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Short communication: Economic impact among 7 reproductive programs for lactating dairy cows, including a sensitivity analysis of the cost of hormonal treatments

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(Article begins on next page)

1 **Short Communication: The reproductive and economic impact among 6 reproductive**
2 **programs for lactating dairy cows including a sensitivity analysis of the cost of hormonal**
3 **treatments**

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7

8

Abstract

9 Hormonal synchronization protocols can dramatically improve the reproductive
10 efficiency of dairy herds, yet some farmers continue to question the economics of these
11 programs based on the cost of hormonal treatments, and hormonal treatment costs vary
12 dramatically among countries. Our objective was to compare the economic impact of
13 reproductive management programs that incorporate varying degrees of detection of estrus and
14 timed AI. A reproductive economic analysis simulation model (the UW-Cornell
15 DairyRepro\$ decision tool) was used to compare the economic impact of pairs of reproductive
16 management programs. We simulated sets of scenarios for 2 analyses. In the first analysis, we
17 calculated the economic impact of switching from a Presynch-Ovsynch program to a Double-
18 Ovsynch program that included a second PGF_{2α} treatment during the Breeding-Ovsynch
19 portion of the protocol (**Double-Ovsynch+PGF**). In the second analysis, we conducted a
20 breakeven analysis in which we incrementally increased the cost of hormonal treatments within
21 various reproductive management programs. Our analyses revealed that a Double-
22 Ovsynch+PGF protocol, the most intensive program evaluated, was more profitable than other
23 programs including a Presynch-Ovsynch protocol with 100% timed AI or a Presynch-Ovsynch

24 protocol that incorporated detection of estrus, despite the higher up-front cost incurred by using
25 more hormonal treatments. This advantage remained until the cost of hormones were 5 to 14
26 times more than the current US market prices and 2 to 6 times greater than the current
27 European market prices. The cost of GnRH had a greater impact on the net profit gain than the
28 cost of PGF_{2α}.

29 Keywords: reproduction, intensive synchronization program, economic impact

30 **Short Communication**

31 Advances in the understanding of the reproductive physiology of dairy cows have lead
32 to the development of management strategies and technologies that improve reproductive
33 performance. Commercial dairy farms are challenged with making the most profitable
34 management decisions among many options and implementing them correctly. Methods for
35 enhancing fertility and breeding efficiency include: detection of estrus (**ED**) (Xu et al., 1998;
36 Rorie et al., 2002), synchronization of estrus (Folman et al., 1984; Momcilovic et al., 1998),
37 and synchronization of ovulation and timed artificial insemination (**TAI**) (Pursley et al., 1995;
38 Moreira et al., 2001; Souza et al., 2008). The newest TAI protocols for first AI (i.e., Double-
39 Ovsynch and G6G), not only increase the AI service rate, but also increase P/AI to TAI
40 (Carvalho et al., 2018). Strategies that maximize ED, which is widely implemented on dairies,
41 have made significant contributions to the profitability of dairy herds (Pecsok et al. 1994). A
42 major limitation of ED, however, is the presence of anovular cows. The proportion of cows that
43 have not re-initiated cyclicity by the end of the voluntary waiting period varies among herds
44 and among parities within a herd and ranges from 5% to 40% (Walsh et al., 2007; Santos et al.,
45 2009; Bamber et al., 2009). The lack of estrous behavior in anovular cows precludes AI to a
46 detected estrus, and many anovular cows are submitted to hormonal protocols for TAI.

47 Protocols for synchronization of ovulation use sequential treatments of GnRH and
48 PGF_{2α} to control follicular development, luteal regression, and time of ovulation.
49 Synchronization of ovulation allows for more precise timing of AI than relying on detection of
50 estrus alone for the timing of AI (Pursley et al., 1995, Souza et al. 2008; Valenza et al., 2012;
51 Fricke et al., 2014). Furthermore, optimization of the hormonal milieu during the Ovsynch
52 protocol increases P/AI for cows at first TAI (Souza et al., 2008; Carvalho et al., 2014) and for
53 cows resynchronized to receive subsequent TAI (Giordano et al., 2012; Lopes et al., 2013;
54 Carvalho et al., 2015). Many farms combine ED and TAI by observing and inseminating cows
55 detected in estrus after the second PGF_{2α} treatment of a Presynch-Ovsynch protocol
56 (Stangaferro et al. 2018 and 2019), whereas cows not detected in estrus complete the protocol
57 and receive TAI. Although incorporation of estrus into a Presynch-Ovsynch protocol increases
58 AI service rate, it also decreases P/AI by 35% compared to 100% TAI after a Presynch-
59 Ovsynch protocol (Borchardt et al., 2016). By contrast, submission of lactating Holstein cows
60 to a Double-Ovsynch+PGF protocol and TAI for first insemination increases the percentage of
61 cows inseminated within 7 d after the end of the voluntary waiting period and increases P/AI at
62 33 and 63 d after first insemination resulting in 64 and 58% more pregnant cows, respectively,
63 than submission of cows for first AI after detection of estrus at a similar day in milk range
64 (Santos et al., 2017).

