CASE REPORT

Desisting from gender dysphoria after 1,5 years of puberty suppression

Angela Rölver, Manuel Föcker, Georg Romer

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ABSTRACT

Background: Over the last decade medical care for transgender youth has improved. Gonadotropin-Releasing-Hormone-Analogues (GnRHa) stop and delay puberty and can relieve the distress of Gender Dysphoria (GD). Only a few adolescents treated with GnRHa desist from GD, thus systematic and in-depth investigations are missing.

Case Presentation: A case of peri pubertal onset gender dysphoria (12;4 years) is presented. It illustrates the desistance from GD after a 15-month-treatment with GnRHa. The blocking of puberty lead to a reduction of GD, the process of gender identity development still continued, leading to a non-binary outcome.

Conclusion: This case report illustrates that after treatment with puberty blockers desistance from gender dysphoria can occur. Further, GnRHa do not "the ongoing process of gender identity formation in psychosexual development. Thus, puberty suppression is a reversible treatment option that can be seen as a meaningful step to prepare the readiness for partially irreversible gender-affirming hormone treatment.

Key Words: Gender Dysphoria; Gender Incongruence; Gonadotropin-Releasing-Hormone-Analogues; Puberty blocking; Gender identity

INTRODUCTION

According to ICD-11 [1], gender incongruence (GI) refers to the mismatch of an individual's experienced gender and his or her birth-assigned sex which often leads to a desire to live and be accepted as a person of the experienced gender. If people suffer from gender incongruence or experience impairment in social, school or other important areas of functioning, Gender Dysphoria (GD) can arise. According to DSM-5 [2] adolescents with GD experience a distressing incongruence between their birth-assigned sex and the gender they identify with. The identification with the other gender and experienced GD can be accompanied by the desire to change the sexual characteristics of the physical body and to seek gender-affirming medical treatment (e.g. hormonal treatment, surgery).

Based on systematic reviews of past studies mainly based on utilization rates of specialized medical centres offering sex-reassigning interventions, the average prevalence rate of transgender adults has been estimated 6.8 of 100.000 persons [3,4]. However, this rate is likely to be biased by underestimation, because individuals with

gender incongruence who did not seek medical treatments were not detected as cases. In a recent population-based epidemiological survey from Sweden gender incongruence in adults was reported by 0.9% of legally males and 1.0% of legally females, whereas a strong desire to seek medical treatment for sex-reassignment was reported by 0,2% of the adult population across both sexes [5]. This finding represents a rate about 30 times higher than the average rate estimated from past reviews named above. For children and adolescents epidemiological studies are missing. In a New Zealandian study, 1.2% of a sample of high school students identified as transgender [6].

Over the last decades, medical care for transgender youth has changed including an increase of specialized health services. These clinics register an increasing demand accompanied by a shift in the gender-sex-ratio towards natal females [7]. Adolescents with GD often face various associated social, emotional and behavioral difficulties [8,9]. The most prevalent associated problems of gender dysphoric adolescents are bullying, depression, suicide attempts and self-harm [10]. Thus, GD is often associated with mental disorders, mostly

Department of Child and Adolescent Psychiatry, University Hospital Münster, Germany.

Correspondence:Department of Child and Adolescent Psychiatry, University Hospital Münster, Germany, Telephone +491753313993, e-mail angela. bieda@ukmuenster.de

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major depression and eating disorder [11].

The majority of prepubescent children with GI/GD has been reported not to have an outcome of persistent gender dysphoria after puberty [12, 13]. With regard to sexual preference, the mostly reported outcome of childhood GD has been (cis-gendered) homosexuality or bisexuality [14]. Persistence of GD has most likely been predicted by the intensity of GD in childhood ('conviction' vs. 'wish') and the amount of reported cross-gendered behaviors. However, despite likelihood there is no certain predictor to discriminate between later 'desisters' and 'persisters' before the onset of puberty [12]. On the other hand, GD that persists from childhood into adolescence is likely to further persist into adulthood [15]. Dutch longitudinal clinical follow-up studies on adolescents with childhood GD who received puberty suppression and/or gender affirming hormones after comprehensive biopsychosocial assessment demonstrated that none of the participants desisted from gender affirming surgery one year later [16,17]. Some adolescents have not experienced childhood GD, thus any predictions about stability are harder to make [18]. Furthermore, there is a growing number of adolescents who identify as non-binary [19]. Given the various developmental pathways for children with GD and the dynamic nature of gender identity in adolescence an individualized approach in clinical care has to be taken.

