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Gynäkologie • Kontrazeption • Psychosomatik • Reproduktionsmedizin • Urologie

D.I.R-Annual 2019
Blumenauer V, Czeromin U, Fehr D, Fiedler K, Gnoth C
Krüssel JS, Kupka MS, Ott A, Tandler-Schneider A
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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 2019 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 nearly 320,000 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 2019 is like the Yearbook 2018 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 2019 yearbook is based on exploitable exports from 131 centers, for which a number of 110,786 plausible cycles was documented. A total of 62,990 women were treated in 2019, resulting in an average of 1.8 cycles per woman. We are happy to note that the pregnancy rate per transfer in fresh cycles was 31.6%, whereas the pregnancy rate per transfer in cryo cycles was 29.6%. In the previous year, 2018, the documented birth rate per embryo transfer was 23.5% in fresh cycles and 20.0% in cryo cycles, with a very good general pregnancy outcome documentation of 95.0%.

Remarkably, 71.8% of the pregnancies led to a birth. A birth rate of 22.3% 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 [BGBl 2013; 56: 1633–41].

In the last years particular attention was placed on the known problems of multiple pregnancies within ART cycles. In 2018, the rate of twin births was 19.2% in fresh cycles and 12.7% in cryo cycles. This rate is still too high, but we can see a small decrease in multiple birth for the first time!

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

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### Number of Treatments in 2019

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

**Members of the German IVF-Registry 2019**  
*n = 137*

- **Registry Participants 2019***  
  *n = 131*

- **Data Received by Deadline Jun 2nd 2020***  
  *n = 131*

- **Documented Treatment Cycles**  
  *n = 110,786*

- **Number of Women Treated**  
  *n = 62,990*

**Mean Number of Treatment Cycles per Woman**  
**1.8**

### Type of plausible treatment 2015 – 2019

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

<table>
<thead>
<tr>
<th></th>
<th>2015</th>
<th>2016</th>
<th>2017</th>
<th>2018</th>
<th>2019</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>n</strong></td>
<td>15,164</td>
<td>15,879</td>
<td>15,606</td>
<td>17,285</td>
<td>17,690</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>15.8</td>
<td>15.8</td>
<td>15.2</td>
<td>16.4</td>
<td>16.5</td>
</tr>
<tr>
<td><strong>IVF</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>48,674</td>
<td>48,690</td>
<td>47,471</td>
<td>46,604</td>
<td>45,381</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>50.6</td>
<td>48.3</td>
<td>46.3</td>
<td>44.3</td>
<td>42.3</td>
</tr>
<tr>
<td><strong>ICSI</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>1,223</td>
<td>1,203</td>
<td>1,170</td>
<td>1,439</td>
<td>1,330</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>1.3</td>
<td>1.2</td>
<td>1.1</td>
<td>1.4</td>
<td>1.2</td>
</tr>
<tr>
<td><strong>IVF/ICSI</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>1,347</td>
<td>1,347</td>
<td>1,307</td>
<td>1,307</td>
<td>1,856</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>1.3</td>
<td>1.3</td>
<td>1.2</td>
<td>1.2</td>
<td>1.7</td>
</tr>
<tr>
<td><strong>Freeze all - MII</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>517</td>
<td>436</td>
<td>933</td>
<td>436</td>
<td>933</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>0.5</td>
<td>0.4</td>
<td>0.9</td>
<td>0.4</td>
<td>0.9</td>
</tr>
<tr>
<td><strong>Freeze All - PNs and Embryos</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>23,571</td>
<td>25,008</td>
<td>27,234</td>
<td>28,698</td>
<td>30,666</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>24.5</td>
<td>24.8</td>
<td>26.6</td>
<td>27.3</td>
<td>28.6</td>
</tr>
<tr>
<td><strong>Cryo</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>7,492</td>
<td>9,974</td>
<td>8,907</td>
<td>9,047</td>
<td>9,263</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>7.8</td>
<td>9.9</td>
<td>8.7</td>
<td>8.6</td>
<td>8.6</td>
</tr>
<tr>
<td><strong>Mixed Fresh</strong> and <strong>Cryo Cycles</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>235</td>
<td>235</td>
<td>235</td>
<td>235</td>
<td>235</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td>0.2</td>
<td>0.2</td>
<td>0.2</td>
<td>0.2</td>
<td>0.2</td>
</tr>
<tr>
<td><strong>None (= Break-off before oocyte treatment or thawing)</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>n</strong></td>
<td>96,124</td>
<td>100,754</td>
<td>102,487</td>
<td>105,102</td>
<td>107,373</td>
</tr>
<tr>
<td><strong>%</strong></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

*Six centers could not be part of the nationale analyses. All centers used the interface ARTbox®.*

**Base quantity: Total number of women, including implausible treatment cycles**
**Summary of Statistics in Brief 2018 – CoD Jun 2nd 2020**

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

- **Births per Transfer:** 23.6%
- **Singletons:** 80.4%
  - n = 10,143
- **Twins:** 19.2%
  - n = 2,419
- **Triplets:** 0.4%
  - n = 56
- **Quadruplets:** 0.0%
  - n = 0
- **Miscarriage:** 19.8%
  - n = 3,416

**Data for 1982 to 2014 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.

---

**Number of Oocyte Retrievals (Freshcycles) 1982 – 2019**

- **Number of Thawing Cycles 1994 – 2019**
- **Registry Participants 1982 – 2019**

**IVF, ICSI* – Prospective and Retrospective Data**
Quality of Documentation 2018/2019
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 2019
107,373 plausible cycles out of 110,786 documented cycles. 96.9% of documented cycles are plausible. This does unfortunately not imply completeness of data required, thus impeding detailed analyses.

Prospectively Documented Cycles 2019
97,147 prospectively documented cycles out of 107,373 plausible cycles. 90.5% of documented 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 2019
In 81,700 out of 82,242 embryo transfers, the cycle outcome was documented (99.3%).

Documented Pregnancy Outcomes 2018
In 23,966 out of 25,230 clinical pregnancies, the pregnancy outcome was documented (95.0%).

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 second year the percentage of prospectively documented cycles is above 90%.

Furthermore we are especially happy that the percentage of documented pregnancy outcomes has increased again. 72.1% of individual centers have reached the goal of documenting more than 95% of pregnancy outcomes. 48.1% have even reached a documentation rate on >98%, 25 centers even 100%!

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 2017 and 2018

Prospective and Retrospective Data

*With a 95%-probability, the true mean lies within the defined confidence interval.*
D·I·R Statistics in Brief – Fresh Cycles 2019 (CoD Jun 2nd 2020)

German IVF Registry – Prospective and Retrospective Data

### Recorded Cycles 2019

| Recorded Cycles 2019 | 110,786 | 100.0% |

### Plausible Cycles 2019 (Prospective and not Prospective)

| Plausible Cycles 2019 | 107,373 | 96.9% |

### Prospective Cycles

| Prospective Cycles | 97,147 | 90.5% |

### Preparation for OPU

| Preparation for OPU | 74,715 | 69.6% |

#### OPU

| OPU | 69,337 | 92.8% |
| No OPU | 5,378 | 7.2% |

#### Oocytes Aspirated

| Oocytes Aspirated | 67,334 | 97.1% |
| No Oocytes | 2,003 | 2.9% |

#### Oocyte Culture

| Oocyte Culture | 65,334 | 97.0% |
| Freeze All – MII* | 1,856 | 2.8% |
| No Oocyte Culture | 144 | 0.2% |

#### Oocyte for Transfer

| Oocyte for Transfer | 64,401 | 98.6% |
| Freeze All – Oocyte Culture | 933 | 1.4% |

### Total Fresh Cycles

| Total Fresh Cycles | 64,401 | 100.0% |

### Total Fertilization

| Total Fertilization | 60,720 | 94.3% |

### Total Transfer

| Total Transfer | 53,012 | 87.3% |
| Outc. Unknown | 285 | 0.5% |

### Total CPR/ET (doc.)

| Total CPR/ET (doc.) (per Treatment 25.9%) | 16,677 | 31.6% |

### IVF

| IVF | 17,690 | 27.5% |
| Fertilization | 16,320 | 92.3% |
| Transfer | 14,380 | 88.1% |

### ICSI

| ICSI | 45,381 | 70.5% |
| Fertilization | 43,081 | 94.9% |
| Transfer | 37,572 | 87.2% |

### IVF/ICSI

| IVF/ICSI | 1,330 | 2.1% |

### Fertilization

| Fertilization | 1,319 | 99.2% |
| Transfer | 1,060 | 80.4% |

### Transfer

| Transfer | 1,060 | 80.4% |

### CPR/ET (doc.)

| CPR/ET (doc.) (per IVF-Treatment 26.9%) | 4,755 | 33.2% |
| CPR/ET (doc.) (per ICSI-Treatment 25.4%) | 11,549 | 30.9% |
| CPR/ET (doc.) (per IVF/ICSI-Treatment 28.0%) | 373 | 35.5% |

*) Out of 1,856 cycles with freeze-all MII oocytes, 390 cycles were performed for FertiPROTECT (=medical freezing) and 953 cycles for social freezing. 513 cycles could not be assigned to either indication.
Out of 1,307 cycles with freeze-all MII oocytes, 280 cycles were performed for FertiPROTECT (=medical freezing) and 791 cycles for social freezing. 236 cycles could not be assigned to either indication.