65 The assessment of the overall economic value of different reproductive management
66 programs (Cabrera and Giordano, 2013; Cabrera, 2014) can be achieved by simulating
67 reproductive performance along with its costs and benefits on a farm-by-farm basis (Giordano
68 et al., 2011; 2012; Kalantari and Cabrera, 2012), and calculating the expected net return (De
69 Vries et al., 2010; Fricke et al., 2010; Cabrera, 2011). In this study, we used a simulation model

70 to compare the economic impact of current and alternative reproductive programs (the UW-
71 Cornell DairyRepro\$ decision tool; Giordano et al., 2012). This study had three major
72 objectives: 1) to analyze the economic profitability of a more intensive reproductive protocol
73 involving more hormonal treatments; 2) to determine if increased hormonal treatment costs
74 would be compensated for by increased production of calves and increased P/AI; and 3) to
75 estimate how high the cost of hormones would have to be to render intensive synchronization
76 protocols that require more hormonal treatments non-profitable. To answer to these questions,
77 a 1,000-cow commercial dairy herd was simulated using the UW-Cornell
78 DairyRepro\$ decision tool. For comparison consistency and to avoid analysis bias, non-studied
79 variables such as mortality rate, body weight, involuntary culling rate, lactation curves, milk
80 price, among others, were kept constant among scenarios at default levels described by
81 Giordano et al. (2012).

82 Six reproductive management programs for first TAI were simulated. The first program
83 simulated was a PreSynch-Ovsynch protocol with ED incorporated after the second PGF_{2α}
84 treatment (**PreSynch-Ovsynch+ED**). The second program simulated was a PreSynch-Ovsynch
85 protocol for 100% TAI with ED incorporated after first TAI (**PreSynch-Ovsynch+EDpost**). In
86 these first two protocols, CR for PreSynch-Ovsynch was set at 35% (Caraviello et al., 2006),
87 Service Rate (SR) for ED was set at 60%, and CR for ED was set at 30% (Giordano et al., 2012;
88 Fricke et al., 2014). Programs 3 through 5 were Presynch-Ovsynch protocols for 100% TAI
89 which were simulated with varying CR to TAI (35%, 40%, 45%; Caraviello et al., 2006; Sousa
90 et al., 2008; Stangaferro et al., 2018). The sixth protocol simulated was a Double-
91 Ovsynch+PGF protocol (Carvalho et al., 2015; Wiltbank et al., 2015) in which CR was set at
92 50% (Souza et al., 2008; 2013; Santos et al., 2017). In all simulations, Ovsynch was used as the

93 resynch protocol, with a 30% CR to TAI (Lopes et al., 2013). These protocols were chosen to
94 reveal the difference in profitability between the most intensive program (i.e., a Double-
95 Ovsynch+PGF protocol) compared to a PreSynch-Ovsynch protocol. The protocols that
96 incorporated ED were selected to understand if incorporation of AI to a detected estrus during
97 these programs would be profitable if performed with PreSynch-Ovsynch, and this profitability
98 difference was calculated compared to a Double-Ovsynch+PGF protocol.

99 Because our objective was to compare the difference in hormonal costs among the
100 programs, all non-hormonal reproduction costs were set at \$0 in the UW-Cornell
101 DairyRepro\$ decision tool. The cost of each GnRH treatment was set at \$2.6, and the cost of
102 each PGF_{2α} treatment was set to \$2.3 to reflect cost of hormones in the US market (Giordano et
103 al., 2012). In addition, the cost of each GnRH treatment was set at \$6.7 and the cost of each
104 PGF_{2α} treatment was set to \$5.1 to reflect cost of hormones in the European market. Cost of
105 hormonal treatments for the European market were based on 11 values for the most common
106 commercial PGF_{2α} products (Cloprostenol 500 µg) and 5 values of the most common GnRH
107 (Gonadorelin 100 µg) products in the Italian market in November, 2018. Based on these costs,
108 the economic simulations were run to calculate the total net profit (\$/cow per yr) and the
109 aggregated hormonal cost of each program. The UW-Cornell DairyRepro\$ decision tool was
110 also used to calculate the number of hormonal treatments required per cow per yr for the
111 various protocols.

