rösner, Dr. med. Julia Rehnitz, Dr. med. Verena Holschbach, Prof. Dr. Ruben Kuon, Dr. med. Anne Marshall, Dr. Marie Weber, Bianca Schell, Dr. rer. nat. Fikret Gürkan Agircan, Dr. rer. nat. Jens Erik Dietrich
Im Neuenheimer Feld 440
69120 Heidelberg
T: 06221 567921, F: 06221 565999
thomas.strowitzki@med.uni-heidelberg.de
www.klinikum.uni-heidelberg.de

Kinderwunsch-Zentrum Stuttgart Praxis Villa Haag

Dr. med. Dieter B. Mayer-Eichberger, Dr. med. Katharina Mayer-Eichberger, Dr. med. Gunilla Gröger, Dr. med. Rafaela Rangel, Dr. rer. nat. Brigitte Blind-Krosch, Samantha Basile
Herweg 69
70174 Stuttgart
T: 0711 221084, F: 0711 221085
info@kinderwunschpraxis.de
www.kinderwunschpraxis.de

Praxis für Reproduktionsmedizin

Dr. Lisa Beiglböck
Dr. med. Lisa Beiglböck, Dr. med. Roland Grau
Friedrichstraße 45
70174 Stuttgart
T: 0711 2200 9030
info@kwz-stuttgart.de
www.frauenarzt-stuttgart.net

Kinderwunschärzte Stuttgart

Überörtliche Berufsausübungsgemeinschaft Dr. med. Tekesin & Dr. med. Kircher GbR

Dr. med. Nina Kircher, Dr. med. Aynur Tekesin, Klaudia Ott
Reinsburgstraße 82
70178 Stuttgart
T: 0711 351455 30, F: 0711 351455 50
info@kinderwunschaerzte-stuttgart.de
www.kinderwunschaerzte-stuttgart.de

Kinderwunschzentrum Bad Cannstatt

Praxis M. Woriedh
Mohammed Woriedh
König-Karl-Straße 66
70372 Stuttgart / Bad-Cannstatt
T: 0711 290671, F: 0711 292321
info@praxis-woriedh.de
www.praxis-woriedh.de

Kinderwunschzentrum Ludwigsburg

Dr. med. Andreas Ott, Dr. med. Annette Schmid, FÄ Petra Klotz, Dr. med. Irina Kiss, Dr. med. Noemi Welsch, Simone Koch
Leonberger Straße 2
71638 Ludwigsburg
T: 07141 688760, F: 07141 688769
info@kiwu-lb.de
www.kiwu-lb.de

KinderwunschPraxis Dres. Göhring

Dr. med. Ulrich Göhring, Dr. med. Inés Göhring
Hagellocher Weg 63
72070 Tübingen
T: 07071 946630, F: 07071 9466399
info@kinderwunschpraxis.com
www.kinderwunschpraxis.com

IVF-Zentrum der Universitäts-Frauenklinik Tübingen

PD Dr. med. Melanie Henes
Calwerstraße 7
72076 Tübingen
T: 07071 2983117, F: 07071 292250
melanie.henes@med.uni-tuebingen.de
www.uni-frauenklinik-tuebingen.de

Kinderwunschzentrum Aalen

Dr. med. Rainer Rau, Dr. med. Birgit Schröppel,
Dr. rer. nat. Roland Eid
Weidenfelder Straße 1
73430 Aalen
T: 07361 62021, F: 07361 62026
info@kinderwunsch-aalen.de
www.kinderwunsch-aalen.de

IVF Zentrum Esslingen

Dr. med. Johann Emil Costea, Dr. med. Alice
Costea, Dr. med. Marius Albowitz
Martinstraße 15
73728 Esslingen
T: 0711 31059160, F: 0711 31059161
info@ivf-praxis.com
www.ivf-praxis.com

Kinderwunsch Frauenärzte**Kinderwunschzentrum Neckarsulm**

Dr. med. Stefan Eisenhardt, Dr. med. Susanne
Kaiser, Sabrina Schomann, MSc., Franziska
Sennert, MSc.
Heilbronner Straße 1
74172 Neckarsulm
T: 07132 4885600, F: 07132 48856099
www.kinderwunsch-frauenaeerzte.de