Best clinical practice for child and adolescent GI/GD

According to international standards of care, clinicans should explore gender identity openly and exploratively. Treatment strategy focuses on constant and supportive counseling of children and parents, helping to create a safe and accepting environment for children and adolescents where they can unfold their gender identity. The affirmative approach encourages children and adolescents to transition to the gender role that fits with his or her subjective identity. This includes for adolescents the option of medical interventions to facilitate gradual and age-appropriate developmental transitions, particularly the option of endocrine treatment, both by suppressing puberty and by subsequent gender-affirming hormone treatment. Medical treatment options should be offered to patients step by step after careful clinical assessments and re-assessments [20]. Puberty suppression as a fully reversible intervention can be seen as an important step to prepare the readiness for gender-affirming hormone treatment both in the gender dysphoric adolescent and his or her social environment [21]. It can also be helpful in cases when further time is needed for an adolescent with GD to explore his or her gender identity. A shared decision making process for the treatment options is recommended, clinicians should evaluate the adolescent's capacity for consent, helping the adolescent to reach the required capabilities for an informed consent [22].

It has been acknowledged in international standards of care that particularly in adolescent GD 'just waiting' cannot be considered a neutral option, because it can create additional harm due to ongoing and irreversible sexual maturation [8,20]. The German Ethical Council has stated in 2020 that in individuals with GD after the onset of puberty the potential consequences of active medical interventions (e.g. puberty blocking) have to be carefully weighed against the potential consequences of not intervening. Clinicians have to carefully consider potential health benefits and risks of both

medical intervening and not (yet) intervening by balanced reasoning in any individual case [22].

Puberty suppression with Gonadotropin-Releasing-Hormone-Analogues (GnRHa)

Current international treatment guidelines recommend GnRHa as an option to attenuate the distress of GD by reducing the suffering caused by the development of secondary sexual characteristics in puberty [20,23,24]. The temporary halting of pubertal maturation relieves distress caused by masculinizing or feminizing body changes (e.g. puberty vocal change in trans girls or menstruation in trans boys). This offers a limited period of time for gender dysphoric adolescents to reflect their gender identity and for mental health professionals to support self-exploration and reflections on other associated developmental problems [25,26]. Gender dysphoric adolescents showed an improved psychological functioning and better social functioning after temporary puberty suppression [11,27]. In a recent study, Sorbara and colleagues [28] found that late pubertal stage in gender dysphoric youth were associated with more mental health problems. Concerning the biological maturation the treatment with GnRHa is fully reversible.

There is a paucity of studies of patients who detransition after intake of GnRHa. In the study of Brik and colleagues [29], 5 adolescents (3.5%) did not continue with gender affirming treatment due to various reasons. The effect on gender identity development was not further explicated. The percentage of discontinuation of GnRHa is small (1,9%-3,5%), thus systematic studies are missing [30]. These studies were conducted in specialized gender clinics which include a comprehensive assessment before medical treatment is started, thus the number of discontinuation can be underestimated. In an opposing position to current international treatment guidelines, some authors argued that puberty suppression by itself may increase the likelihood of persistence because gender dysphoric adolescents may be alienated from the chance to "reconcile" with their "original" gender by gradually adjusting to their sexually maturing bodies [31,32]. A qualitative study by Vrouenrats and colleagues [33] found that there is a lively debate about pros and cons about puberty suppression among mental health professionals working in GD treatment teams which shows that there a still open questions concerning the use GNRHa and a gap between guidelines and the current practice.

The above mentioned opposing position was endorsed by a recent public debate concerning a decision of the High Court of Justice in England and Wales in 2020 who had ruled that minors under 16 would not be competent to give legal consent to the administration of GnRHa because they were not able to oversee the long-term consequences of this treatment. This court had further ruled that puberty blockers should only be applied between the age of 16 and 18 with the permission by a Family Court. Meanwhile, this court decision has been overruled by the British Court of Appeal, which clarified that decisions for or against endocrine treatment options in gender dysphoric minors should fully stay within the responsibility of physicians given that there is valid informed consent by patients and – if applicable - their legal caregivers. In clinical practice a shared and fully informed co-consent of gender-dysphoric minors and their legal caregivers are recommended both for legal and psychological reasons

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[34]. This informed co-consent is to be developed together with health professionals in a careful process of shared decision making [22].