112 Using the PreSynch-Ovsynch protocol with a 35% P/AI as the baseline, the number of
113 hormonal treatments and net profit gain of the various reproductive management protocols was
114 compared (Table 1). Although PreSynch-Ovsynch protocols use fewer hormonal treatments
115 than a Double-Ovsynch+PGF protocol (1.4 to 3.0 fewer treatments per cow per yr); the

116 Double-Ovsynch+PGF protocol was more profitable. The Double-Ovsynch+PGF protocol
117 attained \$40.4 greater profit per cow per yr than the PreSynch-Ovsynch + ED protocol and
118 \$28.9 more than PreSynch-Ovsynch + EDpost protocol based on hormonal costs in the US
119 market. Also, the Double-Ovsynch+PGF protocol was more profitable than the 100% TAI after
120 a Presynch-Ovsynch protocol (\$21.2 to \$46.2 more per cow per yr, depending on CR; Table 1).
121 These results are directly related to the increased reproductive performance of a Double-
122 Ovsynch+PGF protocol (higher CR to first AI) compared to the other protocols. Furthermore,
123 inclusion of ED after the first TAI (i.e., the PreSych-Ovsynch + EDpost protocol) was a more
124 profitable strategy than using ED either before the first TAI (i.e., the PreSynch-Ovsynch + ED
125 protocol) or not incorporating ED at all based on 35% and 40% CR, respectively. This outcome
126 might result because fewer cows are submitted to a resynch protocol at a lower CR when
127 applying more intensive protocols for first TAI that have an increased conception rate. The
128 PreSynch-Ovsynch protocol with a 45% CR had the second greatest net profit among the
129 programs compared because of its higher CR counteracting the resynch cost. The gain in
130 profitability when switching to a Double-Ovsynch+PGF protocol based on US market prices
131 was greater (Table 1) because of the ratio of the costs between GnRH and PGF_{2α}, which was
132 113% in the US market compared to 131% in the European market.

133 The second comparison we made addressed the concern of whether a more intensive
134 synchronization protocol will still result in higher profitability than the less intensive
135 reproductive management protocols based on higher costs of hormones. We therefore
136 conducted sensitivity analyses by incrementally increasing the cost of hormonal treatments to
137 determine the breakeven point at which hormonal costs offset the net profit of 1) having the
138 costs of GnRH and PGF_{2α} at US and European market costs and raising both costs by multiples;

139 2) setting the cost of GnRH at European market average cost of \$6.7 and increasing the cost of
140 PGF_{2α}; and 3) setting the cost of PGF_{2α} at European market average cost of \$5.1 and increasing
141 the cost of GnRH. The ability to vary hormonal treatment costs in the UW-Cornell
142 DairyRepro\$ decision tool made it possible to simulate these scenarios. For this analysis, all
143 general costs such as labor for administering hormonal treatments, labor for transrectal
144 palpation for pregnancy diagnosis and insemination, and the non-studied variables like
145 lactation curve and other herd parameters, were kept as default in the UW-Cornell
146 DairyRepro\$ decision tool (Giordano et al., 2012). Briefly, labor cost for administering
147 hormonal treatments was set at \$15/hr, and labor for transrectal palpation for pregnancy
148 diagnosis was set at \$105/hr. The cost of insemination included semen cost of \$5/AI and a
149 labor cost of insemination at \$5/AI. Groups of simulations were run to compare the most
150 intensive Double-Ovsynch+PGF protocol with each of the other defined protocols compared in
151 this simulation. In each pair-group comparison, the cost of GnRH and/or PGF_{2α} treatments was
152 increased until the net profit became negative, then the breakeven point was identified as the
153 intercept of the trend line with the x-axis (Figure 1). These x-axis intercepts define the
154 hormonal treatment costs at which a Double-Ovsynch+PGF protocol would have an equal net
155 profit to the protocol it was compared against.

