

Supplementary Table 3. Evidence Table of Primary Observational Studies

Study	Design	Patients (prior anti-TNF exposure/n)	Interventions (n)	Comparators (n)	Primary outcomes
Feagan (2019) <sup>22</sup>	Prospective observational Multinational	Data from GEMINI 1: clinical remission by week 14 (329/769)	Maintenance phase of GEMINI 1 (620) Week 6 responders to VDZ: either q 4 weeks or 8 weeks Week 6 non-responders to VDZ: q 4 weeks	Placebo (149)	<ul style="list-style-type: none"> <li>Weeks 14 to 52 sustained clinical remission: VDZ vs. placebo (66.5% vs. 26.7%; <math>\Delta</math>39.8%; 95% CI, 22.7–56.9)</li> <li>Prior anti-TNF failure: VDZ vs. placebo (62.3% vs. 40.0%; <math>\Delta</math>22.3%; 95% CI, -10.8 to 55.6<sup>a</sup>)</li> </ul>
Adedokun (2020) <sup>23</sup>	Prospective observational Multinational	Moderate-to-severe UC (117/348)	Maintenance: UST 90 mg SC q 8 weeks (176)	Maintenance: UST 90 mg SC q 12 weeks (172)	<ul style="list-style-type: none"> <li>Median trough serum UST concentrations: 90 mg SC q 8 weeks vs. SC q 12 weeks (2.7–3.1 <math>\mu</math>g/mL vs. 0.9–1.2 <math>\mu</math>g/mL)</li> </ul>
Xu (2020) <sup>24</sup>	Prospective observational Multinational	Data from UNIFI (431/823)	UST 130 mg IV, then 90 mg SC q 8 weeks (320)	UST 6 mg/kg IV, then 90 mg SC q 12 weeks (319)	<ul style="list-style-type: none"> <li>Median steady-state trough concentrations: UST 90 mg SC q 12 weeks vs. SC q 8 weeks (0.76 <math>\mu</math>g/mL vs. 2.3 <math>\mu</math>g/mL)</li> </ul>
Hu (2020) <sup>25</sup>	Case-control Multinational	Moderate-to-severe UC (345/363)	UST combination with AZA (120)	UST monotherapy (243)	<ul style="list-style-type: none"> <li>Week 14 clinical response: combination therapy vs. monotherapy (54.6% vs. 65.8%; <math>P=0.08</math>)</li> <li>Week 30 clinical response: combination therapy vs. monotherapy (71.6% vs. 77.4%; <math>P=0.33</math>)</li> <li>Week 54 clinical response: combination therapy vs. monotherapy (62.1% vs. 67.0%; <math>P=0.52</math>)</li> </ul>
Sands (2020) <sup>26</sup>	Prospective Observational Multinational	Data from OCTAVE Open (55/123)	TFC dose de-escalation: 66 TFC dose escalation: 57	NA	<ul style="list-style-type: none"> <li>After TFC de-escalation: 92.4% and 84.1% of patients maintained clinical response and 80.3% and 74.6% maintained remission, at months 2 and 12, respectively.</li> <li>After dose escalation, 57.9% and 64.9% of patients recaptured clinical response and 35.1% and 49.1% were in remission, at months 2 and 12, respectively.</li> </ul>
Honap (2020) <sup>27</sup>	Retrospective observational UK	Moderate-to-severe UC (59/134)	TFC 10 mg b.i.d. at least 8 weeks, followed by 5 mg b.i.d.	NA	<ul style="list-style-type: none"> <li>Week 8 clinical response: 73.9%</li> <li>Week 22 steroid-free remission: 43.5%</li> </ul>
Colombel (2020) <sup>28</sup>	Prospective observational Multinational	Data from OCTAVE Open (60/142)	TFC 5 mg b.i.d. during OCTAVE Open, following a 52-week OCTAVE Sustain	NA	<ul style="list-style-type: none"> <li>12-month clinical remission, endoscopic improvement, and clinical response: 68.3%, 73.9%, and 77.5%, respectively.</li> <li>36-month clinical remission, endoscopic improvement, and clinical response: 50.4%, 55.3%, and 56.0%, respectively.</li> </ul>
Goll (2019) <sup>29</sup>	Prospective observational Norway	NOR-SWITCH extension (380)	Switch to CT-P13 at week 52 through 78 weeks (183; UC 38)	Maintenance CT-P13 (197; UC 42)	<ul style="list-style-type: none"> <li>Disease worsening: maintenance vs. switch (16.8% vs. 11.6%; <math>\Delta</math>5.9%; 95% CI, -1.1 to 12.9)</li> </ul>
Gustavsson (2010) <sup>30</sup>	Retrospective observational Sweden and Denmark	Acute severe UC (0/45)	IFX 5 mg/kg single dose (24)	Placebo (21)	<ul style="list-style-type: none"> <li>3-year colectomy rate: IFX vs. placebo (50.0% vs. 76.2%; <math>P=0.012</math>)</li> <li><b>Colectomy rate in patients with endoscopic remission vs. not in remission at 3-month (0% vs 50.0%; <math>P=0.02</math>)</b></li> </ul>
Nalagatla (2019) <sup>31</sup>	Case-control USA	Acute severe UC (0/213)	Accelerated dose: IFX > 5 mg/kg infliximab at shorter intervals through 24 weeks (81)	Standard dose: IFX 5 mg/kg at weeks 0, 2, and 6 (132)	<ul style="list-style-type: none"> <li>In-hospital colectomy rate: accelerated vs. standard IFX (9% vs. 8%; adjusted OR, 1.35; 95% CI, 0.38–4.82)</li> <li>3-, 6-, 12-, or 24-month colectomy rates: no significant difference (<math>P&gt;0.20</math> for all comparisons)</li> </ul>