<table>
<thead>
<tr>
<th>Recorded Cycles 2018</th>
<th>106,397</th>
<th>100.0%</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Plausible Cycles 2018 (Prospective and not Prospective)</strong></td>
<td>105,102</td>
<td>98.8%</td>
</tr>
<tr>
<td><strong>Prospective Cycles</strong></td>
<td>94,866</td>
<td>90.3%</td>
</tr>
<tr>
<td><strong>Preparation for OPU</strong></td>
<td>74,762</td>
<td>71.1%</td>
</tr>
<tr>
<td><strong>OPU</strong></td>
<td>69,757</td>
<td>93.3%</td>
</tr>
<tr>
<td><strong>No OPU</strong></td>
<td>5,005</td>
<td>6.7%</td>
</tr>
<tr>
<td><strong>Oocytes Aspirated</strong></td>
<td>67,811</td>
<td>97.2%</td>
</tr>
<tr>
<td><strong>No Oocytes</strong></td>
<td>1,946</td>
<td>2.8%</td>
</tr>
<tr>
<td><strong>Oocyte Culture</strong></td>
<td>65,764</td>
<td>97.0%</td>
</tr>
<tr>
<td><strong>Freeze All – MII</strong></td>
<td>1,307</td>
<td>1.9%</td>
</tr>
<tr>
<td><strong>No Oocyte Culture</strong></td>
<td>740</td>
<td>1.1%</td>
</tr>
<tr>
<td><strong>Oocyte for Transfer</strong></td>
<td>65,328</td>
<td>99.3%</td>
</tr>
<tr>
<td><strong>Freeze All – Oocyte Culture</strong></td>
<td>436</td>
<td>0.7%</td>
</tr>
<tr>
<td><strong>Total Fresh Cycles</strong></td>
<td>65,328</td>
<td>100.0%</td>
</tr>
<tr>
<td><strong>Total Fertilization</strong></td>
<td>61,230</td>
<td>93.7%</td>
</tr>
<tr>
<td><strong>Total Transfer</strong></td>
<td>53,627</td>
<td>87.6%</td>
</tr>
<tr>
<td><strong>Outc. Unknown</strong></td>
<td>144</td>
<td>0.3%</td>
</tr>
<tr>
<td><strong>Total CPR/ET (doc.)</strong></td>
<td>17,226</td>
<td>32.2%</td>
</tr>
<tr>
<td>(per Treatment 26.4%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>CPR/ET (doc.)</strong></td>
<td>4,644</td>
<td>33.5%</td>
</tr>
<tr>
<td>(per IVF-Treatment 26.9%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Misc.</strong></td>
<td>3,476</td>
<td>20.2%</td>
</tr>
<tr>
<td><strong>EP</strong></td>
<td>247</td>
<td>1.4%</td>
</tr>
<tr>
<td><strong>Not yet Recorded</strong></td>
<td>885</td>
<td>5.1%</td>
</tr>
<tr>
<td><strong>Birth (doc.)</strong></td>
<td>12,618</td>
<td>73.2%</td>
</tr>
<tr>
<td>(per Transfer 23.5%, per Tream. 19.3%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Births</strong></td>
<td>12,534</td>
<td>76.1%</td>
</tr>
<tr>
<td>(per Transfer 25.4%, per Tream. 20.4%)</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>Birth</strong></td>
<td>8,798</td>
<td>72.2%</td>
</tr>
<tr>
<td>(per Transfer 22.8%, per Tream. 18.9%)</td>
<td></td>
<td></td>
</tr>
</tbody>
</table>

| IVF | 17,285 | 26.5% |
| Fertilization | 15,772 | 91.2% |
| Transfer | 13,926 | 88.3% |
| Outc. Unknown | 54 | 0.4% |
| **ICSI** | 46,604 | 71.3% |
| Fertilization | 44,032 | 94.5% |
| Transfer | 38,572 | 89.2% |
| Outc. Unknown | 88 | 0.2% |
| **IVF/ICSI** | 1,439 | 2.2% |
| Fertilization | 1,426 | 99.1% |
| Transfer | 1,129 | 79.2% |
| Outc. Unknown | 5 | 0.4% |

| Misc. | 2,438 | 20.0% |
| **EP** | 147 | 1.2% |
| **Not yet Recorded** | 811 | 6.7% |
| **Birth (doc.)** | 7,106 | 80.8% |
| (per Transfer 22.8%, per Tream. 18.9%) |
| **Births** | 7,211 | 79.3% |
| (per Transfer 25.4%, per Tream. 20.4%) |
| **Birth** | 1,652 | 18.8% |
| (per Transfer 22.8%, per Tream. 18.9%) |

| Misc. | 70 | 18.0% |
| **EP** | 4 | 1.0% |
| **Not yet Recorded** | 28 | 7.2% |
| **Birth (doc.)** | 286 | 73.7% |
| (per Transfer 23.5%, per Tream. 19.9%) |

| Singletons | 10,143 | 80.4% |
| Twins | 2,419 | 19.2% |
| Triplets | 56 | 0.4% |
| Quadruplets | 0 | - |

| Singletons | 2,801 | 79.3% |
| Twins | 721 | 20.4% |
| Triplets | 12 | 0.3% |
| Quadruplets | 0 | - |

| Singletons | 7,106 | 80.8% |
| Twins | 1,652 | 18.8% |
| Triplets | 40 | 0.5% |
| Quadruplets | 0 | - |
## D·I·R Statistics in Brief – Cryo Cycles 2019 (CoD Jun 2nd 2020)

### German IVF Registry – Prospective and Retrospective Data

**Recorded Cycles 2019**

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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</thead>
<tbody>
<tr>
<td>110,786</td>
<td>100.0%</td>
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</tbody>
</table>

**Plausible Cycles 2019 (Prospective and not Prospective)**

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>107,373</td>
<td>96.9%</td>
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</tbody>
</table>

**Prospective Cycles**

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>97,147</td>
<td>90.5%</td>
</tr>
</tbody>
</table>

**Preparation for Cryo Transfer**

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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</tr>
</thead>
<tbody>
<tr>
<td>32,658</td>
<td>30.4%</td>
</tr>
</tbody>
</table>

### Thawing

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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</tr>
</thead>
<tbody>
<tr>
<td>Thawing</td>
<td>30,666</td>
</tr>
<tr>
<td></td>
<td>93.9%</td>
</tr>
<tr>
<td>No Thawing</td>
<td>1,992</td>
</tr>
<tr>
<td></td>
<td>6.1%</td>
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### Fertilization

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>222</td>
<td></td>
</tr>
<tr>
<td></td>
<td>87.4%</td>
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### Transfer

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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<tbody>
<tr>
<td>211</td>
<td></td>
</tr>
<tr>
<td></td>
<td>95.0%</td>
</tr>
</tbody>
</table>

### Total Thawing

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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</thead>
<tbody>
<tr>
<td>29,230</td>
<td></td>
</tr>
<tr>
<td></td>
<td>95.3%</td>
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### Thawing ex IVF

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<table>
<thead>
<tr>
<th></th>
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</thead>
<tbody>
<tr>
<td>8,155</td>
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</tr>
<tr>
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<td>26.6%</td>
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### Thawing ex ICSI

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<table>
<thead>
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</thead>
<tbody>
<tr>
<td>21,395</td>
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<tr>
<td></td>
<td>69.8%</td>
</tr>
</tbody>
</table>

### Thawing Oocytes

<p>| | |</p>
<table>
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<tr>
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</thead>
<tbody>
<tr>
<td>254</td>
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<tr>
<td></td>
<td>0.8%</td>
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</table>

### Thawing Others*

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
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<tbody>
<tr>
<td>862</td>
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<td></td>
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</table>

### Total CPR/ET (doc.)

<p>| | |</p>
<table>
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</thead>
<tbody>
<tr>
<td>8,573</td>
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</table>

### CPR/ET (doc.)

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<table>
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<th></th>
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</thead>
<tbody>
<tr>
<td>2,428</td>
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<td></td>
<td>31.3%</td>
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</tbody>
</table>

### CPR/ET (doc.)

<p>| | |</p>
<table>
<thead>
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<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>5,959</td>
<td></td>
</tr>
<tr>
<td></td>
<td>29.4%</td>
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</tbody>
</table>

### CPR/ET (doc.)

<p>| | |</p>
<table>
<thead>
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<th></th>
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</thead>
<tbody>
<tr>
<td>128</td>
<td></td>
</tr>
<tr>
<td></td>
<td>16.4%</td>
</tr>
</tbody>
</table>

### CPR/ET (doc.)

<p>| | |</p>
<table>
<thead>
<tr>
<th></th>
<th></th>
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</thead>
<tbody>
<tr>
<td>48</td>
<td></td>
</tr>
<tr>
<td></td>
<td>22.7%</td>
</tr>
</tbody>
</table>

*) 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 2018** (CoD Jun 2nd 2020)

**German IVF Registry – Prospective and Retrospective Data**

<table>
<thead>
<tr>
<th>Recorded Cycles 2018</th>
<th>106,397</th>
<th>100.0%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Plausible Cycles 2018 (Prospective and not Prospective)</td>
<td>105,102</td>
<td>98.8%</td>
</tr>
<tr>
<td>Prospective Cycles</td>
<td>94,866</td>
<td>90.3%</td>
</tr>
</tbody>
</table>

**Preparation for Cryo Transfer**

30,340 | 29.8% |

- **Thawing**
  - 28,698 | 94.6% |
  - 1,642 | 5.4% |

- **Fertilization**
  - 245 | 91.4% |
  - 222 | 90.6% |

- **Misc.**
  - 2,035 | 25.4% |
  - 379 | 4.7% |

| Thawing ex IVF | 7,355 | 25.6% |
| Thawing ex ICSI | 20,041 | 69.8% |
| Thawing Others* | 1,034 | 3.6% |
| Thawing Oocytes | 268 | 0.9% |

| CPR/ET (doc.) | 5,759 | 30.1% |
| CPR/ET (doc.) | 54 | 24.9% |

| Misc. | 1,440 | 25.0% |
| EP | 15 | 9.4% |
| Not yet Rec. | 322 | 5.6% |
| Misc. | 2 | 3.7% |
| EP | 0 | 0.0% |
| Not yet Rec. | 27 | 50.0% |

| Birth (doc.) | 1,396 | 68.7% |
| (per Tr. 19.8%, p. Tr. 19.0%) | | |
| Misc. | 3,941 | 68.4% |
| EP | 137 | 85.6% |
| Not yet Rec. | 54 | 61.2% |
| Misc. | 2 | 3.7% |
| EP | 0 | 0.0% |
| Not yet Rec. | 27 | 50.0% |