CASE PRESENTATION

In 2017, CS (12;4 years) was seen for the first time in our specialized gender outpatient service. See figure 1 for the timeline of episode of care.



Figure 1. Timeline of episode of care

Being female by birth-assigned sex, he introduced himself as a boy and reported gender dysphoric feelings which had started six months ago as well as the desire to live in the male gender role. Follow-up appointments together with his parents were made on a lowfrequency basis (every two months) to jointly explore the course of this recent onset complaints. After 9 months of clinical observation the diagnosis of GD was given according to DSM 5 criteria. In October 2017 CS completed his female-to-male social transition in all important areas of everyday life. In January 2018 he additionally started high-frequent psychotherapy with weekly sessions with a psychotherapist. He suffered from conflicts with his parents and started to injure himself. In March 2018, after a 12-month period of clinical observation and after his completed social transition, treatment with GnRHa was started based on child-parent co-consent to halt female maturation from which CS was suffering enormously. In November 2019 (age 15:0) the attending psychotherapist recommended an inpatient admission for more intensive psychotherapy because of the aggravation of CSs depressive symptoms and upcoming doubts about his transgender transition.

At hospital admission CS reported various depressive symptoms: he experienced permanent low mood and most of the times he felt tired and exhausted and had no motivation to engage in things or hobbies he had enjoyed before. He described a feeling of hopelessness and repetitive thoughts mainly about his gender transition.

Along with these ruminating cognitions he had difficulties in making decisions in his daily life and he had withdrawn almost completely from his social life. He reported self-injuring by cutting and scratching himself over a period of already 2,5 years. He experienced suicidal ideations, however, negated suicide attempts or concrete suicidal plans.

Family, development and social history.

CSs parents are married and are working in the field of public administration. He has a younger sister (8 years) with whom he got along well. CS attended secondary school, in the last year his grades worsened. Psychological disorders are unknown in family history.

Physical examination and laboratory investigations.

The physical examination showed a good state of health. The physical appearance was spruce and body shape was slim (weight $49.5~\mathrm{kg}$;

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height 168.5 cm; BMI 17.4 kg/m 2 [7P, -1.45z, female norms]). Laboratory tests were within normal limits, no somatic or endocrinological diseases.

Psychological tests at admission.

In the Becks Depression Inventory (BDI) [35] CS showed a total score of 22, representing a moderate depression at admission.

The Eating Disorder Inventory-2 (EDI-2) [36] is standard for the assessment of attitudinal and behavioral dimensions relevant to anorexia and bulimia nervosa. CS had elevated scores (PR >/= 75 < 90) on the subscales body dissatisfaction, ineffectiveness, interpersonal distrust and maturity fears and clinically relevant sores (PR >/= 90) on the subscales drive for thinness, bulimia, interoceptive awareness and asceticism.

The Social Phobia and Anxiety Inventory for Children (SPAI-C)[37] is a 26-item self-report instrument that measures a range of potentially anxiety-provoking situations as well as symptoms associated with social phobia. CS showed elevated scores in social interactions and performance situations (PR = 96-99 and t-value= 68-73).

The Life Problems Inventory (LPI) [38] measures borderline features in adolescents. CS showed elevated scores on the subscales identity confusion and emotion dysregulation.

Diagnosis and treatment

At admission the diagnostic criteria of a major depressive disorder, single episode, moderate (F32.1) and GD (according to DSM 5) were met.

CS was admitted at our psychotherapy ward for adolescents (focus on emotion regulation) and received over the course of 14 weeks individual psychotherapy sessions (twice a week), family sessions (every second week), music- and art therapy (every week) as well as multiple pedagogical group and individual interventions focusing on emotional and social competencies. The therapeutic setting was based on the dialectical behavior therapy for adolescents [39].

At the beginning of treatment CS was timid and reluctant, avoided contacts with peers. During the course of treatment he gradually built trust, initiated conversations with the staff and peers and was liked by them.