156 Based on our analysis hormonal treatment costs would need to be 5 to 14 times greater
157 in the US market and 2 to 6 times greater in the European market for any of the Presynch-
158 Ovsynch protocols to be more profitable than the Double-Ovsynch+PGF protocol. The greater
159 P/AI at first insemination after a Double-Ovsynch+PGF protocol compensates for the
160 additional hormonal treatment costs so that the cost of hormonal treatments would need to
161 increase substantially before it becomes less profitable than any of the other programs. When

162 the CR of Presynch-Ovsynch protocols increased, the breakeven point was reached earlier
163 because the advantage of the increased CR of Double-Ovsynch+PGFis was decreased.

164 To investigate the cost of one of the hormonal treatments to reach the breakeven point,
165 we ran hypothetical scenarios in which the cost of one hormone was constant but the cost for
166 the other hormone was incrementally increased. In this simulation, the cost of PGF_{2α} was set at
167 \$5.1/dose, and the cost of GnRH was set at \$6.7/dose (i.e., the average European market price).
168 Breakeven hormonal costs from this analysis are reported in Table 2.

169 Overall, the breakeven cost was more sensitive to the cost of GnRH than to the cost of
170 the PGF_{2α} (Table 2). When the cost of GnRH was fixed, the cost of PGF_{2α} could increase
171 considerably before the breakeven point was reached or no breakeven point could be reached
172 by the model. A complete Presynch-Ovsynch protocol uses 2 GnRH and 3 PGF_{2α} treatments,
173 whereas the Double-Ovsynch+PGF protocol uses 4 GnRH and 3 PGF_{2α} treatments with a
174 higher proportion of GnRH treatments (57% vs. 40%) and a lower proportion of PGF_{2α}
175 treatments (43% vs. 60%). If ED was not incorporated into the reproductive program and the
176 cost of GnRH was fixed, profit increased even when the cost of PGF_{2α} increased. Thus, the
177 breakeven points are not reached by the model when increasing the cost of PGF_{2α}. For example,
178 when compared with PreSyncOv 40 with the cost of PGF_{2α} fixed at \$5.1/dose, the cost of a
179 GnRH treatment was \$22.4/dose at the break-even point (Table 2). By contrast, in the first
180 study, one breakeven point was reached when the price was \$20.7/dose for PGF_{2α} and
181 \$23.4/dose for GnRH (Figure 1(a), 9 times the US market prices), indicating that a greater
182 PGF_{2α} cost determines an even higher cost of GnRH to reach the breakeven point. This can be
183 explained by the higher CR of the Double-Ovsynch+PGF protocol that leads to fewer cows
184 submitted to a resynch protocol. Consequently, fewer GnRH treatments per pregnancy are used

185 on average (Table 1). For protocols that incorporate ED, fewer cows are submitted to hormonal
186 treatments which decreases total hormone costs of the protocol; thus, a breakeven point was
187 reached even when the GnRH cost was fixed, but when the cost of PGF_{2α} was unusually high
188 (Table 2).

189 In conclusion, our economic evaluation found that more intensive reproductive
190 programs that use more hormonal treatments but result in substantially increased reproductive
191 performance are more profitable than less intensive programs and remain superior even if
192 hormonal prices are unusually high. Results from these analyses could be reproduced or
193 adjusted by applying the UW-Cornell DairyRepro\$ decision support tool that is openly
194 available at UW-Madison and Cornell University websites.

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299

300 **Table 1.** Comparison in the number of hormonal treatments and net profit between different

Reproductive Program ³	P/AI (%)	Approximated number of treatments ¹ (#/cow per yr)			Net Profit gain over the baseline ² (\$/cow per yr)	
		Total	GnRH	PGF _{2α}	PGF _{2α} at \$2.3 and GnRH at \$2.6 ⁴	PGF _{2α} at \$5.1 and GnRH at \$6.7 ⁴
PreSynch-Ovsynch (baseline)	35	7.8	3.12	4.68	-	-
PreSynch-Ovsynch	40	7.6	3.04	4.56	12.7	13.7
PreSynch-Ovsynch	45	7.4	2.96	4.44	25	26.7
PreSynch-Ovsynch + ED	35 + 30	6.2	2.48	3.72	5.8	8.2
PreSynch-Ovsynch + EDpost	35 + 30	6.3	2.52	3.78	17.3	22.8
Double-Ovsynch+PGF	50	9.2	5.24	3.96	46.2	32.1

301 reproductive synchronization programs.