<table>
<thead>
<tr>
<th>Briths</th>
<th>n %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Singletons</td>
<td>4,780</td>
</tr>
<tr>
<td>Twins</td>
<td>701</td>
</tr>
<tr>
<td>Triplets</td>
<td>18</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
<tr>
<td>Singletons</td>
<td>1,211</td>
</tr>
<tr>
<td>Twins</td>
<td>182</td>
</tr>
<tr>
<td>Triplets</td>
<td>3</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
<tr>
<td>Singletons</td>
<td>3,419</td>
</tr>
<tr>
<td>Twins</td>
<td>507</td>
</tr>
<tr>
<td>Triplets</td>
<td>15</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
<tr>
<td>Singletons</td>
<td>128</td>
</tr>
<tr>
<td>Twins</td>
<td>9</td>
</tr>
<tr>
<td>Triplets</td>
<td>0</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Briths</th>
<th>n %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Singletons</td>
<td>22</td>
</tr>
<tr>
<td>Twins</td>
<td>3</td>
</tr>
<tr>
<td>Triplets</td>
<td>0</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
</tbody>
</table>

*) 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 2018

Prospective Data

IVF 2018

<table>
<thead>
<tr>
<th>Age in Years</th>
<th>&lt;=29</th>
<th>30 – 34</th>
<th>35 – 39</th>
<th>40</th>
<th>41</th>
<th>42</th>
<th>43</th>
<th>44</th>
<th>&gt;=45 Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>OPU</td>
<td>1,886</td>
<td>4,917</td>
<td>6,629</td>
<td>678</td>
<td>615</td>
<td>417</td>
<td>312</td>
<td>196</td>
<td>115</td>
</tr>
<tr>
<td>Oocytes</td>
<td>11.7</td>
<td>10.5</td>
<td>8.3</td>
<td>6.8</td>
<td>6.3</td>
<td>5.5</td>
<td>5.6</td>
<td>4.7</td>
<td>3.5</td>
</tr>
<tr>
<td>Insemination</td>
<td>11.1</td>
<td>10.3</td>
<td>8.2</td>
<td>6.7</td>
<td>6.2</td>
<td>5.5</td>
<td>5.6</td>
<td>4.6</td>
<td>3.4</td>
</tr>
<tr>
<td>ET</td>
<td>1,488</td>
<td>4,057</td>
<td>5,500</td>
<td>545</td>
<td>494</td>
<td>320</td>
<td>236</td>
<td>152</td>
<td>76</td>
</tr>
<tr>
<td>ET/OPU %</td>
<td>78.9</td>
<td>82.5</td>
<td>83.0</td>
<td>80.4</td>
<td>80.3</td>
<td>76.7</td>
<td>75.6</td>
<td>77.6</td>
<td>66.1</td>
</tr>
<tr>
<td>Trans. Embr.</td>
<td>1.8</td>
<td>1.7</td>
<td>1.8</td>
<td>1.7</td>
<td>1.8</td>
<td>1.8</td>
<td>1.8</td>
<td>1.7</td>
<td>1.7</td>
</tr>
<tr>
<td>CP</td>
<td>607</td>
<td>1,604</td>
<td>1,813</td>
<td>130</td>
<td>100</td>
<td>51</td>
<td>33</td>
<td>13</td>
<td>2</td>
</tr>
<tr>
<td>CP/OPU %</td>
<td>32.2</td>
<td>32.6</td>
<td>27.3</td>
<td>19.2</td>
<td>16.3</td>
<td>12.2</td>
<td>10.6</td>
<td>6.6</td>
<td>1.7</td>
</tr>
<tr>
<td>CP/ET Upper Conf. Limit %</td>
<td>43.4</td>
<td>41.2</td>
<td>34.4</td>
<td>26.3</td>
<td>23.9</td>
<td>19.4</td>
<td>17.9</td>
<td>11.6</td>
<td>3.5</td>
</tr>
<tr>
<td>CP/ET %</td>
<td>40.8</td>
<td>39.5</td>
<td>33.0</td>
<td>23.9</td>
<td>20.2</td>
<td>15.9</td>
<td>14.0</td>
<td>8.6</td>
<td>2.6</td>
</tr>
<tr>
<td>CP/ET Lower Conf. Limit %</td>
<td>38.2</td>
<td>38.1</td>
<td>31.8</td>
<td>21.6</td>
<td>17.0</td>
<td>13.0</td>
<td>10.8</td>
<td>6.2</td>
<td>2.0</td>
</tr>
<tr>
<td>CP/ET %: 2 Embryo Transfer + min. 2 remaining 2-PN Embryo Surplus</td>
<td>44.5</td>
<td>43.7</td>
<td>38.9</td>
<td>29.4</td>
<td>23.9</td>
<td>19.3</td>
<td>16.7</td>
<td>15.3</td>
<td>7.0</td>
</tr>
<tr>
<td>Misc./CP Upper Conf. Limit %</td>
<td>17.6</td>
<td>18.9</td>
<td>23.1</td>
<td>41.4</td>
<td>45.4</td>
<td>56.0</td>
<td>56.8</td>
<td>62.4</td>
<td>100.0</td>
</tr>
<tr>
<td>Misc./CP %</td>
<td>15.8</td>
<td>15.9</td>
<td>22.4</td>
<td>37.7</td>
<td>40.0</td>
<td>49.0</td>
<td>48.3</td>
<td>53.8</td>
<td>100.0</td>
</tr>
<tr>
<td>Misc./CP Lower Conf. Limit %</td>
<td>11.6</td>
<td>15.0</td>
<td>18.6</td>
<td>32.2</td>
<td>29.2</td>
<td>38.0</td>
<td>33.9</td>
<td>30.9</td>
<td>100.0</td>
</tr>
<tr>
<td>Births/ET Upper Conf. Limit %</td>
<td>35.0</td>
<td>33.6</td>
<td>25.0</td>
<td>26.2</td>
<td>23.0</td>
<td>18.2</td>
<td>16.4</td>
<td>9.9</td>
<td>0.0</td>
</tr>
<tr>
<td>Births/ET %</td>
<td>32.1</td>
<td>32.4</td>
<td>24.7</td>
<td>14.9</td>
<td>10.9</td>
<td>6.9</td>
<td>7.6</td>
<td>3.3</td>
<td>0.0</td>
</tr>
<tr>
<td>Births/ET Lower Conf. Limit %</td>
<td>29.8</td>
<td>30.6</td>
<td>22.5</td>
<td>8.7</td>
<td>8.9</td>
<td>4.7</td>
<td>4.7</td>
<td>1.5</td>
<td>0.0</td>
</tr>
</tbody>
</table>

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 2018

Prospective Data

### ICSI 2018

<table>
<thead>
<tr>
<th>Age in Years</th>
<th>&lt;= 29</th>
<th>30 – 34</th>
<th>35 – 39</th>
<th>40</th>
<th>41</th>
<th>42</th>
<th>43</th>
<th>44</th>
<th>&gt;=45</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>OPU</td>
<td>5,596</td>
<td>13,629</td>
<td>18,094</td>
<td>2,059</td>
<td>1,578</td>
<td>1,253</td>
<td>875</td>
<td>541</td>
<td>491</td>
<td>44,116</td>
</tr>
<tr>
<td>Oocytes¹</td>
<td>12.2</td>
<td>11.2</td>
<td>8.8</td>
<td>6.9</td>
<td>6.6</td>
<td>5.9</td>
<td>5.5</td>
<td>5.5</td>
<td>4.2</td>
<td>9.6</td>
</tr>
<tr>
<td>Injection¹</td>
<td>8.9</td>
<td>8.6</td>
<td>6.8</td>
<td>5.4</td>
<td>5.2</td>
<td>4.6</td>
<td>4.1</td>
<td>3.9</td>
<td>3.2</td>
<td>7.3</td>
</tr>
<tr>
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<td>4,653</td>
<td>11,603</td>
<td>15,397</td>
<td>1,657</td>
<td>1,259</td>
<td>991</td>
<td>653</td>
<td>412</td>
<td>327</td>
<td>36,952</td>
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<tr>
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<td>85.1</td>
<td>85.1</td>
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<td>79.1</td>
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<td>1.7</td>
<td>1.7</td>
<td>1.8</td>
<td>1.8</td>
<td>1.8</td>
<td>1.8</td>
<td>1.8</td>
<td>1.8</td>
<td>1.7</td>
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<td>4,712</td>
<td>347</td>
<td>228</td>
<td>153</td>
<td>69</td>
<td>26</td>
<td>8</td>
<td>11,790</td>
</tr>
<tr>
<td>CP/OPU</td>
<td>32.6</td>
<td>32.5</td>
<td>26.0</td>
<td>16.9</td>
<td>14.4</td>
<td>12.2</td>
<td>7.9</td>
<td>4.8</td>
<td>1.6</td>
<td>26.7</td>
</tr>
<tr>
<td>CP/ET</td>
<td>40.7</td>
<td>39.0</td>
<td>31.3</td>
<td>23.1</td>
<td>21.4</td>
<td>18.7</td>
<td>13.5</td>
<td>8.6</td>
<td>3.2</td>
<td>32.4</td>
</tr>
<tr>
<td>CP/ET Upper Limit*%</td>
<td>39.2</td>
<td>38.1</td>
<td>30.6</td>
<td>20.9</td>
<td>18.1</td>
<td>15.4</td>
<td>10.6</td>
<td>6.3</td>
<td>2.4</td>
<td>31.9</td>
</tr>
<tr>
<td>CP/ET Lower Limit*%</td>
<td>37.8</td>
<td>37.2</td>
<td>29.9</td>
<td>19.0</td>
<td>15.2</td>
<td>12.6</td>
<td>8.2</td>
<td>4.6</td>
<td>1.8</td>
<td>31.4</td>
</tr>
<tr>
<td>CP/ET %: 2 Emb. Trans. + min. 2 2-PN Surplus</td>
<td>43.7</td>
<td>42.3</td>
<td>38.1</td>
<td>29.0</td>
<td>25.0</td>
<td>19.5</td>
<td>14.2</td>
<td>11.9</td>
<td>13.8</td>
<td>36.2</td>
</tr>
<tr>
<td>Misc./CP Upper Limit*%</td>
<td>14.4</td>
<td>17.5</td>
<td>23.9</td>
<td>35.5</td>
<td>42.3</td>
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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 2014 – 2018