During individual psychotherapy sessions it was easier for CS to open up because of his prior experiences in psychotherapy. Based on a CBT-approach, first a behavioral analysis was undertaken and then an individual biopsychosocial model for understanding his depression and gender dysphoria was set up with a focus on the evolvement and maintenance of symptoms. From a biological perspective, CS reported that his menstruation began very early (with 10 years) and that he experienced it as a drastic and overwhelming event. He could not accept that his body was changing, he did not like either his breasts nor voice. Furthermore, he reported that he learned in a documentary on television that a low weight can cause amenorrhea which was why he set his personal weight gain limit up at 50kg.

From a social perspective, CS reported that at the age of 11 he had a massive conflict with female friends which led to his exclusion from the girls' peer group. Then he joined a boys peer group and felt relieved from the conflict. One year later in summer vacation 2016 he started to ruminate extensively and concluded for himself that a boy would not have that kind of social problems and a menstruation. After that summer vacation, at the age of 14,8 he disclosed his gender problem to his teacher and parents. He cutted his hair short and wore

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loose clothes. In spring 2017, he started transitioning socially, got himself a male name. The social transition was accompanied by a huge relief, resulting in better mood and motivation. In the course of 2018 he started ruminating about his transition, accused himself for making the wrong decision. The rumination came along with strong feelings of insecurity regarding his further proceeding (beginning of cross-sex hormone treatment or not) and self-hate, which worsened the self-injury. Further, he began to feel insecure about daily decisions and therefore withdraw from family, friends and avoidance of activity, was passive in many life areas.

In the inpatient therapeutic milieu CS was able to talk frankly about the development and maintenance factors of his depressive and gender dysphoric feelings and about his fear of gaining weight and physical maturation. These factors were identified and addressed in therapy. Finally after a shared decision making process CS decided to stop puberty suppression which was recommended from a professional perspective. The decision caused CS major distress because he was afraid not to have any control about the physical change of his body, especially the menses. He also feared a weight gain and for a short period of time started restrictive eating and lost weight. The restrictive eating was treated with psychoeducation and an eating protocol. See figure 2 for the weight curve over the course of treatment from week 1 to week 11.

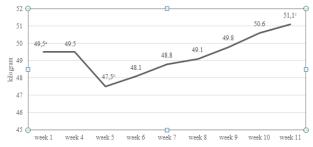


Figure 2. Course of body weight. At admission CS had a BMI 17.4 kg/m² (7P, -1.45z)(a), the lowest BMI was 18.1 kg/m^2 (5P, -1.61z) in week 5 (b), at hospital discharge BMI was 18.1 kg/m^2 [18P, -0.93z] (c).

Over the course of therapy he gained insight in dysfunctional thoughts, emotions and behaviors. At the end of the treatment he accepted his female-sex body, identified himself as non-binary and kept his male name and male pronoun . He did not regret the developmental pathway of the past two years anymore, which he could appreciate as a necessary diversion of finding out about his gender identity. He did not feel easy with his menses, but accepted and coped with it without major suffering from it anymore. He continued psychotherapy on a weekly basis.

CSs parents supported the treatment, in family therapy sessions CS increasingly opened up to his parents, talking more frankly about his emotional problems and his attempts to cope with them.

At hospital discharge the diagnosis of a mild depressive disorder was still present (F32.0). The diagnosis of GD was changed into sexual maturation disorder (F66.0).

Follow-Up- 1,5 years later

At a follow-up assessment CS reported that in summer 2020 she

decided to alter her name and pronouns back to female. She reported that she would define her gender as non-binary. Since hospital discharge she still struggled with her menses, thus she is taking now a gestagen drug to suppress it. CS mentioned that she still had depressive thoughts and behaviors from time to time but could manage her daily life. She could not report her actual weight, however, restrictive eating did not occur anymore. She did not need psychotherapy sessions anymore. Regarding the GnRHa treatment she does not regret anything, she rather sees it as part of her process of finding out her own gender identity. CS's mother confirmed CS's statement.

Psychological tests

In the BDI [35] CS showed a total score of 11, representing a mild depression.

The subscales of the EDI-2 [36] were in the normal range, except the subscale maturity fears ((PR >/= 90).

CS showed on the SPAI-C no elevated scores in social interactions and performance situations (PR = 26-50 and t-value= 44-50).