302 ¹Approximated number of treatments for hormones. GnRH proportion in protocols: Presynch-
303 Ovsynch protocols: 40%; Double-Ovsynch+PGFprotocol: 57%.

304 ²Net profit gain over the baseline is the value of the net profit difference when alternating the
305 baseline protocol to the listed one.

306 ³**PreSynch-Ovsynch** (Presynch-Ovsynch protocol with 35%, 40%, 45% CR), **ED** (Estrus
307 detection performed before first AI service with 60% SR and 30% CR), **EDpost** (Estrus
308 detection performed after first AI protocol with 60% SR and 30% CR), **Double-Ovsynch**
309 **PG2x** (Double Ovsynch with a repeated injection at second prostaglandin with 50% CR).

310 ⁴PGF_{2α} at \$2.3/dose and GnRH at \$2.6/dose representing the US market and PGF_{2α} at \$5.1/dose
311 and GnRH at \$6.7/dose representing the European market.

312

314 **Table 2.** The cost (\$/dose) of GnRH or PGF_{2α} at breakeven profit points (bold numbers), when
 315 the other hormonal cost was set constant at European market price, comparing Presynch-
 316 Ovsynch programs against the most intensive synchronization program, the Double-
 317 Ovsynch+PGF.

Cost (\$/dose) at the breakeven point compared with Double-Ovsynch+PGF ¹										
Hormones	Presynch-Ovsynch ²						Presynch-Ovsynch (35% CR) + ED			
	35% CR		40% CR		45% CR		ED ³		EDpost ⁴	
GnRH	32.8	6.7	22.4	6.7	14.2	6.7	19.0	6.7	13.7	6.7
PGF _{2α}	5.1	-- ⁵	5.1	-- ⁵	5.1	-- ⁵	5.1	97.0	5.1	63.0

318 ¹Double-Ovsynch PG2x (Double-Ovsynch+PGF with 50% CR).

319 ²PreSynch-Ovsynch (Presynch-Ovsynch protocol with 35%, 40%, 45% CR).

320 ³ED (Estrus detection performed before first AI service with 60% SR and 30% CR).

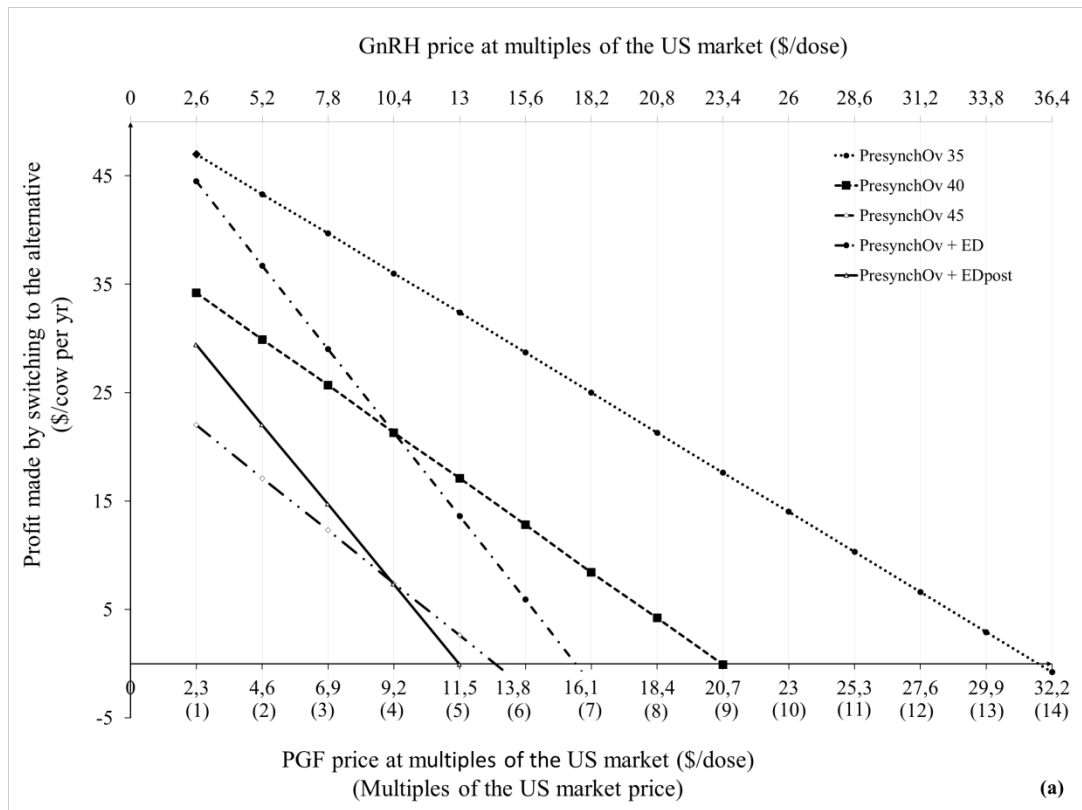
321 ⁴EDpost (Estrus detection performed after first AI protocol with 60% SR and 30% CR).