Prospective Data

### IVF 2014 – 2018

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<thead>
<tr>
<th>Age in Years</th>
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<th>30 – 34</th>
<th>35 – 39</th>
<th>40</th>
<th>41</th>
<th>42</th>
<th>43</th>
<th>44</th>
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<td>3,594</td>
<td>3,962</td>
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<td>3.3</td>
<td>2.8</td>
<td>21.3</td>
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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 2014 – 2018

Prospective Data

### ICSI 2014 – 2018

<table>
<thead>
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<th>Age in Years</th>
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<th>35 – 39</th>
<th>40</th>
<th>41</th>
<th>42</th>
<th>43</th>
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<td>1.9</td>
<td>1.9</td>
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<td>1.9</td>
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<td>17.2</td>
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<tr>
<td>CP/ET %</td>
<td>38.9</td>
<td>38.1</td>
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<td>19.9</td>
<td>16.2</td>
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<td>11.3</td>
<td>6.3</td>
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<td>31.0</td>
<td>23.1</td>
<td>19.0</td>
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<td>12.9</td>
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<td>18.4</td>
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<td>37.4</td>
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<td>48.4</td>
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<td>32.3</td>
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<td>35.6</td>
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<td>7.2</td>
<td>5.2</td>
<td>3.2</td>
<td>21.5</td>
</tr>
<tr>
<td>Births/ET %</td>
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<td>20.7</td>
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<td>10.8</td>
<td>7.7</td>
<td>6.2</td>
<td>4.5</td>
<td>2.7</td>
<td>21.3</td>
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<tr>
<td>Births/ET Lower Confidence Limit %</td>
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<td>26.8</td>
<td>20.4</td>
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<td>9.9</td>
<td>7.1</td>
<td>5.6</td>
<td>3.9</td>
<td>2.3</td>
<td>20.8</td>
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</table>

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 2018

**Prospective Data**

#### IVF 2018

<table>
<thead>
<tr>
<th></th>
<th>n</th>
<th>%</th>
<th>Fertilization %</th>
<th>Embryo %</th>
<th>Transfer %</th>
<th>Clin. Preg. %</th>
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<tr>
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<td>Ectopic Pregnancy</td>
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<td></td>
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<tr>
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* Successful fertilization of at least one oocyte per cycle

#### ICSI 2018

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<th>Fertilization %</th>
<th>Embryo %</th>
<th>Transfer %</th>
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#### IVF and ICSI in Natural Cycles 2018**

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<th>Fertilization %</th>
<th>Embryo %</th>
<th>Transfer %</th>
<th>Clin. Preg. %</th>
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</table>

*) Successful fertilization of at least one oocyte per cycle

**) Natural cycle and natural cycle with low-dose-stimulation were newly defined in the data set. Only clearly allocatable cycles were included into the analyses. The total number is most likely higher.
Results of Thawing-Cycles, TESE, IVF and ICSI with Donor Semen 2018

Prospective Data

### Cryo Transfer Cycles 2018

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<th>PN/Embryo %</th>
<th>Transfer %</th>
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### TESE 2018

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<th>Fertilization %</th>
<th>Embryo %</th>
<th>Transfer %</th>
<th>Clin. Preg. %</th>
</tr>
</thead>
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<td>100.0</td>
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<td>434</td>
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<td>134</td>
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<tr>
<td>Ectopic Pregnancy</td>
<td>12</td>
<td></td>
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<tr>
<td>Not Yet Recorded</td>
<td>51</td>
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### IVF and ICSI with Donor Semen 2018**

<table>
<thead>
<tr>
<th></th>
<th>n</th>
<th>%</th>
<th>Fertilization %</th>
<th>Embryo %</th>
<th>Transfer %</th>
<th>Clin. Preg. %</th>
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<tbody>
<tr>
<td>ART-Treatm. (donor sperm)**</td>
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<td>100.0</td>
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<tr>
<td>Successful Fertilization*</td>
<td>861</td>
<td>76.3</td>
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<tr>
<td>Minimum 1 Embryo</td>
<td>803</td>
<td>71.1</td>
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<td>ET Performed</td>
<td>801</td>
<td>70.9</td>
<td>93.0</td>
<td>99.8</td>
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<tr>
<td>Clin. Pregnancy</td>
<td>272</td>
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<td>34.0</td>
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<tr>
<td>Birth</td>
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<td>16.3</td>
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</tr>
<tr>
<td>Not Yet Recorded</td>
<td>76</td>
<td></td>
<td></td>
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<td>27.9</td>
</tr>
</tbody>
</table>

*) Successful fertilization of at least one oocyte per cycle
**) Currently, the data processing is not completely validated. Only cycles with definite data validation were included in this analysis.
Positive Pregnancy Outcomes 2018
IVF, ICSI – Prospective and Retrospective Data

<table>
<thead>
<tr>
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<tr>
<td>Clinical Pregnancies</td>
<td>17,226</td>
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</tr>
<tr>
<td>Outcome documented</td>
<td>16,341</td>
<td>94.9</td>
</tr>
<tr>
<td>Transfer</td>
<td>53,627</td>
<td></td>
</tr>
<tr>
<td>Life-Birth-Rate per ET</td>
<td>12,449</td>
<td>23.2</td>
</tr>
<tr>
<td>Number of Transfers in Ideal Patients (&lt;=35 Years, &gt;=4 2PN, SET)</td>
<td>6,409</td>
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<tr>
<td>Life-Birth-Rate per Transfer in Ideal Patients (&lt;=35 Years, &gt;=4 2PN, SET)</td>
<td>1,698</td>
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<tr>
<td>Number of Multiple Births After SET</td>
<td>39</td>
<td>2.3</td>
</tr>
<tr>
<td>Number of Transfers in Ideal Patients (&lt;=35 Years, &gt;=4 2PN, DET/TET)</td>
<td>23,922</td>
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</tr>
<tr>
<td>Life-Birth-Rate per Transfer in Ideal Patients (&lt;=35 Years, &gt;=4 2PN, DET/TET)</td>
<td>7,093</td>
<td>29.7</td>
</tr>
<tr>
<td>Number of Multiple Births after DET/TET</td>
<td>1,422</td>
<td>20.0</td>
</tr>
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</table>

Loss of Pregnancy 2018
Prospective and Retrospective Data

<table>
<thead>
<tr>
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<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Clinical Pregnancies</td>
<td>17,226</td>
<td>100.0</td>
</tr>
<tr>
<td>Treatments with Known Cycle Outcome</td>
<td>16,341</td>
<td>94.9</td>
</tr>
<tr>
<td>Miscarriages</td>
<td>3,476</td>
<td>20.2</td>
</tr>
<tr>
<td>Among those:</td>
<td></td>
<td></td>
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<tr>
<td>Induced Abortions</td>
<td>171</td>
<td>4.9</td>
</tr>
<tr>
<td>Stilborn Children</td>
<td>169</td>
<td>1.0</td>
</tr>
</tbody>
</table>

Embryos per Transfer¹ and Children per Birth 1997 – 2018
IVF, ICSI – Prospective and Retrospective Data

<table>
<thead>
<tr>
<th></th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Fresh Cycles</td>
<td>1.90</td>
<td></td>
</tr>
<tr>
<td>Cryo Cycles</td>
<td>1.19</td>
<td></td>
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<tr>
<td>Fresh Cycles</td>
<td>1.88</td>
<td></td>
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<tr>
<td>Cryo Cycles</td>
<td>1.18</td>
<td></td>
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<tr>
<td>Fresh Cycles</td>
<td>1.77</td>
<td></td>
</tr>
<tr>
<td>Cryo Cycles</td>
<td>1.17</td>
<td></td>
</tr>
<tr>
<td>Fresh Cycles</td>
<td>1.66</td>
<td></td>
</tr>
<tr>
<td>Cryo Cycles</td>
<td>1.16</td>
<td></td>
</tr>
<tr>
<td>Fresh Cycles</td>
<td>1.55</td>
<td></td>
</tr>
<tr>
<td>Cryo Cycles</td>
<td>1.15</td>
<td></td>
</tr>
<tr>
<td>Fresh Cycles</td>
<td>1.43</td>
<td></td>
</tr>
<tr>
<td>Cryo Cycles</td>
<td>1.14</td>
<td></td>
</tr>
</tbody>
</table>

¹) Mean

The number of children originating from multiple pregnancies has almost remained stable within the last decades. In 2018, we finally 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!
### Culture According to the "German Middle Way" and Impact on Therapy Outcome – Fresh Cycles 2018

**Prospective and Retrospective Data**

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

| Oocyte Treatment (All) - Culture \( \geq 3 \times 2PN \) | 37,935 |
| Oocyte Treatment - Culture \( \geq 3 \times 2PN \) for Transfer on Day 4-6 | 22,871 |

#### Number of Transfer Cycles with SET Day 4-6

| CP/ET (doc.)* | 6,424 |
| Outcome unknown | 5 |

| Miscarriage | 365 | 15.9% |
| EP | 46 | 2.0% |
| Not yet Recorded | 163 | 7.1% |
| **Birth (doc.)** | 1,725 | 75.0% |
| per Transfer | | 26.9% |

<table>
<thead>
<tr>
<th>Births</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Singletons</td>
<td>1,688</td>
<td>97.9</td>
</tr>
<tr>
<td>Twins</td>
<td>37</td>
<td>2.1</td>
</tr>
<tr>
<td>Triplets</td>
<td>0</td>
<td>0.0</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
<td>0.0</td>
</tr>
</tbody>
</table>