Centrum für Kinderwunsch Pforzheim**Verena Peuten**

Zerrennerstraße 22-24
75172 Pforzheim
T: 07231 2808 280, F: 07231 2808 2888
info@kinderwunsch-pforzheim.de
www.kinderwunsch-pforzheim.de

ivf Baden-Baden GmbH

Prof. Dr. med. Wolfgang Küpker
Beethovenstraße 2
76530 Baden-Baden
T: 07221 7021 701
info@ivf-baden-baden.com
www.ivf-baden-baden.com

Kinderwunsch Bodensee

Dr. med. Andreas Heine, Dr. med. Lena Otto,
Dr. med. Carola Hornung
Maggistraße 5
78224 Singen
T: 07731 9129990, F: 07731 91299999
info@endlichnachwuchs.de
www.endlichnachwuchs.de

CERF Centrum für Gynäkologische Endokrinologie & Reproduktionsmedizin Freiburg

PD Dr. med. Birgit Wetzka, Dr. med. Veronika
Wolk, Dr. med. Aida Hanjalic-Beck, Dr. med.
Stefanie Friebe, Dr. med. Stefanie Meyer
Bismarckallee 7F
79098 Freiburg
T: 0761 207430, F: 0761 2074318
info@kinderwunsch-freiburg.de
www.kinderwunsch-hormone.de

Universitätsklinikum Freiburg**Klinik für Frauenheilkunde, Endokrinologie und Reproduktionsmedizin**

Dr. med. Philipp Wiehle, Dr. med. Katrin Roth,
Dr. Bengi Acar-Perk
Hugstetter Straße 55
79106 Freiburg
T: 0761 27031500, F: 0761 27029120
frk.ivf-ambulanz@uniklinik-freiburg.de
www.uniklinik-freiburg.de/frauenheilkunde/
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Reproduktionsmedizin München im Tal – MVZ Partnergesellschaft mbH

Dr. med. Daniel Noss, Dr. med. Isabel Stoll, Dr.
med. Patrick Mehrle, Dr. med. Kathrin Pohlig
Tal 11
80331 München
T: 089 2422950, F: 089 24229560
info@ivf-tal.de
www.ivf-tal.de

Die Kinderwunschärztin

Dr. med. Corinna Mann, Dr. med. Anja
Albrecht
Herzogspitalstraße 5
80331 München
T: 089 12359565, F: 089 255526819
hallo@kinderwunschaerztin.de
www.kinderwunschaerztin.de

kiz) kinderwunsch im zentrum**Praxis für gynäkologische Endokrinologie und Reproduktionsmedizin**

PD Dr. med. Roxana Popovici, Dr. med. Anja
Kuhlmann
Bayerstraße 3
80335 München
T: 089 4522178 0, F: 089 4522178 45
info@kiiz.de
www.kiiz.de

Kinderwunsch Zentrum an der Oper

Dr. med. Helmut Lacher, Dr. med. Jörg Puchta,
PD Dr. med. Hans-Ulrich Pauer, Dr. med.
Silke Michna
Maximilianstraße 2a
80539 München
T: 089 547041 0, F: 089 547041 34
info@kinderwunschzentrum-an-der-oper.de
www.kinderwunschzentrum-an-der-oper.de

Kinderwunsch Centrum München (MVZ)

Dr. med. Claudia Gaßner, Dr. med. Gottfried
Krüsmann, Prof. Dr. Dr. med. habil. Wolfgang
Würfel, Dr. med. Jan Krüsmann, Dr. med. Irene
von Hertwig, Sabine Völker
Lortzingstraße 26
81241 München
T: 089 2441440, F: 089 24414441
info@ivf-muenchen.de
www.ivf-muenchen.de

Hormon- und Kinderwunschzentrum**Klinik und Poliklinik für Frauenheilkunde und Geburtshilfe der LMU München**

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Hauter M.Sc., Sandra Wissmiller M.Sc., Dimitra
Makri Ph.D., Chaido Ori M.Sc.; Dr. rer. nat.
Viktoria von Schönfeldt
Marchioninistraße 15
81377 München
T: 089 4400 76825, F: 089 4400 73844
T (Sekretariat): 089 4400 76821
Maistraße 11
80337 München
T (standortübergreifend): 089 4400 76876,
F: 089 4400 54918
ivf@med.lmu.de
www.kinderwunsch-LMU.de

