## Supplementary File 2. Overview of key MitraClip™ studies

### RANDOMIZED CONTROLLED TRIALS (RCTs)

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<th>Study Name</th>
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| **EVEREST II**   | (completed) | • 279 patients with significant (≥3+) MR (PMR or SMR) who were suitable candidates for surgery  
• MitraClip™ versus conventional surgery  
• USA and Canada  
• Enrolment: 2005–2008 |
| **COAPT**        | (completed) | • 614 HF patients with SMR (≥3+) unsuitable for mitral valve surgery  
• MitraClip™ + GDMT versus GDMT alone  
• 78 sites in USA  
• Enrolment: 2012–2017 (follow-up ongoing) |
| **MITRA-FR**     | (completed) | • 304 patients with severe SMR  
• MitraClip™ + GDMT versus GDMT alone  
• 37 sites in France  
• Enrolment: 2013–2017 |
| **RESHAPE-HF2**  | (ongoing)   | • 650 patients (including 40 from RESHAPE-HF) with SMR  
• MitraClip™ + GDMT +/-CRT versus GDMT +/-CRT alone  
• CZ, DE, DK, ES, GR, IT, PL, PT, UK  
• 2015–ongoing |
| **MATTERHORN**   | (ongoing)   | • 210 patients with PMR or SMR with left ventricle dysfunction  
• MitraClip™ versus surgery  
• Germany, Switzerland  
• 2015–ongoing |
| **MITRA-HR**     | (ongoing)   | • 330 patients with severe PMR  
• MitraClip™ versus conventional surgery  
• France  
• 2018–ongoing |
| **REPAIR MR**    | (ongoing)   | • ~500 moderate-surgical-risk patients with severe PMR candidates for open heart surgery.  
• MitraClip™ versus open heart mitral surgical repair.  
• 60 sites in the USA, Canada, and Europe  
• Enrolment: 2020–ongoing |

### REGISTRIES

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| **EVEREST II HRS** | (completed) | • Prospective, multicentre study which assessed the efficacy and safety of MitraClip™ in 78 patients with significant (≥3+) MR (PMR or SMR) deemed high risk surgical candidates.  
• USA  
• 2007–2008 |
| **ACCESS-EU**    | (completed) | • Two-phase, prospective, observational study of 567 patients with PMR or SMR who received MitraClip™  
• 14 sites in CH, DE, DK, IT  
• 2008–2011 |
| **EVEREST II REALISM** | (completed) | • Prospective, continued-access study to investigate the real-world use of MitraClip™ in low versus high surgical risk patients (N=965) with PMR or SMR  
• USA  
• 2009–2011 |
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| GRASP                       | • Enrolled 117 patients with grade ≥3+ MR (PMR or SMR) deemed high risk for conventional surgery who received MitraClip™  
                                • Single centre in Italy.  
                                • 2008–2012                                                                  |
| MULTI-CENTRE EXPERIENCE     | • Registry enrolling 173 patients with PMR or SMR treated with MitraClip™  
                                • Three centres in Denmark, Sweden, and UK  
                                • 2009–2012                                                                    |
| FRENCH REGISTRY             | • A study of short- and mid-term efficacy and safety of MitraClip™ in inoperable/high surgical risk patients (N=62) with PMR or SMR  
                                • Seven centres in France  
                                • 2010–2012                                                                    |
| EUROPEAN SENTINEL           | • A real-world overview of 628 patients with PMR or SMR who received MitraClip™  
                                • BE, CH, DE, DK, IT, PL, SE, UK  
                                • 2011–2012                                                                    |
| TRAMI                       | • A study of MitraClip™ efficacy and safety in daily clinical practice based on 1,064 patients with PMR or SMR and cardiac comorbidities  
                                • 21 sites in Germany  
                                • 2010–2013                                                                    |
| EXPAND                      | • A contemporary, prospective study evaluating real-world experience of performance and safety of MitraClip™ (NTR, XTR) in 1,000 patients with PMR, SMR, or mixed MR  
                                • CH, DE, ES, IL, IT, NL, UK, and USA  
                                • 2018–2019                                                                    |
| TVT                         | • Registry enrolling all patients (including those with PMR or SMR; N=12,334 as of September 2017) that have undergone transcatheter valve replacement and repair procedures for the monitoring of safety and real-world outcomes  
                                • Multiple centres throughout the US  
                                • 2011–ongoing                                                                 |
| MITRA SWISS                 | • Prospective, nationwide investigator initiated MitraClip™ registry that continuously enrolls patients with either PMR or SMR undergoing MitraClip™ procedure (N=1,212 as of August 2019)  
                                • 2011–ongoing                                                                 |
| GIOTTO                      | • Prospective, observational patient registry which will enrol up to 1,500 MitraClip™ patients with PMR or SMR for follow-up to 5 years  
                                • Italy  
                                • 2016–ongoing                                                                 |
| EXPAND G4                   | • Prospective patient registry to assess the safety and performance of the fourth generation MitraClip™ device in 100 MitraClip™ G4 recipients with either PMR, SMR, or mixed MR  
                                • 17 sites in the US  
                                • 2020–ongoing                                                                 |