#### Number of Transfer Cycles with DET Day 4-6

| CP/ET (doc.)* | 15,771 |
| Outcome unknown | 63 |

| Miscarriage | 1,140 | 17.5% |
| EP | 88 | 1.4% |
| Not yet Recorded | 481 | 7.4% |
| **Birth (doc.)** | 4,808 | 73.8% |
| per Transfer | | 30.5% |

<table>
<thead>
<tr>
<th>Births</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Singletons</td>
<td>3,444</td>
<td>71.6</td>
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<tr>
<td>Twins</td>
<td>1,334</td>
<td>27.7</td>
</tr>
<tr>
<td>Triplets</td>
<td>30</td>
<td>0.6</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
<td>0.0</td>
</tr>
</tbody>
</table>

*) please note: documented clinical pregnancies per transfer on day <4: 8,410 / 31,432 = 26.8%

### Comparison SET and DET "German Middle Way" Fresh Cycles 2018

![Comparison SET and DET "German Middle Way" Fresh Cycles 2018](image_url)
Culture According to the "German Middle Way" and Impact on Therapy Outcome – Thawing Cycles PNs 2018

Prospective and Retrospective Data

Number of centers thawing >=3 x 2PN for extended culture: n=75

<table>
<thead>
<tr>
<th>Culture &gt;=3 Vital 2PNs After Thawing (All Cycles)</th>
<th>7,898</th>
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</thead>
<tbody>
<tr>
<td>Number of Transfers on Day 2-3</td>
<td>2,621</td>
</tr>
<tr>
<td>Number of Transfer Cycles with SET Day 4-6</td>
<td>1,981</td>
</tr>
<tr>
<td>Culture &gt;=3 Vital 2PNs After Thawing</td>
<td>5,130</td>
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<tr>
<td></td>
<td>Outcome unknown</td>
</tr>
<tr>
<td>CP/ET (doc.)</td>
<td>666</td>
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<tr>
<td>Miscarriage</td>
<td>184</td>
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<tr>
<td>EP</td>
<td>6</td>
</tr>
<tr>
<td>Not yet Recorded</td>
<td>16</td>
</tr>
<tr>
<td>Birth (doc.)</td>
<td>460</td>
</tr>
<tr>
<td>per Transfer</td>
<td></td>
</tr>
<tr>
<td>Births</td>
<td></td>
</tr>
<tr>
<td>Singletons</td>
<td>442</td>
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<tr>
<td>Twins</td>
<td>15</td>
</tr>
<tr>
<td>Triplets</td>
<td>3</td>
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<tr>
<td>Quadruplets</td>
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<tr>
<td>Birth (doc.)/Transfer</td>
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<td>Miscarriage</td>
<td>279</td>
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<tr>
<td>EP</td>
<td>9</td>
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<tr>
<td>Not yet Recorded</td>
<td>45</td>
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<tr>
<td>Birth (doc.)</td>
<td>833</td>
</tr>
<tr>
<td>per Transfer</td>
<td></td>
</tr>
<tr>
<td>Births</td>
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</tr>
<tr>
<td>Singletons</td>
<td>629</td>
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<tr>
<td>Twins</td>
<td>198</td>
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<tr>
<td>Triplets</td>
<td>6</td>
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<tr>
<td>Quadruplets</td>
<td>0</td>
</tr>
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</table>

Comparison SET and DET "German Middle Way" Thawing Cycles PNs
Culture According to the "German Middle Way" and Impact on Therapy Outcome – Thawing Cycles Embryos 2018

Prospective and Retrospective Data

Number of centers transferring previously cryopreserved embryos: n=144

<table>
<thead>
<tr>
<th>Transfer Single ET</th>
<th>Transfer Double ET</th>
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<tbody>
<tr>
<td>CP/ET (doc.) 1,800</td>
<td>CP/ET (doc.) 960</td>
</tr>
<tr>
<td>Outcome unknown 25</td>
<td>Outcome unknown 11</td>
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<tr>
<td>Miscarriage 416</td>
<td>Miscarriage 247</td>
</tr>
<tr>
<td>EP 34</td>
<td>EP 11</td>
</tr>
<tr>
<td>Not yet Recorded 107</td>
<td>Not yet Recorded 24</td>
</tr>
<tr>
<td>Birth (doc.) 1,243 per Transfer</td>
<td>Birth (doc.) 678 per Transfer</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Births</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Singletons</td>
<td>1,219</td>
<td>98.1</td>
</tr>
<tr>
<td>Twins</td>
<td>24</td>
<td>1.9</td>
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<td>Triplets</td>
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<td>0.0</td>
</tr>
<tr>
<td>Quadruplets</td>
<td>0</td>
<td>0.0</td>
</tr>
</tbody>
</table>

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

![Comparison SET and DET "German Middle Way" Thawing Cycles Embryos](image-url)
Pregnancies Cumulative 2016 – 2019
IVF, ICSI, Cryo Cycles – Prospective Data

<table>
<thead>
<tr>
<th>Treatment started in 2016</th>
<th>Retrievals</th>
<th>Fresh Cycles with ET</th>
<th>CP (Fresh Cycles) in %</th>
<th>CP (Fresh Cycles w. Transfer)</th>
<th>CP (Cryo Cycles w. Transfer)</th>
<th>CP/ET (Cryo Cycles) in %</th>
<th>Cumulative Pregnancy Rate in %</th>
</tr>
</thead>
<tbody>
<tr>
<td>1st Treatment</td>
<td>30,718</td>
<td>29,721</td>
<td>30.6</td>
<td>7,678</td>
<td>2,353</td>
<td>30.6</td>
<td>30.6</td>
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<tr>
<td>2nd Treatment</td>
<td>13,554</td>
<td>13,028</td>
<td>27.2</td>
<td>5,377</td>
<td>1,496</td>
<td>27.2</td>
<td>50.4</td>
</tr>
<tr>
<td>3rd Treatment</td>
<td>8,700</td>
<td>8,330</td>
<td>25.9</td>
<td>3,484</td>
<td>291</td>
<td>25.9</td>
<td>62.7</td>
</tr>
<tr>
<td>4th Treatment</td>
<td>4,921</td>
<td>4,982</td>
<td>22.2</td>
<td>6,539</td>
<td>1,696</td>
<td>22.2</td>
<td>70.1</td>
</tr>
<tr>
<td>&gt;4 Treatment</td>
<td>8,194</td>
<td>8,189</td>
<td>22.2</td>
<td>6,539</td>
<td>1,696</td>
<td>22.2</td>
<td>81.9</td>
</tr>
</tbody>
</table>

| 1st Treatment Cycle | 33.3 % | 2nd Treatment Cycle | 63.0 % | 3rd Treatment Cycle | 46.2 % | 4th Treatment Cycle | 73.6 % | >4 Treatment Cycle | 80.8 % |

For one cycle, no plausible data could be correlated to the treatment cycle. This cycle did not result in a transfer.

For 244 cycles, no plausible data could be correlated to the treatment cycles. These cycles resulted in 178 additional transfers with 66 additional clinical pregnancies.

<table>
<thead>
<tr>
<th>Treatment started in 2017</th>
<th>Retrievals</th>
<th>Fresh Cycles with ET</th>
<th>CP (Fresh Cycles) in %</th>
<th>CP (Fresh Cycles w. Transfer)</th>
<th>CP (Cryo Cycles w. Transfer)</th>
<th>CP/ET (Cryo Cycles) in %</th>
<th>Cumulative Pregnancy Rate in %</th>
</tr>
</thead>
<tbody>
<tr>
<td>1st Treatment</td>
<td>24,812</td>
<td>21,422</td>
<td>35.0</td>
<td>6,273</td>
<td>1,925</td>
<td>35.0</td>
<td>30.6</td>
</tr>
<tr>
<td>2nd Treatment</td>
<td>11,784</td>
<td>9,182</td>
<td>33.2</td>
<td>6,273</td>
<td>1,925</td>
<td>30.6</td>
<td>44.0</td>
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<td>3rd Treatment</td>
<td>8,191</td>
<td>6,344</td>
<td>33.1</td>
<td>4,691</td>
<td>1,325</td>
<td>28.2</td>
<td>64.4</td>
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<tr>
<td>4th Treatment</td>
<td>5,917</td>
<td>4,580</td>
<td>33.2</td>
<td>3,277</td>
<td>915</td>
<td>27.9</td>
<td>70.2</td>
</tr>
<tr>
<td>&gt;4 Treatment</td>
<td>14,432</td>
<td>10,640</td>
<td>30.8</td>
<td>10,784</td>
<td>2,834</td>
<td>26.3</td>
<td>79.3</td>
</tr>
</tbody>
</table>

| 1st Treatment Cycle | 35.0 % | 2nd Treatment Cycle | 44.0 % | 3rd Treatment Cycle | 64.4 % | 4th Treatment Cycle | 70.2 % | >4 Treatment Cycle | 79.3 % |

For one cycle, no plausible data could be correlated to the treatment cycle. This cycle did not result in a transfer.