Regarding the LPI [38], all scores were in the normal range for adolescents.

DISCUSSION

Systematic studies and in-depth studies on the phenomenon of desistance from GD after the beginning of puberty suppression in adolescence are lacking. Little is known about the psychological processes behind such desisting outcomes. This case report illustrates that after the beginning of treatment with puberty blockers desistance from GD can occur and that gender identity is often more dynamic and less binary than supposed. This supports the hypothesis that the development of gender identity based on ongoing self-exploration and reflection is not "automatically" impaired or even haltered by suppressing the secretion of sex hormones with GnRHa. Rather psychosexual development including gender identity development can progress despite suppressed sex hormone levels. On the one hand, this case illustrates the potential benefit of puberty suppression as a medically reversible treatment option in cases of GD in early adolescence, in which later desistance may occur in spite of careful and sufficient clinical assessment before starting this treatment. On the other hand, this case underlines the necessity of careful assessment and profound information of young patients with GD and their parents or legal guardians before starting endocrine treatment. This includes information about the possibility of later desistance, as well as diverse outcomes, in particular not only binary ones. A non-binary gender identity appears to be becoming more common among adolescents presenting at gender clinics [40]. Nonbinary youths are highly vulnerable and have important health care needs [19]. For health professionals, it underlines the necessity of careful case monitoring and reassessment after starting treatment with puberty blockers in gender dysphoric youths and before starting partially irreversible interventions, such as gender affirming hormone treatment.

In the case of CS, the ongoing pubertal maturation with increasing feminizing bodily features led to major distress and psychological symptoms. The gender dysphoric and depressive symptoms decreased under the treatment with puberty blockers, CS felt temporarily relieved. This observation is in line with findings that early medical

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interventions improve the psychosocial functioning and wellbeing in gender dysphoric adolescents [16,41]. In the vast majority of cases, in which GD persists into adulthood, the life-long mental health related gain of preventing irreversible masculine or feminine body features can be considered substantial. However, in CS's case, a medically caused "irreversible consolidation" of the transgender pathway by the endocrine intervention of administering puberty blockers was not observed. CS could make use of a reflecting "moratorium" with her maturing body to find out that the perspective of masculinization of her body appearance through gender-affirming hormone treatment would be incongruent with her gender identity. CS expressed feelings of doubt, fear and shame regarding the option of social retransition, however, she was able to reflect this as coping with her process of gender identity development. In her own retrospect the use of GnRHa 'had bought' time which enabled her to further explore her gender identity and to critically question her desire for somatic sexreassigning procedures. Through psychological treatment it was possible to support CS in her individual decisions. She could develop a perspective in which she did not regret her "transitional transition", rather seeing it as a coherent process of her gender identity formation. Thus, the conclusion of this case report is in line with the WPATH standards of care [20] which suggest an affirmative treatment approach for GD in adolescents in which medical treatment options should be offered to patients step by step after careful clinical assessments and re-assessments.

However, the ethical dilemma remains in any treatment decisions with gender diverse youths. Potential long term consequences of medical interventions in cases of later desistance from GD have to be carefully weighed against potentially harmful consequences of irreversible masculinization or feminization of the bodily appearance on the mental health of gender dysphoric youths. To improve security in treatment decisions, longitudinal follow-up studies on larger cohorts of transgender adolescents treated with GnRHa are needed. Furthermore, this case reminds us that gender cannot be understood as a binary concept. In the treatment and counseling of children and adolescents with GI or GD gender variety and multiple pathways should be considered. The primary goal of treatment should be to promote an individual person's mental health and well-being and not a binary identification as male or female.

CONCLUSION

To sum up, this case study demonstrates that under the treatment with puberty blockers, desistance from GD in adolescence can occur. It further demonstrates that GNRH analogues, while blocking sex hormone secretion, do not at the same time "block" the ongoing process of gender identity development. This case report supports the standards in current treatment guidelines [20,23], in which puberty suppression with GnRHa can be offered as a treatment option for minors with GD after the first stages of puberty (minimum Tanner stage 2) have been reached and if gender-affirming hormone treatment cannot be recommended yet. This treatment should be based on a careful comprehensive assessment as well as fully informed consent by both the young patients and their parents or legal guardian.