322 ⁵The breakeven point could not be reached.

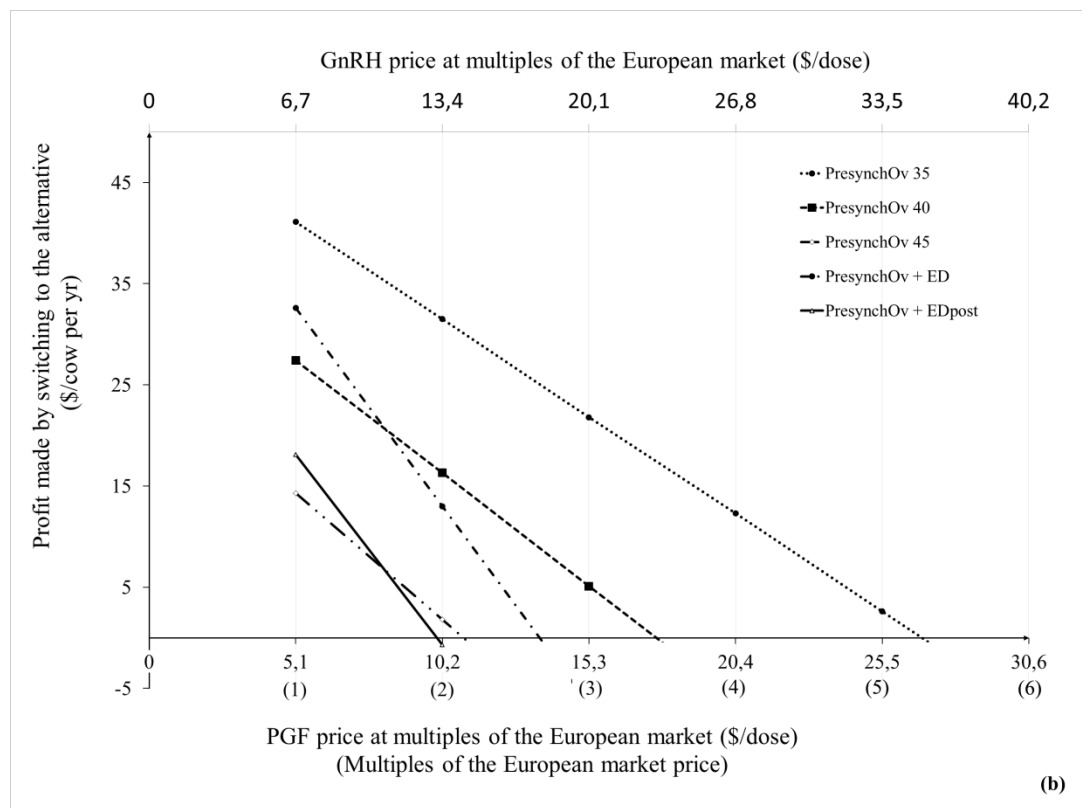
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325 **Figure 1.** Sensitive analysis by identifying the breakeven points when the net profit gain by
326 switching the Presynch-Ovsynch protocols to Double-Ovsynch+PGF protocol become negative
327 with multiples of GnRH and PGF_{2α} for US market price (a) and European market price (b).
328 PreSynchOv 35 (Presync-Ovsynch protocol with 35% CR)
329 PreSynchOv 40 (Presync-Ovsynch protocol with 40% CR)
330 PreSynchOv 45 (Presync-Ovsynch protocol with 45% CR)
331 PreSynchOv + ED (Presync-Ovsynch protocol with 35% CR + estrus detection before first AI
332 service with 60% SR and 30% CR)
333 PreSynchOv + EDpost (Presync-Ovsynch protocol with 35% CR + estrus detection after first AI
334 service with 60% SR and 30% CR)
335 Double-Ovsynch PG2x (Double Ovsynch with a repeated injection at second prostaglandin with
336 50% CR)
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