For 3 cycles, no plausible data could be correlated to the treatment cycles. These cycles resulted in 2 additional transfers with one additional clinical pregnancy.
Oocyte Maturity Depending on Stimulation Protocol 2019
Oocyte Maturity and Development Depending on Downregulation Protocol

ICSI – Prospective and Retrospective Data

<table>
<thead>
<tr>
<th></th>
<th>Cycles</th>
<th>Mean Age</th>
<th>Oocytes Retrieved</th>
<th>Ooc. per Retrieval</th>
<th>Mature Oocytes</th>
<th>Mature Oocytes per Retrieval</th>
<th>2PN</th>
<th>2PN/Oocyte Retrieval</th>
<th>Embryos for ET</th>
<th>Gestational Sacs</th>
<th>Implant. Rate (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Agonist</strong></td>
<td>11,261</td>
<td>35.7</td>
<td>97,950</td>
<td>8.7</td>
<td>80,802</td>
<td>7.2</td>
<td>49,924</td>
<td>4.4</td>
<td>15,472</td>
<td>3,386</td>
<td><strong>21.9</strong></td>
</tr>
<tr>
<td><strong>Antagonist</strong></td>
<td>46,907</td>
<td>35.6</td>
<td>424,450</td>
<td>9.0</td>
<td>354,981</td>
<td>7.6</td>
<td>220,731</td>
<td>4.7</td>
<td>58,933</td>
<td>13,400</td>
<td><strong>22.7</strong></td>
</tr>
</tbody>
</table>

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

Thawing – Prospective and Retrospective Data

<table>
<thead>
<tr>
<th></th>
<th>Freezing on Culture D1</th>
<th>Freezing on Culture D5/6</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Thawing-Cycles resulting in ET</td>
<td>6,682</td>
<td>8,829</td>
<td></td>
</tr>
<tr>
<td>PN/Embryos Thawed</td>
<td>17,864</td>
<td>9,984</td>
<td></td>
</tr>
<tr>
<td><strong>Entities/Thawing-Cycle</strong></td>
<td>2.7</td>
<td>1.1</td>
<td></td>
</tr>
<tr>
<td>Re-Cryo of Embryos</td>
<td>1,510</td>
<td>110</td>
<td></td>
</tr>
<tr>
<td>Embryos Available for Transfer</td>
<td>9,989</td>
<td>9,124</td>
<td></td>
</tr>
<tr>
<td>Re-Cryo/Cycle</td>
<td>0.2</td>
<td>0.0</td>
<td></td>
</tr>
<tr>
<td>Transfer</td>
<td>6,682</td>
<td>8,829</td>
<td></td>
</tr>
<tr>
<td>Gestational Sacs (Intrauterine)</td>
<td>2,627</td>
<td>3,139</td>
<td></td>
</tr>
<tr>
<td><strong>Embryos/ET</strong></td>
<td>1.5</td>
<td>1.0</td>
<td></td>
</tr>
<tr>
<td><strong>Implantation Rate (%)</strong></td>
<td>26.3</td>
<td>34.4</td>
<td></td>
</tr>
</tbody>
</table>
Implantation Rates of D3- and D5-Embryos and Number of Embryos per Transfer

IVF, ICSI – Prospective and Retrospective Data

<table>
<thead>
<tr>
<th></th>
<th>D3 Implantation Rate %</th>
<th>D5 Implantation Rate %</th>
</tr>
</thead>
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Evolution of Retrieved Oocytes (IVF or ICSI)

IVF, ICSI – Prospective and Retrospective Data

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<th>%</th>
<th>ICSI</th>
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<td>14,094</td>
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0 100,000 200,000 300,000 400,000

- IVF
- ICSI

Not Utilizable
Embryos in Cryo
2 PN in Cryo
ET (Fresh)
Clinical Pregnancies (CP)/Fresh Transfer as a Function of Embryo Quality 2019

**IVF, ICSI, IVF/ICSI – Prospective Data**

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<td>CP/ET %</td>
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Clinical Pregnancies (CP)/Frozen Transfer as a Function of Embryo Quality 2019

**Cryo Transfer – Prospective Data**

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<td>CP/ET %</td>
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*In 5,645 cases, resulting in an additional 1,337 pregnancies, no information has been available how many embryos had been transferred.
### Children as a Function of Week of Gestation (WoG) and Birth Weight (BW) 2018

**Prospective and Retrospective Data**

#### IVF, ICSI, IVF/ICSI

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<th>33 - 36</th>
<th>37 - 41</th>
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Percentage of preterm deliveries in singleton pregnancies is 19.7%.
Percentage of preterm deliveries in twin pregnancies is 83.9%.

#### Cryo Transfer

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<th>WoG</th>
<th>20 - 27</th>
<th>28 - 32</th>
<th>33 - 36</th>
<th>37 - 41</th>
<th>&gt;= 42</th>
<th>Total</th>
</tr>
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<tbody>
<tr>
<td><strong>Singletons</strong> (n and %)</td>
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Percentage of preterm deliveries in singleton pregnancies is 18.7%.
Percentage of preterm deliveries in twin pregnancies is 84.2%.
Children Born 1997 – 2018
Prospective and Retrospective Data

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

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<th>Triplets</th>
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Distribution of Indications 2019

IVF and ICSI – Prospective Data

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<th>IVF and ICSI – Prospective Data</th>
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</tr>
<tr>
<td>Homosexuality</td>
<td>345</td>
</tr>
<tr>
<td>Total</td>
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</table>

**) The indication “pathological function test” is included here

| Treatments | Transfers | Clin. Preg. | Shares of Indications (Cycles) 2019 |
| n | n | % | n | % | n | % |
| No Information | 100 | 67 | 67.0 | 21 | 31.3 |
| Idiopathic | 4,780 | 3,590 | 75.1 | 1,188 | 33.1 |
| Male | 25,692 | 22,923 | 89.2 | 7,490 | 32.7 |
| Female | 13,494 | 10,441 | 77.4 | 3,262 | 31.2 |
| Male and Female | 15,338 | 11,769 | 76.7 | 3,335 | 30.0 |
| Missing Male Partner | 261 | 212 | 81.2 | 61 | 28.8 |
| Homosexuality | 345 | 287 | 83.2 | 101 | 35.2 |
| Total | 60,010 | 49,289 | 82.1 | 15,658 | 31.8 |

<table>
<thead>
<tr>
<th>Shares of Indications (Cycles) 2019</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male and Female</td>
</tr>
<tr>
<td>All Others</td>
</tr>
<tr>
<td>Idiopathic</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Male Factor</th>
<th>Normal</th>
<th>Red. Semen Quality</th>
<th>Unknown</th>
<th>Other**</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Female Factor</strong></td>
<td><strong>n</strong></td>
<td><strong>%</strong></td>
<td><strong>n</strong></td>
<td><strong>%</strong></td>
<td><strong>n</strong></td>
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<tr>
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<td>92</td>
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<tr>
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<td>2,076</td>
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<td>517</td>
<td>3.2</td>
<td>70</td>
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<tr>
<td>Endometriosis</td>
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<td>6.8</td>
<td>283</td>
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<td>69</td>
</tr>
<tr>
<td>Hyperandrog./PCO</td>
<td>421</td>
<td>2.6</td>
<td>124</td>
<td>0.8</td>
<td>30</td>
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<tr>
<td>Ovulatory Dysf.</td>
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<td>252</td>
<td>1.5</td>
<td>65</td>
</tr>
<tr>
<td>Psychogen. Factors</td>
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<td>0.1</td>
<td>4</td>
<td>0.0</td>
<td>2</td>
</tr>
<tr>
<td>Age</td>
<td>338</td>
<td>2.1</td>
<td>132</td>
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<td>0</td>
</tr>
<tr>
<td>Other*</td>
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<td>6.7</td>
<td>959</td>
<td>5.8</td>
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<tr>
<td>No Information</td>
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<td>7</td>
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<tr>
<td>Total</td>
<td>7,293</td>
<td>44.4</td>
<td>2,893</td>
<td>17.6</td>
<td>346</td>
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</table>

<table>
<thead>
<tr>
<th>Shares of Indications (Cycles) 2019</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male and Female</td>
</tr>
<tr>
<td>All Others</td>
</tr>
<tr>
<td>Idiopathic</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Male Factor</th>
<th>Normal</th>
<th>Red. Semen Quality</th>
<th>Azoosperma</th>
<th>Unknown</th>
<th>Other**</th>
<th>Total</th>
</tr>
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<tbody>
<tr>
<td><strong>Female Factor</strong></td>
<td><strong>n</strong></td>
<td><strong>%</strong></td>
<td><strong>n</strong></td>
<td><strong>%</strong></td>
<td><strong>n</strong></td>
<td><strong>%</strong></td>
</tr>
<tr>
<td>Normal</td>
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<td>2.2</td>
<td>9,635</td>
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<td>1,131</td>
<td>2.6</td>
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<tr>
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<td>1,865</td>
<td>4.3</td>
<td>62</td>
<td>0.1</td>
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<td>Endometriosis</td>
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<td>1.0</td>
<td>1,767</td>
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<td>76</td>
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<td>Hyperandrog./PCO</td>
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<td>60</td>
<td>0.1</td>
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<tr>
<td>Ovulatory Dysf.</td>
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<td>1,721</td>
<td>3.9</td>
<td>131</td>
<td>0.3</td>
</tr>
<tr>
<td>Psychogen. Factors</td>
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<td>0.0</td>
<td>27</td>
<td>0.1</td>
<td>2</td>
<td>0.0</td>
</tr>
<tr>
<td>Age</td>
<td>237</td>
<td>0.5</td>
<td>1,080</td>
<td>2.5</td>
<td>134</td>
<td>0.3</td>
</tr>
<tr>
<td>Other*</td>
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<td>1.9</td>
<td>6,023</td>
<td>13.8</td>
<td>481</td>
<td>1.1</td>
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<tr>
<td>No Information</td>
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<td>71</td>
<td>0.2</td>
<td>29</td>
<td>0.1</td>
</tr>
<tr>
<td>Total</td>
<td>3,423</td>
<td>7.9</td>
<td>23,240</td>
<td>53.3</td>
<td>2,106</td>
<td>4.8</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Shares of Indications (Cycles) 2019</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male and Female</td>
</tr>
<tr>
<td>All Others</td>
</tr>
<tr>
<td>Idiopathic</td>
</tr>
</tbody>
</table>