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Supplementary Table 3. Continued

Study	Design	Patients (prior anti-TNF exposure/n)	Interventions (n)	Comparators (n)	Primary outcomes
Sebastian (2019) <sup>32</sup>	Case-control propensity score-matched UK	Acute severe UC (0/131)	Accelerated IFX induction (29)	Standard IFX induction (102)	<ul style="list-style-type: none"> <li>• 30-day colectomy rates: accelerated induction vs. standard induction (27% vs. 57%; <math>P=0.048</math>)</li> <li>• Overall colectomy rates: accelerated induction vs. standard induction (31% vs. 57; <math>P=0.09</math>)</li> </ul>
Shah (2018) <sup>33</sup>	Case-control propensity score-matched USA	Acute severe UC (0/146)	High dose IFX: 10 mg/kg (26)	Standard dose IFX: 5 mg/kg (120)	<ul style="list-style-type: none"> <li>• 30-day colectomy rates: standard dose vs. high dose (17.5% vs. 15.4%; <math>P=0.53</math>)</li> </ul>
Gibson (2020) <sup>34</sup>	Case-control Ireland	Acute severe UC (0/145)	Accelerated IFX dose (58)	Standard IFX dose cohort 1 (year 2010-2013) (87) Standard IFX dose cohort 2 (year 2014-2017) (55)	<ul style="list-style-type: none"> <li>• Time to colectomy compares with accelerated dose: standard dose cohort 1, shorter time (log rank <math>P=0.0013</math>); standard dose cohort 2, no significant difference (log rank <math>P=0.32</math>)</li> </ul>
Lau (2015) <sup>35</sup>	Retrospective observational USA	IBD (217; UC 94)	Serum anti-TNF drug level detectable (UC 17)	Serum anti-TNF drug level undetectable (UC 77)	<ul style="list-style-type: none"> <li>• Infectious complications: detectable vs. undetectable (9.3% vs. 11.8%; <math>P=0.78</math>)</li> <li>• Overall postoperative morbidity: detectable vs. undetectable (39.5% vs. 47.1%; <math>P=0.59</math>)</li> </ul>
Zittan (2016) <sup>36</sup>	Retrospective observational Canada	IPAA for UC (758)	Preoperative anti-TNF exposure (196)	Anti-TNF naïve (596)	<ul style="list-style-type: none"> <li>• Postoperative IPAA leak rate: anti-TNF exposure vs. naïve (13.2% vs. 11.7%; <math>P=0.44</math>)</li> </ul>
Abelson (2018) <sup>37</sup>	Retrospective observational USA	Surgery for UC (7,070)	Surgery between year 2006-2013 (3,267)	Surgery between year 1995-2005 (3,803)	<ul style="list-style-type: none"> <li>• Major event: after vs. before the year 2005 (5.3% vs. 7.4%; OR, 1.42; 95% CI, 1.13-1.78)</li> <li>• Procedural complications; after vs. before the year 2005 (9.9% vs. 12.3%; OR, 1.42; 95% CI, 1.20-1.68)</li> </ul>
Ward (2018) <sup>38</sup>	Retrospective observational UK	Subtotal colectomy for UC (6,225)	Anti-TNF exposure within 12 weeks (753) or within 4 weeks (418) prior to surgery	Anti-TNF naïve (5,472)	<ul style="list-style-type: none"> <li>• Postoperative complications: anti TNF exposure within 12 weeks or within 4 weeks vs. naïve (12.4% or 12.9% vs. 12.0%; <math>P=0.696</math> and <math>P=0.571</math>, respectively)</li> </ul>
El-Hussuna (2018) <sup>39</sup>	Prospective observational Denmark	Major abdominal surgery for IBD (46; UC 14)	Anti-TNF exposure (18; UC 5)	Anti-TNF naïve (28; UC 9)	<ul style="list-style-type: none"> <li>• Postoperative complications: anti-TNF exposure vs. naïve (27.8% vs. 28.6%; <math>P&gt;0.05</math>)</li> </ul>
Novello (2020) <sup>40</sup>	Prospective observational case-matched USA	Abdominal surgery for IBD (980)	VDZ exposure (95; UC 59)	VDZ naïve (95; UC 59)	<ul style="list-style-type: none"> <li>• Overall morbidity: VDZ vs. naïve (54.7% vs. 47.4%; OR, 1.3; 95% CI, 0.76-2.3; <math>P=0.32</math>)</li> <li>• Any infection; VDZ vs. naïve (12.6% vs. 20.0%; OR, 0.53; 95% CI, 0.23-1.3; <math>P=0.15</math>)</li> </ul>

<sup>a</sup>Prior anti-TNF exposure or failure.

TNF, tumor necrosis factor; VDZ, vedolizumab; CI, confidence interval; UC, ulcerative colitis; UST, ustekinumab; SC, subcutaneous; IV, intravenous; AZA, azathioprine; TFC, tofacitinib; IFX, infliximab; OR, odds ratio; IBD, inflammatory bowel disease; IPAA, ileal pouch-anal anastomosis; NA, not available; q, every day; b.i.d., twice a day.