Kinderwunschzentrum A.R.T. Bogenhausen**MVZ für gynäkologische Endokrinologie und Reproduktionsmedizin, Akademische Lehrpraxis der TU München**

Prof. Dr. med. Dieter Berg, Dr. med. Bernd
Lesoine, Dr. med. Barbara de Oriol, Dr. med.
Stephanie Ziehr, Dr. rer. nat. Ulrike Berg
Prinzregentenstraße 69
81675 München
T: 089 414240 0, F: 089 414240 11
info@ivf-muenchen.com
www.ivf-muenchen.com

Kinderwunsch Centrum Chiemsee

Dr. med. Susann Böhm, Dr. med. Angelika Stachl
Hochriesstraße 21
83209 Prien am Chiemsee
T: 08051 5050, F: 08051 63499
info@kinderwunsch-chiemsee.de
www.kinderwunsch-chiemsee.de
Rosenheimer Str. 10
83209 Kolbermoor
T: 08031 91506, F: 08031 304817

Kinderwunschpraxis München Nord

Dr. med. Simon Mittenzwei, Dr. med. Judith
Rattenhuber
Schleißheimer Straße 91
85748 Garching b. München
T: 089 45235450, F: 089 452354545
info@kinderwunschpraxis-muenchen-nord.de
www.kinderwunschpraxis-muenchen-nord.de

Kinderwunschzentrum Augsburg – GMP

Dr. med. Daniela Mischitz, Dr. med.
Klaus-Friedrich Hiller, Dr. med. Thomas Bauer,
Dr. med. Harald Kraus, PD Dr. med. Robert
Ochsenkühn, Dr. med. Vera Hepp
Prinzregentenstraße 25
86150 Augsburg
T: 0821 502780, F: 0821 5027878
info@ivf-augsburg.de
www.ivf-augsburg.de

KinderWunschKempten (KWK)

Zentrum für gynäkologische Endokrinologie und Reproduktionsmedizin, Klinik für Frauenheilkunde und Geburtshilfe, Klinikum Kempten, Klinikverbund Allgäu

Prof. Dr. med. Ricardo Felberbaum, Dr. med. Anke Brössner, Dr. med. Esther Kugler, M.Sc., Klin. Embryologin Nadia Jaouad
Robert-Weixler-Straße 50
87439 Kempten
T: 0831 530 3380, F: 0831 530 3378
kinderwunsch@klinikum-kempten.de
www.kv-keoa.de/kinderwunschzentrum/

Praxisklinik Frauenstraße

MVZ für Kinderwunsch und Pränatalmedizin GmbH

Prof. Dr. med. Karl Sterzik, Dr. med. Erwin Strehler, Dr. med. Kerstin Knab
Frauenstraße 51
89073 Ulm
T: 0731 96651-0, F: 0731 96651-320
info@kinderwunsch-ulm.de
www.kinderwunsch-ulm.de

Universitätsfrauenklinik Ulm

UniFee – Kinderwunsch / Fertility and Endocrinology

Prof. Dr. med. Katharina Hancke
Prittwitzstraße 43
89075 Ulm
T: 0731 500 58663, F: 0731 500 58664
unifee.frauenklinik@uniklinik-ulm.de
www.unifee.de

Kinderwunsch-MVZ Ulm GmbH

Dr. med. Friedrich Gagsteiger, Dr. med. Kerstin Eibner, Julia Koglin, Dr. med. Christian Trautmann, Dr. med. Klaus Bühler
Einsteinstraße 59
89077 Ulm
T: 0731 151590, F: 0731 1515915
info@kwz-ulm.de
www.kwz-ulm.de

Kinderwunsch und Frauen-Hormon Centrum Nürnberg

Dr. med. Joachim Neuwinger, Dr. med. Barbara Munzer-Neuwinger, Prof. Dr. med. Peter Licht
Agnesgasse 2-4
90403 Nürnberg
T: 0911 2355500, F: 0911 2355516
aerzte@kinderwunschzentrum-nuernberg.de
www.ivf-nuernberg.de