*) The indications “sperm antibodies” and “cervical factor” are included here

**) The indication “pathological function test” is included here
Mean Age for Women and Men 1997 – 2019

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

<table>
<thead>
<tr>
<th>Year</th>
<th>Man</th>
<th>Woman</th>
</tr>
</thead>
<tbody>
<tr>
<td>1997</td>
<td>38.8</td>
<td>35.6</td>
</tr>
<tr>
<td>1998</td>
<td>38.4</td>
<td>35.2</td>
</tr>
<tr>
<td>1999</td>
<td>38.0</td>
<td>34.8</td>
</tr>
<tr>
<td>2000</td>
<td>37.6</td>
<td>34.4</td>
</tr>
<tr>
<td>2001</td>
<td>37.2</td>
<td>34.0</td>
</tr>
<tr>
<td>2002</td>
<td>36.8</td>
<td>33.6</td>
</tr>
<tr>
<td>2003</td>
<td>36.4</td>
<td>33.2</td>
</tr>
<tr>
<td>2004</td>
<td>36.0</td>
<td>32.8</td>
</tr>
<tr>
<td>2005</td>
<td>35.6</td>
<td>32.4</td>
</tr>
<tr>
<td>2006</td>
<td>35.2</td>
<td>32.0</td>
</tr>
<tr>
<td>2007</td>
<td>34.8</td>
<td></td>
</tr>
<tr>
<td>2008</td>
<td>34.4</td>
<td></td>
</tr>
<tr>
<td>2009</td>
<td>34.0</td>
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<td>2010</td>
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<td>2011</td>
<td>33.2</td>
<td></td>
</tr>
<tr>
<td>2012</td>
<td>32.8</td>
<td></td>
</tr>
<tr>
<td>2013</td>
<td>32.4</td>
<td></td>
</tr>
<tr>
<td>2014</td>
<td>32.0</td>
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</tr>
<tr>
<td>2015</td>
<td></td>
<td></td>
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<tr>
<td>2016</td>
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<td>2017</td>
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<tr>
<td>2018</td>
<td></td>
<td></td>
</tr>
<tr>
<td>2019</td>
<td>38.9</td>
<td>35.5</td>
</tr>
</tbody>
</table>

Pregnancy-Rate per ET Subject to Preexisting Medical- and Lifestyle-Conditions

Prospective and Retrospective Data

<table>
<thead>
<tr>
<th>Hypertension</th>
<th>Nicotine Abuse</th>
<th>BMI &lt;=25</th>
<th>BMI &gt;25 + &lt;30</th>
<th>BMI &gt;=30</th>
</tr>
</thead>
<tbody>
<tr>
<td>Number of Cycles with ET</td>
<td>459</td>
<td>4,748</td>
<td>54,973</td>
<td>10,592</td>
</tr>
<tr>
<td>CP (doc.)</td>
<td>131</td>
<td>1,491</td>
<td>16,970</td>
<td>3,371</td>
</tr>
<tr>
<td>CP (doc.) / ET</td>
<td>28.5</td>
<td>31.4</td>
<td>30.9</td>
<td>31.8</td>
</tr>
</tbody>
</table>

For the first time ever the data evaluation committee has attempted to evaluate anamnestic data and investigate possible relations of pre-existing medical conditions and lifestyle conditions with pregnancy rates per ET.

Due to the nature of the registers data collection, these results must be interpreted with caution.

In only 0.6% of treatment cycles resulting in an ET, hypertension was reported as a preexisting medical condition.

When BMI was automatically calculated from height and weight, nearly 10,000 cycles were performed with a BMI >30. In only 4,700 cycles however, “Adipositas per magne, BMI >30” was actively selected from the drop-down menu. In nearly 7,000 cycles, no data were reported regarding height and weight.
## Clinical Pregnancy Rate as a Function of Stimulation 2019

### Prospective Data

<table>
<thead>
<tr>
<th>Short GnRHa</th>
<th>uFSH</th>
<th>recFSH</th>
<th>hMG</th>
<th>recFSH a. recLH</th>
<th>recFSH a. hMG</th>
<th>Long-Acting recFSH</th>
<th>hrFSH</th>
<th>Other*</th>
<th>No Inform.</th>
<th>Total</th>
<th>Share (%) from total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stimulations (n)</td>
<td>42</td>
<td>2,747</td>
<td>245</td>
<td>2,877</td>
<td>874</td>
<td>218</td>
<td>100</td>
<td>315</td>
<td>176</td>
<td>7,594</td>
<td>12.9</td>
</tr>
<tr>
<td>Transfers (n)</td>
<td>25</td>
<td>2,157</td>
<td>172</td>
<td>2,231</td>
<td>697</td>
<td>167</td>
<td>77</td>
<td>199</td>
<td>91</td>
<td>5,816</td>
<td>13.1</td>
</tr>
<tr>
<td>Transfer (%)</td>
<td>59.5</td>
<td>78.5</td>
<td>70.2</td>
<td>77.5</td>
<td>79.7</td>
<td>76.6</td>
<td>77.0</td>
<td>63.2</td>
<td>51.7</td>
<td>76.6</td>
<td></td>
</tr>
<tr>
<td>CP (n)</td>
<td>9</td>
<td>720</td>
<td>42</td>
<td>740</td>
<td>237</td>
<td>47</td>
<td>33</td>
<td>39</td>
<td>20</td>
<td>1,887</td>
<td>13.2</td>
</tr>
<tr>
<td>CP/ET (%)</td>
<td>36.0</td>
<td>33.4</td>
<td>24.4</td>
<td>33.2</td>
<td>34.0</td>
<td>28.1</td>
<td>42.9</td>
<td>19.6</td>
<td>22.0</td>
<td>32.4</td>
<td></td>
</tr>
<tr>
<td>CP/Stim. (%)</td>
<td>21.4</td>
<td>26.2</td>
<td>17.1</td>
<td>25.7</td>
<td>27.1</td>
<td>21.6</td>
<td>33.0</td>
<td>12.4</td>
<td>11.4</td>
<td>24.8</td>
<td></td>
</tr>
</tbody>
</table>

Mean age of patients with regard to this protocol: 35.4 Jahre

<table>
<thead>
<tr>
<th>Long GnRHa</th>
<th>uFSH</th>
<th>recFSH</th>
<th>hMG</th>
<th>recFSH a. recLH</th>
<th>recFSH a. hMG</th>
<th>Long-Acting recFSH</th>
<th>hrFSH</th>
<th>Other*</th>
<th>No Inform.</th>
<th>Total</th>
<th>Share (%) from total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stimulations (n)</td>
<td>96</td>
<td>3,009</td>
<td>457</td>
<td>2,460</td>
<td>1,008</td>
<td>62</td>
<td>39</td>
<td>28</td>
<td>1,372</td>
<td>8,531</td>
<td>14.5</td>
</tr>
<tr>
<td>Transfers (n)</td>
<td>56</td>
<td>2,446</td>
<td>371</td>
<td>1,952</td>
<td>829</td>
<td>50</td>
<td>30</td>
<td>12</td>
<td>1,110</td>
<td>6,856</td>
<td>15.4</td>
</tr>
<tr>
<td>Transfer (%)</td>
<td>58.3</td>
<td>81.3</td>
<td>81.2</td>
<td>79.3</td>
<td>82.2</td>
<td>80.6</td>
<td>76.9</td>
<td>42.9</td>
<td>80.9</td>
<td>80.4</td>
<td></td>
</tr>
<tr>
<td>CP (n)</td>
<td>13</td>
<td>865</td>
<td>95</td>
<td>631</td>
<td>291</td>
<td>7</td>
<td>7</td>
<td>0</td>
<td>403</td>
<td>2,312</td>
<td>16.1</td>
</tr>
<tr>
<td>CP/ET (%)</td>
<td>23.2</td>
<td>35.4</td>
<td>25.6</td>
<td>32.3</td>
<td>35.1</td>
<td>14.0</td>
<td>23.3</td>
<td>0.0</td>
<td>36.3</td>
<td>33.7</td>
<td></td>
</tr>
<tr>
<td>CP/Stim. (%)</td>
<td>13.5</td>
<td>28.7</td>
<td>20.8</td>
<td>25.7</td>
<td>28.9</td>
<td>11.3</td>
<td>17.9</td>
<td>0.0</td>
<td>29.4</td>
<td>27.1</td>
<td></td>
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</tbody>
</table>

Mean age of patients with regard to this protocol: 34.9 Jahre

<table>
<thead>
<tr>
<th>No GnRHa-Analoga</th>
<th>uFSH</th>
<th>recFSH</th>
<th>hMG</th>
<th>recFSH a. recLH</th>
<th>recFSH a. hMG</th>
<th>Long-Acting recFSH</th>
<th>hrFSH</th>
<th>Other*</th>
<th>No Inform.</th>
<th>Total</th>
<th>Share (%) from total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stimulations (n)</td>
<td>20</td>
<td>1,277</td>
<td>933</td>
<td>1,044</td>
<td>358</td>
<td>208</td>
<td>28</td>
<td>6</td>
<td>57</td>
<td>3,931</td>
<td>6.7</td>
</tr>
<tr>
<td>Transfers (n)</td>
<td>16</td>
<td>1,000</td>
<td>729</td>
<td>844</td>
<td>282</td>
<td>155</td>
<td>20</td>
<td>3</td>
<td>33</td>
<td>3,082</td>
<td>6.9</td>
</tr>
<tr>
<td>Transfer (%)</td>
<td>80.0</td>
<td>78.3</td>
<td>78.1</td>
<td>80.8</td>
<td>78.8</td>
<td>74.5</td>
<td>71.4</td>
<td>50.0</td>
<td>57.9</td>
<td>78.4</td>
<td></td>
</tr>
<tr>
<td>CP (n)</td>
<td>3</td>
<td>318</td>
<td>222</td>
<td>230</td>
<td>68</td>
<td>50</td>
<td>7</td>
<td>0</td>
<td>6</td>
<td>904</td>
<td>6.3</td>
</tr>
<tr>
<td>CP/ET (%)</td>
<td>18.8</td>
<td>31.8</td>
<td>30.5</td>
<td>27.3</td>
<td>24.1</td>
<td>32.3</td>
<td>35.0</td>
<td>0.0</td>
<td>18.2</td>
<td>29.3</td>
<td></td>
</tr>
<tr>
<td>CP/Stim. (%)</td>
<td>15.0</td>
<td>24.9</td>
<td>23.8</td>
<td>22.0</td>
<td>19.0</td>
<td>24.0</td>
<td>25.0</td>
<td>0.0</td>
<td>10.5</td>
<td>23.0</td>
<td></td>
</tr>
</tbody>
</table>