Kinderwunschzentrum Erlangen

Dr. med. Jan van Uem, Dr. med. Madeleine Haas
Michael-Vogel-Straße 1e
91052 Erlangen
T: 09131 80950, F: 09131 809530
info@kinderwunschzentrum-erlangen.de
www.kinderwunschzentrum-erlangen.de

**Zentrum für Reproduktionsmedizin Erlangen
Gemeinschaftspraxis der Frauenärzte Dres.
med. Rolf Behrens und Andreas Hammel**

Dr. med. Rolf Behrens, Dr. med. Andreas Hammel
Nürnberger Straße 35
91052 Erlangen
T: 09131 89530, F: 09131 205410
rezeption@ivf-erlangen.de
www.ivf-erlangen.de

Universitäts-Fortpflanzungszentrum Franken (UFF)

Prof. Dr. med. Matthias W. Beckmann, Prof. Dr. med. Susanne Cupisti, Prof. Dr. rer. nat. Ralf Dittrich
Universitätsstraße 21-23
91054 Erlangen
T: 09131 8533553, F: 09131 8533545
fk-uff@uk-erlangen.de
www.uk-erlangen.de

**Praxis Dr. med. Jürgen Krieg
Kinderwunschzentrum Amberg**

Dr. med. Jürgen Krieg
Emailfabrikstraße 15
92224 Amberg
T: 09621 769370, F: 09621 9601612
info@kinderwunschzentrum-amberg.de
www.kinderwunschzentrum-amberg.de

MVZ KITZ Regensburg GmbH

PD Dr. med. Andreas Schüring, Ina Laubert, Dr. med. Lindihana Saliji-Preniqi, Prof. Dr. med. Bernd Seifert, Dr. med. Janine Suhren
Hemauerstraße 1
93047 Regensburg
T: 0941 9925770, F: 0941 99257723
info@kitz-regensburg.de
www.kitz-regensburg.de

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Fachklinik für Fruchtbarkeitsmedizin

Prof. Dr. med. Monika Bals-Pratsch, M. Sc., Dr. med. Angelika Eder, M. Sc., Dr. med. Tanja Ignatov, Dr. med. Christine Reißmann
Hildegard-von-Bingen-Straße 1
93053 Regensburg
T: 0941 89849944, F: 0941 89849945
praxis@profertilita.de
www.profertilita.de

Kinderwunschzentrum Niederbayern

Dr. med. Hans-Joachim Kroiss, Dr. med. Samuel Dadze, Dr. IM Tem. Elfriede Bernhardt
Stadtfeldstraße 50
94469 Deggendorf
T: 0991 29799332, F: 0991 29799331
dr.kroiss@ivf-bayern.de
www.kinderwunsch-niederbayern.de

MVZ Fertility Center Bayreuth GmbH

Tanja Wissendheit, Dr. med. Miklos Hamori
Friedrich-von-Schiller-Straße 35
95444 Bayreuth
T: 0921 53030210, F: 0921 53030211
info@fertility-center-bt.de
www.fertility-center-bayreuth.de

MainKid

Kinderwunschzentrum am Theater

Prof. Dr. Ursula Zollner, Klaus-Peter Zollner, M.Sc.
Theaterstraße 20
97070 Würzburg
T: 0931 45276630, F: 0931 45276628
info@mainkid.de
www.mainkid-kinderwunsch.de

Zentrum für Reproduktionsmedizin und Pränataldiagnostik

Dr. med. Reinhard Mai, Dr. med. Lore Mulfinger, Dr. med. Florian Jakob (Ang.)
Juliuspromenade 7
97070 Würzburg
T: 0931 321230, F: 0931 3212377
kontakt@drs-mai-mulfinger-jakob.de
www.drs-mai-mulfinger-jakob.de

**Universitätsklinikum Würzburg
Frauenklinik und Poliklinik, Zentrum für gynäkologische Endokrinologie und Reproduktionsmedizin (ZERM)**

Dr. med. Michael Schwab, Dr. med. Anastasia Altides, Dr. rer. nat. Claudia Staib
Josef-Schneider-Straße 4
97080 Würzburg
T: 0931 201 25619, F: 0931 201 25406
kinderwunsch@klinik.uni-wuerzburg.de

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