Mean age of patients with regard to this protocol: 35.6 Jahre

<table>
<thead>
<tr>
<th>GnRHa-Antagonists</th>
<th>uFSH</th>
<th>recFSH</th>
<th>hMG</th>
<th>recFSH a. recLH</th>
<th>recFSH a. hMG</th>
<th>Long-Acting recFSH</th>
<th>hrFSH</th>
<th>Other*</th>
<th>No Inform.</th>
<th>Total</th>
<th>Share (%) from total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Stimulations (n)</td>
<td>411</td>
<td>18,116</td>
<td>1,159</td>
<td>9,706</td>
<td>3,925</td>
<td>1,944</td>
<td>440</td>
<td>1,563</td>
<td>1,392</td>
<td>38,656</td>
<td>65.8</td>
</tr>
<tr>
<td>Transfers (n)</td>
<td>299</td>
<td>14,095</td>
<td>795</td>
<td>7,093</td>
<td>2,977</td>
<td>1,460</td>
<td>308</td>
<td>920</td>
<td>748</td>
<td>28,695</td>
<td>64.6</td>
</tr>
<tr>
<td>Transfer (%)</td>
<td>72.7</td>
<td>77.8</td>
<td>68.6</td>
<td>73.1</td>
<td>75.8</td>
<td>75.1</td>
<td>70.0</td>
<td>58.9</td>
<td>53.7</td>
<td>74.2</td>
<td></td>
</tr>
<tr>
<td>CP (n)</td>
<td>88</td>
<td>4,793</td>
<td>195</td>
<td>2,270</td>
<td>975</td>
<td>383</td>
<td>121</td>
<td>175</td>
<td>227</td>
<td>9,227</td>
<td>64.4</td>
</tr>
<tr>
<td>CP/ET (%)</td>
<td>29.4</td>
<td>34.0</td>
<td>24.5</td>
<td>32.0</td>
<td>32.8</td>
<td>26.2</td>
<td>39.3</td>
<td>19.0</td>
<td>30.3</td>
<td>32.2</td>
<td></td>
</tr>
<tr>
<td>CP/Stim. (%)</td>
<td>21.4</td>
<td>26.5</td>
<td>16.8</td>
<td>23.4</td>
<td>24.8</td>
<td>19.7</td>
<td>27.5</td>
<td>11.2</td>
<td>16.3</td>
<td>23.9</td>
<td></td>
</tr>
</tbody>
</table>

Mean age of patients with regard to this protocol: 35.3 Jahre

**Mean age of patients with regard to all protocols:**

|       | 37.9 | 33.9 | 37.5 | 35.6 | 36.3 | 36.7 | 34.6 | 38.7 | 35.9 | 35.2 |

In 9,451 cycles, either the protocol or the type of gonadotrophin were not clearly reported. These cycles contain a high proportion of hMG, whereby the number of hMG-stimulations is falsely reported too low. This bug is being analyzed, as well as the conspicuously high number of cycles with GnRH-short-protocol.

*) e.g. u-FSH and hMG, Clomifen/rec-FSH, Clomifen/hMG etc.
### Ovarian Hyperstimulation Syndrome (OHSS) as a Function of Stimulation Protocol and Age Cohort 2019

**IVF, ICSI, IVF/ICSI – Prospective Data**

<table>
<thead>
<tr>
<th>Protocol</th>
<th>Cycles</th>
<th>%</th>
<th>Oocytes Retrieved</th>
<th>OHSS III (WHO)</th>
<th>OHSS III/Cycles %</th>
</tr>
</thead>
<tbody>
<tr>
<td>Short GnRHa</td>
<td>7,594</td>
<td>12.9</td>
<td>8.2</td>
<td>26</td>
<td>0.3</td>
</tr>
<tr>
<td>&lt; = 29 Years</td>
<td>867</td>
<td></td>
<td>11.1</td>
<td>2</td>
<td>0.2</td>
</tr>
<tr>
<td>30 – 34 Years</td>
<td>2,365</td>
<td></td>
<td>9.4</td>
<td>9</td>
<td>0.4</td>
</tr>
<tr>
<td>35 – 39 Years</td>
<td>3,278</td>
<td></td>
<td>7.3</td>
<td>11</td>
<td>0.3</td>
</tr>
<tr>
<td>&gt; = 40 Years</td>
<td>1,084</td>
<td></td>
<td>6.4</td>
<td>4</td>
<td>0.4</td>
</tr>
</tbody>
</table>

| Long GnRHa        | 8,531  | 14.5| 9.3              | 29             | 0.3              |
| < = 29 Years      | 1,084  |     | 11.4             | 3              | 0.3              |
| 30 – 34 Years     | 2,811  |     | 10.6             | 11             | 0.4              |
| 35 – 39 Years     | 3,600  |     | 8.4              | 12             | 0.3              |
| > = 40 Years      | 1,036  |     | 6.5              | 3              | 0.3              |

| No GnRHa-Analoga  | 3,931  | 6.7 | 9.3              | 13             | 0.3              |
| < = 29 Years      | 405    |     | 11.7             | 1              | 0.2              |
| 30 – 34 Years     | 1,144  |     | 10.3             | 4              | 0.3              |
| 35 – 39 Years     | 1,697  |     | 8.6              | 6              | 0.4              |
| > = 40 Years      | 685    |     | 6.3              | 2              | 0.3              |

| GnRHa-Antagonisten| 38,656 | 65.8| 9.2              | 133            | 0.3              |
| < = 29 Years      | 4,762  |     | 11.9             | 11             | 0.2              |
| 30 – 34 Years     | 12,481 |     | 10.7             | 48             | 0.4              |
| 35 – 39 Years     | 16,265 |     | 8.3              | 55             | 0.3              |
| > = 40 Years      | 5,148  |     | 5.7              | 19             | 0.4              |

| Total*            | 58,712 | 100.0| 9.1            | 201            | 0.3              |

*) in 9,451 cycles, the protocol could not be reliably determined

### Complications as a Function of Ovum Pick-up (OPU) 2019

**Prospective Data**

<table>
<thead>
<tr>
<th>Total OPU's</th>
<th>62,192</th>
<th>100.0%</th>
</tr>
</thead>
<tbody>
<tr>
<td>No Information</td>
<td>0</td>
<td>0.0%</td>
</tr>
<tr>
<td>No Complications</td>
<td>61,671</td>
<td>99.2%</td>
</tr>
<tr>
<td>Complications</td>
<td>521</td>
<td>0.8%</td>
</tr>
</tbody>
</table>

#### Complications

<table>
<thead>
<tr>
<th>Complications</th>
<th>n</th>
<th>%</th>
</tr>
</thead>
<tbody>
<tr>
<td>Vaginal Bleeding</td>
<td>328</td>
<td>63.0</td>
</tr>
<tr>
<td>Intrabdom. Bleeding</td>
<td>75</td>
<td>14.4</td>
</tr>
<tr>
<td>Intestinal Tract Injury</td>
<td>1</td>
<td>0.2</td>
</tr>
<tr>
<td>Peritonitis</td>
<td>24</td>
<td>4.6</td>
</tr>
<tr>
<td>Hospitalisation</td>
<td>7</td>
<td>1.3</td>
</tr>
<tr>
<td>Surgery</td>
<td>18</td>
<td>3.5</td>
</tr>
<tr>
<td>Other</td>
<td>68</td>
<td>13.1</td>
</tr>
</tbody>
</table>

Total 521 100.0%
For the second year in a row, the 2006 established network FertiPROTEKT e.V. – a consortium of centers from Germany, Austria and Switzerland, that are engaged in fertility preservation in female patients before gonadotoxic therapies – is able to present its yearly registry data within the D-I-R yearbook.

This year, however, the focus is set on the last four years, showing the data from 2016 up to 2019. Since information on ovarian tissue transplantation was scarce in the year 2019, we omit to publish about them this year, since they would not be representative.

Overall, 2019 we registered less counselling visits than in 2018. However, as more interventions are registered, we assume that the number of counselling visits was still high, just not documented.

The distribution of the individual treatments remained the same within the last years.

Last year the number of children <15 years of age counselled increased dramatically. This is not due to the activity of a single center, but rather due to counselling in several university clinics. Furthermore, the reason for the reduced number of patients counselled between 36-40 years remains speculative.
Diagnoses at Consultation

While in former days mainly patients with breast cancer and lymphoma were seen, the incidence of patients with other cancers increased in 2019. We hope that this trend will be kept up, when fertility preservation will be covered by the insurance companies in Germany and Switzerland in the near future.

The ratio of malignant to benign diseases remains stable.

Frequent Diagnosis Groups in Register 2016 – 2019

Underlying Disease when Ovarian Stimulation is Performed

Logarithmic Scale

Underlying Disease when Ovarian Tissue Cryopreservation is Performed

Logarithmic Scale
This year an increased number of GnRHa prescriptions can be noted, most likely due to the more recent articles published in favor of their use.
The entry of the FertiPROTEKT-ID has picked up in 2019, even before it has become a mandatory field for the documentation of ovarian stimulation cycles, and is expected to increase further in the upcoming years. This allows us to measure the quality of the treatment on one hand and to link the underlying indication of the fertility preservation on the other hand.

It is currently assumed that still not all ovarian stimulations are documented within this registry, as the numbers documented differ between the two registries. Furthermore, only two cycles of cryo-embryo-transfers are noted, which is below our expectations.

A goal for the upcoming year will be the definition of indications for recording a cycle at the D-I-R registry as well as to simplify the submission and revision of data within the FertiPROTEKT registry. In addition, an automated creation of a patient document with a FertiPROTEKT-ID is planned, in order to facilitate the communication between the centers regarding one and the same patient.

Therefore, we already integrated the data from the former registries from the years before 2014 into the existing registry. All of them were updated with a FertiPROTEKT-ID, allowing to add this number in the D-I-R registry, when a cyro-embryo-transfer is performed nowadays.
Deutsches IVF-Register e.V. (D·I·R)®
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